

ACCELERON PHARMA INC  
Form 424B4  
January 23, 2014

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Filed Pursuant to Rule 424(b)(4)  
Registration Nos. 333-193252, 333-193495

**PROSPECTUS**

**2,400,000 Shares**

**Common Stock**  
**\$50.00 per share**

We are selling 2,400,000 shares of our common stock.

We have granted the underwriters an option to purchase up to 360,000 additional shares of common stock.

Our common stock is listed on the NASDAQ Global Market under the symbol "XLRN". On January 22, 2014, the last sale price our common stock was \$56.70 per share.

We are an "emerging growth company" as that term is used in the Jumpstart Our Business Startups Act of 2012 and, as such, have elected to comply with certain reduced public company reporting requirements for this prospectus and future filings.

**Investing in our common stock involves risk. See "Risk Factors" beginning on page 11.**

**Neither the Securities and Exchange Commission nor any state securities commission has approved or disapproved of these securities or passed on the adequacy or accuracy of this prospectus. Any representation to the contrary is a criminal offense.**

Per share

Total

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Public Offering Price	\$50.00	\$120,000,000
Underwriting Discounts and Commissions(1)	\$ 3.00	\$ 7,200,000
Proceeds to Acceleron (before expenses)	\$47.00	\$112,800,000

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(1) We refer you to "Underwriting" beginning on page 154 for additional information regarding underwriting compensation.

The underwriters expect to deliver the shares of common stock to investors on or about January 28, 2014 through the book-entry facilities of The Depository Trust Company.

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**Citigroup**

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**Leerink Partners**

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**Piper Jaffray**

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**JMP Securities**

January 22, 2014

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**We are responsible for the information contained in this prospectus and in any free-writing prospectus we prepare or authorize. We have not authorized anyone to provide you with different information, and we take no responsibility for any other information others may give you. We are not, and the underwriters are not, making an offer to sell these securities in any jurisdiction where the offer or sale is not permitted. You should not assume that the information contained in this prospectus is accurate as of any date other than the date on the cover of this prospectus.**

## Trademarks

We own or have rights to trademarks, service marks and trade names that we use in connection with the operation of our business, including our corporate name, logos and website names. Other trademarks, service marks and trade names appearing in this prospectus are the property of their respective owners. The trademarks that we own include Acceleron Pharma®. Solely for convenience, some of the trademarks, service marks and trade names referred to in this prospectus are listed without the ® and ™ symbols, but we will assert, to the fullest extent under applicable law, our rights to our trademarks, service marks and trade names.

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**SUMMARY**

*This summary highlights information contained in other parts of this prospectus. Because it is only a summary, it does not contain all of the information that you should consider before investing in shares of our common stock and it is qualified in its entirety by, and should be read in conjunction with, the more detailed information appearing elsewhere in this prospectus. You should read the entire prospectus carefully, especially "Risk Factors" and "Management's Discussion and Analysis of Financial Condition and Results of Operations", before deciding to buy shares of our common stock. Unless the context requires otherwise, references in this prospectus to "Acceleron", "we", "us" and "our" refer to Acceleron Pharma Inc.*

**Overview**

We are a clinical stage biopharmaceutical company focused on the discovery, development and commercialization of novel protein therapeutics for cancer and rare diseases. Our research focuses on the biology of the Transforming Growth Factor-Beta (TGF- $\beta$ ) protein superfamily, a large and diverse group of molecules that are key regulators in the growth and repair of tissues throughout the human body. We are leaders in understanding the biology of the TGF- $\beta$  superfamily and in targeting these pathways to develop important new medicines. By coupling our discovery and development expertise, including our proprietary knowledge of the TGF- $\beta$  superfamily, with our internal protein engineering and manufacturing capabilities, we have built a highly productive research and development platform that has generated innovative clinical and preclinical protein therapeutic candidates with novel mechanisms of action.

We have three internally discovered protein therapeutic candidates that are currently being studied in numerous ongoing Phase 2 clinical trials, focused on cancer and rare diseases. These differentiated protein therapeutic candidates have the potential to significantly improve clinical outcomes for patients.

**The Acceleron Discovery and Development Platform: Novel Approaches to Potent Biology**

We focus on discovering and developing protein therapeutics that target a group of approximately 30 secreted proteins, or ligands, that are collectively referred to as the TGF- $\beta$  superfamily. These ligands bind to subsets of 12 different receptors on the surface of cells, triggering intracellular changes in gene expression that guide cell growth and differentiation. The TGF- $\beta$  superfamily ligands and their receptors represent a diverse and under-explored set of drug targets with the potential to yield therapeutics that modulate the growth and repair of diseased cells and tissues.

Members of the TGF- $\beta$  superfamily are now recognized as important regulators of red blood cell formation. We have shown that inhibition of members of the TGF- $\beta$  superfamily ameliorates anemia in mouse models of  $\beta$ -thalassemia and myelodysplastic syndromes (MDS). These red blood cell disorders are generally unresponsive to currently approved drugs. Based on our findings, we are developing two protein therapeutic candidates, sotatercept and ACE-536, each of which is currently in Phase 2 clinical trials to treat patients with these diseases.

Members of the TGF- $\beta$  superfamily also play a significant role in regulating blood vessel formation. We and our academic collaborators have shown that mice with a defect in a particular receptor for members of the TGF- $\beta$  superfamily are resistant to tumor growth due to reduced blood vessel formation in the tumor. We have used this insight to design our anti-angiogenic agent, dalantercept, which is currently in Phase 2 clinical trials for the treatment of cancer.

**Sotatercept and ACE-536: Novel Protein Therapeutic Candidates in Phase 2 Clinical Trials for  $\beta$ -thalassemia and MDS**

Together with our collaboration partner, Celgene Corporation, we are developing sotatercept and ACE-536, our lead protein therapeutic candidates, to treat anemia and associated complications in patients with  $\beta$ -thalassemia and MDS. Clinical trials are underway in other diseases as well.

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Sotatercept and ACE-536 have already shown promising biological activity in initial clinical trials. We and Celgene have conducted six clinical trials with sotatercept in over 160 healthy volunteers and cancer patients. We have conducted one clinical trial with ACE-536 in healthy volunteers. In these studies, both sotatercept and ACE-536 caused a dose-dependent increase in the number of red blood cells. Based on these results, we and Celgene have initiated Phase 2 clinical trials with each of these protein therapeutic candidates in  $\beta$ -thalassemia and MDS. We and Celgene plan to initiate Phase 3 clinical trials for one or both of these protein therapeutic candidates in one or both of  $\beta$ -thalassemia and MDS by the end of 2014 or early 2015.

**$\beta$ -thalassemia**

$\beta$ -thalassemia is a hereditary disease arising from defects in genes involved in the production of hemoglobin, the protein responsible for carrying oxygen in red blood cells. During red blood cell formation in the bone marrow, these genetic defects cause most of the cells to die before they mature into fully functional red blood cells. As a consequence, patients with  $\beta$ -thalassemia have anemia, a lower than normal number of red blood cells, and many patients experience a broad array of complications arising from their disease, including an enlarged spleen, skeletal deformities and serious organ damage, such as liver fibrosis and heart failure, resulting from the accumulation of iron. There is no approved drug and no effective drug therapy for the anemia of  $\beta$ -thalassemia. Frequent blood transfusions are used to manage the treatment of anemia in patients with  $\beta$ -thalassemia, but further contribute to the accumulation of iron and associated organ toxicities.

We and Celgene have shown that sotatercept and ACE-536 increase the production of red blood cells by promoting their maturation in the bone marrow. We believe this mechanism of action may be particularly beneficial for patients suffering from diseases, such as  $\beta$ -thalassemia, that are characterized by diminished red blood cell maturation. In a mouse model of  $\beta$ -thalassemia, the mouse version of ACE-536 demonstrated broad disease modifying effects. In this model, the mouse version of ACE-536 increased red blood cell production, reduced spleen size, increased bone density and reduced levels of iron in the kidney and liver.

The Thalassaemia International Federation estimates that there are approximately 300,000 patients worldwide with  $\beta$ -thalassemia, approximately 20,000 of which are in the United States and Europe, who are dependent on frequent blood transfusions. We estimate that there are at least as many  $\beta$ -thalassemia patients who do not receive frequent blood transfusions. Many of these patients have hemoglobin levels that are approximately half that of normal individuals and experience significant complications from the disease.

**Myelodysplastic Syndromes (MDS)**

MDS are a group of heterogeneous hematologic diseases characterized by abnormal proliferation and differentiation of blood precursor cells, including red blood cell precursors, in the bone marrow. This leads to anemia, which is present in the vast majority of MDS patients at the time of diagnosis. Much like the anemia of  $\beta$ -thalassemia, the anemia of MDS is characterized by an over-abundance of early stage red blood cell precursors, a large proportion of which fails to mature into functional red blood cells during the later phases of the red blood cell formation process. Drugs that stimulate the production of early stage red blood cell precursors, such as recombinant erythropoietin, are often used to treat anemia in MDS patients, yet many do not experience a substantial improvement of their anemia with these drugs. Although not approved by the United States Food and Drug Administration (FDA) for use in patients with MDS, these products generate an estimated \$500 to \$700 million in annual U.S. sales from use in these patients, according to our market research.

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**Additional Opportunities for Sotatercept and ACE-536**

Although sotatercept and ACE-536 have similar effects on red blood cells, sotatercept has been shown to increase bone mass and biomarkers of bone growth in humans. Many patients with chronic kidney disease suffer from both anemia and bone loss. Celgene is conducting two Phase 2 clinical trials of sotatercept in patients with chronic kidney disease-mineral and bone disorder. Additionally, we have shown that sotatercept inhibits tumor growth in mouse models of multiple myeloma, a cancer of the bone marrow, and sotatercept is being studied in an investigator-sponsored Phase 2 trial in multiple myeloma patients. Celgene and its collaborators continue to explore sotatercept in additional clinical trials including trials in patients with Diamond-Blackfan anemia and myelofibrosis.

Acceleron and Celgene are exploring the preclinical activity of sotatercept and ACE-536 in other red blood cell disorders including sickle cell disease.

**Our Partnership with Celgene**

We are developing sotatercept and ACE-536 through our exclusive worldwide collaborations with Celgene. As of January 1, 2013, Celgene became responsible for paying 100% of worldwide development costs for both programs. Additionally, we may receive up to \$560.0 million of potential development, regulatory and commercial milestone payments and, if these protein therapeutic candidates are commercialized, we will receive a royalty on net sales in the low-to-mid 20% range. If approved, we also will co-promote sotatercept and ACE-536 in North America, for which our commercialization costs will be entirely funded by Celgene.

**Dalantercept: Novel Protein Therapeutic Candidate in Phase 2 Clinical Trials for Cancer**

Our third clinical stage protein therapeutic candidate, dalantercept, is designed to inhibit blood vessel formation in tumors through a mechanism that is distinct from, and potentially synergistic with, vascular endothelial growth factor (VEGF) pathway inhibitors, the dominant class of cancer drugs that inhibit blood vessel formation. The VEGF pathway inhibitors collectively generate worldwide sales in excess of \$8 billion annually. We are developing dalantercept primarily for use in combination with these successful products to produce better outcomes for cancer patients.

**Inhibiting Angiogenesis to Limit Tumor Growth**

Angiogenesis is a process by which new blood vessels are formed. Angiogenesis can be simplified to two major stages the proliferative stage followed by the maturation stage. During the proliferative stage, vascular endothelial cells, the cells lining the inside of the blood vessels, increase in number. This proliferative stage is followed by the maturation stage during which the endothelial cells coalesce to form tubes which are then stabilized through the recruitment of perivascular cells that form an outer layer of the blood vessels resulting in fully formed, functional vessels.

Tumors depend on angiogenesis to form new blood vessels that supply nutrients and oxygen to feed the rapidly growing malignant cells. The principal molecule driving the proliferative stage of angiogenesis in tumors is a protein called VEGF. Inhibiting VEGF-driven angiogenesis to control tumor growth has become an important and widely-used approach to cancer treatment. There are several FDA-approved cancer drugs that inhibit the VEGF pathway. Despite the success of these drugs, many patients fail to respond or develop resistance to VEGF pathway inhibitor therapy, resulting in an unmet need for new therapies to inhibit angiogenesis by a different mechanism.

We are using our knowledge of the TGF- $\beta$  superfamily to develop dalantercept, a novel protein therapeutic candidate targeting the maturation stage of angiogenesis. Recently, the activin receptor-like kinase 1 (ALK1) has been recognized as an important regulator of the maturation stage of angiogenesis. ALK1 is one of the 12 receptors for ligands in the TGF- $\beta$  superfamily and is found

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primarily on endothelial cells. The importance of the ALK1 pathway in angiogenesis was discovered in part through research into a genetic disease in which patients manifest vascular defects, including a reduced ability to form capillary beds, which are the networks of small blood vessels that connect arteries to veins and are necessary for nutrient and waste exchange in tissues. This research revealed that these patients have only one of two functional copies of the ALK1 gene. The resulting decreased signaling through the ALK1 receptor inhibits blood vessel maturation, leading to the reduced formation of capillary beds.

**Opportunities for Dalantercept**

We reasoned that leveraging the biology of the ALK1 pathway to inhibit maturation of blood vessels could impair the growth of tumors by limiting the development of capillary beds within the tumor. To test this hypothesis, mice with a predisposition to develop tumors were bred to have only one copy, rather than two copies, of the ALK1 gene that normally occur. In response to the loss of half of the ALK1 genes, tumor growth and size and blood vessel density in the tumor were reduced by half. We have also shown in two mouse cancer models that treatment with dalantercept decreases metastases. This is in contrast to VEGF pathway inhibitors, many of which have been shown to increase metastases in mouse cancer models. These results and additional research in the field have established the ALK1 signaling pathway as a promising target for developing a new class of anti-angiogenesis agents, ALK1 pathway inhibitors. We are developing dalantercept to treat cancer by inhibiting the ligands of the TGF- $\beta$  superfamily that signal through the ALK1 receptor.

We believe one promising opportunity for dalantercept will be its use in combination with VEGF pathway inhibitors because these agents target distinct sequential steps in tumor angiogenesis. Moreover, we believe that dalantercept sensitizes blood vessels to increase the effects of treatment with VEGF pathway inhibitors. A combination of ALK1 and VEGF pathway inhibitors could have application in a number of different oncology indications where VEGF pathway inhibitors are currently used, such as liver cancer, brain cancer, non-small cell lung cancer, colorectal cancer and renal cell carcinoma.

With respect to our third clinical stage protein therapeutic candidate, dalantercept, we have conducted a single agent Phase 1 clinical trial in patients with advanced solid tumors. Additionally, we have studied the single agent activity of dalantercept in a Phase 2 clinical trial in patients with advanced head and neck cancer. Our ongoing focus is on the use of dalantercept in combination with an approved VEGF pathway inhibitor where we have both a mechanistic rationale and supportive preclinical data demonstrating dalantercept in combination with a VEGF pathway inhibitor provides enhanced anti-tumor effects in mice bearing human renal cell carcinoma xenographs. In an ongoing Phase 2 clinical trial of dalantercept in combination with axitinib, an approved VEGF pathway inhibitor, in patients with advanced renal cell carcinoma we have completed the dose escalation stage. We have now initiated the dose expansion phase of this study and plan to start the randomized controlled part of the study at the end of Q1 or early Q2 2014. We also intend to initiate a Phase 2 clinical trial of dalantercept in combination with the VEGF pathway inhibitor sorafenib in patients with liver cancer in the first half of 2014.

We have not entered into a partnership for dalantercept and retain worldwide rights to this program.

**ACE-083: Neuromuscular Disorders**

In addition to our clinical stage programs, we are developing a protein therapeutic candidate, ACE-083, for a first-in-human clinical trial that we expect to initiate by the end of 2014. ACE-083 has been designed to promote muscle growth in those muscles in which the drug is injected, with minimal systemic effect. We are focused on the development of ACE-083 for diseases in which increases in the size and function of specific muscles may provide a clinical benefit, including inclusion body myositis, facioscapulohumeral dystrophy (FSHD) and disuse atrophy.

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**Our Development Pipeline**

The status of our three clinical stage protein therapeutic candidates and our most advanced preclinical candidate is summarized below:

**Our Strategy**

Our goal is to be a leader in the discovery, development and commercialization of novel protein therapeutics for cancer and rare diseases. Key components of our strategy are:

***Advance sotatercept and ACE-536 into Phase 3 trials in collaboration with Celgene.*** We and Celgene are jointly developing sotatercept and ACE-536. Assuming successful completion of the ongoing Phase 2 clinical trials in  $\beta$ -thalassemia and MDS, we plan to initiate Phase 3 clinical trials with Celgene for one or both protein therapeutic candidates in one or both diseases by the end of 2014 or early 2015.

***Explore new indications for sotatercept and ACE-536 with Celgene.*** We and Celgene are continuing our preclinical research to assess the opportunity for sotatercept and ACE-536 to treat certain red blood cell disorders known as hemoglobinopathies, which include diseases such as thalassemias and sickle cell disease. Based on our encouraging preclinical and clinical data in  $\beta$ -thalassemia and our emerging understanding of the mechanism of action of these protein therapeutic candidates, we believe there is a potential for activity for sotatercept and ACE-536 in sickle cell disease, and we continue to explore development of these protein therapeutic candidates for this disease.

***Advance dalantercept into Phase 3-enabling clinical trials.*** Beyond our ongoing Phase 2 clinical trials, in 2014, we plan to initiate additional clinical trials of dalantercept in combination with either an approved anti-angiogenesis agent or chemotherapy in advanced solid tumors. One of these trials is expected to be in patients with liver cancer and other trials



may be in patients with brain cancer, lung cancer or colon cancer.

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***Utilize our discovery and development platform to develop additional protein therapeutic candidates.*** In addition to sotatercept, ACE-536 and dalantercept, all of which were internally discovered using our research and development platform, we intend to continue to discover and develop other protein therapeutics that target and regulate various pathways in the TGF- $\beta$  superfamily. We plan to bring an additional protein therapeutic candidate, ACE-083, into the clinic in 2014 targeting diseases involving muscle loss. We are also conducting pre-clinical development of ALK1 pathway inhibitors distinct from dalantercept for the treatment of diseases of the eye including age-related macular degeneration. In addition we are developing new protein therapeutic candidates for the treatment of cancer and diseases involving fibrosis.

***Strategically leverage collaborations to advance our protein therapeutic candidates.*** We have received more than \$250.0 million from our collaboration partners, including Celgene. Our two collaborations with Celgene for sotatercept and ACE-536 provide us with significant funding and access to Celgene's considerable scientific, development, regulatory and commercial capabilities. We will continue to strategically evaluate possible collaborations where doing so could enhance the development or commercialization of other protein therapeutic candidates in our pipeline.

***Establish commercialization and marketing capabilities in North America and potentially other markets.*** We have retained co-promotion rights in North America for sotatercept and ACE-536, which will be entirely funded by Celgene. We intend to build hematology, oncology and neuromuscular disorder focused specialty sales forces and marketing capability to commercialize our protein therapeutic candidates that receive regulatory approval.

### **Risk Factors**

An investment in our common stock involves a high degree of risk. Any of the factors set forth under "Risk Factors" may limit our ability to successfully execute our business strategy. You should carefully consider all of the information set forth in this prospectus and, in particular, should evaluate the specific factors set forth under "Risk Factors" in deciding whether to invest in our common stock. Among these important risks are the following:

We have incurred net operating losses since our inception and anticipate that we will continue to incur substantial operating losses for the foreseeable future. We may never achieve or sustain profitability.

We will require substantial additional financing to achieve our goals, and a failure to obtain this necessary capital when needed could force us to delay, limit, reduce or terminate our development or commercialization efforts of our protein therapeutic candidates.

If Celgene does not devote sufficient resources to the development of sotatercept and ACE-536, is unsuccessful in its efforts or chooses to terminate its agreements with us, our business will be materially harmed.

If our protein therapeutic candidates fail to demonstrate safety and efficacy to the satisfaction of regulatory authorities, we may incur additional costs or experience delays in completing, or ultimately be unable to complete the development and commercialization of our protein therapeutic candidates.

Our future commercial success depends upon attaining significant market acceptance of our protein therapeutic candidates, if approved, among physicians, patients and health care payers and, if we fail to do so, our business will be materially harmed.

We expect to rely on third parties in the manufacturing and clinical development of our protein therapeutic candidates. If they fail to meet deadlines or perform in an unsatisfactory manner our business could be harmed.



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If we are unable to obtain or protect intellectual property rights related to our protein therapeutic candidates, we may not be able to prevent competitors with the same or similar protein therapeutics from entering our markets.

**Implications of Being an Emerging Growth Company**

As a company with less than \$1.0 billion in revenue during our most recently completed fiscal year, we qualify as an "emerging growth company" as defined in Section 2(a) of the Securities Act of 1933, as amended, which we refer to as the Securities Act, as modified by the Jumpstart Our Business Startups Act of 2012, or the JOBS Act. As an emerging growth company, we may take advantage of specified reduced disclosure and other requirements that are otherwise applicable, in general, to public companies that are not emerging growth companies. These provisions include:

Reduced disclosure about our executive compensation arrangements;

No non-binding shareholder advisory votes on executive compensation or golden parachute arrangements;

Exemption from the auditor attestation requirement in the assessment of our internal control over financial reporting; and

Reduced disclosure of financial information in this prospectus, including two years of audited financial information and two years of selected financial information.

We may take advantage of these exemptions for up to five years or such earlier time that we are no longer an emerging growth company. We would cease to be an emerging growth company if we have more than \$1.0 billion in annual revenues as of the end of a fiscal year, if we are deemed to be a large-accelerated filer under the rules of the Securities and Exchange Commission, or if we issue more than \$1.0 billion of non-convertible debt over a three-year-period.

The JOBS Act permits an emerging growth company to take advantage of an extended transition period to comply with new or revised accounting standards applicable to public companies. We are choosing to "opt out" of this provision.

**Corporate Information**

We were incorporated in the state of Delaware in June 2003 as Phoenix Pharma, Inc., and we subsequently changed our name to Acceleron Pharma Inc. and commenced operations in February 2004. Our principal executive offices are located at 128 Sidney Street, Cambridge, Massachusetts 02139, and our telephone number is (617) 649-9200. Our Internet website is [www.acceleronpharma.com](http://www.acceleronpharma.com). The information on, or that can be accessed through, our website is not part of this prospectus, and you should not rely on any such information in making the decision whether to purchase our common stock.

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**THE OFFERING**

Common stock offered by us	2,400,000 shares
Common stock to be outstanding after this offering	30,748,633 shares
Option to purchase additional shares	The underwriters have an option for a period of 30 days to purchase up to 360,000 additional shares of our common stock.
Use of proceeds	The net proceeds from this offering will be approximately \$112.1 million, or approximately \$129.0 million if the underwriters exercise their option to purchase additional shares in full, after deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us. We intend to use the net proceeds of this offering (1) to continue development of dalantercept, (2) to conduct clinical trials and associated activities with a new protein therapeutic candidate, ACE-083, (3) to continue to advance and expand our preclinical research pipeline of protein therapeutic candidates and (4) for working capital and other general corporate purposes, including funding the costs of operating as a public company. See "Use of Proceeds".
Risk factors	You should read the "Risk Factors" section of this prospectus for a discussion of factors to consider carefully before deciding to invest in shares of our common stock.
NASDAQ Global Market symbol	XLRN

The number of shares of common stock to be outstanding after this offering is based on 28,348,633 shares of common stock outstanding as of January 1, 2014 and excludes the following:

3,942,304 shares of common stock issuable upon exercise of stock options outstanding as of January 1, 2014 at a weighted-average exercise price of \$7.05 per share;

979,699 shares of common stock issuable upon the exercise of outstanding warrants as of January 1, 2014 at a weighted-average exercise price of \$6.53 per share;

2,089,945 shares of common stock reserved for future issuance under our 2013 Equity Incentive Plan as of January 1, 2014; and

275,000 shares of common stock reserved for future issuance under our Employee Stock Purchase Plan as of January 1, 2014.

Unless otherwise indicated, all information in this prospectus assumes no issuance or exercise of stock options or warrants on or after January 1, 2014 and no exercise of the underwriters' option to purchase up to an additional 360,000 shares of common stock in this offering.

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The following summary financial data for the years ended December 31, 2011 and 2012 are derived from our audited financial statements included elsewhere in this prospectus. The summary financial data as of September 30, 2013 and for the nine months ended September 30, 2012 and 2013 have been derived from our unaudited financial statements included elsewhere in this prospectus. These unaudited financial statements have been prepared on a basis consistent with our audited financial statements and, in our opinion, contain all adjustments, consisting only of normal and recurring adjustments, necessary for a fair presentation of such financial data. You should read this data together with our audited financial statements and related notes included elsewhere in this prospectus and the information under the captions "Selected Financial Data" and "Management's Discussion and Analysis of Financial Condition and Results of Operations". Our historical results are not necessarily indicative of our future results, and our operating results for the nine-month period ended September 30, 2013 are not necessarily indicative of the results that may be expected for the fiscal year ending December 31, 2013 or any other interim periods or any future year or period.

(in thousands, except per share data)	Year Ended December 31,		Nine Months Ended September 30,	
	2011	2012	2012	2013
Revenue:				
Collaboration revenue:				
License and milestone	\$ 74,406	\$ 9,696	\$ 7,226	\$ 36,044
Cost-sharing, net	4,760	5,558	4,043	9,666
Contract manufacturing	1,745			
Total revenue	80,911	15,254	11,269	45,710
Costs and expenses:				
Research and development	32,713	35,319	25,646	25,834
General and administrative	8,142	8,824	6,318	9,472
Cost of contract manufacturing revenue	1,500			
Total costs and expenses	42,355	44,143	31,964	35,306
Income (loss) from operations	38,556	(28,889)	(20,695)	10,404
Total other expense, net	(2,290)	(3,693)	(1,508)	(14,192)
Net income (loss)	\$ 36,266	\$ (32,582)	\$ (22,203)	\$ (3,788)
Comprehensive income (loss)	\$ 36,266	\$ (32,582)	\$ (22,203)	\$ (3,788)
Net income (loss) per share applicable to common stockholders(1)				
Basic	\$ 0.80	\$ (24.84)	\$ (17.73)	\$ (6.74)
Diluted	\$ 0.78	\$ (24.84)	\$ (17.73)	\$ (6.74)
Weighted-average number of common shares used in computing net income (loss) per share applicable to common stockholders				
Basic	2,328	2,401	2,397	3,100
Diluted	2,716	2,401	2,397	3,100

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(in thousands)	September 30, 2013	
	Actual	As adjusted(2)
<b>Balance Sheet Data:</b>		
Cash and cash equivalents	\$ 116,479	\$ 228,579
Total assets	127,260	239,360
Total current liabilities	16,523	16,523
Long-term deferred revenue	6,205	6,205
Long-term notes payable	10,979	10,979
Warrants to purchase common stock	16,526	16,526
Total stockholders' equity	74,564	186,664

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- (1) See Note 2 within the notes to our financial statements appearing elsewhere in this prospectus for a description of the method used to calculate basic and diluted net income (loss) per common share and pro forma basic and diluted net income (loss) per common share.
- (2) As adjusted to reflect the sale of shares of our common stock in this offering at the public offering price of \$50.00 per share after deducting underwriting discounts and commissions and estimated offering expenses payable by us.

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**RISK FACTORS**

*Investing in our common stock involves a high degree of risk. You should carefully consider the risks and uncertainties described below together with all of the other information contained in this prospectus, including our financial statements and the related notes appearing at the end of this prospectus, before deciding to invest in our common stock. If any of the following risks actually occurs, our business, prospects, operating results and financial condition could suffer materially, the trading price of our common stock could decline and you could lose all or part of your investment. The risks and uncertainties described below are not the only ones we face. Additional risks and uncertainties not presently known to us or that we currently believe to be immaterial may also adversely affect our business.*

**Risks related to our financial position and need for additional capital**

*We have incurred net operating losses since our inception and anticipate that we will continue to incur substantial operating losses for the foreseeable future. We may never achieve or sustain profitability.*

We have incurred net losses during most fiscal periods since our inception. As of September 30, 2013, we had an accumulated deficit of \$174.2 million. We do not know whether or when we will become profitable. To date, we have not commercialized any products or generated any revenues from the sale of products, and we do not expect to generate any product revenues in the foreseeable future. Our losses have resulted principally from costs incurred in our discovery and development activities.

We anticipate that our expenses will increase in the future as we expand our discovery, research, development, manufacturing and commercialization activities. However, we also anticipate that these increased expenses will be partially offset by milestone payments we expect to receive under our agreements with Celgene and potentially by payments we may receive under new collaboration arrangements we may enter into with third parties for dalantercept or other protein therapeutic candidates. If we do not receive the anticipated milestone payments or do not enter into partnerships for dalantercept or other protein therapeutic candidates on acceptable terms, our operating losses will substantially increase over the next several years as we execute our plan to expand our discovery, research, development, manufacturing and commercialization activities. There can be no assurance that we will enter into a new collaboration or achieve milestones and, therefore, no assurance our losses will not increase prohibitively in the future.

To become and remain profitable, we or our partners must succeed in developing our protein therapeutic candidates, obtaining regulatory approval for them, and manufacturing, marketing and selling those products for which we or our partners may obtain regulatory approval. We or they may not succeed in these activities, and we may never generate revenue from product sales that is significant enough to achieve profitability. Even if we achieve profitability in the future, we may not be able to sustain profitability in subsequent periods. Our failure to become or remain profitable would depress our market value and could impair our ability to raise capital, expand our business, discover or develop other protein therapeutic candidates or continue our operations. A decline in the value of our company could cause you to lose all or part of your investment.

*We will require substantial additional financing to achieve our goals, and a failure to obtain this necessary capital when needed could force us to delay, limit, reduce or terminate our product development or commercialization efforts.*

As of September 30, 2013, our cash and cash equivalents were \$116.5 million. We believe that we will continue to expend substantial resources for the foreseeable future developing dalantercept and new protein therapeutic candidates. These expenditures will include costs associated with research and development, potentially acquiring new technologies, conducting preclinical studies and clinical trials, potentially obtaining regulatory approvals and manufacturing products, as well as marketing and selling products approved for sale, if any. In addition, other unanticipated costs may arise. Because the



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outcome of our planned and anticipated clinical trials is highly uncertain, we cannot reasonably estimate the actual amounts necessary to successfully complete the development and commercialization of our protein therapeutic candidates.

Celgene pays development, manufacturing and commercialization and certain patent costs for sotatercept and ACE-536. Other than those costs, our future capital requirements depend on many factors, including:

the scope, progress, results and costs of researching and developing our other protein therapeutic candidates, and conducting preclinical studies and clinical trials;

the timing of, and the costs involved in, obtaining regulatory approvals for our other protein therapeutic candidates if clinical trials are successful;

the cost of commercialization activities for our other protein therapeutics, if any of these protein therapeutics is approved for sale, including marketing, sales and distribution costs;

the cost of manufacturing our other protein therapeutic candidates for clinical trials in preparation for regulatory approval and in preparation for commercialization;

our ability to establish and maintain strategic partnerships, licensing or other arrangements and the financial terms of such agreements;

the costs involved in preparing, filing, prosecuting, maintaining, defending and enforcing patent claims, including litigation costs and the outcome of such litigation; and

the timing, receipt, and amount of sales of, or royalties on, our future products, if any.

Based on our current operating plan, we believe that the net proceeds we receive from this offering, together with receipt of anticipated milestone payments and our existing cash and cash equivalents will be sufficient to fund our projected operating requirements into the first quarter of 2017. However, our operating plan may change as a result of many factors currently unknown to us, and we may need additional funds sooner than planned. In addition, we may seek additional capital due to favorable market conditions or strategic considerations even if we believe we have sufficient funds for our current or future operating plans. Additional funds may not be available when we need them on terms that are acceptable to us, or at all. If adequate funds are not available to us on a timely basis, we may be required to delay, limit, reduce or terminate preclinical studies, clinical trials or other development activities for one or more of our protein therapeutic candidates or delay, limit, reduce or terminate our establishment of sales and marketing capabilities or other activities that may be necessary to commercialize our protein therapeutic candidates.

***Raising additional capital may cause dilution to our existing stockholders, restrict our operations or require us to relinquish rights to our technologies or protein therapeutics on unfavorable terms to us.***

We may seek additional capital through a variety of means, including through private and public equity offerings and debt financings. To the extent that we raise additional capital through the sale of equity or convertible debt securities, your ownership interest will be diluted, and the terms may include liquidation or other preferences that adversely affect your rights as a stockholder. Debt financing, if available, may involve agreements that include covenants limiting or restricting our ability to take certain actions, such as incurring additional debt, making capital expenditures or declaring dividends. If we raise additional funds through strategic partnerships with third parties, we may have to relinquish valuable rights to our technologies or protein therapeutics, or grant licenses on terms that are not favorable to us. If we are unable to raise additional funds through equity or debt financing when needed, we may be required to delay, limit, reduce or terminate our product development or commercialization efforts for dalantercept or any protein therapeutics other than sotatercept or

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ACE-536, or grant rights to develop and market protein therapeutics that we would otherwise prefer to develop and market ourselves.

**Risks Related to Regulatory Review and Approval of Our Protein Therapeutic Candidates**

*If we or our partners do not obtain regulatory approval for our current and future protein therapeutics, our business will be adversely affected.*

Our protein therapeutic candidates will be subject to extensive governmental regulations relating to, among other things, development, clinical trials, manufacturing and commercialization. In order to obtain regulatory approval for the commercial sale of any protein therapeutic candidates, we or our partners must demonstrate through extensive preclinical studies and clinical trials that the protein therapeutic candidate is safe and effective for use in each target indication. Clinical testing is expensive, time-consuming and uncertain as to outcome. We or our partners may gain regulatory approval for sotatercept, ACE-536, dalantercept, or any other protein therapeutic candidate in some but not all of the territories available or some but not all of the target indications, resulting in limited commercial opportunity for the approved protein therapeutics, or we or they may never obtain regulatory approval for these protein therapeutic candidates.

*Delays in the commencement, enrollment or completion of clinical trials of our protein therapeutic candidates could result in increased costs to us as well as a delay or failure in obtaining regulatory approval, or prevent us from commercializing our protein therapeutic candidates on a timely basis, or at all.*

We cannot guarantee that clinical trials will be conducted as planned or completed on schedule, if at all. A failure of one or more clinical trials can occur at any stage of testing. Events that may prevent successful or timely commencement, enrollment or completion of clinical development include:

delays by us or our partners in reaching a consensus with regulatory agencies on trial design;

delays in reaching agreement on acceptable terms with prospective clinical research organizations, or CROs, and clinical trial sites;

delays in obtaining required Institutional Review Board, or IRB, approval at each clinical trial site;

delays in recruiting suitable patients to participate in clinical trials;

imposition of a clinical hold by regulatory agencies for any reason, including safety concerns or after an inspection of clinical operations or trial sites;

failure by CROs, other third parties or us or our partners to adhere to clinical trial requirements;

failure to perform in accordance with the FDA's good clinical practices, or GCP, or applicable regulatory guidelines in other countries;

delays in the testing, validation, manufacturing and delivery of the protein therapeutic candidates to the clinical sites;

delays caused by patients not completing participation in a trial or not returning for post-treatment follow-up;

clinical trial sites or patients dropping out of a trial;

occurrence of serious adverse events in clinical trials that are associated with the protein therapeutic candidates that are viewed to outweigh its potential benefits; or

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changes in regulatory requirements and guidance that require amending or submitting new clinical protocols.

Delays, including delays caused by the above factors, can be costly and could negatively affect our or Celgene's ability to complete a clinical trial. If we or Celgene are not able to successfully complete clinical trials, we will not be able to obtain regulatory approval and will not be able to commercialize our protein therapeutic candidates.

***Clinical failure may occur at any stage of clinical development, and because our protein therapeutic candidates are in an early stage of development, there is a high risk of failure, and we may never succeed in developing marketable products or generating product revenue.***

Our early encouraging preclinical and clinical results for sotatercept, ACE-536 and dalantercept are not necessarily predictive of the results of our ongoing or future clinical trials. Promising results in preclinical studies of a drug candidate may not be predictive of similar results in humans during clinical trials, and successful results from early clinical trials of a drug candidate may not be replicated in later and larger clinical trials or in clinical trials for different indications. If the results of our or our partners' ongoing or future clinical trials are inconclusive with respect to the efficacy of our protein therapeutic candidates or if we or they do not meet the clinical endpoints with statistical significance or if there are safety concerns or adverse events associated with our protein therapeutic candidates, we or our partner may be prevented or delayed in obtaining marketing approval for our protein therapeutic candidates. In addition, data obtained from trials and studies are susceptible to varying interpretations, and regulators may not interpret our data as favorably as we do, which may delay or prevent regulatory approval. Alternatively, even if we or our partners obtain regulatory approval, that approval may be for indications or patient populations that are not as broad as intended or desired or may require labeling that includes significant use or distribution restrictions or safety warnings. We or our partners may also be required to perform additional or unanticipated clinical trials to obtain approval or be subject to additional post-marketing testing requirements to maintain regulatory approval. In addition, regulatory authorities may withdraw their approval of a product or impose restrictions on its distribution, such as in the form of a modified risk evaluation and mitigation strategy.

***If we or any of our partners violate the guidelines pertaining to promotion and advertising of any of our protein therapeutics, if approved, we or they may be subject to disciplinary action by the FDA's Office of Prescription Drug Promotion (OPDP) or other regulatory authorities.***

The FDA's Office of Prescription Drug Promotion, or OPDP, is responsible for reviewing prescription drug advertising and promotional labeling to ensure that the information contained in these materials is not false or misleading. There are specific disclosure requirements, and the applicable regulations mandate that advertisements cannot be false or misleading or omit material facts about the product. Prescription drug promotional materials must present a fair balance between the drug's effectiveness and the risks associated with its use. Most warning letters from OPDP cite inadequate disclosure of risk information.

OPDP prioritizes its actions based on the degree of risk to the public health, and often focuses on newly introduced drugs and those associated with significant health risks. There are two types of letters that OPDP typically sends to companies that violate its drug advertising and promotional guidelines: notice of violation letters, or untitled letters, and warning letters. In the case of an untitled letter, OPDP typically alerts the drug company of the violation and issues a directive to refrain from future violations, but does not typically demand other corrective action. A warning letter is typically issued in cases that are more serious or where the company is a repeat offender. Although we have not received any such letters from OPDP, we or any partner may inadvertently violate OPDP's guidelines in the future and be subject to a OPDP untitled letter or warning letter, which may have a negative impact on our business.

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***If we or our partners fail to obtain regulatory approval in jurisdictions outside the United States, we and they will not be able to market our products in those jurisdictions.***

We and our partners intend to market our protein therapeutic candidates, if approved, in international markets. Such marketing will require separate regulatory approvals in each market and compliance with numerous and varying regulatory requirements. The approval procedures vary from country-to-country and may require additional testing. Moreover, the time required to obtain approval may differ from that required to obtain FDA approval. In addition, in many countries outside the United States, a protein therapeutic must be approved for reimbursement before it can be approved for sale in that country. Approval by the FDA does not ensure approval by regulatory authorities in other countries or jurisdictions, and approval by one foreign regulatory authority does not ensure approval by regulatory authorities in other foreign countries or by the FDA. The foreign regulatory approval process may include all of the risks associated with obtaining FDA approval. We or our partners may not obtain foreign regulatory approvals on a timely basis, if at all. We or our partners may not be able to file for regulatory approvals and may not receive necessary approvals to commercialize our products in any market.

***Even if we or our partners receive regulatory approval for our protein therapeutic candidates, such products will be subject to ongoing regulatory review, which may result in significant additional expense. Additionally, our protein therapeutic candidates, if approved, could be subject to labeling and other restrictions, and we or our partners may be subject to penalties if we fail to comply with regulatory requirements or experience unanticipated problems with our products.***

Any regulatory approvals that we or our partners receive for our protein therapeutic candidates may also be subject to limitations on the approved indicated uses for which the product may be marketed or to conditions of approval, or contain requirements for potentially costly post-marketing testing, including Phase 4 clinical trials, and surveillance to monitor safety and efficacy. In addition, if the FDA approves any of our protein therapeutic candidates, the manufacturing processes, labeling, packaging, distribution, adverse event reporting, storage, advertising, promotion and recordkeeping for the product will be subject to extensive and ongoing regulatory requirements. These requirements include submissions of safety and other post-marketing information and reports, registration, as well as continued compliance with current good manufacturing practice, or cGMP, and GCP, for any clinical trials that we or our partners conduct post-approval.

Later discovery of previously unknown problems with an approved protein therapeutic, including adverse events of unanticipated severity or frequency, or with manufacturing operations or processes, or failure to comply with regulatory requirements, may result in, among other things:

restrictions on the marketing or manufacturing of the product, withdrawal of the product from the market, or voluntary or mandatory product recalls;

finest, warning letters, or holds on clinical trials;

refusal by the FDA to approve pending applications or supplements to approved applications filed by us or our partners, or suspension or revocation of product license approvals;

product seizure or detention, or refusal to permit the import or export of products; and

injunctions or the imposition of civil or criminal penalties.

The FDA's policies may change and additional government regulations may be enacted that could prevent, limit or delay regulatory approval of our protein therapeutic candidates. We cannot predict the likelihood, nature or extent of government regulation that may arise from future legislation or administrative action, either in the United States or abroad. If we or our partners are slow or unable to adapt to changes in existing requirements or the adoption of new requirements or policies, or not able

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to maintain regulatory compliance, we or our partners may lose any marketing approval that may have been obtained and we may not achieve or sustain profitability, which would adversely affect our business.

**Risks Related to Our Reliance on Third Parties**

*We are dependent on Celgene for the successful development and commercialization of two of our three clinical stage protein therapeutic candidates, sotatercept and ACE-536. If Celgene does not devote sufficient resources to the development of these candidates, is unsuccessful in its efforts, or chooses to terminate its agreements with us, our business will be materially harmed.*

We have entered into collaboration agreements with Celgene to develop and commercialize sotatercept and ACE-536. Pursuant to the sotatercept agreement, responsibility for all clinical and other product development activities and for manufacturing sotatercept has been transferred to Celgene. For ACE-536, we are responsible for conducting the two ongoing Phase 2 clinical trials, and we are also responsible for manufacturing supplies for Phase 1 and Phase 2 studies. Celgene will be responsible for all clinical development and manufacturing activities after such studies are completed. As of January 1, 2013, Celgene became responsible for paying 100% of worldwide development costs for sotatercept and ACE-536. We will co-promote sotatercept and ACE-536, if approved by the FDA and its counterparties, in North America. Celgene will be responsible for all commercialization costs, including the cost of our promotion activities.

Celgene is obligated to use commercially reasonable efforts to develop and commercialize sotatercept and ACE-536. Celgene may determine that it is commercially reasonable to develop and commercialize only sotatercept or ACE-536 and discontinue the development or commercialization of the other protein therapeutic candidate, or Celgene may determine that it is not commercially reasonable to continue development of one or both of sotatercept and ACE-536. In the event of any such decision, we may be unable to progress the discontinued candidate or candidates ourselves. In addition, under our collaboration agreements, once Celgene takes over development activities of a protein therapeutic candidate, it may determine the development plan and activities for that protein therapeutic candidate. We may disagree with Celgene about the development strategy it employs, but we will have no rights to impose our development strategy on Celgene. Similarly, Celgene may decide to seek regulatory approval for, and limit commercialization of, either or both of sotatercept and ACE-536 to narrower indications than we would pursue. We would be prevented from developing or commercializing a candidate in an indication that Celgene has chosen not to pursue.

This partnership may not be scientifically or commercially successful due to a number of important factors, including the following:

Celgene has wide discretion in determining the efforts and resources that it will apply to its partnership with us. The timing and amount of any development milestones, and downstream commercial milestones and royalties that we may receive under such partnership will depend on, among other things, the efforts, allocation of resources and successful development and commercialization of these protein therapeutic candidates by Celgene.

Celgene may develop and commercialize, either alone or with others, products that are similar to or competitive with the protein therapeutic candidates that are the subject of its partnerships with us. For example, Celgene is currently commercializing and/or developing certain of its existing products, lenalidomide and azacitidine, for certain MDS patients for which sotatercept and ACE-536 are also being developed.

Celgene may terminate its partnership with us without cause and for circumstances outside of our control, which could make it difficult for us to attract new strategic partners or adversely affect how we are perceived in scientific and financial communities.

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Celgene may develop or commercialize our protein therapeutic candidates in such a way as to elicit litigation that could jeopardize or invalidate our intellectual property rights or expose us to potential liability.

Celgene may not comply with all applicable regulatory requirements, or fail to report safety data in accordance with all applicable regulatory requirements.

If Celgene were to breach its arrangements with us, we may need to enforce our right to terminate the agreement in legal proceedings, which could be costly and cause delay in our ability to receive rights back to the relevant protein therapeutic candidates. If we were to terminate an agreement with Celgene due to Celgene's breach or Celgene terminated the agreement without cause, the development and commercialization of sotatercept and ACE-536 could be delayed, curtailed or terminated because we may not have sufficient financial resources or capabilities to continue development and commercialization of these candidates on our own.

Celgene may enter into one or more transactions with third parties, including a merger, consolidation, reorganization, sale of substantial assets, sale of substantial stock or other change in control, which could divert the attention of its management and adversely affect Celgene's ability to retain and motivate key personnel who are important to the continued development of the programs under the strategic partnership with us. In addition, the third-party to any such transaction could determine to reprioritize Celgene's development programs such that Celgene ceases to diligently pursue the development of our programs and/or cause the respective partnership with us to terminate.

***We and Celgene rely on third parties to conduct preclinical studies and clinical trials for our protein therapeutic candidates, and if they do not properly and successfully perform their obligations to us, we may not be able to obtain regulatory approvals for our protein therapeutic candidates.***

We design the clinical trials for dalantercept and will do so for any future unpartnered protein therapeutic candidates, and we will continue to work with Celgene on trials for sotatercept and ACE-536. However, we and Celgene rely on CROs and other third parties to assist in managing, monitoring and otherwise carrying out many of these trials. We and Celgene compete with many other companies for the resources of these third parties. The third parties on whom we and Celgene rely generally may terminate their engagements at any time, and having to enter into alternative arrangements would delay development and commercialization of our protein therapeutic candidates.

The FDA and foreign regulatory authorities require compliance with regulations and standards, including GCP, for designing, conducting, monitoring, recording, analyzing, and reporting the results of clinical trials to assure that the data and results are credible and accurate and that the rights, integrity and confidentiality of trial participants are protected. Although we and Celgene rely on third parties to conduct many of our and their clinical trials, we and Celgene are responsible for ensuring that each of these clinical trials is conducted in accordance with its general investigational plan, protocol and other requirements.

If these third parties do not successfully carry out their duties under their agreements, if the quality or accuracy of the data they obtain is compromised due to their failure to adhere to clinical trial protocols or to regulatory requirements, or if they otherwise fail to comply with clinical trial protocols or meet expected deadlines, the clinical trials of our protein therapeutic candidates may not meet regulatory requirements. If clinical trials do not meet regulatory requirements or if these third parties need to be replaced, preclinical development activities or clinical trials may be extended, delayed, suspended or terminated. If any of these events occur, we or Celgene may not be able to obtain regulatory approval of our protein therapeutic candidates on a timely basis or at all.

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***We and Celgene intend to rely on third-party manufacturers to make our protein therapeutics, and any failure by a third-party manufacturer may delay or impair our and Celgene's ability to complete clinical trials or commercialize our protein therapeutic candidates.***

Manufacturing biologic drugs is complicated and is tightly regulated by the FDA, the European Medicines Agency, or EMA, and comparable regulatory authorities around the world. We currently manufacture drug substance for our preclinical studies, Phase 1 clinical trials and Phase 2 clinical trials of ACE-536 and dalantercept. For Phase 3 and commercial supply of our products that we have not partnered, we expect to use contract manufacturing organizations. Successfully transferring complicated manufacturing techniques to contract manufacturing organizations and scaling up these techniques for commercial quantities will be time consuming and we may not be able to achieve such transfer. Moreover, the market for contract manufacturing services for protein therapeutics is highly cyclical, with periods of relatively abundant capacity alternating with periods in which there is little available capacity. If any need we or Celgene have for contract manufacturing services increases during a period of industry-wide tight capacity, we or Celgene may not be able to access the required capacity on a timely basis or on commercially viable terms.

In addition, we contract with fill & finishing providers with the appropriate expertise, facilities and scale to meet our needs. Failure to maintain cGMP can result in a contractor receiving FDA sanctions, which can impact our and Celgene's contractors' ability to operate or lead to delays in our clinical development programs. We believe that our current fill & finish contractors are operating in accordance with cGMP, but we can give no assurance that FDA or other regulatory agencies will not conclude that a lack of compliance exists. In addition, any delay in contracting for fill & finish services, or failure of the contract manufacturer to perform the services as needed, may delay clinical trials, registration and launches. Any such issues may have a substantial negative effect on our business.

***For our two lead products, sotatercept and ACE-536, we rely on our collaboration partner Celgene to produce, or contract for the production of, bulk drug substance and finished drug product for late stage clinical trials and for commercial supplies of any approved candidates. Any failure by Celgene or by third-parties with which Celgene contracts may delay or impair the ability to complete late stage clinical trials or commercialize either or both of sotatercept and ACE-536, if approved.***

We produced drug substance for preclinical and Phase 1 and 2 clinical trials for sotatercept and ACE-536. Celgene is now responsible for manufacturing sotatercept and will be responsible for manufacturing ACE-536 for future late-stage clinical trials. Celgene generally does not perform the manufacture of the drug substance or drug product for either sotatercept or ACE-536 itself. Celgene has used and may continue to use contract manufacturers for the manufacture of drug substance and drug product for sotatercept and we have no expectation that Celgene plans to perform the manufacture of bulk drug substance or drug product for either sotatercept or ACE-536 in the future. However, Celgene would have the right to manufacture sotatercept or ACE-536, itself or through the use of contract manufacturers. We understand that they have entered into a manufacturing arrangement for Phase 2 supplies of sotatercept bulk drug substance with contract manufacturers with considerable biotherapeutics manufacturing experience, including manufacturing monoclonal antibodies through processes similar to those used for sotatercept. To date Celgene has not entered into an arrangement with a third party to manufacture supplies of sotatercept or ACE-536 for Phase 3 trials or commercial sales. If they are unable to contract at the appropriate time with a manufacturer willing and able to manufacture sufficient quantities of sotatercept and ACE-536 to meet Phase 3 and commercial demand, either for technical or business reasons, the development and commercialization of sotatercept and ACE-536 may be delayed.



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*We may not be successful in establishing and maintaining additional strategic partnerships, which could adversely affect our ability to develop and commercialize products, negatively impacting our operating results.*

In addition to our current collaborations with Celgene, a part of our strategy is to strategically evaluate and, as deemed appropriate, enter into additional partnerships in the future when strategically attractive, including potentially with major biotechnology or pharmaceutical companies. We face significant competition in seeking appropriate partners for our protein therapeutic candidates, and the negotiation process is time-consuming and complex. In order for us to successfully partner our protein therapeutic candidates, potential partners must view these protein therapeutic candidates as economically valuable in markets they determine to be attractive in light of the terms that we are seeking and other available products for licensing by other companies. Even if we are successful in our efforts to establish new strategic partnerships, the terms that we agree upon may not be favorable to us, and we may not be able to maintain such strategic partnerships if, for example, development or approval of a protein therapeutic is delayed or sales of an approved product are disappointing. Any delay in entering into new strategic partnership agreements related to our protein therapeutic candidates could delay the development and commercialization of our protein therapeutic candidates and reduce their competitiveness even if they reach the market.

If we fail to establish and maintain additional strategic partnerships related to our protein therapeutic candidates, we will bear all of the risk and costs related to the development of any such protein therapeutic candidate, and we may need to seek additional financing, hire additional employees and otherwise develop expertise for which we have not budgeted. This could negatively affect the development of any unpartnered protein therapeutic candidate.

**Risks Related to Our Intellectual Property**

*If we are unable to obtain or protect intellectual property rights related to our protein therapeutic candidates, we may not be able to compete effectively.*

We rely upon a combination of patents, trade secret protection and confidentiality agreements to protect the intellectual property related to our platform technology and protein therapeutic candidates. The patent position of biotechnology companies is generally uncertain because it involves complex legal and factual considerations. The standards applied by the United States Patent and Trademark Office, or USPTO, and foreign patent offices in granting patents are not always applied uniformly or predictably. For example, there is no uniform worldwide policy regarding patentable subject matter or the scope of claims allowable in biotechnology patents. The patent applications that we own or in-license may fail to result in issued patents with claims that cover our protein therapeutic candidates in the United States or in other countries. There is no assurance that all potentially relevant prior art relating to our patents and patent applications has been found. We may be unaware of prior art that could be used to invalidate an issued patent or prevent our pending patent applications from issuing as patents. Even if patents do successfully issue and even if such patents cover our protein therapeutic candidates, third parties may challenge their validity, enforceability or scope, which may result in such patents being narrowed or invalidated. Furthermore, even if they are unchallenged, our patents and patent applications may not adequately protect our intellectual property, provide exclusivity for our protein therapeutic candidates or prevent others from designing around our claims. Any of these outcomes could impair our ability to prevent competition from third parties, which may have an adverse impact on our business.

If patent applications we hold or have in-licensed with respect to our platform or protein therapeutic candidates fail to issue, if their breadth or strength of protection is threatened, or if they fail to provide meaningful exclusivity for our protein therapeutic candidates, it could dissuade companies from collaborating with us. Several patent applications covering our protein therapeutic candidates have been filed recently. We cannot offer any assurances about which, if any, patents will issue, the breadth of any such patents or whether any issued patents will be found invalid and unenforceable or will be threatened

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by third parties. Any successful challenge to these patents or any other patents owned by or licensed to us could deprive us of rights necessary for the successful commercialization of any protein therapeutic candidate that we or our partners may develop. Since patent applications in the United States and most other countries are confidential for a period of time after filing, and some remain so until issued, we cannot be certain that we were the first to file any patent application related to a protein therapeutic candidate. Furthermore, if third parties have filed such patent applications, an interference proceeding in the United States can be initiated by the USPTO or a third party to determine who was the first to invent any of the subject matter covered by the patent claims of our applications. In addition, patents have a limited lifespan. In the United States, the natural expiration of a patent is generally 20 years after it is filed. Various extensions may be available; however, the life of a patent and the protection it affords is limited. If we encounter delays in obtaining regulatory approvals, the period of time during which we could market a protein therapeutic under patent protection could be reduced. Even if patents covering our protein therapeutic candidates are obtained, once the patent life has expired for a product, we may be open to competition from biosimilar products.

Any loss of patent protection could have a material adverse impact on our business. We and our partner may be unable to prevent competitors from entering the market with a product that is similar to or the same as our protein therapeutics. In addition, the royalty we would receive under our collaboration agreements with Celgene for sotatercept and ACE-536 would be reduced by 50% if such product ceases to be covered by a valid claim of our patents even if no competitor with a similar product has entered the market.

***Third-party claims of intellectual property infringement or misappropriation may prevent or delay our development and commercialization efforts.***

Our commercial success depends in part on us and our partners not infringing the patents and proprietary rights of third parties. There is a substantial amount of litigation, both within and outside the United States, involving patent and other intellectual property rights in the biotechnology and pharmaceutical industries, including patent infringement lawsuits, interferences, oppositions and inter partes reexamination proceedings before the USPTO and corresponding foreign patent offices. Numerous U.S. and foreign issued patents and pending patent applications owned by third parties exist in the fields in which we and our partners are developing and may develop our protein therapeutic candidates. As the biotechnology and pharmaceutical industries expand and more patents are issued, the risk increases that our protein therapeutic candidates may be subject to claims of infringement of the patent rights of third parties.

Third parties may assert that we are employing their proprietary technology without authorization. There may be third-party patents or patent applications with claims to materials, formulations, methods of manufacture or methods for treatment related to the use or manufacture of our protein therapeutic candidates, that we failed to identify. For example, applications filed before November 29, 2000 and certain applications filed after that date that will not be filed outside the United States remain confidential until issued as patents. Except for the preceding exceptions, patent applications in the United States and elsewhere are generally published only after a waiting period of approximately 18 months after the earliest filing. Therefore, patent applications covering our platform technology or our protein therapeutic candidates could have been filed by others without our knowledge. Additionally, pending patent applications which have been published can, subject to certain limitations, be later amended in a manner that could cover our platform technologies, our protein therapeutic candidates or the use or manufacture of our protein therapeutic candidates.

If any third-party patents were held by a court of competent jurisdiction to cover aspects of our materials, formulations, methods of manufacture or methods for treatment, the holders of any such patents would be able to block our ability to develop and commercialize the applicable protein therapeutic candidate until such patent expired or unless we or our partners obtain a license. These

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licenses may not be available on acceptable terms, if at all. Even if we or our partners were able to obtain a license, the rights may be nonexclusive, which could result in our competitors gaining access to the same intellectual property. Ultimately, we or our partners could be prevented from commercializing a product, or be forced to cease some aspect of our business operations, if, as a result of actual or threatened patent infringement claims, we or our partners are unable to enter into licenses on acceptable terms. If Celgene is required to enter a license agreement with a third party in order to import, develop, manufacture or commercialize sotatercept or ACE-536, the royalty rate and sales milestone payments that we could receive may be reduced by up to 50%. This could harm our business significantly.

Parties making claims against us or our partners may obtain injunctive or other equitable relief, which could effectively block our or our partners' ability to further develop and commercialize one or more of our protein therapeutic candidates. Defending against claims of patent infringement or misappropriation of trade secrets could be costly and time consuming, regardless of the outcome. Thus, even if we were to ultimately prevail, or to settle at an early stage, such litigation could burden us with substantial unanticipated costs. In addition, litigation or threatened litigation could result in significant demands on the time and attention of our management team, distracting them from the pursuit of other company business. In the event of a successful claim of infringement against us or our partners, we may have to pay substantial damages, including treble damages and attorneys' fees for willful infringement, pay royalties, redesign our infringing products or obtain one or more licenses from third parties, which may be impossible or require substantial time and monetary expenditure.

We may face a claim of misappropriation if a third party believes that we inappropriately obtained and used trade secrets of such third party. If we are found to have misappropriated a third party's trade secrets, we may be prevented from further using such trade secrets, limiting our ability to develop our protein therapeutic candidates, and we may be required to pay damages.

During the course of any patent or other intellectual property litigation, there could be public announcements of the results of hearings, rulings on motions, and other interim proceedings in the litigation. If securities analysts or investors regard these announcements as negative, the perceived value of our protein therapeutics, programs, or intellectual property could be diminished. Accordingly, the market price of our common stock may decline.

***We have in-licensed a portion of our intellectual property, and, if we fail to comply with our obligations under these arrangements, we could lose such intellectual property rights or owe damages to the licensor of such intellectual property.***

We are a party to a number of license agreements that are important to our business, and we may enter into additional license agreements in the future. Our discovery and development platform is built, in part, around patents exclusively in-licensed from academic or research institutions. Certain of our in-licensed intellectual property also covers sotatercept and dalantercept. See "Business Intellectual Property In-Licenses" for a description of our license agreements with the Beth Israel Deaconess Medical Center, the Ludwig Institute for Cancer Research and the Salk Institute for Biological Studies.

Our existing license agreements impose, and we expect that future license agreements will impose, various diligence, milestone payment, royalty and other obligations on us. If there is any conflict, dispute, disagreement or issue of non-performance between us and our licensing partners regarding our rights or obligations under the license agreements, including any such conflict, dispute or disagreement arising from our failure to satisfy payment obligations under any such agreement, we may owe damages, our licensor may have a right to terminate the affected license, and our and our partners' ability to utilize the affected intellectual property in our drug discovery and development efforts, and our ability to enter into collaboration or marketing agreements for an affected protein therapeutic candidate, may be adversely affected.

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For example, the Salk Institute for Biological Studies recently filed a lawsuit against us alleging under one of our license agreements with them, which pertains to ActRIIB, its entitlement to a further share of certain payments received by us under our now-terminated agreement with Shire AG and a share of certain payments received by us under our on-going collaboration agreement with Celgene in connection with ACE-536. Although we and Salk have agreed that ACE-536 is not covered by any patent rights licensed from Salk, an unfavorable outcome in this litigation may lead to us owing significant damages to Salk and higher-than-anticipated future payments under this license in connection with development milestone payments that we may receive from Celgene. It is possible that Salk may seek to terminate the license covering the receptor. We do not believe that such a termination would have a material impact on our business or the development of any of our products. The patents under this license covered only one of our protein therapeutic candidates, ACE-031, the development of which has been discontinued. See "Business Legal Proceedings" for a description of this proceeding.

***Confidentiality agreements with employees and third parties may not prevent unauthorized disclosure of trade secrets and other proprietary information.***

In addition to the protection afforded by patents, we rely on trade secret protection and confidentiality agreements to protect proprietary know-how that is not patentable or that we elect not to patent, processes for which patents are difficult to enforce and any other elements of our platform technology and discovery and development processes that involve proprietary know-how, information or technology that is not covered by patents. However, trade secrets can be difficult to protect. We seek to protect our proprietary technology and processes, in part, by entering into confidentiality agreements with our employees, consultants, and outside scientific advisors, contractors and collaborators. Although we use reasonable efforts to protect our trade secrets, our employees, consultants, contractors, or outside scientific advisors might intentionally or inadvertently disclose our trade secret information to competitors. In addition, competitors may otherwise gain access to our trade secrets or independently develop substantially equivalent information and techniques.

Enforcing a claim that a third party illegally obtained and is using any of our trade secrets is expensive and time consuming, and the outcome is unpredictable. In addition, courts outside the United States sometimes are less willing than U.S. courts to protect trade secrets. Misappropriation or unauthorized disclosure of our trade secrets could impair our competitive position and may have a material adverse effect on our business

**Risks Related to Commercialization of Our Protein Therapeutic Candidates**

***Our future commercial success depends upon attaining significant market acceptance of our protein therapeutic candidates, if approved, among physicians, patients, health care payers and, in cancer markets, acceptance by the operators of major cancer clinics.***

Even if we or our partners obtain regulatory approval for sotatercept, ACE-536, dalantercept or any other protein therapeutics that we may develop or acquire in the future, the product may not gain market acceptance among physicians, health care payors, patients and the medical community. Market acceptance of any approved products depends on a number of factors, including:

the efficacy and safety of the product, as demonstrated in clinical trials;

the clinical indications for which the product is approved and the label approved by regulatory authorities for use with the product, including any warnings that may be required on the label;

acceptance by physicians and patients of the product as a safe and effective treatment;

the cost, safety and efficacy of treatment in relation to alternative treatments;

the availability of adequate reimbursement and pricing by third party payers and government authorities;

the continued projected growth of drug markets in our various indications;

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relative convenience and ease of administration;

the prevalence and severity of adverse side effects; and

the effectiveness of our, and our partners' sales and marketing efforts.

Market acceptance is critical to our ability to generate significant revenue. Any protein therapeutic candidate, if approved and commercialized, may be accepted in only limited capacities or not at all. If any approved products are not accepted by the market to the extent that we expect, we may not be able to generate significant revenue and our business would suffer.

***Reimbursement may be limited or unavailable in certain market segments for our protein therapeutic candidates, which could make it difficult for us to sell our products profitably.***

Market acceptance and sales of any approved protein therapeutics will depend significantly on the availability of adequate coverage and reimbursement from third-party payers and may be affected by existing and future health care reform measures. Government authorities and third-party payors, such as private health insurers and health maintenance organizations, decide which drugs they will pay for and establish reimbursement levels. Reimbursement by a third-party payer may depend upon a number of factors, including the third-party payer's determination that use of a product is:

a covered benefit under its health plan;

safe, effective and medically necessary;

appropriate for the specific patient;

cost-effective; and

neither experimental nor investigational.

Obtaining coverage and reimbursement approval for a product from a government or other third party payer is a time consuming and costly process that could require us to provide supporting scientific, clinical and cost-effectiveness data for the use of our products to the payer. We or our partners may not be able to provide data sufficient to gain acceptance with respect to coverage and reimbursement. We cannot be sure that coverage or adequate reimbursement will be available for any of our protein therapeutic candidates. Also, we cannot be sure that reimbursement amounts will not reduce the demand for, or the price of, our products. If reimbursement is not available or is available only to limited levels, we may not be able to commercialize certain of our products. In addition in the United States, third-party payers are increasingly attempting to contain health care costs by limiting both coverage and the level of reimbursement of new drugs. As a result, significant uncertainty exists as to whether and how much third-party payers will reimburse patients for their use of newly approved drugs, which in turn will put pressure on the pricing of drugs.

***Price controls may be imposed in foreign markets, which may adversely affect our future profitability.***

In some countries, particularly member states of the European Union, the pricing of prescription drugs is subject to governmental control. In these countries, pricing negotiations with governmental authorities can take considerable time after receipt of marketing approval for a product. In addition, there can be considerable pressure by governments and other stakeholders on prices and reimbursement levels, including as part of cost containment measures. Political, economic and regulatory developments may further complicate pricing negotiations, and pricing negotiations may continue after reimbursement has been obtained. Reference pricing used by various European Union member states and parallel distribution, or arbitrage between low-priced and high-priced member states, can further reduce prices. In some countries, we or our partners may be required to conduct a clinical trial or other studies that compare the cost-effectiveness of our protein therapeutic candidates



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to other available therapies in order to obtain or maintain reimbursement or pricing approval. Publication of discounts by third-party payers or authorities may lead to further pressure on the prices or reimbursement levels within the country of publication and other countries. If reimbursement of our products is unavailable or limited in scope or amount, or if pricing is set at unsatisfactory levels, our business could be adversely affected.

***The impact of recent healthcare reform legislation and other changes in the healthcare industry and in healthcare spending on us is currently unknown, and may adversely affect our business model.***

Our revenue prospects could be affected by changes in healthcare spending and policy in the United States and abroad. We operate in a highly regulated industry and new laws, regulations or judicial decisions, or new interpretations of existing laws, regulations or decisions, related to health care availability, the method of delivery or payment for health care products and services could negatively impact our business, operations and financial condition. There is significant interest in promoting health care reform, as evidenced by the enactment in the United States of the Patient Protection and Affordable Care Act and the Health Care and Education Reconciliation Act in 2010. It is likely that federal and state legislatures within the United States and foreign governments will continue to consider changes to existing health care legislation. We cannot predict the reform initiatives that may be adopted in the future or whether initiatives that have been adopted will be repealed or modified. The continuing efforts of the government, insurance companies, managed care organizations and other payers of healthcare services to contain or reduce costs of healthcare may adversely affect:

the demand for any drug products for which we may obtain regulatory approval;

our ability to set a price that we believe is fair for our products;

our ability to obtain coverage and reimbursement approval for a product;

our ability to generate revenues and achieve or maintain profitability; and

the level of taxes that we are required to pay.

***We face substantial competition, which may result in others discovering, developing or commercializing products before, or more successfully, than we do.***

Our future success depends on our or our partners' ability to demonstrate and maintain a competitive advantage with respect to the design, development and commercialization of our protein therapeutic candidates. Our objective is to design, develop and commercialize new products with superior efficacy, convenience, tolerability and safety. In many cases, the protein therapeutics that we commercialize with our strategic partners or on our own will compete with existing, market-leading products.

There are products currently approved to treat patients with MDS, including iron chelation therapy, immunomodulators and various chemotherapeutic agents. In addition, erythropoiesis stimulating agents and red blood cell transfusions are extensively used to treat anemia in MDS. ACE-536 or sotatercept, if approved, will compete with these therapies. In addition, one or more products not currently approved for the treatment of anemia in MDS may in the future be granted marketing approval for the treatment of anemia in MDS or other conditions for which ACE-536 or sotatercept might be approved, including Aranesp®, being developed by Amgen, which is in Phase 3 trials. While there are currently no drug products approved for the treatment of anemia in  $\beta$ -thalassemia, red blood cell transfusions are extensively used and sotatercept or ACE-536, if approved, would compete with this therapy. In addition, HQK-1001, a fetal hemoglobin stimulating agent being developed by HemaQuest Pharmaceuticals, has completed a Phase 1/2 clinical trial and an investigator sponsored Phase 2 clinical trial in patients with  $\beta$ -thalassemia. Further, the future approval, in one or

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more regions, of a biosimilar product to one of our products could create substantial competition and have a material impact on our business.

Sotatercept or ACE-536, if approved for the treatment of chemotherapy-induced anemia or anemia of chronic kidney disease, would compete with erythropoiesis-stimulating agents, such as Epogen® and Aranesp®, marketed by Amgen, and Procrit®, marketed by Johnson & Johnson, that are currently approved for the treatment of chemotherapy-induced anemia or anemia of chronic kidney disease and other therapies in development including oral, small molecule treatments being developed by Astellas Pharma and Fibrogen designed to increase the body's production of erythropoietin.

While we anticipate that dalantercept, if approved for the treatment of cancer, would likely be approved in combination with certain VEGF pathway inhibitors that are currently approved for the treatment of various cancer types, dalantercept would compete with other products, including other angiogenesis inhibitors, approved for the treatment of these cancers.

Many of our potential competitors have significantly greater financial, manufacturing, marketing, drug development, technical and human resources than we do. Large pharmaceutical companies, in particular, have extensive experience in clinical testing, obtaining regulatory approvals, recruiting patients and in manufacturing pharmaceutical products. These companies also have significantly greater research and marketing capabilities than we do and may also have products that have been approved or are in late stages of development, and have collaborative arrangements in our target markets with leading companies and research institutions. Established pharmaceutical companies may also invest heavily to accelerate discovery and development of novel compounds or to in-license novel compounds that could make the protein therapeutics that we develop obsolete. As a result of all of these factors, our competitors may succeed in obtaining patent protection and/or FDA approval or discovering, developing and commercializing protein therapeutics before we do. In addition, any new product that competes with an approved product must demonstrate compelling advantages in efficacy, convenience, tolerability and safety in order to overcome price competition and to be commercially successful. If we are not able to compete effectively against potential competitors, our business will not grow and our financial condition and operations will suffer.

***Our protein therapeutics may cause undesirable side effects or have other properties that delay or prevent their regulatory approval or limit their commercial potential.***

Undesirable side effects caused by our protein therapeutics could cause us, Celgene or regulatory authorities to interrupt, delay or halt clinical trials and could result in the denial of regulatory approval by the FDA or other regulatory authorities and potential products liability claims. We and Celgene are currently conducting a number of Phase 2 trials for our clinical stage protein therapeutic candidates. Serious adverse events deemed to be caused by our protein therapeutics could have a material adverse effect on the development of our protein therapeutic candidates and our business as a whole. The most common adverse event to date in the clinical trials evaluating the safety and efficacy of our protein therapeutic candidates has been hypertension in our clinical trials for sotatercept and fluid retention in our clinical trials for dalantercept. Our understanding of the relationship between our protein therapeutic candidates and these events may change as we gather more information, and additional unexpected adverse events may occur. There can be no assurance that additional adverse events associated with our protein therapeutic candidates will not be observed. Recently, we discontinued the development of ACE-031, another of our clinical stage protein therapeutics that binds to ligands of the TGF- $\beta$  superfamily. Clinical development of ACE-031 was put on clinical hold in 2011 due to preliminary safety observations in patients. After gathering further information from animal toxicology studies, we and our ACE-031 partner, Shire AG, determined that the risk-benefit profile of ACE-031 was not appropriate for the intended patient population, boys aged four and older with a genetic muscle wasting disease, and we discontinued development of this protein therapeutic candidate. As is typical in drug development, we have a program of ongoing toxicology studies in animals for our other clinical stage protein therapeutics and cannot provide assurance that the findings from such studies or any ongoing or future clinical trials will not adversely affect our clinical development activities.



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If we or others identify undesirable side effects caused by our protein therapeutic candidates either before or after receipt of marketing approval, a number of potentially significant negative consequences could result, including:

our clinical trials may be put on hold;

we or our partners may be unable to obtain regulatory approval for our protein therapeutic candidates;

regulatory authorities may withdraw approvals of our protein therapeutic candidates;

regulatory authorities may require additional warnings on the label;

a medication guide outlining the risks of such side effects for distribution to patients may be required;

we could be sued and held liable for harm caused to patients; and

our reputation may suffer.

Any of these events could prevent us from achieving or maintaining market acceptance of our protein therapeutics and could substantially increase commercialization costs.

**Risks Related to Our Business and Industry**

*If we fail to attract and keep senior management and key scientific personnel, we may be unable to successfully develop our protein therapeutics, conduct our clinical trials and commercialize our protein therapeutic candidates.*

We are highly dependent on members of our senior management, including John L. Knopf, Ph.D., our Chief Executive Officer and President and one of our founders. The loss of the services of any of these persons could impede the achievement of our research, development and commercialization objectives. We do not maintain "key person" insurance for any of our executives or other employees.

Recruiting and retaining qualified scientific, clinical, manufacturing, sales and marketing personnel will also be critical to our success. We may not be able to attract and retain these personnel on acceptable terms given the competition among numerous pharmaceutical and biotechnology companies for similar personnel. We also experience competition for the hiring of scientific and clinical personnel from universities and research institutions. In addition, we rely on consultants and advisors, including scientific and clinical advisors, to assist us in formulating our research and development and commercialization strategy. Our consultants and advisors, including our scientific co-founders, may be employed by employers other than us and may have commitments under consulting or advisory contracts with other entities that may limit their availability to us.

*Our employees may engage in misconduct or other improper activities, including noncompliance with regulatory standards and requirements and insider trading.*

We are exposed to the risk of employee fraud or other misconduct. Misconduct by employees could include intentional failures to comply with FDA regulations, to provide accurate information to the FDA, to comply with manufacturing standards we have established, to comply with federal and state healthcare fraud and abuse laws and regulations, to report financial information or data accurately or to disclose unauthorized activities to us. In particular, sales, marketing and business arrangements in the healthcare industry are subject to extensive laws and regulations intended to prevent fraud, kickbacks, self-dealing and other abusive practices. These laws and regulations may restrict or prohibit a wide range of pricing, discounting, marketing and promotion, sales commission, customer incentive programs and other business arrangements. Employee misconduct could also involve the improper use of information obtained in the course of clinical trials, which could result in regulatory sanctions and



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serious harm to our reputation. It is not always possible to identify and deter employee misconduct, and the precautions we take to detect and prevent this activity may not be effective in controlling unknown or unmanaged risks or losses or in protecting us from governmental investigations or other actions or lawsuits stemming from a failure to be in compliance with such laws or regulations. If any such actions are instituted against us, and we are not successful in defending ourselves or asserting our rights, those actions could have a significant impact on our business, including the imposition of significant fines or other sanctions.

***We may encounter difficulties in managing our growth and expanding our operations successfully.***

As we seek to advance our protein therapeutic candidates through clinical trials and commercialization, we will need to expand our development, regulatory, manufacturing, marketing and sales capabilities or contract with third parties to provide these capabilities for us. As our operations expand, we expect that we will need to manage additional relationships with various strategic partners, suppliers and other third parties. Future growth will impose significant added responsibilities on members of management. Our future financial performance and our ability to commercialize our protein therapeutic candidates and to compete effectively will depend, in part, on our ability to manage any future growth effectively. To that end, we must be able to manage our development efforts and clinical trials effectively and hire, train and integrate additional management, administrative and, if necessary, sales and marketing personnel. We may not be able to accomplish these tasks, and our failure to accomplish any of them could prevent us from successfully growing our company.

***If product liability lawsuits are brought against us, we may incur substantial liabilities and may be required to limit commercialization of our protein therapeutics.***

We face an inherent risk of product liability as a result of the clinical testing of our protein therapeutic candidates and will face an even greater risk if we commercialize any products. For example, we may be sued if any product we develop allegedly causes injury or is found to be otherwise unsuitable during product testing, manufacturing, marketing or sale. Any such product liability claims may include allegations of defects in manufacturing, defects in design, a failure to warn of dangers inherent in the product, negligence, strict liability, and a breach of warranties. Claims could also be asserted under state consumer protection acts. If we cannot successfully defend ourselves against product liability claims, we may incur substantial liabilities or be required to limit commercialization of our protein therapeutic candidates. Even a successful defense would require significant financial and management resources. Regardless of the merits or eventual outcome, liability claims may result in:

injury to our reputation;

withdrawal of clinical trial participants;

costs to defend the related litigations;

a diversion of management's time and our resources;

substantial monetary awards to trial participants or patients;

product recalls, withdrawals, or labeling, marketing or promotional restrictions;

loss of revenue;

the inability to commercialize our protein therapeutic candidates; and

a decline in our stock price.

Failure to obtain and retain sufficient product liability insurance at an acceptable cost to protect against potential product liability claims could prevent or inhibit the commercialization of products we develop. We currently carry product liability insurance covering our clinical trials in the amount of

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\$10 million in the aggregate. Although we maintain such insurance, any claim that may be brought against us could result in a court judgment or settlement in an amount that is not covered, in whole or in part, by our insurance or that is in excess of the limits of our insurance coverage. Our insurance policies also have various exclusions, and we may be subject to a product liability claim for which we have no coverage. We will have to pay any amounts awarded by a court or negotiated in a settlement that exceed our coverage limitations or that are not covered by our insurance, and we may not have, or be able to obtain, sufficient capital to pay such amounts.

***We must comply with environmental laws and regulations, and failure to comply with these laws and regulations could expose us to significant liabilities.***

We use hazardous chemicals and radioactive and biological materials in certain aspects of our business and are subject to a variety of federal, state and local laws and regulations governing the use, generation, manufacture, distribution, storage, handling, treatment and disposal of these materials. Although we believe our safety procedures for handling and disposing of these materials and waste products comply with these laws and regulations, we cannot eliminate the risk of accidental injury or contamination from the use, manufacture, distribution, storage, handling, treatment or disposal of hazardous materials. In the event of contamination or injury, or failure to comply with environmental, occupational health and safety and export control laws and regulations, we could be held liable for any resulting damages and any such liability could exceed our assets and resources. We are uninsured for third-party contamination injury.

**Risks Related to Our Common Stock and This Offering**

***We are eligible to be treated as an "emerging growth company" as defined in the Jumpstart Our Business Startups Act of 2012, and we cannot be certain if the reduced disclosure requirements applicable to emerging growth companies will make our common stock less attractive to investors.***

We are an "emerging growth company", as defined in the Jumpstart Our Business Startups Act of 2012, or the JOBS Act. For as long as we continue to be an emerging growth company, we may take advantage of exemptions from various reporting requirements that are applicable to other public companies that are not emerging growth companies, including (1) not being required to comply with the auditor attestation requirements of Section 404 of the Sarbanes-Oxley Act of 2002, which we refer to as the Sarbanes-Oxley Act, (2) reduced disclosure obligations regarding executive compensation in this prospectus and our periodic reports and proxy statements and (3) exemptions from the requirements of holding a nonbinding advisory vote on executive compensation and stockholder approval of any golden parachute payments not previously approved. In addition, as an emerging growth company, we are only required to provide two years of audited financial statements and two years of selected financial data in this prospectus. We could be an emerging growth company for up to five years, although circumstances could cause us to lose that status earlier, including if the market value of our common stock held by non-affiliates exceeds \$700.0 million as of any June 30 before that time or if we have total annual gross revenue of \$1.0 billion or more during any fiscal year before that time, in which cases we would no longer be an emerging growth company as of the following December 31 or, if we issue more than \$1.0 billion in non-convertible debt during any three-year period before that time, we would cease to be an emerging growth company immediately. We cannot predict if investors will find our common stock less attractive because we may rely on these exemptions. If some investors find our common stock less attractive as a result, there may be a less active trading market for our common stock and our stock price may be more volatile.

Under the JOBS Act, emerging growth companies can also delay adopting new or revised accounting standards until such time as those standards apply to private companies. We have irrevocably elected not to avail ourselves of this exemption from new or revised accounting standards and, therefore, will be subject to the same new or revised accounting standards as other public companies that are not emerging growth companies.

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***Our stock price could be highly volatile, and, as a result you may not be able to resell your shares at or above the public offering price.***

Since our initial public offering in September 2013, the price of our common stock as reported on The NASDAQ Global Market has ranged from a low of \$16.78 on November 6 and 8, 2013 to a high of \$57.89 on January 22, 2014. The market price of shares of our common stock could be subject to wide fluctuations in response to many risk factors listed in this section, and others beyond our control, including:

results of clinical trials of our protein therapeutic candidates, including sotatercept, ACE-536 and dalantercept;

the timing of the release of results of our clinical trials that are being conducted by Celgene;

results of clinical trials of our competitors' products;

regulatory actions with respect to our products or our competitors' products;

actual or anticipated fluctuations in our financial condition and operating results;

publication of research reports by securities analysts about us or our competitors or our industry;

our failure or the failure of our competitors to meet analysts' projections or guidance that we or our competitors may give to the market;

additions and departures of key personnel;

strategic decisions by us or our competitors, such as acquisitions, divestitures, spin-offs, joint ventures, strategic investments or changes in business strategy;

the passage of legislation or other regulatory developments affecting us or our industry;

fluctuations in the valuation of companies perceived by investors to be comparable to us;

sales of our common stock by us, our insiders or our other stockholders;

speculation in the press or investment community;

announcement or expectation of additional financing efforts;

changes in accounting principles;

terrorist acts, acts of war or periods of widespread civil unrest;

natural disasters and other calamities;

changes in market conditions for biopharmaceutical stocks; and

changes in general market and economic conditions.

In addition, the stock market has recently experienced significant volatility, particularly with respect to pharmaceutical, biotechnology and other life sciences company stocks. The volatility of pharmaceutical, biotechnology and other life sciences company stocks often does not relate to the operating performance of the companies represented by the stock. As we operate in a single industry, we are especially vulnerable to these factors to the extent that they affect our industry or our products, or to a lesser extent our markets. In the past, securities class action litigation has often been initiated against companies following periods of volatility in their stock price. This type of litigation could result in substantial costs and divert our management's attention and resources, and could also require us to make substantial payments to satisfy judgments or to settle litigation.

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***Our principal stockholders and management own a significant percentage of our stock and will be able to exercise significant influence over matters subject to stockholder approval.***

As of January 1, 2014, our executive officers, directors and principal stockholders, together with their respective affiliates, beneficially owned approximately 60.7% of our common stock, including shares subject to outstanding options and warrants that are exercisable within 60 days after such date, and we expect that upon completion of this offering, that same group will continue to beneficially hold at least 56.3% of our outstanding common stock. Accordingly, even after this offering, these stockholders will be able to exert a significant degree of influence over our management and affairs and over matters requiring stockholder approval, including the election of our board of directors and approval of significant corporate transactions. This concentration of ownership could have the effect of entrenching our management and/or the board of directors, delaying or preventing a change in our control or otherwise discouraging a potential acquirer from attempting to obtain control of us, which in turn could have a material and adverse effect on the fair market value of our common stock.

***A significant portion of our total outstanding shares may be sold into the public market in the near future, which could cause the market price of our common stock to drop significantly, even if our business is doing well.***

Sales of a substantial number of shares of our common stock in the public market, or the market perception that the holders of a large number of shares intend to sell shares, could reduce the market price of our common stock. As of January 1, 2014, based on the number of shares of our common stock then outstanding, assuming (1) the closing of this offering, (2) no exercise of the underwriters' option to purchase additional shares of common stock, and (3) no exercise of outstanding options or warrants, we would have had outstanding an aggregate of 30,748,633 shares of common stock. Of these shares, 7,482,723 shares of common stock, including the 6,417,000 shares sold in our initial public offering, and all of the shares of common stock to be sold in this offering, will be freely tradable in the public market without restriction or further registration under the Securities Act of 1933, as amended, or the Securities Act, unless the shares are held by any of our "affiliates" as such term is defined in Rule 144 of the Securities Act. The remaining shares of common stock are "restricted securities" as such term is defined in Rule 144 or are subject to lock up agreements in effect in connection with the initial public offering or entered into in connection with this offering and will be available for sale in the public market are as follows:

<b>Number of Shares and % of Total Outstanding</b>	<b>Date Available for Sale into Public Market</b>
5,896,337 shares, or 19%	March 17, 2014 due to lock up agreements in effect in connection with our initial public offering. However, the representatives of the underwriters can waive the provisions of these lock-up agreements and allow these stockholders to sell their shares at any time.
15,269,537 shares, or 50%	90 days after the date of this prospectus, due to lock-up agreements between the holders of these shares and the underwriters. However, the representatives of the underwriters can waive the provisions of these lock-up agreements and allow these stockholders to sell their shares at any time.

In addition, as of January 1, 2014, there were 979,699 shares subject to outstanding warrants, 3,942,304 shares subject to outstanding options and an additional 2,089,945 shares reserved for future issuance under our employee benefit plans that will become eligible for sale in the public market to the extent permitted by any applicable vesting requirements, the lock-up agreements and Rules 144 and 701 under the Securities Act of 1933, as amended. Moreover, after this offering, holders of an aggregate of approximately 14.8 million shares of our common stock and holders of warrants to purchase 540,097



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shares of our common stock will have rights, subject to some conditions, to require us to file registration statements covering their shares or to include their shares in registration statements that we may file for ourselves or other stockholders. If such holders, by exercising their registration rights, cause a large number of securities to be registered and sold into the public market, these sales could have an adverse effect on the market price for our common stock. These rights have been waived in connection with this offering pursuant to the terms of the registration rights agreement. We have also registered all shares of common stock that we may issue under our 2013 Equity Incentive Plan, and intend to register annual increases pursuant to this plan on a post effective amendment to the registration statement. Once these shares are issued in accordance with the terms of the plans, they can be freely sold in the public market upon issuance, subject to the lock-up agreements and the restrictions imposed on our affiliates under Rule 144. For more information, see "Shares Eligible for Future Sale Rule 144".

***You will incur immediate and substantial dilution as a result of this offering.***

If you purchase common stock in this offering, you will incur immediate and substantial dilution of \$43.87 per share, representing the difference between the public offering price of \$50.00 per share and our as adjusted net tangible book value per share as of September 30, 2013 after giving effect to this offering. Moreover, in the past we issued warrants and options to acquire common stock at prices significantly below the public offering price. As of January 1, 2014, there were 979,699 shares subject to outstanding warrants and 3,942,304 shares subject to outstanding options. To the extent that these outstanding warrants or options are ultimately exercised, you will incur further dilution.

***We have broad discretion in the use of net proceeds from this offering and may not use them effectively.***

We currently intend to use the net proceeds from this offering to fund the continued development of dalantercept and ACE-083 and to continue to discover and develop other protein therapeutics in our pipeline and for working capital and other general corporate purposes, including funding the costs of operating a public company. See "Use of Proceeds." Any remaining amounts will be used for general corporate purposes, general and administrative expenses, capital expenditures, working capital and prosecution and maintenance of our intellectual property. Although we currently intend to use the net proceeds from this offering in such a manner, we will have broad discretion in the application of the net proceeds. Our failure to apply these funds effectively could affect our ability to continue to develop and commercialize our protein therapeutic candidates.

***We are incurring significant increased costs as a result of operating as a public company, and our management is required to devote substantial time to new compliance initiatives.***

As a newly public company, we are incurring significant legal, accounting and other expenses that we did not incur as a private company. In addition, the Sarbanes-Oxley Act, and rules of the SEC and those of The NASDAQ Global Market, or NASDAQ, have imposed various requirements on public companies including requiring establishment and maintenance of effective disclosure and financial controls. Our management and other personnel will need to devote a substantial amount of time to these compliance initiatives. Moreover, these rules and regulations have increased and will continue to increase our legal and financial compliance costs and will make some activities more time-consuming and costly.

The Sarbanes-Oxley Act requires, among other things, that we maintain effective internal control over financial reporting and disclosure controls and procedures. In particular, we must perform system and process evaluation and testing of our internal control over financial reporting to allow management to report on the effectiveness of our internal control over financial reporting, as required by Section 404 of the Sarbanes-Oxley Act, beginning with our annual report on Form 10-K for the fiscal year ended December 31, 2013. In addition, we will be required to have our independent registered

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public accounting firm attest to the effectiveness of our internal control over financial reporting beginning with our annual report on Form 10-K following the date on which we are no longer an emerging growth company. Our compliance with Section 404 of the Sarbanes-Oxley Act will require that we incur substantial accounting expense and expend significant management efforts. We currently do not have an internal audit group, and we will need to hire additional accounting and financial staff with appropriate public company experience and technical accounting knowledge. If we are not able to comply with the requirements of Section 404 in a timely manner, or if we or our independent registered public accounting firm identify deficiencies in our internal control over financial reporting that are deemed to be material weaknesses, the market price of our stock could decline and we could be subject to sanctions or investigations by NASDAQ, the SEC or other regulatory authorities, which would require additional financial and management resources.

Our ability to successfully implement our business plan and comply with Section 404 requires us to be able to prepare timely and accurate financial statements. We expect that we will need to continue to improve existing, and implement new operational and financial systems, procedures and controls to manage our business effectively. Any delay in the implementation of, or disruption in the transition to, new or enhanced systems, procedures or controls, may cause our operations to suffer and we may be unable to conclude that our internal control over financial reporting is effective and to obtain an unqualified report on internal controls from our auditors as required under Section 404 of the Sarbanes-Oxley Act. This, in turn, could have an adverse impact on trading prices for our common stock, and could adversely affect our ability to access the capital markets.

***We do not expect to pay any cash dividends for the foreseeable future.***

You should not rely on an investment in our common stock to provide dividend income. We do not anticipate that we will pay any cash dividends to holders of our common stock in the foreseeable future. Instead, we plan to retain any earnings to maintain and expand our operations. In addition, our ability to pay cash dividends is currently prohibited by the terms of our debt financing arrangement, and any future debt financing arrangement may contain terms prohibiting or limiting the amount of dividends that may be declared or paid on our common stock. Accordingly, investors must rely on sales of their common stock after price appreciation, which may never occur, as the only way to realize any return on their investment. As a result, investors seeking cash dividends should not purchase our common stock.

***Provisions in our restated certificate of incorporation, our amended and restated by-laws and Delaware law may have anti-takeover effects that could discourage an acquisition of us by others, even if an acquisition would be beneficial to our stockholders, and may prevent attempts by our stockholders to replace or remove our current management.***

Our restated certificate of incorporation, amended and restated by-laws and Delaware law contain provisions that may have the effect of delaying or preventing a change in control of us or changes in our management. Our restated certificate of incorporation and by-laws include provisions that:

authorize "blank check" preferred stock, which could be issued by our board of directors without stockholder approval and may contain voting, liquidation, dividend and other rights superior to our common stock;

create a classified board of directors whose members serve staggered three-year terms;

specify that special meetings of our stockholders can be called only by our board of directors;

prohibit stockholder action by written consent;

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establish an advance notice procedure for stockholder approvals to be brought before an annual meeting of our stockholders, including proposed nominations of persons for election to our board of directors;

provide that our directors may be removed only for cause;

provide that vacancies on our board of directors may be filled only by a majority of directors then in office, even though less than a quorum;

specify that no stockholder is permitted to cumulate votes at any election of directors;

expressly authorize our board of directors to modify, alter or repeal our amended and restated by-laws; and

require supermajority votes of the holders of our common stock to amend specified provisions of our restated certificate of incorporation and amended and restated by-laws

These provisions, alone or together, could delay or prevent hostile takeovers and changes in control or changes in our management.

In addition, because we are incorporated in the state of Delaware, we are governed by the provisions of Section 203 of the Delaware General Corporation Law, which limits the ability of stockholders owning in excess of 15% of our outstanding voting stock to merge or combine with us.

Any provision of our restated certificate of incorporation or amended and restated by-laws or Delaware law that has the effect of delaying or deterring a change in control could limit the opportunity for our stockholders to receive a premium for their shares of our common stock, and could also affect the price that some investors are willing to pay for our common stock.

***Our restated certificate of incorporation designates the Court of Chancery of the State of Delaware and federal court within the State of Delaware as the exclusive forum for certain types of actions and proceedings that may be initiated by our stockholders, which could limit our stockholders' ability to obtain a favorable judicial forum for disputes with us or our directors, officers or employees.***

Our restated certificate of incorporation provides that, subject to limited exceptions, the Court of Chancery of the State of Delaware and federal court within the State of Delaware will be exclusive forums for (1) any derivative action or proceeding brought on our behalf, (2) any action asserting a claim of breach of a fiduciary duty owed by any of our directors, officers or other employees to us or our stockholders, (3) any action asserting a claim against us arising pursuant to any provision of the Delaware General Corporation Law, our restated certificate of incorporation or our amended and restated by-laws, or (4) any other action asserting a claim against us that is governed by the internal affairs doctrine. Any person or entity purchasing or otherwise acquiring any interest in shares of our capital stock shall be deemed to have notice of and to have consented to the provisions of our restated certificate of incorporation described above. This choice of forum provision may limit a stockholder's ability to bring a claim in a judicial forum that it finds favorable for disputes with us or our directors, officers or other employees, which may discourage such lawsuits against us and our directors, officers and employees. Alternatively, if a court were to find these provisions of our restated certificate of incorporation inapplicable to, or unenforceable in respect of, one or more of the specified types of actions or proceedings, we may incur additional costs associated with resolving such matters in other jurisdictions, which could adversely affect our business and financial condition.

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**Cautionary Note Regarding Forward-Looking Statements**

This prospectus contains forward-looking statements. Forward-looking statements are neither historical facts nor assurances of future performance. Instead, they are based on our current beliefs, expectations and assumptions regarding the future of our business, future plans and strategies, our clinical results and other future conditions. The words "anticipate", "believe", "estimate", "expect", "forecast", "goal", "intend", "may", "plan", "predict", "project", "target", "potential", "will", "would", "could", "should", "continue", "contemplate", or the negative of these terms or other similar expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words.

The forward-looking statements in this prospectus include, among other things, statements about:

the timing of results of our ongoing clinical trials;

our plans to develop and commercialize dalantercept and ACE-083 and our and Celgene's plans to develop and commercialize sotatercept and ACE-536;

the potential benefits of strategic partnership agreements and our ability to enter into selective strategic partnership arrangements;

the timing of, and our and Celgene's ability to, obtain and maintain regulatory approvals for our protein therapeutic candidates;

the rate and degree of market acceptance and clinical utility of any approved protein therapeutic candidate;

our ability to quickly and efficiently identify and develop protein therapeutic candidates;

our commercialization, marketing and manufacturing capabilities and strategy;

our intellectual property position; and

our estimates regarding expenses, future revenues, capital requirements, the sufficiency of our current and expected cash resources and our need for additional financing.

We may not actually achieve the plans, intentions or expectations disclosed in our forward-looking statements, and you should not place undue reliance on our forward-looking statements. Actual results or events could differ materially from the plans, intentions and expectations disclosed in the forward-looking statements we make. We have included important factors in the cautionary statements included in this prospectus, particularly in the "Risk Factors" section, that we believe could cause actual results or events to differ materially from the forward-looking statements that we make. Our forward-looking statements do not reflect the potential impact of any future acquisitions, mergers, dispositions, joint ventures or investments we may make.

The forward-looking statements in this prospectus represent our views as of the date of this prospectus. We anticipate that subsequent events and developments will cause our views to change. However, while we may elect to update these forward-looking statements at some point in the future, we have no current intention of doing so except to the extent required by applicable law. You should, therefore, not rely on these forward-looking statements as representing our views as of any date subsequent to the date of this prospectus.



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**USE OF PROCEEDS**

The net proceeds of the sale of 2,400,000 shares of common stock in this offering will be approximately \$112.1 million at the public offering price of \$50.00 per share, after deducting underwriting discounts and commissions and estimated offering expenses payable by us. If the underwriters exercise their option to purchase additional shares of common stock in full, we estimate that the net proceeds will be approximately \$129.0 million, after deducting underwriting discounts and commissions and estimated offering expenses payable by us.

We intend to use the net proceeds from this offering as follows:

approximately \$57.0 million to continue development of dalantercept, including initiation of additional Phase 2 clinical trials of dalantercept in combination with either an approved anti-angiogenesis agent or chemotherapy in advanced solid tumors, and obtaining the supply of dalantercept for Phase 3 clinical studies;

approximately \$8.0 million to conduct clinical trials and associated activities with a new protein therapeutic candidate ACE-083;

approximately \$15.0 million to continue to advance and to expand our preclinical research pipeline of protein therapeutic candidates; and

use the remainder for general and administrative expenses (including personnel-related costs), potential future development programs, early-stage research and development, capital expenditures and working capital and other general corporate purposes.

The expected use of the net proceeds from this offering represents our intentions based upon our current plans and business conditions, which could change in the future as our plans and business conditions evolve. The amounts and timing of our actual expenditures depend on numerous factors, including the ongoing status of and results from our clinical trials and other studies, the progress of our preclinical development efforts and any unforeseen cash needs. As a result, our management will have broad discretion in applying the net proceeds of this offering. Although we may use a portion of the net proceeds of this offering for the acquisition or licensing, as the case may be, of product candidates, technologies, compounds, other assets or complementary businesses, we have no current understandings, agreements or commitments to do so.

Pending the use of the proceeds from this offering, we intend to invest the net proceeds in short-term, interest-bearing, investment-grade securities, certificates of deposit or government securities.

Table of Contents**MARKET PRICE OF OUR COMMON STOCK**

Our common stock has been listed on The NASDAQ Global Market under the symbol "XLRN" since September 19, 2013. Prior to that, there was no public market for our common stock. The following table sets forth for the periods indicated the high and low sales prices per share of our common stock as reported on The NASDAQ Global Market:

<b>Year ended December 31, 2013:</b>	<b>High</b>	<b>Low</b>
Third quarter(1)	\$ 23.41	\$ 18.50
Fourth quarter	\$ 40.02	\$ 16.78
<b>Year ending December 31, 2014:</b>		
First quarter (through January 22, 2014)	\$ 57.89	\$ 36.86

(1) Represents the period from September 19, 2013, the date on which our common stock first began to trade on The NASDAQ Global Market after the pricing of our initial public offering, through September 30, 2013, the end of our third fiscal quarter.

A recent reported closing price for our common stock is set forth on the cover of this prospectus. Computershare Trust Company, N.A. is the transfer agent and registrar for our common stock. As of January 2, 2014, there were 174 holders of record of our common stock.

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**DIVIDEND POLICY**

We have never declared or paid cash dividends on our common stock. We currently intend to retain all available funds and any future earnings, if any, to fund the development and expansion of our business and we do not anticipate paying any cash dividends in the foreseeable future. In addition, our ability to pay cash dividends is currently prohibited by the terms of our debt financing arrangements, and any future debt financing arrangement may contain terms prohibiting or limiting the amount of dividends that may be declared or paid on our common stock. Any future determination to pay dividends will be made at the discretion of our board of directors.



Table of Contents**CAPITALIZATION**

The following table sets forth our cash and cash equivalents and capitalization as of September 30, 2013:

on an actual basis;

on an as adjusted basis to reflect the sale of shares of our common stock offered in this offering at the public offering price of \$50.00 per share, after deducting underwriting discounts and commissions and estimated offering expenses payable by us.

You should read this information together with our audited financial statements and related notes appearing elsewhere in this prospectus and the information set forth under the heading "Selected Financial Data" and "Management's Discussion and Analysis of Financial Condition and Results of Operations".

	<b>As of September 30, 2013</b>	
	<b>Actual</b>	<b>As adjusted</b>
	<b>(in thousands, except share and per share data)</b>	
Cash and cash equivalents	\$ 116,479	\$ 228,579
Notes payable, net of current portion	\$ 10,979	\$ 10,979
Warrants to purchase common stock	16,526	16,526
Stockholders' equity:		
Undesignated preferred stock, \$0.001 par value: 25,000,000 shares authorized and no shares issued or outstanding		
Common stock, \$0.001 par value; 175,000,000 shares authorized actual and as adjusted; 28,069,579 shares issued and outstanding, actual, and 30,469,579 shares issued and outstanding, as adjusted(1)	35	37
Additional paid-in capital	248,750	360,848
Accumulated deficit	(174,221)	(174,221)
Total stockholders' equity	74,564	186,664
Total capitalization	\$ 102,069	\$ 214,169

(1)

The actual and as adjusted information set forth in the table excludes (i) 3,655,968 shares of common stock issuable upon exercise of stock options outstanding as of September 30, 2013 at a weighted-average exercise price of \$4.18 per share, (ii) 1,011,590 shares of common stock issuable upon the exercise of warrants to purchase shares of common stock outstanding as of September 30, 2013 at a weighted-average exercise price of \$6.56 per share, (iii) 1,500,000 shares of common stock reserved for future issuance under our 2013 Equity Incentive Plan as of September 30, 2013, and (iv) 275,000 shares of common stock reserved for future issuance under our Employee Stock Purchase Plan as of September 30, 2013.

Table of Contents**SELECTED FINANCIAL DATA**

The information set forth below should be read in conjunction with the "Management's Discussion and Analysis of Financial Condition and Results of Operations" section of this prospectus and with our financial statements and notes thereto included elsewhere in this prospectus. The selected financial data in this section are not intended to replace the financial statements and are qualified in their entirety by the financial statements and related notes included elsewhere in this prospectus.

The selected statements of operations and comprehensive income (loss) data for the years ended December 31, 2011 and 2012 and the balance sheet data as of December 31, 2011 and 2012 have been derived from our audited financial statements included elsewhere in this prospectus. The selected statements of operations and comprehensive income (loss) data for the nine months ended September 30, 2012 and 2013 and the balance sheet data as of September 30, 2013 have been derived from our unaudited financial statements included elsewhere in this prospectus. In our opinion, these unaudited financial statements have been prepared on a basis consistent with our audited financial statements and contain all adjustments, consisting only of normal and recurring adjustments, necessary for a fair presentation of such financial data. Our historical results for any prior period are not necessarily indicative of results to be expected in any future period, and our interim period results are not necessarily indicative of results to be expected for a full year or any other interim period.

(in thousands, except per share data)	Year Ended December 31,		Nine Months Ended September 30,	
	2011	2012	2012	2013
Revenue:				
Collaboration revenue:				
License and milestone	\$ 74,406	\$ 9,696	\$ 7,226	\$ 36,044
Cost-sharing, net	4,760	5,558	4,043	9,666
Contract manufacturing	1,745			
Total revenue	80,911	15,254	11,269	45,710
Costs and expenses:				
Research and development	32,713	35,319	25,646	25,834
General and administrative	8,142	8,824	6,318	9,472
Cost of contract manufacturing revenue	1,500			
Total costs and expenses	42,355	44,143	31,964	35,306
Income (loss) from operations	38,556	(28,889)	(20,695)	10,404
Total other expense, net	(2,290)	(3,693)	(1,508)	(14,192)
Net income (loss)	\$ 36,266	\$ (32,582)	\$ (22,203)	\$ (3,788)
Comprehensive income (loss)	\$ 36,266	\$ (32,582)	\$ (22,203)	\$ (3,788)
Net income (loss) per share applicable to common stockholders(1)				
Basic	\$ 0.80	\$ (24.84)	\$ (17.73)	\$ 6.74
Diluted	\$ 0.78	\$ (24.84)	\$ (17.73)	\$ 6.74
Weighted-average number of common shares used in computing net income (loss) per share applicable to common stockholders				
Basic	2,328	2,401	2,397	3,100
Diluted	2,716	2,401	2,397	3,100

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(in thousands)	December 31,		September 30,
	2011	2012	2013
<b>Balance Sheet Data:</b>			
Cash and cash equivalents	\$ 65,037	\$ 39,611	\$ 116,479
Total assets	73,789	49,212	127,260
Total current liabilities	23,853	38,802	16,523
Long term deferred revenue	33,350	6,760	6,205
Long-term notes payable		16,525	10,979
Warrants to purchase redeemable convertible preferred stock	1,046	1,422	
Warrants to purchase common stock	3,347	5,229	16,526
Redeemable convertible preferred stock	241,549	268,610	
Total stockholder's (deficit) equity	(232,691)	(290,973)	74,564

(1)

See Note 2 within the notes to our financial statements appearing elsewhere in this prospectus for a description of the method used to calculate basic and diluted net income (loss) per common share and pro forma basic and diluted net income (loss) per common share.

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**MANAGEMENT'S DISCUSSION AND ANALYSIS OF  
FINANCIAL CONDITION AND RESULTS OF OPERATIONS**

*You should read the following discussion and analysis of financial condition and results of operations together with the section entitled "Selected Financial Data" and our financial statements and related notes included elsewhere in this prospectus. This discussion and other parts of this prospectus contain forward-looking statements that involve risk and uncertainties, such as statements of our plans, objectives, expectations and intentions. Our actual results could differ materially from those discussed in these forward-looking statements. Factors that could cause or contribute to such differences include, but are not limited to, those discussed in the "Risk Factors" section.*

**Overview**

We are a clinical stage biopharmaceutical company focused on the discovery, development and commercialization of novel protein therapeutics for cancer and rare diseases. Our research focuses on the biology of the Transforming Growth Factor-Beta (TGF- $\beta$ ) protein superfamily, a large and diverse group of molecules that are key regulators in the growth and repair of tissues throughout the human body. We are leaders in understanding the biology of the TGF- $\beta$  superfamily and in targeting these pathways to develop important new medicines. By coupling our discovery and development expertise, including our proprietary knowledge of the TGF- $\beta$  superfamily, with our internal protein engineering and manufacturing capabilities, we have built a highly productive research & development platform that has generated innovative protein therapeutic candidates with novel mechanisms of action. These differentiated protein therapeutic candidates have the potential to significantly improve clinical outcomes for patients with cancer and rare diseases.

We have three internally discovered protein therapeutic candidates that are currently being studied in numerous ongoing Phase 2 clinical trials, focused on cancer and rare diseases. Our two most advanced protein therapeutic candidates, sotatercept and ACE-536, promote red blood cell production through a novel mechanism. Together with our collaboration partner, Celgene Corporation, which we refer to as Celgene, we are developing sotatercept and ACE-536 to treat anemia and associated complications in patients with  $\beta$ -thalassemia and myelodysplastic syndromes (MDS), red blood cell disorders that are generally unresponsive to currently approved drugs. Our third clinical stage protein therapeutic candidate, dalantercept, is designed to inhibit blood vessel formation through a mechanism that is distinct from, and potentially synergistic with, the dominant class of cancer drugs that inhibit blood vessel formation, the vascular endothelial growth factor (VEGF) pathway inhibitors. We are developing dalantercept primarily for use in combination with these successful products to produce better outcomes for cancer patients.

We are developing sotatercept and ACE-536 through our exclusive worldwide collaborations with Celgene. As of January 1, 2013, Celgene became responsible for paying 100% of worldwide development costs for both programs. We may receive up to \$560.0 million of potential development, regulatory and commercial milestone payments still outstanding and, if these protein therapeutic candidates are commercialized, we will receive a royalty on net sales in the low-to-mid 20% range. We also will co-promote sotatercept and ACE-536 in North America, if approved, for which our commercialization costs will be entirely funded by Celgene. We have not entered into a partnership for dalantercept and retain worldwide rights to this program.

As of September 30, 2013, our operations have been primarily funded by \$105.1 million in equity investments from venture investors, \$86.8 million in net proceeds from our initial public offering, \$49.2 million in equity investments from our partners and \$192.6 million in upfront payments, milestones, and net research and development payments from our strategic partners.

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We expect to continue to incur significant expenses and increasing operating losses over at least the next several years. We expect our expenses will increase substantially in connection with our ongoing activities, as we:

conduct clinical trials for dalantercept;

continue our preclinical studies and potential clinical development efforts of our existing preclinical protein therapeutic candidates;

continue research activities for the discovery of new protein therapeutics;

manufacture protein therapeutics for our preclinical studies and clinical trials;

seek regulatory approval for our protein therapeutics; and

operate as a public company.

We will not generate revenue from product sales unless and until we or a partner successfully complete development and obtain regulatory approval for one or more of our protein therapeutic candidates, which we expect will take a number of years and is subject to significant uncertainty. All current and future development and commercialization costs for sotatercept and ACE-536 are paid by Celgene. If we obtain regulatory approval for dalantercept or any future protein therapeutic candidate, we expect to incur significant commercialization expenses related to product sales, marketing, manufacturing and distribution to the extent that such costs are not paid by future partners. We will seek to fund our operations through the sale of equity, debt financings or other sources, including potential additional collaborations. However, we may be unable to raise additional funds or enter into such other arrangements when needed on favorable terms, or at all. If we fail to raise capital or enter into such other arrangements as, and when, needed, we may have to significantly delay, scale back or discontinue the development or commercialization of one or more of our protein therapeutics.

Our ability to generate product revenue and become profitable depends upon our and our partners' ability to successfully commercialize products. We expect to incur losses for the foreseeable future, and we expect these losses to increase as we continue our development of, and seek regulatory approvals for, our protein therapeutics and potentially begin to commercialize any approved products. For a description of the numerous risks and uncertainties associated with product development, see "Risk Factors".

## **Financial Operations Overview**

### **Revenue**

#### *Collaboration Revenue*

We have not generated any revenue from the sale of products. Our revenue to date has been predominantly derived from collaboration revenue, which includes license and milestone revenues and cost sharing revenue, generated through collaboration and license agreements with partners for the development and commercialization of our protein therapeutics. Cost sharing revenue represents amounts reimbursed by our collaboration partners for expenses incurred by us for research and development activities and, potentially, co-promotion activities, under our collaboration agreements. Cost sharing revenue is recognized in the period that the related activities are performed. To the extent that we reimburse collaborators for costs incurred in connection with activities performed by them, we record these costs as a reduction of cost-sharing revenue.

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***Contract Manufacturing Revenue***

We have generated contract manufacturing revenue in the past but have no current contract manufacturing arrangements. Contract manufacturing revenue consists of revenue received for producing bulk drug substance for third parties other than our partners.

**Costs and Expenses**

***Research and Development Expenses***

Research and development expenses consist primarily of costs directly incurred by us for the development of our protein therapeutic candidates, which include:

direct employee-related expenses, including salaries, benefits, travel and stock-based compensation expense of our research and development personnel;

expenses incurred under agreements with clinical research organizations, or CROs, and investigative sites that will conduct our clinical trials;

the cost of acquiring and manufacturing preclinical and clinical study materials and developing manufacturing processes;

allocated facilities, depreciation, and other expenses, which include rent and maintenance of facilities, insurance and other supplies;

expenses associated with obtaining and maintaining patents; and

costs associated with preclinical activities and regulatory compliance.

Research and development costs are expensed as incurred. Costs for certain development activities are recognized based on an evaluation of the progress to completion of specific tasks using information and data provided to us by our vendors and our clinical sites.

We cannot determine with certainty the duration and completion costs of the current or future clinical trials of our protein therapeutic candidates or if, when, or to what extent we will generate revenues from the commercialization and sale of any of our protein therapeutic candidates for which we or any partner obtain regulatory approval. We or our partners may never succeed in achieving regulatory approval for any of our protein therapeutic candidates. The duration, costs and timing of clinical trials and development of our protein therapeutic candidates will depend on a variety of factors, including:

the scope, rate of progress, and expense of our ongoing, as well as any additional, clinical trials and other research and development activities;

future clinical trial results;

potential changes in government regulation; and

the timing and receipt of any regulatory approvals.

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A change in the outcome of any of these variables with respect to the development of a protein therapeutic candidate could mean a significant change in the costs and timing associated with the development of that protein therapeutic candidate. For example, if the FDA, or another regulatory authority were to require us to conduct clinical trials beyond those that we currently anticipate will be required for the completion of the clinical development of protein therapeutics, or if we experience significant delays in the enrollment in any clinical trials, we could be required to expend significant additional financial resources and time on the completion of clinical development.

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From inception through September 30, 2013, we have incurred \$277.0 million in research and development expenses. We plan to increase our research and development expenses for the foreseeable future as we continue the development of our TGF- $\beta$  platform protein therapeutics, the discovery and development of preclinical protein therapeutics, including ACE-083, and the development of sotatercept, ACE-536 and dalantercept. Beginning January 1, 2013, expenses associated with sotatercept and ACE-536 are reimbursed 100% by Celgene. These reimbursements are recorded as revenue. Of the Phase 2 clinical trials that are underway for sotatercept, ACE-536 and dalantercept, we are expensing the costs of six clinical trials of ACE-536 and dalantercept, of which the two for ACE-536 are reimbursed by Celgene.

We manage certain activities such as clinical trial operations, manufacture of protein therapeutic candidates, and preclinical animal toxicology studies through third-party CROs. The only costs we track by each protein therapeutic candidate are external costs such as services provided to us by CROs, manufacturing of preclinical and clinical drug substance, and other outsourced research and development expenses. We do not assign or allocate to individual development programs internal costs such as salaries and benefits, facilities costs, lab supplies and the costs of preclinical research and studies. Our external research and development expenses for sotatercept, ACE-536, dalantercept and ACE-031 (for which development was suspended in April 2013) during the years ended December 31, 2011 and 2012 and the nine months ended September 30, 2012 and 2013 are as follows:

(in thousands)	Year Ended December 31,		Nine Months Ended September 30,	
	2011	2012	2012	2013
Sotatercept(1)	\$ 974	\$ 6	\$ 6	\$ 1
ACE-536(1)	681	2,885	2,047	3,182
Dalantercept	1,323	3,422	2,220	3,413
ACE-031(2)	4,240	3,453	2,442	997
Total direct research and development expenses	7,218	9,766	6,715	7,593
Other expenses(3)	25,495	25,553	18,931	18,241
Total research and development expenses	\$ 32,713	\$ 35,319	\$ 25,646	\$ 25,834

(1) Beginning January 1, 2013, expenses associated with sotatercept and ACE-536 are reimbursed 100% by Celgene. These reimbursements are recorded as revenue and are presented as cost-sharing, net.

(2) In April 2013, we and Shire AG, which we refer to as Shire, determined not to further advance the development of ACE-031, and Shire terminated our collaboration agreement, effective as of June 30, 2013.

(3) Other expenses include unallocated employee and contractor-related expenses, facility expenses and miscellaneous expenses.

***Contract Manufacturing Expenses***

Contract manufacturing expenses consist primarily of costs incurred for the production of bulk drug substance for third parties other than our partners. The costs generally include employee-related expenses including salary and benefits, direct materials and overhead costs including rent, depreciation, utilities, facility maintenance and insurance. We do not have any current contract manufacturing arrangements.



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***General and Administrative Expenses***

General and administrative expenses consist primarily of salaries and related costs for personnel, including stock-based compensation and travel expenses for our employees in executive, operational, finance and human resource functions and other general and administrative expenses including directors' fees and professional fees for accounting and legal services.

Since the completion of our initial public offering in September 2013, we have experienced increased expenses related to audit, legal, regulatory and tax-related services associated with maintaining compliance with exchange listing and Securities and Exchange Commission requirements, director and officer insurance premiums, and investor relations costs associated with being a public company. We anticipate that our general and administrative expenses will increase in the future as we increase our headcount to support our continued research and development and potential commercialization of our protein therapeutics. Additionally, if and when we believe regulatory approval of a protein therapeutic candidate appears likely, to the extent that we are undertaking commercialization of such protein therapeutic candidate ourselves, we anticipate an increase in payroll and related expenses as a result of our preparation for commercial operations.

***Other Expense, Net***

Other expense, net consists primarily of interest expense from our venture debt facility, interest income earned on cash and cash equivalents, and the re-measurement gain or loss associated with the change in the fair value of our preferred stock and common stock warrant liabilities.

We use the Black-Scholes option pricing model to estimate the fair value of the warrants. We base the estimates in the Black-Scholes option pricing model, in part, on subjective assumptions, including stock price volatility, risk-free interest rate, dividend yield, and the fair value of the preferred stock or common stock underlying the warrants.

**Critical Accounting Policies and Significant Judgments and Estimates**

Our management's discussion and analysis of our financial condition and results of operations are based on our financial statements, which have been prepared in accordance with U.S. generally accepted accounting principles. The preparation of these financial statements requires us to make estimates and judgments that affect the reported amounts of assets, liabilities, and expenses and the disclosure of contingent assets and liabilities in our financial statements. On an ongoing basis, we evaluate our estimates and judgments, including those related to revenue recognition, accrued expenses and stock-based compensation. We also utilize significant estimates and assumptions in determining the fair value of our common stock and the fair value of our liability-classified warrants to purchase preferred stock and common stock. We base our estimates on historical experience, known trends and events, and various other factors that are believed to be reasonable under the circumstances, the results of which form the basis for making judgments about the carrying values of assets and liabilities that are not readily apparent from other sources. Actual results may differ from these estimates under different assumptions or conditions.

While our significant accounting policies are described in more detail in the notes to our financial statements appearing elsewhere in this prospectus, we believe the following accounting policies to be most critical to the judgments and estimates used in the preparation of our financial statements.

**Revenue Recognition**

We have primarily generated revenue through collaboration arrangements with strategic partners for the development and commercialization of our protein therapeutics.

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We recognize revenue in accordance with Accounting Standards Codification (ASC) Topic 605, *Revenue Recognition*. Accordingly, revenue is recognized for each unit of accounting when all of the following criteria are met: (1) persuasive evidence of an arrangement exists; (2) delivery has occurred or services have been rendered; (3) the fee is fixed or determinable; and (4) collectability is reasonably assured.

Amounts received prior to satisfying the revenue recognition criteria are recorded as deferred revenue on our balance sheets. Amounts expected to be recognized as revenue within the 12 months following the balance sheet date are classified as deferred revenue, current portion and amounts not expected to be recognized as revenue within the 12 months following the balance sheet date are classified as deferred revenue, net of current portion.

Under collaboration agreements, we may receive payments for non-refundable up-front fees, milestone payments upon achieving significant development events, research and development reimbursements and royalties on future product sales. These payments are received in connection with the deliverables contained in the arrangements which may include (1) licenses, or options to obtain licenses, to our technology, (2) research and development activities performed for the collaboration partner, (3) participation on joint committees and (4) manufacturing clinical or preclinical material.

Effective January 1, 2011, we adopted Accounting Standards Update (ASU) No. 2009-13, *Multiple-Deliverable Revenue Arrangements*, which amends ASC Topic 605-25, *Revenue Recognition - Multiple Element Arrangements*. This guidance applies to new arrangements as well as existing agreements that are significantly modified after January 1, 2011.

The application of the multiple element guidance requires subjective determinations, and requires management to make judgments about the individual deliverables, and whether such deliverables are separable from the other aspects of the contractual relationship. Deliverables are considered separate units of accounting provided that: (1) the delivered item(s) has value to the customer on a stand-alone basis and (2) if the arrangement includes a general right of return relative to the delivered item(s), delivery or performance of the undelivered item(s) is considered probable and substantially in our control. In determining the units of accounting, management evaluates certain criteria, including whether the deliverables have stand-alone value, based on the consideration of the relevant facts and circumstances for each arrangement, such as the research, manufacturing and commercialization capabilities of the collaboration partner and the availability of the associated expertise in the general marketplace. In addition, we consider whether the collaboration partner can use the other deliverable(s) for their intended purpose without the receipt of the remaining element(s), whether the value of the deliverable is dependent on the undelivered item(s), and whether there are other vendors that can provide the undelivered element(s).

Arrangement consideration that is fixed or determinable is allocated among the separate units of accounting using the relative selling price method, and the applicable revenue recognition criteria, as described above, are applied to each of the separate units of accounting in determining the appropriate period or pattern of recognition. We determine the estimated selling price for deliverables within each agreement using vendor-specific objective evidence (VSOE) of selling price, if available, third-party evidence (TPE) of selling price if VSOE is not available, or management's best estimate of selling price (BESP) if neither VSOE nor TPE is available. Subsequent to the adoption of ASU 2009-13, we typically use BESP to estimate the selling price of the deliverables. Determining the BESP for a unit of accounting requires significant judgment. In developing the BESP for a unit of accounting, we consider applicable market conditions and relevant entity-specific factors, including factors that were contemplated in negotiating the agreement with the customer and estimated costs. We validate the BESP for units of accounting by evaluating whether changes in the key assumptions used to determine the BESP will have a significant effect on the allocation of arrangement consideration between multiple units of accounting.

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Our agreements may contain options which provide the collaboration partner the right to obtain additional licenses. Options are considered substantive if, at the inception of the arrangement, we are at risk as to whether the collaboration partner will choose to exercise the option. Factors that we consider in evaluating whether an option is substantive include the overall objective of the arrangement, the benefit the collaborator might obtain from the arrangement without exercising the option, the cost to exercise the option and the likelihood that the option will be exercised. For arrangements under which an option is considered substantive, we do not consider the item underlying the option to be a deliverable at the inception of the arrangement and the associated option fees are not included in allocable arrangement consideration, assuming the option is not priced at a significant and incremental discount. Conversely, for arrangements under which an option is not considered substantive or if an option is priced at a significant and incremental discount, we would consider the item underlying the option to be a deliverable at the inception of the arrangement and a corresponding amount would be included in allocable arrangement consideration.

We typically receive up-front, non-refundable payments when licensing our intellectual property in conjunction with a research and development agreement. When we believe the license to our intellectual property has stand-alone value, we generally recognize revenue attributed to the license upon delivery. When we believe the license to our intellectual property does not have stand-alone value from the other deliverables to be provided in the arrangement, we generally recognize revenue attributed to the license on a straight-line basis over our contractual or estimated performance period, which is typically the term of our research and development or manufacturing obligations. We continually evaluate these periods, and will adjust the period of revenue recognition if circumstances change.

Research and development funding is recognized as revenue in the period that the related services are performed. When we act as the principal under our collaboration arrangements, we record payments received for the reimbursement of research and development costs as cost-sharing revenue. To the extent that we reimburse the collaborator for costs incurred, we record these costs as a reduction of cost-sharing revenue.

We periodically review the basis for our estimates, and we may change the estimates if circumstances change. These changes can significantly increase or decrease the amount of revenue recognized. As we apply our policy to our collaboration arrangements we make judgments which affected the pattern of revenue recognition. For instance, in our arrangement with Celgene, we are obligated to provide research and development services. We are recognizing revenue over the estimated period of our performance of the research and development services, which was estimated to end in December 2014, the expected completion date of the proof-of-concept trials for ACE-536 under the Celgene collaboration. Another instance relates to our arrangement with Shire AG, where in April 2013, we and Shire determined not to further advance the development of ACE-031 or back-up compounds and Shire terminated our collaboration agreement effective as of June 30, 2013.

In addition to up-front payments and research and development funding, we may also be entitled to milestone payments that are contingent upon achievement of a predefined objective. At the inception of each arrangement that includes milestone payments, we evaluate whether the milestone is substantive and at-risk. This evaluation includes an assessment of whether (1) the consideration is commensurate with either the entity's performance to achieve the milestone, or the enhancement of the value of the delivered item(s) as a result of a specific outcome resulting at least in part from the entity's performance to achieve the milestone, (2) the consideration relates solely to past performance, and (3) the consideration is reasonable relative to all of the deliverables and payment terms within the arrangement. We evaluate factors such as the scientific, regulatory, commercial and other risks that must be overcome to achieve the respective milestone, the level of effort and investment required to achieve the respective milestone, and whether the milestone consideration is reasonable relative to all deliverables and payment terms in the arrangement in making this assessment. On the milestone

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achievement date, assuming all other revenue recognition criteria are met and the milestone is deemed substantive and at-risk, we recognize the payment as license and milestone revenue. For milestones that are not deemed substantive and at-risk, where payment is reasonably assured, we recognize the milestone payment over the remaining service period.

Sales and commercial milestones and royalties will be recognized when and if earned, provided collectability is reasonably assured.

**Clinical Trial Accruals and Related Expenses**

We accrue and expense costs for clinical trial activities performed by third parties, including CROs and clinical investigators, based upon estimates made as of the reporting date of the work completed over the life of the individual study in accordance with agreements established with CROs and clinical trial sites. Some CROs invoice us on a monthly basis, while others invoice upon achievement of milestones and the expense is recorded as services are rendered. We determine the estimates of clinical activities incurred at the end of each reporting period through discussion with internal personnel and outside service providers as to the progress or stage of completion of trials or services, as of the end of each reporting period, pursuant to contracts with numerous clinical trial centers and CROs and the agreed upon fee to be paid for such services. The significant factors considered in estimating accruals include the number of patients enrolled and the percentage of work completed to date. Costs of setting up clinical trial sites for participation in the trials that are paid for in advance are expensed over the estimated set-up period. While the set-up periods vary from one arrangement to another, such set-up periods generally take approximately three months. Set-up activities include clinical site identification, institutional review board, or IRB, submissions, regulatory submissions, clinical investigator kick-off meetings and pre-study site visits. Clinical trial site costs related to patient enrollments are accrued as patients are entered into the trial.

**Stock-Based Compensation**

We account for our stock-based awards in accordance with ASC Topic 718, *Compensation - Stock Compensation*, or ASC 718, which requires all stock-based payments to employees, including grants of employee stock options and modifications to existing stock options, to be recognized in the statements of operations and comprehensive income (loss) based on their fair values. We recognize the compensation cost of awards subject to service-based vesting conditions over the requisite service period, which is generally equal to the vesting term. For awards subject to both performance and service-based vesting conditions, we recognize compensation cost using an accelerated recognition method when it is probable that the performance condition will be achieved. We account for stock-based awards to non-employees using the fair value method. Stock options granted to non-employees are subject to periodic revaluation over their vesting terms and stock-based compensation cost is recognized using an accelerated recognition method.

We estimate the fair value of our stock-based awards to employees and non-employees using the Black-Scholes option pricing model, which requires the input of highly subjective assumptions, including (1) the expected volatility of our stock, (2) the expected term of the award, (3) the risk-free interest rate and (4) expected dividends. Due to the lack of a public market for our common stock prior to the completion of our initial public offering in September 2013, and resulting lack of company-specific historical and implied volatility data, we have based our estimate of expected volatility on the historical volatility of a group of similar companies that are publicly traded. For these analyses, we have selected companies with characteristics that we believe are comparable to ours, including enterprise value, risk profiles, position within the industry, and with historical share price information sufficient to meet the expected life of the stock-based awards. We compute the historical volatility data using the daily closing prices for the selected companies' shares during the equivalent period as the calculated expected term of our stock-based awards. We will continue to apply this process until a sufficient amount of historical

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information regarding the volatility of our own stock price becomes available. We have estimated the expected life of our employee stock options using the "simplified" method, whereby, the expected life equals the average of the vesting term and the original contractual term of the option. The risk-free interest rates for periods within the expected life of the option are based on the U.S. Treasury yield curve in effect during the period the options were granted.

We also estimate forfeitures at the time of grant, and revise those estimates in subsequent periods if actual forfeitures differ from estimates. We use historical data to estimate pre-vesting option forfeitures to the extent that actual forfeitures differ from our estimates, the difference is recorded as a cumulative adjustment in the period the estimates were revised. Stock-based compensation expense recognized in the financial statements is based on awards that are ultimately expected to vest.

We have computed the estimated fair value of stock options at the date of grant using the following weighted-average assumptions:

	Year Ended		Nine Months Ended	
	December 31,	December 31,	September 30,	September 30,
	2011	2012	2012	2013
Expected volatility	66.0%	69.0%	66.9%	70.3%
Expected term (in years)	6.0	6.0	6.0	6.0
Risk-free interest rate	1.1%	0.9%	0.9%	1.4%
Expected dividend yield				

Stock-based compensation totaled approximately \$1.2 million for the year ended December 31, 2012 and \$1.4 million for the nine months ended September 30, 2013. As of September 30, 2013, we had \$3.3 million of unrecognized compensation expense, net of related forfeiture estimates, which is expected to be recognized over a weighted-average remaining vesting period of approximately 2.2 years. We expect the impact of our stock-based compensation expense for stock-based awards granted to employees and non-employees to grow in future periods due to the potential increases in the value of our common stock and headcount.

The following table summarizes by grant date the number of shares of common stock underlying stock options granted from January 1, 2012 through the date we became a public company, as well as the associated per share exercise price and the retrospective estimated fair value per share of our common stock on the date of grant:

Date of Grant	Number of Shares Subject to Awards	Exercise Price Per Share(1)	Retrospective Fair Value Per Share on Date of Grant(2)
March 1, 2012	22,750	\$ 5.28	\$ 5.80
June 7, 2012	238,500	\$ 5.28	\$ 6.12
September 6, 2012	20,250	\$ 5.28	\$ 6.12
November 13, 2012	250,000	\$ 5.28	\$ 7.88
December 12, 2012	190,500	\$ 7.12	\$ 7.88
June 6, 2013	8,750	\$ 9.64	n/a

(1)

Due to the absence of a public market for our common stock prior to September 2013, the exercise price per share was the estimated fair value of common stock and represents the determination by our board of directors of the fair value of our common stock as of the date of each grant, taking into consideration various objective and subjective factors, as discussed more fully below.

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- (2) The fair value of common stock at the grant date was adjusted in connection with a retrospective fair value assessment for financial reporting purposes, as discussed more fully below.

***Determination of the Fair Value of Common Stock on Grant Dates***

For grants made prior to the consummation of our initial public offering in September 2013, our audit committee recommended, and our board of directors determined, the fair value of our common stock considering, in part, the work of an independent third party valuation specialist. Due to the absence of a public market for our common stock, the board determined the estimated per share fair value of our common stock at various dates considering contemporaneous valuations performed in accordance with the guidance outlined in the American Institute of Certified Public Accountants Practice Aid, *Valuation of Privately-Held Company Equity Securities Issued as Compensation*, also known as the Practice Aid. We engaged the valuation firm to perform contemporaneous valuations as of December 21, 2011, December 12, 2012, March 31, 2013 and June 6, 2013. In conducting the contemporaneous valuations, the valuation firm considered all objective and subjective factors that we believed to be relevant for each valuation conducted, including our best estimate of our business condition, prospects and operating performance at each valuation date. Within the contemporaneous valuations performed, a range of factors, assumptions and methodologies were used. The significant factors included:

the prices of our preferred stock sold to or exchanged between outside investors in arm's length transactions, and the rights, preferences and privileges of our preferred stock as compared to those of our common stock, including the liquidation preferences of our preferred stock;

our results of operations, financial position and the status of research and development efforts;

the composition of, and changes to, our management team and board of directors;

the lack of liquidity of our common stock;

our stage of development and business strategy and the material risks related to our business and industry;

the achievement of enterprise milestones, including entering into collaboration and license agreements;

the valuation of publicly traded companies in the life sciences and biotechnology sectors, as well as recently completed mergers and acquisitions of peer companies;

any external market conditions affecting the life sciences and biotechnology industry sectors;

the likelihood of achieving a liquidity event for the holders of our common stock and stock options, such as an initial public offering, or IPO, or a sale of our company, given prevailing market conditions; and

the state of the IPO market for similarly situated privately held biotechnology companies.

The dates of our contemporaneous valuations have not always coincided with the dates of our stock option grants. In determining the exercise prices of the stock options set forth in the table above, our board of directors considered, among other things, the most recent contemporaneous valuations of our common stock and our assessment of additional objective and subjective factors we believed were relevant as of the grant date. The additional factors considered when determining any changes in fair value between the most recent contemporaneous valuation and the grant dates included our stage of research and preclinical development, our operating and financial performance and current business conditions.



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There are significant judgments and estimates inherent in the determination of the fair value of our common stock. These judgments and estimates include assumptions regarding our future operating performance, the time to completing an IPO or other liquidity event, the related company valuations associated with such events, and the determinations of the appropriate valuation methods at each valuation date. If we had made different assumptions, our stock-based compensation expense, net income (loss) and net income (loss) per share applicable to common stockholders could have been different.

In early May 2013, based on the progress of our clinical pipeline, overall capital market conditions and the improving market for biopharmaceutical IPOs, our board of directors determined and directed management to begin preparation and submission of a confidential draft registration statement for an IPO. We selected underwriters and held an organizational meeting in June 2013. We believe these events increased the probability of an early IPO scenario and therefore in connection with the preparation of our financial statements for the year ended December 31, 2012, we retrospectively re-assessed the estimated fair value of our common stock for financial reporting purposes at interim dates between the contemporaneous valuations where there were stock option grants. For these interim periods, we adjusted the fair value based on market conditions, progress made in our development programs and whether we achieved company milestones.

***Common Stock Valuation Methodologies***

These contemporaneous and retrospective valuations were prepared in accordance with the guidelines in the Practice Aid, which prescribes several valuation approaches for setting the value of an enterprise, such as the cost, market and income approaches, and various methodologies for allocating the value of an enterprise to its common stock. We generally used the market approach, in particular the guideline company and precedent transaction methodologies, based on inputs from comparable public companies' equity valuations and comparable acquisition transactions, to estimate the enterprise value of our company.

***Methods Used to Allocate Our Enterprise Value to Classes of Securities***

In accordance with the Practice Aid, we considered the various methods for allocating the enterprise value across our classes and series of capital stock to determine the fair value of our common stock at each valuation date. The methods we considered consisted of the following:

*Current Value Method.* Under the current value method, once the fair value of the enterprise is established, the value is allocated to the various series of preferred and common stock based on their respective seniority, liquidation preferences or conversion values, whichever is greatest.

*Option Pricing Method.* Under the option pricing method, shares are valued by creating a series of call options with exercise prices based on the liquidation preferences and conversion terms of each equity class. The values of the preferred and common stock are inferred by analyzing these options.

*Probability-Weighted Expected Return Method, or PWERM.* The PWERM is a scenario-based analysis that estimates the value per share based on the probability-weighted present value of expected future investment returns, considering each of the possible outcomes available to us, as well as the economic and control rights of each share class.

We used the PWERM to allocate the enterprise values to the common stock for each valuation date. Under this method, the value of the common stock is estimated based upon an analysis of future values for our company assuming various investment outcomes, the timing of which is based, in part, on the plans of our board of directors and management. Under this approach, share value is derived from the probability-weighted present value of expected future investment returns, considering each of the



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possible outcomes available to us, as well as the economic and control rights of each share class. The fair value of our common stock was estimated using a probability-weighted analysis of the present value of the returns afforded to common stockholders under several future stockholder exit or liquidity event scenarios, either through (1) an IPO; (2) an acquisition or sale of our company at a premium to the cumulative liquidation preference of the preferred stockholders; or (3) a sale of our company at a value below the cumulative liquidation preference of the preferred stockholders.

The individual stockholder exit or liquidity scenarios considered in each analysis depended on the specific facts and circumstances, internal and external, present as of each valuation date. The future projected enterprise value used to value our common stock in the IPO scenarios and the sale scenarios were estimated by application of the market approach based on certain key assumptions, including the following:

valuations of companies prior to the receipt of proceeds from initial public offerings completed within three years of the valuation date;

estimated third-party sale values based on recent transactions involving biotechnology or biopharmaceutical companies; and

expected dates for a future IPO or sale of our company.

The present values of our common stock under each scenario were then calculated by applying a risk-adjusted discount rate and then probability-weighting those present values based on our estimate of the relative probability of each scenario.

Finally, the estimated fair value of our common stock was reduced by a discount for lack of marketability. A discount is appropriate because our common stock is unregistered, and the holder of a minority interest in the common stock may not influence the timing of a liquidity event for our company. Our estimate of the appropriate discount for lack of marketability took into consideration put option methodologies consistent with the Practice Aid. We selected a smaller discount after taking into account empirical studies of restricted stock issued by publicly-traded companies.

*March 1, 2012 Common Stock Valuation*

We performed a retrospective valuation of our common stock as of March 1, 2012, and determined the fair value to be \$5.80 per share as of that date. For the retrospective valuation at March 1, 2012, significant assumptions for the PWERM included the probability of occurrence of each scenario, timing to the liquidity event, discount rate and discount for lack of marketability. The specific facts and circumstances considered in assessing these key valuation assumptions included those noted in the following table:

<b>March 1, 2012 Major Assumptions</b>	<b>IPO Short Term</b>	<b>IPO Long Term</b>	<b>Sale High</b>	<b>Sale Low</b>	<b>Sales Below Liquidation Preference</b>
Probability of scenario	20%	20%	25%	25%	10%
Discount for lack of marketability			5%	5%	n/a
Timeline to liquidity (in years)	1.8	2.3	2.5	2.5	3.0
Discount rate common stock	30%	30%	30%	30%	n/a

In applying the market approach to estimate our future enterprise values under the IPO exit scenarios, as described previously, it was assumed that a liquidity event would occur in 1.8 years under a short term scenario or 2.3 years in the IPO long term scenario. We considered our development pipeline and our collaborations as of the valuation date. The selected enterprise value in the short-term scenario was based on the pre-money IPO market data for transactions between the third quartile and the maximum of the observed range. The selected aggregate enterprise value in the long-term scenario

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was based on consideration of the high-end of the observed range of transaction values and assumed our most advanced development projects would continue their positive clinical progression.

In applying the market approach to estimate our aggregate future enterprise values under the two sale scenarios, as described previously, it was assumed that a liquidity event would occur in 2.5 years for the high-case scenario and the low-case scenario. The selected enterprise value utilized in the low-case scenario considered the median of the observed range of comparable transaction values. The selected enterprise value for the high-case scenario was based on the comparable transaction values between the third quartile and the high-end of the observed range. We assumed we would make significant progress and achieve certain key milestones with respect to our development pipeline by the time a sale was consummated, including assumptions that our three most advanced development projects would continue their positive clinical progression, one or more additional compounds would enter Phase 1 trials and several other compounds would be nominated for pre-Investigational New Drug activities.

In the sale at a price below liquidation preference scenario, a sale of our existing research and intellectual property was assumed in 3.0 years, at a value that would not allow the preferred stockholders to realize their full liquidation preference resulting in no value to common stockholders.

Under all the exit scenarios considered in the PWERM, the fair value of our common stock was calculated using the estimated future enterprise valuations, a risk-adjusted discount rate of 30.0% based on the inherent risk of a hypothetical investment in our common stock, and a discount for lack of marketability which was 0% in the IPO scenarios and 5% in all other assumed liquidity events. The risk-adjusted discount rate was based on consideration of the weighted-average cost of capital for comparable biotechnology companies adjusted for company specific risk factors, the venture capital rates of return detailed in the Practice Aid, and an analysis of other quantitative and qualitative factors considered pertinent to estimating the discount rate. The resulting value, which represented the estimated fair value of our common stock as of March 1, 2012, was \$5.80 per share.

*June 7, 2012 and September 6, 2012 Common Stock Valuation*

We performed a retrospective valuation of our common stock as of June 7, 2012, and determined the fair value to be \$6.12 per share as of that date. For the retrospective valuation at June 7, 2012, significant assumptions for the PWERM included the probability of occurrence of each scenario, timing to the liquidity event, discount rate and discount for lack of marketability. The specific facts and circumstances in assessing these key valuation assumptions included those noted in the following table:

<b>June 7, 2012 Major Assumptions</b>	<b>IPO</b>		<b>Sale High</b>	<b>Sale Low</b>	<b>Sales Below Liquidation Preference</b>
	<b>Short Term</b>	<b>Long Term</b>			
Probability of scenario	25%	25%	25%	20%	5%
Discount for lack of marketability			5%	5%	n/a
Timeline to liquidity (in years)	1.5	2.0	2.5	2.5	3.0
Discount rate common stock	30%	30%	30%	30%	n/a

In applying the market approach to estimate our future enterprise values under the IPO exit scenarios, as described previously, it was assumed that a liquidity event would occur in 1.5 years under a short term scenario or 2.0 years in the IPO long term scenario. We considered our development pipeline and our collaborations as of the valuation date. The selected enterprise value in the short-term scenario was based on the pre-money IPO market data for transactions between the third quartile and the maximum of the observed range. The selected aggregate enterprise value in the long-term scenario was based on consideration of the high-end of the observed range of transaction values and assumed our most advanced development projects would continue their positive clinical progression.

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In applying the market approach to estimate our aggregate future enterprise values under the two sale scenarios, as described previously, it was assumed that a liquidity event would occur in 2.5 years for the high-case scenario and the low-case scenario. The selected enterprise value utilized in the low-case scenario considered the median of the observed range of comparable transaction values. The selected enterprise value for the high-case scenario was based on the comparable transaction values between the third quartile and the high-end of the observed range. We assumed we would make significant progress and achieve certain key milestones with respect to our development pipeline by the time a sale was consummated, including assumptions that our three most advanced development projects would continue their positive clinical progression, one or more additional compounds would enter Phase 1 trials and several other compounds would be nominated for pre-IND activities.

In the sale at a price below liquidation preference scenario, a sale of our existing research and intellectual property was assumed in 3.0 years, at a value that would not allow the preferred stockholders to realize their full liquidation preference resulting in no value to common stockholders.

Under all the exit scenarios considered in the PWERM, the fair value of our common stock was calculated using the estimated future enterprise valuations, a risk-adjusted discount rate of 30% based on the inherent risk of a hypothetical investment in our common stock, and a discount for lack of marketability which was decreased to 0% in the long-term IPO scenarios and remained 5% in all other assumed liquidity events. The risk-adjusted discount rate was based on consideration of the weighted-average cost of capital for comparable biotechnology companies adjusted for company specific risk factors, the venture capital rates of return detailed in the Practice Aid, and an analysis of other quantitative and qualitative factors considered pertinent to estimating the discount rate. The resulting value, which represented the estimated fair value of our common stock as of June 7, 2012, was \$6.12 per share.

The estimated per share fair value of our common stock calculated in our valuation as of June 7, 2012 of \$6.12 per share increased from the March 1, 2012 valuation of \$5.80 per share primarily due to the following factors:

timing to a prospective liquidity event had decreased; and

likelihood of an IPO had increased.

As a result of the fact that the number of stock option grants were not significant on September 6, 2012, we utilized the June 7, 2012 valuation to determine the retrospective fair value of our common stock in September 2012.

*November 13, 2012 and December 12, 2012 Common Stock Valuation*

We performed a retrospective valuation of our common stock as of November 13, 2012, and determined the fair value to be \$7.88 per share as of that date. For the retrospective valuation at November 13, 2012, significant assumptions for the PWERM included the probability of occurrence of each scenario, timing to the liquidity event, discount rate and discount for lack of marketability. The specific facts and circumstances considered in assessing these key valuation assumptions included those noted in the following table:

<b>November 13, 2012 Major Assumptions</b>	<b>IPO Short Term</b>	<b>IPO Long Term</b>	<b>Sale High</b>	<b>Sale Low</b>	<b>Sales Below Liquidation Preference</b>
Probability of scenario	30%	25%	25%	15%	5%
Discount for lack of marketability			5%	5%	n/a
Timeline to liquidity (in years)	1.0	1.5	2.0	2.0	2.5
Discount rate common stock	30%	30%	30%	30%	n/a

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In applying the market approach to estimate our future enterprise values under the IPO exit scenarios, as described previously, it was assumed that a liquidity event would occur in 1.0 years under a short term scenario or 1.5 years in the IPO long term scenario due to improvement in IPO market conditions for companies in our industry. We considered our development pipeline and our collaborations as of the valuation date. The selected enterprise value in the short-term scenario was based on the pre-money IPO market data for transactions between the third quartile and the maximum of the observed range. The selected aggregate enterprise value in the long-term scenario was based on consideration of the high-end of the observed range of transaction values and assumed our most advanced development projects would continue their positive clinical progression.

In applying the market approach to estimate our aggregate future enterprise values under the two sale scenarios, as described previously, it was assumed that a liquidity event would occur in 2.0 years for the high-case scenario and the low-case scenario. The selected enterprise value utilized in the low-case scenario considered the median of the observed range of comparable transaction values. The selected enterprise value for the high-case scenario was based on the comparable transaction values between the third quartile and the high-end of the observed range. We assumed we would make significant progress and achieve certain key milestones with respect to our development pipeline by the time a sale was consummated, including assumptions that our three most advanced development projects would continue their positive clinical progression, one or more additional compounds would enter Phase 1 trials and several other compounds would be nominated for pre-IND activities.

In the sale at a price below liquidation preference scenario, a sale of our existing research and intellectual property was assumed in 2.5 years, at a value that would not allow the preferred stockholders to realize their full liquidation preference resulting in no value to common stockholders.

Under all the exit scenarios considered in the PWERM, the fair value of our common stock was calculated using the estimated future enterprise valuations, a lower risk-adjusted discount rate of 25% based on a reduction in the inherent risk of a hypothetical investment in our common stock, and a discount for lack of marketability which remained 0% in the IPO scenarios and remained 5% in all other assumed liquidity events. The risk-adjusted discount rate was based on consideration of the weighted-average cost of capital for comparable biotechnology companies adjusted for company specific risk factors, the venture capital rates of return detailed in the Practice Aid, and an analysis of other quantitative and qualitative factors considered pertinent to estimating the discount rate. The resulting value, which represented the estimated fair value of our common stock as of November 13, 2012, was \$7.88 per share.

The estimated per share fair value of our common stock calculated in our valuation as of November 13, 2012 of \$7.88 per share increased from the June 7, 2012 retrospective valuation estimate of \$6.12 per share primarily due to the following factors:

timing to a prospective liquidity event has decreased since June 2012;

increased likelihood of an IPO; and

initiation of a Phase 2 clinical trial of dalantercept in endometrial cancer.

As a result of the fact that there were no material changes to our business from November 13, 2012 to December 12, 2012, we utilized the November 13, 2012 valuation to determine the exercise price of option grants in December.

*March 31, 2013 Common Stock Valuation*

We performed a contemporaneous valuation of our common stock as of March 31, 2013, and determined the fair value to be \$8.68 per share as of that date. For the valuation at March 31, 2013, significant assumptions for the PWERM included the probability of occurrence of each scenario, timing

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to the liquidity event, discount rate and discount for lack of marketability. The specific facts and circumstances considered in assessing these key valuation assumptions included those noted in the following table:

<b>March 31, 2013 Major Assumptions</b>	<b>IPO</b>		<b>Sale High</b>	<b>Sale Low</b>	<b>Sales Below Liquidation Preference</b>
	<b>Short Term</b>	<b>Long Term</b>			
Probability of scenario	50%	10%	20%	15%	5%
Discount for lack of marketability			5%	5%	n/a
Timeline to liquidity (in years)	0.6	1.0	1.8	2.0	2.3
Discount rate common stock	25%	25%	25%	25%	n/a

In applying the market approach to estimate our future enterprise values under the IPO exit scenarios, as described previously, it was assumed that a liquidity event would occur in 7 months under a short term scenario and 1.0 years in the IPO long term scenario. We considered our development pipeline and our collaborations as of the valuation date. The selected enterprise value in the short-term scenario was based on the pre-money IPO market data for transactions between the third quartile and the maximum of the observed range. The selected aggregate enterprise value in the long-term scenario was based on consideration of the high-end of the observed range of transaction values and assumed our most advanced development projects would continue their positive clinical progression.

In applying the market approach to estimate our aggregate future enterprise values under the two trade sale scenarios, as described previously, it was assumed that a liquidity event would occur in 1.8 years for the high-case scenario and 2.0 years the low-case scenario. The selected enterprise value utilized in the low-case scenario considered the median of the observed range of comparable transaction values. The selected enterprise value for the high-case scenario was based on the comparable transaction values between the third quartile and the high-end of the observed range. We assumed we would make significant progress and achieve certain key milestones with respect to our development pipeline by the time a trade sale was consummated, including assumptions that our three most advanced development projects would continue their positive clinical progression, one or more additional compounds would enter Phase 1 trials and several other compounds would be nominated for pre-IND activities.

In the sale at a price below liquidation preference scenario, a sale of our existing research and intellectual property was assumed in 2.3 years, at a value that would not allow the preferred stockholders to realize their full liquidation preference resulting in no value to common stockholders.

Under all the exit scenarios considered in the PWERM, the fair value of our common stock was calculated using the estimated future enterprise valuations, a lower risk-adjusted discount rate of 25% based on a reduction to the inherent risk of a hypothetical investment in our common stock, and a discount for lack of marketability which remained at 0% in the IPO scenarios and remained 5% in all other assumed liquidity events. The risk-adjusted discount rate was based on consideration of the weighted-average cost of capital for comparable biotechnology companies adjusted for company specific risk factors, the venture capital rates of return detailed in the Practice Aid, and an analysis of other quantitative and qualitative factors considered pertinent to estimating the discount rate. The resulting value, which represented the estimated fair value of our common stock as of March 31, 2013, was \$8.68 per share.

The estimated per share fair value of our common stock calculated in our valuation as of March 31, 2013 of \$8.68 per share increased from the November 13, 2012 valuation of \$7.88 per share primarily due to the following factors:

NASDAQ Biotechnology index increasing 20.8% from November 13, 2012 to March 31, 2013;

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improved capital market conditions for biotechnology companies as evidenced by a recent increase in the number of IPOs and their valuations;

increased likelihood of our board of directors recommending that we pursue an IPO;

decreased timing to a prospective liquidity event; and

initiation of several Phase 2 clinical trials for ACE-536 and dalantercept.

*June 6, 2013 Common Stock Valuation*

We performed a contemporaneous valuation of our common stock as of June 6, 2013, and determined the fair value to be \$9.64 per share as of that date.

For the contemporaneous valuation at June 6, 2013, significant assumptions for the PWERM included the probability of occurrence of each scenario, timing to the liquidity event, discount rate and discount for lack of marketability. The specific facts and circumstances considered in assessing these key valuation assumptions included those noted in the following table:

<b>June 6, 2013 Major Assumptions</b>	<b>IPO Short Term</b>	<b>IPO Long Term</b>	<b>Sale High</b>	<b>Sale Low</b>	<b>Sales Below Liquidation Preference</b>
Probability of scenario	60%	10%	15%	10%	5%
Discount for lack of marketability			5%	5%	n/a
Timeline to liquidity (in years)	0.4	0.8	1.6	1.8	2.1
Discount rate common stock	25%	25%	25%	25%	n/a

In applying the market approach to estimate our future enterprise values under the IPO exit scenarios, as described previously, it was assumed that a liquidity event would occur in 5 months under a short term scenario and 10 months in the IPO long term scenario. We considered our development pipeline and our collaborations as of the valuation date. The selected enterprise value in the short-term scenario was based on the pre-money IPO market data for transactions between the third quartile and the maximum of the observed range. The selected aggregate enterprise value in the long-term scenario was based on consideration of the high-end of the observed range of transaction values and assumed our most advanced development projects would continue their positive clinical progression.

In applying the market approach to estimate our aggregate future enterprise values under the two trade sale scenarios, as described previously, it was assumed that a liquidity event would occur in 1.6 years for the high-case scenario and 1.8 years the low-case scenario. The selected enterprise value utilized in the low-case scenario considered the median of the observed range of comparable transaction values. The selected enterprise value for the high-case scenario was based on the comparable transaction values between the third quartile and the high-end of the observed range. We assumed we would make significant progress and achieve certain key milestones with respect to our development pipeline by the time a trade sale was consummated, including assumptions that our three most advanced development projects would continue their positive clinical progression, one or more additional compounds would enter Phase 1 trials and several other compounds would be nominated for pre-IND activities.

In the sale at a price below liquidation preference scenario, a sale of our existing research and intellectual property was assumed in 2.1 years, at a value that would not allow the preferred stockholders to realize their full liquidation preference resulting in no value to common stockholders.

Under all the exit scenarios considered in the PWERM, the fair value of our common stock was calculated using the estimated future enterprise valuations, a lower risk-adjusted discount rate of 25% based on a reduction to the inherent risk of a hypothetical investment in our common stock, and a discount for lack of marketability which remained at 0% in the IPO scenarios and remained 5% in all

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other assumed liquidity events. The risk-adjusted discount rate was based on consideration of the weighted-average cost of capital for comparable biotechnology companies adjusted for company specific risk factors, the venture capital rates of return detailed in the Practice Aid, and an analysis of other quantitative and qualitative factors considered pertinent to estimating the discount rate. The resulting value, which represented the estimated fair value of our common stock as of June 6, 2013, was \$9.64 per share.

The estimated per share fair value of our common stock calculated in our valuation as of June 6, 2013 of \$9.64 per share increased from the March 31, 2013 valuation of \$8.68 per share primarily due to the following factors:

timing to a prospective liquidity event has decreased since March 2013;

NASDAQ Biotechnology (^NBI) index increasing 9.9% from April 1, 2013 to June 6, 2013;

improved capital market conditions for biotechnology companies as evidenced by a recent increase in the number of public offerings and their initial public offering valuations;

the occurrence of the organizational meeting for our potential IPO on June 5, 2013;

received two FDA Orphan Designations for ACE-536; and

initiated Phase 2 trial of ACE-536 in with  $\beta$ -thalassemia.

*Initial public offering price*

The initial public offering price of \$15.00 per share was determined as a result of negotiations between us and the underwriters. In comparison, our estimate of the fair value of our common stock was \$9.64 per share as of June 6, 2013. As is typical in initial public offerings, the initial public offering price was not derived using a formal determination of fair value, but was determined by negotiation between us and the underwriters. Among the factors that were considered in setting this price were the following:

an analysis of the typical valuation ranges seen in recent initial public offerings for companies in our industry;

the general condition of the securities markets and the recent market prices of, and the demand for, publicly traded common stock of generally comparable companies;

an assumption that there would be a receptive public trading market for pre-commercial biotechnology companies such as us; and

an assumption that there would be sufficient demand for our common stock to support an offering of the size contemplated in the initial public offering.

The initial public offering price reflected a significant increase over the estimated valuation as of June 6, 2013 of \$9.64 per share. We believe the difference is due to the following factors:

The contemporaneous valuation prepared as of June 6, 2013 contained multiple liquidity scenarios, including an initial public offering with an anticipated completion date of mid-September 2013 to which we assigned a probability weighting of 60%. However, the consideration of different scenarios accounts for some but not all of the difference between the initial public offering price and the valuation as of June 6, 2013;

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Advancement in the dose escalation phases of the on-going sotatercept and ACE-536 clinical trials in MDS and  $\beta$ -thalassemia;

Advancement in the treatment of patients in the on-going dalantercept clinical trials in renal cell carcinoma and squamous cell carcinoma of the head and neck;



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Improved capital market conditions for companies in our industry, as evidenced by a recent increase in the number of public offerings by such companies and in the initial public offering valuations of such companies compared to the valuations in their most recent pre-IPO equity financing;

The initial offering price necessarily assumed that the initial public offering had occurred, a public market for our common stock had been created and that our preferred stock had converted into common stock in connection with the initial public offering and, therefore, excluded the marketability or illiquidity discounts associated with the timing or likelihood of an initial public offering, the superior rights and preferences of our preferred stock and the alternative scenarios considered in the contemporaneous valuations over the past two years. Our June 6, 2013 valuation included an illiquidity discount of 0% in the IPO scenarios and 5% for the trade sale and liquidation scenarios;

In the public markets we believe there are investors who may apply more qualitative and subjective valuation criteria to certain of our clinical assets than the valuation methods applied in our valuations, although there can be no assurance that this will in fact be the case. As described above, as a private company we used a more quantitative methodology to determine the fair value of our common stock and this methodology differs from the methodology used to determine the initial public offering price. The initial public offering price was not derived using a formal determination of fair value, but rather was determined by negotiation between us and the underwriters. In particular, the estimate of fair value of our common stock as of June 6, 2013 was not a factor in setting the initial public offering price; and

The price that investors were willing to pay in the initial public offering may have taken into account other things that were not been expressly considered in our prior valuations, were not objectively determinable and that valuation models were not able to quantify.

There are significant additional judgments and estimates inherent in the determination of these valuations. These judgments and estimates include assumptions regarding our future performance, including the successful enrollment and completion of our clinical studies as well as the determination of the appropriate valuation methods. If we had made different assumptions, our stock-based compensation expense could have been different. The foregoing valuation methodologies are not the only methodologies available and they have not been used by us since the completion of our initial public offering. We cannot make assurances as to any particular valuation for our common stock. Accordingly, investors are cautioned not to place undue reliance on the foregoing valuation methodologies as an indicator of future stock prices.

**Warrants to Purchase Preferred Stock and Common Stock**

As of September 30, 2013, we had warrants outstanding to purchase 1,011,590 shares of common stock, of which warrants to purchase 857,586 shares of our common stock contain a provision requiring an adjustment to the number of shares in the event we issue common stock, or securities convertible into or exercisable for common stock, at a price per share lower than the warrant exercise price. The anti-dilution feature requires the warrants to be classified as liabilities and measured at fair value, with changes in fair value recognized as a component of other income (expense). The fair value of the warrants to purchase common stock on the date of issuance and on each re-measurement date for those warrants to purchase common stock are classified as liabilities and are estimated using the Monte Carlo simulation framework. The Company estimated that there would be up to three future financing events over the remaining life of the warrants to purchase common stock. Any modifications to the warrant liabilities are recorded in earnings during the period of the modification. The significant assumptions used in estimating the fair value of our warrant liabilities include the exercise price, volatility of the stock underlying the warrant, risk-free interest rate, estimated fair value of the stock underlying the warrant, and the estimated life of the warrant.

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Additionally, prior to the completion of our initial public offering in September 2013, we had warrants outstanding to purchase shares of Series B, Series C-1 and Series D-1 preferred stock. Freestanding warrants that are related to the purchase of redeemable preferred stock were classified as liabilities and recorded at fair value regardless of the timing of the redemption feature or the redemption price or the likelihood of redemption. The warrants were subject to re-measurement at each balance sheet date and any change in fair value was recognized as a component of other income (expense), net. We measured the fair value of our warrants to purchase preferred stock using a Black-Scholes option pricing model. In connection with the closing of our initial public offering on September 24, 2013, the outstanding warrants to purchase Series B Preferred Stock, Series C-1 Preferred Stock, and Series D-1 Preferred Stock were converted into warrants to purchase common stock and are now classified as a component of equity and are no longer subject to remeasurement. The exercise prices for each of these warrants remained unchanged.

**Emerging Growth Company Status**

The Jumpstart our Business Startups Act of 2012, or the JOBS Act, permits an "emerging growth company" such as us to take advantage of an extended transition period to comply with new or revised accounting standards applicable to public companies. We are choosing to "opt out" of this provision and, as a result, we will comply with new or revised accounting standards as required when they are adopted. This decision to opt out of the extended transition period under the JOBS Act is irrevocable.

**Results of Operations****Comparison of the Nine Months Ended September 30, 2012 and 2013**

(in thousands)	Nine Months Ended September 30,		Increase (Decrease)
	2012	2013	
Revenue:			
Collaboration revenue:			
License and milestone	\$ 7,226	\$ 36,044	\$ 28,818
Cost-sharing, net	4,043	9,666	5,623
Total revenue	11,269	45,710	34,441
Costs and expenses:			
Research and development	25,646	25,834	188
General and administrative	6,318	9,472	3,154
Total costs and expenses	31,964	35,306	3,342
Income (loss) from operations	(20,695)	10,404	31,099
Other income (expense), net	(1,508)	(14,192)	(12,684)
Net income (loss)	\$ (22,203)	\$ (3,788)	\$ 18,415

**Revenue.** We recognized revenue of \$45.7 million in the nine months ended September 30, 2013, compared to \$11.3 million in the same period in 2012. The \$34.4 million increase was primarily due to the \$10.0 million milestone payment received in connection with our Celgene collaboration for the first patient dosed in a Phase 2 trial in ACE-536 and recognizing an additional \$18.6 million of deferred revenue because Shire ended our collaboration as of June 30, 2013. The remaining increase of \$5.8 million was primarily due to an increase in net cost-sharing revenue from Celgene of \$6.9 million due to Celgene assuming 100% of the costs of development for these protein therapeutic candidates as of January 1, 2013, and recognition of \$0.2 million deferred revenue from Celgene, offset by a decrease

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in net cost-sharing revenue from Shire of \$1.3 million due to the end of the collaboration as of June 30, 2013.

The following table shows revenue from all sources for the periods presented.

(in thousands)	Nine Months Ended		Increase (Decrease)
	September 30, 2012	2013	
Collaboration revenue:			
Celgene:			
License and milestone	\$ 1,491	\$ 11,721	\$ 10,230
Cost-sharing, net	2,106	8,961	6,855
Total Celgene	3,597	20,682	17,085
Shire:			
License and milestone	5,735	24,323	18,588
Cost-sharing, net	1,937	705	(1,232)
Total Shire	7,672	25,028	17,356
Total collaboration revenue	11,269	45,710	34,441
Total revenue	\$ 11,269	\$ 45,710	\$ 34,441

**Research and Development Expenses.** Research and development expenses were \$25.8 million in the nine months ended September 30, 2013, compared to \$25.6 million in the same period in 2012. This \$0.2 million increase was primarily due to an increase in expenses associated with clinical activity totaling \$2.8 million, partially offset by a reduction in preclinical animal studies totaling \$2.5 million.

**General and Administrative Expenses.** General and administrative expenses were \$9.5 million in the nine months ended September 30, 2013, compared to \$6.3 million in the same period in 2012. This \$3.2 million increase was primarily related to higher professional fees for legal services in connection with our litigation with the Salk Institute and for increased professional fees and financial consulting services in connection with business development activities totaling \$2.3 million and higher total compensation expenses totaling \$0.9 million.

**Other Expense, Net.** Other expense, net was \$14.2 million in the nine months ended September 30, 2013, compared to \$1.5 million in the same period in 2012. This \$12.7 million increase was primarily due to higher expense associated with the increase in fair value of the liability for warrants of \$12.0 million and an increase in interest expense of \$0.7 million due to a higher average outstanding debt balance in the first half of 2013.

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(in thousands)	Year Ended		Increase (Decrease)
	2011	December 31, 2012	
Revenue:			
Collaboration revenue:			
License and milestone	\$ 74,406	\$ 9,696	\$ (64,710)
Cost-sharing, net	4,760	5,558	798
Contract manufacturing	1,745		(1,745)
Total revenue	80,911	15,254	(65,657)
Costs and operating expenses:			
Research and development	32,713	35,319	2,606
General and administrative	8,142	8,824	682
Cost of contract manufacturing revenue	1,500		(1,500)
Total costs and expenses	42,355	44,143	1,788
Income (loss) from operations	38,556	(28,889)	(67,445)
Other expense, net	(2,290)	(3,693)	(1,403)
Net income (loss)	\$ 36,266	\$ (32,582)	\$ (68,848)

**Revenue.** We recognized revenue of \$15.3 million for the year ended December 31, 2012, compared to \$80.9 million for the year ended December 31, 2011. The \$65.6 million decrease in revenue in 2012 was primarily due to a decrease of \$64.7 million in license and milestone revenue, because during 2011, upon signing the ACE-536 Celgene collaboration and amending the sotatercept agreement, we recognized upfront payments and deferred revenue totaling \$54.8 million. During 2011, we also recognized the remaining \$2.4 million of deferred revenue from the Alkermes collaboration. Also, in 2012 we did not recognize any milestone payments compared to \$7.0 million during 2011. The decrease in license and milestone revenue was offset by higher 2012 cost-sharing revenue due primarily to lower reimbursements paid to Shire for ACE-031. We also recognized \$1.7 million for a contract manufacturing project during 2011.

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The following table shows revenue from all sources for the periods presented.

(in thousands)	Year Ended December 31,		Increase (Decrease)
	2011	2012	
Collaboration revenue:			
Celgene:			
License and milestone	\$ 63,607	\$ 2,035	\$ (61,572)
Cost-sharing, net	(121)	2,879	3,000
Total Celgene	63,486	4,914	(58,572)
Shire:			
License and milestone	8,392	7,661	(731)
Cost-sharing, net	4,148	2,679	(1,469)
Total Shire	12,540	10,340	(2,200)
Alkermes:			
License and milestone	2,407		(2,407)
Cost-sharing, net	733		(733)
Total Alkermes	3,140		(3,140)
Total collaboration revenue	79,166	15,254	(63,912)
Contract manufacturing revenue	1,745		(1,745)
Total revenue	\$ 80,911	\$ 15,254	\$ (65,657)

**Research and Development Expenses.** Research and development expenses were \$35.3 million in the year ended December 31, 2012, compared to \$32.7 million for the year ended December 31, 2011. This \$2.6 million increase was primarily due to increases in expenses related to preclinical animal toxicology studies of \$2.6 million, patent-related legal services of \$0.9 million, external testing of \$0.6 million, clinical trial activities of \$0.5 million, contract labor of \$0.5 million, outsourced research of \$0.3 million and management bonuses of \$0.3 million partially offset by decreases in expenses related to depreciation of \$1.3 million, contract manufacturing of \$0.7 million, supplies of \$0.4 million, and in-licensing of \$0.5 million.

**General and Administrative Expenses.** General and administrative expenses were \$8.8 million in the year ended December 31, 2012, compared to \$8.1 million for the year ended December 31, 2011. This \$0.7 million increase was primarily related to higher professional fees for legal costs of \$0.4 million in connection with litigation activities and higher compensation costs of \$0.3 million.

**Cost of Contract Manufacturing Revenue.** There was no cost of contract manufacturing revenue for the year ended December 31, 2012, compared to \$1.5 million for the year ended December 31, 2011. This decrease was due to there being no contract manufacturing services provided during 2012.

**Other Expense, Net.** Other expense, net was \$3.7 million in the year ended December 31, 2012, compared to \$2.3 million for the year ended December 31, 2011. The increase was primarily due to a \$1.8 million increase in fair value of the liability for warrants to purchase redeemable convertible preferred stock and common stock.

### Liquidity and Capital Resources

We have incurred losses and cumulative negative cash flows from operations since our inception in June 2003, and as of September 30, 2013, we had an accumulated deficit of \$174.2 million. We anticipate that we will continue to incur losses for at least the next several years. We expect that our research and development and general and administrative expenses will continue to increase and, as a

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result, we will need additional capital to fund our operations, which we may raise through a combination of the sale of equity, debt financings or other sources, including potential additional collaborations.

As of September 30, 2013, our operations have been funded by \$105.1 million in equity investments from venture investors, \$49.2 million in equity investments from our partners, and \$192.6 million in upfront payments, milestones, and net research and development payments from our partners.

In September 2013, we completed the sale of 6,417,000 shares of our common stock, including 837,000 shares of common stock sold pursuant to the underwriters' full exercise of their option to purchase additional shares at a public offering price of \$15.00 per share, resulting in net proceeds to us of \$86.8 million, after deducting underwriting discounts and offering expenses. Also in September 2013, we completed a private placement of \$10 million of our common stock at a price of \$15.00 per share.

As of September 30, 2013, we had \$116.5 million in cash and cash equivalents. Cash in excess of immediate requirements is invested in accordance with our investment policy, primarily with a view to liquidity and capital preservation. Currently, our funds are held in money market mutual funds consisting of U.S. government-backed securities.

We entered into a new venture debt facility on June 7, 2012 and, as of September 30, 2013 we had \$18.2 million in venture debt outstanding. After an interest-only period, we began paying down principal on the debt facility in July 2013. Interest accrues at a rate of 8.5% per annum and is payable monthly. The debt facility also included a closing fee of \$0.2 million and is also subject to an additional deferred payment of \$1.2 million which is due at the time of the final payment. We are amortizing the cost over the 42 months of the loan resulting in an effective interest rate of approximately 11.8%. We are not subject to any financial covenants and the debt facility is secured by a lien on all of our property as of, or acquired after, June 7, 2012, except for intellectual property. The debt facility matures in December 2015.

**Cash Flows**

The following table sets forth the primary sources and uses of cash for each of the periods set forth below:

(in thousands)	Year Ended December 31,		Nine Months Ended September 30,	
	2011	2012	2012	2013
Net cash provided by (used in):				
Operating activities	\$ 9,056	\$ (38,884)	\$ (29,435)	\$ (18,286)
Investing activities	(27)	(441)	(322)	(187)
Financing activities	21,092	13,899	13,801	95,341
Net increase (decrease) in cash and cash equivalents	\$ 30,121	\$ (25,426)	\$ (15,956)	\$ 76,868

**Operating Activities.** The significant decrease in net cash used in operating activities for the nine months ended September 30, 2013, compared to the nine months ended September 30, 2012, is primarily due to the receipt of a \$10.0 million milestone payment from Celgene in the first quarter of 2013. The significant decrease in cash provided by operating activities for the year ended December 31, 2012, compared to the year ended December 31, 2011, is primarily due to the upfront and milestone payments of \$32.5 million related to the ACE-536 Agreement received during 2011.

Net cash used in operating activities was \$18.3 million for the nine months ended September 30, 2013, and consisted primarily of a net loss of \$3.8 million adjusted for non-cash items including an increase in fair value of warrants of \$12.6 million, stock-based compensation expense of \$1.4 million,

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depreciation and amortization of \$0.7 million, forgiveness of the related party receivable of \$0.2 million, accretion of deferred interest of \$0.3 million, and amortization of deferred debt issuance costs of \$0.2 million, and a net decrease due to changes in operating assets and liabilities of \$29.9 million. The significant items in the change in operating assets and liabilities include a decrease in deferred revenue of \$26.0 million due primarily to the recognition of \$24.3 million of deferred revenue for the Shire collaboration agreement which was terminated effective June 30, 2013. Other components of the change in operating assets and liabilities include a decrease in accrued expenses of \$1.6 million, an increase in collaboration receivables of \$1.3 million, an increase in prepaid expenses of \$0.8 million, a decrease in deferred rent of \$0.4 million and an increase in accounts payable of \$0.2 million.

Net cash used in operating activities was \$29.4 million for the nine months ended September 30, 2012 and consisted primarily of a net loss of \$22.2 million adjusted for non-cash items including an increase in fair value of warrants of \$0.6 million, stock-based compensation expense of \$0.9 million, depreciation and amortization of \$1.1 million, accretion of deferred interest of \$0.3 million, and amortization of deferred debt issuance costs of \$0.1 million, and a net decrease due to changes in operating assets and liabilities of \$10.1 million. The significant items in the change in operating assets and liabilities include a decrease in deferred revenue of \$7.2 million due to the ongoing recognition of revenue deferred in connection with up-front payments for the Celgene and Shire collaboration agreements, a decrease in accounts payable of \$0.9 million and an increase in prepaid expenses and other current assets of \$1.3 million. Other components of the change in operating assets and liabilities include an increase in collaboration receivables of \$1.0 million, an increase in accrued expenses of \$0.7 million and a decrease in deferred rent of \$0.4 million.

Net cash used in operating activities was \$38.9 million for the year ended December 31, 2012 and is primarily due to a net loss of \$32.6 million adjusted for non-cash items including an increase in the fair value of warrants of \$2.3 million, stock-based compensation of \$1.2 million, depreciation and amortization of \$1.3 million, and accretion of deferred interest of \$0.3 million and a net decrease in operating assets and liabilities of \$11.5 million. The significant items in the change in operating assets and liabilities include a decrease in deferred revenue of \$9.7 million due to the ongoing recognition of revenue deferred in connection with up-front payments for the Celgene and Shire collaboration agreements, a decrease in accounts payable of \$1.3 million and an increase in collaboration receivables of \$1.1 million, offset in part by an increase in accrued expenses of \$1.6 million. Other components of the change in operating assets and liabilities include an increase in prepaid expenses and other current assets of \$0.6 million and a decrease in deferred rent of \$0.5 million.

Net cash provided by operating activities was \$9.1 million for the year ended December 31, 2011 and is primarily due to net income of \$36.3 million, which was impacted by non-cash items including depreciation and amortization of \$3.1 million, stock-based compensation of \$1.4 million, an increase in the fair value of warrants of \$0.5 million, accretion of deferred interest of \$0.3 million and amortization of debt discount of \$0.2 million and a net decrease in operating assets and liabilities of \$32.8 million. The significant items in the change in operating assets and liabilities include a decrease in deferred revenue of \$35.1 million due primarily to the acceleration of deferred revenue associated with the Celgene collaboration upfront payments as a result of the modification of the collaboration agreement, as well as a decrease in accrued expenses of \$2.8 million, offset in part by a decrease in prepaid expenses and other current assets of \$2.6 million and a decrease in collaboration receivables of \$1.8 million. Other components of the change in operating assets and liabilities include an increase in accounts payable of \$0.3 million and an increase in deferred rent of \$0.2 million.

**Investing Activities.** Net cash used in investing activities was \$0.2 million for the nine months ended September 30, 2013 and \$0.3 million for the nine months ended September 30, 2012 and consisted of purchases of property and equipment.

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Net cash used in investing activities was \$27,000 for the year ended December 31, 2011 and \$0.4 million for the year ended December 31, 2012 and consisted of purchases of property and equipment.

**Financing Activities.** Net cash provided by financing activities was \$95.3 million for the nine months ended September 30, 2013 and consisted of \$97.4 million in net proceeds received from the company's initial public offering and concurrent private placement, offset by \$1.8 million of principal payments made to pay down our venture debt line and \$0.3 million paid to repurchase and retire redeemable convertible preferred stock, common stock and warrants to purchase common stock. Net cash provided by financing activities was \$13.8 million for the nine months ended September 30, 2012 and consisted primarily of \$19.9 million in net proceeds received from the drawdown of our new venture debt line in June 2012, offset by \$6.2 million of principal payments made to pay down our previous venture debt line.

Net cash provided by financing activities was \$21.1 million for the year ended December 31, 2011 and consisted primarily of \$30.4 million of net proceeds received from the sale of 9,704,756 shares of our Series F preferred stock, as well as \$0.2 million received from the exercise of stock options and warrants to purchase common stock, offset in part by \$9.5 million of principal payments made to pay down a previous venture debt facility.

Net cash provided by financing activities was \$13.9 million for the year ended December 31, 2012 and consisted of \$19.9 million in net proceeds received from the drawdown of our new venture debt line in June 2012, as well as \$0.2 million received from the exercise of stock options and warrants to purchase common stock, offset by \$6.2 million of principal payments made to pay down our previous venture debt line.

**Operating Capital Requirements**

To date, we have not generated any revenue from product sales. We do not know when, or if, we will generate any revenue from product sales. We will not generate revenue from product sales unless and until we or our partners obtain regulatory approval of and commercialize one of our current or future protein therapeutics. We anticipate that we will continue to generate losses for the foreseeable future, and we expect the losses to increase as we continue the development of, and seek and obtain regulatory approvals for, dalantercept and any future protein therapeutics, and begin to commercialize any approved products. We are subject to all of the risks incident in the development of protein therapeutics, and we may encounter unforeseen expenses, difficulties, complications, delays and other unknown factors that may adversely affect our business. Since the closing of our initial public offering, we have incurred, and expect to continue to incur, additional costs associated with operating as a public company. We anticipate that we will need additional funding in connection with our continuing operations.

We believe that the net proceeds we receive from this offering, together with receipt of anticipated milestone payments and our existing cash and cash equivalents will be sufficient to fund our projected operating requirements into the first quarter of 2017. However, we will require additional capital for the further development of our existing protein therapeutic candidates and may also need to raise additional funds sooner to pursue other development activities related to additional protein therapeutic candidates.

Until we can generate a sufficient amount of revenue from our products, if ever, we expect to fund our operations through a combination of equity offerings, or debt financings or other sources including potential additional collaborations. Additional capital may not be available on favorable terms, if at all. If we are unable to raise additional capital in sufficient amounts or on terms acceptable to us, we may have to significantly delay, scale back or discontinue the development or commercialization of one or more of our protein therapeutic candidates. If we raise additional funds through the issuance of



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additional debt or equity securities, it could result in dilution to our existing stockholders and increased fixed payment obligations, and these securities may have rights senior to those of our common stock. If we incur indebtedness, we could become subject to covenants that would restrict our operations and potentially impair our competitiveness, such as limitations on our ability to incur additional debt, limitations on our ability to acquire, sell or license intellectual property rights and other operating restrictions that could adversely impact our ability to conduct our business. We may not be able to enter into new collaboration arrangements for any of our proprietary protein therapeutic candidates. Any of these events could significantly harm our business, financial condition and prospects.

Our forecast of the period of time through which our financial resources will be adequate to support our operations is a forward-looking statement and involves risks and uncertainties, and actual results could vary as a result of a number of factors. We have based this estimate on assumptions that may prove to be wrong, and we could utilize our available capital resources sooner than we currently expect. Our future funding requirements, both near and long-term, will depend on many factors, including, but not limited to:

the achievement of milestones under our agreement with Celgene;

the terms and timing of any other collaborative, licensing and other arrangements that we may establish;

the initiation, progress, timing and completion of preclinical studies and clinical trials for our protein therapeutic candidates and potential protein therapeutic candidates;

the number and characteristics of protein therapeutic candidates that we pursue;

the progress, costs and results of our clinical trials;

the outcome, timing and cost of regulatory approvals;

delays that may be caused by changing regulatory requirements;

the cost and timing of hiring new employees to support our continued growth;

the costs involved in filing and prosecuting patent applications and enforcing and defending patent claims;

the costs and timing of procuring clinical and commercial supplies of our protein therapeutic candidates;

the extent to which we acquire or invest in businesses, products or technologies; and

the costs involved in defending and prosecuting litigation regarding in-licensed intellectual property including our litigation with the Salk Institute. See "Business Litigation".

Table of Contents**Contractual Obligations and Commitments**

The following is a summary of our long-term contractual cash obligations as of December 31, 2012.

(in thousands)	Total	Less than 1 Year	1 to 3 Years	3 to 5 Years	More than 5 Years
Operating lease obligations(1)	\$ 23,979	\$ 4,522	\$ 8,628	\$ 7,876	\$ 2,953
Less: sublease income(2)	(1,407)	(583)	(824)		
Venture debt facility(3)	24,320	5,304	19,016		
Total	\$ 46,892	\$ 9,243	\$ 26,820	\$ 7,876	\$ 2,953

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- (1) We lease office space at 128 Sidney Street and 149 Sidney Street in Cambridge, Massachusetts under noncancelable operating leases that expire in September 2018, and at 12 Emily Street in Cambridge, Massachusetts under a noncancelable operating lease that expires in May 2015.
- (2) In February 2011, we entered into a sublease for 14,214 square feet of office space at 12 Emily Street in Cambridge, Massachusetts.
- (3) In June 2012, we entered into a \$20.0 million venture debt facility to provide working capital to fund operating activities. The loans under this debt facility are secured by our assets and are being repaid over 42 months beginning with a 12 month interest only period. Interest rates were fixed at the time of drawdown, with an effective rate of 11.8%.

We also have obligations to make future payments to third party licensors that become due and payable on the achievement of certain development, regulatory and commercial milestones. We have not included these commitments on our balance sheet or in the table above because the achievement and timing of these milestones is not fixed or determinable. These commitments include the following:

Under our license agreement with the Beth Israel Deaconess Medical Center, or BIDMC, in respect of BIDMC's joint interest in patent rights related to the treatment of renal cell cancer by combination therapy with dalantercept and VEGF-receptor tyrosine kinase inhibitors, we agreed to pay BIDMC specified development and sales milestone payments aggregating up to \$1.0 million. In addition, we are required to pay BIDMC royalties in the low single digits on worldwide net product sales of drug labeled for treatment regimens that are claimed in the licensed patents.

Under our license agreement with the Ludwig Institute for Cancer Research, or LICR, in respect of patent rights relating to the first cloning of the type I activin receptors, as well as the treatment of pancreatic tumors with dalantercept, we agreed to pay LICR specified development and sales milestone payments aggregating up to \$1.6 million relating to the development and commercialization of dalantercept. In addition, we are required to pay LICR royalties in the low single-digits on worldwide net product sales of dalantercept, with royalty obligations continuing at a 50% reduced rate for a period of time after patent expiration. If we sublicense the LICR patent rights, we will owe LICR a percentage of sublicensing revenue, excluding payments based on the level of sales, profits or other levels of commercialization.

Under our two license agreements with the Salk Institute for Biological Studies, or Salk, relating to the first cloning of the type II activin receptors, if we sublicense the Salk patent rights, we will owe Salk a percentage of sublicensing revenue, excluding payments based on sales. Under one agreement we also agreed to pay Salk specified development milestone payments totaling up to \$2.0 million for sotatercept. Under the other agreement we also agreed to pay Salk specified development milestone payments of up to \$0.7 million for ACE-536. In addition, under both agreements, we are required to pay Salk royalties in the low single-digits on worldwide net product sales by us or our sublicensees under the licensed patent rights of products claimed in



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the licensed patents, or products derived from use of the licensed patent rights, with royalty obligations for sotatercept continuing at a reduced rate for a period of time after patent expiration.

We enter into contracts in the normal course of business with CROs for clinical trials and clinical supply manufacturing and with vendors for preclinical safety and research studies, research supplies and other services and products for operating purposes. These contracts generally provide for termination on notice, and therefore are cancelable contracts and not included in the table of contractual obligations and commitments.

**Net Operating Loss (NOL) Carryforwards**

We have deferred tax assets of approximately \$68.2 million as of December 31, 2012, which have been fully offset by a valuation allowance due to uncertainties surrounding our ability to realize these tax benefits. The deferred tax assets are primarily composed of federal and state tax net operating loss, or NOL, carryforwards and research and development tax credit carryforwards. As of December 31, 2012, we had federal NOL carryforwards of approximately \$93.3 million and state NOL carryforwards of \$75.4 million available to reduce future taxable income, if any. These federal NOL carryforwards expire at various times through 2032 and the state NOL carryforwards expire at various times through 2032. In general, if we experience a greater than 50 percent aggregate change in ownership of certain significant stockholders over a three-year period, or a Section 382 ownership change, utilization of our pre-change NOL carryforwards are subject to an annual limitation under Section 382 of the Internal Revenue Code of 1986, as amended, and similar state laws. Such limitations may result in expiration of a portion of the NOL carryforwards before utilization and may be substantial. If we experience a Section 382 ownership change in connection with this offering or as a result of future changes in our stock ownership, some of which changes are outside our control, the tax benefits related to the NOL carryforwards may be limited or lost.

**Off-Balance Sheet Arrangements**

We did not have during the periods presented, and we do not currently have, any off-balance sheet arrangements, as defined in the rules and regulations of the Securities and Exchange Commission.

**Quantitative and Qualitative Disclosures About Market Risks**

We are exposed to market risk related to changes in interest rates. As of September 30, 2013, we had cash and cash equivalents of \$116.5 million. Our cash equivalents are invested in money market mutual funds consisting of U.S. government-backed securities. Our primary exposure to market risk is interest rate sensitivity, which is affected by changes in the general level of U.S. interest rates, particularly because our investments are in short-term securities. Due to the short-term duration of our investment portfolio and the low risk profile of our investments, an immediate 100 basis point change in interest rates would not have a material effect on the fair market value of our portfolio.

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**BUSINESS**

**Overview**

We are a clinical stage biopharmaceutical company focused on the discovery, development and commercialization of novel protein therapeutics for cancer and rare diseases. Our research focuses on the biology of the Transforming Growth Factor-Beta (TGF- $\beta$ ) protein superfamily, a large and diverse group of molecules that are key regulators in the growth and repair of tissues throughout the human body. We are leaders in understanding the biology of the TGF- $\beta$  superfamily and in targeting these pathways to develop important new medicines. By coupling our discovery and development expertise, including our proprietary knowledge of the TGF- $\beta$  superfamily, with our internal protein engineering and manufacturing capabilities, we have built a highly productive discovery and development platform that has generated innovative protein therapeutic candidates with novel mechanisms of action. These differentiated protein therapeutic candidates have the potential to significantly improve clinical outcomes for patients with cancer and rare diseases.

We focus on discovering and developing protein therapeutics that target a group of approximately 30 secreted proteins, or ligands, that are collectively referred to as the TGF- $\beta$  superfamily. These ligands bind to subsets of 12 different receptors on the surface of cells, triggering intra-cellular changes in gene expression that guide cell growth and differentiation. The TGF- $\beta$  superfamily ligands and their receptors represent an under-explored and diverse set of drug targets with the potential to yield therapeutics that modulate the growth and repair of diseased cells and tissues.

We have three internally discovered protein therapeutic candidates that are currently being studied in numerous ongoing Phase 2 clinical trials, focused on cancer and rare diseases. Our two most advanced protein therapeutic candidates, sotatercept and ACE-536, promote red blood cell production through a novel mechanism. Together with our collaboration partner, Celgene Corporation, we are developing sotatercept and ACE-536 to treat anemia and associated complications in patients with  $\beta$ -thalassemia and myelodysplastic syndromes (MDS). These red blood cell disorders are generally unresponsive to currently approved drugs. Our third clinical stage protein therapeutic candidate, dalantercept, is designed to inhibit blood vessel formation through a mechanism that is distinct from, and potentially synergistic with, the dominant class of cancer drugs that inhibit blood vessel formation, the vascular endothelial growth factor (VEGF) pathway inhibitors. We are developing dalantercept primarily for use in combination with these products to produce better outcomes for cancer patients. We estimate that we have spent approximately \$142.1 million on research and development from 2010 through September 30, 2013.

Sotatercept and ACE-536 have already shown promising biological activity in our initial clinical trials. We and Celgene have conducted six human clinical trials with sotatercept in over 160 healthy volunteers and cancer patients. We have conducted one clinical trial with ACE-536 in healthy volunteers. In these studies, both sotatercept and ACE-536 caused a dose-dependent increase in the number of red blood cells. Based on these results, we and Celgene have initiated Phase 2 clinical trials with each of these protein therapeutic candidates in  $\beta$ -thalassemia and MDS. In the ongoing trials of sotatercept and ACE-536 in patients with  $\beta$ -thalassemia, we have observed encouraging, dose-dependent increases in hemoglobin in non-transfusion dependent patients at the three dose levels tested. We and Celgene plan to initiate Phase 3 clinical trials for one or both of these protein therapeutic candidates in one or both of  $\beta$ -thalassemia and MDS by the end of 2014 or early 2015.

With respect to our third clinical stage protein therapeutic candidate, dalantercept, we have conducted a single agent Phase 1 clinical trial in patients with advanced solid tumors. Of the 29 evaluable patients treated in this clinical trial, one had a partial response and 13 had stable disease, according to RECIST criteria. Additionally, we have studied the single agent activity of dalantercept in a Phase 2 clinical trial in patients with advanced head and neck cancer. Of the 29 evaluable patients at the 1.2 mg/kg dose level in this Phase 2 clinical trial, one had a partial response and ten had stable disease, according to RECIST criteria. Our ongoing focus is on the use of dalantercept in combination

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with an approved VEGF pathway inhibitor where we have provided both a mechanistic rationale and supportive preclinical data demonstrating dalantercept in combination with a VEGF pathway inhibitor provides enhanced anti-tumor effects in mice bearing human renal cell carcinoma xenographs. In an ongoing Phase 2 clinical trial of dalantercept in combination with axitinib, an approved VEGF pathway inhibitor, in patients with advanced renal cell carcinoma we have completed the dose escalation stage and found that dalantercept administered at a dose level of 1.2 mg/kg is well tolerated in combination with the FDA approved dose level of axitinib. We have now initiated the dose expansion phase of this study and plan to start the randomized controlled part of the study at the end of Q1 or early Q2 2014. We also intend to initiate a Phase 2 clinical trial of dalantercept in combination with the VEGF pathway inhibitor sorafenib in patients with liver cancer in the first half of 2014.

In addition to our clinical stage programs, we are developing a novel protein therapeutic candidate, ACE-083, for a first-in-human clinical trial that we expect to initiate by the end of 2014. ACE-083 has been designed to promote muscle growth in those muscles in which the drug is injected, with minimal systemic effect. We are focused on the development of ACE-083 for diseases in which increases in the size and function of specific muscles may provide a clinical benefit, including inclusion body myositis, facioscapulohumeral dystrophy (FSHD) and disuse atrophy.

We are developing sotatercept and ACE-536 through our exclusive worldwide collaborations with Celgene. As of January 1, 2013, Celgene became responsible for paying 100% of worldwide development costs for both programs. We may receive up to an additional \$560.0 million of potential development, regulatory and commercial milestone payments and, if these protein therapeutic candidates are commercialized, we will receive a royalty on net sales in the low-to-mid 20% range. We will co-promote sotatercept and ACE-536, if approved, in North America for which our commercialization costs will be entirely funded by Celgene.

We have not entered into a partnership for dalantercept and retain worldwide rights to this program.

As of September 30, 2013, our operations have been funded primarily by \$105.1 million in equity investments from venture investors, \$86.8 million from investors in our initial public offering, \$49.2 million in equity investments from our collaboration partners Celgene and Alkermes, Inc. (Alkermes) and \$192.6 million in upfront payments, milestones, and net research and development payments from our collaboration partners.

**Our Strategy**

Our goal is to be a leader in the discovery, development and commercialization of novel protein therapeutics for cancer and rare diseases. Key components of our strategy are:

***Advance sotatercept and ACE-536 into Phase 3 trials in collaboration with Celgene.*** We and Celgene are jointly developing sotatercept and ACE-536. Assuming successful completion of the ongoing Phase 2 clinical trials in  $\beta$ -thalassemia and MDS, we plan to initiate Phase 3 clinical trials with Celgene for one or both protein therapeutics in one or both diseases by the end of 2014 or early 2015.

***Explore new indications for sotatercept and ACE-536 with Celgene.*** We and Celgene are continuing our preclinical research to assess the opportunity for sotatercept and ACE-536 to treat certain red blood cell disorders known as hemoglobinopathies, which include diseases such as thalassemias and sickle cell disease. Based on our encouraging preclinical and clinical data in  $\beta$ -thalassemia and our emerging understanding of the mechanism of action of these protein therapeutic candidates, we believe there is a potential for activity for sotatercept and ACE-536 in sickle cell disease, and we continue to explore development of these protein therapeutic candidates for this disease.

***Advance dalantercept into Phase 3-enabling clinical trials.*** Beyond our ongoing Phase 2 clinical trials, in 2014, we plan to initiate additional clinical trials of dalantercept in combination with

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either an approved anti-angiogenesis agent or chemotherapy in advanced solid tumors. One of these trials is expected to be in patients with liver cancer and other trials may be in patients with brain cancer, lung cancer or colon cancer.

***Utilize our discovery and development platform to develop additional protein therapeutic candidates.*** In addition to sotatercept, ACE-536 and dalantercept, all of which were internally discovered using our research and development platform, we intend to continue to discover and develop other protein therapeutics that target and regulate various pathways in the TGF- $\beta$  superfamily. We plan to bring an additional protein therapeutic candidate, ACE-083, into the clinic in 2014 targeting diseases involving muscle loss. We are also conducting pre-clinical development of ALK1 pathway inhibitors distinct from dalantercept for the treatment of diseases of the eye including age-related macular degeneration. In addition we are developing new protein therapeutic candidates for the treatment of cancer and diseases involving fibrosis.

***Strategically leverage collaborations to advance our protein therapeutic candidates.*** To date, we have received more than \$250.0 million from our corporate partners, including Celgene. Our two collaborations with Celgene for sotatercept and ACE-536 provide us with significant funding and access to Celgene's considerable scientific, development, regulatory and commercial capabilities. We will continue to strategically evaluate possible collaborations where doing so could enhance the development or commercialization of other protein therapeutic candidates in our pipeline.

***Establish commercialization and marketing capabilities in North America and potentially other markets.*** We have retained co-promotion rights in North America for sotatercept and ACE-536, which will be entirely funded by Celgene. We intend to build hematology, oncology and neuromuscular disorder focused specialty sales forces and marketing capability to commercialize our protein therapeutic candidates that receive regulatory approval.

**The Acceleron Discovery Platform: Novel Approaches to Potent Biology**

Since our founding, we have focused on developing protein therapeutics that target a group of approximately 30 secreted proteins, or ligands, that are collectively referred to as the TGF- $\beta$  superfamily. These ligands bind to subsets of 12 different receptors on the surface of cells, triggering intra-cellular changes in gene expression that guide cell growth and differentiation. The TGF- $\beta$  superfamily ligands and their receptors represent a diverse and underexplored set of drug targets with the potential to yield potent therapeutics for the growth and repair of diseased cells and tissues. Applying our proprietary discovery and development platform, including our knowledge of the biology of the TGF- $\beta$  superfamily and its receptors, we have generated a robust pipeline of innovative clinical and preclinical protein therapeutic candidates targeting key mechanisms underlying cancer and rare diseases.

***Our Focus The TGF- $\beta$  Superfamily***

On a daily basis, the human body must orchestrate the growth and differentiation of cells to maintain and repair its cells and organ systems. Stem cells and precursor cells are undifferentiated cell types that reside in most tissues of the body. When tissue growth or regeneration is required, these undifferentiated cells divide and, through a series of intermediate stages, give rise to new, fully differentiated cells that build or repair the affected tissue. Decades of research have identified the TGF- $\beta$  superfamily and its associated receptors as key regulators of the growth and differentiation of stem and precursor cells.

Until recently, regulation of the erythropoietin pathway was the primary therapeutic approach to stimulate red blood cell formation. Members of the TGF- $\beta$  superfamily are now recognized as important regulators of red blood cell formation. We have shown that inhibition of members of the TGF- $\beta$  superfamily ameliorates anemia in mouse models of  $\beta$ -thalassemia and MDS. Based on our findings, we are developing two protein therapeutic candidates, sotatercept and ACE-536, each of which is currently in Phase 2 clinical trials to treat patients with these diseases.

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Members of the TGF- $\beta$  superfamily also play a significant role in regulating blood vessel formation. We and our academic collaborators have shown that mice with a genetic defect in a particular receptor for members of the TGF- $\beta$  superfamily are resistant to tumor growth due to reduced blood vessel formation in the tumor. We have used this insight to design our Phase 2 anti-angiogenic agent, dalantercept, for the treatment of cancer.

Members of the family are also significant regulators of muscle development. A genetic defect in a TGF- $\beta$  superfamily ligand, known as myostatin, causes profound increases in skeletal muscle. A naturally occurring mutation in myostatin has been identified in animals, such as "double-muscled" breeds of cattle and in the "bully whippet" offspring of whippet racing dogs, which have been selectively bred to have increased muscle mass or function. Furthermore, a mutation in myostatin has been identified in a human family, members of which exhibit exceptional musculature and strength. We are actively working on preclinical programs to increase muscle mass and strength.

Ligands of the TGF- $\beta$  superfamily cause these profound biological effects by altering gene expression in target cells. As shown in the illustration below, a ligand of the superfamily initiates intracellular signaling by binding to a receptor that is located on the surface of a target cell. Upon binding to the ligand, the receptor activates specific transcription factors inside the target cell, which are called Smad proteins. The activated Smad proteins regulate gene expression and guide cellular growth and differentiation.

The TGF- $\beta$  superfamily ligands are divided into subgroups termed the activins, the Growth and Differentiation Factors (GDFs), the Bone Morphogenetic Proteins (BMPs) and the TGF- $\beta$  subgroup (for which the superfamily is named). Our clinical stage protein therapeutic candidates focus on the activin, GDF and BMP subgroups.

We believe that, by employing our proprietary discovery and development platform, we can design protein therapeutic candidates that alter TGF- $\beta$  superfamily signaling and unlock the therapeutic potential of this group of proteins.

***Acceleron Approach***

By combining the powerful biology of the TGF- $\beta$  superfamily with our discovery and development expertise and our internal protein engineering and manufacturing capabilities, we have built a robust clinical and preclinical pipeline of protein therapeutic candidates targeting key mechanisms underlying cancer and rare diseases.

We have taken a comprehensive, receptor-focused approach to access the biology of the TGF- $\beta$  superfamily. We recognized that the 12 receptors for the superfamily act as control points for the ligands and therefore represent an attractive approach for pharmacological intervention. We have in-licensed patent rights for nine of the 12 receptors and systematically evaluated interactions between each receptor and a comprehensive panel of ligands. In the body, these ligands are naturally regulated by trap proteins that bind to the ligands thereby blocking ligand-receptor interactions and diminishing signaling in the cell. To mimic this natural regulatory approach, we have built our protein therapeutic



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candidates using the ligand-binding part of the receptors, depicted in the upper part of the figure below, as traps that capture the relevant groups of ligands in each biological process. We link the ligand-binding portion, the extracellular domain, of these receptors to the portion of a human antibody known as the Fc domain, depicted in the lower part of the figure below, which confers favorable pharmaceutical properties. The resulting "fused" proteins can be administered by simple intravenous or subcutaneous injection and reside in the blood for sufficient periods of time to permit dosing on a weekly or monthly basis.

Protein therapeutics constructed this way are referred to as "receptor fusion proteins" or "ligand traps". Some of the most successful protein therapeutics on the market belong to this category including Enbrel® (etanercept), Eylea® (aflibercept) and Orencia® (abatacept).

As shown in the figure below, our receptor fusion proteins act as ligand traps by binding to ligands of the TGF- $\beta$  superfamily, preventing those ligands from binding to the cell surface receptors, and thereby preventing activation of Smad proteins in the target cell.

To take full advantage of our proprietary discovery and development platform, we have developed an integrated set of technologies and capabilities to rapidly and cost-effectively create, test and advance multiple protein therapeutic candidates. Our protein engineering expertise allows us to create and optimize our receptor fusion proteins. We have developed the capability to generate recombinant cell lines that produce our protein therapeutic candidates, and assess the activity of these molecules in



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animals using our internal animal pharmacology facility or the capabilities of our academic collaborators. We have also invested in infrastructure to manufacture Phase 1 and Phase 2 clinical material quickly and flexibly using our internal current good manufacturing practices, or cGMP, compliant protein production facility to support clinical development of our protein therapeutic candidates.

We use our integrated platform of research, development and manufacturing technologies to rapidly and cost-effectively create, test and advance our protein therapeutic candidates. Our robust clinical and preclinical pipeline is focused on areas of high-unmet medical need, particularly in the areas of cancer and rare diseases.

**Our Product Pipeline**

We have four development stage protein therapeutic candidates, of which three are currently in numerous ongoing clinical trials and the fourth we expect to begin human clinical trials by the end of 2014. Celgene is currently conducting four Phase 2 clinical trials and overseeing three investigator-sponsored trials with sotatercept. We are conducting two Phase 2 clinical trials with ACE-536, two Phase 2 clinical trials with dalantercept and overseeing a collaborative group-sponsored Phase 2 clinical trial of dalantercept. We expect to initiate a Phase 2 clinical trial with dalantercept in patients with hepatocellular carcinoma in the first half of 2014 and to initiate a Phase 1 clinical trial with ACE-083 by the end of 2014.

**Sotatercept and ACE-536**

*Anemia in Patients with  $\beta$ -thalassemia and MDS*

Erythropoiesis, the process by which precursor cells proliferate and differentiate to give rise to red blood cells, is one of the most important and active processes in human biology. The primary role of red blood cells is to carry and deliver oxygen to other cells throughout the body. At any given time, there are approximately 25 trillion red blood cells in normal adult circulation which account for roughly 25% of the body's total number of cells. The human body produces 2.4 million new red blood cells each second. Red blood cell formation starts in the bone marrow with cells referred to as red blood cell precursors. These precursor cells go through many rounds of cellular proliferation, combined with cellular differentiation, to become more specialized cells to carry out their role as mature, functional red blood cells. We believe this highly active process of red blood cell production is normally tightly controlled by positive and negative regulators of the erythropoietic process. Erythropoietin is a positive regulator that stimulates proliferation of early red blood cell precursor cells, the BFU-E and CFU-E

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cells depicted in the figure below. Based on our research, it is now recognized that certain ligands in the TGF- $\beta$  superfamily are negative regulators of red blood cell precursors, starting with the Pro-E cells and those that follow, as depicted in the figure below. These members of the TGF- $\beta$  superfamily restrain the maturation of these precursors into later stage precursors and ultimately into functional red blood cells (RBCs).

**Depiction of Normal Erythropoiesis**

In certain diseases, the highly active process of red blood cell production does not function properly, leading to a reduction in the number of functional red blood cells, a condition known as anemia. Anemia in some disease settings is currently treated by the use of erythropoiesis stimulating agents, such as recombinant erythropoietin, that stimulate proliferation of early stage precursors of red blood cells. However, in certain diseases, such as  $\beta$ -thalassemia and MDS, anemia is caused by defects in the production of late stage red blood cell precursors, which is known as ineffective erythropoiesis.

Anemias caused by ineffective erythropoiesis are not well-treated by current therapies. As shown in the illustration below, ineffective erythropoiesis is characterized by an over-abundance of early stage red blood cell precursors and a decreased ability of late stage precursor cells to properly differentiate into healthy, functional red blood cells. The resulting anemia stimulates the body's overproduction of erythropoietin, which exacerbates the over-abundance of early stage precursors. Because the defective step in ineffective erythropoiesis lies downstream of the early stage precursors, the increase in the number of these cells fails to resolve the anemia.

**Depiction of Ineffective Erythropoiesis**



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Based on our preclinical research, we believe that TGF- $\beta$  superfamily ligands function as negative regulators of erythropoiesis by inhibiting the maturation of these early stage red blood cell precursors. Both sotatercept and ACE-536 are ligand traps designed to inhibit these negative regulators of late stage red blood cell precursors and promote their maturation into functional red blood cells.

We are developing sotatercept and ACE-536, through our collaborations with Celgene, as treatments for anemia in diseases in which erythropoiesis-stimulating agents are either not approved or are not well-suited to treat the underlying anemia. In diseases such as  $\beta$ -thalassemia and MDS in which anemia is caused by ineffective erythropoiesis, we believe both sotatercept and ACE-536 may help correct this defective process. Although similar in terms of their effects on red blood cells, there are differences in how these two protein therapeutic candidates bind to and inhibit ligands. Unlike ACE-536, sotatercept binds to and inhibits activin A, a TGF- $\beta$  superfamily ligand, and has been shown to increase bone mass and biomarkers of bone formation in clinical trials. Given its effects on bone, sotatercept is being studied in patients with chronic kidney disease, where it has the potential to treat both anemia and mineral and bone disorder. In addition, in preclinical studies, sotatercept inhibits the growth of myeloma cells. Therefore, sotatercept is also being studied in multiple myeloma patients to inhibit tumor growth and improve the anemia and the bone loss associated with the disease.

*$\beta$ -thalassemia*

The thalassemias comprise a heterogeneous group of disorders arising from defects in the genes that encode the proteins that comprise hemoglobin. Hemoglobin is a four-subunit protein complex formed of two  $\alpha$ -subunits and two  $\beta$ -subunits, each with an iron-containing heme group that binds to and carries oxygen molecules within red blood cells. There are two main classifications of thalassemia,  $\alpha$ -thalassemia and  $\beta$ -thalassemia, depending on whether the genetic defect lies in the gene encoding the  $\alpha$ -subunit or the  $\beta$ -subunit.  $\beta$ -thalassemia is particularly prevalent throughout the Mediterranean region, Middle East, and Southeast Asia, and, due to migration and immigration, is now a global disease. The Thalassaemia International Federation estimates that there are approximately 300,000 patients worldwide with  $\beta$ -thalassemia, approximately 20,000 of which are in the United States and Europe, who are dependent on frequent blood transfusions. We estimate that there are at least as many  $\beta$ -thalassemia patients in the same regions who are not transfusion dependent and not included in these estimates. Many of these patients have hemoglobin levels that are approximately half that of normal individuals and experience significant complications of the disease.

Anemia of  $\beta$ -thalassemia is primarily a result of ineffective erythropoiesis. The genetic defect leads to decreased production of the  $\beta$ -subunits of hemoglobin resulting in an excess amount of the  $\alpha$ -subunits. In normal erythropoiesis, excess unpaired  $\alpha$ -subunits are eliminated by a cellular component called the proteasome. The proteasome is normally required for effective red blood cell maturation to selectively remove cellular components and organelles such as mitochondria which are replaced by hemoglobin, which constitutes 90% of the protein in a mature red blood cell. In thalassemia, the proteasome becomes saturated with the abnormally high levels of unpaired  $\alpha$ -subunits and is unable to remove other cellular components and participate in the maturation process; this causes the block in maturation. Moreover, those free  $\alpha$ -subunits that are not eliminated by the proteasome form aggregates, called hemichromes, which damage the maturing red blood cells. These hemichromes, along with the saturation of the proteasome by unpaired  $\alpha$ -subunits, contribute to the ineffective erythropoiesis of  $\beta$ -thalassemia. The damaged red blood cells are filtered out by the spleen and have a reduced life span, resulting in anemia and enlargement of the spleen.

Patients with the most severe form of  $\beta$ -thalassemia produce few, if any,  $\beta$ -subunits, resulting in an increased amount of free  $\alpha$ -subunits and consequently a high number of hemichromes. These patients typically present with life-threatening anemia within the first year of life and require regular and lifelong red blood cell transfusions, usually every 2 to 4 weeks. Because red blood cells contain significant amounts of iron, this intensive transfusion regimen contributes to a condition known as iron

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overload, which is the principal cause of mortality. Consequently, therapy to reduce iron overload, called iron chelation therapy, is also part of standard treatment in these patients and typically begins after patients have received approximately 20 transfusions during their lifetime. Iron chelation therapy alone costs between \$25,000 and \$40,000 per year and yet does not treat the underlying anemia. The course of the disease depends largely on whether patients are maintained on an adequate transfusion and iron chelation regimen. Poor compliance with transfusion and/or iron chelation is associated with a poor prognosis and shortened survival. However, even with the standard of care, patients are at risk of infection from transfusions as well as toxicities related to iron chelation therapy.

Patients with an intermediate form of  $\beta$ -thalassemia, who are not necessarily dependent on frequent transfusions early in life, nevertheless suffer from a wide range of debilitating conditions. The ongoing ineffective erythropoiesis leads to various complications affecting a wide range of organ systems. By the second decade of life, most of these patients' hemoglobin levels have declined to the 6-8 g/dL range, or approximately half that of normal individuals. In an attempt to correct this chronic anemia, the body produces high levels of erythropoietin resulting in a continued stimulation of the early red blood cell precursors in the bone marrow. The number of these precursors grows to such an extent in the bone marrow that it leads to skeletal deformities, porosity of the long bones, and bone fractures. Splenomegaly, or enlargement of the spleen, is the result in part of continuous clearance by the spleen of the malformed red blood cells damaged by hemichromes. This commonly leads patients to require removal of their spleen, which in turn leads to worsening of other complications, such as blood clots. Iron overload is another significant complication even in the absence of red blood cell transfusions. This is due to increased intestinal iron absorption as a result of the ongoing ineffective erythropoiesis. Patients also suffer from various endocrine disorders due, in large part, to the accumulation of iron in the endocrine glands. Importantly, iron can also accumulate in the liver and heart, leading to severe complications such as liver fibrosis and heart failure.

No drug is approved to treat the anemia of  $\beta$ -thalassemia. Hematopoietic stem cell transplantation is viewed as the only curative approach for  $\beta$ -thalassemia, although this option is limited by the availability of appropriate donors and by risks, including death, associated with the bone marrow transplant procedure. Consequently this treatment is used only in the most severely affected patients.

***Myelodysplastic Syndromes***

Myelodysplastic syndromes, or MDS, are a group of heterogeneous hematologic diseases characterized by abnormal proliferation and differentiation of blood precursor cells, including red blood cell precursors, in the bone marrow. This leads to peripheral reductions in red blood cells, often accompanied by decreases in white blood cells and platelets, as well as a risk of disease progression to acute myeloid leukemia. Anemia is present in the vast majority of MDS patients at the time of diagnosis. MDS is primarily a disease of the elderly, with 88% of cases diagnosed in individuals 60 years of age or older. Cancer surveillance databases estimate the annual incidence of MDS in the United States at 10,000 to 15,000 cases and the overall U.S. prevalence at approximately 30,000 to 60,000 patients.

Hematopoietic stem cell transplantation represents the only treatment modality with curative potential, although the relatively high morbidity and mortality of this approach limits its use. Approximately 23% of MDS patients are categorized as intermediate-2 to high risk. These patients are typically treated with inhibitors of DNA methyltransferase such as Vidaza® (2012 U.S. sales of \$324 million for MDS) or Dacogen® (2012 U.S. sales of \$233 million for MDS). Of the remaining 77% of patients categorized as low to intermediate-1 risk, approximately 10% have a specific chromosomal mutation and are typically treated with Revlimid® (2012 U.S. sales of \$257 million for MDS). The remaining 67% of patients typically receive red blood cell transfusions or erythropoiesis stimulating agents, though erythropoiesis stimulating agents are not approved by the FDA or the EMA for the

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treatment of anemia in MDS patients. Our internal market research estimates that erythropoiesis stimulating agents generate \$500 to \$700 million in annual U.S. sales from their use in this disease.

The anemia in MDS is primarily due to ineffective erythropoiesis, and a significant number of MDS patients have serum erythropoietin levels substantially above the normal range, indicating that the anemia in these MDS patients is not a consequence of erythropoietin deficiency. The ineffective erythropoiesis of MDS may be caused by excess signaling by members of the TGF- $\beta$  superfamily, which signaling inhibits red blood cell maturation. For this reason we believe that blocking this excess signaling by sotatercept or ACE-536 may reverse this inhibition. Approximately 50% of MDS patients are unresponsive to the administration of recombinant erythropoietin and instead require red blood cell transfusions, which can increase the risk of infection and iron-overload related toxicities. Treatment-resistant anemia resulting from ineffective erythropoiesis is a major cause of morbidity in MDS patients.

***Chronic Kidney Disease***

Anemia is a common complication of chronic kidney disease. Because erythropoietin is produced primarily in the kidney and to a lesser extent in the liver, patients with chronic kidney disease produce sub-optimal amounts of erythropoietin, which leads to anemia. Additional serious complications of chronic kidney disease include a condition known as chronic kidney disease mineral and bone disorder that occurs when the diseased kidneys fail to maintain proper levels of calcium and phosphorous in the blood, leading to abnormal bone hormone levels, weakened bones and vascular calcification. Bone and vascular disorders are common complications in people with chronic kidney disease and bone disorders affect almost all patients receiving dialysis. According to the United States Renal Data System, there are over 400,000 chronic kidney disease patients receiving dialysis in the United States. Erythropoiesis stimulating agents have been approved for this indication for over twenty years. Sotatercept has the potential to differentiate itself from erythropoiesis stimulating agents in this patient population because of its positive effects on bone metabolism observed following the administration of sotatercept in preclinical models, healthy volunteers and cancer patients. Additionally, in mouse models of vascular calcification, sotatercept caused a reduction of calcified deposits in the aorta.

**Sotatercept Clinical and Preclinical Development**

Sotatercept is a soluble receptor fusion protein consisting of the extracellular domain of the activin receptor type IIA (ActRIIA) linked to the Fc domain of human IgG1. Sotatercept acts as a protein trap for TGF- $\beta$  superfamily ligands that signal through the ActRIIA receptor. Sotatercept has increased red blood cells in multiple clinical trials.

***Ongoing Phase 2 Clinical Trials of Sotatercept***

Our collaboration partner, Celgene, is currently conducting four Phase 2 clinical trials of sotatercept in patients with  $\beta$ -thalassemia, MDS and chronic kidney disease. The FDA has granted orphan designation for sotatercept for the treatment of  $\beta$ -thalassemia. We understand that Celgene plans to submit an application for orphan drug designation of sotatercept for treatment of MDS. Through collaborations with leading academic institutions, Celgene is also overseeing three investigator-sponsored trials.

***Celgene-Sponsored Clinical Trials***

***$\beta$ -thalassemia.*** Celgene is conducting a Phase 2 clinical trial of sotatercept designed as an ascending dose study to determine the safety and efficacy of sotatercept in adults with  $\beta$ -thalassemia. The dose levels to be studied are 0.1, 0.3, 0.5 mg/kg and 0.75 mg, with the possibility of further dose



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escalation up to 1.0 and 1.5 mg/kg, given subcutaneously once every three weeks for a period of 6 cycles with continued treatment at the discretion of the investigator for up to 22 months. Each cohort includes six or more patients during the dose escalation phase, followed by an expansion phase at a selected dose level in up to ten additional patients. The first patient in the trial was first dosed in November 2012. Celgene has completed enrolling the 0.1, 0.3, and 0.5 mg/kg cohorts and is now enrolling patients in the 0.75 mg/kg cohort. The primary outcome measure of the trial is to identify a safe dose level and to measure efficacy (1) in transfusion dependent patients by a reduction of transfusion burden by  $\geq 20\%$  compared to the pretreatment transfusion burden for each patient and (2) in non-transfusion dependent patients by an increase in hemoglobin level by  $\geq 1$  g/dL compared to the baseline hemoglobin, sustained for 12 weeks. This trial will also evaluate as exploratory endpoints the effects of sotatercept on iron overload, which is an important cause of morbidity and mortality associated with  $\beta$ -thalassemia, and bone metabolism. The trial is being conducted in six sites in Italy, France, Greece, and the United Kingdom and may enroll approximately 65 patients.

Sotatercept has generated encouraging preliminary data in the ongoing Phase 2 clinical trial of sotatercept in  $\beta$ -thalassemia patients.

As shown in the figure below, sotatercept has generated encouraging dose-dependent increases in hemoglobin levels in patients in the Phase 2 clinical trial who are non-transfusion dependent based on preliminary data from the three lowest dose levels in that trial.

**Mean Change in Hemoglobin Level From Baseline by Dose Cohort in  
Non-Transfusion Dependent  $\beta$ -Thalassemia Patients Treated With Sotatercept**

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Another analysis of the data from this trial also shows dose-dependent increases in hemoglobin levels. In the analysis shown below, within the first two months of receiving the first dose of sotatercept:

84% of non-transfusion dependent patients in each of the 0.5 and 0.3 mg/kg dose levels achieved at least a 1 g/dL increase in hemoglobin, while none of the non-transfusion dependent patients at the lowest dose level (0.1 mg/kg) achieved this threshold.

33%, 16% and 0% of non-transfusion dependent patients achieved a hemoglobin increase of at least 2 g/dL in the 0.5, 0.3, and 0.1 mg/kg dose levels, respectively.

**Maximum Change in Hemoglobin Level From Baseline in Non-Transfusion Dependent  $\beta$ -Thalassemia Patients During the First 3 Cycles (Day 64) of Sotatercept Treatment**

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The figure below shows that there is a statistically significant relationship ( $p < 0.001$ ) between drug exposure and the maximum increase in hemoglobin during the first three cycles across the three lowest dose levels of sotatercept. The x-axis shows the levels of sotatercept in patients' serum and the y-axis shows the patients' maximum change in hemoglobin. The figure illustrates that as the drug exposure in patients increases, so does the maximum increase in hemoglobin.

**Relationship Between Drug Exposure and Hemoglobin Level in Non-Transfusion Dependent  $\beta$ -Thalassemia Patients Through the First 3 Cycles (Day 64) of Sotatercept Treatment**

*Only patients completing the first 3 planned treatment cycles are included.  
(AUC) area under the curve; HgB, hemoglobin*

We expect Celgene to establish a range of recommended sotatercept dose levels, based on these data and additional data to be gathered as the clinical trials continue. We expect that in future clinical trials, patients will begin treatment at a recommended starting dose level and to undergo individualized dose titration based on hemoglobin response and tolerability to achieve and maintain an appropriate hemoglobin level. We expect Celgene to continue to dose escalate in this trial with the objective to determine the dose range for evaluation in the expansion stage of this trial. If this activity is confirmed with an acceptable safety profile, we and Celgene plan to initiate pivotal trial(s) in  $\beta$ -thalassemia by the end of 2014 or early 2015. At the dose levels that have been studied to date, we have not yet observed an effect in the transfusion dependent patients. Based on currently projected timelines, which are subject to change, we expect additional data from this clinical trial to become available as follows: data from additional dose levels and extended treatment in patients from the dose escalation portion of the clinical trial in the second quarter of 2014, and additional data in the fourth quarter of 2014.

**MDS.** Celgene is conducting a Phase 2 clinical trial of sotatercept for the treatment of anemia in patients with low-or intermediate-1 risk MDS. The dose levels to be studied are 0.1, 0.3, 0.5 and 1.0 and up to 2.0 mg/kg given subcutaneously once every three weeks for five cycles, and up to three additional cycles for late responders, with continued treatment at the discretion of the investigator. Each cohort may include up to 20 patients receiving a single dose level during the dose escalation

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phase, followed by an expansion phase at a selected dose level in up to 15 additional patients. The first patient in the trial was first dosed in December 2012. Celgene has currently completed the 0.1, 0.3 and 0.5 mg/kg cohorts and is now enrolling patients in the 1.0 mg/kg cohort. Dose escalation may go up to 2.0 mg/kg. The primary outcome measure is erythroid hematological improvement (HI-E). For patients who require transfusions of <4 units of red blood cells in the eight weeks prior to dosing, HI-E is an increase in hemoglobin of  $\geq 1.5$  g/dL sustained over a period  $\geq 8$  weeks in the absence of red blood cell transfusions. For subjects that require transfusions of  $\geq 4$  units of red blood cells in the eight weeks prior to dosing, HI-E is a decrease of  $\geq 4$  units of red blood cells transfused over a period of eight weeks compared to the number of units transfused in the eight weeks prior to treatment. This trial will also evaluate the effects of sotatercept on iron overload and bone metabolism. The trial is being conducted at up to 23 sites in the United States and France and may enroll up to 115 patients. Based on currently projected timelines, which are subject to change, we expect additional data from this clinical trial to become available as follows: data from the dose escalation portion of the clinical trial in the second quarter of 2014, and additional data in the fourth quarter of 2014.

**Chronic Kidney Disease.** Celgene is conducting two Phase 2 clinical trials with sotatercept in patients with chronic kidney disease. The first is a Phase 2 clinical trial with sotatercept designed as a randomized, placebo-controlled dose escalation study to evaluate the pharmacokinetics, safety, efficacy, tolerability and pharmacodynamics of sotatercept for the correction of anemia in patients with chronic kidney disease on hemodialysis. The first patient in the trial was first dosed in August 2010. The first dose level was 0.1 mg/kg administered subcutaneously as a single dose. Subsequent dose levels to be studied are 0.3, 0.5 and 0.7 mg/kg administered subcutaneously once every four weeks for up to eight cycles. Each cohort will include up to 12 (nine sotatercept-treated and three placebo-treated) patients receiving a single dose level during the dose escalation phase, followed by an additional cohort at a selected dose level. Celgene has completed enrollment in the 0.1, 0.3 and 0.5 mg/kg cohorts and is now enrolling patients in the 0.7 mg/kg cohort. The primary endpoints are pharmacokinetics and safety. Other endpoints include effects on hemoglobin and serum markers of bone metabolism. The trial is being conducted at up to 21 sites in the United States and may enroll up to 56 patients.

Early data from this trial are encouraging. An interim analysis from this clinical trial indicates that sotatercept produces dose dependent increases in hemoglobin in end stage renal disease patients on hemodialysis. The data will be presented at the National Kidney Foundation Spring Clinical Meeting in April 2014.

Based in part on these interim data, and previously observed effects of sotatercept on bone biomarkers, Celgene has initiated a second phase 2 clinical trial in Europe with sotatercept in patients with end stage renal disease (ESRD) who are on hemodialysis. The first patient in this trial was first dosed in December 2013. The study is designed as a two-part study to assess the safety and efficacy of sotatercept as a therapy to treat anemia and to control the adverse manifestations of chronic kidney disease-mineral and bone disorder (CKD-MBD). Patients in both parts of the study must first be on a stable dose of an erythropoiesis stimulating agent (ESA) to maintain hemoglobin levels and, after an ESA treatment free period of approximately five days, will then be switched to treatment with sotatercept.

The first part is a dose-escalation study of intravenous and subcutaneous routes of administration of sotatercept in approximately 60 patients to evaluate pharmacokinetics, safety and tolerability. Patients in the dose escalation part of the study will be given sotatercept once every two weeks up to a total of eight doses and followed for approximately four months after their last dose. The first part of the study is designed to inform the dosing regimens to be tested in the second part of the clinical trial. The second part will be a randomized, controlled study of approximately 230 patients to evaluate the efficacy and safety of sotatercept versus an erythropoiesis stimulating agent. Efficacy measures for part two of the study include the change in mean hemoglobin concentration from baseline and the ability of sotatercept to maintain patients' hemoglobin levels within a target range after switching from an ESA

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to sotatercept. Measures of biomarkers for bone formation and bone resorption and for mineral metabolism also will be studied, along with imaging of vascular calcification.

***Sotatercept Investigator Sponsored Trials***

Through collaborations with leading academic institutions, Celgene is overseeing investigator-sponsored trials in multiple myeloma, Diamond-Blackfan anemia and myelofibrosis.

Multiple myeloma is a cancer of the bone marrow that leads to the uncontrolled growth of certain white blood cells, causing bone marrow failure, bone pain, bone fractures and kidney problems. Nearly all multiple myeloma patients suffer from anemia. Investigators at the Massachusetts General Hospital are conducting a trial to explore the possibility that the combination of anti-myeloma therapies Revlimid® and dexamethasone together with sotatercept may reduce the growth of cancer cells along with improving anemia as well as bone lesions that often occur in patients with multiple myeloma.

Diamond-Blackfan anemia is a rare and severe anemia that is present at birth in affected individuals. Investigators at North Shore Long Island Jewish Health System are conducting a trial to determine the safety and efficacy of sotatercept in adults with Diamond-Blackfan anemia who are red blood cell transfusion-dependent.

Myelofibrosis is an acquired disease of the bone marrow that results in replacement of the bone marrow with fibrotic tissue leading to bone marrow failure and inability to make new blood cells, including red blood cells, which leads to anemia. Investigators at the MD Anderson Cancer Center are conducting a trial to determine the safety and efficacy of sotatercept in patients with myeloproliferative neoplasm-associated myelofibrosis and anemia.

***Completed Clinical Trials***

Six human clinical trials of sotatercept, including Phase 1 clinical trials in healthy volunteers and Phase 2 clinical trials of patients with multiple myeloma, breast cancer, and non-small cell lung cancer, collectively involving over 160 patients have been conducted to date. In healthy volunteers, we observed increases in red blood cells and hemoglobin. The mean change in hemoglobin for the patients who received a single dose of 1.0 mg/kg was almost 3 g/dL, which is similar to receiving a transfusion of three units of blood. We have also shown that in a randomized, placebo-controlled trial in patients with multiple myeloma receiving melphalan, prednisolone and thalidomide, sotatercept produced dose-dependent increases in hemoglobin. In the placebo and 0.1 mg/kg sotatercept cohorts, none of the patients achieved at least a 1.5 g/dL increase in hemoglobin at day 29 of the trial compared to their baseline levels. In the 0.3 and 0.5 mg/kg sotatercept cohorts, 13% and 38% of the patients, respectively, achieved at least a 1.5 g/dL increase in hemoglobin at day 29 of the trial compared to their baseline levels. In a randomized, placebo-controlled clinical trial in breast cancer patients who had anemia due to myelosuppressive chemotherapy, sotatercept produced dose-dependent increases in hemoglobin levels. In both the placebo and 0.1 mg/kg sotatercept cohorts, 20% of the patients had their hemoglobin levels increase to at least 11 g/dL maintained for 28 days in the absence of a red blood cell transfusion or use of an erythropoiesis stimulating agent. In the 0.3 mg/kg cohort, 22% of the patients achieved this outcome and in the 0.5 mg/kg cohort, 75% of the patients achieved this threshold. In a randomized, dose-ranging Phase 2 trial of sotatercept in patients with metastatic non-small cell lung cancer, sotatercept, administered at a fixed dose of 15 or 30 mg given subcutaneously every six weeks, produced increases in hemoglobin. In patients who did not receive red blood cell transfusions within the first four weeks, the change from baseline was at least 1 g/dL of hemoglobin for 40% of patients at week 2 and 16% of patients at week four. Given the results of these trials, we and Celgene may decide to pursue further clinical development in the future in one or more of these indications.

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***Safety***

Across the completed clinical trials, sotatercept has been generally well-tolerated. In studies with healthy volunteers, the only treatment-related serious adverse event was a report of persistent, progressive high blood pressure in one subject. While the precise cause of elevated blood pressure cannot be determined, it was an expected consequence of elevated red blood cell levels that occurred in this subject. Commonly observed adverse events included headache, infection, dizziness, hypertension, hot flush, tingling, muscle spasms, limb injury, fatigue and asthenia. In three studies of patients with cancer (myeloma, breast and lung cancer), one sudden death was reported in a myeloma patient. The event was evaluated as probably related to the concurrent anti-myeloma therapy of melphalan, prednisolone and thalidomide and possibly related to sotatercept. One patient with advanced breast cancer experienced serious adverse events of perforated gastric ulcer and peptic ulcer disease that were evaluated as possibly related to sotatercept. One patient with advanced lung cancer experienced a serious adverse event of a cerebrovascular accident (blockage of a blood vessel in the brain) that was suspected as related to treatment.

Among the ongoing clinical trials managed by Celgene, as of December 23, 2013, no treatment-related serious adverse events have been reported in the MDS trial. In the  $\beta$ -thalassemia trial as of December 9, 2013, two patients have exhibited serious adverse events that were suspected as related to sotatercept: bone pain and superficial thrombophlebitis (an inflamed blood clot in a superficial vein). One patient at the 0.5 mg/kg dose level had a treatment-related Grade 3 adverse event of ventricular extrasystoles (ventricular heart contractions) and discontinued treatment.

***Sotatercept Investigational New Drug (IND) Applications***

Sotatercept is the subject of three separate company-sponsored U.S. IND applications. We submitted the first IND to the FDA on March 13, 2006 for the treatment of postmenopausal osteoporosis. There are currently no studies being conducted under this IND. We submitted the second IND to the FDA on March 27, 2009 to assess the use of sotatercept for the treatment of anemia in various cancer-related indications. We transferred sponsorship of both INDs to Celgene on January 19, 2010. Under the second IND, sotatercept is currently being studied in patients with lower-risk MDS. A third IND was submitted by Celgene to the FDA on January 25, 2010 to assess sotatercept for the treatment of anemia in patients with end-stage renal disease. In addition, sotatercept is being studied in Europe under three separate Clinical Trial Applications (CTAs). The first CTA is for a Phase 2 study for the treatment of anemia in adult patients with  $\beta$ -thalassemia, submitted to France on December 28, 2011, to the United Kingdom on July 26, 2012, to Italy on July 27, 2012, and to Greece on November 23, 2012. The second CTA is for a Phase 2 study for the treatment of anemia in patients with lower-risk MDS, submitted to France on October 10, 2012. The third CTA is for a Phase 2 study for the treatment of anemia in patients with chronic kidney disease, with Belgium, Germany, Portugal, Spain, and the UK joining under the Voluntary Harmonization Procedure on June 17, 2013. Sotatercept is also being studied in the United States under three investigator-sponsored INDs.

***Preclinical Studies***

In preclinical studies, RAP-011 (the mouse equivalent of sotatercept) was evaluated in a broad range of animal pharmacology studies to assess its biological effects. RAP-011 has been shown to increase red blood cell counts in mice, rats, and monkeys. RAP-011 showed increased hemoglobin and red blood cell counts in mouse models of  $\beta$ -thalassemia and MDS, demonstrating decreased ineffective erythropoiesis in these models. RAP-011 was also able to prevent chemotherapy-induced anemia in a mouse model of this condition and was able to correct anemia in a mouse model of chronic kidney disease. RAP-011 increased bone mineral density in ovariectomized mice and has demonstrated positive effects in mice on bone lesions and bone metastases in a number of cancer models including models of

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multiple myeloma. The preclinical activity of sotatercept is also being evaluated in a mouse model of sickle cell disease.

**ACE-536 Clinical and Preclinical Development**

ACE-536 is a soluble receptor fusion protein consisting of a modified extracellular domain of the activin receptor type IIB (ActRIIB) linked to the Fc domain of human IgG1.

***Ongoing Phase 2 Clinical Trials of ACE-536***

We are conducting Phase 2 clinical trials of ACE-536 in patients with  $\beta$ -thalassemia and in patients with MDS. The FDA has granted orphan designation for ACE-536 for the treatment of  $\beta$ -thalassemia and for the treatment of MDS.

***$\beta$ -thalassemia.*** We are conducting a Phase 2 clinical trial of ACE-536, designed as an ascending dose trial to evaluate the safety and efficacy in patients with  $\beta$ -thalassemia. The dose levels to be studied are 0.2, 0.4, 0.6, 0.8 and 1.0 mg/kg given subcutaneously once every three weeks for up to 85 days. Each cohort will include three to six patients receiving a single dose level during the dose escalation phase. This will be followed by an expansion phase at a selected dose level in up to 20 patients. We are in the process of amending the protocol to include additional types of transfusion-dependent patients, to increase the size of the expansion cohort of the trial and to potentially study higher dose levels of ACE-536. The first patient in the trial was first dosed in March 2013. We have completed enrollment of the 0.2, 0.4 and 0.6 mg/kg cohorts and are currently enrolling patients in the 0.8 mg/kg cohort. The primary outcome measure is the proportion of patients who have an increase in hemoglobin of  $\geq 1.5$  g/dL from baseline for  $\geq 14$  days (in the absence of red blood cell transfusions) in non-transfusion dependent patients or a  $\geq 20\%$  reduction in red blood cell transfusion burden compared to the pretreatment transfusion burden in transfusion dependent patients. This trial will also examine the effects of ACE-536 on iron overload, an important cause of morbidity and mortality in  $\beta$ -thalassemia patients. Secondary endpoints include markers of serum iron and hemolysis. The trial is being conducted at up to seven sites in Italy and Greece, and we plan to include additional sites in Europe and may enroll up to 72 patients.

Initial data from this clinical trial is encouraging. As of January 3, 2014, preliminary data after three cycles (approximately two months) of treatment with ACE-536 show that non-transfusion dependent  $\beta$ -thalassemia patients in the 0.6 mg/kg dose group achieved a mean increase in hemoglobin of approximately 1.5 g/dL, while patients in the 0.2 and 0.4 mg/kg dose groups achieved a mean increase in hemoglobin of approximately 0.0 and 0.8 g/dL, respectively. Based on these data and additional data to be gathered during the clinical trial, we expect to establish a range of recommended ACE-536 dose levels. We expect that in future clinical trials, patients will begin treatment at a recommended starting dose level and to undergo individualized dose titration based on hemoglobin response and tolerability to achieve and maintain an appropriate hemoglobin level.

Based on currently projected timelines, which are subject to change, we expect data from this clinical trial to become available as follows: data from the dose escalation portion of the clinical trial during the second quarter of 2014, and additional data in the fourth quarter of 2014.

***MDS.*** We are conducting a Phase 2 clinical trial of ACE-536 designed as an ascending dose trial in patients with low or intermediate-1 risk MDS. The dose levels to be studied are 0.125, 0.25, 0.5, 0.75, 1.0, 1.33 and 1.75 mg/kg given subcutaneously once every three weeks for up to 85 days. Each cohort will include three to six patients receiving a single dose level during the dose escalation phase. This will be followed by an expansion phase at a selected dose level in up to 30 patients. The first patient in the trial was first dosed in January 2013. We have currently completed enrollment in the 0.125, 0.25, 0.5, 0.75 and 1.0 mg/kg cohorts and are now enrolling patients in the 1.33 mg/kg cohort. The primary outcome measure is the proportion of patients who have an increase of hemoglobin  $\geq 1.5$  g/dL from

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baseline for 14 days in the absence of red blood cell transfusions in non-transfusion dependent patients or a  $\geq 50\%$  or  $\geq 4$  unit reduction of red blood cell transfusions over a period of eight weeks compared to pretreatment transfusion burden in transfusion-dependent patients. This trial will also examine the effects of ACE-536 on iron overload. The trial is being conducted at up to nine sites in Germany and may enroll up to 72 patients. Based on currently projected timelines, which are subject to change, we expect additional data from this clinical trial to become available as follows: data from the dose escalation portion of the clinical trial during the second quarter of 2014, and additional data in the fourth quarter of 2014.

***Completed Phase 1 Clinical Trial***

ACE-536 was studied in a double-blind, placebo-controlled, randomized, ascending dose Phase 1 clinical trial in 32 healthy volunteers. ACE-536 produced dose-dependent increases in hemoglobin and red blood cells. The proportion of subjects with a hemoglobin increase of  $\geq 1.0$  g/dL increased on a dose-dependent basis, with approximately 80% of subjects in the 0.25 mg/kg dose level achieving this threshold.

***Safety***

In the completed Phase 1 clinical trial in healthy volunteers, ACE-536 was well-tolerated. No ACE-536 related serious adverse events were reported in the completed Phase 1 clinical trial. Commonly observed possibly or probably treatment-related adverse events included injection site bruising, injection site blemish, dry skin, numbness, muscle spasms, muscle pain, generalized itchiness and raised rash. In the ongoing Phase 2 clinical trials, there have been no ACE-536 related serious adverse events reported as of January 3, 2014. One patient in the ACE-536 thalassemia trial who was treated at the 0.8 mg/kg dose level had dose-limiting toxicity of worsening lumbar spine bone pain. The patient had a dose reduction to 0.6 mg/kg for the second cycle and subsequently withdrew from the study.

***ACE-536 Investigational New Drug (IND) Applications***

ACE-536 is being studied in the United States under an IND that we submitted to the FDA on June 14, 2011. The indication identified in the IND is for the treatment of anemia in patients with MDS. No studies are being conducted under this IND at this time. In addition, ACE-536 is being studied in Europe under two separate Clinical Trial Applications (CTAs). The first is for a Phase 2 study for the treatment of anemia in adult patients with  $\beta$ -thalassemia, submitted to Italy on August 29, 2012, to Turkey on June 14, 2013, and to Greece on July 2, 2013. The second is for a Phase 2 study for the treatment of anemia in patients with low- or intermediate-1 risk MDS, submitted to Germany on August 21, 2012.

***Preclinical Studies***

A number of preclinical pharmacology studies have been conducted with ACE-536 or its mouse version, RAP-536, that demonstrate its effects on red blood cells, hemoglobin and hematocrit. Collectively ACE-536 and RAP-536 have shown activity in mouse models of  $\beta$ -thalassemia, MDS, chemotherapy-induced anemia, acute blood loss and renal anemia.

***$\beta$ -thalassemia.*** RAP-536 has been evaluated in a series of studies using a mouse model of  $\beta$ -thalassemia. These mice carry deletion mutations in the  $\beta$ -globin genes, resulting in a deficiency of  $\beta$ -globin protein and hematologic abnormalities very similar to those seen in human  $\beta$ -thalassemia patients, including severe anemia and the formation of hemichromes resulting in ineffective erythropoiesis. These mice also exhibit severe complications common in patients with thalassemia, such as an enlarged spleen, bone loss and iron overload. In these mice, RAP-536 treatment improved



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numerous hematologic parameters, including significant increases in red blood cell count, hemoglobin levels, and hematocrit, decreased serum erythropoietin, normalized red blood cell size, and reduced red blood cell breakdown, as measured by serum bilirubin.

Representative blood smears were taken from the  $\beta$ -thalassemia mouse studies for both the placebo-treated animals and the RAP-536 treated animals. As shown in the image below, RAP-536 improved red blood cell morphology by reducing the number of poorly formed and damaged red blood cells, and reducing the amount of cellular debris that results from dying red blood cells.

Importantly, RAP-536 improved the maturation of later stage red blood cell precursor populations, in the bone marrow and spleen, with concomitant reductions in the earlier-stage red blood cell precursor populations. We believe RAP-536 reduced the ineffective erythropoiesis by decreasing the formation of harmful hemichromes. It appears that RAP-536 may achieve this effect in part by stimulating the proteasome, thus promoting the removal of unpaired  $\alpha$ -hemoglobin and stimulating red blood cell maturation.

This reduction in ineffective erythropoiesis reduced severe and common complications of the disease in mice, evidenced by reduced iron deposition in organs, reduced spleen weights and normalized bone density. Based on the numerous beneficial effects of RAP-536 in this mouse model of  $\beta$ -thalassemia, we believe that it is modifying the disease and has the potential to do so in human patients.

**MDS.** In a mouse model of MDS, RAP-536 treated animals had statistically significant increases in red blood cell count, hemoglobin levels and hematocrit compared to controls. Additionally, RAP-536 reduced the ineffective erythropoiesis as evidenced by the improvement in the ratio of red blood cell precursors to other cells in the bone marrow.

**Sickle Cell Disease.** We and Celgene are exploring the preclinical activity of ACE-536 in a mouse model of sickle cell disease.

Taken together, our clinical and preclinical results suggest that ACE-536 could be a meaningful novel therapy to treat anemia.

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**Dalantercept**

*Inhibiting Angiogenesis to Limit Tumor Growth*

Angiogenesis is a process by which new blood vessels are formed. Angiogenesis can be simplified to two major stages the proliferative stage followed by the maturation stage. During the proliferative stage, vascular endothelial cells, the cells lining the inside of the blood vessels, multiply in number and migrate to the site where a new vessel will be formed. This proliferative stage is followed by the maturation stage during which the endothelial cells coalesce to form tubes which are then stabilized through the recruitment of perivascular cells that form an outer layer of the blood vessels resulting in fully formed, functional vessels.

Tumors depend on angiogenesis to form new blood vessels to supply nutrients and oxygen to feed the rapidly growing malignant cells. The principal molecule driving the proliferative stage of angiogenesis in tumors is a protein called vascular endothelial growth factor (VEGF). Inhibiting VEGF-driven angiogenesis to control tumor growth has become an important and widely-used approach to cancer treatment. There are several FDA-approved cancer drugs that inhibit the VEGF pathway, with over \$8 billion in aggregate worldwide sales. Despite the success of these drugs, many patients fail to respond or develop resistance to VEGF pathway inhibitor therapy, resulting in an unmet need for new therapies to inhibit angiogenesis by a different mechanism.

We are using our knowledge of the TGF- $\beta$  superfamily to develop dalantercept, a novel protein therapeutic candidate targeting the maturation stage of angiogenesis. Recently, the activin receptor-like kinase 1 (ALK1) has been recognized as a key regulator of the maturation stage of angiogenesis. ALK1 is one of the 12 receptors for ligands in the TGF- $\beta$  superfamily and is found primarily on endothelial cells. The importance of the ALK1 pathway in angiogenesis was discovered, in part, through research into the genetic basis of the disease hereditary hemorrhagic telangiectasia 2 (HHT-2) in which patients manifest vascular defects including reduced ability to form capillary beds, which are the networks of small blood vessels that connect arteries to veins and are necessary for nutrient and waste exchange in tissues. This research revealed that these patients have only one of two functional copies of the ALK1 gene.

We reasoned that leveraging the biology of the ALK1 pathway to inhibit maturation of blood vessels could impair the growth of tumors by limiting the development of capillary beds within the tumor. To test this hypothesis, mice with a predisposition to develop tumors were bred to have only one, rather than two copies, of the ALK1 gene. In response to the loss of half of the ALK1 genes, tumor growth and size and blood vessel density in the tumor were reduced by half. These results and additional research in the field have established the ALK1 signaling pathway as a promising target for developing a new class of anti-angiogenesis agents ALK1 pathway inhibitors.

We believe one promising opportunity for dalantercept will be its use in combination with VEGF pathway inhibitors because these agents target distinct sequential steps in angiogenesis. Moreover, we and others have hypothesized that agents, such as dalantercept, that inhibit vessel maturation are able to sensitize the tumor vasculature to the anticancer effects of VEGF pathway inhibition. We believe that newly formed blood vessels become more resistant to VEGF pathway inhibitors as they mature. Therefore we believe that by preventing blood vessel maturation, dalantercept may maintain newly formed vessels in an immature state that increases their susceptibility to VEGF pathway inhibitors.

We and our academic collaborators have also shown in two mouse cancer models that treatment with dalantercept decreases metastases. This is in contrast to VEGF pathway inhibitors that increase metastases in mouse cancer models.

We believe that a combination of ALK1 and VEGF pathway inhibitors could have application in a number of different oncology indications where VEGF pathway inhibitors are currently used. The currently approved VEGF pathway inhibitors include Avastin® (bevacizumab), Nexavar® (sorafenib),

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Sutent® (sunitinib), Inlyta® (axitinib), and Votrient® (pazopanib). Four large markets for which these drugs have been approved are non-small cell lung cancer, colorectal cancer, renal cell carcinoma and liver cancer.

*Non-Small Cell Lung Cancer (NSCLC).* The National Cancer Institute estimates there will be 228,190 new cases of lung cancer in the United States in 2013 with 159,480 deaths. In 2012, sales of Avastin® in NSCLC were \$1.2 billion in the United States and \$1.7 billion worldwide.

*Colorectal Cancer.* The National Cancer Institute estimates there will be 142,820 new cases of colon cancer or rectal cancer in the United States in 2013 with 50,830 deaths. In 2012, sales of Avastin® for colorectal cancer were \$1.2 billion in the United States and \$3.6 billion worldwide.

*Renal Cell Carcinoma.* The National Cancer Institute estimates there will be 65,150 new cases of renal cell carcinoma in the United States in 2013 with 13,680 deaths. In 2012, U.S. sales of drugs for renal cell carcinoma were \$1.2 billion, of which \$800 million were anti-angiogenesis drugs that target the VEGF pathway, principally Sutent®, Nexavar® and Avastin®. Worldwide sales in 2012 of drugs for renal cell carcinoma were \$3.1 billion, of which \$2.3 billion were drugs that target the VEGF pathway.

*Liver Cancer.* The National Cancer Institute estimates there will be 30,640 new cases of liver cancer in the United States in 2013 with 21,670 deaths. The only drug approved in the United States for the treatment of liver cancer is the VEGF pathway inhibitor Nexavar®. In 2012, sales of Nexavar® for liver cancer were \$189 million in the United States and \$756 million worldwide.

*Other Tumors.* One or more anti-angiogenesis agents are also approved as treatments for neuroendocrine tumors, thyroid cancer and glioblastoma.

*Developing Indications.* It is believed that angiogenesis is important in the growth and spread of a number of additional highly-vascularized cancers, including endometrial cancer (cancer of the uterus), ovarian cancer, and head and neck cancer. While no anti-angiogenesis agents are approved in the U.S. for these indications, Avastin® is approved in Europe for the treatment of ovarian cancer.

The first two cancers for which we are studying the combination of dalantercept plus a VEGF pathway inhibitor are renal cell carcinoma and liver cancer. In renal cell cancer, sunitinib and axitinib are the most prescribed VEGF pathway inhibitors for first and second line patients, respectively. In the first line setting, sunitinib results in progression-free survival rates of approximately 11 months. In the second line setting, for patients whose disease had progressed despite receiving sunitinib in the first line setting, axitinib produced a progression-free survival rate of approximately 4.8 months. We believe the combination of dalantercept plus axitinib in the second line setting has the potential to increase the rate of progression-free survival greater than axitinib alone. In liver cancer, the VEGF pathway inhibitor, sorafenib, is approved in the first line setting yet the unmet medical need remains significant. In the first line setting, sorafenib results in time to progression of approximately 5.5 months.

**Dalantercept Clinical and Preclinical Development**

Dalantercept is comprised of the extracellular domain of the ALK1 receptor linked to the Fc domain of IgG1. Dalantercept acts as a ligand trap for ligands in the TGF- $\beta$  superfamily that signal through the ALK1 receptor. We have completed a Phase 1 trial of dalantercept and are pursuing a program of ongoing and planned Phase 2 trials seeking to demonstrate single agent activity of dalantercept for advanced solid tumors and activity of dalantercept in combination with approved VEGF pathway inhibitors or chemotherapy in advanced solid tumors.

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***Ongoing and Planned Phase 2 Clinical Trials of Dalantercept***

We are currently conducting two Phase 2 clinical trials of dalantercept in head and neck cancer and renal cell carcinoma. Additionally, through collaborations with a National Cancer Institute funded collaborative research group, the Gynecologic Oncology Group, we are overseeing an additional Phase 2 clinical trial in ovarian cancer. We plan to initiate a trial of dalantercept in hepatocellular carcinoma in the first half of 2014. We plan to submit applications for orphan designation of dalantercept for those indications or subsets of indications that meet FDA requirements for orphan status.

*Acceleron Sponsored Clinical Trials*

***Squamous Cell Carcinoma of the Head and Neck.*** We are conducting a single agent Phase 2 clinical trial of dalantercept in an ascending dose trial in patients with recurrent or metastatic squamous cell carcinoma of the head and neck. The first patient in the trial was first dosed in October 2011. After an initial cohort of two patients treated at a fixed dose level of 80 mg, we amended the trial and began recruitment of patients under the amended protocol in the first quarter of 2012 to study the dose level of 0.6 mg/kg given subcutaneously once every three weeks. The protocol was subsequently amended to increase the dose level of dalantercept to 1.2 mg/kg. Patients continue to receive dalantercept until there is disease progression (either clinically or as measured by analysis of radiographic imaging according to RECIST criteria) or dalantercept is no longer tolerated. The primary outcome measure is objective response rate as measured by RECIST criteria, and there are a number of secondary outcome measures of tumor response. The trial is being conducted in 12 centers in the United States, and we completed enrollment in July 2013 with a total of 46 patients, including two patients treated at the 80 mg dose, 13 at the 0.6 mg/kg dose, and 31 at the 1.2 mg/kg dose. Of these 46 patients, 41 patients (one patient at 80 mg, 13 patients at 0.6 mg/kg, and 27 patients at 1.2 mg/kg) were evaluable for radiological response according to RECIST criteria as of December 20, 2013. The preliminary data for these 41 patients are as follows: no patients at 80 mg, three patients (23%) at 0.6 mg/kg and ten patients (37%) at 1.2 mg/kg achieved stable disease as their best response at the beginning of their third cycle and, at the 1.2 mg/kg dose level, one patient (2.4%) achieved a partial response. None of these patients achieved a complete response. These preliminary data suggest dalantercept has dose dependent but modest single agent activity in patients with advanced squamous cell carcinoma of the head and neck. We believe the greatest potential for dalantercept will be in combination with VEGF pathway inhibitors or in combination with cytotoxic chemotherapy.

***Renal Cell Carcinoma.*** We are conducting a two-part Phase 2 clinical trial of dalantercept in combination with axitinib, an approved VEGF pathway inhibitor, in patients with advanced renal cell carcinoma. Part one of this trial is designed as a single-arm dose escalation and expansion stage with the primary endpoint of evaluating the safety and tolerability of various dose levels of dalantercept in combination with axitinib to select a dose level of dalantercept (in combination with axitinib) for further study if merited. The dose levels of dalantercept studied in the dose escalation stage were 0.6, 0.9, and 1.2 mg/kg given subcutaneously once every three weeks. The number of patients enrolled at the 0.6, 0.9 and 1.2 mg/kg dose levels were four, four and five patients, respectively. The dose level of 1.2 mg/kg was selected for use in the expansion phase and we are now enrolling up to 20 additional patients in this stage of the study. The first patient in the trial was first dosed in January 2013. Patients continue to receive dalantercept and axitinib until there is disease progression (either clinically or as measured by RECIST criteria) or the combination is no longer tolerated. Up to a total of 44 patients may be enrolled in the dose escalation and expansion part of the trial. Part two of the trial will be a randomized comparison of the selected dose of dalantercept in combination with axitinib versus axitinib alone with a total of 112 patients. The primary endpoint of part two of the trial will be progression-free survival. The trial is currently being conducted in 7 sites in the United States.

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We believe that early preliminary data from this trial are encouraging. No dose-limiting toxicities have been observed during the 29 day assessment period for each dose cohort. Of the 13 patients enrolled in the dose escalation part of the trial, seven (54%) had received one prior therapy and six (46%) had received  $\geq 2$  prior therapies, including two patients with  $\geq 2$  prior VEGF-pathway inhibitors. As of December 24, 2013, six of eleven (55%) evaluable patients completed  $\geq 6$  cycles (approximately 4.5 months) of treatment, including two of four patients at the 0.6 mg/kg dose level, two of four patients at the 0.9 mg/kg dose level and two of three patients at the 1.2 mg/kg dose level. These data provide preliminary evidence that dalantercept can be safely combined with the VEGF-pathway inhibitor axitinib in patients with renal cell carcinoma. As of the most recent analysis (August 28, 2013) of tumor response according to RECIST criteria in the first two cohorts, one patient had achieved a partial response and one patient had achieved stable disease at the 0.6 mg/kg dose level, and one patient had achieved a partial response and three patients had achieved stable disease at the 0.9 mg/kg dose level. Data from patients at the 1.2 mg/kg dose level have not yet been analyzed.

Based on currently projected timelines, which are subject to change, we expect to initiate part two of this trial by the end of the first quarter or the beginning of the second quarter of 2014, which, if successful, we expect would enable a Phase 3 trial. We expect additional dose escalation data to be available when we initiate part two of the trial.

***Hepatocellular Carcinoma***

In the first half of 2014, we plan to initiate a Phase 2 single-arm dose escalation and expansion clinical trial of dalantercept in combination with sorafenib, an approved VEGF pathway inhibitor, in patients with hepatocellular carcinoma. The primary endpoint will be to evaluate the safety and tolerability of various dose levels of dalantercept in combination with sorafenib. Secondary endpoints are expected to include time to progression, progression-free survival, disease control rate, and overall survival. The dose levels planned are 0.9 mg/kg of dalantercept given subcutaneously once every three weeks in combination with 400 mg sorafenib given once daily, 1.2 mg/kg of dalantercept given subcutaneously once every three weeks in combination with 400 mg sorafenib given once daily, and 1.2 mg/kg of dalantercept given subcutaneously once every three weeks in combination with 400 mg sorafenib given twice daily. Each cohort will include up to six patients receiving a single dose level during the dose escalation phase, followed by an expansion phase in up to 20 additional patients. Patients will continue to receive dalantercept and sorafenib until there is disease progression (either clinically or as measured by RECIST criteria) or the combination is no longer tolerated.

***Gynecologic Oncology Group (GOG) Sponsored Trials***

The Gynecologic Oncology Group, one of the National Cancer Institute's funded collaborative cancer research groups, sponsored two Phase 2 clinical trials to study the activity of dalantercept at a dose level of 1.2 mg/kg given as a single agent via subcutaneous injection every three weeks. The first trial was in patients with recurrent or persistent endometrial cancer and the second trial is in patients with recurrent or persistent ovarian cancer. Both of these clinical trials were designed as two part studies to assess the activity of dalantercept based on either of two endpoints: RECIST-defined response rate or progression free survival at 6 months. If there is sufficient activity in the first part of the trial, additional patients will be enrolled in the second, expanded part of the trial. The endometrial cancer study enrolled 28 patients in part one. Of the 28 patients, 16 (57.1%) achieved stable disease and 5 (17.9%) were alive and progression free for at least 6 months. Of these 5 patients, 3 were alive and progression free without receiving non-protocol therapy for at least 6 months. Fatigue, anemia, constipation, and edema were the most commonly reported toxicities regardless of attribution. The GOG has notified us that there was not sufficient activity to enroll additional patients in the second part of the endometrial cancer trial. The GOG ovarian cancer study enrolled 30 patients to part 1. We

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anticipate that in the middle of 2014, we may receive notification from the GOG if there is sufficient activity to enroll additional patients in the second part of the ovarian cancer trial.

***Phase 1 Trial Results***

A Phase 1 ascending dose trial evaluated the safety, tolerability, pharmacokinetics and anti-tumor activity of dalantercept in patients with advanced solid tumors. Dalantercept was given subcutaneously approximately once every three weeks until disease progression. Thirty-seven patients were enrolled in dose groups at 0.1, 0.2, 0.4, 0.8, 1.6, 3.2 and 4.8 mg/kg. In this trial, dalantercept demonstrated anti-tumor activity based on decreases or stabilization of tumor size. As shown in the figure below, out of the 29 evaluable patients treated, one (3%) had a partial response and 13 (45%) had stable disease according to RECIST criteria. Of the 13 who experienced stable disease, eight experienced stable disease for at least three months. Treatment continued until the patient experienced progressive disease.

The figure below displays each patient's best overall response by the maximum percent change decrease in target lesion size. The dose level each patient received is shown below their bar.

**Best Overall Response by the Maximum Percent Change Decrease in Target Lesion Size According to RECIST v1.1 Criteria**

In addition to these effects on tumor size, dalantercept demonstrated likely anti-angiogenic activity evidenced by a reduction of tumor metabolic activity as well as decreases in tumor blood flow. Lastly, some patients were observed to have dilated blood vessels in the skin, similar to those in HHT-2 patients, suggesting ALK1 pathway inhibition.

*Safety*

In clinical trials to date, dalantercept has been generally well-tolerated. In the initial Phase 1 clinical trial in advanced cancer patients, five patients out of 37 experienced serious adverse events deemed treatment-related that were reported as left ventricular dysfunction, fatigue, fluid overload, and



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congestive heart failure. Two of these patients had prior coronary artery disease. In subsequent trials fluid overload has been successfully managed with diuretics. As of December 24, 2013, the following treatment related adverse events have been observed in our ongoing clinical trials. Two patients in the head and neck cancer clinical trial have experienced serious adverse events of fluid accumulation around the lungs that were determined to be possibly related to dalantercept. Another patient in the head and neck trial has experienced serious adverse events of tracheal obstruction and pulmonary edema that were determined to be possibly related to dalantercept. In the clinical trial of patients with endometrial cancer, seven patients have experienced treatment-related serious adverse events reported as fluid accumulation in the abdominal cavity, fluid accumulation around the lungs, rectal fistula, gastric bleeding, vomiting, anemia, and shortness of breath. In the clinical trial of patients with ovarian cancer, one patient has experienced treatment related serious adverse events reported as hypokalemia (decreased potassium), anorexia, dehydration and increased creatinine. In the renal cell carcinoma trial, there have been no dalantercept-related serious adverse events. Adverse events associated with axitinib did not occur with higher than expected frequency or severity.

*Dalantercept Investigational New Drug (IND) Applications*

Dalantercept is being studied in the United States under an IND that we submitted to the FDA on July 29, 2009 for the treatment of patients with advanced solid tumors or multiple myeloma. Dalantercept is also being studied in the United States under two INDs sponsored by the Gynecologic Oncology Group: the first was submitted on August 2, 2012 for the treatment of recurrent/persistent endometrial carcinoma and the second submitted on September 25, 2012 for the treatment of recurrent/persistent ovarian carcinoma.

*Preclinical Studies*

We have demonstrated that dalantercept as a single agent inhibits tumor growth and angiogenesis in a variety of mouse models of cancer. Importantly, we have shown that dalantercept is a potent inhibitor of the maturation stage of angiogenesis. This is in contrast with VEGF pathway inhibitors that target the proliferative stage of angiogenesis.

We also demonstrated that dalantercept in combination with a VEGF pathway inhibitor provides enhanced anti-tumor effects. In mice bearing human renal cell carcinoma xenografts, we and our academic collaborators have shown that simultaneous administration of both dalantercept and sunitinib, a VEGF-receptor tyrosine kinase inhibitor, had substantially greater efficacy than either agent alone. In another mouse model of human renal cell carcinoma that develops resistance to sunitinib, tumor



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growth was blocked by the simultaneous administration of dalantercept. The figures below summarize those results.

**Dalantcept/Sunitinib Combination  
Exceeds Activity of Either Alone  
(Mouse Model of Renal Cell Carcinoma (A498))**

**Dalantcept/Sunitinib Combination  
Slows Tumor Growth in a Sunitinib Resistant Model  
(Mouse Model of Renal Cell Carcinoma (786O))**

*Collaboration with Drs. Wang, Bhatt, Mier, Atkins; Beth Israel Deaconess Medical Center, Boston*

**Development Objectives**

For sotatercept and ACE-536, our development strategy, determined in collaboration with Celgene, for both  $\beta$ -thalassemia and MDS is to conduct similar clinical trials with each protein therapeutic candidate in each disease essentially in parallel. For each disease, we and Celgene will review the data from both studies and determine which, if either, protein therapeutic candidate to move forward into subsequent, pivotal studies. It is our goal to initiate the Phase 3 clinical trials for one or both protein therapeutic candidates in one or both of these diseases by the end of 2014 or early 2015.

In addition, we and Celgene are performing preclinical research to assess the opportunity for sotatercept and ACE-536 to treat additional red blood cell disorders known as hemoglobinopathies, which include diseases such as thalassemias and sickle cell disease. Based on our encouraging preclinical and clinical data in  $\beta$ -thalassemia and our emerging understanding of the mechanism of action of these protein therapeutic candidates, we believe there is a potential for activity in other diseases such as sickle cell disease and preclinical studies are currently underway.

For dalantercept, our development strategy is to continue the renal cell carcinoma trial and to initiate during 2014 part two of the trial that compares the combination of dalantercept and axitinib to axitinib alone. We plan to initiate the hepatocellular carcinoma trial in the first half of 2014. We will also work toward completion of the ongoing single agent trial in head and neck cancer. We expect to initiate during the second half of 2014, at least one additional trial of dalantercept in cancer patients in combination with an approved VEGF pathway inhibitor. We are currently considering trials of dalantercept in combination with bevacizumab in glioblastoma (a form of brain cancer), and in colorectal cancer or lung cancer in combination with chemotherapy and/or a VEGF pathway inhibitor.

**Our Preclinical Pipeline**

We are using our discovery platform and knowledge of the TGF- $\beta$  superfamily to design and evaluate promising new protein therapeutic candidates that inhibit ligands of the TGF- $\beta$  superfamily.



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We have preclinical stage protein therapeutic candidates in our pipeline that have shown promising activity in animal models such as:

inhibition of liver fibrosis in mouse models of this condition;

improvement of cardiovascular function in a mouse model of a fibrotic disorder of the lungs; and

improvement in diseases of the eye such as in a mouse laser-induced neovascularization model of age-related macular degeneration (AMD).

***ACE-083***

We are developing a novel protein therapeutic candidate, ACE-083, for a first-in-human clinical trial that we expect to initiate by the end of 2014. ACE-083 acts as a trap for ligands in the TGF- $\beta$  superfamily that are known to be involved in the regulation of muscle mass. By inhibiting these ligands, ACE-083 can increase muscle mass, as we have demonstrated in animal studies. ACE-083 has been designed to affect those muscles in which the drug is injected. In preclinical animal studies, ACE-083 has shown dose dependent increases in muscle mass in the injected muscles but no systemic increases in muscle mass. We are focused on the development of ACE-083 for diseases in which increases in the size and function of specific muscles may provide a clinical benefit, including inclusion body myositis, facioscapulohumeral dystrophy (FSHD) and disuse atrophy.

**ACE-083 Selectively Doubles Muscle Mass in  
Injected Muscle with One Month of Treatment**

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p<0.05 vs. PBS (placebo) & vs. non-injected leg

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*ALK1 Pathway Inhibitors for the Treatment of Diseases of the Eye*

Although VEGF pathway inhibitors are the standard of care for age-related macular degeneration, or AMD, there is a need for improved therapies. Perivascular cell coverage may protect endothelial cells and limit the effectiveness of VEGF pathway inhibitors in AMD. By reducing the number of perivascular cells, the activity of VEGF pathway inhibitors on unprotected endothelial cells may be potentially enhanced. Human genetic evidence indicates that patients with hereditary hemorrhagic telangiectasia (HHT-2) who lack expression of a single ALK1 gene results in defective vasculature with reduced perivascular cell coverage. We are evaluating ALK1 pathway inhibitors, distinct from dalantercept, for use in the treatment of diseases of the eye. The combination of a VEGF pathway inhibitor and an ALK1 pathway inhibitor may lead to more effective treatment of neovascularization diseases of the eye including AMD.

**Our Strategic Partnerships**

Collaborations with corporate partners have provided us with significant funding and access to our partners' scientific, development, regulatory and commercial capabilities. We have received more than \$250.0 million from our collaborations with Celgene, Alkermes and Shire.

*Celgene*

On February 20, 2008 we entered into an agreement, which we refer to as the Sotatercept Agreement, with Celgene Corporation, under which we granted to Celgene worldwide rights to sotatercept. On August 2, 2011 we entered into a second agreement with Celgene for ACE-536, which we refer to as the ACE-536 Agreement under which we granted to Celgene worldwide rights to ACE-536 and also amended certain terms of the Sotatercept Agreement. These agreements provide Celgene exclusive licenses for these protein therapeutic candidates in all indications, as well as exclusive rights to obtain a license to certain future compounds.

**Sotatercept Agreement.** Under the terms of the Sotatercept Agreement, we and Celgene are collaborating on the development and commercialization of sotatercept. We also granted Celgene an option to license discovery stage compounds against three specified targets. Celgene paid \$45.0 million and bought \$5.0 million of equity upon execution of the Sotatercept Agreement and, as of September 30, 2013, we have received \$34.5 million in research and development funding and milestone payments for the sotatercept program.

We retained responsibility for research, development through the end of Phase 2a clinical trials, as well as manufacturing the clinical supplies for these trials. These activities are substantially complete. Celgene is conducting the current Phase 2 trials for  $\beta$ -thalassemia, MDS and chronic kidney disease and will be responsible for any future clinical trials for sotatercept as well as for all future manufacture of sotatercept. We are eligible to receive future development, regulatory and commercial milestones of up to \$360.0 million for the sotatercept program and up to an additional \$348.0 million for each of the three discovery stage programs. None of the three discovery stage programs has advanced to the stage to achieve payment of a milestone, nor do we expect any such milestone payments in the near future.

**ACE-536 Agreement.** Under the terms of the ACE-536 Agreement, we and Celgene are collaborating in the development and commercialization of ACE-536. We also granted Celgene an option to license products for which Acceleron files an investigational new drug application for the treatment of anemia. Celgene paid \$25.0 million to us upon execution of the ACE-536 Agreement in August 2011 and, as of September 30, 2013, we have received \$28.3 million in research and development funding and milestone payments for the ACE-536 program.

Under this agreement, we retained responsibility for research, development through the end of Phase 1 and the two ongoing Phase 2 clinical trials in MDS and  $\beta$ -thalassemia, as well as

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manufacturing the clinical supplies for these studies. Celgene will conduct subsequent Phase 2 and Phase 3 clinical trials. Acceleron will manufacture ACE-536 for all Phase 1 and Phase 2 clinical trials, and Celgene will have responsibility for the manufacture of ACE-536 for Phase 3 clinical trials and commercial supplies. We are eligible to receive future development, regulatory and commercial milestones of up to \$200.0 million for the ACE-536 program.

**Both Agreements.** Under each agreement, the conduct of the collaboration is managed by a Joint Development Committee and Joint Commercialization Committee. Other than with respect to certain matters related to our conduct of Phase 2 trials, in the event of a deadlock of a committee, the resolution of the relevant issue is determined by Celgene. Prior to January 1, 2013, Celgene paid the majority of development costs under the Sotatercept and ACE-536 Agreements. As of January 1, 2013, Celgene became responsible for paying 100% of worldwide development costs for both programs. Celgene will be responsible for all commercialization costs worldwide. We are obligated to co-promote sotatercept, ACE-536 and future products, in each case if approved, under both agreements in North America, and Celgene will pay all costs related thereto. We will receive tiered royalties in the low-to-mid 20% range on net sales of sotatercept and ACE-536. The royalty schedules for sotatercept and ACE-536 are the same. Celgene is obligated to use commercially reasonable efforts to develop and commercialize sotatercept and ACE-536. Celgene may determine that it is commercially reasonable to develop and commercialize only sotatercept or ACE-536 and discontinue the development or commercialization of the other protein therapeutic candidate, or Celgene may determine that it is not commercially reasonable to continue development of one or both of sotatercept and ACE-536. In the event of any such decision, we may be unable to progress the discontinued candidate or candidates ourselves. The agreements are terminable by either party upon a breach that is uncured and continuing or by Celgene for convenience on a country by country or product by product basis, or in its entirety. Celgene may also terminate the agreement, in its entirety or on a product by product basis, for failure of a product to meet a development or clinical trial endpoint. Termination for cause by us or termination by Celgene for convenience or failure to meet an endpoint will have the effect of terminating the applicable license, while termination for cause by Celgene will have the effect of reducing remaining royalties by a certain percentage.

***Other Collaborations***

**Alkermes.** On December 3, 2009, we entered into a Collaboration and License Agreement with Alkermes relating to a proprietary technology platform for extending the circulating half-life of certain proteins. Under the terms of the agreement, we granted Alkermes worldwide rights to apply this technology to proteins outside of the TGF- $\beta$  superfamily in return for an upfront license payment. We are entitled to future development, regulatory and sales milestones and mid-single digit royalties on product sales for each drug developed and commercialized by Alkermes using this technology. To our knowledge, Alkermes is not currently developing any products using this technology.

**Shire.** On September 8, 2010, we entered into an agreement with Shire AG for the joint development and commercialization of ACE-031, a clinical stage protein therapeutic candidate. We granted Shire an exclusive license in markets outside of North America. Under the terms of the agreement, Shire made an upfront cash payment of \$45.0 million. We received \$8.7 million in research and development payments from Shire during the term of the agreement. In April 2013, we and Shire determined not to further advance the development of ACE-031, and Shire terminated our collaboration agreement, effective as of June 30, 2013 and all rights reverted to us. We currently have no plans to continue the development of ACE-031.

**Competition**

The development and commercialization of new drugs is highly competitive. We and our collaborators will face competition with respect to all protein therapeutics we may develop or

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commercialize in the future from pharmaceutical and biotechnology companies worldwide. Many of the entities developing and marketing potentially competing products have significantly greater financial resources and expertise than we do in research and development, manufacturing, preclinical testing, conducting clinical trials, obtaining regulatory approvals and marketing. Our commercial opportunity will be reduced or eliminated if our competitors develop and commercialize products that are more effective, have fewer side effects, are more convenient or are less expensive than any products that we may develop.

If our clinical stage protein therapeutics are approved, they will compete with currently marketed drugs and therapies used for treatment of the following indications, and potentially with drug candidates currently in development for the same indications:

*β-thalassemia*

If either sotatercept or ACE-536 is approved for the treatment of patients with β-thalassemia, it would compete with:

Red blood cell transfusions and iron chelation therapy, such as Novartis's oral iron chelating agent, Exjade®. We are also aware that Shire is studying a new oral iron chelator in clinical trials.

Fetal hemoglobin stimulating agents, such as hydroxyurea, which are primarily used to treat patients with anemia from sickle cell disease, are sometimes used to treat patients with β-thalassemia. In addition, HQK-1001, a fetal hemoglobin stimulating agent being developed by HemaQuest Pharmaceuticals, Inc., has completed a Phase 1/2 clinical trial and an investigator sponsored Phase 2 clinical trial in patients with β-thalassemia.

Hematopoietic stem cell transplant treatment is given to a small percentage of patients with β-thalassemia, since it requires a sufficiently well-matched source of donor cells. Certain academic centers around the world are seeking to develop improvements to this approach.

Other therapies in development, including gene therapy are being developed by several different groups, including bluebird bio, Inc., Memorial Sloan Kettering Cancer Center, GlaxoSmithKline plc, and Sangamo BioSciences Inc. in collaboration with Biogen Idec.

**MDS**

If either sotatercept or ACE-536 is approved for the treatment of patients with MDS, it would compete with the following:

Recombinant erythropoietin and other erythropoiesis stimulating agents. Although these agents are not approved to treat anemia in MDS, current practice guidelines include the use of erythropoiesis stimulating agents and granulocyte colony stimulating factor agents (G-CSF) to treat patients with MDS. Additionally, Amgen's erythropoiesis stimulating agent, Aranesp®, is currently in Phase 3 clinical trials for treatment of anemia in patients with MDS.

Red blood cell transfusion and iron chelation therapy, including Exjade®, which is used to treat anemia in patients with MDS.

Immunomodulators, including Celgene's approved product, Revlimid® (lenalidomide), for the treatment of anemia of certain MDS patients.

Other therapies in development, including: an oral form of the hypomethylating agent azacitidine, known as CC-486, being developed by Celgene to treat patients with transfusion dependent anemia and thrombocytopenia due to lower risk MDS, which is currently in Phase 3

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clinical trials in the United States and Europe; and an anti-cancer therapy being developed by Onconova to treat patients with MDS.

***Chronic Kidney Disease***

If either sotatercept or ACE-536 is approved for the treatment of anemia in patients with chronic kidney disease, it would compete primarily with erythropoiesis stimulating agents that have been approved to treat these patients for over 20 years. In 2011, the Centers for Medicare and Medicaid Services (CMS) changed the reimbursement practice for erythropoiesis stimulating agents in chronic kidney disease patients on dialysis, which has led to changes in the way erythropoiesis stimulating agents are used in clinical practice, including decreasing the number of patients treated with erythropoiesis stimulating agents as well as decreasing the average dose and duration of therapy. These changes and the anticipated future introduction of biosimilar erythropoiesis stimulating agents are expected to generate additional price pressure in this market. Additionally, we are aware that Astellas Pharma and Fibrogen are developing oral, small molecule treatments that increase the production of erythropoietin to treat patients with anemia.

***Oncology Therapies***

We are developing dalantercept to be used in combination with VEGF pathway inhibitors for the treatment of cancer. If dalantercept is approved, it would compete with:

Other non-VEGF angiogenesis inhibitors in development, which also have the potential to be combined with VEGF pathway inhibitors or used independently of VEGF pathway inhibitors to inhibit angiogenesis. Amgen, Regeneron, MedImmune, OncoMed Pharmaceuticals, Pfizer and Tracoon are each developing non-VEGF angiogenesis inhibitors.

Pfizer's fully human monoclonal antibody to the ALK1 receptor, which is in Phase 2 trials in malignant pleural mesothelioma.

In addition to the therapies mentioned above, there are many generic chemotherapy agents and other regimens commonly used to treat various types of cancer, including renal cell carcinoma, head and neck, endometrial and ovarian cancer.

***Therapies for Treating Muscle Loss***

We are in the process of moving our lead pre-clinical protein therapeutic candidate, ACE-083, into its initial clinical trial. We plan to develop ACE-083 for the treatment of neuromuscular disorders and other diseases characterized by a loss of muscle function. We are aware of other approaches to treating muscle loss that are in clinical trials. Novartis is developing bimagrumab (BYM338), a monoclonal antibody targeting the activin receptor type IIB (ActRIIB), in various phase 2 clinical trials to treat pathological muscle loss and weakness. Nationwide Children's Hospital, in collaboration with The Myositis Association and Parent Project Muscular Dystrophy, is conducting a phase 1 clinical trial of a gene therapy delivery of follistatin (FS344) to muscle in patients with Becker muscular dystrophy (BMD) and sporadic inclusion body myositis (sIBM). Eli Lilly is developing, LY2495655, a myostatin monoclonal antibody in phase 2 clinical trials for disuse muscle atrophy and cancer-related cachexia. Regeneron and Sanofi are developing a myostatin monoclonal antibody, REGN1033 (SAR391786), which is in phase 1 clinical development for treatment of sarcopenia. Biogen Idec is conducting a phase 1 clinical trial of BIIB023, a monoclonal antibody targeting the tumor necrosis factor (TNF)-like weak inducer of apoptosis (TWEAK), to evaluate its effects on muscle atrophy in healthy male volunteers. Atara Bio is developing PINTA 745, a myostatin inhibiting peptibody to treat protein energy wasting in patients with end stage renal disease who are on dialysis.



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The key competitive factors affecting the success of any approved product will be its efficacy, safety profile, price, method of administration and level of promotional activity.

**Commercialization**

We retain co-promotion rights with our collaboration partner, Celgene, for both sotatercept and ACE-536 in North America, and under the terms of our agreements with Celgene, our commercialization costs will be entirely funded by Celgene. We also currently retain worldwide commercialization rights for our oncology protein therapeutic candidate, dalantercept, and our muscle protein therapeutic candidate, ACE-083. We intend to build hematology, oncology and neuromuscular disorder focused, specialty sales forces in North America and, possibly, other markets to effectively support the commercialization of these and future products. We believe that a specialty sales force will be sufficient to target key prescribing physicians in these areas. We currently do not have any sales or marketing capabilities or experience. We will establish the required capabilities within an appropriate time frame ahead of any product approval and commercialization to support a product launch. If we are not able to establish sales and marketing capabilities or are not successful in commercializing our future products, either on our own or through collaborations with Celgene, any future product revenue will be materially adversely affected.

**Intellectual Property**

Our commercial success depends in part on our ability to obtain and maintain proprietary protection for our protein therapeutics, novel biological discoveries, screening and drug development technology, to operate without infringing on the proprietary rights of others and to prevent others from infringing our proprietary rights. Our policy is to seek to protect our proprietary position by, among other methods, filing U.S. and foreign patent applications related to our proprietary technology, inventions and improvements that are important to the development and implementation of our business. We also rely on trade secrets, know-how, continuing technological innovation and potential in-licensing opportunities to develop and maintain our proprietary position. Additionally, we expect to benefit from a variety of statutory frameworks in the United States, Europe and other countries that relate to the regulation of biosimilar molecules and orphan drug status. These statutory frameworks provide periods of non-patent-based exclusivity for qualifying molecules. See "Government Regulations".

Our patenting strategy is focused on our protein therapeutics. We seek composition-of-matter and method-of-treatment patents for each such protein in key therapeutic areas. We also seek patent protection with respect to companion diagnostic methods and compositions and treatments for targeted patient populations. We have sought patent protection alone or jointly with our collaborators, as dictated by our collaboration agreements.

Our patent estate, on a worldwide basis, includes approximately 75 issued patents and approximately 300 pending patent applications, with pending and issued claims relating to all of our current clinical stage protein therapeutic candidates, sotatercept, ACE-536 and dalantercept. Of these, approximately 20 issued patents cover the nine receptors for the TGF- $\beta$  superfamily that we have selected as the core focus of our discovery approach. These figures include in-licensed patents and patent applications to which we hold exclusive commercial rights.

Individual patents extend for varying periods of time depending on the date of filing of the patent application or the date of patent issuance and the legal term of patents in the countries in which they are obtained. Generally, patents issued from applications filed in the United States are effective for twenty years from the earliest non-provisional filing date. In addition, in certain instances, a patent term can be extended to recapture a portion of the term effectively lost as a result of the FDA regulatory review period, however, the restoration period cannot be longer than five years and the total

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patent term including the restoration period must not exceed 14 years following FDA approval. The duration of foreign patents varies in accordance with provisions of applicable local law, but typically is also twenty years from the earliest international filing date. Our issued patents with respect to our receptor-focused platform will expire on dates ranging from 2013 to 2018, and, our issued patents and pending applications with respect to our protein therapeutic candidates will expire on dates ranging from 2026 to 2033, exclusive of possible patent term extensions. However, the actual protection afforded by a patent varies on a product by product basis, from country to country and depends upon many factors, including the type of patent, the scope of its coverage, the availability of extensions of patent term, the availability of legal remedies in a particular country and the validity and enforceability of the patent.

National and international patent laws concerning protein therapeutics remain highly unsettled. No consistent policy regarding the patent-eligibility or the breadth of claims allowed in such patents has emerged to date in the United States, Europe or other countries. Changes in either the patent laws or in interpretations of patent laws in the United States and other countries can diminish our ability to protect our inventions and enforce our intellectual property rights. Accordingly, we cannot predict the breadth or enforceability of claims that may be granted in our patents or in third-party patents. The biotechnology and pharmaceutical industries are characterized by extensive litigation regarding patents and other intellectual property rights. Our ability to maintain and solidify our proprietary position for our drugs and technology will depend on our success in obtaining effective claims and enforcing those claims once granted. We do not know whether any of the patent applications that we may file or license from third parties will result in the issuance of any patents. The issued patents that we own or may receive in the future, may be challenged, invalidated or circumvented, and the rights granted under any issued patents may not provide us with sufficient protection or competitive advantages against competitors with similar technology. Furthermore, our competitors may be able to independently develop and commercialize similar drugs or duplicate our technology, business model or strategy without infringing our patents. Because of the extensive time required for clinical development and regulatory review of a drug we may develop, it is possible that, before any of our drugs can be commercialized, any related patent may expire or remain in force for only a short period following commercialization, thereby reducing any advantage of any such patent. The patent positions for our most advanced programs are summarized below:

*Sotatercept Patent Coverage*

We hold two issued patents covering the sotatercept composition of matter in the United States, one issued patent in Europe (registered in most countries of the European Patent Convention) and additional patents issued or pending in many other major jurisdictions worldwide, including Japan, China, South Korea, Brazil, Mexico, Russia, Israel and India. The expected expiration date for these composition of matter patents is 2026 plus any extensions of term available under national law.

We hold two issued patents covering the treatment of anemia by administration of sotatercept in the United States and similar patents issued or pending in many other major jurisdictions worldwide, including Europe, Japan, China, South Korea, Brazil, Mexico, Russia, Israel and India. The expected expiration date for these composition of matter patents is 2027 exclusive of possible patent term extensions.

We also hold patents and patent applications directed to a variety of other uses for sotatercept, including the reduction of tumor cell burden in multiple myeloma.

*ACE-536 Patent Coverage*

We hold two issued patents covering the ACE-536 composition of matter in the United States, and additional patents issued or pending in many other major jurisdictions worldwide, including Europe,

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Japan, China, South Korea, Brazil, Mexico, Russia and India. The expected expiration dates for these composition of matter patents are 2028 and 2029, exclusive of possible patent term extensions.

We hold one issued patent covering the treatment of anemia by administration of ACE-536 in the United States and similar patents issued or pending in other major jurisdictions worldwide, including Europe, Japan, China, South Korea, Brazil, Mexico, Russia and India. The expected expiration date for these method of treatment patents is 2029 exclusive of possible patent term extensions.

*Dalantercept Patent Coverage*

We hold one issued patent covering the dalantercept composition of matter in the United States, which is expected to expire in 2029, exclusive of possible patent term extensions, and we hold additional pending patent applications. We hold additional issued patents and pending patent applications covering composition of matter in many other major jurisdictions worldwide, including Europe, Japan, China, South Korea, Brazil, Mexico, Russia and India. The expected expiration dates for these patent filings claiming the dalantercept composition of matter, if issued, are either 2027 or 2029, exclusive of possible patent term extensions.

We hold one issued patent covering the treatment of tumor angiogenesis by administration of dalantercept in the United States and similar patents issued or pending in other major jurisdictions worldwide, including Europe, Japan, China, South Korea, Brazil, Mexico, Russia and India. The expected expiration date for these method of treatment patents is 2027, exclusive of possible patent term extensions.

We also hold patent applications directed to a variety of other uses for dalantercept, including the treatment of renal cell carcinoma with a combination of dalantercept and a VEGF-targeted tyrosine kinase inhibitor. This patent application is jointly invented and owned with the Beth Israel Deaconess Medical Center, or BIDMC, and we have secured an exclusive license to the BIDMC rights. The expected expiration date for these patent applications, should they issue as patents, is 2033 plus any extensions of term available under national law.

*Trade Secrets*

In addition to patents, we rely upon unpatented trade secrets and know-how and continuing technological innovation to develop and maintain our competitive position. We seek to protect our proprietary information, in part, using confidentiality agreements with our commercial partners, collaborators, employees and consultants and invention assignment agreements with our employees. These agreements are designed to protect our proprietary information and, in the case of the invention assignment agreements, to grant us ownership of technologies that are developed through a relationship with a third party. These agreements may be breached, and we may not have adequate remedies for any breach. In addition, our trade secrets may otherwise become known or be independently discovered by competitors. To the extent that our commercial partners, collaborators, employees and consultants use intellectual property owned by others in their work for us, disputes may arise as to the rights in related or resulting know-how and inventions.

*In-Licenses*

Effective June 21, 2012, we entered into a license agreement with the Beth Israel Deaconess Medical Center, or BIDMC, to obtain worldwide, exclusive rights under patent filings jointly invented by us and BIDMC. The patent rights relate to the treatment of renal cell cancer by combination therapy with dalantercept and VEGF-receptor tyrosine kinase inhibitors (TKIs). The intellectual property includes one pending U.S. patent filing and one pending PCT (international) patent filing. If issued, the patents are predicted to expire in 2033. Under the agreement, BIDMC retained rights, on behalf of itself and other non-profit academic institutions, to practice under the licensed rights for

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non-profit purposes. The license rights granted to us are further subject to any rights the United States Government may have in such licensed rights due to its sponsorship of research that led to the creation of the licensed rights. We agreed to pay BIDMC specified development and sales milestone payments aggregating up to \$1.0 million. In addition, we are required to pay BIDMC royalties in the low single-digits on worldwide net product sales of drug labeled for treatment regimens that are claimed in the licensed patents. The agreement terminates upon the expiration of the last valid claim of the licensed patent rights. We may terminate the agreement at any time by giving BIDMC advance written notice. The agreement may also be terminated by BIDMC in the event of a material breach by us or in the event we become subject to specified bankruptcy or similar circumstances. In any termination event, we retain our joint ownership of the patent rights and a worldwide non-exclusive license with right to sublicense.

In August 6, 2010, we entered into an amended and restated license agreement with the Ludwig Institute for Cancer Research, or LICR, to obtain worldwide, exclusive rights under patent filings solely owned by LICR and patent rights jointly invented by us and LICR. The LICR-owned patent rights relate to the first cloning of the type I activin receptors, ALK1, ALK2, ALK3, ALK4, ALK5 and ALK6, and include claims to nucleic acids, proteins and antibodies with respect to each of the foregoing. These patent rights expire between the years 2013 and 2018. The license excludes the rights with regard to anti-ALK2 antibodies. The joint patent rights relate to the treatment of pancreatic tumors with dalantercept and, if issued, such patent rights are expected to expire in 2029. Under the agreement, LICR retained rights, on behalf of itself and other non-profit academic institutions, to practice under the licensed rights for non-profit purposes. We agreed to pay LICR specified development and sales milestone payments aggregating up to \$1.6 million for dalantercept. In addition, we are required to pay LICR royalties in the low single-digits on worldwide net product sales of products claimed in the licensed patents, with royalty obligations continuing at a 50% reduced rate for eight years after patent expiration. If we sublicense the LICR patent rights, we will owe LICR a percentage of sublicensing revenue, excluding payments based on the level of sales, profits or other levels of commercialization. The agreement terminates upon the expiration of royalty obligations. We may terminate the agreement at any time by giving LICR advance written notice. The agreement may also be terminated by LICR in the event of a material breach by us or in the event we become subject to specified bankruptcy or similar circumstances. In any termination we retain our joint ownership right in the jointly owned patent filings.

In August 2010, we entered into two amended and restated license agreements with the Salk Institute for Biological Studies, or Salk, providing rights under U.S. patent filings solely owned by Salk. The agreements for the licensed patent rights relate to the first cloning of the type II activin receptors, human ActRIIA and frog ActRIIB, respectively, and include claims to vertebrate homolog nucleic acids and proteins with respect to each of the foregoing. These patent rights expire between the years 2016 and 2017. One of these agreements relates to ActRIIA and sotatercept; the other agreement relates to ActRIIB, ACE-536 and the discontinued program ACE-031. The licenses granted are exclusive as to the therapeutic products that are covered by the patents and non-exclusive as to diagnostic products and other products that are developed using the Salk patent rights. If we sublicense the Salk patent rights, we will owe Salk a percentage of sublicensing revenue, excluding payments based on sales. Under the agreements, Salk retained rights, on behalf of itself and other non-profit academic institutions, to practice under the licensed rights for non-profit purposes. We agreed to pay Salk specified development milestone payments totaling up to \$2.0 million for sotatercept and \$0.7 million for ACE-536. In addition, we are required to pay Salk royalties in the low single-digits on worldwide net product sales by us or our sublicensees of products claimed in the licensed patents, or derived from use of the licensed patent rights, with royalty obligations continuing at a reduced rate for a period of time after patent expiration. The agreements terminate upon the expiration of royalty obligations. We may terminate either agreement at any time by giving Salk advance written notice. Either agreement

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may also be terminated by Salk in the event of a material breach by us or in the event we become subject to bankruptcy or similar circumstances.

**Government Regulation**

The preclinical studies and clinical testing, manufacture, labeling, storage, record keeping, advertising, promotion, export, marketing and sales, among other things, of our protein therapeutic candidates and future products, are subject to extensive regulation by governmental authorities in the United States and other countries. In the United States, pharmaceutical products are regulated by the FDA under the Federal Food, Drug, and Cosmetic Act and other laws, including, in the case of biologics, the Public Health Service Act. We expect sotatercept, ACE-536, and dalantercept to be regulated by the FDA as biologics and to be reviewed by the Center for Drug Evaluation and Research (CDER) as proteins intended for therapeutic use. Protein therapeutics require the submission of a Biologics License Application, or BLA, and approval by the FDA prior to being marketed in the U.S. Manufacturers of protein therapeutics may also be subject to state regulation. Failure to comply with FDA requirements, both before and after product approval, may subject us or our partners, contract manufacturers, and suppliers to administrative or judicial sanctions, including FDA refusal to approve applications, warning letters, product recalls, product seizures, total or partial suspension of production or distribution, fines and/or criminal prosecution.

The steps required before a biologic may be approved for marketing of an indication in the United States generally include:

completion of preclinical laboratory tests, animal studies and formulation studies conducted according to Good Laboratory Practices, or GLPs, and other applicable regulations;

submission to the FDA of an Investigational New Drug application or IND, which must become effective before human clinical trials may commence;

completion of adequate and well-controlled human clinical trials in accordance with Good Clinical Practices, or GCPs, to establish that the biological product is "safe, pure and potent", which is analogous to the safety and efficacy approval standard for a chemical drug product for its intended use;

submission to the FDA of a BLA;

satisfactory completion of an FDA pre-approval inspection of the manufacturing facility or facilities at which the product is produced to assess compliance with applicable current Good Manufacturing Practice requirements, or cGMPs; and

FDA review of the BLA and issuance of a biologics license which is the approval necessary to market a protein therapeutic.

Preclinical studies include laboratory evaluation of product chemistry, toxicity and formulation as well as animal studies to assess the potential safety and efficacy of the biologic candidate. Preclinical studies must be conducted in compliance with FDA regulations regarding GLPs. The results of the preclinical tests, together with manufacturing information and analytical data, are submitted to the FDA as part of an IND. Some preclinical testing may continue even after the IND is submitted. In addition to including the results of the preclinical testing, the IND will also include a protocol detailing, among other things, the objectives of the clinical trial, the parameters to be used in monitoring safety and the effectiveness criteria to be evaluated if the first phase or phases of the clinical trial lends themselves to an efficacy determination. The IND will automatically become effective 30 days after receipt by the FDA, unless the FDA within the 30-day time period places the IND on clinical hold because of its concerns about the drug candidate or the conduct of the trial described in the clinical protocol included in the IND. The IND sponsor and the FDA must resolve any outstanding concerns before clinical trials can proceed.

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All clinical trials must be conducted under the supervision of one or more qualified principal investigators in accordance with GCPs. They must be conducted under protocols detailing the objectives of the applicable phase of the trial, dosing procedures, research subject selection and exclusion criteria and the safety and effectiveness criteria to be evaluated. Each protocol must be submitted to the FDA as part of the IND, and progress reports detailing the status of the clinical trials must be submitted to the FDA annually. Sponsors also must timely report to the FDA serious and unexpected adverse reactions, any clinically important increase in the rate of a serious suspected adverse reaction over that listed in the protocol or investigator's brochure, or any findings from other studies or animal or in vitro testing that suggest a significant risk in humans exposed to the drug. An institutional review board, or IRB, at each institution participating in the clinical trial must review and approve the protocol before a clinical trial commences at that institution, approve the information regarding the trial and the consent form that must be provided to each research subject or the subject's legal representative, and monitor the study until completed.

Clinical trials are typically conducted in three sequential phases, but the phases may overlap and different trials may be initiated with the same drug candidate within the same phase of development in similar or differing patient populations. Phase 1 trials may be conducted in a limited number of patients, but are usually conducted in healthy volunteer subjects. The drug candidate is initially tested for safety and, as appropriate, for absorption, metabolism, distribution, excretion, pharmacodynamics and pharmacokinetics.

Phase 2 usually involves trials in a larger, but still limited, patient population to evaluate preliminarily the efficacy of the drug candidate for specific, targeted indications to determine dosage tolerance and optimal dosage and to identify possible short-term adverse effects and safety risks.

Phase 3 trials are undertaken to further evaluate clinical efficacy of a specific endpoint and to test further for safety within an expanded patient population at geographically dispersed clinical trial sites. Phase 1, Phase 2, or Phase 3 testing might not be completed successfully within any specific time period, if at all, with respect to any of our protein therapeutic candidates. Results from one trial are not necessarily predictive of results from later trials. Furthermore, the FDA or the sponsor may suspend clinical trials at any time on various grounds, including a finding that the subjects or patients are being exposed to an unacceptable health risk. Similarly, an IRB can suspend or terminate approval of a clinical trial at its institution if the clinical trial is not being conducted in accordance with the IRB's requirements or if the drug candidate has been associated with unexpected serious harm to patients.

The results of the preclinical studies and clinical trials, together with other detailed information, including information on the manufacture and composition of the product, are submitted to the FDA as part of a BLA requesting approval to market the drug candidate for a proposed indication. Under the Prescription Drug User Fee Act, as re-authorized most recently in July 2012, the fees payable to the FDA for reviewing a BLA, as well as annual fees for commercial manufacturing establishments and for approved products, can be substantial. The fee for review of an application that requires clinical data, such as a BLA, for the one year period ending September 30, 2013, is almost \$2.0 million, subject to certain limited deferrals, waivers, and reductions that may be available. The fees typically increase each year. Each BLA submitted to the FDA for approval is reviewed for administrative completeness and reviewability within 60 days following receipt by the FDA of the application. If the BLA is found complete, the FDA will file the BLA, triggering a full review of the application. The FDA may refuse to file any BLA that it deems incomplete or not properly reviewable at the time of submission. The FDA's established goal is to review 90% of priority BLA applications within six months after the application is accepted for filing and 90% of standard BLA applications within 10 months of the acceptance date, whereupon a review decision is to be made. The FDA, however, may not approve a drug candidate within these established goals and its review goals are subject to change from time to time. Further, the outcome of the review, even if generally favorable, may not be an actual approval

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but a "complete response letter" that describes additional work that must be done before the application can be approved. Before approving a BLA, the FDA may inspect the facility or facilities at which the product is manufactured and will not approve the product unless the facility complies with cGMPs. The FDA may deny approval of a BLA if applicable statutory or regulatory criteria are not satisfied, or may require additional testing or information, which can extend the review process. FDA approval of any application may include many delays or never be granted. If a product is approved, the approval may impose limitations on the uses for which the product may be marketed, may require that warning statements be included in the product labeling, may require that additional studies be conducted following approval as a condition of the approval, and may impose restrictions and conditions on product distribution, prescribing, or dispensing in the form of a Risk Evaluation and Mitigation Strategy, or REMS, or otherwise limit the scope of any approval. The FDA must approve a BLA supplement or a new BLA before a product may be marketed for other uses or before certain manufacturing or other changes may be made. Further post-marketing testing and surveillance to monitor the safety or efficacy of a product is required. Also, product approvals may be withdrawn if compliance with regulatory standards is not maintained or if safety or manufacturing problems occur following initial marketing. In addition, new government requirements may be established that could delay or prevent regulatory approval of our protein therapeutic candidates under development.

As part of the recently-enacted Patient Protection and Affordable Care Act of 2010, under the subtitle of Biologics Price Competition and Innovation Act of 2009, or the BPCI, a statutory pathway has been created for licensure, or approval, of biological products that are biosimilar to, and possibly interchangeable with, earlier biological products licensed under the Public Health Service Act. Also under the BPCI, innovator manufacturers of original reference biological products are granted 12 years of exclusivity before biosimilars can be approved for marketing in the United States. The objectives of the BPCI are conceptually similar to those of the Drug Price Competition and Patent Term Restoration Act of 1984, commonly referred to as the "Hatch-Waxman Act", which established abbreviated pathways for the approval of drug products. The implementation of an abbreviated approval pathway for biological products is under the direction of the FDA and is currently being developed. In late 2010, the FDA held a hearing to receive comments from a broad group of stakeholders regarding the implementation of the BPCI. Since that hearing in 2010, the FDA, in February 2012 and February 2013, has issued several draft guidances for industry related to the BPCI, addressing scientific, quality and procedural issues relevant to an abbreviated application for a biosimilar product. The approval of a biologic product biosimilar to one of our products could have a material adverse impact on our business as it may be significantly less costly to bring to market and may be priced significantly lower than our products.

Both before and after the FDA approves a product, the manufacturer and the holder or holders of the BLA for the product are subject to comprehensive regulatory oversight. For example, quality control and manufacturing procedures must conform, on an ongoing basis, to cGMP requirements, and the FDA periodically inspects manufacturing facilities to assess compliance with cGMPs. Accordingly, manufacturers must continue to spend time, money and effort to maintain cGMP compliance.

***Orphan Drug Act***

The Orphan Drug Act provides incentives to manufacturers to develop and market drugs for rare diseases and conditions affecting fewer than 200,000 persons in the United States at the time of application for orphan drug designation. Orphan drug designation must be requested before submitting a BLA. Orphan drug designation does not convey any advantage in, or shorten the duration of, the regulatory review and approval process. If a product that has orphan drug designation subsequently receives the first FDA approval for the disease for which it has such designation, the holder of the approval is entitled to a seven-year exclusive marketing period in the United States for that product except in very limited circumstances. For example, a drug that the FDA considers to be clinically

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superior to, or different from, another approved orphan drug, even though for the same indication, may also obtain approval in the United States during the seven-year exclusive marketing period. In addition, holders of exclusivity for orphan drugs are expected to assure the availability of sufficient quantities of their orphan drugs to meet the needs of patients. Failure to do so could result in the withdrawal of marketing exclusivity for the drug. ACE-536 has orphan drug designation in the United States for the treatment of  $\beta$ -thalassemia and for the treatment of MDS. The FDA has granted orphan designation for sotatercept for the treatment of  $\beta$ -thalassemia.

Legislation similar to the Orphan Drug Act has been enacted outside the U.S., including in the EU. The orphan legislation in the EU is available for therapies addressing chronic debilitating or life-threatening conditions that affect five or fewer out of 10,000 persons or are financially not viable to develop. The market exclusivity period is for ten years, although that period can be reduced to six years if, at the end of the fifth year, available evidence establishes that the product is sufficiently profitable not to justify maintenance of market exclusivity. The market exclusivity may be extended to 12 years if sponsors complete a pediatric investigation plan agreed upon with the relevant committee of the EMA.

*Expedited Review and Approval*

The FDA has various programs, including Fast Track, priority review, and accelerated approval, which are intended to expedite or simplify the process for reviewing drugs, and/or provide for the approval of a drug on the basis of a surrogate endpoint. Even if a drug qualifies for one or more of these programs, the FDA may later decide that the drug no longer meets the conditions for qualification or that the time period for FDA review or approval will be shortened. Generally, drugs that are eligible for these programs are those for serious or life-threatening conditions, those with the potential to address unmet medical needs and those that offer meaningful benefits over existing treatments. For example, Fast Track is a process designed to facilitate the development and expedite the review of drugs to treat serious or life-threatening diseases or conditions and fill unmet medical needs. Priority review is designed to give drugs that offer major advances in treatment or provide a treatment where no adequate therapy exists an initial review within six months as compared to a standard review time of ten months. Although Fast Track and priority review do not affect the standards for approval, the FDA will attempt to facilitate early and frequent meetings with a sponsor of a Fast Track designated drug and expedite review of the application for a drug designated for priority review. Accelerated approval provides for an earlier approval for a new drug that is intended to treat a serious or life-threatening disease or condition and that fills an unmet medical need based on a surrogate endpoint. A surrogate endpoint is a laboratory measurement or physical sign used as an indirect or substitute measurement representing a clinically meaningful outcome. As a condition of approval, the FDA may require that a sponsor of a drug candidate receiving accelerated approval perform post-marketing clinical trials to confirm the clinically meaning full outcome as predicted by the surrogate marker trial.

In the Food and Drug Administration Safety and Innovation Act, or FDASIA, which was signed into law in July 2012, Congress encouraged the FDA to utilize innovative and flexible approaches to the assessment of products under accelerated approval. The law required the FDA to issue related draft guidance within a year after the law's enactment and also promulgate confirming regulatory changes. In June 2013, the FDA published a draft Guidance for Industry entitled, "Expedited Programs for Serious Conditions Drugs and Biologics" which provides guidance on FDA programs that are intended to facilitate and expedite development and review of new drugs as well as threshold criteria generally applicable to concluding that a drug is a candidate for these expedited development and review programs. In addition to the Fast Track, accelerated approval and priority review programs discussed above, the FDA also provided guidance on a new program for Breakthrough Therapy designation. A request for Breakthrough Therapy designation should be submitted concurrently with, or



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as an amendment to an IND. FDA has already granted this designation to over 30 new drugs and recently approved a couple of Breakthrough Therapy designated drugs.

***Pediatric Exclusivity and Pediatric Use***

Under the Best Pharmaceuticals for Children Act, or BPCA, certain drugs may obtain an additional six months of exclusivity, if the sponsor submits information requested in writing by the FDA, or a Written Request, relating to the use of the active moiety of the drug in children. The FDA may not issue a Written Request for studies on unapproved or approved indications or where it determines that information relating to the use of a drug in a pediatric population, or part of the pediatric population, may not produce health benefits in that population.

We have not received a Written Request for such pediatric studies, although we may ask the FDA to issue a Written Request for such studies in the future. To receive the six-month pediatric market exclusivity, we would have to receive a Written Request from the FDA, conduct the requested studies in accordance with a written agreement with the FDA or, if there is no written agreement, in accordance with commonly accepted scientific principles, and submit reports of the studies. A Written Request may include studies for indications that are not currently in the labeling if the FDA determines that such information will benefit the public health. The FDA will accept the reports upon its determination that the studies were conducted in accordance with and are responsive to the original Written Request or commonly accepted scientific principles, as appropriate, and that the reports comply with the FDA's filing requirements.

In addition, the Pediatric Research Equity Act, or PREA, requires a sponsor to conduct pediatric studies for most drugs and biologicals, for a new active ingredient, new indication, new dosage form, new dosing regimen or new route of administration. Under PREA, original NDAs, BLAs and supplements thereto must contain a pediatric assessment unless the sponsor has received a deferral or waiver. The required assessment must include the evaluation of the safety and effectiveness of the product for the claimed indications in all relevant pediatric subpopulations and support dosing and administration for each pediatric subpopulation for which the product is safe and effective. The sponsor or FDA may request a deferral of pediatric studies for some or all of the pediatric subpopulations. A deferral may be granted for several reasons, including a finding that the drug or biologic is ready for approval for use in adults before pediatric studies are complete or that additional safety or effectiveness data needs to be collected before the pediatric studies begin. After April 2013, the FDA must send a non-compliance letter to any sponsor that fails to submit the required assessment, keep a deferral current or fails to submit a request for approval of a pediatric formulation.

As part of the FDASIA, Congress made a few revisions to BPCA and PREA, which were slated to expire on September 30, 2012, and made both laws permanent.

***Reimbursement***

In both domestic and foreign markets, sales and reimbursement of any approved products will depend, in part, on the extent to which the costs of such products will be covered by third-party payors, such as government health programs, commercial insurance and managed healthcare organizations. These third-party payors are increasingly challenging the prices charged for medical products and services and imposing controls to manage costs. The containment of healthcare costs has become a priority of federal and state governments and the prices of drugs have been a focus in this effort. Governments have shown significant interest in implementing cost-containment programs, including price controls, restrictions on reimbursement and requirements for substitution of generic products. Adoption of price controls and cost-containment measures, and adoption of more restrictive policies in jurisdictions with existing controls and measures, could further limit our net revenue and results. In addition, there is significant uncertainty regarding the reimbursement status of newly approved

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healthcare products. We may need to conduct expensive pharmacoeconomic studies in order to demonstrate the cost-effectiveness of our products. If third-party payors do not consider our products to be cost-effective compared to other therapies, the payors may not cover our products after approved as a benefit under their plans or, if they do, the level of payment may not be sufficient to allow us to sell our products on a profitable basis.

Within the United States, if we obtain appropriate approval in the future to market any of our current protein therapeutic candidates, we may seek approval and coverage for those products under Medicaid, Medicare and the Public Health Service (PHS) pharmaceutical pricing program and also seek to sell the products to federal agencies.

Medicaid is a joint federal and state program that is administered by the states for low income and disabled beneficiaries. Under the Medicaid Drug Rebate Program, manufacturers are required to pay a rebate for each unit of product reimbursed by the state Medicaid programs. The amount of the rebate for each product is set by law and may be subject to an additional discount if certain pricing increases more than inflation.

Medicare is a federal program administered by the federal government that covers individuals age 65 and over as well as those with certain disabilities. Medicare Part D provides coverage to enrolled Medicare patients for self-administered drugs (i.e., drugs that do not need to be administered by a physician). Medicare Part D is administered by private prescription drug plans approved by the U.S. government and each drug plan establishes its own Medicare Part D formulary for prescription drug coverage and pricing, which the drug plan may modify from time-to-time.

Medicare Part B covers most injectable drugs given in an in-patient setting, and some drugs administered by a licensed medical provider in hospital outpatient departments and doctors offices. Medicare Part B is administered by Medicare Administrative Contractors, which generally have the responsibility of making coverage decisions. Subject to certain payment adjustments and limits, Medicare generally pays for Part B covered drugs based on a percentage of manufacturer-reported average sales price.

Drug products are subject to discounted pricing when purchased by federal agencies via the Federal Supply Schedule (FSS). FFS participation is required for a drug product to be covered and paid for by certain federal agencies and for coverage under Medicaid, Medicare Part B and the PHS pharmaceutical pricing program. FSS pricing is negotiated periodically with the Department of Veterans Affairs. FSS pricing is intended to not exceed the price that a manufacturer charges its most-favored non-federal customer for its product. In addition, prices for drugs purchased by the Veterans Administration, Department of Defense (including drugs purchased by military personnel and dependents through the TRICARE retail pharmacy program), Coast Guard, and PHS are subject to a cap on pricing (known as the "federal ceiling price") and may be subject to an additional discount if pricing increases more than inflation.

To maintain coverage of drugs under the Medicaid Drug Rebate Program, manufacturers are required to extend discounts to certain purchasers under the PHS pharmaceutical pricing program. Purchasers eligible for discounts include hospitals that serve a disproportionate share of financially needy patients, community health clinics and other entities that receive health services grants from the PHS.

The American Recovery and Reinvestment Act of 2009 provides funding for the federal government to compare the effectiveness of different treatments for the same illness. A plan for the research will be developed by the Department of Health and Human Services, the Agency for Healthcare Research and Quality and the National Institutes for Health, and periodic reports on the status of the research and related expenditures will be made to Congress. Although the results of the comparative effectiveness studies are not intended to mandate coverage policies for public or private

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payors, it is not clear what effect, if any, the research will have on the sales of any product, if any such product or the condition that it is intended to treat is the subject of a study. It is also possible that comparative effectiveness research demonstrating benefits in a competitor's product could adversely affect the sales of any of our approved protein therapeutics. If third-party payors do not consider our products to be cost-effective compared to other available therapies, they may not cover our products as a benefit under their plans or, if they do, the level of payment may not be sufficient to allow us to sell our products on a profitable basis.

The United States and state governments continue to propose and pass legislation designed to reduce the cost of healthcare. In March 2010, the United States Congress enacted the Patient Protection and Affordable Care Act and the Health Care and Education Reconciliation Act which includes changes to the coverage and payment for drug products under government health care programs. Adoption of other new legislation at the federal or state level could further limit reimbursement for pharmaceuticals.

Outside the United States, ensuring adequate coverage and payment for our products will face challenges. Pricing of prescription pharmaceuticals is subject to governmental control in many countries. Pricing negotiations with governmental authorities can extend well beyond the receipt of regulatory marketing approval for a product and may require us to conduct a clinical trial that compares the cost effectiveness of our protein therapeutic candidates or products to other available therapies. The conduct of such a clinical trial could be expensive and result in delays in our commercialization efforts. Third-party payors are challenging the prices charged for medical products and services, and many third-party payors limit reimbursement for newly-approved health care products. Recent budgetary pressures in many European Union countries are also causing governments to consider or implement various cost-containment measures, such as price freezes, increased price cuts and rebates. If budget pressures continue, governments may implement additional cost-containment measures. Cost-control initiatives could decrease the price we might establish for products that we may develop or sell, which would result in lower product revenues or royalties payable to us. There can be no assurance that any country that has price controls or reimbursement limitations for pharmaceutical products will allow favorable reimbursement and pricing arrangements for any of our products.

***Foreign Regulation***

In addition to regulations in the United States, we will be subject to a variety of foreign regulations governing clinical trials and commercial sales and distribution of our protein therapeutic candidates. Whether or not we obtain FDA approval for a protein therapeutic candidate, we must obtain approval from the comparable regulatory authorities of foreign countries or economic areas, such as the European Union, before we may commence clinical trials or market products in those countries or areas. The approval process and requirements governing the conduct of clinical trials, product licensing, pricing and reimbursement vary greatly from place to place, and the time may be longer or shorter than that required for FDA approval.

Certain countries outside of the United States have a process that requires the submission of a clinical trial application much like an IND prior to the commencement of human clinical trials. In Europe, for example, a clinical trial application, or CTA, must be submitted to the competent national health authority and to independent ethics committees in each country in which a company intends to conduct clinical trials. Once the CTA is approved in accordance with a country's requirements, clinical trial development may proceed in that country. In all cases, the clinical trials must be conducted in accordance with good clinical practices, or GCPs and other applicable regulatory requirements.

Under European Union regulatory systems, a company may submit marketing authorization applications either under a centralized or decentralized procedure. The centralized procedure is compulsory for medicinal products produced by biotechnology or those medicinal products containing

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new active substances for specific indications such as the treatment of AIDS, cancer, neurodegenerative disorders, diabetes, viral diseases and designated orphan medicines, and optional for other medicines which are highly innovative. Under the centralized procedure, a marketing application is submitted to the European Medicines Agency where it will be evaluated by the Committee for Medicinal Products for Human Use and a favorable opinion typically results in the grant by the European Commission of a single marketing authorization that is valid for all European Union member states within 67 days of receipt of the opinion. The initial marketing authorization is valid for five years, but once renewed is usually valid for an unlimited period. The decentralized procedure provides for approval by one or more "concerned" member states based on an assessment of an application performed by one member state, known as the "reference" member state. Under the decentralized approval procedure, an applicant submits an application, or dossier, and related materials to the reference member state and concerned member states. The reference member state prepares a draft assessment and drafts of the related materials within 120 days after receipt of a valid application. Within 90 days of receiving the reference member state's assessment report, each concerned member state must decide whether to approve the assessment report and related materials. If a member state does not recognize the marketing authorization, the disputed points are eventually referred to the European Commission, whose decision is binding on all member states.

As in the United States, we may apply for designation of a product as an orphan drug for the treatment of a specific indication in the European Union before the application for marketing authorization is made. Orphan drugs in Europe enjoy economic and marketing benefits, including up to 10 years of market exclusivity for the approved indication unless another applicant can show that its product is safer, more effective or otherwise clinically superior to the orphan designated product.

***Additional Regulation***

We are also subject to regulation under the Occupational Safety and Health Act, the Environmental Protection Act, the Toxic Substances Control Act, the Resource Conservation and Recovery Act and other present and potential federal, state or local regulations. These and other laws govern our use, handling and disposal of various biological and chemical substances used in, and waste generated by our operations. Our research and development involves the controlled use of hazardous materials, chemicals and viruses. Although we believe that our safety procedures for handling and disposing of such materials comply with the standards prescribed by state and federal regulations, the risk of accidental contamination or injury from these materials cannot be completely eliminated. In the event of such an accident, we could be held liable for any damages that result and any such liability could exceed our resources.

There have been a number of federal and state proposals during the last few years regarding the pricing of pharmaceutical and biological products, government control and other changes to the healthcare system of the U.S. It is uncertain what legislative proposals will be adopted or what actions federal, state or private payers for medical goods and services may take in response to any healthcare reform proposals or legislation. We cannot predict the effect medical or healthcare reforms may have on our business, and no assurance can be given that any such reforms will not have a material adverse effect.

**Manufacturing**

We currently manufacture drug substance for our preclinical studies, Phase 1 clinical trials and Phase 2 clinical trials of ACE-536 and dalantercept. We manufacture material compliant to U.S. and European cGMP at our 12,000 square foot multi-product facility located at our corporate headquarters in Cambridge, Massachusetts. We have the capabilities to manufacture receptor fusion proteins, monoclonal antibodies, and other protein therapeutics.

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Our manufacturing facility is based on single use, disposable technology to maximize the focus of personnel and other resources on the production process, minimizing the need for cleaning and sterilization while optimizing the efficiency of product change-over. The facility consists of four independent clean rooms totaling 4,000 square feet. The facility includes one 250 liter and one 1,000 liter single use bioreactor and has space for two additional 1,000 liter bioreactors.

Approximately 20 fulltime employees focus on our process development and manufacturing activities. We believe that our strategic investment in manufacturing capabilities allows us to advance our protein therapeutic candidates at a more rapid pace and provides us with more portfolio flexibility than if we used a contract manufacturer. The facility produces drug substance in a cost-effective manner while allowing us to retain control over the process and provides an ability to balance the requirements of multiple programs and avoid costly commitments of funds before clinical data are available.

Our manufacturing capabilities encompass the full manufacturing process through quality control and quality assurance. These groups are integrated with our project teams from discovery through development. This structure enables us to efficiently transfer research stage lead molecules into manufacturing. We have designed our manufacturing facility and processes to provide maximum flexibility and rapid change over for the manufacture of different protein therapeutic candidates. We outsource fill-finish, packaging, labeling, shipping, and distribution.

We manufacture our protein therapeutic candidates using readily available raw materials and well established manufacturing procedures based on a standardized process modified for each of our protein therapeutic candidates. We produce our proteins in bioreactors using Chinese hamster ovary cells that have been genetically engineered to produce our specific protein therapeutic candidates. We then purify the proteins using industry standard methods, which include affinity chromatography and ultrafiltration operations. Processes developed within our facility have been successfully transferred to commercial facilities based on stainless steel bioreactors. We have conducted comparability characterization on sotatercept between our Phase 2 material and material made at a commercial manufacturer and found them to be comparable.

We believe that we can scale our manufacturing processes to support our clinical development programs and the potential commercialization of our protein therapeutic candidates. For our early phase protein therapeutic candidates, we intend to continue to manufacture drug substance for preclinical testing and Phase 1 and Phase 2 clinical development at our current facilities. As ACE-536 progresses to Phase 3 clinical trials, we intend to transfer the process for Phase 3 production to Celgene, under the terms of our collaboration agreements. We have already successfully transferred the manufacturing process for sotatercept to Celgene, and we expect Celgene will use a contract manufacturer for Phase 3 and commercial supply of sotatercept and ACE-536. We intend to contract with a third party manufacturer for the supply of dalantercept for Phase 3 clinical trials.

**Employees**

As of September 30, 2013, we had 78 full-time employees, 62 of whom are involved in research, development or manufacturing, and 20 of whom have Ph.D. or M.D. degrees. We have no collective bargaining agreements with our employees and we have not experienced any work stoppages. We consider our relations with our employees to be good.

**Facilities**

Our corporate, research and development, manufacturing, and clinical trial operations are located in Cambridge, Massachusetts. We lease approximately 94,500 square feet of office and laboratory space in three adjacent buildings with aggregate monthly net-rent expense of approximately \$0.4 million. We have sublet approximately 20,000 square feet of space in one of our leased buildings. Two leases expire

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in September 2018 and one lease expires in May 2015. We believe our facilities are adequate for our current needs and that suitable additional substitute space would be available if needed.

**Legal Proceedings**

On October 18, 2012, the Salk Institute for Biological Studies, which we refer to as Salk, filed a complaint in the Massachusetts Superior Court for Suffolk County, alleging that we breached one of our two licensing agreements with Salk. The licensing agreement in dispute provides us with a license with respect to certain of Salk's U.S. patents related to the ActRIIB activin receptor proteins. Salk contends that, under the licensing agreement, we owed Salk a greater share of the upfront payment that we received under our now-terminated agreement with Shire AG regarding ACE-031 and a share of the upfront payment and development milestone payments that we have received under our ongoing collaboration agreement with Celgene regarding ACE-536. Salk is seeking a total of approximately \$10.5 million plus interest in payment and a 15% share of future development milestone payments received under our agreement with Celgene regarding ACE-536. We contend that no additional amounts are due to Salk and that we have complied with all of our payment obligations under the applicable Salk license agreement.

We moved to dismiss the complaint on December 3, 2012. The Court denied our motion on February 28, 2013. On March 14, 2013, Acceleron answered the complaint and asserted patent invalidity counterclaims. On the basis of those counterclaims, Acceleron removed the action on March 28, 2013 to the United States District Court for the District of Massachusetts. The parties have since reached an agreement on a stipulation as to certain patent issues raised in the action, and Acceleron has dismissed its counterclaims. The Court held an initial scheduling conference on May 30, 2013, and the parties are in the process of fact discovery. The case is currently scheduled for trial in September 2014. We intend to defend our position vigorously.

Table of Contents**MANAGEMENT****Executive Officers, Significant Employees and Directors**

Below is a list of the names, ages as of January 1, 2014 and positions, and a brief account of the business experience of the individuals who serve as our executive officers and directors as of the date of this prospectus. Our certificate of incorporation will provide that our board of directors will be divided into three class of directors, with the classes as nearly equal in number as possible. Each of our directors identified below serves in the class indicated. Subject to any earlier resignation or removal in accordance with the terms of our restated certificate of incorporation and by-laws, our Class I directors will serve until the 2014 annual meeting of stockholders; our Class II directors will serve until the 2015 annual meeting of stockholders; and our Class III directors will serve until the 2016 annual meeting of stockholders.

<b>Name</b>	<b>Age</b>	<b>Position</b>
John L. Knopf, Ph.D.	61	Chief Executive Officer and President; Director (Class II)
Kevin F. McLaughlin	57	Senior Vice President, Chief Financial Officer and Treasurer
Matthew L. Sherman, M.D.	58	Senior Vice President and Chief Medical Officer
Steven D. Ertel	44	Senior Vice President and Chief Business Officer
Ravindra Kumar, Ph.D.	53	Vice President and Chief Scientific Officer
John D. Quisel, J.D., Ph.D.	42	Vice President, General Counsel and Secretary
Anthony B. Evin, Ph.D.	72	Director (Class II)
Jean M. George	55	Director (Class I)
George Golumbeski, Ph.D.	56	Director (Class I)
Edwin M. Kania, Jr.	56	Director (Class I)
Tom Maniatis, Ph.D.	70	Director (Class III)
Terrance G. McGuire	57	Director (Class II)
Richard F. Pops	51	Director (Class III)
Joseph S. Zakrzewski	51	Director (Class III)

**John L. Knopf, Ph.D.** co-founded Acceleron in 2003 and is our Chief Executive Officer and President. Dr. Knopf served on our board of directors from 2003 to 2004, and has served from 2007 to the present. Prior to founding Acceleron, Dr. Knopf served as Site Head of the Wyeth Research facilities in Cambridge, MA and Vice President of Metabolic and Respiratory Disease. Dr. Knopf was an early key scientist at Genetics Institute (GI) from 1982 to 1998, where he participated in the development of pioneering biopharmaceutical products including the first treatment for hemophilia, recombinant factor VIII *Recombinate*® and helped establish GI as a premier biopharmaceutical company. While at GI, he established a structure-based small molecule discovery group. Dr. Knopf is the author of several key scientific manuscripts in the area of signal transduction, and is named as an inventor of several patents. Dr. Knopf received a BS in biology from SUNY Stonybrook and his Ph.D. in biology at SUNY Buffalo. We believe Dr. Knopf's extensive experience and knowledge of biopharmaceuticals and our company qualifies him to serve as a member of our board of directors.

**Kevin F. McLaughlin** joined Acceleron in November 2010 and is our Senior Vice President, Chief Financial Officer and Treasurer. He most recently served, from 2009 through 2010, as Senior Vice President and Chief Financial Officer of Qteros, Inc. He was a co-founder of Aptius Education, Inc. and from 2007 through 2009 he worked as the Chief Operating Officer and a director. From 1996 through 2007, Mr. McLaughlin held several executive positions with PRAECIS Pharmaceuticals, Inc. He joined PRAECIS as their first Chief Financial Officer where he had responsibility for private financings, partnership financings, the company's initial public offering and subsequent stock offering. Later, Mr. McLaughlin became COO, and then President and CEO, and he served as a member of the board of directors. In this capacity he was responsible for negotiating the sale of the company to GlaxoSmithKline. He began his career in senior financial roles at Prime Computer and Computervision

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Corporation. Mr. McLaughlin received a BS in business from Northeastern University and an MBA from Babson College.

**Matthew L. Sherman, M.D.** joined Acceleron in May 2006 and is our Senior Vice President and Chief Medical Officer. Previously, he served as Senior Vice President and Chief Medical Officer at Synta Pharmaceuticals where he was responsible for clinical research, clinical operations, biostatistics, data management, regulatory affairs, quality assurance and program management. Prior to that, Dr. Sherman worked at Genetics Institute and Wyeth Pharmaceuticals in various capacities including Therapeutic Area Director for Oncology. While at Wyeth, Dr. Sherman provided senior oncology and hematology leadership for worldwide clinical development for both small molecule and biologic therapeutics, including the submission and approval of Mylotarg® by the FDA. He has published numerous papers and book chapters in the field of oncology and clinical development and is named as an inventor of several patents. Dr. Sherman is board certified in Medical Oncology and Internal Medicine and held various clinical positions at Harvard Medical School with corresponding hospital appointments at the Dana-Farber Cancer Institute and Brigham and Women's Hospital. Dr. Sherman received an SB in chemistry from the Massachusetts Institute of Technology and an MD from Dartmouth Medical School.

**Steven D. Ertel** joined Acceleron in January 2006 and is our Senior Vice President and Chief Business Officer. Mr. Ertel established our business development function and currently leads our business development, commercial strategy and program management functions. Mr. Ertel has over 20 years of experience in the biotechnology industry at Vivus, Inc., Genentech, Inc., Biogen Idec, Inc., and Synta. His responsibilities at these companies included program management for preclinical and clinical stage programs, commercial strategy for clinical stage programs, market launch of a novel biologic agent, and business development. Mr. Ertel began his career in the venture capital industry at Oxford Bioscience Partners. Mr. Ertel received a BSE in biomedical engineering from Duke University and an MBA from the Wharton School at the University of Pennsylvania.

**Ravindra Kumar, Ph.D.** joined Acceleron in March 2004 and is currently our Vice President and Chief Scientific Officer. Dr. Kumar established and currently leads our discovery research. Previously, Dr. Kumar worked for 12 years at Genetics Institute and Wyeth Pharmaceuticals. At Genetics Institute, Dr. Kumar was a key member of the Small Molecule Drug Discovery group and was responsible for cell biology. Following the integration of discovery functions from GI and Wyeth Pharmaceuticals, Dr. Kumar served as Senior Scientist in the Biological Chemistry group. Dr. Kumar is the author of several key scientific manuscripts in the area of protein glycosylation and is named as an inventor of several patents. Dr. Kumar received his BS in chemistry from Rohilkhand University, his MS in chemistry from Meerut University, his Ph.D. from University of New Brunswick and he completed his post-doctoral fellowship at Albert Einstein College of Medicine, in Bronx, NY.

**John D. Quisel, J.D., Ph.D.** joined Acceleron in October 2006 and is our Vice President and General Counsel and Secretary. Prior to joining us, Dr. Quisel worked at the Boston office of Ropes & Gray LLP and, prior to that, the Boston office of Foley Hoag LLP. In his work at law firms, Dr. Quisel has, through strategic in-licensing and protection of internal research programs, assembled and licensed product and platform focused intellectual property portfolios for numerous biotechnology ventures. Over his entire career, Dr. Quisel's experience spans many aspects of biotechnology law, including the negotiation of intellectual property licenses and product development collaborations, patent prosecution and litigation. Dr. Quisel received an AB in biology from Harvard University, an MS in biology from Stanford University, a Ph.D. in biology from the Massachusetts Institute of Technology and a J.D. from Harvard Law School.

**Anthony B. Evnin, Ph.D.** has served as a member of our board of directors since 2004. He has been a Partner at Venrock, a venture capital firm, where he focuses largely on life sciences investments and, in particular, biotechnology investments since 1975. Before this, he served as a manager of



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business development at Story Chemical Corporation and a research scientist at Union Carbide Corporation. Dr. Evnin currently serves on the boards of AVEO Pharmaceuticals, Infinity Pharmaceuticals, Inc., Constellation Pharmaceuticals, Inc., and Juno Therapeutics, Inc. During the last five years, Dr. Evnin served as a director of Altea Therapeutics Corporation, Boston-Power, Inc., Celladon Corporation, CymaBay Therapeutics, Inc., Icagen, Inc., Memory Pharmaceuticals Corp., Pharms Corporation, and Sunesis Pharmaceuticals, Inc. Dr. Evnin is a Trustee of The Rockefeller University and of The Jackson Laboratory, Trustee Emeritus of Princeton University, a Member of the Boards of Overseers and Managers of Memorial Sloan-Kettering Cancer Center, a Director of the New York Genome Center, and a member of the Board of Directors of the Albert and Mary Lasker Foundation. Dr. Evnin received an AB in chemistry from Princeton University and a Ph.D. in chemistry from the Massachusetts Institute of Technology. We believe Dr. Evnin's substantial experience as an investor in, and director of, early stage biopharmaceutical companies, as well as his expertise in corporate strategy at a publicly traded biopharmaceutical company qualify him to serve as a member of our board of directors.

**Jean M. George** has served as a member of our board of directors since 2005. Since 2002, Ms. George has been a Managing Director at Advanced Technology Ventures (ATV), and, concurrently since April 2013, Ms. George has been a Managing Director at LSV Capital Management. She joined ATV in 2002 and serves as the firm's East Coast lead partner for healthcare investments. Prior to joining ATV, Ms. George was a Director at BancBoston Ventures, where she led the health care team's investment activity in NuGenesis Technologies Corp., Microbia, Inc., Syntonix Pharmaceuticals, Inc. and Neurometrix, Inc. Before BancBoston Ventures, she worked at Genzyme Corporation from 1988 to 1998, where she held a variety of operational roles in marketing, product development, and business development, including Vice President of Global Sales and Marketing. She also worked as a Vice President and Founder of Genzyme's Tissue Repair Division. She is currently a Director of Calithera Biosciences, Hydra Biosciences, Inc., Zeltiq Aesthetics, Inc., Portola Pharmaceuticals, Inc., Thrasos Therapeutics and Catabasis Pharmaceuticals, Inc. Ms. George was a Director of Hypnion, Inc., Critical Therapeutics, Inc. and Proteolix, Inc. She was named a member of the Scientific Advisory Board for the Massachusetts Life Sciences Center. Ms. George received a BS in biology from the University of Maine and an MBA from Simmons College Graduate School of Management. We believe that Ms. George's executive experience in the life sciences and therapeutic device industries qualifies her to serve as a member of our board of directors.

**George Golumbeski, Ph.D.** has served as a member of our board of directors since 2011. Since 2009, Dr. Golumbeski has been a Senior Vice President of Business Development for Celgene Corporation, where he is responsible for the full array of business development activities, including identification and evaluation of opportunities, structuring and negotiating transactions, in-licensing, M&A, out-licensing, and alliance management. At Celgene, these activities are focused primarily within the therapeutic areas of oncology and inflammation. From 2008 to 2009, Dr. Golumbeski served as the CEO of Nabriva Therapeutics AG. Prior to Nabriva, Dr. Golumbeski served as Vice President of Business Development, Licensing and Strategy for Novartis-Oncology. During his tenure at Novartis, Dr. Golumbeski's group closed a significant number of collaboration agreements which bolstered the development pipeline. Earlier in his career, Dr. Golumbeski held senior positions at Elan Pharmaceuticals and at Schwarz Pharma, where he led the effort to in-license rotigotine and lacosamide (now both approved agents). Dr. Golumbeski received a BA in biology from the University of Virginia and a Ph.D. in genetics from the University of Wisconsin-Madison. We believe that Dr. Golumbeski's experience as an officer of other pharmaceutical companies, as well as Dr. Golumbeski's extensive experience in research and development and corporate leadership positions, qualify him to serve as a member of our board of directors.

**Edwin M. Kania, Jr.** has served as a member of our board of directors since 2004. Since 2000, Mr. Kania has been a Managing Partner and the Chairman of Flagship Ventures, a Boston-based

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venture capital firm that he co-founded and that also manages the Applied Genomic Technology Capital Fund, L.P. (AGTC Fund) as well as funds raised by OneLiberty Ventures. Prior to co-founding Flagship Ventures, Mr. Kania was a Managing Partner at OneLiberty Ventures and its predecessor firm, Morgan Holland Ventures which he joined in 1985. Mr. Kania currently serves on the boards of several private companies. Mr. Kania has also served on the boards of Aspect Medical, EXACT Sciences and other public and private companies. Mr. Kania's direct investment experience covers over 100 companies, and he has been intimately involved in the launch and development of more than a dozen companies as the founding, lead or co-lead investor, and has on occasion assumed operating roles in support of management. Mr. Kania received a BS in physics from Dartmouth College and an MBA from Harvard Business School. We believe that Mr. Kania's extensive experience investing in, guiding and leading start-up and early phase companies qualifies him to serve as a member of our board of directors.

**Tom Maniatis, Ph.D.** co-founded Acceleron in 2003 and has served as a member of our board of directors and chairman of our Scientific Advisory Board since 2003. Since 2010 he has been a Professor and Chair of the Department of Biochemistry & Molecular Biophysics at the Columbia University College of Physicians and Surgeons. Prior to working at Columbia, Dr. Maniatis was a professor at Harvard University where he studied the mechanisms of transcription and RNA splicing in eukaryotes. Dr. Maniatis currently serves on the board of Constellation Pharmaceuticals, Inc. Dr. Maniatis is also a co-founder of Genetics Institute and ProScript Inc., where he served on the board of directors. Dr. Maniatis is a member of the U.S. National Academy of Sciences, and has received numerous awards for his research contributions, including the Eli Lilly Research Award in Microbiology and Immunology, the Richard Lounsbery Award for Biology and Medicine from the U.S. and French National Academies of Science, and the 2012 Lasker-Koshland Special Achievement Award in Medical Science. Dr. Maniatis received a BA in biology, an MS in chemistry from the University of Colorado at Boulder, and a Ph.D. in molecular biology from Vanderbilt University. We believe Dr. Maniatis's extensive experience and knowledge of biopharmaceuticals and the biopharmaceutical industry qualify him to serve as a member of our board of directors.

**Terrance G. McGuire** has served as a member of our board of directors since 2005. Mr. McGuire co-founded Polaris Partners in 1996 and is currently one of their general partners. Prior to starting Polaris Partners, Mr. McGuire spent seven years at Burr, Egan, Deleage & Co., investing in early stage medical and information technology companies. He currently serves on the board of directors of Adimab/Arsanis, Alector, Quantum Designs, Arsenal Medical/480 Biomedical, Editas, Iora Health, Ironwood Pharmaceuticals, Life Line Screening, MicroCHIPS, NextCode, Pulmatrix, SustainX, and Trevena. He has served on the board of directors of numerous companies, including Remon Medical Technologies, GlycoFi, Akamai Technologies, Aspect Medical Systems, Cubist Pharmaceuticals, Transform Pharma, and deCODE genetics. Mr. McGuire is the former chairman of the National Venture Capital Association, chairman of the board of the Thayer School of Engineering at Dartmouth College, and a member of the boards of The David H. Koch Institute for Integrative Cancer Research at the Massachusetts Institute of Technology and The Arthur Rock Center for Entrepreneurship at Harvard Business School. Mr. McGuire earned a BS in physics and economics from Hobart College, an MS in engineering from The Thayer School at Dartmouth College, and an MBA from Harvard Business School. We believe that Mr. McGuire's extensive experience as a venture capitalist focused on the biotechnology industry, as well as Mr. McGuire's many years of experience helping companies evolve from the start-up phase to successful public companies qualify him to serve as a member of our board of directors.

**Richard F. Pops** has served as a member of our board of directors since 2004. Since 2011, Mr. Pops has served as Chief Executive Officer and Chairman of the board of Alkermes plc, the parent company of Alkermes. From 2009 to 2011, Mr. Pops served as Chief Executive Officer and Chairman of the Board of Alkermes, from 2007 to 2009 he served as the Chairman of the board of Alkermes,

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and from 1991 through 2007 he served as the Chief Executive Officer of Alkermes. Mr. Pops also serves on the board of directors of Neurocrine Biosciences, Inc., Epizyme Inc., the Biotechnology Industry Organization (BIO) and Pharmaceutical Researcher and Manufacturers of America (PhRMA). He has previously served on the board of directors of Sirtris Pharmaceuticals from 2004 to 2008, and CombinatoRx, Inc. from 2001 to 2009. Mr. Pops also served on the board of directors of Reliant Pharmaceuticals, a privately held pharmaceutical company purchased by GlaxoSmithKline in 2007, and on the advisory board of Polaris Venture Partners. He was a member of the Harvard Medical School Board of Fellows from 2002 through June 2012. Mr. Pops received a BA in economics from Stanford University. We believe that Mr. Pops leadership experience and industry knowledge qualify him to serve as a member of our board of directors.

**Joseph S. Zakrzewski** has served as a member of our board of directors since 2011. Since 2010 and through 2013, Mr. Zakrzewski had been Chairman and Chief Executive Officer of Amarin Corporation. From 2007 to 2010, Mr. Zakrzewski served as President and Chief Executive Officer of Xcellerex. From 2005 to 2007, Mr. Zakrzewski served as the Chief Operating Officer of Reliant Pharmaceuticals, overseeing the launch of Omacor®, a drug to treat elevated triglyceride levels. From 1988 to 2004, Mr. Zakrzewski served in a variety of positions at Eli Lilly and Company including as Vice President, Corporate Business Development from 2003 through 2004. In addition, Mr. Zakrzewski served as a Venture Partner with OrbiMed in 2010 and 2011. Mr. Zakrzewski also currently serves on the board of directors of Amarin and Insulet Corporation and has also served on the board of directors of Xcellerex, Azelon/Zelos Therapeutics and Promedior. Mr. Zakrzewski received a BS in Chemical Engineering and an MS in Biochemical Engineering from Drexel University as well as an MBA in Finance from Indiana University. We believe that Mr. Zakrzewski's substantial experience as an executive officer of other pharmaceutical companies, as well as Mr. Zakrzewski's service on boards of directors of other pharmaceutical companies qualify him to serve as a member of our board of directors.

**Board Composition**

Our board of directors is currently comprised of 9 members. Our board of directors has determined that each of Ms. George and Messrs Evnin, Golumbeski, Kania, Maniatis, McGuire, Pops and Zakrzewski is independent for NASDAQ purposes. The members of our board of directors were elected in compliance with the provisions of the voting agreement among us and our major stockholders. The voting agreement terminated upon the closing of our initial public offering on September 24, 2013, and at present we do not have any contractual obligations regarding the election of our directors. See "Certain Relationships and Related Party Transactions". Our directors hold office until their successors have been elected and qualified or until their earlier death, resignation or removal. There are no family relationships among any of our directors or executive officers.

**Board Committees**

Our board of directors has three standing committees: the audit committee, the compensation committee and the nominating and corporate governance committee.

*Audit Committee*

Our audit committee is composed of Anthony B. Evnin, Ph.D., Jean M. George and Joseph S. Zakrzewski, with Dr. Evnin serving as chairman of the committee. Our board of directors has determined that Dr. Evnin, Ms. George, and Mr. Zakrzewski meet the independence requirements of Rule 10A-3 under the Exchange Act and the applicable listing standards of NASDAQ. Our board of directors has determined that Joseph S. Zakrzewski is an "audit committee financial expert" within the

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meaning of the Securities and Exchange Commission, or SEC, regulations and applicable listing standards of NASDAQ. The audit committee's responsibilities include:

appointing, approving the compensation of, and assessing the qualifications, performance and independence of our independent registered public accounting firm;

pre-approving audit and permissible non-audit services, and the terms of such services, to be provided by our independent registered public accounting firm;

reviewing the internal audit plan with the independent registered public accounting firm and members of management responsible for preparing our financial statements;

reviewing and discussing with management and the independent registered public accounting firm our annual and quarterly financial statements and related disclosures as well as critical accounting policies and practices used by us;

reviewing the adequacy of our internal control over financial reporting;

establishing policies and procedures for the receipt and retention of accounting-related complaints and concerns;

recommending, based upon the audit committee's review and discussions with management and the independent registered public accounting firm, whether our audited financial statements shall be included in our Annual Report on Form 10-K;

monitoring our compliance with legal and regulatory requirements as they relate to our financial statements and accounting matters;

preparing the audit committee report required by the rules of the SEC to be included in our annual proxy statement;

viewing all related party transactions for potential conflict of interest situations and approving all such transactions; and

reviewing and discussing with management and our independent registered public accounting firm our earnings releases and scripts.

***Compensation Committee***

Our compensation committee is composed of Edwin M. Kania, Jr., Tom Maniatis, Ph.D. and Joseph S. Zakrzewski, with Mr. Kania serving as chairman of the committee. Our board of directors has determined that each member of the compensation committee is "independent" as defined under the applicable listing standards of NASDAQ. The compensation committee's responsibilities include:

annually reviewing and approving corporate goals and objectives relevant to the compensation of our chief executive officer;

evaluating the performance of our chief executive officer in light of such corporate goals and objectives and determining and approving the compensation of our chief executive officer;

reviewing and approving the compensation of our other executive officers;

appointing, compensating and overseeing the work of any compensation consultant, legal counsel or other advisor retained by the compensation committee;

conducting the independence assessment outlined in NASDAQ rules with respect to any compensation consultant, legal counsel or other advisor retained by the compensation committee;

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annually reviewing and reassessing the adequacy of the committee charter in its compliance with the listing requirements of NASDAQ;

reviewing and establishing our overall management compensation, philosophy and policy;

overseeing and administering our equity compensation and other compensatory plans;

reviewing and approving our equity and incentive policies and procedures for the grant of equity-based awards and approving the grant of such equity-based awards;

reviewing and making recommendations to the board of directors with respect to director compensation; and

reviewing and discussing with management the compensation discussion and analysis to be included in our annual proxy statement or Annual Report on Form 10-K.

***Nominating and Corporate Governance Committee***

Our nominating and corporate governance committee is composed of Anthony B. Evnin, Ph.D., Jean M. George and Terrance G. McGuire, with Ms. George serving as chairman of the committee. Our board of directors has determined that each member of the nominating and corporate governance committee is "independent" as defined under the applicable listing standards of NASDAQ. The nominating and corporate governance committee's responsibilities include:

developing and recommending to the board of directors criteria for board and committee membership;

establishing procedures for identifying and evaluating board of director candidates, including nominees recommended by stockholders;

identifying individuals qualified to become members of the board of directors;

recommending to the board of directors the persons to be nominated for election as directors and to each of the board's committees;

developing and recommending to the board of directors a set of corporate governance principles;

articulating to each director what is expected, including reference to the corporate governance principles and directors' duties and responsibilities;

reviewing and recommending to the board of directors practices and policies with respect to directors;

reviewing and recommending to the board of directors the functions, duties and compositions of the committees of the board of directors;

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reviewing and assessing the adequacy of the committee charter and submitting any changes to the board of directors for approval;

consider and report to the board of directors any questions of possible conflicts of interest of board of directors members;

provide for new director orientation and continuing education for existing directors on a periodic basis;

performing an evaluation of the performance of the committee; and

overseeing the evaluation of the board of directors and management.

Our board of directors may establish other committees from time to time.

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***Compensation Committee Interlocks and Insider Participation***

None of the members of our compensation committee has at any time during the prior three years been one of our officers or employees. None of our executive officers currently serves, or in the past fiscal year has served, as a member of the board of directors or compensation committee of any entity that has one or more executive officers serving on our board of directors or compensation committee. For a description of transactions between us and members of our compensation committee and affiliates of such members, please see "Certain Relationships and Related Party Transactions."

***Code of Business Conduct and Ethics***

We have adopted a code of business conduct and ethics that applies to all of our employees, officers and directors, including those officers responsible for financial reporting. Our code of business conduct and ethics is available on our website. We intend to disclose any amendments to the code, or any waivers of its requirements, on our website.



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**EXECUTIVE OFFICER AND DIRECTOR COMPENSATION**

**Overview**

The following discussion relates to the compensation of our founder, Chief Executive Officer and President, John L. Knopf, Ph.D., and our two most highly compensated executive officers (other than our chief executive officer), Matthew L. Sherman, M.D., our Chief Medical Officer and Senior Vice President, and John D. Quisel, J.D., Ph.D., our General Counsel, Vice President and Secretary, who are collectively referred to in this prospectus as our named executive officers. Each year, our compensation committee reviews and determines the compensation of our executive officers, including our named executive officers. Our executive compensation program is designed to attract and retain a highly skilled team of key executives and to align the compensation of our executives with the interests of our shareholders by rewarding the achievement of short- and long-term strategic financial goals, which we believe serves to enhance short- and long-term value creation for our shareholders.

**Elements of Executive Compensation**

The compensation of our named executive officers consists of base salary, annual cash bonuses, equity awards and employee benefits that are made available to all salaried employees. Our named executive officers are also entitled to certain compensation and benefits upon certain terminations of employment and certain change in control transactions pursuant to employment agreements.

**Base Salaries.** Base salaries for our named executive officers are determined based on the scope of each officer's responsibilities along with his respective experience and contributions to the company. Base salaries for our named executive officers are determined annually by our compensation committee, subject to review by our board of directors. When reviewing base salaries for increase, our compensation committee takes factors into account such as each officer's experience and individual performance, the company's performance as a whole, data from surveys of compensation paid by comparable companies, and general industry conditions, but does not assign any specific weighting to any factor. For fiscal 2013, our board of directors approved a base salary of \$380,662 for Dr. Sherman and \$319,300 for Dr. Quisel, representing an increase of 4.5% and 3.0%, respectively, from the base salary for each such executive in 2012. Dr. Knopf's base salary did not increase in 2013.

**Annual Cash Bonuses.** Our annual cash bonus program promotes and rewards the achievement of key strategic and business goals. For fiscal 2013, the target annual bonus as a percentage of base salary for each of Dr. Knopf, Dr. Sherman and Dr. Quisel remained at 50%, 30% and 30%, respectively.

At the beginning of fiscal 2013, our compensation committee established corporate performance goals, each having a designated weighting, that included key strategic and financial goals of the company relating to research and the company's clinical pipeline, process development and manufacturing, business development and the achievement of financial objectives. At the end of the 2013 fiscal year, our compensation committee met and evaluated the performance of the company against the specified performance goals. Based on its evaluation, our compensation committee recommended payment of 2013 annual bonuses at the target level for all employees covered under this program and above the target level for members of our management team, including our named executive officers, after considering the outstanding performance of the company, the achievement of the company performance goals above the target level in fiscal 2013 and the successful completion of important financing activities in fiscal 2013, including the successful completion of our initial public offering. Our board of directors approved the recommendations of our compensation committee and each of Drs. Knopf, Sherman and Quisel received a cash bonus for 2013 equal to \$300,000, \$171,315 and \$143,680, respectively, representing a payment of 75%, 45% and 45%, respectively, of each executive's base salary.

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**Equity Awards.** Our named executive officers participate in our Acceleron Pharma Inc. 2003 Stock Option and Restricted Stock Plan, which we refer to as the "2003 Plan". See " 2003 Plan" below for additional details about the 2003 Plan. Our named executive officers also participate in our Acceleron Pharma Inc. 2013 Equity Incentive Plan, which we refer to as the "2013 Equity Incentive Plan". See " 2013 Equity Incentive Plan" below for additional details about the 2013 Equity Incentive Plan. Grants under the 2003 Plan and the 2013 Equity Incentive Plan, including those made to our named executive officers, generally consist of stock option awards subject to time-based vesting. Awards that are subject to time-based vesting generally vest either in quarterly installments over four years or vest as to 25% of the shares subject to the option after one year and thereafter continue to vest in quarterly installments over the following three years, generally subject to continued employment. During fiscal 2013, each of Drs. Knopf, Sherman and Quisel was awarded a stock option under the 2013 Equity Incentive Plan to purchase 110,000, 29,000 and 29,000 shares of our common stock, respectively, in each case vesting in equal quarterly installments over four years. Stock option awards serve to align the interests of our named executive officers with our shareholders, because no value is created unless the value of our common stock appreciates after grant. They also encourage retention through the use of time-based vesting. Pursuant to agreements with certain members of senior management, including our named executive officers, a portion of the executive's stock option and restricted stock awards will vest automatically as of the date of a change of control and all or a portion of the executive's stock option and restricted stock awards may vest upon certain terminations of employment.

**Benefits.** We provide modest benefits to our named executive officers. These benefits and perquisites, such as participation in our 401(k) plan and basic health and welfare benefit coverage, are available to all of our eligible employees.

**Employment Agreements and Change of Control Agreements.** Drs. Knopf, Sherman and Quisel have entered into employment agreements with us that include severance, change of control, and restrictive covenant provisions. We believe that change of control arrangements provide our executives with security that will likely reduce any reluctance they may have to pursue a change of control transaction that could be in the best interests of our stockholders. We also believe that reasonable severance and change of control benefits are necessary in order to attract and retain high-quality executive officers.

**Summary Compensation Table**

The following table sets forth information about certain compensation awarded or paid to our named executive officers for the 2013 fiscal year.

Name and Principal Position	Year	Salary \$(1)	Option Awards \$(2)	Non-Equity Incentive Plan		Total (\$)
				Compensation \$(3)	All Other Compensation	
<b>John L. Knopf, Ph.D.</b> Chief Executive Officer and President	2013	400,000	1,696,423	300,000	237,461(4)	2,633,884
	2012	400,000	1,355,300	200,000		1,955,300
<b>Matthew L. Sherman, M.D.</b> Chief Medical Officer & Senior Vice President	2013	380,662	447,239	171,315		999,216
	2012	364,270	124,550	109,281		598,101
<b>John D. Quisel, J.D., Ph.D.</b> General Counsel, Vice President and Secretary	2013	319,300	447,239	143,680		910,219
	2012	310,000	158,415	93,000		561,415

(1)

Salaries include amounts contributed by the named executive officer to our 401(k) plan.

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- (2) Amounts shown reflect the grant date fair value of options awarded during each of fiscal 2012 and 2013 determined in accordance with the Financial Accounting Standards Board, Accounting Standards Codification Topic 718, *Compensation Stock Compensation*. These amounts exclude the value of estimated forfeitures. With respect to the performance-based option granted to Dr. Knopf in 2012, the amount included in the table assumes the highest level of performance is achieved. Assumptions used in the calculation of these amounts are included in Note 11 to our financial statements included elsewhere in this prospectus.
- (3) Amounts shown reflect the cash bonus amount paid to the named executive officer for each of fiscal 2012 and 2013 that was earned based on company performance.
- (4) Represents income imputed to Dr. Knopf in connection with the forgiveness of the entire principal amount of a promissory note entered into by him and the Company, plus accrued interest. See " Executive Loan" below.

**Outstanding Equity Awards at Fiscal Year-End**

The following table sets forth information regarding equity awards held by our named executive officers as of December 31, 2013.

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## OPTION AWARDS

Name	Equity Incentive Plan Awards:			Option Exercise Price (\$)(5)	Option Expiration Date(6)
	Number of Securities Underlying Unexercised Options (#) Exercisable	Number of Securities Underlying Unexercised Options (#) Unexercisable	Number of Securities Underlying Unexercised Unearned Options (#)		
<b>John L. Knopf</b>	12,500(1)			0.40	2/2/2015
	12,500(2)			0.40	3/29/2016
	100,000(2)			1.80	1/31/2017
	262,500(2)			5.08	3/27/2018
	46,876(2)	3,124(2)		5.88	2/4/2020
	65,625(2)	21,875(2)		3.88	12/2/2020
	25,000(2)	25,000(2)		5.28	12/16/2021
	42,969(2)	94,531(2)		5.28	11/13/2022
		37,500(3)		5.28	11/13/2022
		37,500(3)		5.28	11/13/2022
	3,125(4)	34,375(4)		5.28	11/13/2022
		110,000(2)		24.11	12/5/2023
<b>Matthew L. Sherman</b>	59,766(1)			0.48	5/31/2016
	32,812(2)			1.80	1/31/2017
	35,000(2)			5.08	3/27/2018
	28,125(2)	9,375(2)		3.88	12/2/2020
	6,250(2)	6,250(2)		5.28	12/16/2021
	6,250(2)	18,750(2)		7.12	12/12/2022
		29,000(2)		24.11	12/5/2023
<b>John D. Quisel</b>	27,500(1)			0.92	11/15/2016
	2,500(2)			1.80	6/12/2017
	7,500(2)			5.08	3/27/2018
	12,500(2)			5.88	12/17/2018
	25,000(2)			5.88	12/2/2019
	46,875(2)	15,625(2)		3.88	12/2/2020
	9,374(2)	9,376(2)		5.28	12/16/2021
	9,375(2)	15,625(2)		5.28	6/7/2022
	3,125(2)	9,375(2)		7.12	12/12/2022
	29,000(2)		24.11	12/5/2023	

- (1) Reflects time-based options to purchase shares of our common stock that vested as to 25% of the shares subject to the option on the first anniversary of the vesting commencement date and thereafter vested in equal quarterly installments over the following three years, subject to the executive's continued employment.
- (2) Reflects time-based options to purchase shares of our common stock that vest in equal quarterly installments over four years generally subject to the executive's continued employment.
- (3) Reflects time- and performance-based options granted to Dr. Knopf to purchase shares of our common stock that vest as to 1/12th of the shares subject to the option on the date that is three months from the date on which the company achieves a specified performance condition related either to a financial goal or clinical study milestone and that continue to vest thereafter on a quarterly basis over three years on each three-month anniversary of the initial vesting date. To the



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extent unvested, the option will fully vest on September 6, 2016, generally subject to Dr. Knopf's continued employment.

- (4) Reflects time- and performance-based options granted to Dr. Knopf to purchase shares of our common stock that vest as to 1/12th of the shares subject to the option on the date that is three months from the date on which the company achieves a specified performance condition related to a financial goal and that continue to vest thereafter on a quarterly basis over three years on each three-month anniversary of the initial vesting date. To the extent unvested, the option will fully vest on September 6, 2016, generally subject to Dr. Knopf's continued employment. On December 5, 2013, our board of directors determined that the performance goal had been met on September 24, 2013. As a result, 1/12<sup>th</sup> of the shares subject to the stock option vested on December 24, 2013, and the remainder of the shares will vest on each of the subsequent eleven three-month anniversaries of such date.
- (5) The exercise price of the stock options was not less than the fair market value of a share of our common stock on the date of grant, as determined by our board of directors based, in part, on an independent third party valuation with respect to the period prior to our initial public offering. Stock options granted in fiscal 2013 subsequent to us becoming a public company were granted with an exercise price equal to the closing price of a share of our common stock on the date the stock option was granted.
- (6) All stock options have a 10-year term measured from the date of grant.

***Retirement Benefits***

We do not maintain any qualified or non-qualified defined benefit plans or supplemental executive retirement plans that cover our named executive officers. We offer a tax-qualified retirement plan, which we refer to as our 401(k) plan, to eligible employees, including our named executive officers. Our 401(k) plan permits eligible employees to defer up to 100% of their annual eligible compensation, subject to certain limitations imposed by the Internal Revenue Service. Employees' elective deferrals are immediately fully vested and non-forfeitable under this plan. We may, but are not required to, make discretionary matching contributions and other employer contributions on behalf of eligible employees under this plan. We made matching contributions on behalf of eligible employees in fiscal year 2013, but made no such contributions on behalf of our named executive officers.

**Employment Agreements**

We have entered into amended and restated employment agreements with each of Drs. Knopf, Sherman and Quisel, each providing for a 2013 base salary of \$400,000, \$380,662 and \$319,300, respectively, in each case subject to discretionary annual increases and a discretionary bonus based on performance goals in accordance with our annual cash bonus program. Each agreement also provides for certain payments and benefits upon a qualifying termination of the executive's employment and a change of control, as described below.

***Change of Control.*** At the time of the consummation of a change of control, 50%, in the case of Dr. Knopf, and 25%, in the case of Drs. Sherman and Quisel, of any unvested stock options then held by the executive that were granted on or prior to the effective date of the executive's amended and restated employment agreement (referred to as the amendment date) will vest.

***Termination of Employment Without Cause or for Good Reason Following a Change of Control.*** If, within one year after the consummation of a change of control, the executive's employment is terminated by us (or our successor) other than for cause or the executive terminates his employment for good reason (as such terms are defined in the executive's employment agreement): (1) we will pay the executive a lump sum payment equal to the product of 1.5 times, in the case of Dr. Knopf, or one

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times, in the case of Drs. Sherman and Quisel, (x) the sum of executive's then-current annual base salary plus 100% of the executive's target bonus for the year in which the termination occurs, (2) 100% of any unvested equity and equity-based awards held by the executive at the time of such termination will fully vest, and (3) if the executive elects under COBRA or any successor law to continue participation in our group health and/or dental plans in which the executive was participating prior to such termination, we will pay or, at our option reimburse the executive for, the full premium cost of that participation for 18 months, in the case of Dr. Knopf, or 12 months, in the case of Drs. Sherman and Quisel, following the date the executive's employment terminates or, if earlier, until the date the executive becomes eligible to enroll in such plans of a new employer. We will also pay the executive any base salary earned but not paid and any vacation time accrued but not used, in each case as of the termination date.

***Termination of Employment Without Cause or for Good Reason.*** If the executive's employment is terminated by us other than for cause or the executive terminates his employment for good reason (as such terms are defined in the executive's employment agreement) under circumstances other than as described in the preceding paragraph: (1) we will continue to pay the executive his base salary for a period of 18 months, in the case of Dr. Knopf, or 12 months, in the case of Drs. Sherman and Quisel, in accordance with our payroll policy, (2) any unvested stock option awards the executive holds at the time of such termination of employment that are subject only to time-based vesting and that were granted on or prior to the amendment date will vest to the extent such stock option awards would have otherwise vested over the 12 months, nine months or six months, in the case of each of Dr. Knopf, Dr. Sherman, or Dr. Quisel, respectively, following such termination of employment, and (3) if the executive elects under COBRA or any successor law to continue participation in our group health and/or dental plans in which the executive was participating prior to such termination, we will pay or, at our option reimburse the executive for, the full premium cost of that participation for 18 months, in the case of Dr. Knopf, or 12 months, in the case of Drs. Sherman and Quisel. We will also pay the executive any base salary earned but not paid and any vacation time accrued but not used, in each case as of the termination date.

***Termination of Employment Due to Death or Disability.*** If the executive's employment terminates due to the executive's death or disability, all unvested stock options then held by the executive that were granted on or prior to the amendment date will vest as of the date of termination. In the event of such a termination of employment due to disability, to the extent we do not maintain a disability plan providing for continuation of the executive's base salary for one year following the date of such termination, during this period we will pay the executive, at the time the executive's base salary would otherwise have been paid, an amount equal to the excess of 100% of the executive's base salary over the disability insurance benefits, if any, actually paid to the executive. We will also pay the executive any base salary earned but not paid and any vacation time accrued but not used, in each case as of the termination date.

***Severance Subject to Release of Claims and Compliance With Restrictive Covenants.*** Our obligation to provide the executive with any severance payments or other benefits under the employment agreement is conditioned on the executive signing an effective release of claims in our favor and the executive's continued full performance of his obligations under the Employee Confidentiality, Non-Compete and Proprietary Information Agreement relating to confidentiality, noncompetition and nonsolicitation.

***Other Termination of Employment.*** If the executive's employment is terminated for any reason other than by us without cause, by the executive for good reason, or due to the executive's death or disability, the executive will only be entitled to receive earned but unpaid base salary and any accrued but not used vacation as of the termination date.

***280G Gross-up.*** In the event that a change in ownership or control of the company under Section 280G of the Internal Revenue Code of 1986, as amended, or the Code, and the regulations

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thereunder, occurs on or before the second anniversary of the amendment date, if any portion of the payments made pursuant to the executive's employment agreement or otherwise constitutes an "excess parachute payment" within the meaning of Section 280G of the Code, we will pay the executive an additional amount that, after the imposition of all taxes with respect to such gross-up payment equals the excise tax with respect to the excess parachute payment. If the change in ownership or control occurs after the second anniversary of the amendment date and any portion of the payments made pursuant to the employment agreement or otherwise constitutes an excess parachute payment, the executive will be entitled to receive an amount of such payments reduced so that no portion of the payments would constitute an excess parachute payment, or the amount otherwise payable to the executive under the employment agreement or otherwise reduced by all applicable taxes, including the excise tax, whichever amount results in the greater amount payable to the executive.

**Executive Loan**

We and Dr. Knopf are parties to a Secured Promissory Note dated January 28, 2008 and amended on November 13, 2012, pursuant to which we made a loan to Dr. Knopf with a principal balance of \$200,000 and an interest rate of 3.11% per annum. The outstanding balance of \$237,461, including principal and accrued and unpaid interest on the note, was forgiven immediately prior to the public filing of the registration statement in connection with our initial public offering.

**2013 Director Compensation**

The following table sets forth information concerning the compensation earned by our directors during 2013. Dr. Knopf receives no additional compensation for his service as a director, and, consequently, is not included in this table. The compensation received by Dr. Knopf as an employee during 2013 is included in the "Summary Compensation Table" above.

Name	Fees Paid in Cash \$(1)	Option Awards \$(2)(3)	Total (\$)
Anthony B. Evnin	13,438	308,441	321,878
Jean George	12,500	308,441	320,941
Edwin M. Kania	11,250	308,441	319,691
Tom Maniatis	25,000	308,441	333,441
Terrance G. McGuire	9,688	308,441	318,128
Richard F. Pops	31,250	308,441	339,691
Joseph S. Zakrzewski	30,625	308,441	339,066

(1) Amounts represent annual cash compensation for services rendered by each member of the board of directors.

(2) As of December 31, 2013, our directors held the following aggregate number of stock options: Dr. Evnin, 20,000; Ms. George, 20,000; Mr. Kania, 20,000; Mr. McGuire, 20,000; Dr. Maniatis, 47,500; Mr. Pops, 70,000; and Mr. Zakrzewski, 38,750. Dr. Golumbeski did not hold any stock options or other stock awards as of December 31, 2013.

(3) Amounts shown reflect the grant date fair value of options awarded during fiscal 2013 determined in accordance with the Financial Accounting Standards Board, Accounting Standards Codification Topic 718, *Compensation Stock Compensation*. These amounts exclude the value of estimated forfeitures. Assumptions used in the calculation of these amounts are included in Note 11 to our financial statements included elsewhere in this prospectus.

Our board of directors has adopted a non-employee director compensation policy that is designed to provide a total compensation package that enables us to attract and retain, on a long-term basis,



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high caliber non-employee directors. Under the policy, all non-employee directors will be paid cash compensation as set forth below:

	<b>Annual Retainer</b>
<b>Board of Directors:</b>	
All non-employee members	\$ 35,000
Additional retainer for Lead Independent Director	\$ 15,000
<b>Audit Committee:</b>	
Chairman	\$ 15,000
Non-Chairman members	\$ 7,500
<b>Compensation Committee:</b>	
Chairman	\$ 10,000
Non-Chairman members	\$ 5,000
<b>Nominating and Corporate Governance Committee:</b>	
Chairman	\$ 7,500
Non-Chairman members	\$ 3,750

Under our non-employee director compensation policy, each person who is initially appointed or elected to the board of directors will be eligible to receive a grant of stock options to purchase 20,000 shares of our common stock under our 2013 Equity Incentive Plan on the date he or she first becomes a non-employee director, which will vest quarterly in equal installments over a three-year period. In addition, each continuing non-employee director will be eligible to receive an annual option grant to purchase 10,000 shares of our common stock, which will vest in full on the first anniversary of the grant date. The options will be granted with an exercise price equal to the fair market value of a share of our common stock on the date of grant. In connection with the completion of our initial public offering in September 2013 and in recognition of new and increased duties of the board, in fiscal year 2013, each board member other than Dr. Golumbeski was granted a stock option to purchase 20,000 shares of our common stock.

**Equity and Incentive Plans***2003 Plan*

The 2003 Plan, which became effective December 17, 2003, provides for the grant of options to purchase shares of our common stock and for the grant or sale of restricted stock to key employees and directors of, and consultants and advisors to, us and our affiliates who, in the opinion of the compensation committee, are in a position to contribute significantly to our success and the success of our affiliates. The summary of the 2003 Plan is not a complete description of all provisions of the 2003 Plan and is qualified in its entirety by reference to the 2003 Plan.

The 2003 Plan is administered by our compensation committee, which has authority to determine eligibility for and grant awards and to determine the terms and conditions of all awards, including the time or times upon which awards vest or become exercisable and remain exercisable.

Following our initial public offering in September 2013, all equity-based awards will be granted under the 2013 Equity Incentive Plan described below.

**Authorized Shares.** Subject to adjustment, the maximum number of shares of our common stock that may be delivered in satisfaction of awards under the 2003 Plan is 4,937,500 shares. Shares of our common stock to be issued under the 2003 Plan may be authorized but unissued shares of our common stock or previously issued shares acquired by the company.

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**Stock Options.** The compensation committee determines the exercise price of each stock option, which will not be less than the fair market value of a share of our common stock on the date of grant. Unless otherwise provided by our compensation committee, a participant's unvested stock options will immediately terminate upon the participant's cessation of employment, and vested stock options will remain outstanding for three months or one year (in the case of death) (or, in each case, until the applicable expiration date, if earlier). If the compensation committee determines that the cessation of a participant's employment resulted for reasons that cast such discredit on the participant as to justify immediate termination of stock options, all stock options then held by the participant will immediately terminate upon such termination of employment, whether or not vested.

**Restricted Stock.** The compensation committee may grant or sell restricted stock to any participant (including, but not limited to, upon the exercise of options to purchase common stock) subject to the conditions and restrictions and for such purchase price, if any, as determined by the compensation committee. A participant will have all the rights of a shareholder with respect to shares of restricted stock granted or sold to the participant under the 2003 Plan. Unless the compensation committee determines otherwise, upon the cessation of a participant's employment for any reason, including death, the company will have the right (but not the obligation) to reacquire any shares of restricted stock outstanding at the participant's original purchase price, if any, for such shares. If there is no purchase price, restricted stock will be forfeited upon such termination of employment.

**Covered Transactions.** Unless an award agreement provides otherwise, in the event of a covered transaction (as defined in the 2003 Plan) in which there is an acquiring or surviving entity, the compensation committee may provide for the assumption or substitution of some or all outstanding awards by the acquiring or surviving entity. If the awards are not so assumed or substituted, and except as otherwise provided in an award agreement, each stock option will vest and become fully exercisable prior to the covered transaction and the stock option will terminate upon consummation of the covered transaction. In the case of restricted stock, the compensation committee may require that any amounts delivered, exchanged or otherwise paid in respect of such stock in connection with the covered transaction be placed in escrow or otherwise made subject to such restrictions as the compensation committee deems appropriate. Under the 2003 Plan, a covered transaction generally includes a consolidation, merger or similar transaction in which the company is not the surviving entity, a sale or transfer of all or substantially all of our assets or a dissolution or liquidation of our company.

**2013 Equity Incentive Plan**

Our board of directors adopted the 2013 Equity Incentive Plan and, following our initial public offering in September 2013, all equity-based awards will be granted under the 2013 Equity Incentive Plan. The following summary describes the material terms of the 2013 Equity Incentive Plan. This summary of the 2013 Equity Incentive Plan is not a complete description of all provisions of the 2013 Equity Incentive Plan and is qualified in its entirety by reference to the 2013 Equity Incentive Plan.

**Plan Administration.** The 2013 Equity Incentive Plan is administered by our compensation committee. Our compensation committee has the authority to, among other things, interpret the 2013 Equity Incentive Plan, determine eligibility for, grant and determine the terms of awards under the 2013 Equity Incentive Plan, and to do all things necessary to carry out the purposes of the 2013 Equity Incentive Plan. Our compensation committee's determinations under the 2013 Equity Incentive Plan are conclusive and binding.

**Authorized Shares.** Subject to adjustment, the maximum number of shares of our common stock that may be delivered in satisfaction of awards under the 2013 Equity Incentive Plan is currently 2,633,945 shares, which includes 155,884 shares of our common stock that were available for grant under the 2003 Plan on the date the 2013 Equity Incentive Plan was adopted and an increase of 1,133,945 shares on January 1, 2014 pursuant to the annual adjustment described below. The number of

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shares of our common stock available for issuance under the 2013 Equity Incentive Plan will be automatically increased annually on each January 1<sup>st</sup>, from January 1, 2014 through January 1, 2023, in an amount equal to the least of

3,150,000 shares;

4.0% of the outstanding shares of our common stock as of the close of business on the immediately preceding December 31<sup>st</sup>; and

such other amount as our board of directors may determine.

On January 1, 2014, the number of shares of our common stock reserved for issuance increased to 2,633,945 shares. As of January 1, 2014, there were 2,089,945 shares of our common stock available for issuance under the 2013 Equity Incentive Plan. Shares of our common stock to be issued under the 2013 Equity Incentive Plan may be authorized but unissued shares of our common stock or previously issued shares acquired by us. Any shares of our common stock underlying awards that are settled in cash or otherwise expire, terminate, or are forfeited prior to the issuance of stock will again be available for issuance under the 2013 Equity Incentive Plan.

**Individual Limits.** The maximum number of shares of our common stock subject to stock options and the maximum number of shares of our common stock subject to stock appreciation rights that may be granted to any person in any calendar year is each 1,000,000 shares. The maximum number of shares of our common stock subject to other awards that may be granted to any person in any calendar year is 500,000 shares.

**Eligibility.** Our compensation committee will select participants from among our key employees, directors, consultants and advisors and of our affiliates who are in a position to contribute significantly to the success of the company and its affiliates. Eligibility for options intended to be incentive stock options, or ISOs, is limited to employees of the company or certain affiliates.

**Types of Awards.** The 2013 Equity Incentive Plan provides for grants of stock options, stock appreciation rights, restricted and unrestricted stock, stock units, performance awards and other awards convertible into or otherwise based on shares of our common stock. Dividend equivalents may also be provided in connection with an award under the 2013 Equity Incentive Plan.

Stock options and stock appreciation rights. The exercise price of an option, and the base price against which a stock appreciation right is to be measured, may not be less than the fair market value (or, in the case of an ISO granted to a ten percent shareholder, 110% of the fair market value) of a share of our common stock on the date of grant. Our compensation committee will determine the time or times at which stock options or stock appreciation rights become exercisable and the terms on which such awards remain exercisable.

Restricted and unrestricted stock. A restricted stock award is an award of common stock subject to forfeiture restrictions, while an unrestricted stock award is not subject to restrictions.

Stock units. A stock unit award is denominated in shares of our common stock and entitles the participant to receive stock or cash measured by the value of the shares in the future. The delivery of stock or cash under a stock unit may be subject to the satisfaction of performance conditions or other vesting conditions.

Performance awards. A performance award is an award the vesting, settlement or exercisability of which is subject to specified performance criteria.

**Vesting.** Our compensation committee has the authority to determine the vesting schedule applicable to each award, and to accelerate the vesting or exercisability of any award.

**Termination of Employment.** Our compensation committee will determine the effect of termination of employment or service on an award. Unless otherwise provided by our compensation committee or



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in an award agreement, upon a termination of a participant's employment all unvested options then held by the participant and other awards requiring exercise will terminate and all other unvested awards will be forfeited and all vested stock options and stock appreciation rights then held by the participant will remain outstanding for three months or one year in the case of death or, in each case, until the applicable expiration date, if earlier. All stock options and stock appreciation rights held by a participant immediately prior to the participant's termination of employment will immediately terminate upon termination of employment if the termination is for cause as defined in the 2013 Equity Incentive Plan or occurs in circumstances that would have constituted grounds for the participant's employment to be terminated for cause, in the determination of the Administrator.

**Performance Criteria.** The 2013 Equity Incentive Plan provides for the grant of performance awards that are made based upon, and subject to achieving, "performance objectives". Performance objectives with respect to those awards that are intended to qualify as "performance-based compensation" for purposes of Section 162(m) of the Code, or Section 162(m) are limited to an objectively determinable measure or measures of performance relating to any or any combination of the following (measured either absolutely or by reference to an index or indices and determined either on a consolidated basis or, as the context permits, on a divisional, subsidiary, line of business, project or geographical basis or in combinations thereof): sales; revenues; assets; expenses; earnings before or after deduction for all or any portion of interest, taxes, depreciation, amortization or equity expense, whether or not on a continuing operations or an aggregate or per share basis; return on equity, investment, capital, capital employed or assets; one or more operating ratios; operating income or profit, including on an after-tax basis; net income; borrowing levels, leverage ratios or credit rating; market share; capital expenditures; cash flow; stock price; stockholder return; sales of particular products or services; customer acquisition or retention; acquisitions and divestitures (in whole or in part); joint ventures, strategic alliances, licenses or collaborations; spin-offs, split-ups and the like; reorganizations; recapitalizations, restructurings, financings (issuance of debt or equity) or refinancings; manufacturing or process development; or achievement of clinical trial or research objectives, regulatory or other filings or approvals or other product development milestones.

To the extent consistent with the requirements for satisfying the performance-based compensation exception under Section 162(m), our compensation committee may provide in the case of any award intended to qualify for such exception that one or more of the performance objectives applicable to an award will be adjusted in an objectively determinable manner to reflect events (for example, the impact of charges for restructurings, discontinued operations, mergers, acquisitions, extraordinary items, and other unusual or non-recurring items, and the cumulative effects of tax on accounting changes, each as defined by U.S. generally accepted accounting principles) occurring during the performance period of such award that affect the applicable performance objectives.

**Transferability.** Awards under the 2013 Equity Incentive Plan may not be transferred except by will or by the laws of descent and distribution, unless (for awards other than ISOs) otherwise provided by our compensation committee.

**Covered Transactions.** In the event of a consolidation, merger or similar transaction, a sale or transfer of all or substantially all of our assets or our dissolution or liquidation, our compensation committee may, among other things, provide for continuation or assumption of outstanding awards, for new grants in substitution of outstanding awards, for the accelerated vesting or delivery of shares under awards or for a cash-out of outstanding awards, in each case on such terms and with such restrictions as it deems appropriate. Except as our compensation committee may otherwise determine, awards not assumed will automatically terminate and in the case of outstanding shares of restricted stock, will automatically be forfeited upon the consummation of such covered transaction.

**Adjustment.** In the event of a stock dividend, stock split or combination of shares including a reverse stock split, recapitalization or other change in our capital structure that constitutes an equity

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restructuring within the meaning of the Financial Accounting Standards Board, Accounting Standards Codification Topic 718, *Compensation Stock Compensation*, our compensation committee will make appropriate adjustments to the maximum number of shares that may be delivered under, and the individual share limits included in, the 2013 Equity Incentive Plan, and will also make appropriate adjustments to the number and kind of shares of stock or securities subject to awards, the exercise prices of such awards or any other terms of awards affected by such change. Our compensation committee will also make the types of adjustments described above to take into account distributions and other events other than those listed above if it determines that such adjustments are appropriate to avoid distortion and preserve the value of awards.

***Amendment and Termination.*** Our compensation committee will be able to amend the 2013 Equity Incentive Plan or outstanding awards, or terminate the 2013 Equity Incentive Plan as to future grants of awards, except that our compensation committee will not be able to alter the terms of an award if it would affect materially and adversely a participant's rights under the award without the participant's consent (unless expressly provided in the 2013 Equity Incentive Plan or the right to alter the terms of an award was expressly reserved by our compensation committee at the time the award was granted). Stockholder approval will be required for any amendment to the 2013 Equity Incentive Plan to the extent such approval is required by law, including the Code or applicable stock exchange requirements.

***Employee Stock Purchase Plan***

Our board of directors adopted an Employee Stock Purchase Plan (the "ESPP") as a means of permitting our eligible employees, including our named executive officers, to acquire shares of our common stock. Two hundred seventy five thousand (275,000) shares of our common stock are available for issuance under the ESPP. Under the ESPP, eligible employees of the company may purchase shares of our common stock during pre-specified purchase periods at a price equal to the lesser of 85% of the fair market value of a share of our common stock at the beginning of the purchase period or 85% of the fair market value of a share of our common stock at the end of the purchase period. As of the date of this prospectus, the initial purchase period under the ESPP has not yet commenced and our board of directors has not determined the date on which such initial purchase period will commence.

***Acceleron Pharma Inc. Cash Incentive Plan***

Our board of directors adopted the Acceleron Pharma Inc. Cash Incentive Plan, or the Cash Incentive Plan. Starting with our 2014 fiscal year, annual award opportunities for executive officers, including our named executive officers, and other key employees will be granted under the Cash Incentive Plan. The following summary describes the material terms of the Cash Incentive Plan. This summary is not a complete description of all provisions of the Cash Incentive Plan and is qualified in its entirety by reference to the Cash Incentive Plan.

***Administration.*** The Cash Incentive Plan will be administered by our compensation committee. Our compensation committee has authority to interpret the Cash Incentive Plan, and any interpretation or decision by the compensation committee with regard to any questions arising under the Cash Incentive Plan will be final and conclusive on all participants.

***Eligibility.*** Executive officers and other key employees of the Company and our subsidiaries will be selected from time to time by the compensation committee to participate in the Cash Incentive Plan.

***Awards.*** Award opportunities under the Cash Incentive Plan will be granted by our compensation committee prior to, or within a specified period of time following the beginning of, the fiscal year of the Company (or other performance period selected by the compensation committee). The compensation committee will establish the performance criteria applicable to the award, the amount or amounts payable if the performance criteria are achieved, and such other terms and conditions as the

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compensation committee deems appropriate. The Cash Incentive Plan permits the grant of awards that are intended to qualify as exempt performance-based compensation under Section 162(m) as well as awards that are not intended to so qualify. Any awards that are intended to qualify as performance-based compensation will be administered in accordance with the requirements of Section 162(m).

**Performance Criteria.** Awards under the Cash Incentive Plan will be made based on, and subject to achieving, "performance criteria" established by our compensation committee. Performance criteria for awards intended to qualify as performance-based compensation for purposes of Section 162(m) are limited to the objectively determinable measures of performance relating to any or any combination of the following (measured either absolutely or by reference to an index or indices or the performance of one or more companies and determined either on a consolidated basis or, as the context permits, on a divisional, subsidiary, line of business, project or geographical basis or in combinations thereof): sales; revenues; assets; expenses; earnings before or after deduction for all or any portion of interest, taxes, depreciation, amortization or equity expense, whether or not on a continuing operations or an aggregate or per share basis; return on equity, investment, capital, capital employed or assets; one or more operating ratios; operating income or profit, including on an after-tax basis; net income; borrowing levels, leverage ratios or credit rating; market share; capital expenditures; cash flow; stock price; stockholder return; sales of particular products or services; customer acquisition or retention; acquisitions and divestitures (in whole or in part); joint ventures, strategic alliances, licenses or collaborations; spin-offs, split-ups and the like; reorganizations; recapitalizations, restructurings, financings (issuance of debt or equity) or refinancings; manufacturing or process development; or achievement of clinical trial or research objectives, regulatory or other filings or approvals or other product development milestones.

To the extent consistent with the requirements of Section 162(m), the compensation committee may establish that in the case of any award intended to qualify as exempt performance-based compensation under Section 162(m), that one or more of the performance criteria applicable to such award be adjusted in an objectively determinable manner to reflect events (for example, the impact of charges for restructurings, discontinued operations, mergers, acquisitions, extraordinary items, and other unusual or non-recurring items, and the cumulative effects of tax on accounting changes, each as defined by U.S. generally accepted accounting principles) occurring during the performance period of such award that affect the applicable performance criteria.

**Payment.** A participant will be entitled to payment under an award only if all conditions to payment have been satisfied in accordance with the Cash Incentive Plan and the terms of the award. Following the close of the performance period, our compensation committee will determine (and, to the extent required by Section 162(m), certify) whether and to what extent the applicable performance criteria have been satisfied. Our compensation committee will then determine the actual payment, if any, under each award. Our compensation committee has the sole and absolute discretion to reduce (including to zero) the actual payment to be made under any award. Our compensation committee will determine the payment dates for awards under the Cash Incentive Plan. Our compensation committee may permit a participant to defer payment of an award.

**Payment Limits.** The maximum payment to any participant under the Cash Incentive Plan in any fiscal year will in no event exceed \$1 million.

**Recovery of Compensation.** Awards under the Cash Incentive Plan will be subject to forfeiture, termination and rescission, and a participant who receives a payment pursuant to the Cash Incentive Plan will be obligated to return to us such payment, to the extent provided by our compensation committee in an award agreement, pursuant to Company policy relating to the recovery of erroneously-paid incentive compensation, or as otherwise required by law or applicable stock exchange listing standards.

**Amendment and Termination.** Our compensation committee may amend the Cash Incentive Plan at any time, provided that any amendment will be approved by our stockholders if required by Section 162(m). Our compensation committee may terminate the Cash Incentive Plan at any time.

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**CERTAIN RELATIONSHIPS AND RELATED PARTY TRANSACTIONS**

The following is a description of transactions, since January 1, 2012, to which we have been a party, in which the amount involved exceeded or will exceed \$120,000, and in which any related person had a direct or indirect material interest.

**Indemnification Agreements**

We have entered into indemnification agreements with each of our directors and executive officers. These agreements will require us to indemnify these individuals and, in certain cases, affiliates of such individuals, to the fullest extent permissible under Delaware law against liabilities that may arise by reason of their service to us or at our direction, and to advance expenses incurred as a result of any proceeding against them as to which they could be indemnified.

**Registration Rights Agreement**

In connection with our Series F preferred stock financing, on December 22, 2011, we entered into an amended and restated registration rights agreement with the holders of all of our then-outstanding shares of preferred stock including certain of our named executive officers and entities with which certain of our directors are affiliated. The agreement provides that these holders, for so long as they may hold registrable securities, as defined in the agreement, have the right to demand that we file a registration statement with respect to the common stock issued upon conversion of our preferred stock. These holders may also request that shares of common stock held by them be included in certain registration statements that we are otherwise filing. See "Description of Capital Stock Registration Rights".

**Right of First Refusal and Co-Sale Agreement**

In connection with our Series F preferred stock financing, on December 22, 2011, we entered into an amended and restated right of first refusal and co-sale agreement with the holders of all of our then-outstanding shares of preferred stock including certain of our named executive officers and entities with which certain of our directors are affiliated. Pursuant to the terms of this agreement, in the event of a proposed sale of shares of our common or preferred stock, the seller was required to first offer such shares to the company and to the other investors, subject to certain conditions and restrictions. This agreement terminated upon the completion of our initial public offering on September 24, 2013.

**Voting Agreement**

In connection with our Series F preferred stock financing on December 22, 2011, we entered into an amended and restated voting agreement with the holders of all of our then-outstanding shares of preferred stock including certain of our named executive officers and entities with which certain of our directors are affiliated, with respect to the election of directors and certain other matters. All of our current directors were elected pursuant to the terms of this agreement. This agreement terminated upon the completion of our initial public offering on September 24, 2013.

**Investor Rights Agreement**

In connection with our Series F preferred stock financing, on December 22, 2011, we entered into an amended and restated investor rights agreement with the holders of all of our then-outstanding shares of preferred stock including certain of our named executive officers and entities with which certain of our directors are affiliated. Pursuant to the terms of this agreement, we granted our investors certain information rights as well as the right to participate pro rata in any future private financing rounds. This agreement terminated upon the completion of our initial public offering on September 24, 2013.



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**Transactions with Our Executive Officers, Directors and 5% Stockholders**

On March 13, 2013, we repurchased shares of our common and preferred stock, as well as warrants to purchase shares of our common stock, from UBS Juniper Crossover Fund, LLC, an affiliate of OrbiMed Advisors, for \$300,000 in aggregate.

On January 28, 2008, we entered into a secured promissory note with our chief executive officer, which was amended on November 13, 2012, with a principal balance of \$200,000 and an interest rate of 3.11% per annum. The note was secured by a pledge of shares of the company's stock under a pledge agreement dated January 28, 2008. The terms of the note provided that it would mature on January 28, 2014, or be forgiven upon the occurrence of certain corporate events, including the filing of a registration statement with the SEC. The outstanding principal balance and accrued and unpaid interest on the note was forgiven on August 7, 2013. See "Executive Compensation Executive Loan".

**Related Person Transactions Policy**

We have adopted a related person transaction approval policy that governs the review of related person transactions. Pursuant to this policy, if we want to enter into a transaction with a related person or an affiliate of a related person, our General Counsel will review the proposed transaction to determine, based on applicable NASDAQ and SEC rules, if such transaction requires pre-approval by the audit committee and/or board of directors. If pre-approval is required, such matters will be reviewed at the next regular or special audit committee and/or board of directors meeting. We may not enter into a related person transaction unless our General Counsel has either specifically confirmed in writing that no further reviews are necessary or that all requisite corporate reviews have been obtained.

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**PRINCIPAL STOCKHOLDERS**

The following table sets forth information relating to the beneficial ownership of our common stock as of January 1, 2014, by: each person, or group of affiliated persons, known by us to beneficially own more than 5% of our outstanding shares of common stock; each of our directors; each of our named executive officers; and all directors and executive officers as a group.

The number of shares beneficially owned by each entity, person, director or executive officer is determined in accordance with the rules of the SEC, and the information is not necessarily indicative of beneficial ownership for any other purpose. Under such rules, beneficial ownership includes any shares over which the individual has sole or shared voting power or investment power as well as any shares that the individual has the right to acquire within 60 days of January 1, 2014 through the exercise of any stock option, warrants or other rights. Except as otherwise indicated, and subject to applicable community property laws, the persons named in the table have sole voting and investment power with respect to all shares of common stock held by that person.

The percentage of shares beneficially owned is computed on the basis of 28,348,633 shares of our common stock outstanding as of January 1, 2014. Shares of our common stock that a person has the right to acquire within 60 days of January 1, 2014 are deemed outstanding for purposes of computing the percentage ownership of the person holding such rights, but are not deemed outstanding for purposes of computing the percentage ownership of any other person, except with respect to the percentage ownership of all directors and executive officers as a group. Unless otherwise indicated

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below, the address for each beneficial owner listed is c/o John Quisel, 128 Sidney Street, Cambridge, MA 02139.

Name and address of beneficial owner	Number of shares beneficially owned	Percentage of shares beneficially owned	
		Before offering	After offering
<b>5% or greater stockholders:</b>			
Polaris Venture Partners, and related funds(1) 650 East Kendall Street, 4 <sup>th</sup> Floor Cambridge, MA 02142	3,317,136	11.6%	10.7%
Venrock Partners, and related funds(2) 55 Cambridge Parkway, Suite 100 Cambridge, MA 02142	2,622,435	9.2%	8.5%
Advanced Technology Ventures, and related funds(3) 500 Boylston Street, Suite 1380 Boston, MA 02116	2,662,092	9.3%	8.6%
Celgene Corporation(4) 86 Morris Avenue Summit, NJ 07901	3,211,866	11.3%	11.4%
Flagship Ventures(5) 1 Memorial Drive Cambridge, MA 02142	2,276,479	8.0%	7.4%
OrbiMed Advisors LLC(6) 601 Lexington Avenue, 54 <sup>th</sup> Floor New York, NY 10022	2,272,819	8.0%	7.4%
<b>Directors and named executive officers:</b>			
John L. Knopf, Ph.D.(7)	801,094	2.8%	2.6%
Anthony B. Evnin, Ph.D.(8)	2,624,102	9.2%	8.5%
Jean M. George(9) George Golumbeski, Ph.D.	2,663,759	9.3%	8.6%
Edwin M. Kania, Jr.(10)	2,297,614	8.1%	7.5%
Terrance G. McGuire(11)	3,318,803	11.6%	10.7%
Tom Maniatis, Ph.D.(12)	291,681	1.0%	*
Richard F. Pops(13)	76,250	*	*
Joseph S. Zakrzewski(14)	18,855	*	*
Matthew L. Sherman, M.D.(15)	227,438	*	*
John Quisel, J.D., Ph.D.(16)	145,562	*	*
All executive officers and directors as a group (15 persons)(17)	13,021,202	43.7%	40.4%

\* Represents beneficial ownership of less than one percent of our outstanding common stock.

(1) Consists of (i) 3,077,388 shares of common stock and warrants to purchase 180,518 shares of common stock held by Polaris Venture Partners IV, L.P. and (ii) 55,846 shares of common stock and warrants to purchase 3,384 shares of common stock held by Polaris Venture Partners Entrepreneurs' Fund IV, L.P. (together with Polaris Venture Partners IV, L.P., the Polaris Funds). North Star Venture Management 2000, LLC directly or indirectly provides investment advisory services to various venture capital funds, including the Polaris Funds. Each of the Polaris Funds has the sole voting and investment power with respect to the shares of the Company directly held by the applicable Polaris Fund. The respective general partners of the Polaris Funds may be deemed to have sole voting and investment power with respect to the shares held by such funds.

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The respective general partners disclaim beneficial ownership of all the shares held by the Polaris Funds except to the extent of their proportionate pecuniary interests therein. The members of North Star Venture Management 2000, LLC (the Polaris Management Members) are also members of Polaris Venture Management Co., IV, L.L.C. (the general partner of each of the Polaris Funds). As members of the general partner and North Star Venture Management 2000, LLC, the Polaris Management Members may be deemed to share voting and investment powers for the shares held by the Polaris Funds. The Polaris Management Members disclaim beneficial ownership of all such shares held by the funds except to the extent of their proportionate pecuniary interests therein. Mr. Terrance G. McGuire, a director of the Company, has an assignee interest in Polaris Venture Management Co. IV, L.L.C. To the extent that he is deemed to share voting and investment powers with respect to the shares held by the Polaris Funds, Mr. McGuire disclaims beneficial ownership of all the shares held by the funds except to the extent of his proportionate pecuniary interest therein.

- (2) Consists of (i) 414,360 shares of common stock and warrants to purchase 20,936 shares of common stock held by Venrock Partners, L.P. (Venrock Partners), (ii) 2,032,352 shares of common stock and warrants to purchase 102,795 shares of common stock held by Venrock Associates IV, L.P. (Venrock IV) and (iii) 49,440 shares of common stock and warrants to purchase 2,525 shares of common stock held by Venrock Entrepreneurs Fund IV, L.P. (Venrock Entrepreneurs, and together with Venrock Partners and Venrock IV, the Venrock Entities). The sole general partner of Venrock IV is Venrock Management IV, LLC (VM4). The sole general partner of Venrock Partners is Venrock Partners Management, LLC (VPM). The sole general partner of Venrock Entrepreneurs is VEF Management IV, LLC (VEFM4). VM4, VPM and VEFM4 disclaim beneficial ownership over all shares held by the Venrock Entities, except to the extent of their indirect pecuniary interests therein. Anthony B. Evnin, Ph.D., a director of the Company, is a member of VM4, VPM and VEFM4 and as such, he may be deemed to have voting and investment power with respect to these shares. Dr. Evnin disclaims beneficial ownership of these shares except to the extent of his indirect pecuniary interest therein.
- (3) Consists of (i) 2,018,586 shares of common stock and warrants to purchase 119,322 shares of common stock held by Advanced Technology Ventures VII, L.P. (ATV VII), (ii) 81,002 shares of common stock and warrants to purchase 4,788 shares of common stock held by Advanced Technology Ventures VII (B), L.P. (ATV VII-B), (iii) 38,934 shares of common stock and warrants to purchase 2,301 shares of common stock held by Advanced Technology Ventures VII (C), L.P. (ATV VII-C), (iv) 12,025 shares of common stock and warrants to purchase 711 shares of common stock held by ATV Entrepreneurs VII, L.P. (ATV VII-E), (v) 337,565 shares of common stock and warrants to purchase 19,916 shares of common stock held by Advanced Technology Ventures VI, L.P. (ATV VI), (vi) 21,543 shares of common stock and warrants to purchase 1,271 shares of common stock held by ATV Entrepreneurs VI, L.P. (ATV VI-E) and (vii) 4,128 shares of common stock held by ATV Alliance 2003, L.P. (ATV A 2003). ATV Associates VII, L.L.C (ATV A VII) is the general partner of ATV VII, ATV VII-B, ATV VII-C and ATV VII-E and exercises voting and dispositive authority over the shares held by ATV VII, ATV VII-B, ATV VII-C and ATV VII-E. Voting and dispositive decisions of ATV A VII are made collectively by Michael A. Carusi, Jean M. George (one of our directors), Steven N. Baloff, Robert C. Hower and William C. Wiberg (collectively, the ATV A VII Managing Directors). ATV A VII and each of the ATV A VII Managing Directors disclaim beneficial ownership of the shares held by ATV VII, ATV VII-B, ATV VII-C and ATV VII-E except to the extent of their pecuniary interest therein. ATV Associates VI, L.L.C (ATV A VI) is the general partner of ATV VI and ATV VI-E and exercises voting and dispositive authority over the shares held by ATV VI and ATV VI-E. Voting and dispositive decisions of ATV A VI are made collectively by Michael A. Carusi, Steven N. Baloff, Pieter J. Schiller, Robert C. Hower and William C. Wiberg (collectively, the ATV A VI Managing Directors). ATV A VI and each of the ATV A VI Managing Directors disclaim

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beneficial ownership of the shares held by ATV VI and ATV VI-E except to the extent of their pecuniary interest therein. ATV Alliance Associates, L.L.C. (ATV Alliance, LLC) is the general partner of ATV A 2003 and exercises voting and dispositive authority over the shares held by ATV A 2003. Voting and dispositive decisions of ATV Alliance, LLC are made by Jean M. George (one of our directors). ATV Alliance, LLC and Jean M. George disclaim beneficial ownership of the shares held by ATV A 2003 except to the extent of their pecuniary interest therein.

- (4) Includes 38,979 shares of common stock that can be acquired upon the exercise of warrants to purchase shares of our common stock. Percentage of shares owned after the offering includes 300,000 shares purchased in the offering.
- (5) Consists of (i) 2,146,720 shares of common stock held by Applied Genomic Technology Capital Fund, L.P. (AGTC Fund) and (ii) 129,759 shares of common stock held by AGTC Advisors Fund, L.P. (AGTC). NewcoGen Group, Inc., or NewcoGen Inc., is the general partner of AGTC Partners, L.P., which is the general partner of each of AGTC and AGTC Fund. NewcoGen Inc. is a wholly-owned subsidiary of Flagship Ventures Management, Inc. Noubar B. Afeyan Ph.D. and Edwin M. Kania, Jr. are the directors of Flagship Ventures Management, Inc. and may be deemed to have beneficial ownership with respect to all shares held by AGTC and AGTC Fund. Mr. Kania, one of our directors, and Dr. Afeyan disclaim beneficial ownership over shares held by AGTC and AGTC Fund except to the extent of their pecuniary interest therein.
- (6) Consists of (i) 509,989 shares of common stock and warrants to purchase 33,229 shares of common stock held by OrbiMed Private Investments II, LP (OPI II), (ii) 1,362,080 shares of common stock and warrants to purchase 88,749 shares of common stock held by OrbiMed Private Investments II (QP), LP (OPI QP), (iii) 178,679 shares of common stock and warrants to purchase 24,151 shares of common stock held by OPI II, and (iv) 66,900 shares of common stock and warrants to purchase 9,042 shares of common stock held by OPI QP. OrbiMed Advisors LLC, or OrbiMed, a registered investment adviser under the Investment Advisers Act of 1940, as amended, is the managing member of OrbiMed Capital GP II LLC, which is the general partner of OPI II and OPI QP (collectively, the OrbiMed Funds). Mr. Samuel D. Isaly is the managing member of and owner of a controlling interest in OrbiMed. Accordingly, OrbiMed and Mr. Isaly may be deemed to have voting and investment power over the shares held by OrbiMed Funds noted above. OrbiMed and Mr. Isaly disclaim beneficial ownership with respect to such shares, except to the extent of their pecuniary interest therein, if any.
- (7) Includes 581,094 shares of common stock that can be acquired upon the exercise of outstanding options.
- (8) Consists of shares held by the Venrock Entities. By virtue of the relationships described in footnote 2 above, Dr. Evnin may be deemed to share beneficial ownership in the shares held by the Venrock Entities. Dr. Evnin disclaims beneficial ownership of the shares referred to in footnote 2 above. Includes 1,666 shares of common stock that can be acquired upon the exercise of outstanding options.
- (9) Consists of shares held by the ATV Entities. By virtue of the relationships described in footnote 3 above, Ms. George may be deemed to share beneficial ownership in the shares held by the ATV Entities. Ms. George disclaims beneficial ownership of the shares referred to in footnote 3 above. Includes 1,666 shares of common stock that can be acquired upon the exercise of outstanding options.
- (10) Consists of shares held by AGTC or AGTC Fund. By virtue of the relationships described in footnote 4 above, Mr. Kania may be deemed to share beneficial ownership in the shares held by AGTC or AGTC Fund. Mr. Kania disclaims beneficial ownership of the shares referred to in

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footnote 4 above. Includes 1,666 shares of common stock that can be acquired upon the exercise of outstanding options.

- (11) Consists of shares held by Polaris Venture Partners or related funds. By virtue of the relationships described in footnote 1 above, Mr. McGuire may be deemed to share beneficial ownership in the shares held by Polaris Venture Partners or related funds. Mr. McGuire disclaims beneficial ownership of the shares referred to in footnote 1 above.
- (12) Includes 1,666 shares of common stock that can be acquired upon the exercise of outstanding options.
- (13) Includes 26,250 shares of common stock that can be acquired upon the exercise of outstanding options.
- (14) Includes 47,500 shares of common stock that can be acquired upon the exercise of outstanding options.
- (15) Includes 18,855 shares of common stock that can be acquired upon the exercise of outstanding options.
- (16) Includes 170,016 shares of common stock that can be acquired upon the exercise of outstanding options.
- (17) Includes 145,562 shares of common stock that can be acquired upon the exercise of outstanding options.
- (17) Includes 1,476,439 shares of common stock that can be acquired upon the exercise of outstanding options.

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**DESCRIPTION OF CAPITAL STOCK**

**General**

The following description of our capital stock is intended as a summary only and is qualified in its entirety by reference to our restated certificate of incorporation and amended and restated by-laws, which are filed as exhibits to the registration statement of which this prospectus is a part, and to the applicable provisions of the Delaware General Corporation Law. We refer in this section to our restated certificate of incorporation as our certificate of incorporation, and we refer to our amended and restated by-laws as our by-laws.

Our authorized capital stock consists of 175,000,000 shares of our common stock, par value \$0.001 per share, and 25,000,000 shares of our preferred stock, par value \$0.001 per share, all of which preferred stock is undesignated.

As of January 1, 2014, we had issued and outstanding:

28,348,633 shares of our common stock;

options to purchase a total of 3,942,304 shares of our common stock with a weighted-average exercise price of \$7.05 per share;

warrants to purchase a total of 979,699 shares of our common stock with a weighted-average exercise price of \$6.53 per share;

As of January 1, 2014, we had 174 stockholders of record.

**Common Stock**

**Dividend Rights.** Subject to preferences that may apply to shares of preferred stock outstanding at the time, holders of outstanding shares of common stock will be entitled to receive dividends out of assets legally available at the times and in the amounts as the board of directors may from time to time determine.

**Voting Rights.** Each outstanding share of common stock will be entitled to one vote on all matters submitted to a vote of stockholders. Holders of shares of our common stock shall have no cumulative voting rights.

**Preemptive Rights.** Our common stock will not be entitled to preemptive or other similar subscription rights to purchase any of our securities.

**Conversion or Redemption Rights.** Our common stock will be neither convertible nor redeemable.

**Liquidation Rights.** Upon our liquidation, the holders of our common stock will be entitled to receive pro rata our assets which are legally available for distribution, after payment of all debts and other liabilities and subject to the prior rights of any holders of preferred stock then outstanding.

**Listing.** Our common stock is listed on the NASDAQ Global Market under the symbol XLRN.

**Preferred Stock**

Our board of directors may, without further action by our stockholders, from time to time, direct the issuance of shares of preferred stock in series and may, at the time of issuance, determine the designations, powers, preferences, privileges, and relative participating, optional or special rights as well as the qualifications, limitations or restrictions thereof, including dividend rights, conversion rights, voting rights, terms of redemption and liquidation preferences, any or all of which may be greater than the rights of the common stock. Satisfaction of any dividend preferences of outstanding shares of preferred stock would reduce the amount of funds available for the payment of dividends on shares of





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our common stock. Holders of shares of preferred stock may be entitled to receive a preference payment in the event of our liquidation before any payment is made to the holders of shares of our common stock. Under certain circumstances, the issuance of shares of preferred stock may render more difficult or tend to discourage a merger, tender offer or proxy contest, the assumption of control by a holder of a large block of our securities or the removal of incumbent management. Upon the affirmative vote of a majority of the total number of directors then in office, our board of directors, without stockholder approval, may issue shares of preferred stock with voting and conversion rights which could adversely affect the holders of shares of our common stock and the market value of our common stock. There are no shares of preferred stock outstanding, and we have no present intention to issue any shares of preferred stock.

**Registration Rights**

We are party to an amended and restated registration rights agreement with the holders of approximately 14.8 million shares of our common stock.

Under the amended and restated registration rights agreement, holders of registrable shares can demand that we file a registration statement or request that their shares be included on a registration statement that we are otherwise filing, in either case, registering the resale of their shares of common stock. These registration rights are subject to conditions and limitations, including the right, in certain circumstances, of the underwriters of an offering to limit the number of shares included in such registration and our right, in certain circumstances, not to effect a requested S-1 or S-3 registration within 60 days before or six months following the Company's estimated date of filing of a registration statement pertaining to an underwritten public offering of securities for the account of the Company offering of our securities, including this offering.

*Demand Registration Rights*

Following the expiration of any applicable lock up periods to which they are subject, the holders of at least a majority of the registrable shares may require us to file a registration statement under the Securities Act at our expense with respect to the resale of their registrable shares, and we are required to use our best efforts to effect the registration.

*Piggyback Registration Rights*

If we propose to register any of our securities under the Securities Act for our own account or the account of any other holder, the holders of registrable shares are entitled to notice of such registration and to request that we include registrable shares for resale on such registration statement, subject to the right of any underwriter to limit the number of shares included in such registration. We will pay all registration expenses, other than underwriting discounts and commissions, related to any demand or piggyback registration. The amended and restated registration rights agreement contains customary cross- indemnification provisions, pursuant to which we are obligated to indemnify the selling stockholders, in the event of misstatements or omissions in the registration statement attributable to us except in the event of fraud and they are obligated to indemnify us for misstatements or omissions attributable to them. The registration rights will not terminate until all registrable shares have been sold or no longer qualify as registrable shares.

**Anti-Takeover Effects of Our Certificate of Incorporation and Our By-Laws**

Our certificate of incorporation and by-laws contain certain provisions that are intended to enhance the likelihood of continuity and stability in the composition of the board of directors and which may have the effect of delaying, deferring or preventing a future takeover or change in control of the company unless such takeover or change in control is approved by the board of directors.

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These provisions include:

**Classified Board.** Our certificate of incorporation provides that our board of directors be divided into three classes of directors, with the classes as nearly equal in number as possible. As a result, approximately one-third of our board of directors will be elected each year. The classification of directors has the effect of making it more difficult for stockholders to change the composition of our board. Our certificate of incorporation also provides that, subject to any rights of holders of preferred stock to elect additional directors under specified circumstances, the number of directors are fixed exclusively pursuant to a resolution adopted by our board of directors.

**Action by Written Consent; Special Meetings of Stockholders.** Our certificate of incorporation provides that stockholder action can be taken only at an annual or special meeting of stockholders and cannot be taken by written consent in lieu of a meeting. Our certificate of incorporation and the by-laws also provide that, except as otherwise required by law, special meetings of the stockholders can only be called pursuant to a resolution adopted by a majority of the board of directors. Except as described above, stockholders are not permitted to call a special meeting or to require the board of directors to call a special meeting.

**Removal of Directors.** Our certificate of incorporation provides that our directors may be removed only for cause by the affirmative vote of at least 75% of the voting power of our outstanding shares of capital stock, voting together as a single class. This requirement of a supermajority vote to remove directors could enable a minority of our stockholders to prevent a change in the composition of our board.

**Advance Notice Procedures.** Our by-laws establish an advance notice procedure for stockholder proposals to be brought before an annual meeting of our stockholders, including proposed nominations of persons for election to the board of directors. Stockholders at an annual meeting will only be able to consider proposals or nominations specified in the notice of meeting or brought before the meeting by or at the direction of the board of directors or by a stockholder who was a stockholder of record on the record date for the meeting, who is entitled to vote at the meeting and who has given our Secretary timely written notice, in proper form, of the stockholder's intention to bring that business before the meeting. Although the by-laws do not give the board of directors the power to approve or disapprove stockholder nominations of candidates or proposals regarding other business to be conducted at a special or annual meeting, the by-laws may have the effect of precluding the conduct of certain business at a meeting if the proper procedures are not followed or may discourage or deter a potential acquiror from conducting a solicitation of proxies to elect its own slate of directors or otherwise attempting to obtain control of the company.

**Super Majority Approval Requirements.** The Delaware General Corporation Law generally provides that the affirmative vote of a majority of the shares entitled to vote on any matter is required to amend a corporation's certificate of incorporation or by-laws, unless either a corporation's certificate of incorporation or by-laws requires a greater percentage. Our certificate of incorporation and by-laws provide that the affirmative vote of holders of at least 75% of the total votes eligible to be cast in the election of directors are required to amend, alter, change or repeal specified provisions. This requirement of a supermajority vote to approve amendments to our certificate of incorporation and by-laws could enable a minority of our stockholders to exercise veto power over any such amendments.

**Authorized but Unissued Shares.** Our authorized but unissued shares of common stock and preferred stock will be available for future issuance without stockholder approval. These additional shares may be utilized for a variety of corporate purposes, including future public offerings to raise additional capital, corporate acquisitions and employee benefit plans. The existence of authorized but unissued shares of common stock and preferred stock could render more difficult or discourage an

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attempt to obtain control of a majority of our common stock by means of a proxy contest, tender offer, merger or otherwise.

***Exclusive Forum.*** Our certificate of incorporation provides that, subject to limited exceptions, the state or federal court located within the State of Delaware is the sole and exclusive forum for (1) any derivative action or proceeding brought on our behalf, (2) any action asserting a claim of breach of a fiduciary duty owed by any of our directors, officers or other employees to us or our stockholders, (3) any action asserting a claim against us arising pursuant to any provision of the Delaware General Corporation Law, our certificate of incorporation or our by-laws, or (4) any other action asserting a claim against us that is governed by the internal affairs doctrine. Any person or entity purchasing or otherwise acquiring any interest in shares of our capital stock shall be deemed to have notice of and to have consented to the provisions of our certificate of incorporation described above. Although we believe these provisions benefit us by providing increased consistency in the application of Delaware law for the specified types of actions and proceedings, the provisions may have the effect of discouraging lawsuits against our directors and officers. The enforceability of similar choice of forum provisions in other companies' certificates of incorporation has been challenged in legal proceedings, and it is possible that, in connection with one or more actions or proceedings described above, a court could find the choice of forum provisions contained in our certificate of incorporation to be inapplicable or unenforceable.

***Section 203 of the Delaware General Corporation Law***

We are subject to the provisions of Section 203 of the Delaware General Corporation Law. In general, Section 203 prohibits a publicly-held Delaware corporation from engaging in a "business combination" with an "interested stockholder" for a three-year period following the time that this stockholder becomes an interested stockholder, unless the business combination is approved in a prescribed manner. A "business combination" includes, among other things, a merger, asset or stock sale or other transaction resulting in a financial benefit to the interested stockholder. An "interested stockholder" is a person who, together with affiliates and associates, owns, or did own within three years prior to the determination of interested stockholder status, 15% or more of the corporation's voting stock.

Under Section 203, a business combination between a corporation and an interested stockholder is prohibited unless it satisfies one of the following conditions: before the stockholder became interested, the board of directors approved either the business combination or the transaction which resulted in the stockholder becoming an interested stockholder; upon consummation of the transaction which resulted in the stockholder becoming an interested stockholder, the interested stockholder owned at least 85% of the voting stock of the corporation outstanding at the time the transaction commenced, excluding for purposes of determining the voting stock outstanding, shares owned by persons who are directors and also officers, and employee stock plans, in some instances; or at or after the time the stockholder became interested, the business combination was approved by the board of directors of the corporation and authorized at an annual or special meeting of the stockholders by the affirmative vote of at least two-thirds of the outstanding voting stock which is not owned by the interested stockholder.

A Delaware corporation may "opt out" of these provisions with an express provision in its original certificate of incorporation or an express provision in its certificate of incorporation or by-laws resulting from a stockholders' amendment approved by at least a majority of the outstanding voting shares. We have not opted out of these provisions. As a result, mergers or other takeover or change in control attempts of us may be discouraged or prevented.

***Transfer Agent and Registrar***

The transfer agent and registrar for our common stock is Computershare Trust Company, N.A. The transfer agent and registrar's address is 250 Royall Street Canton, Massachusetts 02021.

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**SHARES ELIGIBLE FOR FUTURE SALE**

Future sales of our common stock, including shares issued upon the exercise of outstanding options or warrants, in the public market after this offering, or the perception that those sales may occur, could cause the prevailing market price for our common stock to fall or impair our ability to raise equity capital in the future.

As of January 1, 2014, based on the number of shares of our common stock then outstanding, assuming (1) the closing of this offering, (2) no exercise of the underwriters' option to purchase additional shares of common stock, and (3) no exercise of outstanding options or warrants, we would have had outstanding an aggregate of 30,748,633 shares of common stock. Of these shares, 7,482,723 shares of common stock, including the 6,417,000 shares sold in our initial public offering, and all of the shares of common stock to be sold in this offering, and any shares sold upon exercise of the underwriters' option to purchase additional shares, will be freely tradable in the public market without restriction or further registration under the Securities Act of 1933, as amended, or the Securities Act, unless the shares are held by any of our "affiliates" as such term is defined in Rule 144 of the Securities Act. The remaining shares of common stock are "restricted securities" as such term is defined in Rule 144 or subject to lock up agreements in effect in connection with the initial public offering or entered into in connection with this offering (as described below) and will be available for sale in the public market as follows:

**Approximate Number of Shares**

5,896,337 shares, or 19%

**First Date Available for Sale into Public Market**

March 17, 2014 due to lock up agreements in effect in connection with our initial public offering. However, the representatives of the underwriters can waive the provisions of these lock-up agreements and allow these stockholders to sell their shares at any time.

15,269,537 shares, or 50%

90 days after the date of this prospectus upon expiration of the lock-up agreements referred to below, subject in some cases to applicable volume limitations under Rule 144.

**Lock-up Agreements**

In connection with this offering, we, our officers and directors and certain of their affiliates holding approximately 53% of our shares of common stock outstanding as of January 1, 2014 have agreed, subject to certain exceptions, with the underwriters not to dispose of or hedge any shares of our common stock or securities convertible into or exchangeable for shares of common stock during the period from the date of the lock-up agreement continuing through the date 90 days after the date of this prospectus, except with the prior written consent of Citigroup Global Markets Inc. and Leerink Partners LLC, the representatives of the underwriters. The representatives of the underwriters have advised us that they have no current intent or arrangement to release any of the shares subject to the lock-up agreements prior to the expiration of the lock-up period.

Following the lock-up periods set forth in the agreements described above, and assuming that the representatives of the underwriters do not release any parties from these agreements, all of the shares of our common stock that are restricted securities or are held by our affiliates as of the date of this prospectus will be eligible for sale in the public market in compliance with Rule 144 under the Securities Act.

In addition, pursuant to each of our amended and restated investors' rights agreement and amended and restated right of first refusal and co-sale agreement, the parties thereto have agreed that,

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if requested in writing by the representatives of the underwriters of the initial public offering of our securities, they will not sell, make any short sale of, grant any option for the purchase of, or otherwise dispose of any shares of our stock during the same 90-day restricted period referred to above. The representatives of the underwriters have invoked this written request and, accordingly, that the parties to these agreements will be subject to the related transaction restrictions. Holders of approximately 20.9 million shares of common stock, or 73.6% of our outstanding shares of common stock outstanding as of January 1, 2014, are collectively subject to lock-up restrictions as parties to these agreements, lock-up agreements with the underwriters in our initial public offering or lock-up agreements with the underwriters.

**Rule 144**

In general, under Rule 144, as currently in effect, once we have been subject to the public company reporting requirements of the Securities Exchange Act of 1934, as amended, or the Exchange Act, for at least 90 days, a person (or persons whose shares are required to be aggregated) who is not deemed to have been one of our "affiliates" for purposes of Rule 144 at any time during the three months preceding a sale, and who has beneficially owned restricted securities within the meaning of Rule 144 for at least six months, including the holding period of any prior owner other than one of our "affiliates," is entitled to sell those shares in the public market (subject to the lock-up agreement referred to above, if applicable) without complying with the manner of sale, volume limitations or notice provisions of Rule 144, but subject to compliance with the public information requirements of Rule 144. If such a person has beneficially owned the sales proposed to be sold for at least one year, including the holding period of any prior owner other than "affiliates", then such person is entitled to sell such shares in the public market without complying with any of the requirements of Rule 144 (subject to the lock-up agreement referred to above, if applicable). In general, under Rule 144, as currently in effect, once we have been subject to the public company reporting requirements of the Exchange Act for at least 90 days, our "affiliates", as defined in Rule 144, who have beneficially owned the shares proposed to be sold for at least six months are entitled to sell in the public market, upon expiration of any applicable lock-up agreements and within any three-month period, a number of those shares of our common stock that does not exceed the greater of: 1% of the number of common shares then outstanding, which will equal approximately 307,486 shares of common stock immediately after this offering (calculated on the basis of the number of shares of our common stock outstanding as of January 1, 2014 and the assumptions described above); or the average weekly trading volume of our common stock on the NASDAQ Global Market during the four calendar weeks preceding the filing of a notice on Form 144 with respect to such sale.

Such sales under Rule 144 by our "affiliates" or persons selling shares on behalf of our "affiliates" are also subject to certain manner of sale provisions, notice requirements and to the availability of current public information about us. Notwithstanding the availability of Rule 144, the holders of substantially all of our restricted securities have entered into lock-up agreements as referenced above and their restricted securities will become eligible for sale (subject to the above limitations under Rule 144) upon the expiration of the restrictions set forth in those agreements.

**Rule 701**

In general, under Rule 701 as currently in effect, any of our employees, directors, officers, consultants or advisors who acquired common stock from us in connection with a written compensatory stock or option plan or other written agreement in compliance with Rule 701 under the Securities Act before the effective date of the registration statement of which this prospectus is a part (to the extent such common stock is not subject to a lock-up agreement) is entitled to rely on Rule 701 to resell such shares beginning 90 days after we become subject to the public company reporting requirements of the Exchange Act in reliance on Rule 144, but without compliance with the holding period requirements

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contained in Rule 144. Accordingly, subject to any applicable lock-up agreements, beginning 90 days after we become subject to the public company reporting requirements of the Exchange Act, under Rule 701 persons who are not our "affiliates", as defined in Rule 144, may resell those shares without complying with the minimum holding period or public information requirements of Rule 144, and persons who are our "affiliates" may resell those shares without compliance with Rule 144's minimum holding period requirements (subject to the terms of the lock-up agreement referred to below, if applicable).

**Equity Incentive Plans**

We have filed with the SEC a registration statement under the Securities Act covering the shares of common stock that we may issue upon exercise of outstanding options reserved for issuance under our 2003 Stock Option and Restricted Stock Plan and 2013 Equity Incentive Plan. Accordingly, shares registered under such registration statement were available for sale in the open market following its effective date, subject to Rule 144 volume limitations and the lock-up agreements described above, if applicable.

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**MATERIAL UNITED STATES FEDERAL INCOME TAX  
CONSIDERATIONS FOR NON-U.S. HOLDERS**

The following is a summary of the material U.S. federal income and estate tax considerations relating to the purchase, ownership and disposition of our common stock by Non-U.S. Holders (defined below). This summary does not purport to be a complete analysis of all the potential tax considerations relevant to Non-U.S. Holders of our common stock. This summary is based upon the Internal Revenue Code, the Treasury regulations promulgated or proposed thereunder and administrative and judicial interpretations thereof, all as of the date hereof and all of which are subject to change at any time, possibly on a retroactive basis.

This summary assumes that shares of our common stock are held as "capital assets" within the meaning of Section 1221 of the Internal Revenue Code (generally, property held for investment). This summary does not purport to deal with all aspects of U.S. federal income and estate taxation that might be relevant to particular Non-U.S. Holders in light of their particular investment circumstances or status, nor does it address specific tax considerations that may be relevant to particular persons (including, for example, financial institutions, broker-dealers, insurance companies, partnerships or other pass-through entities, certain U.S. expatriates, tax-exempt organizations, pension plans, "controlled foreign corporations", "passive foreign investment companies", corporations that accumulate earnings to avoid U.S. federal income tax, persons in special situations, such as those who have elected to mark securities to market or those who hold common stock as part of a straddle, hedge, conversion transaction, synthetic security or other integrated investment, or holders subject to the alternative minimum or the newly effective 3.8% Medicare tax on net investment income). In addition, except as explicitly addressed herein with respect to estate tax, this summary does not address estate and gift tax considerations or considerations under the tax laws of any state, local or non-U.S. jurisdiction.

For purposes of this summary, a "Non-U.S. Holder" means a beneficial owner of common stock that for U.S. federal income tax purposes is not classified as a partnership and is not:

an individual who is a citizen or resident of the United States;

a corporation or any other organization taxable as a corporation for U.S. federal income tax purposes, created or organized in or under the laws of the United States, any state thereof or the District of Columbia;

an estate, the income of which is included in gross income for U.S. federal income tax purposes regardless of its source; or

a trust if (1) a U.S. court is able to exercise primary supervision over the trust's administration and one or more U.S. persons have the authority to control all of the trust's substantial decisions or (2) the trust has a valid election in effect under applicable U.S. Treasury regulations to be treated as a U.S. person.

If an entity that is classified as a partnership for U.S. federal income tax purposes holds our common stock, the tax treatment of persons treated as its partners for U.S. federal income tax purposes will generally depend upon the status of the partner and the activities of the partnership. Partnerships and other entities that are classified as partnerships for U.S. federal income tax purposes and persons holding our common stock through a partnership or other entity classified as a partnership for U.S. federal income tax purposes are urged to consult their own tax advisors.

There can be no assurance that the Internal Revenue Service (IRS) will not challenge one or more of the tax consequences described herein, and we have not obtained, nor do we intend to obtain a ruling from the IRS with respect to the U.S. federal income or estate tax consequences to a Non-U.S. Holder of the purchase, ownership or disposition of our common stock.

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**THIS SUMMARY IS FOR GENERAL INFORMATION ONLY AND IS NOT INTENDED TO BE TAX ADVICE. NON-U.S. HOLDERS ARE URGED TO CONSULT THEIR TAX ADVISORS CONCERNING THE U.S. FEDERAL INCOME AND ESTATE TAXATION, STATE, LOCAL AND NON-U.S. TAXATION AND OTHER TAX CONSEQUENCES TO THEM OF THE PURCHASE, OWNERSHIP AND DISPOSITION OF OUR COMMON STOCK.**

**Distributions on Our Common Stock**

As discussed under "Dividend Policy" above, we do not currently expect to pay dividends. In the event that we do make a distribution of cash or property with respect to our common stock, any such distributions generally will constitute dividends for U.S. federal income tax purposes to the extent of our current and accumulated earnings and profits, if any, as determined under U.S. federal income tax principles. If a distribution exceeds our current and accumulated earnings and profits, the excess will constitute a return of capital and will first reduce the holder's adjusted tax basis in our common stock, but not below zero. Any remaining excess will be treated as capital gain, subject to the tax treatment described below in "Gain on Sale, Exchange or Other Taxable Disposition of Our Common Stock". Any such distribution would also be subject to the discussion below under the section titled "Additional Withholding and Reporting Requirements".

Dividends paid to a Non-U.S. Holder generally will be subject to a 30% U.S. federal withholding tax unless such Non-U.S. Holder provides us or our agent, as the case may be, with the appropriate IRS Form W-8, such as:

IRS Form W-8BEN (or successor form) certifying, under penalties of perjury, a reduction in withholding under an applicable income tax treaty, or

IRS Form W-8ECI (or successor form) certifying that a dividend paid on common stock is not subject to withholding tax because it is effectively connected with a trade or business in the United States of the Non-U.S. Holder (in which case such dividend generally will be subject to regular graduated U.S. tax rates as described below).

The certification requirement described above must be provided to us or our agent prior to the payment of dividends and must be updated periodically. The certification also may require a Non-U.S. Holder that provides an IRS form or that claims treaty benefits to provide its U.S. taxpayer identification number. Special certification and other requirements apply in the case of certain Non-U.S. Holders that hold shares of our common stock through intermediaries or are pass-through entities for U.S. federal income tax purposes.

Each Non-U.S. Holder is urged to consult its own tax advisor about the specific methods for satisfying these requirements. A claim for exemption will not be valid if the person receiving the applicable form has actual knowledge or reason to know that the statements on the form are false.

If dividends are effectively connected with a trade or business in the United States of a Non-U.S. Holder (and, if required by an applicable income tax treaty, attributable to a U.S. permanent establishment), the Non-U.S. Holder, although exempt from the withholding tax described above (provided that the certifications described above are satisfied), generally will be subject to U.S. federal income tax on such dividends on a net income basis in the same manner as if it were a resident of the United States. In addition, if a Non-U.S. Holder is treated as a corporation for U.S. federal income tax purposes, the Non-U.S. Holder may be subject to an additional "branch profits tax" equal to 30% (unless reduced by an applicable income treaty) of its earnings and profits in respect of such effectively connected dividend income.

Non-U.S. Holders that do not timely provide us or our agent with the required certification, but which are eligible for a reduced rate of U.S. federal withholding tax pursuant to an income tax treaty, may obtain a refund or credit of any excess amount withheld by timely filing an appropriate claim for refund with the IRS.



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**Gain on Sale, Exchange or Other Taxable Disposition of Our Common Stock**

Subject to the discussion below under the section titled " Additional Withholding and Reporting Requirements", in general, a Non-U.S. Holder will not be subject to U.S. federal income tax or withholding tax on gain realized upon such holder's sale, exchange or other taxable disposition of shares of our common stock unless (1) such Non-U.S. Holder is an individual who is present in the United States for 183 days or more in the taxable year of disposition, and certain other conditions are met, (2) we are or have been a "United States real property holding corporation", as defined in the Internal Revenue Code (a USRPHC), at any time within the shorter of the five-year period preceding the disposition and the Non-U.S. Holder's holding period in the shares of our common stock, and certain other requirements are met, or (3) such gain is effectively connected with the conduct by such Non-U.S. Holder of a trade or business in the United States (and, if required by an applicable income tax treaty, is attributable to a permanent establishment maintained by such Non-U.S. Holder in the United States).

If the first exception applies, the Non-U.S. Holder generally will be subject to U.S. federal income tax at a rate of 30% (or at a reduced rate under an applicable income tax treaty) on the amount by which such Non-U.S. Holder's capital gains allocable to U.S. sources exceed capital losses allocable to U.S. sources during the taxable year of the disposition. If the third exception applies, the Non-U.S. Holder generally will be subject to U.S. federal income tax with respect to such gain on a net income basis in the same manner as if it were a resident of the United States and a Non-U.S. Holder that is a corporation for U.S. federal income tax purposes may also be subject to a branch profits tax with respect to any earnings and profits attributable to such gain at a rate of 30% (or at a reduced rate under an applicable income tax treaty).

Generally, a corporation is a USRPHC only if the fair market value of its U.S. real property interests (as defined in the Internal Revenue Code) equals or exceeds 50% of the sum of the fair market value of its worldwide real property interests plus its other assets used or held for use in a trade or business. Although there can be no assurance in this regard, we believe that we are not, and do not anticipate becoming, a USRPHC. However, because the determination of whether we are a USRPHC depends on the fair market value of our U.S. real property interests relative to the fair market value of other business assets, there can be no assurance that we will not become a USRPHC in the future. Even if we became a USRPHC, a Non-U.S. Holder would not be subject to U.S. federal income tax on a sale, exchange or other taxable disposition of our common stock by reason of our status as USRPHC so long as our common stock is regularly traded on an established securities market at any time during the calendar year in which the disposition occurs and such Non-U.S. Holder does not own and is not deemed to own (directly, indirectly or constructively) more than 5% of our common stock at any time during the shorter of the five year period ending on the date of disposition and the holder's holding period. However, no assurance can be provided that our common stock will be regularly traded on an established securities market for purposes of the rules described above. Prospective investors are encouraged to consult their own tax advisors regarding the possible consequences to them if we are, or were to become, a USRPHC.

**Additional Withholding and Reporting Requirements**

Legislation enacted in March 2010 and related Treasury guidance (commonly referred to as FATCA) will impose, in certain circumstances, U.S. federal withholding at a rate of 30% on payments of (1) dividends on our common stock on or after January 1, 2014, and (2) gross proceeds from the sale or other disposition of our common stock on or after January 1, 2017. In the case of payments made to a "foreign financial institution" as defined under FATCA (including, among other entities, an investment fund), as a beneficial owner or as an intermediary, the tax generally will be imposed, subject to certain exceptions, unless such institution (1) enters into (or is otherwise subject to) and complies with an agreement with the U.S. government (a "FATCA Agreement") or (2) complies with applicable foreign law enacted in connection with an intergovernmental agreement between the United States and

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a foreign jurisdiction (an "IGA"), in either case to, among other things, collect and provide to the U.S. or other relevant tax authorities certain information regarding U.S. account holders of such institution. In the case of payments made to a foreign entity that is not a foreign financial institution (as a beneficial owner), the tax generally will be imposed, subject to certain exceptions, unless such foreign entity provides the withholding agent with a certification that it does not have any "substantial U.S. owner" (generally, any specified U.S. person that directly or indirectly owns more than a specified percentage of such entity) or that identifies its substantial U.S. owners. If our common stock is held through a foreign financial institution that enters into (or is otherwise subject to) a FATCA Agreement, such foreign financial institution (or, in certain cases, a person paying amounts to such foreign financial institution) generally will be required, subject to certain exceptions, to withhold such tax on payments of dividends and proceeds described above made to (1) a person (including an individual) that fails to comply with certain information requests or (2) a foreign financial institution that has not entered into (and is not otherwise subject to) a FATCA Agreement and is not required to comply with FATCA pursuant to applicable foreign law enacted in connection with an IGA.

Prospective investors should consult their own tax advisors regarding the possible impact of these rules on their investment in our common stock, and the entities through which they hold our common stock, including, without limitation, the process and deadlines for meeting the applicable requirements to prevent the imposition of this 30% withholding tax under FATCA.

**Backup Withholding and Information Reporting**

We must report annually to the IRS and to each Non-U.S. Holder the gross amount of the distributions on our common stock paid to the holder and the tax withheld, if any, with respect to the distributions. Non-U.S. Holders may have to comply with specific certification procedures to establish that the holder is not a United States person (as defined in the Internal Revenue Code) in order to avoid backup withholding at the applicable rate, currently 28%, with respect to dividends on our common stock. Dividends paid to Non-U.S. Holders subject to the U.S. withholding tax, as described above under the section titled "Distributions on Our Common Stock", generally will be exempt from U.S. backup withholding.

Information reporting and backup withholding will generally apply to the proceeds of a disposition of our common stock by a Non-U.S. Holder effected by or through the U.S. office of any broker, U.S. or foreign, unless the holder certifies its status as a Non-U.S. Holder and satisfies certain other requirements, or otherwise establishes an exemption. Generally, information reporting and backup withholding will not apply to a payment of disposition proceeds to a Non-U.S. Holder where the transaction is effected outside the United States through a non-U.S. office of a broker. However, for information reporting purposes, dispositions effected through a non-U.S. office of a broker with substantial U.S. ownership or operations generally will be treated in a manner similar to dispositions effected through a U.S. office of a broker. Prospective investors should consult their own tax advisors regarding the application of the information reporting and backup withholding rules to them.

Copies of information returns may be made available to the tax authorities of the country in which the Non-U.S. Holder resides or, in which the Non-U.S. Holder is incorporated, under the provisions of a specific treaty or agreement.

Backup withholding is not an additional tax. Any amounts withheld under the backup withholding rules from a payment to a Non-U.S. Holder can be refunded or credited against the Non-U.S. Holder's U.S. federal income tax liability, if any, provided that an appropriate claim is timely filed with the IRS.

**Federal Estate Tax**

Common stock owned (or treated as owned) by an individual who is not a citizen or a resident of the United States (as defined for U.S. federal estate tax purposes) at the time of death will be included in the individual's gross estate for U.S. federal estate tax purposes unless an applicable estate or other tax treaty provides otherwise, and therefore, may be subject to U.S. federal estate tax.

Table of Contents**UNDERWRITING**

Citigroup Global Markets Inc. and Leerink Partners LLC are acting as joint book-running managers of the offering and as representatives of the underwriters named below. Subject to the terms and conditions stated in the underwriting agreement dated the date of this prospectus, each underwriter named below has severally agreed to purchase, and we have agreed to sell to that underwriter, the number of shares set forth opposite the underwriter's name.

<b>Underwriter</b>	<b>Number of Shares</b>
Citigroup Global Markets Inc.	960,000
Leerink Partners LLC	720,000
Piper Jaffray & Co.	420,000
JMP Securities LLC	300,000
 Total	 2,400,000

The underwriting agreement provides that the obligations of the underwriters to purchase the shares included in this offering are subject to approval of legal matters by counsel and to other conditions. The underwriters are obligated to purchase all the shares (other than those covered by the option to purchase additional shares described below) if they purchase any of the shares.

Shares sold by the underwriters to the public will initially be offered at the public offering price set forth on the cover of this prospectus. Any shares sold by the underwriters to securities dealers may be sold at a discount from the public offering price not to exceed \$1.80 per share. If all the shares are not sold at the initial offering price, the underwriters may change the offering price and the other selling terms. The representatives have advised us that the underwriters do not intend to make sales to discretionary accounts.

We have granted to the underwriters an option, exercisable for 30 days from the date of this prospectus, to purchase up to 360,000 additional shares at the public offering price less the underwriting discount. To the extent the option is exercised, each underwriter must purchase a number of additional shares approximately proportionate to that underwriter's initial purchase commitment. Any shares issued or sold under the option will be issued and sold on the same terms and conditions as the other shares that are the subject of this offering.

We, and our officers and directors have agreed that, for a period of 90 days from the date of this prospectus, we and they will not, without the prior written consent of Citigroup and Leerink, dispose of or hedge any shares or any securities convertible into or exchangeable for our common stock. Citigroup and Leerink in their sole discretion may release any of the securities subject to these lock-up agreements at any time, which, in the case of officers and directors, shall be with notice.

Our common stock is listed on the NASDAQ Global Market under the symbol XLRN.

The following table shows the underwriting discounts and commissions that we are to pay to the underwriters in connection with this offering. These amounts are shown assuming both no exercise and full exercise of the underwriters' option to purchase additional shares of common stock.

	<b>Paid by Acceleron</b>	
	<b>No Exercise</b>	<b>Full Exercise</b>
Per share	\$ 3.00	\$ 3.00
Total	\$ 7,200,000	\$ 8,280,000

We estimate that our portion of the total expenses of this offering will be approximately \$700,000.

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We have also agreed to reimburse the underwriters for certain of their expenses in an amount up to \$30,000 as set forth in the underwriting agreement.

In connection with the offering, the underwriters may purchase and sell shares in the open market. Purchases and sales in the open market may include short sales, purchases to cover short positions, which may include purchases pursuant to the underwriters' option to purchase additional shares, and stabilizing purchases.

Short sales involve secondary market sales by the underwriters of a greater number of shares than they are required to purchase in the offering.

"Covered" short sales are sales of shares in an amount up to the number of shares represented by the underwriters' option to purchase additional shares.

"Naked" short sales are sales of shares in an amount in excess of the number of shares represented by the underwriters' option to purchase additional shares.

Covering transactions involve purchases of shares either pursuant to the underwriters' option to purchase additional shares or in the open market in order to cover short positions.

To close a naked short position, the underwriters must purchase shares in the open market. A naked short position is more likely to be created if the underwriters are concerned that there may be downward pressure on the price of the shares in the open market after pricing that could adversely affect investors who purchase in the offering.

To close a covered short position, the underwriters must purchase shares in the open market or must exercise the option to purchase additional shares. In determining the source of shares to close the covered short position, the underwriters will consider, among other things, the price of shares available for purchase in the open market as compared to the price at which they may purchase shares through the option to purchase additional shares.

Stabilizing transactions involve bids to purchase shares so long as the stabilizing bids do not exceed a specified maximum.

Purchases to cover short positions and stabilizing purchases, as well as other purchases by the underwriters for their own accounts, may have the effect of preventing or retarding a decline in the market price of the shares. They may also cause the price of the shares to be higher than the price that would otherwise exist in the open market in the absence of these transactions. The underwriters may conduct these transactions on the NASDAQ Global Market, in the over-the-counter market or otherwise. If the underwriters commence any of these transactions, they may discontinue them at any time.

**Other Relationships**

Some of the underwriters and their affiliates have engaged in, and may in the future engage in, investment banking and other commercial dealings in the ordinary course of business with us or our affiliates, including serving as underwriters for our initial public offering in September 2013, for which they received, or may in the future receive, customary fees and commissions for these transactions.

**Conflicts of Interest**

The underwriters are full service financial institutions engaged in various activities, which may include securities trading, commercial and investment banking, financial advisory, investment management, principal investment, hedging, financing and brokerage activities. Certain of the underwriters and their respective affiliates may, from time to time, engage in transactions with and perform services for us in the ordinary course of their business for which they may receive customary



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fees and reimbursement of expenses. In the ordinary course of their various business activities, the underwriters and their respective affiliates may make or hold a broad array of investments and actively trade debt and equity securities (or related derivative securities) and financial instruments (which may include bank loans and/or credit default swaps) for their own account and for the accounts of their customers and may at any time hold long and short positions in such securities and instruments. Such investments and securities activities may involve securities and/or instruments of ours or our affiliates. The underwriters and their affiliates may also make investment recommendations and/or publish or express independent research views in respect of such securities or financial instruments and may hold, or recommend to clients that they acquire, long and/or short positions in such securities and instruments.

We have agreed to indemnify the underwriters against certain liabilities, including liabilities under the Securities Act, or to contribute to payments the underwriters may be required to make because of any of those liabilities.

**Notice to Prospective Investors in the European Economic Area**

In relation to each member state of the European Economic Area that has implemented the Prospectus Directive (each, a relevant member state), with effect from and including the date on which the Prospectus Directive is implemented in that relevant member state (the relevant implementation date), an offer of shares described in this prospectus may not be made to the public in that relevant member state other than:

to any legal entity which is a qualified investor as defined in the Prospectus Directive;

to fewer than 100 or, if the relevant member state has implemented the relevant provision of the 2010 PD Amending Directive, 150 natural or legal persons (other than qualified investors as defined in the Prospectus Directive), as permitted under the Prospectus Directive, subject to obtaining the prior consent of the relevant Dealer or Dealers nominated by us for any such offer; or

in any other circumstances falling within Article 3(2) of the Prospectus Directive,

provided that no such offer of shares shall require us or any underwriter to publish a prospectus pursuant to Article 3 of the Prospectus Directive.

For purposes of this provision, the expression an "offer of securities to the public" in any relevant member state means the communication in any form and by any means of sufficient information on the terms of the offer and the shares to be offered so as to enable an investor to decide to purchase or subscribe for the shares, as the expression may be varied in that member state by any measure implementing the Prospectus Directive in that member state, and the expression "Prospectus Directive" means Directive 2003/71/EC (and amendments thereto, including the 2010 PD Amending Directive, to the extent implemented in the relevant member state) and includes any relevant implementing measure in the relevant member state. The expression 2010 PD Amending Directive means Directive 2010/73/EU.

The sellers of the shares have not authorized and do not authorize the making of any offer of shares through any financial intermediary on their behalf, other than offers made by the underwriters with a view to the final placement of the shares as contemplated in this prospectus. Accordingly, no purchaser of the shares, other than the underwriters, is authorized to make any further offer of the shares on behalf of the sellers or the underwriters.

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**Notice to Prospective Investors in the United Kingdom**

This prospectus is only being distributed to, and is only directed at, persons in the United Kingdom that are qualified investors within the meaning of Article 2(1)(e) of the Prospectus Directive that are also (i) investment professionals falling within Article 19(5) of the Financial Services and Markets Act 2000 (Financial Promotion) Order 2005 (the Order) or (ii) high net worth entities, and other persons to whom it may lawfully be communicated, falling within Article 49(2)(a) to (d) of the Order (each such person being referred to as a "relevant person"). This prospectus and its contents are confidential and should not be distributed, published or reproduced (in whole or in part) or disclosed by recipients to any other persons in the United Kingdom. Any person in the United Kingdom that is not a relevant person should not act or rely on this document or any of its contents.

**Notice to Prospective Investors in France**

Neither this prospectus nor any other offering material relating to the shares described in this prospectus has been submitted to the clearance procedures of the *Autorité des Marchés Financiers* or of the competent authority of another member state of the European Economic Area and notified to the *Autorité des Marchés Financiers*. The shares have not been offered or sold and will not be offered or sold, directly or indirectly, to the public in France. Neither this prospectus nor any other offering material relating to the shares has been or will be:

released, issued, distributed or caused to be released, issued or distributed to the public in France; or

used in connection with any offer for subscription or sale of the shares to the public in France.

Such offers, sales and distributions will be made in France only:

to qualified investors (*investisseurs qualifiés*) and/or to a restricted circle of investors (*cercle restreint d'investisseurs*), in each case investing for their own account, all as defined in, and in accordance with articles L.411-2, D.411-1, D.411-2, D.734-1, D.744-1, D.754-1 and D.764-1 of the French *Code monétaire et financier*;

to investment services providers authorized to engage in portfolio management on behalf of third parties; or

in a transaction that, in accordance with article L.411-2-II-1<sup>o</sup>-or-2<sup>o</sup>-or 3<sup>o</sup> of the French *Code monétaire et financier* and article 211-2 of the General Regulations (*Règlement Général*) of the *Autorité des Marchés Financiers*, does not constitute a public offer (*appel public à l'épargne*).

The shares may be resold directly or indirectly, only in compliance with articles L.411-1, L.411-2, L.412-1 and L.621-8 through L.621-8-3 of the French *Code monétaire et financier*.

**Notice to Prospective Investors in Australia**

No prospectus or other disclosure document (as defined in the Corporations Act 2001 (Cth) of Australia (Corporations Act)) in relation to the common stock has been or will be lodged with the Australian Securities & Investments Commission (ASIC). This document has not been lodged with ASIC and is only directed to certain categories of exempt persons. Accordingly, if you receive this document in Australia:

- (a) you confirm and warrant that you are either:
  - (i) a "sophisticated investor" under section 708(8)(a) or (b) of the Corporations Act;
  - (ii) a "sophisticated investor" under section 708(8)(c) or (d) of the Corporations Act and that you have provided an accountant's certificate to us which complies with the requirements





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of section 708(8)(c)(i) or (ii) of the Corporations Act and related regulations before the offer has been made;

(iii)

a person associated with the company under section 708(12) of the Corporations Act; or

(iv)

a "professional investor" within the meaning of section 708(11)(a) or (b) of the Corporations Act, and to the extent that you are unable to confirm or warrant that you are an exempt sophisticated investor, associated person or professional investor under the Corporations Act any offer made to you under this document is void and incapable of acceptance; and

(b)

you warrant and agree that you will not offer any of the common stock for resale in Australia within 12 months of that common stock being issued unless any such resale offer is exempt from the requirement to issue a disclosure document under section 708 of the Corporations Act.

**Notice to Prospective Investors in Hong Kong**

The shares may not be offered or sold in Hong Kong by means of any document other than (i) in circumstances which do not constitute an offer to the public within the meaning of the Companies Ordinance (Cap. 32, Laws of Hong Kong), or (ii) to "professional investors" within the meaning of the Securities and Futures Ordinance (Cap. 571, Laws of Hong Kong) and any rules made thereunder, or (iii) in other circumstances which do not result in the document being a "prospectus" within the meaning of the Companies Ordinance (Cap. 32, Laws of Hong Kong) and no advertisement, invitation or document relating to the shares may be issued or may be in the possession of any person for the purpose of issue (in each case whether in Hong Kong or elsewhere), which is directed at, or the contents of which are likely to be accessed or read by, the public in Hong Kong (except if permitted to do so under the laws of Hong Kong) other than with respect to shares which are or are intended to be disposed of only to persons outside Hong Kong or only to "professional investors" within the meaning of the Securities and Futures Ordinance (Cap. 571, Laws of Hong Kong) and any rules made thereunder.

**Notice to Prospective Investors in Japan**

The shares offered in this prospectus have not been and will not be registered under the Financial Instruments and Exchange Law of Japan. The shares have not been offered or sold and will not be offered or sold, directly or indirectly, in Japan or to or for the account of any resident of Japan (including any corporation or other entity organized under the laws of Japan), except (i) pursuant to an exemption from the registration requirements of the Financial Instruments and Exchange Law and (ii) in compliance with any other applicable requirements of Japanese law.

**Notice to Prospective Investors in Singapore**

This prospectus has not been registered as a prospectus with the Monetary Authority of Singapore. Accordingly, this prospectus and any other document or material in connection with the offer or sale, or invitation for subscription or purchase, of the shares may not be circulated or distributed, nor may the shares be offered or sold, or be made the subject of an invitation for subscription or purchase, whether directly or indirectly, to persons in Singapore other than (i) to an institutional investor under Section 274 of the Securities and Futures Act, Chapter 289 of Singapore (the SFA), (ii) to a relevant person pursuant to Section 275(1), or any person pursuant to Section 275(1A), and in accordance with the conditions specified in Section 275 of the SFA or (iii) otherwise pursuant to, and in accordance with the conditions of, any other applicable provision of the SFA, in each case subject to compliance with conditions set forth in the SFA.

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Where the shares are subscribed or purchased under Section 275 of the SFA by a relevant person which is:

a corporation (which is not an accredited investor (as defined in Section 4A of the SFA)) the sole business of which is to hold investments and the entire share capital of which is owned by one or more individuals, each of whom is an accredited investor; or

a trust (where the trustee is not an accredited investor) whose sole purpose is to hold investments and each beneficiary of the trust is an individual who is an accredited investor,

shares, debentures and units of shares and debentures of that corporation or the beneficiaries' rights and interest (howsoever described) in that trust shall not be transferred within six months after that corporation or that trust has acquired the shares pursuant to an offer made under Section 275 of the SFA except:

to an institutional investor (for corporations, under Section 274 of the SFA) or to a relevant person defined in Section 275(2) of the SFA, or to any person pursuant to an offer that is made on terms that such shares, debentures and units of shares and debentures of that corporation or such rights and interest in that trust are acquired at a consideration of not less than \$0.2 million (or its equivalent in a foreign currency) for each transaction, whether such amount is to be paid for in cash or by exchange of securities or other assets, and further for corporations, in accordance with the conditions specified in Section 275 of the SFA;

where no consideration is or will be given for the transfer; or

where the transfer is by operation of law.

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**LEGAL MATTERS**

The validity of the issuance of our common stock offered in this prospectus will be passed upon for us by Ropes & Gray LLP, Boston, Massachusetts. Certain legal matters in connection with this offering will be passed upon for the underwriters by Mintz, Levin, Cohn, Ferris, Glovsky and Popeo, P.C., Boston, Massachusetts.

**EXPERTS**

The financial statements of Acceleron Pharma Inc. at December 31, 2011 and 2012, and for the years then ended, appearing in this Prospectus and Registration Statement have been audited by Ernst & Young LLP, independent registered public accounting firm, as set forth in their report thereon appearing elsewhere herein, and are included in reliance upon such report given on the authority of such firm as experts in accounting and auditing.

**WHERE YOU CAN FIND MORE INFORMATION**

We have filed with the SEC a registration statement on Form S-1 under the Securities Act with respect to the shares of common stock offered hereby. This prospectus, which constitutes a part of the registration statement, does not contain all of the information set forth in the registration statement or the exhibits and schedules filed therewith. For further information with respect to us and the common stock offered hereby, reference is made to the registration statement and the exhibits and schedules filed therewith. Statements contained in this prospectus regarding the contents of any contract or any other document that is filed as an exhibit to the registration statement are not necessarily complete, and each such statement is qualified in all respects by reference to the full text of such contract or other document filed as an exhibit to the registration statement. A copy of the registration statement and the exhibits and schedules filed therewith may be inspected without charge at the public reference room maintained by the SEC, located at 100 F Street N.E., Washington, D.C. 20549, and copies of all or any part of the registration statement may be obtained from such offices upon the payment of the fees prescribed by the SEC. Please call the SEC at 1-800-SEC-0330 for further information about the public reference room. The SEC also maintains a website that contains reports, proxy and information statements and other information regarding registrants that file electronically with the SEC. The address is [www.sec.gov](http://www.sec.gov).

We are subject to the information and periodic reporting requirements of the Exchange Act and, in accordance therewith, file periodic reports, proxy statements and other information with the SEC. Such periodic reports, proxy statements and other information are available for inspection and copying at the public reference room and website of the SEC referred to above.

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**Acceleron Pharma Inc.**

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<u>Statements of Operations and Comprehensive Income (Loss) for the Years Ended December 31, 2011 and 2012</u>	<u>F-4</u>
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**Report of Independent Registered Public Accounting Firm**

The Board of Directors and Stockholders of  
Acceleron Pharma Inc.

We have audited the accompanying balance sheets of Acceleron Pharma Inc. (the Company) as of December 31, 2011 and 2012, and the related statements of operations and comprehensive income (loss), redeemable convertible preferred stock and stockholders' deficit, and cash flows for the years then ended. These financial statements are the responsibility of the Company's management. Our responsibility is to express an opinion on these financial statements based on our audits.

We conducted our audits in accordance with the standards of the Public Company Accounting Oversight Board (United States). Those standards require that we plan and perform the audit to obtain reasonable assurance about whether the financial statements are free of material misstatement. We were not engaged to perform an audit of the Company's internal control over financial reporting. Our audits included consideration of internal control over financial reporting as a basis for designing audit procedures that are appropriate in the circumstances, but not for the purpose of expressing an opinion on the effectiveness of the Company's internal control over financial reporting. Accordingly, we express no such opinion. An audit also includes examining, on a test basis, evidence supporting the amounts and disclosures in the financial statements, assessing the accounting principles used and significant estimates made by management, and evaluating the overall financial statement presentation. We believe that our audits provide a reasonable basis for our opinion.

In our opinion, the financial statements referred to above present fairly, in all material respects, the financial position of Acceleron Pharma Inc. at December 31, 2011 and 2012, and the results of its operations and its cash flows for the years then ended, in conformity with U.S. generally accepted accounting principles.

/s/ Ernst & Young LLP

Boston, Massachusetts  
July 3, 2013, except for Note 16,  
as to which the date is  
September 5, 2013

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## Acceleron Pharma Inc.

## Balance Sheets

(amounts in thousands except share and per share data)

	December 31,	
	2011	2012
<b>Assets</b>		
Current assets:		
Cash and cash equivalents	\$ 65,037	\$ 39,611
Collaboration receivables (includes related party amounts of \$1,024 and \$1,840 at December 31, 2011 and 2012, respectively)	1,660	2,776
Related party receivable		
Prepaid expenses and other current assets	1,044	1,474
<b>Total current assets</b>	<b>67,741</b>	<b>43,861</b>
Property and equipment, net	4,911	4,059
Restricted cash	912	913
Related party receivables	225	233
Other assets		146
<b>Total assets</b>	<b>\$ 73,789</b>	<b>\$ 49,212</b>
<b>Liabilities, redeemable convertible preferred stock and stockholders' deficit</b>		
Current liabilities:		
Accounts payable	\$ 1,914	\$ 642
Accrued expenses (includes related party amounts of \$833 and \$861 at December 31, 2011 and 2012, respectively)	4,513	6,153
Deferred revenue	10,946	27,840
Deferred rent	483	499
Notes payable, net of discount	5,997	3,668
<b>Total current liabilities</b>	<b>23,853</b>	<b>38,802</b>
Deferred revenue, net of current portion	33,350	6,760
Deferred rent, net of current portion	3,335	2,837
Notes payable, net of current portion and discount		16,525
Warrants to purchase redeemable convertible preferred stock	1,046	1,422
Warrants to purchase common stock	3,347	5,229
<b>Total liabilities</b>	<b>64,931</b>	<b>71,575</b>
Commitments and contingencies (Note 7)		
Redeemable convertible preferred stock (Note 8)	241,549	268,610
Stockholders' deficit:		
Common stock, \$0.001 par value: 104,013,161 shares authorized; 2,393,458 and 2,432,155 shares issued and outstanding at December 31, 2011 and 2012, respectively	3	3
Additional paid-in capital		
Accumulated deficit	(232,694)	(290,976)
<b>Total stockholders' deficit</b>	<b>(232,691)</b>	<b>(290,973)</b>
<b>Total liabilities, redeemable convertible preferred stock and stockholders' deficit</b>	<b>\$ 73,789</b>	<b>\$ 49,212</b>

See accompanying notes to financial statements.

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## Acceleron Pharma Inc.

## Statements of Operations and Comprehensive Income (Loss)

(amounts in thousands except per share data)

	Year Ended December 31,	
	2011	2012
<b>Revenue:</b>		
Collaboration revenue:		
License and milestone	\$ 74,406	\$ 9,696
Cost-sharing, net	4,760	5,558
Contract manufacturing	1,745	
Total revenue <sup>(1)</sup>	80,911	15,254
<b>Costs and expenses:</b>		
Research and development	32,713	35,319
General and administrative	8,142	8,824
Cost of contract manufacturing revenue	1,500	
Total costs and expenses	42,355	44,143
Income (loss) from operations	38,556	(28,889)
<b>Other (expense) income:</b>		
Other expense, net	(485)	(2,255)
Interest income	17	91
Interest expense	(1,822)	(1,529)
Total other expense, net	(2,290)	(3,693)
Net income (loss)	\$ 36,266	\$ (32,582)
Comprehensive income (loss)	\$ 36,266	\$ (32,582)
<b>Reconciliation of net income (loss) to net income (loss) applicable to common stockholders:</b>		
Net income (loss)	\$ 36,266	\$ (32,582)
Accretion of dividends, interest, redemption value and issuance costs on redeemable convertible preferred stock	(21,757)	(27,061)
Net income (loss) applicable to participating securities	(12,645)	
Net income (loss) applicable to common stockholders basic	\$ 1,864	\$ (59,643)
Net income (loss)	\$ 36,266	\$ (32,582)
Accretion of dividends, interest, redemption value and issuance costs on redeemable convertible preferred stock	(21,757)	(27,061)
Net income (loss) applicable to participating securities	(12,395)	
Net income (loss) applicable to common stockholders diluted	\$ 2,114	\$ (59,643)
<b>Net income (loss) per share applicable to common stockholders: (Note 2)</b>		
Basic	\$ 0.80	\$ (24.84)
Diluted	\$ 0.78	\$ (24.84)



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Weighted-average number of common shares used in computing net income (loss) per share applicable to common stockholders:

Basic	2,328	2,401
Diluted	2,716	2,401

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(1) Includes related party revenue (Note 15) \$ 64,220 \$ 4,914

See accompanying notes to financial statements.

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**Acceleron Pharma Inc.**  
**Statements of Redeemable Convertible Preferred Stock and Stockholders' Deficit**  
(amounts in thousands except share data)

	Series A Redeemable Convertible Preferred Stock		Series B Redeemable Convertible Preferred Stock		Series C Redeemable Convertible Preferred Stock		Series C-1 Redeemable Convertible Preferred Stock		Series D Redeemable Convertible Preferred Stock		Series D-1 Redeemable Convertible Preferred Stock	
	Number of		Number of		Number of		Number of		Number of		Number of	
	Shares	Value	Shares	Value	Shares	Value	Shares	Value	Shares	Value	Shares	Value
Balance at December 31, 2010	6,410,976	\$ 57,433	4,204,185	\$ 55,880	2,978,062	\$ 48,726	457,875	\$ 7,571	234,940	\$ 2,989	636,942	\$ 8,392
Sale of Series F redeemable convertible preferred stock net of issuance costs of \$92												
Accretion of dividends, interest, redemption value and issuance costs related to redeemable convertible preferred stock		4,616		5,584		5,594		908		668		1,736
Compensation expense associated with stock options												
Grant of stock options to nonemployees												
Exercise of stock options												
Exercise of common warrants												
Net loss												
Balance at December 31, 2011	6,410,976	62,049	4,204,185	61,464	2,978,062	54,320	457,875	8,479	234,940	3,657	636,942	10,128
Accretion of dividends, interest, redemption value and issuance costs related to redeemable convertible preferred stock		4,616		5,580		5,589		908		668		1,736
Compensation expense associated with stock options												
Exercise of stock options												
Net loss												
Balance at December 31, 2012	6,410,976	\$ 66,665	4,204,185	\$ 67,044	2,978,062	\$ 59,909	457,875	\$ 9,387	234,940	\$ 4,325	636,942	\$ 11,864

See accompanying notes to financial statements.

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**Acceleron Pharma Inc.**  
**Statements of Redeemable Convertible Preferred Stock and Stockholders' Deficit (continued)**  
(amounts in thousands except share data)

	Series E Redeemable Convertible Preferred Stock		Series F Redeemable Convertible Preferred Stock		Total Redeemable Convertible Preferred Stock	Common Stock			Total Stockholders' Deficit	
	Number of Shares	Value	Number of Shares	Value		Number of Shares	Par Value	Additional Paid-In Capital		Accumulated Deficit
Balance at December 31, 2010	816,060	\$ 8,423			\$ 189,414	2,261,461	\$ 3	\$	\$ (248,820)	\$ (248,817)
Sale of Series F redeemable convertible preferred stock net of issuance costs of \$92			2,426,171	30,378	30,378					
Accretion of dividends, interest, redemption value and issuance costs related to redeemable convertible preferred stock		2,511		140	21,757			(1,617)	(20,140)	(21,757)
Compensation expense associated with stock options								1,212		1,212
Grant of stock options to nonemployees								215		215
Exercise of stock options						94,748		190		190
Exercise of common warrants						37,249				
Net loss									36,266	36,266
Balance at December 31, 2011	816,060	10,934	2,426,171	30,518	241,549	2,393,458	3		(232,694)	(232,691)
Accretion of dividends, interest, redemption value and issuance costs related to redeemable convertible preferred stock		2,459		5,505	27,061			(1,361)	(25,700)	(27,061)
Compensation expense associated with stock options								1,206		1,206
Exercise of stock options						38,697		155		155
Net loss									(32,582)	(32,582)
Balance at December 31, 2012	816,060	\$ 13,393	2,426,171	\$ 36,023	\$ 268,610	2,432,155	\$ 3	\$	\$ (290,976)	\$ (290,973)

See accompanying notes to financial statements.

Table of Contents**Acceleron Pharma Inc.****Statements of Cash Flows****(amounts in thousands)**

	<b>Year Ended December 31,</b>	
	<b>2011</b>	<b>2012</b>
<b>Operating Activities</b>		
Net income (loss)	\$ 36,266	\$ (32,582)
Adjustments to reconcile net income (loss) to net cash provided by (used in) operating activities:		
Depreciation and amortization	3,134	1,293
Stock-based compensation	1,427	1,206
Amortization of debt discount	162	51
Accretion of deferred interest	271	335
Amortization of deferred debt issuance costs	79	84
Change in fair value of warrants	481	2,258
Changes in assets and liabilities:		
Prepaid expenses and other current assets	2,564	(594)
Collaboration receivables	1,836	(1,116)
Related party receivable	(6)	(8)
Accounts payable	334	(1,272)
Accrued expenses	(2,773)	1,640
Deferred revenue	(35,130)	(9,696)
Deferred rent	211	(482)
Restricted cash	200	(1)
Net cash provided by (used in) operating activities	9,056	(38,884)
<b>Investing Activities</b>		
Purchases of property and equipment	(27)	(441)
Net cash used in investing activities	(27)	(441)
<b>Financing Activities</b>		
Proceeds from issuance of redeemable convertible preferred stock, net of issuance costs	30,378	
Proceeds from long-term debt, net of issuance costs		19,935
Payments of long-term debt	(9,476)	(6,191)
Proceeds from exercise of stock options and warrants to purchase common stock	190	155
Net cash provided by financing activities	21,092	13,899
Net increase (decrease) in cash and cash equivalents	30,121	(25,426)
Cash and cash equivalents at beginning of year	34,916	65,037
Cash and cash equivalents at end of year	\$ 65,037	\$ 39,611
<b>Supplemental Disclosure of Cash Flow Information:</b>		
Cash paid for interest	\$ 1,461	\$ 1,065
<b>Supplemental Disclosure of Non-Cash Investing and Financing Activities:</b>		
Accretion of dividends, interest, redemption value, and issuance costs on preferred stock	\$ 21,757	\$ 27,061

See accompanying notes to financial statements.



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**Acceleron Pharma Inc.**

**Notes to Financial Statements**

**Years Ended December 31, 2011 and 2012**

**1. Nature of Business**

Acceleron Pharma Inc. (Acceleron or the Company) was incorporated in the state of Delaware on June 13, 2003, as Phoenix Pharma, Inc. The Company subsequently changed its name to Acceleron Pharma Inc. and commenced operations in February 2004. The Company is a Cambridge, Massachusetts-based biopharmaceutical company focused on the discovery, development and commercialization of novel protein therapeutics for cancer and rare diseases. The Company's research focuses on the biology of the Transforming Growth Factor-Beta (TGF- $\beta$ ) protein superfamily, a large and diverse group of molecules that regulate the growth and repair of tissues throughout the human body. By coupling its discovery and development expertise, including its proprietary knowledge of the TGF- $\beta$  superfamily, with internal protein engineering and manufacturing capabilities, the Company has built a highly productive research and development platform that has generated numerous innovative protein therapeutics with novel mechanisms of action. The Company has internally discovered three protein therapeutics that are currently being studied in 12 ongoing Phase 2 clinical trials, focused on the areas of cancer and rare diseases.

The Company is subject to risks common to companies in the biotechnology industry, including, but not limited to, risk that the Company never achieves profitability, the need for substantial additional financing, risk of relying on third parties, risks of clinical trial failures, dependence on key personnel, protection of proprietary technology and compliance with government regulations.

***Liquidity***

As of December 31, 2012, the Company had an accumulated deficit of \$291.0 million, and will require substantial additional capital to fund its research and development. The Company believes that its cash resources of \$39.6 million at December 31, 2012 will be sufficient to allow the Company to fund its current operating plan through January 1, 2014; however, the Company will be required to raise additional capital to fund operations beyond this time. As the Company continues to incur losses, a transition to profitability is dependent upon the successful development, approval and commercialization of its product candidates and the achievement of a level of revenues adequate to support the Company's cost structure. The Company may never achieve profitability, and unless and until it does, the Company will continue to need to raise additional capital. Management intends to fund future operations through the sale of equity, debt financings or other sources, including potential additional collaborations. There can be no assurances, however, that additional funding will be available on terms acceptable to the Company, or at all.

**2. Summary of Significant Accounting Policies**

The accompanying financial statements reflect the application of certain significant accounting policies as described below and elsewhere in these notes to the financial statements. The Company believes that a significant accounting policy is one that is both important to the portrayal of the Company's financial condition and results, and requires management's most difficult, subjective, or complex judgments, often as the result of the need to make estimates about the effect of matters that are inherently uncertain.

***Basis of Presentation***

The accompanying financial statements have been prepared in conformity with accounting principles generally accepted in the United States of America (GAAP). Any reference in these notes to applicable guidance is meant to refer to the authoritative United States generally accepted accounting

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**2. Summary of Significant Accounting Policies (continued)**

principles as found in the Accounting Standards Codification (ASC) and Accounting Standards Update (ASU) of the Financial Accounting Standards Board (FASB).

*Use of Estimates*

The preparation of financial statements in conformity with GAAP requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities, the disclosure of contingent assets and liabilities at the date of the financial statements, and the reported amounts expensed during the reporting period. Actual results could materially differ from those estimates.

Management considers many factors in selecting appropriate financial accounting policies and controls, and in developing the estimates and assumptions that are used in the preparation of these financial statements. Management must apply significant judgment in this process. In addition, other factors may affect estimates, including: expected business and operational changes, sensitivity and volatility associated with the assumptions used in developing estimates, and whether historical trends are expected to be representative of future trends. The estimation process often may yield a range of potentially reasonable estimates of the ultimate future outcomes and management must select an amount that falls within that range of reasonable estimates. This process may result in actual results differing materially from those estimated amounts used in the preparation of the financial statements if these results differ from historical experience, or other assumptions do not turn out to be substantially accurate, even if such assumptions are reasonable when made. In preparing these financial statements, management used significant estimates in the following areas, among others: revenue recognition, stock-based compensation expense, the determination of the fair value of stock-based awards, the fair value of liability-classified warrants, accrued expenses, and the recoverability of the Company's net deferred tax assets and related valuation allowance.

The Company utilizes significant estimates and assumptions in determining the fair value of its common stock. The Company's board of directors (the Board) determined the estimated fair value of the Company's common stock based on a number of objective and subjective factors, including external market conditions affecting the biotechnology industry sector and the prices at which the Company sold shares of redeemable convertible preferred stock, the superior rights and preferences of securities senior to the Company's common stock at the time, and the likelihood of achieving a liquidity event, such as an initial public offering (IPO) or sale of the Company.

The Company utilized various valuation methodologies in accordance with the framework of the American Institute of Certified Public Accountants' Technical Practice Aid, *Valuation of Privately-Held Company Equity Securities Issued as Compensation*, to estimate the fair value of its common stock. Each valuation methodology includes estimates and assumptions that require the Company's judgment. These estimates and assumptions include a number of objective and subjective factors, including external market conditions affecting the biotechnology industry sector, the prices at which the Company sold shares of preferred stock, the superior rights and preferences of securities senior to the Common Stock at the time and the likelihood of achieving a liquidity event, such as an IPO or sale. Significant changes to the key assumptions used in the valuations could result in different fair values of common stock at each valuation date.

*Reclassifications*

The Company has reclassified certain prior period amounts in the balance sheet as of December 31, 2011, totaling \$0.5 million related to deferred rent from long-term to short-term to

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**2. Summary of Significant Accounting Policies (continued)**

conform to the current period presentation. This reclassification had no impact on the previously reported results of operations or cash flows for the year ended December 31, 2011.

***Collaboration Receivable***

Credit is extended to customers based upon an evaluation of the customer's financial condition. Collaboration receivables are recorded at net realizable value. The Company does not charge interest on past due balances. Collaboration receivables are determined to be past due when the payment due date is exceeded. The Company utilizes a specific identification accounts receivable reserve methodology based on a review of outstanding balances and previous activities to determine the allowance for doubtful accounts. The Company charges off uncollectible receivables at the time the Company determines the receivable is no longer collectible. The Company did not have an allowance for doubtful accounts at December 31, 2011 or 2012.

***Segment Information***

Operating segments are identified as components of an enterprise about which separate discrete financial information is available for evaluation by the chief operating decision maker, or decision making group, in making decisions on how to allocate resources and assess performance. The Company's chief operating decision maker is the chief executive officer. The Company and the chief executive officer view the Company's operations and manage its business as one operating segment. All material long-lived assets of the Company reside in the United States. The Company does use contract research organizations (CROs) and research institutions located outside the United States. Some of these expenses are subject to collaboration reimbursement which is presented as a component of cost sharing, net in the statement of operations and comprehensive income (loss).

***Cash and Cash Equivalents***

The Company considers all highly liquid investments purchased with original maturities of 90 days or less at acquisition to be cash equivalents. Cash and cash equivalents include cash held in banks and amounts held in interest-bearing money market accounts. Cash equivalents are carried at cost, which approximates their fair market value.

***Concentrations of Credit Risk and Off-Balance Sheet Risk***

The Company has no off-balance sheet risk, such as foreign exchange contracts, option contracts, or other foreign hedging arrangements. Financial instruments that potentially subject the Company to concentrations of credit risk are primarily cash, cash equivalents, restricted cash and accounts receivable. The Company maintains its cash and cash equivalent balances in the form of money market accounts with financial institutions that management believes are creditworthy. The Company's investment policy includes guidelines on the quality of the institutions and financial instruments and defines allowable investments that the Company believes minimizes the exposure to concentration of credit risk.

The Company routinely assesses the creditworthiness of its customers and collaboration partners. The Company has not experienced any material losses related to receivables from individual customers and collaboration partners, or groups of customers. The Company does not require collateral. Due to these factors, no additional credit risk beyond amounts provided for collection losses is believed by management to be probable in the Company's accounts receivable.



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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**2. Summary of Significant Accounting Policies (continued)**

*Deferred IPO Issuance Costs*

Deferred issuance costs, which primarily consist of direct incremental legal and accounting fees relating to the IPO, are capitalized. The deferred issuance costs will be offset against IPO proceeds upon the consummation of the offering. In the event the offering is terminated, or delayed more than 90 days, deferred offering costs will be expensed. No amounts were deferred as of December 31, 2011 or 2012.

*Disclosure of Fair Value of Financial Instruments*

The carrying amounts of the Company's financial instruments, which include cash, cash equivalents, collaboration receivables, accounts payable, accrued expenses and notes payable, approximated their fair values at December 31, 2011 and 2012, due to the short-term nature of these instruments, and for the notes payable, the interest rates the Company believes it could obtain for borrowings with similar terms. See discussion below on the determination of the fair value of the Company's preferred and common stock warrants.

The Company has evaluated the estimated fair value of financial instruments using available market information and management's estimates. The use of different market assumptions and/or estimation methodologies could have a significant effect on the estimated fair value amounts.

*Fair Value Measurements*

ASC Topic 820, *Fair Value Measurement* (ASC 820), establishes a fair value hierarchy for instruments measured at fair value that distinguishes between assumptions based on market data (observable inputs) and the Company's own assumptions (unobservable inputs). Observable inputs are inputs that market participants would use in pricing the asset or liability based on market data obtained from sources independent of the Company. Unobservable inputs are inputs that reflect the Company's assumptions about the inputs that market participants would use in pricing the asset or liability, and are developed based on the best information available in the circumstances.

ASC 820 identifies fair value as the exchange price, or exit price, representing the amount that would be received to sell an asset or paid to transfer a liability in an orderly transaction between market participants. As a basis for considering market participant assumptions in fair value measurements, ASC Topic 820 establishes a three-tier fair value hierarchy that distinguishes between the following:

Level 1 Quoted market prices in active markets for identical assets or liabilities.

Level 2 Inputs other than Level 1 inputs that are either directly or indirectly observable, such as quoted market prices, interest rates, and yield curves.

Level 3 Unobservable inputs developed using estimates of assumptions developed by the Company, which reflect those that a market participant would use.

To the extent that the valuation is based on models or inputs that are less observable or unobservable in the market, the determination of fair value requires more judgment. Accordingly, the degree of judgment exercised by the Company in determining fair value is greatest for instruments categorized in Level 3. A financial instrument's level within the fair value hierarchy is based on the lowest level of any input that is significant to the fair value measurement.

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****2. Summary of Significant Accounting Policies (continued)**

Items measured at fair value on a recurring basis include warrants to purchase redeemable convertible preferred stock and warrants to purchase common stock (Note 6). During the periods presented, the Company has not changed the manner in which it values assets and liabilities that are measured at fair value using Level 3 inputs.

The following tables set forth the Company's financial instruments carried at fair value using the lowest level of input applicable to each financial instrument as of December 31, 2011 and 2012 (in thousands):

	Quoted Prices in Active Markets for Identical Items (Level 1)	December 31, 2011 Significant Other Observable Inputs (Level 2)	December 31, 2011 Significant Unobservable Inputs (Level 3)	Total
<b>Assets:</b>				
Money market funds	\$ 61,269	\$	\$	\$ 61,269
Restricted cash	912			912
<b>Total assets</b>	<b>\$ 62,181</b>	<b>\$</b>	<b>\$</b>	<b>\$ 62,181</b>
<b>Liabilities:</b>				
Warrants to purchase redeemable convertible preferred stock	\$	\$	\$ 1,046	\$ 1,046
Warrants to purchase common stock			3,347	3,347
<b>Total liabilities</b>	<b>\$</b>	<b>\$</b>	<b>\$ 4,393</b>	<b>\$ 4,393</b>

	Quoted Prices in Active Markets for Identical Items (Level 1)	December 31, 2012 Significant other Observable Inputs (Level 2)	December 31, 2012 Significant Unobservable Inputs (Level 3)	Total
<b>Assets:</b>				
Money market funds	\$ 36,847	\$	\$	\$ 36,847
Restricted cash	913			913
<b>Total assets</b>	<b>\$ 37,760</b>	<b>\$</b>	<b>\$</b>	<b>\$ 37,760</b>
<b>Liabilities:</b>				
Warrants to purchase redeemable convertible preferred stock	\$	\$	\$ 1,422	\$ 1,422
Warrants to purchase common stock			5,229	5,229
<b>Total liabilities</b>	<b>\$</b>	<b>\$</b>	<b>\$ 6,651</b>	<b>\$ 6,651</b>

The following table sets forth a summary of changes in the fair value of the Company's preferred and common stock warrant liability, which represents a recurring measurement that is classified within

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****2. Summary of Significant Accounting Policies (continued)**

Level 3 of the fair value hierarchy, wherein fair value is estimated using significant unobservable inputs (in thousands):

	Year Ended December 31,	
	2011	2012
Beginning balance	\$ 3,912	\$ 4,393
Change in fair value	481	2,258
Exercises		
Repurchases		
Ending balance	\$ 4,393	\$ 6,651

The money market funds noted above are included in cash and cash equivalents in the accompanying balance sheets. The Company recognizes transfers between levels of the fair value hierarchy as of the end of the reporting period. There were no transfers within the hierarchy during the years ended December 31, 2011 or 2012.

The fair value of the warrants on the date of issuance and on each re-measurement date for those warrants classified as liabilities is estimated using the Black-Scholes option pricing model. This method of valuation involves using inputs such as the fair value of the Company's various classes of preferred stock, stock price volatility, the contractual term of the warrants, risk free interest rates, and dividend yields. Due to the nature of these inputs, the valuation of the warrants is considered a Level 3 measurement. See Note 6 for further discussions of the accounting for the warrants, as well as for a summary of the significant inputs and assumptions used to determine the fair value of the warrants.

The Company measures eligible assets and liabilities at fair value, with changes in value recognized in earnings. Fair value treatment may be elected either upon initial recognition of an eligible asset or liability or, for an existing asset or liability, if an event triggers a new basis of accounting. The Company did not elect to remeasure any of its existing financial assets or liabilities, and did not elect the fair value option for any financial assets and liabilities transacted in the years ended December 31, 2011 or 2012.

***Property and Equipment***

Property and equipment is stated at cost. Maintenance and repairs that do not improve or extend the lives of the respective assets are expensed to operations as incurred. Upon disposal, retirement or sale the related cost and accumulated depreciation is removed from the accounts and any resulting gain or loss is included in the results of operations. Depreciation and amortization is calculated using the straight-line method over the estimated useful lives of the assets, which are as follows:

Asset	Estimated Useful Life
Computer equipment and software	3 years
Office and laboratory equipment	3 years
Leasehold improvements	Shorter of the useful life or remaining lease term

The Company reviews long-lived assets when events or changes in circumstances indicate the carrying value of the assets may not be recoverable. Recoverability is measured by comparison of the

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**2. Summary of Significant Accounting Policies (continued)**

book values of the assets to future net undiscounted cash flows that the assets are expected to generate. If such assets are considered to be impaired, the impairment to be recognized is measured by the amount by which the book value of the assets exceed their fair value, which is measured based on the projected discounted future net cash flows arising from the assets. No impairment losses have been recorded during the years ended December 31, 2011 or 2012.

***Revenue Recognition***

The company has primarily generated revenue through collaboration, license and research arrangements with collaboration partners for the development and commercialization of protein therapeutics.

The Company recognizes revenue in accordance with FASB ASC Topic 605, *Revenue Recognition*. Accordingly, revenue is recognized for each unit of accounting when all of the following criteria are met: (1) persuasive evidence of an arrangement exists; (2) delivery has occurred or services have been rendered; (3) the fee is fixed or determinable; and (4) collectability is reasonably assured.

Amounts received prior to satisfying the revenue recognition criteria are recorded as deferred revenue in the Company's balance sheets. Amounts expected to be recognized as revenue within the 12 months following the balance sheet date are classified as deferred revenue, current portion. Amounts not expected to be recognized as revenue within the 12 months following the balance sheet date are classified as deferred revenue, net of current portion.

***Multiple Element Revenue Arrangements***

The Company enters into collaboration agreements from time to time, which are more fully described in Note 10. The arrangements generally contain multiple elements or deliverables, which may include (1) licenses, or options to obtain licenses, to the Company's technology, (2) research and development activities performed for the collaboration partner, (3) participation on Joint Development Committees, and (4) manufacturing clinical or preclinical material. Payments pursuant to these arrangements typically include non-refundable, up-front payments, milestone payments upon achieving significant development events, research and development reimbursements, sales milestones, and royalties on future product sales.

Effective January 1, 2011, the Company adopted ASU No. 2009-13, *Multiple-Deliverable Revenue Arrangements* (ASU 2009-13), which amends Topic 605-25, *Revenue Recognition - Multiple Element Arrangements* (ASC 605-25). The Company applies this guidance to new arrangements as well as existing agreements that are significantly modified after January 1, 2011. For agreements that are significantly modified, the Company determines the estimated selling price for the remaining undelivered elements as of the date of the material modification and allocates arrangement consideration based upon the estimated selling price to the undelivered elements.

The application of the multiple element guidance requires subjective determinations, and requires management to make judgments about the individual deliverables, and whether such deliverables are separable from the other aspects of the contractual relationship. Deliverables are considered separate units of accounting provided that: (1) the delivered item(s) has value to the customer on a stand-alone basis and (2) if the arrangement includes a general right of return relative to the delivered item(s),

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**2. Summary of Significant Accounting Policies (continued)**

delivery or performance of the undelivered item(s) is considered probable and substantially in the control of the Company. In determining the units of accounting, management evaluates certain criteria, including whether the deliverables have stand-alone value, based on the consideration of the relevant facts and circumstances for each arrangement, such as the research, manufacturing and commercialization capabilities of the collaboration partner and the availability of the associated expertise in the general marketplace. In addition, the Company considers whether the collaboration partner can use the other deliverable(s) for their intended purpose without the receipt of the remaining element(s), whether the value of the deliverable is dependent on the undelivered item(s) and whether there are other vendors that can provide the undelivered element(s). Arrangement consideration that is fixed or determinable is allocated among the separate units of accounting using the relative selling price method, and the applicable revenue recognition criteria, as described above, are applied to each of the separate units of accounting in determining the appropriate period or pattern of recognition.

The Company determines the estimated selling price for deliverables within each agreement using vendor-specific objective evidence (VSOE) of selling price, if available, third-party evidence (TPE) of selling price if VSOE is not available, or management's best estimate of selling price (BESP) if neither VSOE nor TPE is available. Subsequent to the adoption of ASU 2009-13, the Company typically uses BESP to estimate the selling price of the deliverables. Determining the BESP for a unit of accounting requires significant judgment. In developing the BESP for a unit of accounting, the Company considers applicable market conditions and relevant entity-specific factors, including factors that were contemplated in negotiating the agreement with the customer and estimated costs. The Company validates the BESP for units of accounting by evaluating whether changes in the key assumptions used to determine the BESP will have a significant effect on the allocation of arrangement consideration between multiple units of accounting.

The Company typically receives up-front, non-refundable payments when licensing its intellectual property in conjunction with a collaboration agreement. When management believes the license to its intellectual property does not have stand-alone value from the other deliverables to be provided in the arrangement, the Company generally recognizes revenue attributed to the license on a straight-line basis over the contractual or estimated performance period, which is typically the term of the Company's research and development or manufacturing obligations. The Company continually evaluates these periods, and will adjust the period of revenue recognition if circumstances change. When management believes the license to its intellectual property has stand-alone value, the Company generally recognizes revenue attributed to the license upon delivery.

Research and development funding is recognized as revenue in the period that the related services are performed. When the Company acts as the principal under its collaboration agreements, it records payments received for the reimbursement of research and development costs as cost-sharing revenue in the statements of operations and comprehensive income (loss). To the extent that the Company reimburses the collaborator for costs incurred, the Company records these costs as a reduction of cost-sharing revenue.

The Company's agreements may contain options which provide the collaboration partner the right to obtain additional licenses. Options are considered substantive if, at the inception of the arrangement, the Company is at risk as to whether the collaboration partner will choose to exercise the option. Factors considered in evaluating whether an option is substantive include the overall objective of the

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**2. Summary of Significant Accounting Policies (continued)**

arrangement, the benefit the collaborator might obtain from the arrangement without exercising the option, the cost to exercise the option and the likelihood that the option will be exercised. For arrangements under which an option is considered substantive, the Company does not consider the item underlying the option to be a deliverable at the inception of the arrangement and the associated option fees are not included in allocable arrangement consideration, assuming the option is not priced at a significant and incremental discount. Conversely, for arrangements under which an option is not considered substantive or if an option is priced at a significant and incremental discount, the Company would consider the item underlying the option to be a deliverable at the inception of the arrangement and a corresponding amount would be included in allocable arrangement consideration.

Effective January 1, 2011, the Company adopted ASU No. 2010-17, *Revenue Recognition Milestone Method* (ASU 2010-17). At the inception of each arrangement that includes milestone payments, the Company evaluates, with respect to each milestone, whether the milestone is substantive and at-risk. This evaluation includes an assessment of whether (a) the consideration is commensurate with either (1) the entity's performance to achieve the milestone, or (2) the enhancement of the value of the delivered item(s) as a result of a specific outcome resulting at least in part from the entity's performance to achieve the milestone, (b) the consideration relates solely to past performance, and (c) the consideration is reasonable relative to all of the deliverables and payment terms within the arrangement. The Company evaluates factors such as the scientific, regulatory, commercial, and other risks that must be overcome to achieve the respective milestone, the level of effort and investment required to achieve the respective milestone, and whether the milestone consideration is reasonable relative to all deliverables and payment terms in the arrangement in making this assessment. On the milestone achievement date, assuming all other revenue recognition criteria are met and the milestone is deemed substantive and at-risk, the Company recognizes the payment as license and milestone revenue. For milestones that are not deemed substantive and at-risk, where payment is reasonably assured, the Company recognizes the milestone payment over the remaining service period.

Sales and commercial milestones and royalties will be recognized when and if earned, provided collectability is reasonably assured.

***Contract Manufacturing Revenue***

Contract manufacturing revenue is recognized upon delivery of the product in accordance with the terms of the contract, which specifies when transfer of title and risk of loss occurs.

***Research and Development Expenses***

Research and development costs are charged to expense as costs are incurred in performing research and development activities. Research and development costs include all direct costs, including salaries, stock compensation and benefits for research and development personnel, outside consultants, costs of clinical trials, sponsored research, clinical trials insurance, other outside costs, depreciation and facility costs related to the development of drug candidates. The Company records up-front, non-refundable payments made to outside vendors, or other payments made in advance of services performed or goods being delivered, as prepaid expenses, which are expensed as services are performed or the goods are delivered.

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**2. Summary of Significant Accounting Policies (continued)**

Certain research and development projects are, or have been, partially funded by collaboration agreements, and the expenses related to these activities are included in research and development costs. The Company records the related reimbursement of research and development under these agreements as revenue, as more fully described above and in Note 10.

***Stock-Based Compensation***

At December 31, 2012, the Company had one stock-based compensation plan, which is more fully described in Note 11. The Company accounts for stock-based compensation in accordance with the provisions of ASC Topic 718, *Compensation - Stock Compensation* (ASC 718), which requires the recognition of expense related to the fair value of stock-based compensation awards in the statements of operations and comprehensive income (loss).

For stock options issued to employees and members of the Board for their services on the Board, the Company estimates the grant date fair value of each option using the Black-Scholes option-pricing model. The use of the Black-Scholes option-pricing model requires management to make assumptions with respect to the expected term of the option, the expected volatility of the common stock consistent with the expected life of the option, risk-free interest rates and expected dividend yields of the common stock. For awards subject to service-based vesting conditions, the Company recognizes stock-based compensation expense, net of estimated forfeitures, equal to the grant date fair value of stock options on a straight-line basis over the requisite service period, which is generally the vesting term. For awards subject to both performance and service-based vesting conditions, the Company recognizes stock-based compensation expense using an accelerated recognition method when it is probable that the performance condition will be achieved. Forfeitures are required to be estimated at the time of grant and revised, if necessary, in subsequent periods if actual forfeitures differ from those estimates.

Share-based payments issued to non-employees are recorded at their fair values, and are periodically revalued as the equity instruments vest and are recognized as expense over the related service period in accordance with the provisions of ASC 718 and ASC Topic 505, *Equity*. For stock-based awards granted to non-employees, the Company recognizes stock-based compensation expense using an accelerated recognition method.

See Note 11 for a discussion of the assumptions used by the Company in determining the grant date fair value of options granted under the Black-Scholes option pricing model, as well as a summary of the stock option activity under the Company's stock-based compensation plan for the year ended December 31, 2012.

***Income Taxes***

Income taxes are recorded in accordance with ASC Topic 740, *Income Taxes* (ASC 740), which provides for deferred taxes using an asset and liability approach. The Company recognizes deferred tax assets and liabilities for the expected future tax consequences of events that have been included in the financial statements or tax returns. Deferred tax assets and liabilities are determined based on the difference between the financial statement and tax bases of assets and liabilities using enacted tax rates in effect for the year in which the differences are expected to reverse. Valuation allowances are provided, if based upon the weight of available evidence, it is more likely than not that some or all of the deferred tax assets will not be realized.

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****2. Summary of Significant Accounting Policies (continued)**

The Company accounts for uncertain tax positions in accordance with the provisions of ASC 740. When uncertain tax positions exist, the Company recognizes the tax benefit of tax positions to the extent that the benefit will more likely than not be realized. The determination as to whether the tax benefit will more likely than not be realized is based upon the technical merits of the tax position as well as consideration of the available facts and circumstances. As of December 31, 2011 and 2012, the Company does not have any significant uncertain tax positions.

***Net Income (Loss) Per Share***

Net income (loss) per share information is determined using the two-class method, which includes the weighted-average number of common stock outstanding during the period and other securities that participate in dividends (a participating security). The Company's redeemable convertible preferred stock are participating securities as defined by ASC 260-10, *Earnings Per Share*.

Under the two-class method, basic net income (loss) per share applicable to common stockholders is computed by dividing the net income (loss) applicable to common stockholders by the weighted-average number of common shares outstanding for the reporting period. Diluted net income (loss) per share is computed using the more dilutive of (1) the two-class method or (2) the if-converted method. The Company allocates net income first to preferred stockholders based on dividend rights under the Company's articles of incorporation and then to preferred and common stockholders based on ownership interests. Net losses are not allocated to preferred stockholders as they do not have an obligation to share in the Company's net losses.

Diluted net income (loss) per share gives effect to all potentially dilutive securities, including redeemable convertible preferred stock, and shares issuable upon the exercise of outstanding warrants and stock options, using the treasury stock method. For the year ended December 31, 2012, the Company has excluded the effects of all potentially dilutive shares, which include redeemable convertible preferred stock, warrants for redeemable convertible preferred stock, warrants for common stock and outstanding common stock options, from the weighted-average number of common shares outstanding as their inclusion in the computation for all periods would be anti-dilutive due to net losses.

The following common stock equivalents were excluded from the calculation of diluted net income (loss) per share for the periods indicated because including them would have had an anti-dilutive effect (in thousands):

	<b>Year Ended December 31,</b>	
	<b>2011</b>	<b>2012</b>
Outstanding stock options		3,730
Common stock warrants	874	884
Preferred stock		18,166
Preferred stock warrants		248
	874	23,028

***Comprehensive Income (Loss)***

Comprehensive income (loss) is defined as the change in equity of a business enterprise during a period from transactions, other events, and circumstances from non-owner sources. Comprehensive



Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****2. Summary of Significant Accounting Policies (continued)**

income (loss) consists of net income (loss) and other comprehensive income (loss), which includes certain changes in equity that are excluded from net income (loss). Comprehensive income (loss) has been disclosed in the accompanying statements of operations and comprehensive income (loss) and equals the Company's net income (loss) for all periods presented.

***Subsequent Events***

The Company considers events or transactions that occur after the balance sheet date but prior to the issuance of the financial statements to provide additional evidence relative to certain estimates or to identify matters that require additional disclosure. See Note 16.

***Application of New or Revised Accounting Standards***

On April 5, 2012, the Jump-Start Our Business Startups Act (the JOBS Act) was signed into law. The JOBS Act contains provisions that, among other things, reduce certain reporting requirements for an "emerging growth company." As an emerging growth company the Company has elected to not take advantage of the extended transition period afforded by the JOBS Act for the implementation of new or revised accounting standards, and as a result, will comply with new or revised accounting standards on the relevant dates on which adoption of such standards is required for non-emerging growth companies.

***Recently Adopted Accounting Pronouncements***

From time to time, new accounting pronouncements are issued by the FASB or other standard setting bodies and adopted by the Company as of the specified effective date. Unless otherwise discussed, the Company believes that the impact of recently issued standards that are not yet effective will not have a material impact on its financial position or results of operations upon adoption.

**3. Property and Equipment, Net**

Property and equipment, net, consists of the following (in thousands):

	<b>December 31,</b>	
	<b>2011</b>	<b>2012</b>
Computer equipment and software	\$ 728	\$ 919
Office equipment	179	179
Laboratory equipment	11,692	11,815
Leasehold improvements	10,060	10,088
Construction in progress	162	8
Total property and equipment	22,821	23,009
Accumulated depreciation and amortization	(17,910)	(18,950)
Property and equipment, net	\$ 4,911	\$ 4,059

Depreciation and amortization expense was \$3.1 million and \$1.3 million, for the years ending December 31, 2011 and 2012, respectively.

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****4. Restricted Cash**

As of December 31, 2011 and 2012, the Company maintained letters of credit totaling \$0.9 million held in the form of a money market account as collateral for the Company's facility lease obligations and its credit cards.

**5. Accrued Expenses**

Accrued expenses consist of the following (in thousands):

	December 31,	
	2011	2012
Collaboration expense	\$ 1,042	\$ 1,000
Research and development related	570	1,282
Employee compensation	1,963	2,448
Professional services	368	607
Other	570	816
	\$ 4,513	\$ 6,153

**6. Warrants**

Below is a summary of the number of shares issuable upon exercise of outstanding warrants and the terms and accounting treatment for the outstanding warrants (in thousands, except per share data):

	Warrants as of		Weighted- Average Exercise Price Per Share	Expiration	Balance Sheet Classification	
	December 31, 2011	December 31, 2012			December 31, 2011	2012
Warrant to purchase Series A Preferred Stock	107	107	\$ 4.00	February 28, 2013	Liability	Liability(1)
Warrants to purchase Series B Preferred Stock	32	32	7.40	December 21, 2013	Liability	Liability
Warrants to purchase Series C-1 Preferred Stock	46	46	10.92	June 25, 2019	Liability	Liability
Warrants to purchase Series D-1 Preferred Stock	64	64	12.56	March 18, 2020	Liability	Liability
Warrants to purchase common stock	872	872	5.88	June 10, 2020 - July 9, 2020	Liability	Liability
Warrants to purchase common stock	13	13	4.00 - 7.40	March 31, 2015 - December 31, 2017	Equity	Equity(2)
All warrants	1,134	1,134	\$ 6.56			

(1) On February 6, 2013, the warrant holder exercised a warrant to purchase 107 shares of Series A Preferred Stock on a net basis, resulting in the issuance of 47 shares of Series A Preferred Stock.

(2)

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Warrants to purchase common stock were issued in connection with various debt financing transactions that were consummated in periods prior to December 31, 2011. See discussion below for further details.

In connection with various financing transactions that were consummated in periods prior to December 31, 2011, the Company issued warrants for the purchase of up to 106,500 shares of the Company's Series A redeemable convertible preferred stock (Series A Preferred Stock), 31,891 shares of the Company's Series B redeemable convertible preferred stock (Series B Preferred Stock), 45,786 shares of the Company's Series C-1 redeemable convertible preferred stock (Series C-1 Preferred Stock), and 63,693 shares of the Company's Series D-1 redeemable convertible preferred stock (Series D-1 Preferred Stock). Each warrant was immediately exercisable. The warrants to purchase Series A and Series B Preferred Stock expire seven years from the original date of issuance, while the warrants to purchase Series C-1 and Series D-1 Preferred Stock expire ten years from the original date of issuance. The warrants to purchase shares of the Company's preferred stock have an exercise price

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Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****6. Warrants (continued)**

equal to the original issuance price of the underlying instrument. Each warrant is exercisable on either a physical settlement or net share settlement basis and the redemption provisions are outside the control of the Company. Upon the conversion of the Series A Preferred Stock and/or Series B Preferred Stock and/or Series C-1 Preferred Stock and/or Series D-1 Preferred Stock into shares of common stock, the associated warrants to purchase shares of the Company's preferred stock are will become exercisable for shares of common stock.

The Company follows the provisions of ASC Topic 480, *Issuer's Accounting for Freestanding Warrants and Other Similar Instruments on Shares that Are Redeemable*, which requires that warrants to purchase redeemable preferred stock be classified as liabilities. In addition, the value of the warrants is remeasured to the then-current fair value at each reporting date. Changes in fair value are recorded to other income (expense), net. For the years ended December 31, 2011 and 2012, the Company remeasured the fair value of all of its outstanding warrants to purchase shares of the Company's preferred stock, using current assumptions, resulting in an increase in fair value of \$0.1 million and \$0.4 million, respectively, which was recorded in other expense net in the accompanying statements of operations and comprehensive income (loss). The Company will continue to re-measure the fair value of the liability associated with the warrants to purchase shares of Series B Preferred Stock, Series C-1 Preferred Stock, and Series D-1 Preferred Stock at the end of each reporting period until the earlier of the exercise or expiration of the applicable warrants or until such time that the underlying preferred stock is reclassified to permanent equity.

In December 2012, the Company modified the warrant to purchase 106,500 shares of Series A Preferred Stock and extended the expiration date from December 21, 2012 to February 28, 2013. On February 6, 2013, the holder of the warrant exercised the warrant on a net basis, resulting in the issuance of 46,668 shares of Series A Preferred Stock. Upon exercise, the Company re-measured the fair value of the warrant and recorded the resulting increase in fair value of \$0.1 million as other expense in the accompanying statement of operations and comprehensive income.

In connection with the Series E redeemable convertible preferred stock (Series E Preferred Stock) financing transactions that took place in June 2010 and July 2010, the Company issued warrants to purchase up to 871,580 shares of common stock. Each warrant was immediately exercisable and expires ten years from the original date of issuance. The warrants to purchase shares of the Company's common stock have an exercise price equal to the estimated fair value of the underlying instrument as of the initial date such warrants were issued. Each warrant is exercisable on either a physical settlement or net share settlement basis from the date of issuance. The warrant agreement contains a provision requiring an adjustment to the number of shares in the event the Company issues common stock, or securities convertible into or exercisable for common stock, at a price per share lower than the warrant exercise price. The Company concluded the anti-dilution feature required the warrants to be classified as liabilities under ASC Topic 815, *Derivatives and Hedging - Contracts in Entity's Own Equity* (ASC 815). The warrants are measured at fair value, with changes in fair value recognized as a gain or loss to other income (expense) in the statements of operations and comprehensive income (loss) for each reporting period thereafter. The fair value of the common stock warrants were recorded as a discount to the preferred stock issued of \$3.0 million, and the preferred stock is being accreted to the redemption value. On December 31, 2011 and 2012, the Company remeasured the fair value of the outstanding warrants, using current assumptions, resulting in an increase in fair value of \$0.3 million and \$1.9 million, respectively, which was recorded in other expense in the accompanying statements of operations and comprehensive income (loss) for the years ended December 31, 2011 and 2012. The Company will continue to re-measure the fair value of the liability associated with the warrants to

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****6. Warrants (continued)**

purchase common stock at the end of each reporting period until the earlier of the exercise or the expiration of the applicable warrants. On March 31, 2013, the Company retired 13,994 warrants to purchase common stock as a consequence of a repurchase of shares from an investor. All remaining outstanding warrants were fully vested and exercisable as of December 31, 2011 and 2012.

In connection with various financing transactions that were consummated in periods prior to December 31, 2011, the Company issued warrants to purchase up to 12,634 shares of common stock. The awards of warrants to purchase shares of common stock are accounted for as equity instruments. The warrants are exercisable at any time through their respective expiration dates. The fair value at issuance was calculated using the Black-Scholes option-pricing model, and was charged to interest expense during the periods the related debt was outstanding.

The Company issued warrants to purchase up to 41,388 shares of common stock in periods prior to December 31, 2011 in exchange for consulting services provided by a third party pursuant to stand-alone award agreements that are independent of an equity incentive plan. The warrants vested upon achievement of four milestones and were outstanding for approximately seven years from the date of issuance. During the year ended December 31, 2011, the holder exercised 41,388 warrants to purchase common stock on a net basis resulting in the issuance of 37,249 shares of common stock. There were no exercises, cancellations, or expirations of warrants during the year ended December 31, 2012.

***Fair Value***

The fair value of the warrants to purchase preferred stock on the date of issuance and on each re-measurement date for those warrants to purchase preferred stock classified as liabilities, is estimated using the Black-Scholes option pricing model. This method of valuation involves using inputs such as the fair value of the Company's various classes of preferred stock and common stock, stock price volatility, contractual term of the warrants, risk free interest rates, and dividend yields. The fair value of the warrants to purchase common stock on the date of issuance and on each re-measurement date for those warrants to purchase common stock are classified as liabilities and are estimated using the Monte Carlo simulation framework, which incorporated three future financing events over the remaining life of the warrants to purchase common stock. Due to the nature of these inputs and the valuation techniques utilized, the valuation of the warrants to purchase preferred stock and common stock are considered a Level 3 measurement (Note 2).

The fair value of each warrant to purchase shares of the Company's Series A Preferred Stock was estimated using the Black-Scholes option pricing model with the following assumptions:

	<b>Year Ended</b>	
	<b>December 31,</b>	
	<b>2011</b>	<b>2012(1)</b>
Fair value of underlying instrument	\$ 6.76	\$ 9.24
Expected volatility	66.0%	69.1%
Expected term (in years)	1.16	0.16
Risk-free interest rate	0.12%	0.04%
Expected dividend yield	%	%

(1)

During December 2012, the expiration date of the warrant to purchase Series A Preferred Stock was extended from December 21, 2012 to February 28, 2013. The warrant to purchase Series A Preferred Stock was exercised during the three months ended March 31, 2013.

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****6. Warrants (continued)**

The fair value of each warrant to purchase shares of the Company's Series B Preferred Stock was estimated using the Black-Scholes option pricing model with the following assumptions:

	Year Ended December 31,	
	2011	2012
Fair value of underlying instrument	\$ 7.56	\$ 9.96
Expected volatility	66.0%	69.1%
Expected term (in years)	1.98	0.97
Risk-free interest rate	0.25%	0.16%
Expected dividend yield	%	%

The fair value of each warrant to purchase shares of the Company's Series C-1 Preferred Stock was estimated using the Black-Scholes option pricing model with the following assumptions:

	Year Ended December 31,	
	2011	2012
Fair value of underlying instrument	\$ 8.84	\$ 11.04
Expected volatility	66.0%	69.1%
Expected term (in years)	7.46	6.46
Risk-free interest rate	1.35%	0.95%
Expected dividend yield	%	%

The fair value of each warrant to purchase shares of the Company's Series D-1 Preferred Stock was estimated using the Black-Scholes option pricing model with the following assumptions:

	Year Ended December 31,	
	2011	2012
Fair value of underlying instrument	\$ 8.84	\$ 10.52
Expected volatility	66.0%	69.1%
Expected term (in years)	8.22	7.22
Risk-free interest rate	1.62%	1.18%
Expected dividend yield	%	%

*Fair Value of Underlying Instrument*

The Company estimated the fair value of its shares of Series A Preferred Stock, Series B-1 Preferred Stock, Series C-1 Preferred Stock and Series D-1 Preferred Stock as of December 31, 2011 and 2012 using the PWERM.

*Expected Volatility*

The Company estimated the expected volatility based on actual historical volatility of the stock price of similar companies with publicly-traded equity securities. The Company calculated the historical volatility of the selected companies by using daily closing prices over a period of the expected term of the associated award. The companies were selected based on their enterprise value, risk profiles, position within the industry, and with historical share price information sufficient to meet the expected term of the associated award. A decrease in the selected volatility would decrease the fair value of the underlying instrument.

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****6. Warrants (continued)***Expected Term*

The Company based the expected term on the actual remaining contractual term of each respective warrant. A decrease in the expected term would decrease the fair value of the underlying instrument.

*Risk-Free Interest Rate*

The Company estimated the risk-free interest rate in reference to the yield on U.S. Treasury securities with a maturity date commensurate with the expected term of the associated award. A decrease in the selected risk-free rate would decrease the fair value of the underlying instrument.

*Expected Dividend Yield*

The Company estimated the expected dividend yield based on consideration of its historical dividend experience and future dividend expectations. The Company has not historically declared or paid dividends to stockholders. Moreover, it does not intend to pay dividends in the future, but instead expects to retain any earnings to invest in the continued growth of the business. Accordingly, the Company assumed an expected dividend yield of 0.0%.

**7. Commitments and Contingencies***Operating Leases*

The Company leases its facilities under non-cancelable operating leases that expire at various dates through May 2018. All of the Company's leases contain escalating rent clauses, which require higher rent payments in future years. The Company expenses rent on a straight-line basis over the term of the lease, including any rent-free periods. In addition, the Company received certain leasehold improvement incentives, and recorded these incentives as deferred rent, which is amortized as a reduction of rent expense over the life of the lease. Rent expense of approximately \$3.6 million and \$3.5 million were incurred during the years ended December 31, 2011 and 2012, respectively.

Future annual minimum lease payments as of December 31, 2012, are as follows (in thousands):

2013	\$	4,522
2014		4,522
2015		4,106
2016		3,938
2017		3,938
2018		2,953
<b>Total</b>	<b>\$</b>	<b>23,979</b>

In February 2011, the Company entered into a sublease agreement for a portion of one of its facility leases. The tenant will pay rent on the lease from February 28, 2011 until May 30, 2015. The Company will continue to utilize the remaining portion of the leased property.

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****7. Commitments and Contingencies (continued)**

Future annual minimum sublease payments as of December 31, 2012, are as follows (in thousands):

2013	\$	583
2014		583
2015		241
Total	\$	1,407

***Legal Proceedings***

On October 18, 2012, the Salk Institute for Biological Studies (Salk) filed a complaint in the Massachusetts Superior Court for Suffolk County, alleging that the Company breached one of the Company's two licensing agreements with Salk. The licensing agreement in dispute provides the Company with a license with respect to certain of Salk's U.S. patents related to the ActRIIB activin receptor proteins. Salk contends that, under the licensing agreement, the Company owed Salk a greater share of the upfront payment that it received under its now-terminated agreement with Shire AG regarding ACE-031 and a share of the upfront payment and development milestone payments that the Company has received under its ongoing collaboration agreement with Celgene regarding ACE-536. Salk is seeking a total of approximately \$10.5 million plus interest in payment and a 15% share of future development milestone payments received under the agreement with Celgene regarding ACE-536. The Company contends that no additional amounts are due to Salk and that it has complied with all of its payment obligations under the applicable Salk license agreement.

The Company moved to dismiss the complaint on December 3, 2012. The Court denied the Company's motion on February 28, 2013. On March 14, 2013, Acceleron answered the complaint and asserted patent invalidity counterclaims. On the basis of those counterclaims, Acceleron removed the action on March 28, 2013 to the United States District Court for the District of Massachusetts. The parties have since reached an agreement on a stipulation as to certain patent issues raised in the action, and Acceleron has dismissed its counterclaims. The Court held an initial scheduling conference on May 30, 2013, and the parties have begun fact discovery. The case is currently scheduled for trial in September 2014. The Company intends to defend its position vigorously.

The Company evaluated the suit under ASC Topic 450, *Contingencies*, as a loss contingency. The estimated loss from a loss contingency shall be accrued if information available before the financial statements are issued indicates that it is probable a liability had been incurred at the date of the financial statements, and the amount of loss can be reasonably estimated. Because the Company believes that the potential for an unfavorable outcome is not probable, it has not established a reserve with respect to the dispute as of December 31, 2012.

The Company's estimates can be affected by various factors. As of December 31, 2012, management has determined a loss is reasonably possible. Although the Company believes it would successfully defend the lawsuit, the Company has in the past participated in settlement discussions with Salk. Accordingly, the Company has estimated the range of possible losses as of December 31, 2012 and to be between \$0 and \$10.5 million plus interest.



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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**7. Commitments and Contingencies (continued)**

*Other*

The Company is also party to various agreements, principally relating to licensed technology, that require future payments relating to milestones not met at December 31, 2012, or royalties on future sales of specified products. No milestone or royalty payments under these agreements are expected to be payable in the immediate future. See Note 10 for discussion of these arrangements.

The Company enters into standard indemnification agreements in the ordinary course of business. Pursuant to the agreements, the Company indemnifies, holds harmless, and agrees to reimburse the indemnified party for losses suffered or incurred by the indemnified party, generally the Company's business partners or customers, in connection with any U.S. patent or any copyright or other intellectual property infringement claim by any third party with respect to the Company's products. The term of these indemnification agreements is generally perpetual any time after execution of the agreement. The maximum potential amount of future payments the Company could be required to make under these indemnification agreements is unlimited. The Company has never incurred costs to defend lawsuits or settle claims related to these indemnification agreements.

**8. Redeemable Convertible Preferred Stock**

As of December 31, 2012 the authorized capital stock of the Company included 74,077,227 shares of preferred stock, par value \$0.001 per share, of which: (1) 26,069,980 shares have been designated as Series A Preferred Stock, (2) 16,944,378 shares have been designated as Series B Preferred Stock, (3) 11,923,077 shares have been designated as Series C redeemable convertible (Series C Preferred Stock), (4) 2,014,652 shares have been designated as Series C-1 Preferred Stock, (5) 955,414 shares have been designated as Series D redeemable convertible preferred stock (Series D Preferred Stock), (6) 2,802,548 shares have been designated as Series D-1 Preferred Stock, (7) 3,662,422 shares have been designated as Series E Preferred Stock, and (8) 9,704,756 shares have been designated as Series F redeemable convertible preferred stock (Series F Preferred Stock, and all collectively the Preferred Stock).

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****8. Redeemable Convertible Preferred Stock (continued)**

The Company's Preferred Stock consisted of the following (in thousands, except share and per share data):

	December 31,	
	2011	2012
Series A Preferred Stock, \$0.001 par value: 26,069,980 shares authorized, 6,410,976 shares issued and outstanding at December 31, 2011 and 2012 and 6,457,644 shares at June 30, 2013, at redemption value(1)	\$ 62,049	\$ 66,665
Series B Preferred Stock, \$0.001 par value: 16,944,378 shares authorized, 4,204,185 shares issued and outstanding at December 31, 2011 and 2012, at redemption value(2)	61,464	67,044
Series C Preferred Stock, \$0.001 par value: 11,923,077 shares authorized, 2,978,062 shares issued, and outstanding at December 31, 2011 and 2012, at redemption value(2)	54,320	59,909
Series C-1 Preferred Stock, \$0.001 par value: 2,014,652 shares authorized, 457,875 issued, and outstanding at December 31, 2011 and 2012, at redemption value	8,479	9,387
Series D Preferred Stock, \$0.001 par value: 955,414 shares authorized, 234,940 shares issued, and outstanding at December 31, 2011 and 2012, at redemption value(2)	3,657	4,325
Series D-1 Preferred Stock, \$0.001 par value: 2,802,548 shares authorized, 636,942 issued and outstanding at December 31, 2011 and 2012, at redemption value	10,128	11,864
Series E Preferred Stock, \$0.001 par value: 3,662,422 shares authorized, 816,060 shares issued and outstanding at December 31, 2011 and 2012, at redemption value(2)	10,934	13,393
Series F Preferred Stock, \$0.001 par value: 9,704,756 shares authorized, 2,426,171 issued and outstanding at December 31, 2011 and 2012, at redemption value(2)	30,518	36,023
<b>Total redeemable convertible preferred stock</b>	<b>\$ 241,549</b>	<b>\$ 268,610</b>

(1) On February 6, 2013, the warrant holder exercised a warrant to purchase 106,500 shares of Series A Preferred Stock on a net basis, resulting in the issuance of 46,668 shares of Series A Preferred Stock.

(2) On March 13, 2013, the Company retired 139,741 shares of Series B Preferred Stock, 21,744 shares of Series C Preferred Stock, 2,906 shares of Series D Preferred Stock, 13,103 shares of Series E Preferred Stock and 4,825 shares of Series F Preferred Stock as a consequence of a repurchase of shares from an investor.

The holders of the Company's Preferred Stock have rights and preferences as specified below.

**Dividends**

The holders of the Company's Preferred Stock are entitled to receive, when and as declared by the Board, preferential cumulative dividends at the rate of 8% per share per annum of the stated value thereof. Such dividends shall be cumulative and shall accrue from the original issue date, whether or not earned or declared, and whether or not in any fiscal year there shall be net profits or surplus available for the payment of dividends in such fiscal year. No dividends or other distributions will be made with respect to the common stock until all declared dividends on the Preferred Stock have been paid. Additionally, if the Board declares a dividend with respect to the common stock, the Board must also declare at the same time, a dividend to the holders of the Preferred equal to the dividend that would have been payable if the outstanding Preferred Stock had been converted into shares of common stock. No dividends have been declared or paid by the Company through December 31, 2012.

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**8. Redeemable Convertible Preferred Stock (continued)**

**Liquidation**

In the event of any liquidation, dissolution, or winding up of the Company, whether voluntary or involuntary, the holders of Series F Preferred Stock are entitled to receive an amount equal to the greater of (a) \$12.56 per share, subject to appropriate adjustment, plus all dividends accrued or declared but unpaid, or (b) an amount per share as would have been payable had each share been converted to common stock immediately prior to the liquidation event. No payment shall be made to the holders of Series A, Series B, Series C, Series C-1, Series D, Series D-1 and Series E Preferred Stock or common stock unless and until full payment has been made to the holders of Series F Preferred Stock.

After payment has been made to the holders of Series F Preferred Stock, the holders of Series E Preferred Stock are entitled to receive an amount equal to the greater of (a) the Special Series E Liquidation Payment (as defined below), plus all dividends accrued or declared but unpaid, or (b) an amount per share as would have been payable had each share been converted to common stock immediately prior to the liquidation event. No payment shall be made to the holders of Series A, Series B, Series C, Series C-1, Series D and Series D-1 Preferred Stock or common stock unless and until full payment has been made to the holders of Series E Preferred Stock.

The Special Series E Liquidation Payment is equal to a preference calculated as a 22% annually compounded return on the initial per share investment amount of \$12.56 per share from the date of issuance to the date of an initial public offering, subject to appropriate adjustments.

After payment has been made to the holders of Series F and Series E Preferred Stock, the holders of Series D and Series D-1 Preferred Stock are entitled to receive an amount equal to the greater of (a) \$12.56 per share, subject to appropriate adjustment, plus any dividends accrued or declared but unpaid, or (b) an amount per share as would have been payable had each share been converted to common stock immediately prior to the liquidation event. No payment shall be made to the holders of Series A, Series B, Series C and Series C-1 Preferred Stock or common stock unless and until full payment has been made to the holders of Series D and Series D-1 Preferred Stock.

After payment has been made to the holders of Series F, Series E, Series D, and Series D-1 Preferred Stock, the holders of Series C and Series C-1 Preferred Stock are entitled to receive an amount equal to the greater of (a) \$10.92 per share, subject to appropriate adjustment for Series C-1 Preferred Stock and \$10.40 per share, subject to appropriate adjustment, for Series C Preferred Stock, plus any dividends accrued or declared but unpaid, or (b) an amount per share as would have been payable had each share been converted to common stock immediately prior to the liquidation event. No payment shall be made to the holders of Series A and Series B Preferred Stock or common stock unless and until full payment has been made to the holders of Series C and Series C-1 Preferred Stock.

After payment has been made to the holders of Series F, Series E, Series D, Series D-1, Series C and Series C-1 Preferred Stock, the holders of Series B Preferred Stock are entitled to receive, an amount equal to the greater of (a) \$7.40 per share, subject to appropriate adjustment, plus any dividends accrued or declared but unpaid, or (b) an amount per share as would have been payable had each share been converted to common stock immediately prior to the liquidation event. No payment shall be made to the holders of Series A Preferred Stock or of common stock unless and until full payment has been made to the holders of Series B Preferred Stock.

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**8. Redeemable Convertible Preferred Stock (continued)**

After payment has been made to the holders of Series F, Series E, Series D, Series D-1, Series C, Series C-1 and Series B Preferred Stock, the holders of Series A Preferred Stock are entitled to receive, in preference to the holders of common stock, an amount equal to the greater of (a) \$4.00 per share, subject to appropriate adjustment, plus any dividends accrued or declared but unpaid, or (b) an amount per share as would have been payable had each share been converted to common stock immediately prior to the liquidation event. No payment shall be made to the holders of common stock unless and until full payment has been made to the holders of Series A Preferred Stock.

The remaining assets of the Company available for distribution, after distribution to the holders of Series F, Series E, Series D, Series D-1, Series C, Series C-1, Series B and Series A Preferred Stock shall be distributed ratably among the holders of common stock.

**Voting**

The holders of the Preferred Stock are entitled to vote, together with the holders of common stock, on all matters submitted to stockholders for a vote. The holders of the Preferred Stock are entitled to the number of votes equal to the number of shares of common stock into which each share of the Preferred Stock is convertible at the time of such vote. On various matters submitted to the stockholders for a vote, certain series of Preferred Stock are entitled to separate votes.

**Conversion**

*Voluntary*

Each share of Preferred Stock is convertible at the option of the holder into such number of shares of common stock as is determined by dividing \$4.00 in the case of Series A Preferred Stock, \$7.40 in the case of Series B Preferred Stock, \$10.40 in the case of Series C Stock, \$10.92 in the case of Series C-1 Stock, \$12.56 in the case of Series D and D-1 Preferred Stock, \$12.56 in the case of Series E Stock, and \$12.56 in the case of Series F Stock by the conversion prices in effect at the time of conversion. As of December 31, 2012, the conversion rate for all series of Preferred Stock is 1:1, but is subject to adjustment in the future upon the occurrence of certain events.

*Mandatory*

Each share of Preferred Stock shall be automatically converted into shares of common stock at the conversion price in effect at the time of conversion, upon (1) the closing of an IPO of the Company's common stock in which the price is greater than \$37.68 per share, adjusted for certain dilutive events, and which results in gross proceeds of at least \$50.0 million.

Each share of Preferred Stock shall be automatically converted into shares of common stock at the conversion price in effect at the time of conversion, upon (1) the closing of an IPO of the Company's common stock in which the price is between \$14.80 and \$37.68 per share, adjusted for certain dilutive events, and which results in gross proceeds of at least \$50.0 million, and (2) the election by the holders of at least two-thirds of the outstanding shares of the respective series of Series B Preferred Stock, Series C Preferred Stock, Series D Preferred Stock, Series E Preferred Stock, and Series F Preferred Stock, voting as a single class.

Each share of Preferred Stock shall be automatically converted into shares of common stock at the conversion price in effect at the time of conversion, upon (1) the closing of an IPO of the Company's common stock in which the price is less than \$14.80 per share, adjusted for certain dilutive events, and

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**8. Redeemable Convertible Preferred Stock (continued)**

which results in gross proceeds of at least \$50.0 million, and (2) the election by the holders of at least two-thirds of the outstanding shares of Series B Preferred Stock, and (3) the election by the holders of at least 60% of the outstanding shares of the respective series of Series C Preferred Stock, Series D Preferred Stock, Series E Preferred Stock, and Series F Preferred Stock voting as a single class.

In the event of a closing of an IPO of the Company's common stock not meeting the criteria discussed above, and the election by the holders of at least two thirds of the outstanding shares of the respective series of Series B Preferred Stock, Series C Preferred Stock, Series D Preferred Stock, Series E Preferred Stock, and Series F Preferred Stock voting as a single class, each share of Preferred Stock shall be automatically converted into shares of common stock at the conversion price in effect at the time of conversion, upon.

In the event of an automatic conversion of the Preferred Stock upon the closing of an IPO in which the per-share price is less than \$37.68, adjusted for certain dilutive events, each share of Series E Preferred stock will be converted into common stock at the greater of (1) the number of shares which would be received under the conversion price in effect at the time of the offering based upon the conversion features noted above, or (2) a ratio determined by dividing the Special Series E Liquidation Payment, accrued from the issuance date through the date of an IPO, by the price per share of the Company's common stock in an IPO.

***Special Mandatory***

In the event that any holder of shares of Preferred Stock does not participate in a Qualified Financing (as defined) by purchasing, in the aggregate, in such Qualified Financing and within the time period specified by the Company, such holder's pro rata amount, then such holder's shares of preferred stock will automatically convert into common stock at the respective Conversion Price (as defined).

The Company evaluated each series of its Preferred Stock and determined that each individual series is considered an equity host under ASC 815. In making this determination, the Company's analysis followed the whole instrument approach which compares an individual feature against the entire preferred stock instrument which includes that feature. The Company's analysis was based on a consideration of the economic characteristics and risks of each series of Preferred Stock. More specifically, the Company evaluated all of the stated and implied substantive terms and features, including: (1) whether the Preferred Stock included redemption features, (2) how and when any redemption features could be exercised, (3) whether the holders of Preferred Stock were entitled to dividends, (4) the voting rights of the Preferred Stock and (5) the existence and nature of any conversion rights. As a result of the Company's conclusion that the Preferred Stock represents an equity host, the conversion feature of all series of Preferred Stock is considered to be clearly and closely related to the associated Preferred Stock host instrument. Accordingly, the conversion feature of all series of Preferred Stock is not considered an embedded derivative that requires bifurcation.

The Company accounts for potential beneficial conversion features under FASB ASC Topic 470-20, *Debt with Conversion and Other Options*. At the time of each of the issuances of Preferred Stock, the Company's common stock into which each series of the Company's Preferred Stock is convertible had an estimated fair value less than the effective conversion prices of the Preferred Stock. Therefore, there was no intrinsic value on the respective commitment dates.

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**8. Redeemable Convertible Preferred Stock (continued)**

As noted above, in certain events, the Series E Preferred Stock may convert to common stock on a basis higher than 1-to-1, based on a formula driven by the date on which the Company completes an IPO and the price of such offering. The Company concluded, in accordance with the provisions of ASC 470, that as the changes to the conversion terms would be triggered by a future event that is outside of the Company's control, this represents a contingent conversion option, and, therefore, should not be recognized until and unless the triggering event occurs. The Company evaluated whether a beneficial conversion feature may be required to be recorded with respect to the Series E Preferred Stock when the triggering event occurs, measured based on the number of shares of common stock assumed to be issuable upon the resolution of the contingency multiplied by the commitment date fair value of the common stock less the proceeds of the Series E Preferred Stock financing. Based upon this assessment, the Company determined that no beneficial conversion feature is required to be recognized.

**Redemption**

The Company shall be required to redeem all, but not less than all, of the outstanding shares of the Series F Preferred Stock, as applicable, in three equal installments at the written election of holders of 83% of the outstanding shares of Series F Preferred Stock at any time on or after the date that is 90 days before the fifth anniversary of the original issue date of the Company's Series F Preferred Stock, which is September 22, 2016. The redemption price per share of Series F Preferred Stock shall be equal to (1) \$12.56 for the Series F Preferred Stock (the Series F Base Redemption Price) adjusted for certain dilutive events, plus all dividends accrued or declared but unpaid on such share on the applicable redemption date, plus (2) an additional amount computed similar to interest payable on the Series F Base Redemption Price at the rate equal to simple interest of 10% per annum from the date of issuance of such shares.

After full redemption of the Series F Preferred Stock, the Company shall be required to redeem all, but not less than all, of the outstanding shares of the Series E Preferred Stock, in three equal installments at the written election of holders of 75% of the outstanding shares of Series E Preferred Stock at any time on or after the date that is 90 days before the fifth anniversary of the original issue date of the Company's Series F Preferred Stock. The redemption price per share of Series E Preferred Stock shall be equal to (1) \$12.56 for the Series E Preferred Stock (the Series E Base Redemption Price) adjusted for certain dilutive events, plus all dividends accrued or declared but unpaid on such share on the applicable redemption date, plus (2) an additional amount computed similar to interest payable on the Series E Base Redemption Price at the rate equal to simple interest of 10% per annum from the date of issuance of such shares.

After full redemption of the Series F and Series E Preferred Stock, the Company shall be required to redeem all, but not less than all, of the outstanding shares of the Series D and Series D-1 Preferred Stock, as applicable, in three equal installments at the written election of holders of 85% of the outstanding shares of Series D and Series D-1 Preferred Stock at any time on or after the date that is 90 days before the fifth anniversary of the original issue date of the Company's Series F Preferred Stock. The redemption price per share of Series D Preferred Stock shall be equal to (1) \$12.56 for the Series D and D-1 Preferred Stock (the Series D Base Redemption Price) adjusted for certain dilutive events, plus all dividends accrued or declared but unpaid on such share on the applicable redemption date, plus (2) an additional amount computed similar to interest payable on the Series D Base

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**8. Redeemable Convertible Preferred Stock (continued)**

Redemption Price at the rate equal to simple interest of 10% per annum from the date of issuance of such shares.

After full redemption of the Series F, Series E, Series D, and Series D-1 Preferred Stock, the Company shall be required to redeem all, but not less than all, of the outstanding shares of the Series C Preferred Stock, as applicable, in three equal installments at the written election of holders of two-thirds of the outstanding shares of Series C and Series C-1 Preferred Stock at any time on or after the date that is 90 days before the fifth anniversary of the original issue date of the Company's Series F Preferred Stock. The redemption price per share shall be equal to (1) \$10.40 in the case of Series C Preferred Stock and \$10.92 in the case of Series C-1 Preferred Stock (the Series C Base Redemption Price), adjusted for certain dilutive events, plus all dividends accrued or declared but unpaid on such share on the applicable redemption date, plus (2) an additional amount computed similar to interest payable on the Series C Base Redemption Price at the rate equal to simple interest of 10% per annum from the date of issuance of such shares.

After full redemption of the Series F, Series E, Series D, Series D-1, Series C, and Series C-1 Preferred Stock, the Company shall be required to redeem all, but not less than all, of the outstanding shares of the Series B Preferred Stock, as applicable, in three equal installments at the written election of holders of two-thirds of the outstanding shares of Series B Preferred Stock at any time on or after the date that is 90 days before the fifth anniversary of the original issue date of the Company's Series F Preferred Stock. The redemption price per share shall be equal to (1) \$7.40 for the Series B Preferred Stock (the Series B Base Redemption Price), adjusted for certain dilutive events, plus all dividends accrued or declared but unpaid on such share on the applicable redemption date, plus (2) an additional amount computed similar to interest payable on the Series B Base Redemption Price at the rate equal to simple interest of 10% per annum from the date of issuance of such shares.

After full redemption of the Series F, Series E, Series D, Series D-1, Series C, Series C-1, and Series B Preferred Stock, the Company shall be required to redeem all, but not less than all, of the outstanding shares of the Series A Preferred Stock, as applicable, in three equal installments at the written election of holders of two-thirds of the outstanding shares of Series A Preferred Stock at any time on or after the date that is 90 days before the fifth anniversary of the original issue date of the Company's Series F Preferred Stock. The redemption price per share shall be equal to (1) \$4.00 for the Series A Preferred Stock (the Series A Base Redemption Price), adjusted for certain dilutive events, plus all dividends accrued or declared but unpaid on such share on the applicable redemption date, plus (2) an additional amount computed similar to interest payable on the Series A Base Redemption Price at the rate equal to simple interest of 10% per annum from the date of issuance of such shares.

As the Preferred Stock may become redeemable upon an event that is outside of the control of the Company, the Preferred Stock has been classified outside of permanent equity.

**9. Common Stock**

As of December 31, 2012, the authorized capital stock of the Company included 104,013,161 shares of common stock, par value \$0.001 per share.

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****9. Common Stock (continued)****General**

The voting, dividend and liquidation rights of the holders of shares of common stock are subject to and qualified by the rights, powers and preferences of the holders of the Preferred Stock. The common stock has the following characteristics:

*Voting*

The holders of shares of common stock are entitled to one vote for each share of common stock held at all meetings of stockholders and written actions in lieu of meetings.

*Dividends*

The holders of shares of common stock are entitled to receive dividends, if and when declared by the Board. Cash dividends may not be declared or paid to holders of shares of common stock until paid on each series of outstanding Preferred Stock in accordance with their respective terms. No dividends have been declared or paid by the Company through December 31, 2012.

*Liquidation*

After payment to the holders of shares of Preferred Stock of their liquidation preferences, the holders of shares of common stock are entitled to share ratably in the Company's remaining assets available for distribution to stockholders, in the event of any voluntary or involuntary liquidation, dissolution or winding up of the Company or upon the occurrence of a deemed liquidation event.

**Reserved for Future Issuance**

There were 2,393,458 and 2,432,155 common shares issued and outstanding as of December 31, 2011 and 2012, respectively. The Company has reserved for future issuance the following number of shares of common stock (in thousands):

	<b>December 31, 2012</b>
Conversion of Series A Preferred Stock	6,411
Conversion of Series B Preferred Stock	4,204
Conversion of Series C Preferred Stock	2,978
Conversion of Series C-1 Preferred Stock	458
Conversion of Series D Preferred Stock	235
Conversion of Series D-1 Preferred Stock	637
Conversion of Series E Preferred Stock	816
Conversion of Series F Preferred Stock	2,426
Warrants to purchase Preferred Stock	248
Outstanding stock options to purchase common stock	3,730
Shares available for future issuance under stock option plan	120
Warrants to purchase common stock	884
Additional shares reserved for unissued, but designated, Preferred Stock	55,912
Total shares of authorized common stock reserved for future issuance	79,059



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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**10. Significant Agreements**

**Celgene**

*Overview*

On February 20, 2008 the Company entered into a collaboration, license, and option agreement (the Sotatercept Agreement) with Celgene Corporation (Celgene) relating to sotatercept. On August 2, 2011, the Company entered into a second collaboration, license and option agreement with Celgene for ACE-536 (the ACE-536 Agreement), and also amended certain terms of the Sotatercept Agreement. These agreements provide Celgene exclusive licenses for Sotatercept and ACE-536 in all indications, as well as exclusive rights to obtain a license to certain future compounds. Celgene is a global biopharmaceutical company primarily engaged in the discovery, development and commercialization of innovative therapies designed to treat cancer and immune-inflammatory related diseases.

*Sotatercept Agreement*

Under the terms of the Sotatercept Agreement, the Company and Celgene collaborate worldwide for the joint development and commercialization of sotatercept. The Company also granted Celgene an option to license three discovery stage compounds. Under the terms of the agreement, the Company and Celgene will jointly develop, manufacture and commercialize sotatercept. Celgene paid \$45.0 million of nonrefundable, upfront license and option payments to the Company upon the closing of the Sotatercept Agreement.

The Company retained responsibility for research, development through the end of Phase 2a clinical trials, as well as manufacturing the clinical supplies for these trials. These activities were substantially completed in 2011. Celgene is conducting the ongoing Phase 2 trials for myelodysplastic syndromes (MDS), chronic kidney disease, and  $\beta$ -thalassemia and will be responsible for any Phase 3 clinical trials, as well as additional Phase 2 clinical trials, and will be responsible for overseeing the manufacture of Phase 3 and commercial supplies by third party contract manufacturing organizations. Under the agreement, the Company was eligible to receive clinical milestones of up to \$88.0 million, regulatory milestones of up to \$272.0 million, and commercial milestones of up to \$150.0 million for sotatercept. Clinical milestone payments are triggered upon initiation of a defined phase of clinical research for a product candidate. Regulatory milestone payments are triggered upon the acceptance of the marketing application and upon the approval to market a product candidate by the Food and Drug Administration (FDA) or other global regulatory authorities. Commercial milestone payments are triggered when an approved pharmaceutical product reaches certain defined levels of net sales by Celgene in countries outside of North America. In addition, to the extent sotatercept is commercialized, the Company would be entitled to receive tiered royalty payments in the low-to-mid twenty percent range of net sales from sales generated from all geographies. Royalty payments are subject to certain reductions, including for entry of a generic product onto the market.

Additionally, for three named discovery-stage option programs the Company was eligible to receive option fees of up to \$30.0 million, clinical milestones of up to \$53.3 million, regulatory milestones of up to \$204.0 million, and commercial milestones of up to \$150.0 million for each option program. Clinical milestone payments are triggered upon initiation of a defined phase of clinical research for a product candidate. Regulatory milestone payments are triggered upon the acceptance of the marketing application and upon the approval to market a product candidate by the FDA or other global regulatory authorities. Commercial milestone payments are triggered when an approved pharmaceutical

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****10. Significant Agreements (continued)**

product reaches certain defined levels of net sales by Celgene in countries outside of North America. Option fee payments are triggered upon license of any of the option programs by Celgene. In addition, to the extent an option compound is commercialized, the Company would be entitled to receive tiered royalty payments in the low-to-mid twenty percent range of net sales from sales generated from all geographies. Royalty payments are subject to certain reductions, including for entry of a generic product onto the market. None of the three discovery stage programs has advanced to the stage to achieve payment of a milestone.

In connection with entering into the Sotatercept Agreement, Celgene purchased 457,875 shares of Series C-1 Preferred Stock at the aggregate purchase price of \$5.0 million. The Series C-1 Preferred Stock was purchased at an amount that was deemed to represent fair value at the time of purchase. In the event that the Company's IPO results in gross proceeds of at least \$35.0 million, Celgene has committed to purchase, in a private offering concurrently with the IPO, shares of common stock equal to \$10.0 million at the issuance price per share at the IPO if the gross proceeds from the IPO are greater than \$50.0 million or twenty percent (20%) of the gross proceeds if the IPO raises less than \$50.0 million.

Commensurate with the execution of the ACE-536 Agreement described below, the Company and Celgene agreed to modify the terms of the Sotatercept Agreement. The modified terms included: (1) a change to the responsibility for development costs to align with the ACE-536 Agreement, with Celgene responsible for more than half of the worldwide costs through December 31, 2012, and 100% of the development costs thereafter, (2) future contingent development milestones for sotatercept were amended to a two-category (oncology and non-oncology) structure with potential future clinical milestones of \$27.0 million and regulatory milestones of \$190.0 million from a four-category (various cancer indications) structure and, (3) future contingent development milestones for option compounds were amended to a two-category (oncology and non-oncology) structure with potential future clinical milestones of \$25.5 million and regulatory milestones of \$142.5 million from a four-category (various cancer indications) structure, and (4) an option to buy down tiered royalty payments on both Sotatercept and ACE-536 with a one-time \$25.0 million payment on or prior to January 1, 2013. The potential commercial milestones remained unchanged. Through December 31, 2012, the Company has received \$34.2 million in research and development funding and milestone payments for sotatercept under the original and modified agreements. The next likely clinical milestone payment would be \$7.0 million and result from Celgene's start of a Phase 2b clinical trial in chronic kidney disease.

The Sotatercept Agreement will expire on a country-by-country basis on the occurrence of both of the following: (1) the expiration of the royalty term with respect to all license products in such country, and (2) the exercise or forfeiture by Celgene of its option with regard to each option compound. The royalty term for each licensed product in each country outside North America is the period commencing with first commercial sale of the applicable licensed product in the applicable country and ending on the latest of expiration of specified patent coverage or a specified period of years. The royalty term for each licensed product in North America is the period commencing with the first commercial sale in North America and ending, on a licensed product and country-by-country basis on the date which commercialization of such licensed product has ceased. The term for each option compound runs for a specified period of years unless Celgene exercises its option, in which case the compound becomes a licensed product, or forfeits its option by failing to make certain payments following the achievement of certain milestones in early clinical development of the option compound.

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**10. Significant Agreements (continued)**

Celgene has the right to terminate the agreement with respect to one or more licensed targets or in its entirety, upon 180 days' notice (or 45 days' notice if the licensed product has failed to meet certain end point criteria with respect to clinical trials or other development activities). The agreement may also be terminated in its entirety by either Celgene or the Company in the event of a material breach by the other party or in the event of a bankruptcy filing of the other party. There are no cancellation, termination or refund provisions in this arrangement that contain material financial consequences to the Company.

***ACE-536 Agreement***

Under the terms of the ACE-536 Agreement, the Company and Celgene collaborate worldwide for the joint development and commercialization of ACE-536. The Company also granted Celgene an option for future products Acceleron files an Investigational New Drug application for the treatment of anemia. Celgene paid \$25.0 million on the closing of the ACE-536 Agreement in August, 2011.

The Company retains responsibility for research, development through the end of Phase 1 and initial Phase 2 clinical trials, as well as manufacturing the clinical supplies for these studies. Celgene will conduct subsequent Phase 2 and Phase 3 clinical studies. Acceleron will manufacture ACE-536 for the Phase 1 and Phase 2 clinical trials and Celgene will be responsible for overseeing the manufacture of Phase 3 and commercial supplies by third party contract manufacturing organizations. The Company is eligible to receive clinical milestones of up to \$32.5 million, regulatory milestones of up to \$105.0 million and commercial milestones of up to \$80.0 million for ACE-536. The Company will receive additional, lower development, regulatory, and commercial milestones for any additional products for the treatment of anemia on which Celgene exercises an option. Clinical milestone payments are triggered upon initiation of a defined phase of clinical research for a product candidate. Regulatory milestone payments are triggered upon the acceptance of the marketing application and upon approval to market a protein therapeutic candidate by the FDA or other global regulatory authorities. Commercial milestone payments are triggered when an approved pharmaceutical product reaches certain defined levels of net sales by Celgene in countries outside of North America. In addition, to the extent ACE-536 is commercialized, the Company would be entitled to receive tiered royalty payments in the low-to-mid twenty percent range of net sales from sales generated from all geographies. Royalty payments are subject to certain reductions, including for entry of a generic product onto the market. Through December 31, 2012, the Company has received \$13.3 million in research and development funding and milestone payments for ACE-536. The next likely clinical milestone payment would be \$15.0 million and result from the start of a Phase 3 study in MDS or  $\beta$ -thalassemia. The Company has not yet identified additional compounds for the treatment of anemia. Accordingly, there is no assurance that the Company will generate future value from additional programs.

The ACE-536 Agreement will expire on a country-by-country basis on the occurrence of both of the following: (1) the expiration of the royalty term with respect to all license products in such country, and (2) the end of the option term. The royalty term for each licensed product in each country outside North America is the period commencing with first commercial sale of the applicable licensed product in the applicable country and ending on the latest of expiration of specified patent coverage or a specified period of years. The royalty term for each licensed product in North America is the period commencing with the first commercial sale in North America and ending, on a licensed product and

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****10. Significant Agreements (continued)**

country-by-country basis on the date which commercialization of such licensed product has ceased. The option term runs until the later of (1) the date on which no development or commercialization activities are ongoing or are expected to commence for any licensed products under the ACE-536 Agreement; (2) the date on which no development or commercialization activities are ongoing or are expected to commence for any licensed products under the Sotatercept Agreement and all option rights under the Sotatercept Agreement have been forfeited with respect to each option compound where Celgene has made a payment with respect to such compound; and (3) the royalty term for all licensed products under the ACE-536 Agreement and the Sotatercept Agreement has ended; provided that if at the time the option term would otherwise end any option compounds under the ACE-536 Agreement are in clinical development the option term shall continue until Celgene's rights to such compound are either exercised or forfeited.

Celgene has the right to terminate the ACE-536 Agreement with respect to one or more licensed targets or in its entirety, upon 180 days' notice (or 45 days' notice if the licensed product has failed to meet certain end point criteria with respect to clinical trials or other development activities), provided that Celgene may not terminate the ACE-536 Agreement prior to the completion of the on-going ACE-536  $\beta$ -thalassemia and ACE-536 MDS Phase 2 clinical trials, except under certain conditions. The agreement may also be terminated in its entirety by either Celgene or the Company in the event of a material breach by the other party or in the event of a bankruptcy filing of the other party. There are no cancellation, termination or refund provisions in this arrangement that contain material financial consequences to the Company.

***Both Agreements***

The Company and Celgene shared development costs under the Sotatercept and ACE-536 Agreements through December 31, 2012. As of January 1, 2013, Celgene is responsible for paying 100% of worldwide development costs under both agreements. Celgene will be responsible for all commercialization costs worldwide. The Company has the right to co-promote sotatercept, ACE-536 and future products under both agreements in North America. Celgene's option to buy down royalty rates for sotatercept and ACE-536 expired unexercised and, therefore, the Company will receive tiered royalties in the low-to-mid twenty percent range on net sales of sotatercept and ACE-536. The royalty schedules for sotatercept and ACE-536 are the same.

***Accounting Analysis***

Prior to 2011, the Company accounted for the Sotatercept Agreement, as a multiple element arrangement under ASC 605-25 (prior to the amendments of ASU 2009-13). The Company identified the following deliverables under the arrangement; (1) the license to the ActRIIA compound, (2) right to license option program compounds, (3) participation in the joint development committee, (4) participation in the joint commercialization committee and (5) research and development activities. Under the provisions of ASC 605-25, applicable to the arrangement, since the Company could not establish VSOE for the undelivered elements, the Company was required to recognize the initial consideration, consisting of the \$45.0 million of nonrefundable upfront license and option payments, over the period the undelivered elements were to be delivered, which was initially estimated to be 15 years. As of the date of the modification of the agreement, there was approximately \$34.7 million of deferred revenue under the arrangement.

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**10. Significant Agreements (continued)**

As a result of the material modifications to the cost sharing obligations, milestone payments structure and royalty payment structure, the Company concluded the modification represented a significant modification under ASU 2009-13, which required the Company to apply the updated provisions of ASU 2009-13 subsequent to the modification.

Because the ACE-536 Agreement and the amendment to the Sotatercept Agreement were negotiated in contemplation of each other, and the Company had not yet completed all of its obligations pursuant to the Sotatercept Agreement, the agreements were considered one arrangement for accounting purposes. The deliverables under the combined arrangement include: (1) licenses to develop and commercialize sotatercept and ACE-536, (2) performance of research and development services, (3) participation on the joint development committees, and (4) the performance of manufacturing services to provide clinical material to Celgene. The Company has determined the option to future products related to the treatment of anemia represents a substantive option. The Company is under no obligation to discover, develop or deliver any new compounds that modulate anemia and Celgene is not contractually obligated to exercise the option. As a result, the Company is at risk as to whether Celgene will exercise the option.

All of these deliverables identified in the arrangement were deemed to have stand-alone value and to meet the criteria to be accounted for as separate units of accounting under ASC 605-25. Factors considered in making this determination included, among other things, the subject of the licenses, the nature of the research and development services, and the capabilities of Celgene.

The total arrangement consideration of \$77.7 million under the ACE-536 Agreement and amended Sotatercept Agreement consisted of (1) the \$25.0 million up-front payment for the license of ACE-536, (2) the remaining deferred revenue from the Sotatercept Agreement of \$34.7 million, and (3) estimated payments for development activities and manufacturing services of \$18.0 million. The Company used its BEBP for each of the undelivered elements as the Company did not have VSOE or TPE of selling price for each deliverable. The Company's BEBP considered its development plan for the compounds, expected manufacturing services, and reimbursement from Celgene (reimbursement of more than half of development expenses through December 31, 2012 and 100% thereafter). The Company determined its BEBP for each of the undelivered elements under the arrangements as of the arrangement execution date as follows:

\$18.8 million for research and development services

\$2.9 million for the sotatercept joint development committee

\$3.7 million for the ACE 536 joint development committee

\$2.8 million for the manufacturing services

After determining BEBP of the undelivered elements, the remaining consideration of \$49.5 million was recognized upon execution of the arrangements. The difference between the estimated payments of \$18.0 million and the estimated selling prices which totaled \$28.2 million, using BEBP, for undelivered elements was \$10.2 million. This amount was deferred at inception and will be recognized as the undelivered elements are delivered, using the proportional performance method, or ratably in the case of performance on the Joint Development Committee.

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****10. Significant Agreements (continued)**

During 2011, the Company achieved a \$7.5 million clinical milestone under its ACE-536 Agreement, related to the dosing of the first patient in a multiple-dose clinical trial. The Company evaluated the milestone and determined that it was not substantive, as there was no significant uncertainty to achieving the milestone upon execution of the ACE-536 Agreement. As such, the Company allocated the \$7.5 million payment based on the allocation of arrangement consideration determined at the execution date of the ACE-536 Agreement and amended Sotatercept Agreement. Based on this allocation, the Company recognized \$4.8 million of the payment upon achievement, with the remaining \$2.7 million recognized as revenue as the undelivered elements are delivered, consistent with the treatment of the up-front payment. During 2011, the Company achieved a \$7.0 million clinical milestone under its Sotatercept Agreement, related to the dosing of the first patient for a Phase 2b clinical trial. The Company evaluated the milestone and deemed it to be substantive and consistent with the definition of a milestone included in ASU 2010-17 and, accordingly, recognized the \$7.0 million payment in revenue during the year ended in December 31, 2011. During January 2013, the Company achieved a \$10.0 million clinical milestone under its ACE-536 Agreement, related to the dosing of the first patient for a Phase 2 clinical trial. The Company evaluated the milestone and deemed it to be substantive and consistent with the definition of a milestone included in ASU 2010-17. The remaining development milestones under the ACE-536 and Sotatercept Agreements were deemed to be substantive and consistent with the definition of a milestone included in ASU 2010-17 and, accordingly, the Company will recognize payments related to the achievement of such milestones, if any, when such milestone is achieved. Factors considered in this determination included scientific and regulatory risks that must be overcome to achieve the milestones, the level of effort and investment required to achieve each milestone, and the monetary value attributed to each milestone. During the years ended December 31, 2011 and 2012, the Company recognized \$54.8 million and \$2.0 million, respectively, of the total deferred revenue as license and milestone revenue in the accompanying statements of operations and comprehensive income (loss).

Pursuant to the terms of the agreement, Celgene and the Company share development costs, with Celgene responsible for substantially more than half of the costs for sotatercept and ACE-536 until December 31, 2012 and 100% of the costs from January 1, 2013 and thereafter. Payments from Celgene with respect to research and development costs incurred by the Company are recorded as cost-sharing revenue. Payments by the Company to Celgene for research and development costs incurred by Celgene are recorded as a reduction to cost-sharing revenue. During the years ended December 31, 2011 and 2012 the Company recorded net cost-sharing revenue of \$(0.1) million and \$2.9 million, respectively, which includes payments to Celgene of \$2.8 million and \$2.8 million, respectively which were recorded as contra-revenue.

**Other Agreements***Shire License*

In September 2010, the Company entered into a license and collaboration agreement granting Shire AG the exclusive right to develop, manufacture and commercialize ActRIIB compounds in territories outside North America. Shire also received the right to conduct research and manufacture commercial supplies in North America for ActRIIB compounds. The lead ActRIIB compound was designated ACE-031. Under the initial development plan, the companies share the costs associated with developing and commercializing ACE-031, in Duchenne Muscular Dystrophy. In September 2010, Shire

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****10. Significant Agreements (continued)**

made a nonrefundable, up-front license payment to the Company of \$45.0 million. In accordance with the Company's revenue recognition policy prior to the adoption of ASU 2009-13, the up-front license payment of \$45.0 million was deferred, and will be recognized as revenue ratably over three years, which represented the original period over which the Company expected to perform and deliver research and development and manufacturing services. On February 8, 2011, the FDA placed ACE-031 on clinical hold. The Company re-assessed the duration of its deliverables under the license agreement and estimated the new term to be approximately five years. The adjustment was treated as a change in accounting estimate with the remaining deferred revenue of \$38.8 million at February 8, 2011, recognized prospectively over the new period of research and development and manufacturing services. In April 2013, the Company and Shire determined not to further pursue development of ACE-031 and Shire sent the Company a notice of termination for the ACE-031 collaboration. The collaboration terminated effective June 30, 2013. At December 31, 2012, the Company had classified the remaining deferred revenue as current in the balance sheet. Upon the effectiveness of the termination of the Shire Agreement in the second quarter of 2013, the Company accelerated the recognition of \$22.4 million of remaining deferred revenue from upfront non-refundable payments received under the Shire Agreement as it had no further obligation for deliverables under the Shire Agreement. During the years ended December 31, 2011 and 2012, the Company recognized \$8.4 million and \$7.7 million, respectively, of the up-front, non-refundable payments as license and milestone revenue in the accompanying statements of operations and comprehensive income (loss).

The agreement also included contingent milestone payments, based on the achievement of development milestones totaling \$223.8 million and commercial milestones of \$228.8 million for ActRIIB compounds. The milestones under the Shire Agreement were deemed to be substantive and consistent with the definition of a milestone included in ASU 2010-17 and, accordingly, the Company recognized payments related to the achievement of such milestones, if any, when such milestone was achieved. Factors considered in this determination included scientific and regulatory risks that must be overcome to achieve the milestones, the level of effort and investment required to achieve each milestone, and the monetary value attributed to each milestone.

Pursuant to the terms of the agreement, Shire and the Company shared development costs, with Shire responsible for 65% of the costs for ACE-031 and 55% of the costs for licensed compounds other than ACE-031. Payments from Shire with respect to research and development costs incurred by the Company are recorded as cost-sharing revenue. Payments by the Company to Shire for research and development costs incurred by Shire are recorded as a reduction to cost-sharing revenue. During the years ended December 31, 2011 and 2012, the Company recorded net cost-sharing revenue of \$4.1 million and \$2.7 million, respectively, which includes payments to Shire of \$2.0 million and \$0.7 million, respectively, which are recorded as contra-revenue in the accompanying statements of operations and comprehensive income (loss).

***Alkermes License***

In December 2009, the Company entered into a Collaboration and License Agreement with Alkermes, Inc. (Alkermes) relating to a proprietary technology platform for extending the circulating half-life of certain proteins. Under the terms of the agreement, Alkermes paid the Company an up-front cash payment of \$2.0 million in December 2009, which was deferred and recognized as license revenue ratably over the estimated research and development term. In addition, Alkermes purchased

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****10. Significant Agreements (continued)**

636,942 shares of Series D-1 Preferred Stock at a per share price of \$12.56, resulting in gross proceeds to the Company of \$8.0 million. The Company determined that the price of \$12.56 paid by Alkermes included a premium of \$2.32 per share over the fair value of the Series D-1 Preferred Stock of \$10.24 as calculated by the Company in its contemporaneous stock valuation. Accordingly, the Company has recognized the premium of \$1.5 million as additional license revenue on a straight-line basis over the term discussed above. In October 2011, Alkermes discontinued development of the compounds being investigated under the license agreement, and as a result, the Company recognized the remaining \$2.4 million of the up-front payment as revenue, as it had no further obligations under the arrangement, though the license continues.

As the principal in the collaboration, Acceleron recognized cost-sharing revenue for reimbursement payments from Alkermes. During the year ended December 31, 2011, the Company recognized net cost-sharing revenue of \$0.7 million. No amounts were recognized in subsequent periods.

***ImmunoGen Services Agreement***

In October 2010, the Company entered into a Biopharmaceutical Services Agreement with ImmunoGen, Inc. Acceleron agreed to develop and manufacture an ImmunoGen product. The Company determined the arrangement should be accounted for as a service arrangement, using the proportional performance method. Accordingly, the Company recognized revenue as the underlying performance criteria were met. The costs associated with the services were charged to operations as incurred. As of December 31, 2011, the work was completed, and the Company recorded revenue of \$1.7 million for the year ended December 31, 2011.

***Other***

The Company entered into a license agreement with a non-profit institution for an exclusive, sublicensable, worldwide, royalty-bearing license to certain patents developed by the institution (Primary Licensed Products). In addition, the Company was granted a non-exclusive, non-sub-licensable license for Secondary Licensed Products. As compensation for the licenses, the Company issued 250,000 shares of its common stock to the institution, the fair value of which was \$25,000, and was expensed during 2004, to research and development expense. The Company also agreed to pay specified development milestone payments totaling up to \$2.0 million for sotatercept and \$0.7 million for ACE-536. In addition, the Company is obligated to pay milestone fees based on the Company's research and development progress, and U.S. sublicensing revenue ranging from 10%-25%, as well as a royalty ranging from 1.0%-3.5% of net sales on any products developed under the licenses. During 2011 and 2012, the Company paid and expensed milestones and fees defined under the agreement totaling \$0.1 million and zero, respectively. The Company also paid \$0.5 million and zero in 2011 and 2012, respectively, based on the receipt of U.S. sublicensing revenue, which is recorded as research and development expense.

The Company entered into another license agreement with certain individuals for an exclusive, sublicensable, worldwide, royalty-bearing license to certain patents developed by the individuals. The Company agreed to pay specified development and sales milestone payments aggregating up to \$1.0 million relating to the development and commercialization of dalantercept. In addition, the Company is required to pay royalties in the low single-digits on worldwide net product sales of dalantercept, with royalty obligations continuing at a 50% reduced rate for a period of time after patent expiration. If the Company sublicenses its patent rights, the Company will owe a percentage of



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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**10. Significant Agreements (continued)**

sublicensing revenue, excluding payments based on the level of sales, profits or other levels of commercialization. During the years ended December 31, 2011 and 2012, the Company did not reach any milestones defined under the agreement and, therefore, no amounts have been paid or expensed.

During 2012, the Company executed a license agreement with a research institution for an exclusive, sublicensable, worldwide, royalty-bearing license. The Company is obligated to pay development milestones and commercial milestone fees totaling up to \$1.0 million. Under the agreement, if the Company uses the inventors in the clinical research, the development milestones are waived and commercial milestones shall change to \$0.8 million plus any waived milestones. The Company will also pay \$25,000 annually upon first commercial sale as well as royalties of 1.5% of net sales on any products developed under the patents. During the year ended December 31, 2012, the Company did not reach any milestones defined under the agreement and, therefore, no amounts have been paid or expensed.

**11. Stock-Based Compensation**

The Company's 2003 Stock Option and Restricted Stock Plan (the 2003 Plan) provides for the issuance of stock options, restricted stock awards, and restricted stock to employees, officers, directors, consultants, and key personnel of the Company as determined by the Board. As of December 31, 2012, the total number of shares of common stock which may be issued under the 2003 Plan was 4,937,500. The number of options available for future grant was 119,542 at December 31, 2012. This number can be increased by the Board, subject to the approval of the shareholders

The Company has not granted unrestricted stock awards under the 2003 Plan since its inception. Stock options carry an exercise price equal to the estimated fair value of the Company's common stock on the date of grant. Options generally expire ten years following the date of grant. Stock options and restricted stock awards typically vest over four years, but vesting provisions can vary based on the discretion of the Board.

Shares of the Company's common stock underlying any awards that are forfeited, canceled, withheld upon exercise of an option, or settlement of an award to cover the exercise price or tax withholding, reacquired by the Company prior to vesting, satisfied without the issuance of shares of the Company's common stock, or otherwise terminated other than by exercise will be added back to the shares of common stock available for issuance under the 2003 Plan. Shares available for issuance under the 2003 Plan may be authorized but unissued shares of the Company's common stock or shares of the Company's common stock that have been reacquired by the Company.

During 2010, the Company modified the awards of three employees that left the Company. The modifications all related to the term of vested options post termination. The changes ranged from 3.5 years to the remaining life of the option. Awards were reviewed under ASC 718, and the fair value of the unvested options that were modified will be re-measured and the expense adjusted at each reporting period. During the years ended December 31, 2011 and 2012, non-employee stock compensation expense of \$0.2 million and \$36,000 respectively, was recorded.

The Company recognized stock-based compensation expense totaling \$1.4 million and \$1.2 million, during the years ended December 31, 2011 and 2012, respectively.

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****11. Stock-Based Compensation (continued)**

Total compensation cost recognized for all stock-based compensation awards in the statements of operations and comprehensive income (loss) is as follows (in thousands):

	<b>Year Ended December 31,</b>	
	<b>2011</b>	<b>2012</b>
Research and development	\$ 686	\$ 514
General and administrative	741	692
	<b>\$ 1,427</b>	<b>\$ 1,206</b>

The fair value of each option issued to employees was estimated at the date of grant using the Black-Scholes option pricing model with the following weighted-average assumptions (in thousands):

	<b>Year Ended December 31,</b>	
	<b>2011</b>	<b>2012</b>
Expected volatility	66.0%	69.0%
Expected term (in years)	6.0	6.0
Risk-free interest rate	1.1%	0.9%
Expected dividend yield	%	%

***Fair Value of Underlying Instrument***

The Company estimates the fair value of its stock-based awards to employees using the Black-Scholes option pricing model.

***Expected Volatility***

The Company estimated the expected volatility based on actual historical volatility of the stock price of similar companies with publicly-traded equity securities. The Company calculated the historical volatility of the selected companies by using daily closing prices over a period of the expected term of the associated award. The companies were selected based on their enterprise value, risk profiles, position within the industry, and with historical share price information sufficient to meet the expected term of the associated award. A decrease in the selected volatility would decrease the fair value of the underlying instrument.

***Expected Term***

The Company estimates the expected life of its employee stock options using the "simplified" method, as prescribed in Staff Accounting Bulletin (SAB) No. 107, whereby, the expected life equals the arithmetic average of the vesting term and the original contractual term of the option due to its lack of sufficient historical data.

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****11. Stock-Based Compensation (continued)*****Risk-Free Interest Rate***

The Company estimated the risk-free interest rate in reference to the yield on U.S. Treasury securities with a maturity date commensurate with the expected term of the associated award. A decrease in the selected risk-free rate would decrease the fair value of the underlying instrument.

***Expected Dividend Yield***

The Company estimated the expected dividend yield based on consideration of its historical dividend experience and future dividend expectations. The Company has not historically declared or paid dividends to stockholders. Moreover, it does not intend to pay dividends in the future, but instead expects to retain any earnings to invest in the continued growth of the business. Accordingly, the Company assumed an expected dividend yield of 0.0%.

**Stock Options**

The following table summarizes the stock option activity under the 2003 Plan during the year ended December 31, 2012 (in thousands):

	Number of Grants	Weighted- Average Exercise Price Per Share	Weighted- Average Contractual Life (in years)	Aggregate Intrinsic Value(1)
Outstanding at December 31, 2011	3,151	\$ 3.76	6.88	\$ 4,968
Granted	722	\$ 5.76		
Exercised	(39)	\$ 4.04		
Canceled or forfeited	(104)	\$ 4.32		
Outstanding at December 31, 2012	3,730	\$ 4.16	6.62	\$ 13,946
Exercisable at December 31, 2012	2,379	\$ 3.56	5.32	\$ 10,250
Vested and expected to vest at December 31, 2012(2)	3,637	\$ 4.12	6.55	\$ 13,722

(1) The aggregate intrinsic value is calculated as the difference between the exercise price of the underlying options and the estimated fair value of the common stock for the options that were in the money at December 31, 2011 and 2012.

(2) This represents the number of vested options at December 31, 2012, plus the number of unvested options expected to vest at December 31, 2012, based on the unvested options outstanding at December 31, 2012, adjusted for the estimated forfeiture rate.

During the years ended December 31, 2011 and 2012, the Company granted stock options to purchase an aggregate of 334,175 and 722,000, shares of its common stock, respectively, with a weighted-average grant date fair value of options granted of \$5.12 and \$7.20, respectively.

During the years ended December 31, 2011 and 2012, current and former employees of the Company exercised a total of 94,748 and 38,697, options, respectively, resulting in total proceeds of \$0.2 million and \$0.2 million, respectively.

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The aggregate intrinsic value of options exercised during the year ended December 31, 2012, was \$47,000.

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Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****11. Stock-Based Compensation (continued)**

As of December 31, 2012, there was \$4.4 million of unrecognized compensation expense related to unvested stock options that is expected to be recognized over a weighted-average period of 2.9 years.

**12. 401(k) Savings Plan**

In 2004, the Company established a defined-contribution savings plan under Section 401(k) of the Internal Revenue Code (the 401(k) Plan). The 401(k) Plan covers all employees who meet defined minimum age and service requirements, and allows participants to defer a portion of their annual compensation on a pretax basis. The Company has not made any contributions to the 401(k) Plan through December 31, 2012.

**13. Income Taxes**

The Company provides for income taxes under ASC 740. Under ASC 740, the liability method is used in accounting for income taxes. Under this method, deferred tax assets and liabilities are determined based on differences between financial reporting and tax bases of assets and liabilities, and are measured using the enacted tax rates and laws that will be in effect when the differences are expected to reverse.

For the years ended December 31, 2011 and 2012, the Company did not record a current or deferred income tax expense or benefit.

The Company's income (loss) before income taxes was \$36.3 million and \$(32.6) million for the years ended December 31, 2011 and 2012, respectively, and was generated entirely in the United States.

Deferred taxes are recognized for temporary differences between the basis of assets and liabilities for financial statement and income tax purposes. The significant components of the Company's deferred tax assets are comprised of the following (in thousands):

	<b>Year Ended December 31,</b>	
	<b>2011</b>	<b>2012</b>
Deferred tax assets:		
U.S. and state net operating loss carryforwards	\$ 20,016	\$ 35,584
Research and development credits	5,383	5,384
Deferred revenue	25,690	21,882
Accruals and other temporary differences	5,889	5,333
Total deferred tax assets	56,978	68,183
Less valuation allowance	(56,978)	(68,183)
Net deferred tax assets	\$	\$

The Company has evaluated the positive and negative evidence bearing upon the realizability of its deferred tax assets. Based on the Company's history of operating losses, the Company has concluded that it is more likely than not that the benefit of its deferred tax assets will not be realized. Accordingly, the Company has provided a full valuation allowance for deferred tax assets as of December 31, 2011 and 2012. The valuation allowance increased by \$11.2 million during the year ended December 31, 2012, due primarily to the generation of net operating losses during the period. The valuation allowance decreased by \$14.3 million during the year ended December 31, 2011, due primarily to the utilization of net operating losses during the period.

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****13. Income Taxes (continued)**

A reconciliation of income tax expense computed at the statutory federal income tax rate to income taxes as reflected in the financial statements is as follows:

	Year Ended December 31,	
	2011	2012
Federal income tax expense at statutory rate	34.0%	34.0%
State income tax, net of federal benefit	5.0%	4.2%
Permanent differences	1.5%	(3.4)%
Research and development credit	(1.0)%	%
Other	%	(0.4)%
Change in valuation allowance	(39.5)%	(34.4)%
<b>Effective income tax rate</b>	<b>0.0%</b>	<b>0.0%</b>

As of December 31, 2011 and 2012, the Company had U.S. federal net operating loss carryforwards of \$53.6 million and \$93.3 million, respectively, which may be available to offset future income tax liabilities and expire at various dates through 2032. As of December 31, 2011 and 2012, the Company also had U.S. state net operating loss carryforwards of \$35.8 million and \$75.4 million, respectively, which may be available to offset future income tax liabilities and expire at various dates through 2032. As a result of the up-front payment pursuant to the Company's collaboration agreement with Celgene, the Company expects that it will use a significant portion of its net operating loss carryforwards in 2011.

As of December 31, 2011 and 2012, the Company had federal research and development tax credit carryforwards of \$3.8 million and \$3.8 million, respectively, available to reduce future tax liabilities which expire at various dates through 2032. As of December 31, 2011 and 2012, the Company had state research and development tax credit carryforwards of approximately \$2.4 million and \$2.4 million, respectively, available to reduce future tax liabilities which expire at various dates through 2027.

Under the provisions of the Internal Revenue Code, the net operating loss and tax credit carryforwards are subject to review and possible adjustment by the Internal Revenue Service and state tax authorities. Net operating loss and tax credit carryforwards may become subject to an annual limitation in the event of certain cumulative changes in the ownership interest of significant shareholders over a three-year period in excess of 50 percent, as defined under Sections 382 and 383 of the Internal Revenue Code, respectively, as well as similar state provisions. This could limit the amount of tax attributes that can be utilized annually to offset future taxable income or tax liabilities. The amount of the annual limitation is determined based on the value of the Company immediately prior to the ownership change. Subsequent ownership changes may further affect the limitation in future years. The Company has completed several financings since its inception which may have resulted in a change in control as defined by Sections 382 and 383 of the Internal Revenue Code, or could result in a change in control in the future.

The Company will recognize interest and penalties related to uncertain tax positions in income tax expense. As of December 31, 2011 and 2012, the Company had no accrued interest or penalties related to uncertain tax positions and no amounts have been recognized in the Company's statements of operations and comprehensive income (loss).

For all years through December 31, 2012, the Company generated research credits but has not conducted a study to document the qualified activities. This study may result in an adjustment to the

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**13. Income Taxes (continued)**

Company's research and development credit carryforwards; however, until a study is completed and any adjustment is known, no amounts are being presented as an uncertain tax position for these two years. A full valuation allowance has been provided against the Company's research and development credits and, if an adjustment is required, this adjustment would be offset by an adjustment to the deferred tax asset established for the research and development credit carryforwards and the valuation allowance.

The Company files income tax returns in the United States, and various state jurisdictions. The federal and state income tax returns are generally subject to tax examinations for the tax years ended December 31, 2009 through December 31, 2012. To the extent the Company has tax attribute carryforwards, the tax years in which the attribute was generated may still be adjusted upon examination by the Internal Revenue Service, state or foreign tax authorities to the extent utilized in a future period.

**14. Long-Term Debt**

On June 26, 2009, the Company entered into a Senior Loan Agreement (the 2009 Senior Loan Agreement) with three lenders that provides for a total funding commitment of \$10.0 million. The Company was required to make payments over 36 months, the first 6 payments of which were interest only, and the principal balance plus accrued interest was payable over the remaining 30 months. Interest accrued at 12.70% per annum. The Company was not subject to any financial covenants under this arrangement. The 2009 Senior Loan Agreement was secured by substantially all of the assets of the Company other than intellectual property and certain permanent capital improvements to the leased facilities. In accordance with the 2009 Senior Loan Agreement, the Company issued warrants to purchase 45,786 shares of Series C-1 Preferred Stock with a fair value at issuance of \$0.3 million. The fair value of the warrants, which was determined using the Black-Scholes option pricing model on the date of issue was treated as a discount to the debt and accreted to interest expense using the effective interest method. As of December 31, 2011 and 2012, the outstanding balance under the 2009 Senior Loan Agreement was \$2.3 million and zero, respectively.

On March 18, 2010, the Company entered into a loan modification agreement (the 2010 Loan Modification Agreement) with the same three lenders as the 2009 Senior Loan Agreement. The 2010 Loan Modification Agreement provides for an additional funding commitment of \$10.0 million. As of December 31, 2011 and 2012, the outstanding balance under the 2010 Loan Modification Agreement was \$3.2 million and zero, respectively. The Company was required to make payments over 27 months, the first 3 payments of which were interest only, and the principal balance plus accrued interest was payable over the remaining 24 months. Interest accrued at 15.00% per annum. The Company was not subject to any financial covenants under this arrangement. The 2010 Loan Modification Agreement was secured by substantially all of the assets of the Company other than intellectual property and certain permanent capital improvements to the leased facilities. In accordance with the 2010 Loan Modification Agreement, the Company issued warrants to purchase 63,693 shares of Series D-1 Preferred Stock with a fair value at issuance of \$0.5 million. The fair value of the warrants, which was determined using the Black-Scholes option pricing model, on the date of issue was treated as a discount to the debt and accreted to interest expense using the effective interest method.

On June 7, 2012, the Company entered into a loan and security agreement (the Loan Agreement) with the same three lenders, pursuant to which the Company received a loan in the aggregate principal amount of \$20.0 million. The Company is required to repay the aggregate principal balance under the Loan Agreement in 42 months. The first 12 payments are interest only and the remaining 30 payments

Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****14. Long-Term Debt (continued)**

are equal monthly installments of principal plus interest. The Loan Agreement provided that the interest only period could be extended under certain circumstances. The Company did not trigger the requirements and will begin paying principal in July 2013.

Per annum interest is payable at the 8.5%. The Loan Agreement also included a closing fee of \$0.2 million. The Company is amortizing the cost over the 42 months of loan. The Loan Agreement is also subject to an additional deferred payment of \$1.2 million due with the final payment. The Company is recording the deferred payment to interest expense over the term of the Loan Agreement. The resulting effective interest rate is approximately 11.8%. The company is not subject to any financial covenants and the Loan Agreement is secured by a lien on all of the Company's personal property as of, or acquired after, the date of the Loan Agreement, except for intellectual property.

The Loan Agreement defines events of default, including the occurrence of an event that results in a material adverse effect upon the Company's business operations, properties, assets or condition (financial or otherwise), its ability to perform its obligations under and in accordance with the terms of the Loan Agreement, or upon the ability of the lenders to enforce any of their rights or remedies with respect to such obligations, or upon the collateral under the Loan Agreement or upon the liens of the lenders on such collateral or upon the priority of such liens. The lenders also received a right, to purchase at fair value, up to \$2.0 million of equity of the Company sold in any sale by the Company to third parties of equity securities resulting in at least \$5.0 million in net cash proceeds to the Company, subject to certain exceptions. As of December 31, 2012, there have been no events of default under the loan. As of December 31, 2012, the principal balance outstanding was \$20.0 million.

At December 31, 2012, future minimum payments related to long-term debt were as follows (in thousands):

Year ending December 31:	
2013	\$ 5,304
2014	8,908
2015	10,108
Less amounts representing interest	(3,120)
Less Deferred Fee	(1,200)
Future minimum principal payments	20,000
Less current portion	3,668
Noncurrent financing obligations	\$ 16,332

**15. Related Party Transactions****Celgene Corporation (Celgene)**

In connection with its entry into the collaboration agreement with Celgene, on February 2008, the Company sold Celgene 457,875 shares of its Series C-1 Preferred Stock. As part of the Company's June 2010 Series E financing, Celgene purchased 36,496 shares of Series E Preferred Stock and received warrants to purchase 38,979 shares of common stock. As part of the Company's December 2011 Series F financing, Celgene purchased 1,990,446 shares of Series F Preferred Stock. As a result of these transactions, Celgene owned 9.9% of the Company's fully diluted equity as of December 31, 2012. Refer to Note 10 for additional information regarding this collaboration agreement.



Table of Contents**Acceleron Pharma Inc.****Notes to Financial Statements (continued)****15. Related Party Transactions (continued)**

During the year ended December 31, 2012, the Company recognized \$4.9 million in collaboration revenue under the Celgene collaboration arrangement and, as of December 31, 2012, had \$10.3 million of deferred revenue related to the Celgene collaboration arrangement.

The Company recognized revenue from Celgene during the years ended December 31, 2011 and 2012 as follows (in thousands):

	Year Ended December 31,	
	2011	2012
License and milestone	\$ 63,607	\$ 2,035
Cost sharing, net	(121)	2,879
	\$ 63,486	\$ 4,914

**Alkermes**

One of the Company's directors is also the Chairman, President, and Chief Executive Officer of Alkermes plc, the parent company of Alkermes, with which the Company entered into a collaboration agreement during 2009.

As of December 31, 2012, Alkermes held 695,250 shares of the Company's Preferred Stock and warrants to purchase 42,624 shares of common stock. For the year ended December 31, 2011, Alkermes paid the Company \$0.7 million for research services. No such fees were paid to the Company during 2012.

***Related-Party Receivable***

On January 28, 2008, the Company issued a secured promissory note (the Note Receivable) in the amount of \$0.2 million to the current chief executive officer of the Company (the CEO). The Note Receivable bears interest at an annual interest rate of 3.11% and was initially repayable on the earlier of January 28, 2011, or the date prior to the date that the Company files a registration statement with the SEC, covering shares of its common stock. The Note Receivable is secured by shares of the Company's common stock owned by the CEO. On December 22, 2010, the term was extended until January 28, 2014, or the date prior to the date that the Company files a registration statement with the SEC covering shares of its common stock.

In November 2012, the Company further modified the terms of the Note Receivable, such that in the event that an acquisition event occurs or the company files a registration statement with the SEC on or before the maturity date, the unpaid principal and interest will be forgiven. The Company evaluated the forgiveness provisions and determined that forgiveness was not probable as of December 31, 2012, and as such, continued to record the Note Receivable as an asset at December 31, 2012. As a result of the Company's filing of a registration statement with the SEC on August 6, 2013 which triggered the forgiveness of the Note Receivable, the Company expensed the unpaid principal and interest expense totaling \$0.2 million as compensation expense during 2013.

**16. Subsequent Events**

The Company has completed an evaluation of all subsequent events after the audited balance sheet date of December 31, 2012 through September 5, 2013, to ensure that this filing includes appropriate disclosure of events both recognized in the financial statements as of December 31, 2012,

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**Acceleron Pharma Inc.**

**Notes to Financial Statements (continued)**

**16. Subsequent Events (continued)**

and events which occurred subsequently but were not recognized in the financial statements. The Company has concluded that no subsequent events have occurred that require disclosure, except as disclosed within these financial statements and except as described below.

On September 4, 2013, the Board approved the following actions, which were approved by the stockholders on the same day:

A 1-for-4 reverse stock split of the Company's common stock and redeemable convertible preferred stock, which was effective on September 5, 2013. All share and per share data shown in the accompanying financial statements and related notes have been retroactively revised to reflect the reverse stock split.

The adoption of the 2013 Equity Incentive Plan (the 2013 Plan). The Company has reserved for issuance an aggregate of 1,500,000 shares of common stock under the 2013 Plan which is comprised of (i) the remaining 155,884 shares reserved for issuance under the 2003 Plan and (ii) an additional 1,344,116 shares. The 2013 Plan provides that the number of shares reserved and available for issuance under the plan will automatically increase each January 1, beginning in 2014, by the lesser of (i) 3,150,000 shares, or (ii) 4% of the outstanding number of shares of the Company's common stock on the immediately preceding December 31st. This number is subject to adjustment in the event of a stock split, stock dividend or other change in the Company's capitalization.

The adoption of the 2013 Employee Stock Purchase Plan (the 2013 ESPP). Under the 2013 ESPP, 275,000 shares of the Company's common stock will be available for issuance and eligible employees of the Company may purchase shares of common stock during pre-specified purchase periods at a price equal to the lesser of 85% of the fair market value of a share of its common stock at the beginning of the purchase period or 85% of the fair market value of a share of its common stock at the end of the purchase period. The Board has not determined the date on which the initial purchase period will commence under the 2013 ESPP, although the initial purchase period will not commence prior to the completion of the Company's IPO.

On September 4, 2013, the Board also approved for filing immediately following the effectiveness of the Company's registration statement in connection with its IPO, the Restated Certificate of Incorporation to increase the authorized number of shares of common stock from 104,013,161 to 175,000,000, to authorize 25,000,000 shares of undesignated preferred stock, par value \$0.001 per share, and to eliminate all references to the previously designated Series Preferred Stock. This Restated Certificate of Incorporation was approved by the stockholders on September 4, 2013.

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**Acceleron Pharma Inc.**

**Index to Unaudited Interim Condensed Financial Statements**

	<b>Pages</b>
<u>Condensed Balance Sheets as of September 30, 2013 and December 31, 2012</u>	<u>F-52</u>
<u>Condensed Statements of Operations and Comprehensive Income (loss) for the Three and Nine Months Ended September 30, 2013 and 2012</u>	<u>F-53</u>
<u>Condensed Statements of Cash Flows for the Nine Months Ended September 30, 2013 and 2012</u>	<u>F-54</u>
<u>Notes to Unaudited Interim Condensed Financial Statements</u>	<u>F-55</u>
F-51	

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Table of Contents**Acceleron Pharma Inc.****Condensed Balance Sheets****(amounts in thousands except share and per share data)****(unaudited)**

	<b>September 30, 2013</b>	<b>December 31, 2012</b>
<b>Assets</b>		
Current assets:		
Cash and cash equivalents	\$ 116,479	\$ 39,611
Collaboration receivables (includes related party amounts of \$3,713 and \$1,840 at September 30, 2013 and December 31, 2012, respectively)	4,103	2,776
Prepaid expenses and other current assets	2,179	1,474
Total current assets	122,761	43,861
Property and equipment, net	3,564	4,059
Restricted cash	913	913
Related party receivables		233
Other assets	22	146
Total assets	\$ 127,260	\$ 49,212
<b>Liabilities, redeemable convertible preferred stock and stockholders' equity (deficit)</b>		
Current liabilities:		
Accounts payable	\$ 891	\$ 642
Accrued expenses (includes related party amounts of \$0 and \$861 at September 30, 2013 and December 31, 2012, respectively)	5,126	6,153
Deferred revenue	2,351	27,840
Deferred rent	499	499
Notes payable, net of discount	7,656	3,668
Total current liabilities	16,523	38,802
Deferred revenue, net of current portion	6,205	6,760
Deferred rent, net of current portion	2,463	2,837
Notes payable, net of current portion and discount	10,979	16,525
Warrants to purchase redeemable convertible preferred stock		1,422
Warrants to purchase common stock	16,526	5,229
Total liabilities	52,696	71,575
Commitments and contingencies (Note 13)		
Redeemable convertible preferred stock		268,610
Stockholders' equity (deficit):		
Undesignated preferred stock, \$0.001 par value: 25,000,000 shares authorized and no shares issued or outstanding at September 30, 2013; No shares authorized, issued or outstanding at December 2012		
Common stock, \$0.001 par value: 175,000,000 and 104,013,161 shares authorized at September 30, 2013 and December 31, 2012, respectively; 28,069,579, and 2,432,155 shares issued and outstanding at September 30, 2013 and December 31, 2012, respectively	35	3
Additional paid-in capital	248,750	
Accumulated deficit	(174,221)	(290,976)
Total stockholders' equity (deficit)	74,564	(290,973)

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Total liabilities, redeemable convertible preferred stock and stockholders' equity (deficit)	\$	127,260	\$	49,212
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See accompanying notes to these condensed financial statements.

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## Acceleron Pharma Inc.

## Condensed Statements of Operations and Comprehensive Loss

(amounts in thousands except per share data)

(unaudited)

	Three Months Ended September 30,		Nine Months Ended September, 30,	
	2013	2012	2013	2012
<b>Revenue:</b>				
Collaboration revenue:				
License and milestone	\$ 638	\$ 2,461	\$ 36,044	\$ 7,226
Cost-sharing, net	3,632	1,444	9,666	4,043
<b>Total revenue(1)</b>	<b>4,270</b>	<b>3,905</b>	<b>45,710</b>	<b>11,269</b>
<b>Costs and expenses:</b>				
Research and development	8,143	8,722	25,834	25,646
General and administrative	3,011	2,041	9,472	6,318
<b>Total costs and expenses</b>	<b>11,154</b>	<b>10,763</b>	<b>35,306</b>	<b>31,964</b>
(Loss) income from operations	(6,884)	(6,858)	10,404	(20,695)
<b>Other (expense) income:</b>				
Other (expense) income, net	(11,149)	132	(12,571)	(565)
Interest income	5	22	25	75
Interest expense	(485)	(511)	(1,646)	(1,018)
<b>Total other expense, net</b>	<b>(11,629)</b>	<b>(357)</b>	<b>(14,192)</b>	<b>(1,508)</b>
<b>Net loss</b>	<b>\$ (18,513)</b>	<b>\$ (7,215)</b>	<b>\$ (3,788)</b>	<b>\$ (22,203)</b>
<b>Comprehensive loss</b>	<b>\$ (18,513)</b>	<b>\$ (7,215)</b>	<b>\$ (3,788)</b>	<b>\$ (22,203)</b>
<b>Reconciliation of net loss to net loss applicable to common stockholders:</b>				
Net loss	\$ (18,513)	\$ (7,215)	\$ (3,788)	\$ (22,203)
Accretion of dividends, interest, redemption value and issuance costs on redeemable convertible preferred stock	(6,272)	(6,747)	(19,870)	(20,293)
Gain on extinguishment of redeemable convertible preferred stock			2,765	
<b>Net loss applicable to common stockholders basic and diluted</b>	<b>\$ (24,785)</b>	<b>\$ (13,962)</b>	<b>\$ (20,893)</b>	<b>\$ (42,496)</b>
<b>Net loss per share applicable to common stockholders: (Note 8)</b>				
Basic and diluted	\$ (5.62)	\$ (5.82)	\$ (6.74)	\$ (17.73)
<b>Weighted-average number of common shares used in computing net loss per share applicable to common stockholders:</b>				
Basic and diluted	4,406	2,400	3,100	2,397

(1) Includes related party revenue (Note 18) \$ 4,270 \$ 1,381 \$ 20,763 \$ 3,597

See accompanying notes to these condensed financial statements.

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Table of Contents**Acceleron Pharma Inc.****Condensed Statements of Cash Flows**

(amounts in thousands)

(unaudited)

	<b>Nine Months Ended September 30,</b>	
	<b>2013</b>	<b>2012</b>
<b>Operating Activities</b>		
Net loss	\$ (3,788)	\$ (22,203)
Adjustments to reconcile net loss to net cash used in operating activities:		
Depreciation and amortization	681	1,086
Stock-based compensation	1,441	861
Amortization of debt discount		51
Accretion of deferred interest	257	250
Amortization of deferred debt issuance costs	182	64
Change in fair value of warrants	12,649	565
Gain on retirement of warrants	(76)	
Forgiveness of related party receivable	237	
Changes in assets and liabilities:		
Prepaid expenses and other current assets	(762)	(1,323)
Collaboration receivables	(1,327)	(1,014)
Related party receivable	(4)	(6)
Accounts payable	243	(894)
Accrued expenses	(1,602)	712
Deferred revenue	(26,044)	(7,226)
Deferred rent	(373)	(358)
Net cash used in operating activities	(18,286)	(29,435)
<b>Investing Activities</b>		
Purchases of property and equipment	(187)	(322)
Net cash used in investing activities	(187)	(322)
<b>Financing Activities</b>		
Proceeds from issuance of common stock from initial public offering, net issuance costs	87,406	
Proceeds from issuance of common stock from private placement	10,000	
Proceeds from long-term debt, net of issuance costs		19,945
Payments of long-term debt	(1,815)	(6,191)
Payments made to repurchase redeemable convertible preferred stock, common stock and warrants to purchase common stock	(300)	
Proceeds from exercise of stock options and warrants to purchase common stock	50	47
Net cash provided by financing activities	95,341	13,801
Net increase (decrease) in cash and cash equivalents	76,868	(15,956)
Cash and cash equivalents at beginning of period	39,611	65,037
Cash and cash equivalents at end of period	\$ 116,479	\$ 49,081
<b>Supplemental Disclosure of Cash Flow Information:</b>		
Cash paid for interest	\$ 1,262	\$ 640



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**Supplemental Disclosure of Non-Cash Investing and Financing Activities:**

Accretion of dividends, interest, redemption value, and issuance costs on preferred stock	\$ 19,870	\$ 20,293
Cashless exercise of warrants	\$ 678	\$
Initial public offering costs included in accounts payable and accrued expense	\$ 582	\$
Reclassification of warrant liability to additional paid-in capital	\$ 2,013	\$
Conversion of redeemable convertible preferred stock into common stock	\$ 286,094	\$

See accompanying notes to these condensed financial statements.

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements**

**1. Nature of Business**

Acceleron Pharma Inc. (Acceleron or the Company) was incorporated in the state of Delaware on June 13, 2003, as Phoenix Pharma, Inc. The Company subsequently changed its name to Acceleron Pharma Inc. and commenced operations in February 2004. The Company is a Cambridge, Massachusetts-based biopharmaceutical company focused on the discovery, development and commercialization of novel protein therapeutics for cancer and rare diseases. The Company's research focuses on the biology of the Transforming Growth Factor-Beta (TGF- $\beta$ ) protein superfamily, a large and diverse group of molecules that regulate the growth and repair of tissues throughout the human body. By coupling its discovery and development expertise, including its proprietary knowledge of the TGF- $\beta$  superfamily, with internal protein engineering and manufacturing capabilities, the Company has built a highly productive research and development platform that has generated numerous innovative protein therapeutics with novel mechanisms of action. The Company has internally discovered three protein therapeutics that are currently being studied in 12 ongoing Phase 2 clinical trials, focused on the areas of cancer and rare diseases.

The Company is subject to risks common to companies in the biotechnology industry, including, but not limited to, risk that the Company never achieves profitability, the need for substantial additional financing, risk of relying on third parties, risks of clinical trial failures, dependence on key personnel, protection of proprietary technology and compliance with government regulations.

**2. Basis of Presentation**

The accompanying financial statements have been prepared in conformity with accounting principles generally accepted in the United States of America (GAAP). Any reference in these notes to applicable guidance is meant to refer to the authoritative United States generally accepted accounting principles as found in the Accounting Standards Codification (ASC) and Accounting Standards Update (ASU) of the Financial Accounting Standards Board (FASB).

The accompanying interim balance sheet as of September 30, 2013, the statements of operations and comprehensive loss for the three and nine months ended September 30, 2013 and 2012 and statements of cash flows for the nine months ended September 30, 2013 and 2012, and the financial data and other information disclosed in these notes related to the nine months ended September 30, 2013 and 2012 are unaudited. The unaudited interim financial statements have been prepared on the same basis as the audited annual financial statements as of and for the year ended December 31, 2012, and, in the opinion of management, reflect all adjustments, consisting of normal recurring adjustments, necessary for the fair presentation of the Company's financial position as of September 30, 2013, and the results of its operations and its cash flows for the three and nine months ended September 30, 2013 and 2012.

The results for the nine months ended September 30, 2013 are not necessarily indicative of the results to be expected for the year ending December 31, 2013, any other interim periods, or any future year or period. These interim financial statements should be read in conjunction with the audited financial statements as of and for the year ended December 31, 2012, and the notes thereto, which are included in the Company's Prospectus that forms a part of the Company's Registration Statement on Form S-1 (File No. 333-190417), which was filed with the Securities and Exchange Commission (the SEC) pursuant to Rule 424(b) on September 19, 2013 (the Prospectus).

On September 24, 2013 the Company completed its initial public offering (IPO) whereby the Company sold 6,417,000 shares of common stock (including 837,000 shares of common stock sold by

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**2. Basis of Presentation (continued)**

the Company pursuant to the full exercise of an overallotment option by the underwriters in connection with the offering) at a price of \$15.00 per share. The shares began trading on the Nasdaq Global Select Market on September 19, 2013. The aggregate net proceeds received by the Company from the offering were \$86.8 million, net of underwriting discounts and commissions and estimated offering expenses payable by the Company. Upon the closing of the IPO, all outstanding shares of convertible preferred stock converted into 18,516,993 shares of common stock and warrants exercisable for convertible preferred stock were automatically converted into warrants exercisable for 141,370 shares of common stock, resulting in the reclassification of the related convertible preferred stock warrant liability of \$2.0 million to additional paid-in capital. Additionally, the Company is now authorized to issue 175,000,000 shares of common stock and 25,000,000 shares of undesignated preferred stock.

On September 24, 2013 the Company also completed the sale of a private placement of 666,667 shares of common stock to Celgene Corporation at the IPO price of \$15.00 per share concurrent with and at the same offer price as the IPO. The aggregate net proceeds received by the Company from the concurrent private placement were \$10.0 million.

On August 23, 2013, the board of directors (the Board) and the stockholders of the Company approved a one-for-four reverse stock split of the Company's outstanding common stock, which was effected on September 3, 2013. Stockholders entitled to fractional shares as a result of the reverse stock split will receive a cash payment in lieu of receiving fractional shares. The Company's historical share and per share information have been retroactively adjusted to give effect to this reverse stock split. Shares of common stock underlying outstanding stock options and other equity instruments were proportionately reduced and the respective exercise prices, if applicable, were proportionately increased in accordance with the terms of the agreements governing such securities.

The accompanying condensed financial statements reflect the application of certain significant accounting policies as described below and elsewhere in these notes to the financial statements. As of September 30, 2013, the Company's significant accounting policies and estimates, which are detailed in the Company's Prospectus, have not changed.

**3. Use of Estimates**

The preparation of financial statements in conformity with GAAP requires management to make estimates and assumptions that affect the reported amounts of assets and liabilities, the disclosure of contingent assets and liabilities at the date of the financial statements, and the reported amounts expensed during the reporting period. Actual results could materially differ from those estimates.

Management considers many factors in selecting appropriate financial accounting policies and controls, and in developing the estimates and assumptions that are used in the preparation of these financial statements. Management must apply significant judgment in this process. In addition, other factors may affect estimates, including: expected business and operational changes, sensitivity and volatility associated with the assumptions used in developing estimates, and whether historical trends are expected to be representative of future trends. The estimation process often may yield a range of potentially reasonable estimates of the ultimate future outcomes and management must select an amount that falls within that range of reasonable estimates. This process may result in actual results differing materially from those estimated amounts used in the preparation of the financial statements if these results differ from historical experience, or other assumptions do not turn out to be substantially accurate, even if such assumptions are reasonable when made. In preparing these financial statements,

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**3. Use of Estimates (continued)**

management used significant estimates in the following areas, among others: revenue recognition, stock-based compensation expense, the determination of the fair value of stock-based awards, the fair value of liability-classified warrants, accrued expenses, and the recoverability of the Company's net deferred tax assets and related valuation allowance.

The Company utilized significant estimates and assumptions in determining the fair value of its common stock prior to the completion of the IPO. The Board determined the estimated fair value of the Company's common stock based on a number of objective and subjective factors, including external market conditions affecting the biotechnology industry sector and the prices at which the Company sold shares of redeemable convertible preferred stock, the superior rights and preferences of securities senior to the Company's common stock at the time, and the likelihood of achieving a liquidity event, such as an IPO or sale of the Company.

**4. Segment Information**

Operating segments are identified as components of an enterprise about which separate discrete financial information is available for evaluation by the chief operating decision maker, or decision making group, in making decisions on how to allocate resources and assess performance. The Company's chief operating decision maker is the chief executive officer. The Company and the chief executive officer view the Company's operations and manage its business as one operating segment. All material long-lived assets of the Company reside in the United States. The Company does use contract research organizations (CROs) and research institutions located outside the United States. Some of these expenses are subject to collaboration reimbursement which is presented as a component of cost sharing, net in the statement of operations and comprehensive loss.

**5. Cash and Cash Equivalents and Restricted cash**

The Company considers all highly liquid investments purchased with original maturities of 90 days or less at acquisition to be cash equivalents. Cash and cash equivalents include cash held in banks and amounts held in interest-bearing money market accounts. Cash equivalents are carried at cost, which approximates their fair market value. As of September 30, 2013 and December 31, 2012, the Company maintained letters of credit totaling \$0.9 million held in the form of a money market account as collateral for the Company's facility lease obligations and its credit cards.

**6. Concentrations of Credit Risk and Off-Balance Sheet Risk**

The Company has no off-balance sheet risk, such as foreign exchange contracts, option contracts, or other foreign hedging arrangements. Financial instruments that potentially subject the Company to concentrations of credit risk are primarily cash, cash equivalents, restricted cash and accounts receivable. The Company maintains its cash and cash equivalent balances in the form of money market accounts with financial institutions that management believes are creditworthy. The Company's investment policy includes guidelines on the quality of the institutions and financial instruments and defines allowable investments that the Company believes minimizes the exposure to concentration of credit risk.

The Company routinely assesses the creditworthiness of its customers and collaboration partners. The Company has not experienced any material losses related to receivables from individual customers and collaboration partners, or groups of customers. The Company does not require collateral. Due to

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**6. Concentrations of Credit Risk and Off-Balance Sheet Risk (continued)**

these factors, no additional credit risk beyond amounts provided for collection losses is believed by management to be probable in the Company's accounts receivable.

**7. Fair Value Measurements**

ASC Topic 820, *Fair Value Measurement* (ASC 820), establishes a fair value hierarchy for instruments measured at fair value that distinguishes between assumptions based on market data (observable inputs) and the Company's own assumptions (unobservable inputs). Observable inputs are inputs that market participants would use in pricing the asset or liability based on market data obtained from sources independent of the Company. Unobservable inputs are inputs that reflect the Company's assumptions about the inputs that market participants would use in pricing the asset or liability, and are developed based on the best information available in the circumstances.

ASC 820 identifies fair value as the exchange price, or exit price, representing the amount that would be received to sell an asset or paid to transfer a liability in an orderly transaction between market participants. As a basis for considering market participant assumptions in fair value measurements, ASC Topic 820 establishes a three-tier fair value hierarchy that distinguishes between the following:

Level 1 Quoted market prices in active markets for identical assets or liabilities.

Level 2 Inputs other than Level 1 inputs that are either directly or indirectly observable, such as quoted market prices, interest rates, and yield curves.

Level 3 Unobservable inputs developed using estimates of assumptions developed by the Company, which reflect those that a market participant would use.

To the extent that the valuation is based on models or inputs that are less observable or unobservable in the market, the determination of fair value requires more judgment. Accordingly, the degree of judgment exercised by the Company in determining fair value is greatest for instruments categorized in Level 3. A financial instrument's level within the fair value hierarchy is based on the lowest level of any input that is significant to the fair value measurement.

Items measured at fair value on a recurring basis include warrants to purchase redeemable convertible preferred stock and warrants to purchase common stock (Note 7). During the periods presented, the Company has not changed the manner in which it values assets and liabilities that are measured at fair value using Level 3 inputs.

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## Acceleron Pharma Inc.

## Notes to Unaudited Interim Condensed Financial Statements (continued)

## 7. Fair Value Measurements (continued)

The following tables set forth the Company's financial instruments carried at fair value using the lowest level of input applicable to each financial instrument as of September 30, 2013 and December 31, 2012 (in thousands):

	September 30, 2013			
	Quoted Prices in Active Markets for Identical Items (Level 1)	Significant Other Observable Inputs (Level 2)	Significant Unobservable Inputs (Level 3)	Total
<b>Assets:</b>				
Money market funds	\$ 116,382	\$	\$	\$ 116,382
Restricted cash	913			913
Total assets	\$ 117,295	\$	\$	\$ 117,295
<b>Liabilities:</b>				
Warrants to purchase redeemable convertible preferred stock	\$	\$	\$	\$
Warrants to purchase common stock			16,526	16,526
Total liabilities	\$	\$	\$ 16,526	\$ 16,526
	December 31, 2012			
	Quoted Prices in Active Markets for Identical Items (Level 1)	Significant other Observable Inputs (Level 2)	Significant Unobservable Inputs (Level 3)	Total
<b>Assets:</b>				
Money market funds	\$ 36,847	\$	\$	\$ 36,847
Restricted cash	913			913
Total assets	\$ 37,760	\$	\$	\$ 37,760
<b>Liabilities:</b>				
Warrants to purchase redeemable convertible preferred stock	\$	\$	\$ 1,422	\$ 1,422
Warrants to purchase common stock			5,229	5,229
Total liabilities	\$	\$	\$ 6,651	\$ 6,651

Table of Contents**Accelaron Pharma Inc.****Notes to Unaudited Interim Condensed Financial Statements (continued)****7. Fair Value Measurements (continued)**

The following table sets forth a summary of changes in the fair value of the Company's preferred and common stock warrant liability, which have been classified within Level 3 of the fair value hierarchy, wherein fair value is estimated using significant unobservable inputs (in thousands):

	Three Months Ended September 30,		Nine Months Ended September 30,	
	2013	2012	2013	2012
Beginning balance	\$ 7,390	\$ 5,089	\$ 6,651	\$ 4,393
Change in fair value	11,149	(132)	12,649	564
Exercises			(678)	
Repurchases			(83)	
Conversions	(2,013)		(2,013)	
Ending balance	\$ 16,526	\$ 4,957	\$ 16,526	\$ 4,957

The money market funds noted above are included in cash and cash equivalents in the accompanying balance sheets. The Company recognizes transfers between levels of the fair value hierarchy as of the end of the reporting period. There were no transfers within the hierarchy during the nine months ended September 30, 2013 or the year ended December 31, 2012 except for the transfer out of the warrants to purchase redeemable convertible preferred stock as described below.

During the three and nine months ended September 30, 2013, as a result of the closing of the IPO, the warrants to purchase preferred stock were converted to warrants to purchase common stock. The resulting warrants to purchase common stock meet the criteria to be classified as permanent equity and are no longer required to be measured at fair value at each reporting period.

The fair value of the warrants to purchase preferred stock that were classified as liabilities was estimated using the Black-Scholes option pricing model at the date of issuance and on each re-measurement date. This method of valuation involves using inputs such as the fair value of the Company's various classes of preferred stock, stock price volatility, the contractual term of the warrants, risk free interest rates, and dividend yields. Due to the nature of these inputs, the valuation of the warrants is considered a Level 3 measurement. See Note 12 for further discussions of the accounting for the warrants, as well as for a summary of the significant inputs and assumptions used to determine the fair value of the warrants.

The fair value of warrants to purchase common stock that are classified as liabilities is estimated using a Monte Carlo model. This method of valuation involves using inputs such as the fair value of a share of common stock, stock price volatility, and the contractual term of the warrants. Due to the nature of these inputs, the valuation of the warrants is considered a Level 3 measurement.

The Company measures eligible assets and liabilities at fair value, with changes in value recognized in earnings. Fair value treatment may be elected either upon initial recognition of an eligible asset or liability or, for an existing asset or liability, if an event triggers a new basis of accounting. The Company did not elect to remeasure any of its existing financial assets or liabilities, and did not elect the fair value option for any financial assets and liabilities transacted in the nine months ended September 30, 2013 or the year ended December 31, 2012.

Table of Contents**Acceleron Pharma Inc.****Notes to Unaudited Interim Condensed Financial Statements (continued)****7. Fair Value Measurements (continued)**

The Company accounts for uncertain tax positions in accordance with the provisions of ASC 740. When uncertain tax positions exist, the Company recognizes the tax benefit of tax positions to the extent that the benefit will more likely than not be realized. The determination as to whether the tax benefit will more likely than not be realized is based upon the technical merits of the tax position as well as consideration of the available facts and circumstances. As of September 30, 2013 and December 31, 2012, the Company does not have any significant uncertain tax positions.

**8. Net Loss Per Share**

The following common stock equivalents were excluded from the calculation of diluted net loss per share for the periods indicated because including them would have had an anti-dilutive effect (in thousands):

	<b>Three Months Ended September 30,</b>		<b>Nine Months Ended September 30,</b>	
	<b>2013</b>	<b>2012</b>	<b>2013</b>	<b>2012</b>
Outstanding stock options	3,667	3,352	3,690	3,232
Common stock warrants	881	884	874	884
Preferred stock	16,658	18,166	17,609	18,166
Preferred stock warrants	130	248	152	248
	21,336	22,650	22,325	22,530

**9. Comprehensive Income (Loss)**

Comprehensive income (loss) is defined as the change in equity of a business enterprise during a period from transactions, other events, and circumstances from non-owner sources. Comprehensive income (loss) consists of net income (loss) and other comprehensive income (loss), which includes certain changes in equity that are excluded from net income (loss). Comprehensive loss has been disclosed in the accompanying statements of operations and comprehensive income (loss) and equals the Company's net loss for all periods presented.

**10. Subsequent Events**

The Company considers events or transactions that occur after the balance sheet date but prior to the issuance of the financial statements to provide additional evidence relative to certain estimates or to identify matters that require additional disclosure. The Company has evaluated all subsequent events and determined that there are no material recognized or unrecognized subsequent events requiring disclosure.

**11. Recently Adopted Accounting Pronouncements**

From time to time, new accounting pronouncements are issued by the FASB or other standard setting bodies and adopted by the Company as of the specified effective date. Unless otherwise discussed, the Company believes that the impact of recently issued standards that are not yet effective will not have a material impact on its financial position or results of operations upon adoption.



Table of Contents**Accelaron Pharma Inc.****Notes to Unaudited Interim Condensed Financial Statements (continued)****12. Warrants**

Below is a summary of the number of shares issuable upon exercise of outstanding warrants and the terms and accounting treatment for the outstanding warrants (in thousands, except per share data):

	Warrants as of		Weighted-Average Exercise Price Per Share	Expiration	Balance Sheet Classification	
	September 30, 2013	December 31, 2012			September 30, 2013	December 31, 2012
Warrant to purchase Series A Preferred Stock		107	\$ 4.00	February 28, 2013	N/A(1)	Liability
Warrants to purchase Series B Preferred Stock		32	7.40	December 21, 2013	N/A(2)	Liability
Warrants to purchase Series C-1 Preferred Stock		46	10.92	June 25, 2019	N/A(2)	Liability
Warrants to purchase Series D-1 Preferred Stock		64	12.56	March 18, 2020	N/A(2)	Liability
Warrants to purchase Common Stock	32		7.40	December 21, 2013	Equity(2)	N/A
Warrants to purchase Common Stock	46		10.92	June 25, 2019	Equity(2)	N/A
Warrants to purchase Common Stock	64		12.56	March 18, 2020	Equity(2)	N/A
Warrants to purchase Common stock	858	872	5.88	June 10, 2020 - July 9, 2020	Liability	Liability
Warrants to purchase Common stock	13	13	4.00 - 7.40	March 31, 2015 - December 31, 2017	Equity(3)	Equity
All warrants	1,013	1,134	\$ 6.56			

- (1) On February 6, 2013, the warrant holder exercised a warrant to purchase 107 shares of Series A Preferred Stock on a net basis, resulting in the issuance of 47 shares of Series A Preferred Stock.
- (2) Warrants to purchase Series B Preferred Stock, Series C-1 Preferred Stock, and Series D-1 Preferred Stock were converted to warrants to purchase common stock at the closing of the IPO on September 24, 2013.
- (3) Warrants to purchase common stock were issued in connection with various debt financing transactions that were consummated in periods prior to December 31, 2012. See discussion below for further details.

In connection with various financing transactions that were consummated in periods prior to December 31, 2012, the Company issued warrants for the purchase of up to 106,500 shares of the Company's Series A redeemable convertible preferred stock (Series A Preferred Stock), 31,891 shares of the Company's Series B redeemable convertible preferred stock (Series B Preferred Stock), 45,786 shares of the Company's Series C-1 redeemable convertible preferred stock (Series C-1 Preferred Stock), and 63,693 shares of the Company's Series D-1 redeemable convertible preferred stock (Series D-1 Preferred Stock). Each warrant was immediately exercisable. The warrants to purchase Series A and Series B Preferred Stock expire seven years from the original date of issuance, while the warrants to purchase Series C-1 and Series D-1 Preferred Stock expire ten years from the original date of issuance. The warrants to purchase shares of the Company's preferred stock have an exercise price equal to the original issuance price of the underlying instrument. Each warrant is exercisable on either a physical settlement or net share settlement basis and the redemption provisions are outside the control of the Company. In connection with the closing of the Company's IPO on September 24, 2013, the outstanding warrants to purchase Series B Preferred Stock, Series C-1 Preferred Stock, and Series D-1 Preferred

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Stock were converted into warrants to purchase common stock The exercise prices for each of these warrants remained unchanged.

The Company follows the provisions of ASC Topic 480, *Issuer's Accounting for Freestanding Warrants and Other Similar Instruments on Shares that Are Redeemable*, which requires that warrants to purchase redeemable preferred stock be classified as liabilities. In addition, the value of the warrants is remeasured to the then-current fair value at each reporting date. Changes in fair value are recorded to other income (expense), net. For the three months ended September 30, 2013 and 2012 and the nine

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Table of Contents**Acceleron Pharma Inc.****Notes to Unaudited Interim Condensed Financial Statements (continued)****12. Warrants (continued)**

months ended September 30, 2013 and 2012, the Company remeasured the fair value of all of its outstanding warrants to purchase shares of the Company's preferred stock up until the conversion of such warrants on September 24, 2013, using current assumptions, resulting in an increase in fair value of \$1.0 million, \$0.0 million, \$1.3 million and \$0.0 million, respectively, which was recorded in other expense, net in the accompanying statements of operations and comprehensive loss. As a result of the closing of the IPO and the resulting conversion of the warrants to purchase preferred shares into warrants to purchase common stock, the fair value of the warrant liability at September 24, 2013 was reclassified to permanent equity and therefore, is no longer subject to remeasurement.

In December 2012, the Company modified the warrant to purchase 106,500 shares of Series A Preferred Stock and extended the expiration date from December 21, 2012 to February 28, 2013. During the nine months ended September 30, 2013, the holder of the warrant exercised the warrant on a net basis, resulting in the issuance of 46,668 shares of Series A Preferred Stock. Upon exercise, the Company re-measured the fair value of the warrant and recorded the resulting increase in fair value of \$0.1 million as other expense in the accompanying statement of operations and comprehensive loss for the nine months ended September 30, 2013.

In connection with the Series E redeemable convertible preferred stock (Series E Preferred Stock) financing transactions that took place in June 2010 and July 2010, the Company issued warrants to purchase up to 871,580 shares of common stock. Each warrant was immediately exercisable and expires ten years from the original date of issuance. The warrants to purchase shares of the Company's common stock have an exercise price equal to the estimated fair value of the underlying instrument as of the initial date such warrants were issued. Each warrant is exercisable on either a physical settlement or net share settlement basis from the date of issuance. The warrant agreement contains a provision requiring an adjustment to the number of shares in the event the Company issues common stock, or securities convertible into or exercisable for common stock, at a price per share lower than the warrant exercise price. The Company concluded the anti-dilution feature required the warrants to be classified as liabilities under ASC Topic 815, *Derivatives and Hedging - Contracts in Entity's Own Equity* (ASC 815). The warrants are measured at fair value, with changes in fair value recognized as a gain or loss to other income (expense) in the statements of operations and comprehensive income (loss) for each reporting period thereafter. The fair value of the common stock warrants were recorded as a discount to the preferred stock issued of \$3.0 million, and the preferred stock was being accreted to the redemption value. At the end of each reporting period, the Company remeasured the fair value of the outstanding warrants, using current assumptions, resulting in an increase (decrease) in fair value of \$10.1 million, (\$0.1 million), \$11.3 million, and \$0.5 million, respectively, which was recorded in other expense in the accompanying statements of operations and comprehensive loss for the three months ended September 30, 2013 and 2012 and the nine months ended September 30, 2013 and 2012. The Company will continue to re-measure the fair value of the liability associated with the warrants to purchase common stock at the end of each reporting period until the earlier of the exercise or the expiration of the applicable warrants. On March 31, 2013, the Company retired 13,994 warrants to purchase common stock as a consequence of a repurchase of shares from an investor. All remaining outstanding warrants were fully vested and exercisable as of September 30, 2013 and December 31, 2012.

In connection with various financing transactions that were consummated in periods prior to December 31, 2012, the Company issued warrants to purchase up to 12,634 shares of common stock. The awards of warrants to purchase shares of common stock are accounted for as equity instruments.

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**Accelaron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**12. Warrants (continued)**

The warrants are exercisable at any time through their respective expiration dates. The fair value at issuance was calculated using the Black-Scholes option-pricing model, and was charged to interest expense during the periods the related debt was outstanding.

The Company issued warrants to purchase up to 41,388 shares of common stock in periods prior to December 31, 2012 in exchange for consulting services provided by a third party pursuant to stand-alone award agreements that are independent of an equity incentive plan. The warrants vested upon achievement of four milestones and were outstanding for approximately seven years from the date of issuance. There were no exercises, cancellations, or expirations of warrants during the year ended December 31, 2012.

*Fair Value*

The fair value of the warrants to purchase preferred stock on the date of issuance and on each re-measurement date for those warrants to purchase preferred stock classified as liabilities, was estimated using the Black-Scholes option pricing model. This method of valuation involves using inputs such as the fair value of the Company's various classes of preferred stock and common stock, stock price volatility, contractual term of the warrants, risk free interest rates, and dividend yields. The fair value of the warrants to purchase common stock on the date of issuance and on each re-measurement date for those warrants to purchase common stock are classified as liabilities and are estimated using the Monte Carlo simulation framework, which incorporated three future financing events over the remaining life of the warrants to purchase common stock. Due to the nature of these inputs and the valuation techniques utilized, the valuation of the warrants to purchase preferred stock and common stock are considered a Level 3 measurement (Note 7).

**13. Commitments and Contingencies**

*Legal Proceedings*

On October 18, 2012, the Salk Institute for Biological Studies (Salk) filed a complaint in the Massachusetts Superior Court for Suffolk County, alleging that the Company breached one of the Company's two licensing agreements with Salk. The licensing agreement in dispute provides the Company with a license with respect to certain of Salk's U.S. patents related to the ActRIIB activin receptor proteins. Salk contends that, under the licensing agreement, the Company owed Salk a greater share of the upfront payment that it received under its now-terminated agreement with Shire AG regarding ACE-031 and a share of the upfront payment and development milestone payments that the Company has received under its ongoing collaboration agreement with Celgene regarding ACE-536. Salk is seeking a total of approximately \$10.5 million plus interest in payment and a 15% share of future development milestone payments received under the agreement with Celgene regarding ACE-536. The Company contends that no additional amounts are due to Salk and that it has complied with all of its payment obligations under the applicable Salk license agreement.

The Company moved to dismiss the complaint on December 3, 2012. The Court denied the Company's motion on February 28, 2013. On March 14, 2013, Accelaron answered the complaint and asserted patent invalidity counterclaims. On the basis of those counterclaims, Accelaron removed the action on March 28, 2013 to the United States District Court for the District of Massachusetts. The parties have since reached an agreement on a stipulation as to certain patent issues raised in the action, and Accelaron has dismissed its counterclaims. The Court held an initial scheduling conference

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**13. Commitments and Contingencies (continued)**

on May 30, 2013, and the parties have begun fact discovery. The case is currently scheduled for trial in September 2014. The Company intends to defend its position vigorously.

The Company evaluated the suit under ASC Topic 450, *Contingencies*, as a loss contingency. The estimated loss from a loss contingency shall be accrued if information available before the financial statements are issued indicates that it is probable a liability had been incurred at the date of the financial statements, and the amount of loss can be reasonably estimated. Because the Company believes that the potential for an unfavorable outcome is not probable, it has not established a reserve with respect to the dispute as of September 30, 2013 or December 31, 2012.

The Company's estimates can be affected by various factors. As of December 31, 2012 and September 30, 2013, management has determined a loss is reasonably possible. Although the Company believes it would successfully defend the lawsuit, the Company has in the past participated in settlement discussions with Salk. Accordingly, the Company has estimated the range of possible losses as of September 30, 2013 and December 31, 2012 to be between \$0 and \$10.5 million plus interest.

***Other***

The Company is also party to various agreements, principally relating to licensed technology, that require future payments relating to milestones not met at September 30, 2013 and December 31, 2012, or royalties on future sales of specified products. No milestone or royalty payments under these agreements are expected to be payable in the immediate future. See Note 14 for discussion of these arrangements.

The Company enters into standard indemnification agreements in the ordinary course of business. Pursuant to the agreements, the Company indemnifies, holds harmless, and agrees to reimburse the indemnified party for losses suffered or incurred by the indemnified party, generally the Company's business partners or customers, in connection with any U.S. patent or any copyright or other intellectual property infringement claim by any third party with respect to the Company's products. The term of these indemnification agreements is generally perpetual any time after execution of the agreement. The maximum potential amount of future payments the Company could be required to make under these indemnification agreements is unlimited. The Company has never incurred costs to defend lawsuits or settle claims related to these indemnification agreements.

**14. Significant Agreements**

**Celgene**

***Overview***

On February 20, 2008 the Company entered into a collaboration, license, and option agreement (the Sotatercept Agreement) with Celgene Corporation (Celgene) relating to sotatercept. On August 2, 2011, the Company entered into a second collaboration, license and option agreement with Celgene for ACE-536 (the ACE-536 Agreement), and also amended certain terms of the Sotatercept Agreement. These agreements provide Celgene exclusive licenses for Sotatercept and ACE-536 in all indications, as well as exclusive rights to obtain a license to certain future compounds. Celgene is a global biopharmaceutical company primarily engaged in the discovery, development and commercialization of innovative therapies designed to treat cancer and immune-inflammatory related diseases.

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**14. Significant Agreements (continued)**

*Sotatercept Agreement*

Under the terms of the Sotatercept Agreement, the Company and Celgene collaborate worldwide for the joint development and commercialization of sotatercept. The Company also granted Celgene an option to license three discovery stage compounds. Under the terms of the agreement, the Company and Celgene will jointly develop, manufacture and commercialize sotatercept. Celgene paid \$45.0 million of nonrefundable, upfront license and option payments to the Company upon the closing of the Sotatercept Agreement.

The Company retained responsibility for research, development through the end of Phase 2a clinical trials, as well as manufacturing the clinical supplies for these trials. These activities were substantially completed in 2011. Celgene is conducting the ongoing Phase 2 trials for myelodysplastic syndromes (MDS), chronic kidney disease, and  $\beta$ -thalassemia and will be responsible for any Phase 3 clinical trials, as well as additional Phase 2 clinical trials, and will be responsible for overseeing the manufacture of Phase 3 and commercial supplies by third party contract manufacturing organizations. Under the agreement, the Company was eligible to receive clinical milestones of up to \$88.0 million, regulatory milestones of up to \$272.0 million, and commercial milestones of up to \$150.0 million for sotatercept. Clinical milestone payments are triggered upon initiation of a defined phase of clinical research for a product candidate. Regulatory milestone payments are triggered upon the acceptance of the marketing application and upon the approval to market a product candidate by the Food and Drug Administration (FDA) or other global regulatory authorities. Commercial milestone payments are triggered when an approved pharmaceutical product reaches certain defined levels of net sales by Celgene in countries outside of North America. In addition, to the extent sotatercept is commercialized, the Company would be entitled to receive tiered royalty payments in the low-to-mid twenty percent range of net sales from sales generated from all geographies. Royalty payments are subject to certain reductions, including for entry of a generic product onto the market.

Additionally, for three named discovery-stage option programs the Company was eligible to receive option fees of up to \$30.0 million, clinical milestones of up to \$53.3 million, regulatory milestones of up to \$204.0 million, and commercial milestones of up to \$150.0 million for each option program. Clinical milestone payments are triggered upon initiation of a defined phase of clinical research for a product candidate. Regulatory milestone payments are triggered upon the acceptance of the marketing application and upon the approval to market a product candidate by the FDA or other global regulatory authorities. Commercial milestone payments are triggered when an approved pharmaceutical product reaches certain defined levels of net sales by Celgene in countries outside of North America. Option fee payments are triggered upon license of any of the option programs by Celgene. In addition, to the extent an option compound is commercialized, the Company would be entitled to receive tiered royalty payments in the low-to-mid twenty percent range of net sales from sales generated from all geographies. Royalty payments are subject to certain reductions, including for entry of a generic product onto the market. None of the three discovery stage programs has advanced to the stage to achieve payment of a milestone.

In connection with entering into the Sotatercept Agreement, Celgene purchased 457,875 shares of Series C-1 Preferred Stock at the aggregate purchase price of \$5.0 million. The Series C-1 Preferred Stock was purchased at an amount that was deemed to represent fair value at the time of purchase. Concurrent with the IPO, Celgene purchased 666,667 shares of Common Stock at the IPO offer price of \$15.00 per share.

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**14. Significant Agreements (continued)**

Commensurate with the execution of the ACE-536 Agreement described below, the Company and Celgene agreed to modify the terms of the Sotatercept Agreement. The modified terms included: (1) a change to the responsibility for development costs to align with the ACE-536 Agreement, with Celgene responsible for more than half of the worldwide costs through December 31, 2012, and 100% of the development costs thereafter, (2) future contingent development milestones for sotatercept were amended to a two-category (oncology and non-oncology) structure with potential future clinical milestones of \$27.0 million and regulatory milestones of \$190.0 million from a four-category (various cancer indications) structure and, (3) future contingent development milestones for option compounds were amended to a two-category (oncology and non-oncology) structure with potential future clinical milestones of \$25.5 million and regulatory milestones of \$142.5 million from a four-category (various cancer indications) structure, and (4) an option to buy down tiered royalty payments on both Sotatercept and ACE-536 with a one-time \$25.0 million payment on or prior to January 1, 2013. The potential commercial milestones remained unchanged. Through September 30, 2013, the Company has received \$34.5 million in research and development funding and milestone payments for sotatercept under the original and modified agreements. The next likely clinical milestone payment would be \$7.0 million and result from Celgene's start of a Phase 2b clinical trial in chronic kidney disease.

The Sotatercept Agreement will expire on a country-by-country basis on the occurrence of both of the following: (1) the expiration of the royalty term with respect to all license products in such country, and (2) the exercise or forfeiture by Celgene of its option with regard to each option compound. The royalty term for each licensed product in each country outside North America is the period commencing with first commercial sale of the applicable licensed product in the applicable country and ending on the latest of expiration of specified patent coverage or a specified period of years. The royalty term for each licensed product in North America is the period commencing with the first commercial sale in North America and ending, on a licensed product and country-by-country basis on the date which commercialization of such licensed product has ceased. The term for each option compound runs for a specified period of years unless Celgene exercises its option, in which case the compound becomes a licensed product, or forfeits its option by failing to make certain payments following the achievement of certain milestones in early clinical development of the option compound.

Celgene has the right to terminate the agreement with respect to one or more licensed targets or in its entirety, upon 180 days' notice (or 45 days' notice if the licensed product has failed to meet certain end point criteria with respect to clinical trials or other development activities). The agreement may also be terminated in its entirety by either Celgene or the Company in the event of a material breach by the other party or in the event of a bankruptcy filing of the other party. There are no cancellation, termination or refund provisions in this arrangement that contain material financial consequences to the Company.

***ACE-536 Agreement***

Under the terms of the ACE-536 Agreement, the Company and Celgene collaborate worldwide for the joint development and commercialization of ACE-536. The Company also granted Celgene an option for future products for which Acceleron files an Investigational New Drug application for the treatment of anemia. Celgene paid \$25.0 million on the closing of the ACE-536 Agreement in August, 2011.

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**14. Significant Agreements (continued)**

The Company retains responsibility for research, development through the end of Phase 1 and initial Phase 2 clinical trials, as well as manufacturing the clinical supplies for these studies. Celgene will conduct subsequent Phase 2 and Phase 3 clinical studies. Acceleron will manufacture ACE-536 for the Phase 1 and Phase 2 clinical trials and Celgene will be responsible for overseeing the manufacture of Phase 3 and commercial supplies by third party contract manufacturing organizations. The Company is eligible to receive clinical milestones of up to \$32.5 million, regulatory milestones of up to \$105.0 million and commercial milestones of up to \$80.0 million for ACE-536. The Company will receive additional, lower development, regulatory, and commercial milestones for any additional products for the treatment of anemia on which Celgene exercises an option. Clinical milestone payments are triggered upon initiation of a defined phase of clinical research for a product candidate. Regulatory milestone payments are triggered upon the acceptance of the marketing application and upon approval to market a protein therapeutic candidate by the FDA or other global regulatory authorities. Commercial milestone payments are triggered when an approved pharmaceutical product reaches certain defined levels of net sales by Celgene in countries outside of North America. In addition, to the extent ACE-536 is commercialized, the Company would be entitled to receive tiered royalty payments in the low-to-mid twenty percent range of net sales from sales generated from all geographies. Royalty payments are subject to certain reductions, including for entry of a generic product onto the market.

Through September 30, 2013, the Company has received \$28.3 million in research and development funding and milestone payments for ACE-536. The next likely clinical milestone payment would be \$15.0 million and result from the start of a Phase 3 study in MDS or  $\beta$ -thalassemia. The Company has not yet identified additional compounds for the treatment of anemia. Accordingly, there is no assurance that the Company will generate future value from additional programs.

The ACE-536 Agreement will expire on a country-by-country basis on the occurrence of both of the following: (1) the expiration of the royalty term with respect to all license products in such country, and (2) the end of the option term. The royalty term for each licensed product in each country outside North America is the period commencing with first commercial sale of the applicable licensed product in the applicable country and ending on the latest of expiration of specified patent coverage or a specified period of years. The royalty term for each licensed product in North America is the period commencing with the first commercial sale in North America and ending, on a licensed product and country-by-country basis on the date which commercialization of such licensed product has ceased. The option term runs until the later of (1) the date on which no development or commercialization activities are ongoing or are expected to commence for any licensed products under the ACE-536 Agreement; (2) the date on which no development or commercialization activities are ongoing or are expected to commence for any licensed products under the Sotatercept Agreement and all option rights under the Sotatercept Agreement have been forfeited with respect to each option compound where Celgene has made a payment with respect to such compound; and (3) the royalty term for all licensed products under the ACE-536 Agreement and the Sotatercept Agreement has ended; provided that if at the time the option term would otherwise end any option compounds under the ACE-536 Agreement are in clinical development the option term shall continue until Celgene's rights to such compound are either exercised or forfeited.

Celgene has the right to terminate the ACE-536 Agreement with respect to one or more licensed targets or in its entirety, upon 180 days' notice (or 45 days' notice if the licensed product has failed to meet certain end point criteria with respect to clinical trials or other development activities), provided that Celgene may not terminate the ACE-536 Agreement prior to the completion of the on-going



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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**14. Significant Agreements (continued)**

ACE-536  $\beta$ -thalassemia and ACE-536 MDS Phase 2 clinical trials, except under certain conditions. The agreement may also be terminated in its entirety by either Celgene or the Company in the event of a material breach by the other party or in the event of a bankruptcy filing of the other party. There are no cancellation, termination or refund provisions in this arrangement that contain material financial consequences to the Company.

***Both Agreements***

The Company and Celgene shared development costs under the Sotatercept and ACE-536 Agreements through December 31, 2012. As of January 1, 2013, Celgene is responsible for paying 100% of worldwide development costs under both agreements. Celgene will be responsible for all commercialization costs worldwide. The Company has the right to co-promote sotatercept, ACE-536 and future products under both agreements in North America. Celgene's option to buy down royalty rates for sotatercept and ACE-536 expired unexercised and, therefore, the Company will receive tiered royalties in the low-to-mid twenty percent range on net sales of sotatercept and ACE-536. The royalty schedules for sotatercept and ACE-536 are the same.

***Accounting Analysis***

Prior to 2011, the Company accounted for the Sotatercept Agreement, as a multiple element arrangement under ASC 605-25 (prior to the amendments of ASU 2009-13). The Company identified the following deliverables under the arrangement; (1) the license to the ActRIIA compound, (2) right to license option program compounds, (3) participation in the joint development committee, (4) participation in the joint commercialization committee and (5) research and development activities. Under the provisions of ASC 605- 25, applicable to the arrangement, since the Company could not establish VSOE for the undelivered elements, the Company was required to recognize the initial consideration, consisting of the \$45.0 million of nonrefundable upfront license and option payments, over the period the undelivered elements were to be delivered, which was initially estimated to be 15 years. As of the date of the modification of the agreement, there was approximately \$34.7 million of deferred revenue under the arrangement.

As a result of the material modifications to the cost sharing obligations, milestone payments structure and royalty payment structure, the Company concluded the modification represented a significant modification under ASU 2009-13, which required the Company to apply the updated provisions of ASU 2009-13 subsequent to the modification.

Because the ACE-536 Agreement and the amendment to the Sotatercept Agreement were negotiated in contemplation of each other, and the Company had not yet completed all of its obligations pursuant to the Sotatercept Agreement, the agreements were considered one arrangement for accounting purposes. The deliverables under the combined arrangement include: (1) licenses to develop and commercialize sotatercept and ACE-536, (2) performance of research and development services, (3) participation on the joint development committees, and (4) the performance of manufacturing services to provide clinical material to Celgene. The Company has determined the option to future products related to the treatment of anemia represents a substantive option. The Company is under no obligation to discover, develop or deliver any new compounds that modulate anemia and Celgene is not contractually obligated to exercise the option. As a result, the Company is at risk as to whether Celgene will exercise the option.

Table of Contents**Acceleron Pharma Inc.****Notes to Unaudited Interim Condensed Financial Statements (continued)****14. Significant Agreements (continued)**

All of these deliverables identified in the arrangement were deemed to have stand-alone value and to meet the criteria to be accounted for as separate units of accounting under ASC 605-25. Factors considered in making this determination included, among other things, the subject of the licenses, the nature of the research and development services, and the capabilities of Celgene.

The total arrangement consideration of \$77.7 million under the ACE-536 Agreement and amended Sotatercept Agreement consisted of (1) the \$25.0 million up-front payment for the license of ACE-536, (2) the remaining deferred revenue from the Sotatercept Agreement of \$34.7 million, and (3) estimated payments for development activities and manufacturing services of \$18.0 million. The Company used its BEBP for each of the undelivered elements as the Company did not have VSOE or TPE of selling price for each deliverable. The Company's BEBP considered its development plan for the compounds, expected manufacturing services, and reimbursement from Celgene (reimbursement of more than half of development expenses through December 31, 2012 and 100% thereafter). The Company determined its BEBP for each of the undelivered elements under the arrangements as of the arrangement execution date as follows:

\$18.8 million for research and development services

\$2.9 million for the sotatercept joint development committee

\$3.7 million for the ACE 536 joint development committee

\$2.8 million for the manufacturing services

After determining BEBP of the undelivered elements, the remaining consideration of \$49.5 million was recognized upon execution of the arrangements. The difference between the estimated payments of \$18.0 million and the estimated selling prices which totaled \$28.2 million, using BEBP, for undelivered elements was \$10.2 million. This amount was deferred at inception and will be recognized as the undelivered elements are delivered, using the proportional performance method, or ratably in the case of performance on the Joint Development Committee.

As noted above, the total arrangement consideration includes estimated payments for development activities and manufacturing services identified at the outset of the ACE-536 Agreement and amended Sotatercept Agreement. At the end of each reporting period, the Company reassesses the estimated payments to be received related to these services and the BEBP of the undelivered elements based upon the Company's current estimates. The Company accounts for such changes as a change in accounting estimate and the cumulative impact of any change is reflected in the period of change.

During 2011, the Company achieved a \$7.5 million clinical milestone under its ACE-536 Agreement, related to the dosing of the first patient in a multiple-dose clinical trial. The Company evaluated the milestone and determined that it was not substantive, as there was no significant uncertainty to achieving the milestone upon execution of the ACE-536 Agreement. As such, the Company allocated the \$7.5 million payment based on the allocation of arrangement consideration determined at the execution date of the ACE-536 Agreement and amended Sotatercept Agreement. Based on this allocation, the Company recognized \$4.8 million of the payment upon achievement, with the remaining \$2.7 million recognized as revenue as the undelivered elements are delivered, consistent with the treatment of the up-front payment. During January 2013, the Company achieved a \$10.0 million clinical milestone under its ACE-536 Agreement, related to the dosing of the first patient for a Phase 2 clinical trial. The Company evaluated the milestone and deemed it to be substantive and

Table of Contents**Acceleron Pharma Inc.****Notes to Unaudited Interim Condensed Financial Statements (continued)****14. Significant Agreements (continued)**

consistent with the definition of a milestone included in ASU 2010-17 and, accordingly, recognized the \$10.0 million payment in revenue during the nine months ended September 30, 2013. The remaining development milestones under the ACE-536 and Sotatercept Agreements were deemed to be substantive and consistent with the definition of a milestone included in ASU 2010-17 and, accordingly, the Company will recognize payments related to the achievement of such milestones, if any, when such milestone is achieved. Factors considered in this determination included scientific and regulatory risks that must be overcome to achieve the milestones, the level of effort and investment required to achieve each milestone, and the monetary value attributed to each milestone. During the three months ended September 30, 2013 and 2012, and the nine months ended September 30, 2013 and 2012, the Company recognized \$0.6 million, \$0.5 million, \$1.7 million and \$1.5 million, respectively, of the total deferred revenue as license and milestone revenue in the accompanying statements of operations and comprehensive loss.

Pursuant to the terms of the agreement, Celgene and the Company share development costs, with Celgene responsible for substantially more than half of the costs for sotatercept and ACE-536 until December 31, 2012 and 100% of the costs from January 1, 2013 and thereafter. Payments from Celgene with respect to research and development costs incurred by the Company are recorded as cost-sharing revenue. Payments by the Company to Celgene for research and development costs incurred by Celgene are recorded as a reduction to cost-sharing revenue. During the three months ended September 30, 2013 and 2012, and the nine months ended September 30, 2013 and 2012 the Company recorded net cost-sharing revenue of \$3.6 million, \$0.8 million, \$9.0 million and \$2.1 million, respectively, which includes payments to Celgene of, zero, \$0.6 million, zero and \$1.9 million, respectively, which were recorded as contra-revenue.

**Other Agreements***Shire License*

In September 2010, the Company entered into a license and collaboration agreement granting Shire AG the exclusive right to develop, manufacture and commercialize ActRIIB compounds in territories outside North America. Shire also received the right to conduct research and manufacture commercial supplies in North America for ActRIIB compounds. The lead ActRIIB compound was designated ACE-031. Under the initial development plan, the companies share the costs associated with developing and commercializing ACE-031, in Duchenne Muscular Dystrophy. In September 2010, Shire made a nonrefundable, up-front license payment to the Company of \$45.0 million. In accordance with the Company's revenue recognition policy prior to the adoption of ASU 2009-13, the up-front license payment of \$45.0 million was deferred, and will be recognized as revenue ratably over three years, which represented the original period over which the Company expected to perform and deliver research and development and manufacturing services. On February 8, 2011, the FDA placed ACE-031 on clinical hold. The Company re-assessed the duration of its deliverables under the license agreement and estimated the new term to be approximately five years. The adjustment was treated as a change in accounting estimate with the remaining deferred revenue of \$38.8 million at February 8, 2011, recognized prospectively over the new period of research and development and manufacturing services. In April 2013, the Company and Shire determined not to further pursue development of ACE-031 and Shire sent the Company a notice of termination for the ACE-031 collaboration. The collaboration terminated effective June 30, 2013. At December 31, 2012, the Company had classified the remaining deferred revenue as current in the balance sheet. Upon the effectiveness of the termination of the

Table of Contents**Acceleron Pharma Inc.****Notes to Unaudited Interim Condensed Financial Statements (continued)****14. Significant Agreements (continued)**

Shire Agreement in the second quarter of 2013, the Company accelerated the recognition of \$22.4 million of remaining deferred revenue from upfront non-refundable payments received under the Shire Agreement as it had no further obligation for deliverables under the Shire Agreement. During the three months ended September 30, 2013 and 2012, and the nine months ended September 30, 2013 and 2012, the Company recognized zero, \$1.9 million, \$24.3 million and \$5.7 million, respectively of the up-front, non-refundable payments as license and milestone revenue in the accompanying statements of operations and comprehensive loss.

The agreement also included contingent milestone payments, based on the achievement of development milestones totaling \$223.8 million and commercial milestones of \$228.8 million for ActRIIB compounds. The milestones under the Shire Agreement were deemed to be substantive and consistent with the definition of a milestone included in ASU 2010-17 and, accordingly, the Company recognized payments related to the achievement of such milestones, if any, when such milestone was achieved. Factors considered in this determination included scientific and regulatory risks that must be overcome to achieve the milestones, the level of effort and investment required to achieve each milestone, and the monetary value attributed to each milestone.

Pursuant to the terms of the agreement, Shire and the Company shared development costs, with Shire responsible for 65% of the costs for ACE-031 and 55% of the costs for licensed compounds other than ACE-031. Payments from Shire with respect to research and development costs incurred by the Company are recorded as cost-sharing revenue. Payments by the Company to Shire for research and development costs incurred by Shire are recorded as a reduction to cost-sharing revenue. During the three months ended September 30, 2013 and 2012, and the nine months ended September 30, 2013 and 2012, the Company recorded net cost-sharing revenue of zero, \$0.6 million, \$0.6 million, and \$1.9 million, respectively, which includes payments to Shire of zero, \$0.2 million, \$0.2 million, and \$0.6 million, respectively, which are recorded as contra-revenue in the accompanying statements of operations and comprehensive loss.

***Other***

The Company entered into a license agreement with a non-profit institution for an exclusive, sublicensable, worldwide, royalty-bearing license to certain patents developed by the institution (Primary Licensed Products). In addition, the Company was granted a non-exclusive, non-sub-licensable license for Secondary Licensed Products. As compensation for the licenses, the Company issued 62,500 shares of its common stock to the institution, the fair value of which was \$25,000, and was expensed during 2004, to research and development expense. The Company also agreed to pay specified development milestone payments totaling up to \$2.0 million for sotatercept and \$0.7 million for ACE-536. In addition, the Company is obligated to pay milestone fees based on the Company's research and development progress, and U.S. sublicensing revenue ranging from 10%-25%, as well as a royalty ranging from 1.0%-3.5% of net sales on any products developed under the licenses. During the three months ended September 30, 2013 and 2012, and the nine months ended September 30, 2013 and 2012, the Company paid and expensed milestones and fees defined under the agreement totaling \$50,000, zero, \$50,000, and zero respectively.

The Company entered into another license agreement with certain individuals for an exclusive, sublicensable, worldwide, royalty-bearing license to certain patents developed by the individuals. The Company agreed to pay specified development and sales milestone payments aggregating up to

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**14. Significant Agreements (continued)**

\$1.0 million relating to the development and commercialization of dalantercept. In addition, the Company is required to pay royalties in the low single-digits on worldwide net product sales of dalantercept, with royalty obligations continuing at a 50% reduced rate for a period of time after patent expiration. If the Company sublicenses its patent rights, the Company will owe a percentage of sublicensing revenue, excluding payments based on the level of sales, profits or other levels of commercialization. During the nine months ended September 30, 2013 and 2012, the Company did not reach any milestones defined under the agreement and, therefore, no amounts have been paid or expensed.

During 2012, the Company executed a license agreement with a research institution for an exclusive, sublicensable, worldwide, royalty-bearing license. The Company is obligated to pay development milestones and commercial milestone fees totaling up to \$1.0 million. Under the agreement, if the Company uses the inventors in the clinical research, the development milestones are waived and commercial milestones shall change to \$0.8 million plus any waived milestones. The Company will also pay \$25,000 annually upon first commercial sale as well as royalties of 1.5% of net sales on any products developed under the patents. During the nine months ended September 30, 2013 and 2012, the Company did not reach any milestones defined under the agreement and, therefore, no amounts have been paid or expensed.

**15. Stock-Based Compensation**

At September 30, 2013, the Company had two stock-based compensations plans, which are more fully described below.

The Company's 2003 Stock Option and Restricted Stock Plan (the 2003 Plan) provides for the issuance of stock options, restricted stock awards, and restricted stock to employees, officers, directors, consultants, and key personnel of the Company as determined by the Board. In conjunction with the effectiveness of the 2013 Equity Incentive Plan (the 2013 Plan) described below, the Company determined that no further stock options or other equity-based awards may be granted under the 2003 Plan.

On September 4, 2013, the Company adopted the 2013 Plan. The Company has reserved for issuance an aggregate of 1,500,000 shares of common stock under the 2013 Plan, which is comprised of (i) the remaining 155,884 shares reserved for issuance under the 2003 Plan and (ii) an additional 1,344,116 shares. The 2013 Plan provides that the number of shares reserved and available for issuance under the plan will automatically increase each January 1, beginning in 2014, by the lesser of (i) 3,150,000 shares, or (ii) 4% of the outstanding number of shares of the Company's common stock on the immediately preceding December 31st. This number is subject to adjustment in the event of a stock split, stock dividend or other change in the Company's capitalization. No grants were made under the 2013 Plan as of September 30, 2013, and 1,500,000 shares were available for issuance under the 2013 Plan as of September 30, 2013.

The Company has not granted unrestricted stock awards under the 2003 Plan and the 2013 Plan since its inception. Stock options carry an exercise price equal to the estimated fair value of the Company's common stock on the date of grant. Options generally expire ten years following the date of grant. Stock options and restricted stock awards typically vest over four years, but vesting provisions can vary based on the discretion of the Board.

Table of Contents**Accelaron Pharma Inc.****Notes to Unaudited Interim Condensed Financial Statements (continued)****15. Stock-Based Compensation (continued)**

Shares of the Company's common stock underlying any awards that are forfeited, canceled, withheld upon exercise of an option, or settlement of an award to cover the exercise price or tax withholding, reacquired by the Company prior to vesting, satisfied without the issuance of shares of the Company's common stock, or otherwise terminated other than by exercise will be added back to the shares of common stock available for issuance under the 2013 Plan. Shares available for issuance under the 2013 Plan may be authorized but unissued shares of the Company's common stock or shares of the Company's common stock that have been reacquired by the Company.

Additionally, on September 4, 2013, the company adopted the 2013 Employee Stock Purchase Plan (the 2013 ESPP). Under the 2013 ESPP, 275,000 shares of the Company's common stock will be available for issuance and eligible employees of the Company may purchase shares of common stock during pre-specified purchase periods at a price equal to the lesser of 85% of the fair market value of a share of its common stock at the beginning of the purchase period or 85% of the fair market value of a share of its common stock at the end of the purchase period. As of September 30, 2013, the initial purchase period under the 2013 ESPP has not yet commenced.

The Company recognized stock-based compensation expense totaling \$0.5 million, \$0.3 million, \$1.4 million and \$0.9 million during the three months ended September 30, 2013 and 2012 and the nine months ended September 30, 2013 and 2012, respectively.

Total compensation cost recognized for all stock-based compensation awards in the statements of operations and comprehensive income (loss) is as follows (in thousands):

	<b>Three Months Ended September 30,</b>		<b>Nine Months Ended September 30,</b>	
	<b>2013</b>	<b>2012</b>	<b>2013</b>	<b>2012</b>
Research and development	\$ 149	\$ 137	\$ 460	\$ 374
General and administrative	344	196	981	487
	\$ 493	\$ 332	\$ 1,441	\$ 861

The fair value of each option issued to employees was estimated at the date of grant using the Black-Scholes option pricing model with the following weighted-average assumptions (in thousands):

	<b>Three Months Ended September 30,</b>		<b>Nine Months Ended September 30,</b>	
	<b>2013</b>	<b>2012</b>	<b>2013</b>	<b>2012</b>
Expected volatility	% 66.9%	70.3%	66.9%	
Expected term (in years)		6.0	6.0	6.0
Risk-free interest rate	% 0.9%	1.4%	0.9%	
Expected dividend yield	%	%	%	%

***Fair Value of Underlying Instrument***

The Company estimates the fair value of its stock-based awards to employees using the Black-Scholes option pricing model.

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**15. Stock-Based Compensation (continued)**

*Expected Volatility*

The Company estimated the expected volatility based on actual historical volatility of the stock price of similar companies with publicly-traded equity securities. The Company calculated the historical volatility of the selected companies by using daily closing prices over a period of the expected term of the associated award. The companies were selected based on their enterprise value, risk profiles, position within the industry, and with historical share price information sufficient to meet the expected term of the associated award. A decrease in the selected volatility would decrease the fair value of the underlying instrument.

*Expected Term*

The Company estimates the expected life of its employee stock options using the "simplified" method, as prescribed in Staff Accounting Bulletin (SAB) No. 107, whereby, the expected life equals the arithmetic average of the vesting term and the original contractual term of the option due to its lack of sufficient historical data.

*Risk-Free Interest Rate*

The Company estimated the risk-free interest rate in reference to the yield on U.S. Treasury securities with a maturity date commensurate with the expected term of the associated award. A decrease in the selected risk-free rate would decrease the fair value of the underlying instrument.

*Expected Dividend Yield*

The Company estimated the expected dividend yield based on consideration of its historical dividend experience and future dividend expectations. The Company has not historically declared or paid dividends to stockholders. Moreover, it does not intend to pay dividends in the future, but instead expects to retain any earnings to invest in the continued growth of the business. Accordingly, the Company assumed an expected dividend yield of 0.0%.

Table of Contents**Accelaron Pharma Inc.****Notes to Unaudited Interim Condensed Financial Statements (continued)****15. Stock-Based Compensation (continued)****Stock Options**

The following table summarizes the stock option activity for all stock option plans during the nine months ended September 30, 2013 (in thousands):

	Number of Grants	Weighted- Average Exercise Price Per Share	Weighted- Average Contractual Life (in years)	Aggregate Intrinsic Value(1)
Outstanding at December 31, 2012	3,730	\$ 4.16	6.62	
Granted	9	\$ 9.64		
Exercised	(38)	\$ 1.34		
Canceled or forfeited	(45)	\$ 4.31		
Outstanding at September 30, 2013	3,656	\$ 4.18	6.00	\$ 65,987
Exercisable at September 30, 2013	2,665	\$ 3.78	5.12	\$ 49,173
Vested and expected to vest at September 30, 2013(2)	3,604	\$ 4.16	5.96	\$ 65,113

(1) The aggregate intrinsic value is calculated as the difference between the exercise price of the underlying options and the estimated fair value of the common stock for the options that were in the money at September 30, 2013.

(2) This represents the number of vested options at September 30, 2013, plus the number of unvested options expected to vest at September 30, 2013, based on the unvested options outstanding at September 30, 2013, adjusted for the estimated forfeiture rate.

During the nine months ended September 30, 2013, the Company granted stock options to purchase an aggregate of 8,750 shares of its common stock, with a weighted-average grant date fair value of options granted of \$9.64.

During the nine months ended September 30, 2013, current and former employees of the Company exercised a total of 37,532 options, resulting in total proceeds of \$50,000.

The aggregate intrinsic value of options exercised during the nine months ended September 30, 2013 was \$306,000.

As of September 30, 2013, there was \$3.3 million of unrecognized compensation expense related to unvested stock options that is expected to be recognized over a weighted-average period of 2.2 years.

**16. Income Taxes**

The Company provides for income taxes under ASC Topic 740, Accounting for Income Taxes. Under ASC Topic 740, the liability method is used in accounting for income taxes. Under this method, deferred tax assets and liabilities are determined based on differences between financial reporting and tax bases of assets and liabilities, and are measured using the enacted tax rates and laws that will be in effect when the differences are expected to reverse.



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For the three and nine months end September 30, 2013 and 2012, the Company did not record a current or deferred income tax expense or benefit.

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**16. Income Taxes (continued)**

The Company has evaluated the positive and negative evidence bearing upon the realizability of its deferred tax assets. Based on the Company's history of operating losses, the Company has concluded that it is more likely than not that the benefit of its deferred tax assets will not be realized. Accordingly, the Company has provided a full valuation allowance for deferred tax assets as of September 30, 2013 and December 31, 2012.

The Company files income tax returns in the United States, and various state and foreign jurisdictions. The federal, state and foreign income tax returns are generally subject to tax examinations for the tax years ended December 31, 2009 through December 31, 2012. To the extent the Company has tax attribute carryforwards, the tax years in which the attribute was generated may still be adjusted upon examination by the Internal Revenue Service, state or foreign tax authorities to the extent utilized in a future period.

**17. Long-Term Debt**

On June 7, 2012, the Company entered into a loan and security agreement (the Loan Agreement) with three lenders, pursuant to which the Company received a loan in the aggregate principal amount of \$20.0 million. The Company is required to repay the aggregate principal balance under the Loan Agreement in 42 months. The first 12 payments are interest only and the remaining 30 payments are equal monthly installments of principal plus interest. The Loan Agreement provided that the interest only period could be extended under certain circumstances. The Company did not trigger the requirements and began paying principal in July 2013.

Per annum interest is payable at the 8.5%. The Loan Agreement also included a closing fee of \$0.2 million. The Company is amortizing the cost over the 42 months of loan. The Loan Agreement is also subject to an additional deferred payment of \$1.2 million due with the final payment. The Company is recording the deferred payment to interest expense over the term of the Loan Agreement. The resulting effective interest rate is approximately 11.8%. The company is not subject to any financial covenants and the Loan Agreement is secured by a lien on all of the Company's personal property as of, or acquired after, the date of the Loan Agreement, except for intellectual property.

The Loan Agreement defines events of default, including the occurrence of an event that results in a material adverse effect upon the Company's business operations, properties, assets or condition (financial or otherwise), its ability to perform its obligations under and in accordance with the terms of the Loan Agreement, or upon the ability of the lenders to enforce any of their rights or remedies with respect to such obligations, or upon the collateral under the Loan Agreement or upon the liens of the lenders on such collateral or upon the priority of such liens. As of September 30, 2013 and December 31, 2012, there have been no events of default under the loan. As of September 30, 2013 and December 31, 2012, the principal balance outstanding was \$18.2 million and \$20.0 million, respectively.

Table of Contents**Acceleron Pharma Inc.****Notes to Unaudited Interim Condensed Financial Statements (continued)****17. Long-Term Debt (continued)**

The roll-forward of the notes payable balance during the nine months ending September 30, 2013, was as follows (in thousands):

Total notes payable (current and long -term portions) balance as of December 31, 2012	\$ 20,193
Interest accrued	257
Repayment of long-term debt	(1,815)
	18,635
Less current portion	(7,656)
Noncurrent financing obligations as of September 30, 2013	\$ 10,979

**18. Related Party Transactions****Celgene Corporation (Celgene)**

In connection with its entry into the collaboration agreement with Celgene, on February 2008, the Company sold Celgene 457,875 shares of its Series C-1 Preferred Stock. As part of the Company's June 2010 Series E financing, Celgene purchased 36,496 shares of Series E Preferred Stock and received warrants to purchase 38,979 shares of common stock. As part of the Company's December 2011 Series F financing, Celgene purchased 1,990,446 shares of Series F Preferred Stock. In connection with the Company's September 2013 initial public offering, Celgene purchased 666,667 shares of common stock. As a result of these transactions, Celgene owned 9.8% and 9.9% of the Company's fully diluted equity as of September 30, 2013 and December 31, 2012, respectively. Refer to Note 14 for additional information regarding this collaboration agreement.

During the nine months ended September 30, 2013, the Company recognized \$20.8 million in collaboration revenue under the Celgene collaboration arrangement and, as of September 30, 2013, had \$8.6 million of deferred revenue related to the Celgene collaboration arrangement.

The Company recognized revenue from Celgene during the three and nine months ended September 30, 2013 and 2012 as follows (in thousands):

	Three Months Ended September 30,		Nine Months Ended September 30,	
	2013	2012	2013	2012
License and milestone	\$ 638	\$ 535	\$ 11,722	\$ 1,491
Cost sharing, net	3,632	846	9,041	2,106
	\$ 4,270	\$ 1,381	\$ 20,763	\$ 3,597

**Alkermes**

One of the Company's directors is also the Chairman, President, and Chief Executive Officer of Alkermes plc, the parent company of Alkermes, Inc. (Alkermes), with which the Company entered into a collaboration agreement during 2009.

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**Acceleron Pharma Inc.**

**Notes to Unaudited Interim Condensed Financial Statements (continued)**

**18. Related Party Transactions (continued)**

As of December 31, 2012, Alkermes held 695,250 shares of the Company's Preferred Stock and warrants to purchase 42,624 shares of common stock. Upon the closing of the IPO on September 24, 2013, all of the shares of the Company's preferred stock held by Alkermes were converted into 718,655 shares of common stock. No research fees were paid to the Company during 2012 or 2013.

**Related-Party Receivable**

On January 28, 2008, the Company issued a secured promissory note (the Note Receivable) in the amount of \$0.2 million to the current chief executive officer of the Company (the CEO). The Note Receivable bears interest at an annual interest rate of 3.11% and was initially repayable on the earlier of January 28, 2011, or the date prior to the date that the Company files a registration statement with the SEC, covering shares of its common stock. The Note Receivable was secured by shares of the Company's common stock owned by the CEO. On December 22, 2010, the term was extended until January 28, 2014, or the date prior to the date that the Company files a registration statement with the SEC covering shares of its common stock.

In November 2012, the Company further modified the terms of the Note Receivable, such that in the event that an acquisition event occurs or the company files a registration statement with the SEC on or before the maturity date, the unpaid principal and interest will be forgiven. The Company evaluated the forgiveness provisions and determined that forgiveness was not probable as of December 31, 2012, and as such, continued to record the Note Receivable as an asset at December 31, 2012. As a result of the Company's filing of a registration statement with the SEC on August 6, 2013 which triggered the forgiveness of the Note Receivable, the Company expensed the unpaid principal and interest expense totaling \$0.2 million as compensation expense during the nine months ended September 30, 2013.

**19. Supplementary Financial Data**

The following table presents certain unaudited quarterly financial information for the eleven quarters in the period ended September 30, 2013. This information has been prepared on the same basis as the audited financial statements and includes all adjustments (consisting only of normal recurring adjustments) necessary to present fairly the unaudited quarterly results of operations set forth

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## Acceleron Pharma Inc.

## Notes to Unaudited Interim Condensed Financial Statements (continued)

## 19. Supplementary Financial Data (continued)

herein. Net income (loss) per share for all periods presented have been retroactively adjusted to reflect the 1-for-4 reverse stock split effected on September 5, 2013.

	For the Three Months Ended(1)			
	March 31	June 30	September 30	December 31
	(in thousands except per share data)			
(unaudited)				
<b>2013:</b>				
Total revenue	\$ 15,012	\$ 26,427	\$ 4,270	
Total costs and expenses	(11,876)	(12,276)	(11,154)	
Income (loss) from operations	3,136	14,151	(6,884)	
Net income (loss)	1,647	13,078	(18,513)	
Basic net income (loss) per share*	\$ (0.24)	\$ 0.30	\$ (5.62)	
Diluted net income (loss) per share*	\$ (0.24)	\$ 0.28	\$ (5.62)	
<b>2012:</b>				
Total revenue	\$ 3,324	\$ 4,040	\$ 3,905	\$ 3,985
Total costs and expenses	(10,257)	(10,944)	(10,763)	(12,179)
Loss from operations	(6,933)	(6,904)	(6,858)	(8,194)
Net loss	(7,588)	(7,400)	(7,215)	(10,379)
Basic net loss per share*	\$ (1.50)	\$ (5.93)	\$ (5.82)	\$ (7.10)
Diluted net loss per share*	\$ (1.50)	\$ (5.93)	\$ (5.82)	\$ (7.10)
<b>2011:</b>				
Total revenue	\$ 6,260	\$ 12,925	\$ 57,534	\$ 4,192
Total costs and expenses	(11,442)	(11,497)	(9,680)	(9,736)
Income (loss) from operations	(5,182)	1,428	47,854	(5,544)
Net income (loss)	(5,725)	958	47,486	(6,452)
Basic net income (loss) per share*	\$ (4.88)	\$ (1.93)	\$ 2.33	\$ (5.03)
Diluted net income (loss) per share*	\$ (4.88)	\$ (1.93)	\$ 2.30	\$ (5.03)

(1) The amounts were computed independently for each quarter, and the sum of the quarters may not total the annual amounts.

\* Applicable to common stockholders

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**2,400,000 Shares**

**Common Stock**

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**PROSPECTUS**

**January 22, 2014**

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**Citigroup**

**Leerink Partners**

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**Piper Jaffray**

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**JMP Securities**

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