Pacira Pharmaceuticals, Inc. Form S-1 November 07, 2011

As filed with the Securities and Exchange Commission on November 7, 2011

Registration No. 333-

51-0619477

(I.R.S. Employer)

UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

Form S-1

REGISTRATION STATEMENT UNDER THE SECURITIES ACT OF 1933

PACIRA PHARMACEUTICALS, INC.

(Exact Name of Registrant as Specified in Its Charter)

Delaware

(State or Other Jurisdiction of Incorporation or Organization)

2834

(Primary Standard Industrial Classification Code No.) 5 Sylvan Way, Suite 100 Parsippany, New Jersey 07054 (973) 254-3560

(Address, including zip code, and telephone number, including area code, of registrant's principal executive offices)

David M. Stack President and Chief Executive Officer 5 Sylvan Way, Suite 125 Parsippany, New Jersey 07054 (973) 254-3560

(Name, address, including zip code, and telephone number, including area code, of agent for service)

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Approximate date of commencement of proposed sale to the public: As soon as practicable after this Registration Statement is declared effective.

If any of the securities being registered on this form are offered on a delayed or continuous basis pursuant to Rule 415 under the Securities Act of 1933, as amended (the "Securities Act") please check the following box. o

If this Form is filed to register additional securities for an offering pursuant to Rule 462(b) under the Securities Act, please check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering.

If this Form is a post-effective amendment filed pursuant to Rule 462(c) under the Securities Act, please check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering. o

If this Form is a post-effective amendment filed pursuant to Rule 462(d) under the Securities Act, please check the following box and list the Securities Act registration statement number of the earlier effective registration statement for the same offering. o

Indicate by check mark whether the registrant is a large accelerated filer, an accelerated filer, a non-accelerated filer, or a smaller reporting company. See the definitions of "large accelerated filer," "accelerated filer" and "smaller reporting company" in Rule 12b2 of the Exchange Act.

Large accelerated filer o Accelerated filer o Non-accelerated filer ý Smaller reporting company o
(Do not check if a smaller reporting company)

CALCULATION OF REGISTRATION FEE

	Class of Securities to be Registered	Proposed Maximum Aggregate Offering Price ¹	Amount of Registration Fee ²
Comm	on Stock, par value \$0.001 per share	\$53,199,000	\$6,097
1			
2	Estimated solely for the purpose of computing the registration fee in accordance with Rule 457(o)	under the Securities Act.	
2	Calculated pursuant to Rule 457(o) based on an estimate of the proposed maximum aggregate offer	ring price.	

The Registrant hereby amends this Registration Statement on such date or dates as may be necessary to delay its effective date until the Registrant shall file a further amendment which specifically states that this Registration Statement shall thereafter become effective in accordance with Section 8(a) of the Securities Act or until the Registration Statement shall become effective on such date as the Commission, acting pursuant to Section 8(a), may determine.

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The information in this prospectus is not complete and may be changed. We may not sell these securities until the registration statement filed with the Securities and Exchange Commission is effective. This prospectus is not an offer to sell these securities and it is not soliciting an offer to buy these securities in any state where the offer or sale is not permitted.

Subject to Completion, dated November 7, 2011.

PROSPECTUS

6,000,000 Shares

Common Stock

We are offering 6,000,000 shares of our common stock.

Our common stock trades on The NASDAQ Global Market under the symbol "PCRX." On November 4, 2011, the last reported trading price of our common stock was \$7.71 per share.

Investing in our common stock involves risks. See "Risk Factors" beginning on page 9 of this prospectus.

	Per		
	share	Total	
Price to the public	\$	\$	
Underwriting discounts and commissions	\$	\$	
Proceeds to us (before expenses)	\$	\$	

We have granted the underwriters the option to purchase 900,000 additional shares of common stock on the same terms and conditions set forth above if the underwriters sell more than 6,000,000 shares of common stock in this offering.

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.

The underwriters expect to deliver the shares on or about , 2011.

Barclays Capital			Jefferies
Piper Jaffray	Wedbush PacGrow Life Sciences Prospectus dated	Brean Murray, Carret & Co.	

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You should rely only on the information contained in this prospectus and any free writing prospectus prepared by or on behalf of us or to which we have referred you. We have not authorized anyone to provide you with information that is different. We are offering to sell shares of our common stock, and seeking offers to buy shares of our common stock, only in jurisdictions where offers and sales are permitted. The information in this prospectus is accurate only as of the date of this prospectus, regardless of the time of delivery of this prospectus or any sale of our common stock.

For investors outside the United States: neither we nor any of the underwriters have taken any action to permit a public offering of the shares of our common stock or the possession or distribution of this prospectus in any jurisdiction where action for that purpose is required, other than the United States. You are required to inform yourselves about and to observe any restrictions relating to this offering and the distribution of this prospectus.

We obtained the industry, market and competitive position data in this prospectus from our own internal estimates and research as well as from industry and general publications and research, surveys and studies conducted by third parties. Industry publications, studies and surveys generally state that they have been obtained from sources believed to be reliable, although they do not guarantee the accuracy or completeness of such information.

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PROSPECTUS SUMMARY

This summary highlights information contained elsewhere in this prospectus. You should read the following summary together with the more detailed information appearing in this prospectus, including our consolidated financial statements and related notes, and the risk factors beginning on page 9, before deciding whether to purchase shares of our common stock. Unless the context otherwise requires, we use the terms "Pacira," "our company," we," "us" and "our" in this prospectus to refer to Pacira Pharmaceuticals, Inc. and its subsidiaries.

Overview

We are an emerging specialty pharmaceutical company focused on the development, commercialization and manufacture of proprietary pharmaceutical products, based on our proprietary DepoFoam drug delivery technology, for use in hospitals and ambulatory surgery centers.

On October 28, 2011, the United States Food and Drug Administration, or FDA, approved our New Drug Application, or NDA, for our lead product candidate, EXPAREL, a liposome injection of bupivacaine, an amide-type local anesthetic, indicated for administration into the surgical site to produce postsurgical analgesia.

Our clinical data demonstrates that EXPAREL provides analgesia for up to 72 hours post-surgery, compared with approximately eight hours or less for bupivacaine. Bupivacaine and other shorter acting local anesthetics of the amide type such as mepivacaine and lidocaine are commonly used as the first line of treatment, pre- and post-operatively, of a multimodal postsurgical pain treatment regimen. Because bupivacaine, mepivacaine and lidocaine last approximately eight hours or less, administration of these local anesthetics is commonly followed by the systemic administration of opioids, such as morphine. Together, these drugs form the foundation of the multimodal postsurgical pain treatment regimen for the treatment of extended duration pain. Opioids are associated with a variety of significant adverse events leading to healthcare practitioners seeking opioid-sparing strategies for their patients and unfavorable hospital economics.

We believe EXPAREL addresses a significant unmet medical need for a long-acting non-opioid postsurgical analgesic, resulting in simplified postsurgical pain management and reduced opioid consumption, leading to improved patient outcomes and enhanced hospital economics. We estimate there are approximately 39 million opportunities annually in the United States for EXPAREL to be used.

EXPAREL will be launched by certain members of our management team who have successfully launched multiple products in the hospital market. Our commercial team has executed on a full range of pre-launch activities for EXPAREL including: (i) publications and abstracts for the EXPAREL clinical program efficacy and safety, health outcomes studies, and review articles on postsurgical pain management; (ii) health outcomes studies which provide retrospective and prospective analyses for our hospital customers using their own hospital data to demonstrate the true cost of opioid-based postsurgical pain management; (iii) key opinion leader, or KOL, development programs and advisory boards to address topics of best practice techniques, guidelines and protocols for the use of EXPAREL, educational needs of our physician, pharmacist and registered nurse customers, nerve block clinical studies and additional indications for the future development of EXPAREL; and (iv) education initiatives such as center of excellence programs, preceptorship programs, pain protocols and predictive models for enhanced patient care, web-based training and virtual launch programs.

We are developing a sales force entirely dedicated to commercializing EXPAREL comprised of approximately 60 representatives, seven regional managers and a national sales manager. We intend to develop this sales force pursuant to a contract with Quintiles Commercial US, Inc., a division of Quintiles, Inc., or Quintiles, and under the terms of this contract we have the flexibility to hire all or a portion of the sales force dedicated to commercializing EXPAREL as full-time employees of Pacira, upon 60 days notice to Quintiles. We believe that our pre-launch activities including significant personal

interactions with our hospital customers, position us for a successful launch of EXPAREL in the first quarter of 2012.

EXPAREL consists of bupivacaine encapsulated in DepoFoam, both of which are used in FDA-approved products. DepoFoam, our extended release drug delivery technology, is the basis for our two additional FDA-approved commercial products: DepoCyt(e) and DepoDur, which we manufacture for our commercial partners. DepoFoam-based products have been manufactured for over a decade and have an extensive safety record and regulatory approvals in the United States, European countries and other territories. Bupivacaine, a well-characterized, FDA-approved anesthetic/analgesic, has an established safety profile and over 20 years of use in the United States. We currently manufacture clinical supplies of EXPAREL and intend to manufacture and commercialize EXPAREL now that it has been approved.

The FDA-approved label for EXPAREL includes a broad label for postsurgical analgesia by local administration into the surgical site, or infiltration, a procedure commonly employing bupivacaine. The approved indication states "EXPAREL is a liposome injection of bupivacaine, an amide type local anesthetic, indicated for administration into the surgical site to produce postsurgical analgesia". We also plan to expand the indications of EXPAREL to include nerve block and epidural administration, markets where bupivacaine is also used routinely.

Our current product portfolio and product candidate pipeline is summarized in the table below:

Product(s)/Product Candidate(s) EXPAREL	Primary Indication(s) Postsurgical analgesia by infiltration	Status Approved by FDA	Commercialization Rights Pacira (worldwide)
	Postsurgical analgesia nerve block	Phase 2 (completed)	Pacira (worldwide)
	Postsurgical analgesia epidural administration	Phase 1 (completed)	Pacira (worldwide)
DepoCyt(e)	Lymphomatous meningitis	Marketed	Sigma-Tau Pharmaceuticals Mundipharma International
DepoDur	Post-operative pain	Marketed	EKR Therapeutics Flynn Pharmaceuticals
DepoNSAID	Acute pain	Preclinical	Pacira (worldwide)
DepoMethotrexate	Rheumatoid arthritis Oncology	Preclinical Preclinical	Pacira (worldwide) Pacira (worldwide)

Postsurgical Pain Market Overview

According to Thomson Reuters, roughly 45 million surgical procedures were performed in the United States during the twelve months ending in October 2007. We estimate there are approximately 39 million opportunities annually in the United States where EXPAREL could be used to improve patient outcomes and enhance hospital economics. Postsurgical pain is a response to tissue damage during surgery that stimulates peripheral nerves, which signal the brain to produce a sensory and psychological response. Numerous studies reveal that the incidence and severity of postsurgical pain is primarily determined by the type of surgery, duration of surgery and the pain treatment choice following surgery. Postsurgical pain is usually the most severe the first few days after the completion of a surgical procedure.

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Limitations of Current Therapies for Postsurgical Pain

Substantially all surgical patients experience postsurgical pain, with approximately 50% reporting inadequate pain relief according to certain epidemiological studies. Unrelieved acute pain causes patient suffering and can lead to other health problems, which delays recovery from surgery and may result in higher healthcare costs. According to the Agency for Healthcare Research and Quality, aggressive prevention of pain is better than treatment of pain because, once established, pain is more difficult to suppress. Current multimodal therapy for postsurgical pain includes wound infiltration with local anesthetics combined with the systemic administration of opioid and non-steroidal anti-inflammatory drug, or NSAID, analgesics.

Our Solution EXPAREL

EXPAREL provides continuous and extended postsurgical analgesia for up to 72 hours and reduces the consumption of supplemental opioid medications. We believe this will simplify postsurgical pain management, minimize breakthrough episodes of pain and result in improved patient outcomes and enhanced hospital economics.

Our EXPAREL strategy has four principal elements:

Replace the use of bupivacaine in postsurgical infiltration. Based on our clinical data, EXPAREL:

extends postsurgical analgesia for up to 72 hours, from approximately eight hours or less;

utilizes existing postsurgical infiltration administration techniques;

dilutes easily with normal saline to reach desired volume;

is a ready-to-use formulation; and

facilitates treatment of both small and large surgical sites.

Become the foundation of a postsurgical pain management regimen in order to reduce and delay opioid usage. Based on the clinical data from our Phase 3 hemorrhoidectomy trial as well as our retrospective health outcomes studies data, EXPAREL:

significantly delays and reduces opioid usage while improving postsurgical pain management:

delays first opioid usage to approximately 14 hours post-surgery, compared to approximately one hour for placebo;

significantly increases the percentage of patients requiring no opioid rescue medication through 72 hours post-surgery, to 28% compared to 10% for placebo;

results in 45% less opioid usage at 72 hours post-surgery compared to placebo; and

increases the percentage of patients who are pain free at 24 hours post-surgery compared to placebo.

Improve patient satisfaction. We believe EXPAREL:

provides effective pain control without the need for expensive and difficult to use delivery technologies which extend the duration of action for bupivacaine, such as elastomeric bags, or opioids administered through patient controlled analgesia, or PCA, when considered as part of a multimodal postsurgical pain regimen with an NSAID and acetaminophen and morphine rescue;

reduces the need for patients to be constrained by elastomeric bags and PCA systems, which are clumsy, difficult to use and may introduce catheter-related issues, including infection;

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promotes maintenance of early postsurgical pain management, thereby reducing the time spent in the intensive care unit; and

promotes early ambulation, which potentially reduces the risk of life-threatening blood clots, and allows quicker return of bowel function, thereby leading to a faster switch to oral nutrition and medicine, and thus a faster discharge from the hospital.

Develop and seek approval of EXPAREL for nerve block and epidural administration. We believe these additional indications for EXPAREL:

presents a low-risk, low-cost opportunity for clinical development; and

enables us to fully leverage our manufacturing and sales infrastructure.

Maufacturing and Intellectual Property

We manufacture all our DepoFoam-based products, including commercial supplies of DepoCyt(e) and DepoDur for our commercial partners. We currently manufacture clinical supplies of EXPAREL and intend to manufacture and commercialize EXPAREL now that it has been approved.

We have developed significant know-how regarding our manufacturing process and protect our technology through trade secrets and patents. We have over 15 families of patents and patent applications relating to various aspects of DepoFoam delivery technology. Issued U.S. patents protect the composition of EXPAREL and methods for modifying its rate of drug release. We have also submitted additional patent applications related to the composition of, and manufacturing process for, EXPAREL. In addition, in April 2011 we filed a non-provisional patent application relating to a new process to manufacture EXPAREL and other DepoFoam-based products, which, if granted, could prevent others from using this process until 2031.

Our Strategy

Our goal is to be a leading specialty pharmaceutical company focused on the development, commercialization and manufacture of proprietary pharmaceutical products principally for use in hospitals and ambulatory surgery centers. We plan to achieve this by:

commercializing EXPAREL in the United States for postsurgical analgesia by infiltration into the surgical site;

building a streamlined commercial organization concentrating on major hospitals and ambulatory surgery centers in the United States and targeting surgeons, anesthesiologists, pharmacists and nurses;

working directly with managed care payers, quality improvement organizations, KOLs in the field of postsurgical pain management and leading influence hospitals with Phase 3b, Phase 4 retrospective and prospective trials to demonstrate the economic benefits of EXPAREL;

securing commercial partnerships for EXPAREL in regions outside of the United States;

obtaining FDA approval for nerve block and epidural administration indications for EXPAREL;

manufacturing all our DepoFoam-based products, including EXPAREL, DepoCyt(e) and DepoDur, in our current Good Manufacturing Practices, or cGMP, compliant facilities;

continuing to expand our marketed product portfolio through development of additional DepoFoam-based hospital products utilizing a 505(b)(2) strategy, which permits us to rely upon the FDA's previous findings of safety and effectiveness for an approved product. A 505(b)(2) strategy may not succeed if there are successful challenges to the FDA's interpretation of Section 505(b)(2); and

continuing research and development partnerships to provide DepoFoam-based products to enhance the duration of action and patient compliance for partner products.

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Risk Factors

Our business is subject to a number of risks of which you should be aware before making an investment decision. These risks are discussed more fully in the "Risk Factors" section of this prospectus immediately following this prospectus summary. These risks include the following:

Our success depends on our ability to successfully commercialize EXPAREL.

If we are unable to establish effective marketing and sales capabilities to market and sell EXPAREL, we may be unable to generate product revenues.

Our efforts to successfully commercialize EXPAREL are subject to many internal and external challenges and if we cannot overcome these challenges in a timely manner, our future revenues and profits could be materially and adversely impacted.

If we fail to manufacture EXPAREL in sufficient quantities and at acceptable quality and pricing levels, or to fully comply with cGMP regulations, we may face delays in the commercialization of this product or be unable to meet market demand, and may lose potential revenues.

We have doubts about our ability to continue as a going concern, which may hinder our ability to obtain future financing.

We may not be able to manage our business effectively if we are unable to attract and retain key personnel.

Corporate History and Information

We were incorporated in Delaware under the name Blue Acquisition Corp. in December 2006 and changed our name to Pacira, Inc. in June 2007. In October 2010, we changed our name to Pacira Pharmaceuticals, Inc.

Pacira Pharmaceuticals, Inc. is the holding company for our California operating subsidiary of the same name, which we refer to as PPI-California. On March 24, 2007, MPM Capital, Sanderling Ventures, OrbiMed Advisors, HBM BioVentures, the Foundation for Research and their co-investors, through Pacira Pharmaceuticals, Inc., acquired PPI-California, from SkyePharma Holding, Inc., which we refer to as the Acquisition. PPI-California was known as SkyePharma, Inc. prior to the Acquisition. In this prospectus, the term Predecessor refers to SkyePharma, Inc. prior to March 24, 2007, or the Acquisition Date, and the term Successor refers to Pacira Pharmaceuticals, Inc. and its consolidated subsidiaries.

Our principal executive offices are located at 5 Sylvan Way, Suite 100, Parsippany, New Jersey 07054, and our telephone number is (973) 254-3560. Our website address is www.pacira.com. Information contained on our website is not incorporated by reference into this prospectus, and you should not consider information contained on our website to be part of this prospectus or in deciding whether to purchase shares of our common stock.

Pacira®, DepoFoam®, DepoCyt® (U.S. registration), DepoCyte® (EU registration), DepoDur®, EXPAREL , the Pacira logo and other trademarks or service marks of Pacira appearing in this prospectus are the property of Pacira. This prospectus contains additional trade names, trademarks and service marks of other companies. In the prospectus, references to DepoCyt(e) mean DepoCyt when discussed in the context of the United States and Canada and DepoCyte when discussed in the context of Europe.

The Offering

Common stock offered by Pacira 6,000,000 shares

Common stock to be outstanding after this

offering

23,228,827 shares (24,128,827 shares in the event the underwriters elect to exercise their option

to purchase additional shares from us in full)

Use of proceeds We estimate that the net proceeds to us from this offering, after deducting estimated

underwriting discounts and commissions and estimated offering expenses, will be approximately \$43.0 million, or \$49.5 million if the underwriters exercise their option to purchase additional shares from us in full. We intend to use the net proceeds from this offering for the planned manufacture and commercialization of EXPAREL in the United States, and for

working capital and other general corporate purposes.

See "Use of Proceeds."

Risk factors You should read the "Risk Factors" section and other information included in this prospectus

for a discussion of factors to consider carefully before deciding to invest in shares of our

common stock.

"PCRX"

The NASDAQ Global Market symbol

The number of shares of our common stock to be outstanding after this offering is based on 17,228,827 shares of common stock outstanding as of September 30, 2011, and excludes:

2,332,287 shares of common stock issuable upon the exercise of stock options outstanding as of September 30, 2011, at a weighted average exercise price of \$3.72 per share;

61,242 shares of common stock available for future issuance under our equity compensation plans as of September 30, 2011;

527,656 shares of common stock issuable upon the exercise of warrants outstanding and exercisable at a weighted average exercise price of \$10.22 per share; and

36,750 shares of common stock issuable upon the exercise of outstanding stock options that we granted after September 30, 2011, at a weighted average exercise price of \$9.74 per share.

Except as otherwise noted, all information in this prospectus:

assumes no exercise of outstanding options or warrants; and

assumes no exercise by the underwriters of their option to purchase additional shares of common stock to cover over-allotments.

Summary Consolidated Financial Data

The following tables summarize our consolidated financial data as of the dates and for the periods indicated. You should read this data together with our financial statements and related notes included elsewhere in this prospectus and the information under "Selected Consolidated Financial Data" and "Management's Discussion and Analysis of Financial Condition and Results of Operations." The consolidated financial data in this section is not intended to replace our consolidated financial statements and the accompanying notes. The unaudited consolidated financial data include, in the opinion of our management, all adjustments, consisting only of normal recurring adjustments that are necessary for a fair presentation of our financial position and results of operations for these periods. Our historical results for any prior period are not necessarily indicative of results to be expected in any future period, and our results for any interim period are not necessarily indicative of results to be expected for a full fiscal year.

The consolidated financial data for the years ended December 31, 2008, 2009 and 2010 have been derived from our audited consolidated financial statements included elsewhere in this prospectus. The consolidated financial data as of September 30, 2011, and for the nine months ended September 30, 2010 and 2011, have been derived from our unaudited consolidated financial statements included elsewhere in this prospectus.

The as adjusted consolidated balance sheet data at September 30, 2011 gives effect to our sale of shares of common stock offered by this prospectus at an assumed public offering price of \$7.71 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on November 4, 2011, after deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us.

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		Year Ended December 31,				Nine Months Ended September 30,					
		2008 2009 2010			2010	2010		2011			
			(audited)				(unaudited)		(unaudited)	
				(in tho	usa	nds, except	share and per share data)				
Consolidated Statement of								-			
Operations Data:											
Revenues	\$	13,925	\$	15,006	\$	14,562	\$	12,371	\$	11,456	
Operating Expenses											
Cost of revenues		17,463		12,301		12,276		10,168		10,138	
Research and development		33,214		26,233		18,628		14,954		12,237	
Selling, general and administrative		8,611		5,020		6,030		3,948		13,465	
Total operating expenses		59,288		43,554		36,934		29,070		35,840	
				,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,				_,,,,,		22,010	
Loss from operations		(45,363)		(28,548)		(22,372)		(16,699)		(24,384)	
Other income (expense):		(43,303)		(20,540)		(22,372)		(10,099)		(24,304)	
Loss on early extinguishment of debt						(184)					
Interest income		235		77		146		112		111	
Interest expense		233		(1,723)		(3,959)		(2,577)		(4,068)	
Royalty interest obligation		3,490		(1,880)		(930)		(1,048)		235	
Other income (expense)		(224)		367		150		107		61	
other meome (expense)		(221)		307		130		107		01	
Total other income (evenes) not		3,501		(3,159)		(4 777)		(3,406)		(2.661)	
Total other income (expense), net		3,301		(3,139)		(4,777)		(3,400)		(3,661)	
	_		_		_		_		_		
Net loss	\$	(41,862)	\$	(31,707)	\$	(27,149)	\$	(20,105)	\$	(28,045)	
Net loss per common share basic and											
diluted	\$	(79.23)	\$	(55.32)	\$	(47.29)	\$	(35.02)	\$	(1.89)	
Weighted average number of common											
shares-basic and diluted		528,357		573,118		574,072		574,112		14,826,054	

As of	Septembe	er 30, 2011
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	Actual	As	Adjusted	
	(unaudited, in thousands)			
Consolidated Balance Sheet Data:				
Cash and cash equivalents	\$ 16,402	\$	59,386	
Short-term investments	20,666		20,666	
Working capital	25,571		68,555	
Total assets	77,609		120,593	
Long-term debt, including current portion	25,474		25,474	
Common stock	17		23	
Additional paid-in capital	178,821		221,799	
Accumulated deficit	(164,956)		(164,956)	
Total stockholders' equity	13,875		56,859	
		8		

RISK FACTORS

Investing in our common stock involves a high degree of risk. Before you decide to invest in our common stock, you should consider carefully the risks described below, together with the other information contained in this prospectus, including our financial statements and the related notes appearing at the end of this prospectus. We believe the risks described below are the risks that are material to us as of the date of this prospectus. If any of the following risks actually occur, our business, financial condition, results of operations and future growth prospects would likely be materially and adversely affected. In these circumstances, the market price of our common stock could decline, and you may lose all or part of your investment.

Risks Related to the Development and Commercialization of our Product Candidates

Our success depends on our ability to successfully commercialize EXPAREL.

We have invested a significant portion of our efforts and financial resources in the development of EXPAREL. Our success depends on our ability to effectively commercialize EXPAREL, which was approved by the FDA on October 28, 2011, for administration into the surgical site to produce postsurgical analgesia.

We plan to commercially launch EXPAREL in the first quarter of 2012, but our ability to effectively commercialize and generate revenues from EXPAREL will depend on our ability to:

create market demand for EXPAREL through our marketing and sales activities, and any other arrangements to promote this product we may later establish;

train, deploy and support a qualified sales force which will be developed on a contract basis with Quintiles;

secure formaulary approvals for EXPAREL at a substantial number of targeted hospitals;

manufacture EXPAREL in sufficient quantities in compliance with requirements of the FDA and similar foreign regulatory agencies and at acceptable quality and pricing levels in order to meet commercial demand;

implement and maintain agreements with wholesalers, distributors and group purchasing organizations on commercially reasonable terms:

receive adequate levels of coverage and reimbursement for EXPAREL from commercial health plans and governmental health programs;

maintain compliance with regulatory requirements;

ensure that our entire supply chain for EXPAREL efficiently and consistently delivers EXPAREL to our customers; and

maintain and defend our patent protection and regulatory exclusivity for EXPAREL.

Any disruption in our ability to generate revenues from the sale of EXPAREL or lack of success in its commercialization will have a material and adverse impact on our results of operations.

Our efforts to successfully commercialize EXPAREL are subject to many internal and external challenges and if we cannot overcome these challenges in a timely manner, our future revenues and profits could be materially and adversely impacted.

As EXPAREL will be a newly marketed drug, none of the members of the EXPAREL sales force have ever promoted EXPAREL. As a result, we are required to expend significant time and resources to train the sales force to be credible and persuasive in convincing physicians and hospitals to use

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EXPAREL. In addition, we also must train the sales force to ensure that a consistent and appropriate message about EXPAREL is delivered to our potential customers. If we are unable to effectively train the sales force and equip them with effective materials, including medical and sales literature to help them inform and educate potential customers about the benefits and risks of EXPAREL and its proper administration, our efforts to successfully commercialize EXPAREL could be put in jeopardy, which could have a material adverse effect on our future revenues and profits.

In addition to our extensive internal efforts, the successful commercialization of EXPAREL will require many third parties, over whom we have no control, to choose to utilize EXPAREL. These third parties include physicians and hospital pharmacy and therapeutics committees, which we refer to as P&T committees. Generally, before we can attempt to sell EXPAREL in a hospital, EXPAREL must be approved for addition to that hospital's list of approved drugs, or formulary list, by the hospital's P&T committee. A hospital's P&T committee typically governs all matters pertaining to the use of medications within the institution, including the review of medication formulary data and recommendations for the appropriate use of drugs within the institution to the medical staff. The frequency of P&T committee meetings at hospitals varies considerably, and P&T committees often require additional information to aid in their decision-making process. Therefore, we may experience substantial delays in obtaining formulary approvals. Additionally, hospital pharmacists may be concerned that the cost of acquiring EXPAREL for use in their institutions will adversely impact their overall pharmacy budgets, which could cause pharmacists to resist efforts to add EXPAREL to the formulary, or to implement restrictions on the usage of EXPAREL in order to control costs. We cannot guarantee that we will be successful in obtaining the approvals we need from enough P&T committees quickly enough to optimize hospital sales of EXPAREL.

Even if we obtain hospital formulary approval for EXPAREL, physicians must still prescribe EXPAREL for its commercialization to be successful. Because EXPAREL is a new drug with no track record of sales in the United States, any inability to timely supply EXPAREL to our customers, or any unexpected side effects that develop from use of the drug, particularly early in product launch, may lead physicians to not accept EXPAREL as a viable treatment alternative.

If EXPAREL does not achieve broad market acceptance, the revenues that we generate from its sales will be limited. The degree of market acceptance of EXPAREL will also depend on a number of other factors, including:

changes in the standard of care for the targeted indications for EXPAREL, which could reduce the marketing impact of any claims that we could make following FDA approval;

the relative convenience and ease of administration of EXPAREL;

the prevalence and severity of adverse events associated with EXPAREL;

cost of treatment versus economic and clinical benefit in relation to alternative treatments;

the availability of adequate coverage or reimbursement by third parties, such as insurance companies and other healthcare payers, and by government healthcare programs, including Medicare and Medicaid;

the extent and strength of our marketing and distribution of EXPAREL;

the safety, efficacy and other potential advantages over, and availability of, alternative treatments, including, in the case of EXPAREL, a number of products already used to treat pain in the hospital setting; and

distribution and use restrictions imposed by the FDA or to which we agree as part of a mandatory risk evaluation and mitigation strategy or voluntary risk management plan.

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Our ability to effectively promote and sell EXPAREL and any product candidates that we may develop, license or acquire in the hospital marketplace will also depend on pricing and cost effectiveness, including our ability to produce a product at a competitive price and therefore achieve acceptance of the product onto hospital formularies, and our ability to obtain sufficient third-party coverage or reimbursement. Since many hospitals are members of group purchasing organizations, which leverage the purchasing power of a group of entities to obtain discounts based on the collective buying power of the group, our ability to attract customers in the hospital marketplace will also depend on our ability to effectively promote our product candidates to group purchasing organizations. We will also need to demonstrate acceptable evidence of safety and efficacy, as well as relative convenience and ease of administration. Market acceptance could be further limited depending on the prevalence and severity of any expected or unexpected adverse side effects associated with our product candidates.

In addition, the labeling approved by the FDA does not contain claims that EXPAREL is safer or more effective than competitive products and does not permit us to promote EXPAREL as being superior to competing products. Further, the availability of inexpensive generic forms of postsurgical pain management products may also limit acceptance of EXPAREL among physicians, patients and third-party payers. If EXPAREL does not achieve an adequate level of acceptance among physicians, patients and third-party payers, we may not generate meaningful revenues from EXPAREL and we may not become profitable.

We face significant competition from other pharmaceutical and biotechnology companies. Our operating results will suffer if we fail to compete effectively.

The pharmaceutical and biotechnology industries are intensely competitive and subject to rapid and significant technological change. Our major competitors include organizations such as major multinational pharmaceutical companies, established biotechnology companies and specialty pharmaceutical and generic drug companies. Many of our competitors have greater financial and other resources than we have, such as larger research and development staff, more extensive marketing, distribution, sales and manufacturing organizations and experience, more extensive clinical trial and regulatory experience, expertise in prosecution of intellectual property rights and access to development resources like personnel generally and technology. As a result, these companies may obtain regulatory approval more rapidly than we are able to and may be more effective in selling and marketing their products. Smaller or early stage companies may also prove to be significant competitors, particularly through collaborative arrangements with large, established companies. Our competitors may succeed in developing, acquiring or licensing on an exclusive basis technologies and drug products that are more effective or less costly than EXPAREL or any product candidate that we are currently developing or that we may develop, which could render our products obsolete and noncompetitive or significantly harm the commercial opportunity for EXPAREL or our product candidates.

As a result of these factors, our competitors may obtain patent protection or other intellectual property rights that limit our ability to develop other indications for, or commercialize, EXPAREL. Our competitors may also develop drugs that are more effective, useful or less costly than ours and may be more successful than us in manufacturing and marketing their products.

EXPAREL will compete with well-established products with similar indications. Competing products available for postsurgical pain management include opioids such as morphine, fentanyl, meperidine and hydromorphone, each of which is available generically from several manufacturers, and several of which are available as proprietary products using novel delivery systems. Ketorolac, an injectable non-steroidal anti-inflammatory drug, or NSAID, is also available generically in the United States from several manufacturers, and Caldolor (ibuprofen for injection), an NSAID, has been approved by the FDA for pain management and fever in adults. In addition, EXPAREL will compete with non-opioid products such as bupivacaine, Marcaine, ropivacaine and other anesthetics/analgesics, all of which are also used in the treatment of postsurgical pain and are available as either oral tablets,

injectable dosage forms or administered using novel delivery systems. Additional products may be developed for the treatment of acute pain, including new injectable NSAIDs, novel opioids, new formulations of currently available opioids and NSAIDS, long-acting local anesthetics and new chemical entities as well as alternative delivery forms of various opioids and NSAIDs.

We will also compete with an extended release bupivacaine product in development by Durect Corporation which has been licensed to Hospira in North America (Posidur) and to Nycomed for Europe (Optesia). EXPAREL also competes with elastomeric bag/catheter devices intended to provide bupivacaine over several days. I-FLOW Corporation (acquired by Kimberly-Clark Corporation in 2009) has marketed these medical devices in the United States since 2004.

If we are unable to establish effective marketing and sales capabilities or enter into agreements with third parties to market and sell EXPAREL, we may be unable to generate product revenues.

We are currently building our commercial infrastructure for the marketing, sale and distribution of pharmaceutical products. In order to commercialize EXPAREL, we must build our marketing, sales and distribution capabilities. We have entered into an agreement with Quintiles for the outsourcing of our specialty sales force of approximately 60 representatives. We may also seek to commercialize EXPAREL outside the United States, although we currently plan to do so with a marketing and sales collaborator and not with our own sales force.

The establishment, development and training of our sales force and related compliance plans to market EXPAREL is expensive and time consuming and can potentially delay the commercial launch of EXPAREL. In the event we are not successful in developing our marketing and sales infrastructure, we may not be able to successfully commercialize EXPAREL, which would limit our ability to generate product revenues.

We rely on third parties to perform many essential services for EXPAREL and any other products that we commercialize, including services related to customer service support, warehousing and inventory program services, distribution services, contract administration and chargeback processing services, accounts receivable management and cash application services, and financial management and information technology services. If these third parties fail to perform as expected or to comply with legal and regulatory requirements, our ability to commercialize EXPAREL will be significantly impacted and we may be subject to regulatory sanctions.

We have entered into agreements with third-party service providers to perform a variety of functions related to the sale and distribution of EXPAREL, key aspects of which are out of our direct control. These service providers provide key services related to customer service support, warehousing and inventory program services, distribution services, contract administration and chargeback processing services, accounts receivable management and cash application services, and financial management and information technology services. In addition, most of our inventory is stored at a single warehouse maintained by one such service provider. We substantially rely on these providers as well as other third-party providers that perform services for us, including entrusting our inventories of products to their care and handling. If these third-party service providers fail to comply with applicable laws and regulations, fail to meet expected deadlines, or otherwise do not carry out their contractual duties to us, or encounter physical or natural damage at their facilities, our ability to deliver product to meet commercial demand would be significantly impaired. In addition, we may engage third parties to perform various other services for us relating to adverse event reporting, safety database management, fulfillment of requests for medical information regarding our product candidates and related services. If the quality or accuracy of the data maintained by these service providers is insufficient, we could be subject to regulatory sanctions.

Distribution of our DepoFoam-based products, including EXPAREL, requires cold-chain distribution provided by third parties, whereby the product must be maintained between specified

temperatures. We and our partners have utilized similar cold-chain processes for DepoCyt(e) and DepoDur. If a problem occurs in our cold-chain distribution processes, whether through our failure to maintain our products or product candidates between specified temperatures or because of a failure of one of our distributors or partners to maintain the temperature of the products or product candidates, the product or product candidate could be adulterated and rendered unusable. This could have a material adverse effect on our business, financial condition, results of operations and reputation.

We will need to increase the size of our organization and effectively manage our sales force, and we may experience difficulties in managing growth.

As of September 30, 2011, we had 122 employees. We will need to substantially expand our managerial, commercial, financial, manufacturing and other personnel resources in order to manage our operations and prepare for the commercialization of EXPAREL. Our management, personnel, systems and facilities currently in place may not be adequate to support this future growth. In addition, we may not be able to recruit and retain qualified personnel in the future, particularly marketing positions, due to competition for personnel among pharmaceutical businesses, and the failure to do so could have a significant negative impact on our future product revenues and business results. We will also need to effectively manage our sales force that we outsource from Quintiles. Our need to effectively manage our operations, growth and various projects requires that we:

continue the hiring, outsourcing in the case of our sales force, and training of an effective commercial organization for the commercialization of EXPAREL, and establish appropriate systems, policies and infrastructure to support that organization;

ensure that our consultants and other service providers successfully carry out their contractual obligations, provide high quality results, and meet expected deadlines;

continue to carry out our own contractual obligations to our licensors and other third parties; and

continue to improve our operational, financial and management controls, reporting systems and procedures.

We may be unable to successfully implement these tasks on a larger scale and, accordingly, may not achieve our development and commercialization goals.

We are reliant on our contract with Quintiles for the marketing and sale of EXPAREL.

We have entered into an agreement with Quintiles for the outsourcing of a sales force to commercialize EXPAREL. The risks in outsourcing the sales function to any third party include the following:

the third party may not apply the expected financial resources or required expertise to successfully market and sell EXPAREL;

the third party may not invest in the development of a sales and marketing force and the related infrastructure at levels that ensure that sales of EXPAREL reach their full potential;

the third party may not comply with applicable legal requirements, including the requirement to promote drug products only for uses for which they have been approved;

disputes may arise between us and the third party that may delay the commercialization of EXPAREL or adversely affect its sales or profitability; or

the third party may enter into agreements with other parties that have products that could compete with EXPAREL.

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We are substantially dependent on the success of Quintiles in performing its responsibilities and the continued cooperation of Quintiles, including Quintiles' cooperation with our training of the sales and marketing force. Quintiles may not cooperate with us to perform its obligations under our agreement and we cannot control the amount and timing of Quintiles' resources that will be devoted to the marketing and sale of EXPAREL. The occurrence of any of these events could adversely affect the commercialization of EXPAREL and materially harm our business and stock price by slowing the pace of growth of such sales, by reducing the profitability of EXPAREL or by adversely affecting the reputation of EXPAREL in the market.

We may not be able to manage our business effectively if we are unable to attract and retain key personnel.

We may not be able to attract or retain qualified management and commercial, scientific and clinical personnel due to the intense competition for qualified personnel among biotechnology, pharmaceutical and other businesses, particularly in the San Diego, California and Northern New Jersey areas. If we are not able to attract and retain necessary personnel to accomplish our business objectives, we may experience constraints that will significantly impede the achievement of our development objectives, our ability to raise additional capital and our ability to implement our business strategy.

Our industry has experienced a high rate of turnover of management personnel in recent years. We are highly dependent on the development and manufacturing expertise for our DepoFoam delivery technology and the commercialization expertise of certain members of our senior management. In particular, we are highly dependent on the skills and leadership of our management team, including David Stack, our president and chief executive officer. If we lose one or more of these key employees, our ability to successfully implement our business strategy could be seriously harmed. Replacing key employees may be difficult and may take an extended period of time because of the limited number of individuals in our industry with the breadth of skills and experience required to develop, gain regulatory approval of and commercialize products successfully. Competition to hire from this limited pool is intense, and we may be unable to hire, train, retain or motivate additional key personnel.

Mr. Stack, our chief executive officer, is also a managing director at MPM Capital and a managing partner of Stack Pharmaceuticals, Inc. Although Mr. Stack has devoted substantially all of his time to our company over the past 12 months, Mr. Stack's responsibilities at MPM Capital and Stack Pharmaceuticals, Inc. might require that he spend less than all his time managing our company in the future.

Under our consulting agreement with Gary Patou, M.D., our chief medical officer, he is not required to devote all of his time to our company. We cannot assure you that Dr. Patou's time commitment to us will be sufficient to perform the duties of our chief medical officer.

We face potential product liability exposure, and if successful claims are brought against us, we may incur substantial liability for DepoCyt(e), DepoDur, EXPAREL or product candidates that we may develop and may have to limit their commercialization.

The use of DepoCyt(e), DepoDur, EXPAREL and any product candidates that we may develop, license or acquire in clinical trials and the sale of any products for which we obtain regulatory approval expose us to the risk of product liability claims. Product liability claims might be brought against us by consumers, health care providers or others using, administering or selling our products. If we cannot successfully defend ourselves against these claims, we will incur substantial liabilities. Regardless of merit or eventual outcome, liability claims may result in:

loss of revenue from decreased demand for our products and/or product candidates;

impairment of our business reputation or financial stability;

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costs of related litigation;
substantial monetary awards to patients or other claimants;
diversion of management attention;
loss of revenues;
withdrawal of clinical trial participants and potential termination of clinical trial sites or entire clinical programs; and
the inability to commercialize our product candidates.

We have obtained limited product liability insurance coverage for our products and our clinical trials with a \$10.0 million annual aggregate coverage limit. However, our insurance coverage may not reimburse us or may not be sufficient to reimburse us for any expenses or losses we may suffer. Moreover, insurance coverage is becoming increasingly expensive, and, in the future, we may not be able to maintain insurance coverage at a reasonable cost or in sufficient amounts to protect us against losses due to liability. We intend to expand our insurance coverage to include the sale of additional commercial products upon FDA approval for our product candidates in development, but we may be unable to obtain commercially reasonable product liability insurance for any products approved for marketing, or at all. On occasion, large judgments have been awarded in class action lawsuits based on drugs that had unanticipated side effects. A successful product liability claim or series of claims brought against us could cause our stock price to fall and, if judgments exceed our insurance coverage, could decrease our cash and adversely affect our business.

If we fail to manufacture EXPAREL in sufficient quantities and at acceptable quality and pricing levels, or to fully comply with cGMP regulations, we may face delays in the commercialization of this product or be unable to meet market demand, and may lose potential revenues.

The manufacture of EXPAREL requires significant expertise and capital investment, including the development of advanced manufacturing techniques and process controls, and the use of specialized processing equipment. We must comply with federal, state and foreign regulations, including FDA's regulations governing current Good Manufacturing Practices, or cGMP, enforced by the FDA through its facilities inspection program and by similar regulatory authorities in other jurisdictions where we do business. These requirements include, among other things, quality control, quality assurance and the maintenance of records and documentation. The FDA or similar foreign regulatory authorities at any time may implement new standards, or change their interpretation and enforcement of existing standards for manufacture, packaging or testing of our products. Any failure to comply with applicable regulations may result in fines and civil penalties, suspension of production, product seizure or recall, imposition of a consent decree, or withdrawal of product approval, and would limit the availability of our product. Any manufacturing defect or error discovered after products have been produced and distributed also could result in significant consequences, including costly recall procedures, re-stocking costs, damage to our reputation and potential for product liability claims.

We purchase raw materials and components from various suppliers in order to manufacture EXPAREL. If we are unable to source the required raw materials from our suppliers, we may experience delays in manufacturing EXPAREL and may not be able to meet our customers' demands for EXPAREL.

If we are unable to produce the required commercial quantities of EXPAREL to meet market demand for EXPAREL on a timely basis or at all, or if we fail to comply with applicable laws for the manufacturing of EXPAREL, we will suffer damage to our reputation and commercial prospects and we will lose potential revenues.

We are the sole manufacturer of DepoCyt(e) and DepoDur and we only have two FDA approved manufacturing facilities. Our inability to continue manufacturing adequate supplies of DepoCyt(e) and DepoDur could result in a disruption in the supply of DepoCyt(e) and DepoDur to our partners.

We are the sole manufacturer of DepoCyt(e) and DepoDur. We develop and manufacture DepoCyt(e) and DepoDur at our facilities in San Diego, California, which are the only FDA approved sites for manufacturing DepoCyt(e) and DepoDur in the world. Our San Diego facilities are subject to the risks of a natural or man-made disaster, including earthquakes and fires, or other business disruption. There can be no assurance that we would be able to meet our requirements for DepoCyt(e) and DepoDur if there were a catastrophic event or failure of our current manufacturing system. If we are required to change or add a new manufacturer or supplier, the process would likely require prior FDA and/or equivalent foreign regulatory authority approval, and would be very time consuming. An inability to continue manufacturing adequate supplies of DepoCyt(e) and DepoDur at our facility in San Diego, California could result in a disruption in the supply of DepoCyt(e) and DepoDur to our partners and breach of our contractual obligations.

If we fail to manufacture DepoCyt(e) and DepoDur we will lose revenues and be in breach of our licensing obligations.

We have licensed the commercial rights in specified territories of the world to market and sell our products, DepoCyt(e) and DepoDur. Under those licenses we have obligations to manufacture commercial product for our commercial partners. If we are unable to timely fill the orders placed with us by our commercial partners, we will potentially lose revenue and be in breach of our licensing obligations under the agreements. In addition, we would be in breach of our obligations to comply with our supply and distribution agreements for DepoCyt(e) and DepoDur, which would in turn be a breach of our obligations under our amended and restated royalty interests assignment agreement, or the Amended and Restated Royalty Interests Assignment Agreement, with Royalty Securitization Trust I, an affiliate of Paul Capital Advisors, LLC, or Paul Capital. See "Risk Factors Risks Related to Our Financial Condition and Capital Requirements Under our financing arrangement with Paul Capital, upon the occurrence of certain events, Paul Capital may require us to repurchase the right to receive royalty payments that we assigned to it, or may foreclose on certain assets that secure our obligations to Paul Capital. Any exercise by Paul Capital of its right to cause us to repurchase the assigned right or any foreclosure by Paul Capital would adversely affect our results of operations and our financial condition."

We rely on third parties for the timely supply of specified raw materials and equipment for the manufacture of DepoCyt(e) and DepoDur. Although we actively manage these third-party relationships to provide continuity and quality, some events which are beyond our control could result in the complete or partial failure of these goods and services. Any such failure could have a material adverse effect on our financial condition and operations.

The manufacture of pharmaceutical products requires significant expertise and capital investment, including the development of advanced manufacturing techniques and process controls, and the use of specialized processing equipment. We must comply with federal, state and foreign regulations, including current Good Manufacturing Practices, or cGMP, regulations and in the case of the manufacturing of DepoDur required government licenses regarding the storage and use of controlled substances. Any failure to comply with applicable regulations may result in fines and civil penalties, suspension of production, suspension or delay in product approval for sale, product seizure or recall, or withdrawal of product approval, and would limit the availability of our product. Any manufacturing defect or error discovered after products have been produced and distributed could result in even more significant consequences, including costly recall procedures, re-stocking costs, damage to our reputation, product liability claims and litigation.

Our future growth depends on our ability to identify, develop, acquire or in-license products and if we do not successfully identify develop, acquire or in-license related product candidates or integrate them into our operations, we may have limited growth opportunities.

An important part of our business strategy is to continue to develop a pipeline of product candidates by developing, acquiring or in-licensing products, businesses or technologies that we believe are a strategic fit with our focus on the hospital marketplace. However, these business activities may entail numerous operational and financial risks, including:

difficulty or inability to secure financing to fund development activities for such development, acquisition or in-licensed products or technologies;

incurrence of substantial debt or dilutive issuances of securities to pay for development, acquisition or in-licensing of new products;

disruption of our business and diversion of our management's time and attention;

higher than expected development, acquisition or in-license and integration costs;

exposure to unknown liabilities;

difficulty and cost in combining the operations and personnel of any acquired businesses with our operations and personnel;

inability to retain key employees of any acquired businesses;

difficulty in managing multiple product development programs; and

inability to successfully develop new products or clinical failure.

We have limited resources to identify and execute the development, acquisition or in-licensing of products, businesses and technologies and integrate them into our current infrastructure. We may compete with larger pharmaceutical companies and other competitors in our efforts to establish new collaborations and in-licensing opportunities. These competitors likely will have access to greater financial resources than us and may have greater expertise in identifying and evaluating new opportunities. Moreover, we may devote resources to potential development, acquisitions or in-licensing opportunities that are never completed, or we may fail to realize the anticipated benefits of such efforts.

Our business involves the use of hazardous materials and we must comply with environmental laws and regulations, which can be expensive and restrict how we do business.

Our manufacturing activities involve the controlled storage, use and disposal of hazardous materials, including the components of our products, product candidates and other hazardous compounds. We are subject to federal, state and local laws and regulations governing the use, manufacture, storage, handling, release and disposal of, and exposure to, these hazardous materials. Violation of these laws and regulations could lead to substantial fines and penalties. Although we believe that our safety procedures for handling and disposing of these materials comply with the standards prescribed by these laws and regulations, we cannot eliminate the risk of accidental contamination or injury from these materials. In the event of an accident, state or federal authorities may curtail our use of these materials and interrupt our business operations. In addition, we could become subject to potentially material liabilities relating to the investigation and cleanup of any contamination, whether currently unknown or caused by future releases.

Our business and operations would suffer in the event of system failures.

Despite the implementation of security measures, our internal computer systems are vulnerable to damage from computer viruses, unauthorized access, natural disasters, terrorism, war and

telecommunication and electrical failures. Any system failure, accident or security breach that causes interruptions in our operations could result in a material disruption of our product development programs. For example, the loss of clinical trial data from completed clinical trials for EXPAREL could result in delays in our regulatory approval efforts and significantly increase our costs to recover or reproduce the data. To the extent that any disruption or security breach results in a loss or damage to our data or applications, or inappropriate disclosure of confidential or proprietary information, we may incur liability and the further development of our product candidates may be delayed.

Any collaboration arrangements that we may enter into in the future may not be successful, which could adversely affect our ability to develop and commercialize our product candidates.

Our business model is to commercialize our product candidates in the United States and generally to seek collaboration arrangements with pharmaceutical or biotechnology companies for the development or commercialization of our product candidates in the rest of the world. Accordingly, we may enter into collaboration arrangements in the future on a selective basis. Any future collaboration arrangements that we enter into may not be successful. The success of our collaboration arrangements will depend heavily on the efforts and activities of our collaborators. Collaborators generally have significant discretion in determining the efforts and resources that they will apply to these collaboration arrangements.

Disagreements between parties to a collaboration arrangement regarding clinical development and commercialization matters can lead to delays in the development process or commercializing the applicable product candidate and, in some cases, termination of the collaboration arrangement. These disagreements can be difficult to resolve if neither of the parties has final decision making authority.

Collaborations with pharmaceutical companies and other third parties often are terminated or allowed to expire by the other party. Any such termination or expiration would adversely affect us financially and could harm our business reputation.

Regulatory Risks

We may not receive regulatory approval for any of our product candidates, or the approval may be delayed for various reasons, including successful challenges to the FDA's interpretation of Section 505(b)(2), which would have a material adverse effect on our business and financial condition.

We may experience delays in our efforts to obtain regulatory approval from the FDA for any of our product candidates, and there can be no assurance that such approval will not be delayed, or that the FDA will ultimately approve these product candidates.

The FDA, as a condition of the EXPAREL approval on October 28, 2011, has required us to study EXPAREL in pediatric patients. We have agreed to a trial timeline where, over several years, we will study pediatric patient populations in descending order starting with 12 - 18 year olds and ending with children under two years of age. These trials will be expensive and time consuming and we will be required to meet the timelines for completion as agreed with the FDA.

The FDA may determine that EXPAREL or any of our product candidates have undesirable side effects.

If concerns are raised regarding the safety of a new drug as a result of undesirable side effects identified during clinical testing, the FDA may decline to approve the drug at the end of the NDA review period or issue a letter requesting additional data or information prior to making a final decision regarding whether or not to approve the drug. The number of such requests for additional data or information issued by the FDA in recent years has increased, and resulted in substantial delays in the approval of several new drugs. Undesirable side effects caused by EXPAREL or any product candidate could also result in the inclusion of unfavorable information in our product labeling,

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imposition of distribution or use restrictions, a requirement to conduct post-market studies, denial of regulatory approval by the FDA or other regulatory authorities for any or all targeted indications, and in turn prevent us from commercializing and generating revenues from the sale of EXPAREL or any product candidate.

For example, the side effects observed in the EXPAREL clinical trials completed to date include nausea and vomiting. In addition, the class of drugs that EXPAREL belongs to has been associated with nervous system and cardiovascular toxicities at high doses. We cannot be certain that these side effects and others will not be observed in the future, or that the FDA will not require additional trials or impose more severe labeling restrictions due to these side effects or other concerns. The active component of EXPAREL is bupivacaine and bupivacaine infusions have been associated with the destruction of articular cartilage, or chondrolysis. Chondrolysis has not been observed in clinical trials of EXPAREL, but we cannot be certain that this side effect will not be observed in the future.

Following approval of EXPAREL or any of our product candidates, if we or others later identify undesirable side effects caused by such products:

regulatory authorities may require the addition of unfavorable labeling statements, specific warnings or a contraindication;

regulatory authorities may suspend or withdraw their approval of the product, or require it to be removed from the market;

regulatory authorities may impose restrictions on the distribution or use of the product;

we may be required to change the way the product is administered, conduct additional clinical trials or change the labeling of the product;

we may be subject to product liability claims and litigation; and

our reputation may suffer.

Any of these events could prevent us from achieving or maintaining market acceptance of EXPAREL or any of our product candidates and could substantially increase our commercialization costs and expenses, which in turn could delay or prevent us from generating significant revenues from its sale.

Regulatory approval for any approved product is limited by the FDA to those specific indications and conditions for which clinical safety and efficacy have been demonstrated.

Any regulatory approval is limited to those specific diseases and indications for which a product is deemed to be safe and effective by the FDA. For example, the FDA-approved label for EXPAREL does not include an indication in obstetrical paracervical block anesthesia. In addition to the FDA approval required for new formulations, any new indication for an approved product also requires FDA approval. If we are not able to obtain FDA approval for any desired future indications for our products and product candidates, our ability to effectively market and sell our products may be reduced and our business may be adversely affected.

While physicians may choose to prescribe drugs for uses that are not described in the product's labeling and for uses that differ from those tested in clinical studies and approved by the regulatory authorities, our ability to promote the products is limited to those indications that are specifically approved by the FDA. These "off-label" uses are common across medical specialties and may constitute an appropriate treatment for some patients in varied circumstances. Regulatory authorities in the United States generally do not regulate the behavior of physicians in their choice of treatments. Regulatory authorities do, however, restrict communications by pharmaceutical companies on the subject of off-label use. If our promotional activities fail to comply with these regulations or guidelines, we may be subject to warnings from, or enforcement action by, these authorities. In addition, our

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failure to follow FDA rules and guidelines relating to promotion and advertising may cause the FDA to issue warning letters or untitled letters, suspend or withdraw an approved product from the market, require a recall or institute fines or civil fines, or could result in disgorgement of money, operating restrictions, injunctions or criminal prosecution, any of which could harm our business.

EXPAREL and any other products we may market, including DepoCyt(e) and DepoDur, will remain subject to substantial regulatory scrutiny.

EXPAREL, DepoCyt(e) and DepoDur and any product candidates that we may develop, license or acquire will also be subject to ongoing FDA requirements with respect to the manufacturing, labeling, packaging, storage, distribution, advertising, promotion, record-keeping and submission of safety and other post-market information on the drug. In addition, the subsequent discovery of previously unknown problems with a product may result in restrictions on the product, including withdrawal of the product from the market.

If EXPAREL, DepoCyt(e) and DepoDur or any other product that we may develop, license or acquire fails to comply with applicable regulatory requirements, such as cGMP regulations, a regulatory agency may:

issue warning letters or untitled letters;
require us to enter into a consent decree, which can include imposition of various fines, reimbursements for inspection costs, required due dates for specific actions and penalties for noncompliance;
impose fines and other civil or criminal penalties;
suspend regulatory approval;
suspend any ongoing clinical trials;
refuse to approve pending applications or supplements to approved applications filed by us;
impose restrictions on operations, including costly new manufacturing requirements; or
seize or detain products or require a product recall.

For example, the FDA informed us that certain adverse event reports related to DepoCyt(e) and DepoDur submitted to us during the previous two years were not submitted by us to the FDA within the required 15-day timeframe for reporting such events. In response to the FDA's observations, we enhanced our reporting procedures and hired additional personnel to support our pharmacovigilance efforts.

If we fail to comply with federal and state healthcare laws, including fraud and abuse and health information privacy and security laws, we could face substantial penalties and our business, results of operations, financial condition and prospects could be adversely affected.

As a pharmaceutical company, even though we do not provide healthcare services or receive payments directly from or bill directly to Medicare, Medicaid or other third-party payers for our products, certain federal and state healthcare laws and regulations pertaining to fraud and abuse and patients' rights are and will be applicable to our business. We would be subject to healthcare fraud and abuse and patient privacy regulation by both the federal government and the states in which we conduct our business. The laws that may affect our ability to operate include:

the federal Anti-Kickback Statute, which constrains our marketing practices, educational programs, pricing policies, and relationships with healthcare providers or other entities, by prohibiting, among other things, soliciting, receiving, offering or paying remuneration, directly or indirectly, to induce, or in return for, the purchase or recommendation of an item or service

reimbursable under a federal healthcare program, such as the Medicare and Medicaid programs;

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federal civil and criminal false claims laws and civil monetary penalty laws, which prohibit, among other things, individuals or entities from knowingly presenting, or causing to be presented, claims for payment from Medicare, Medicaid, or other third-party payers that are false or fraudulent;

the federal Health Insurance Portability and Accountability Act of 1996, or HIPAA, which created new federal criminal statutes that prohibit executing a scheme to defraud any healthcare benefit program and making false statements relating to healthcare matters;

federal physician self-referral laws, such as the Stark law, which prohibit a physician from making a referral to a provider of certain health services with which the physician or the physician's family member has a financial interest, and prohibit submission of a claim for reimbursement pursuant to a prohibited referral;

HIPAA, as amended by the Health Information Technology and Clinical Health Act, or HITECH, and its implementing regulations, which imposes certain requirements relating to the privacy, security and transmission of individually identifiable health information; and

state law equivalents of each of the above federal laws, such as anti-kickback and false claims laws which may apply to items or services reimbursed by any third-party payer, including commercial insurers, and state laws governing the privacy and security of health information in certain circumstances, many of which differ from each other in significant ways and may not have the same effect, thus complicating compliance efforts.

Because of the breadth of these laws and the narrowness of the statutory exceptions and safe harbors available under the U.S. federal Anti-Kickback Statute, it is possible that some of our business activities could be subject to challenge under one or more of such laws. Recently, several pharmaceutical and other healthcare companies have been prosecuted under the federal false claims laws for allegedly inflating drug prices they report to pricing services, which in turn are used by the government to set Medicare and Medicaid reimbursement rates, and for allegedly providing free product to customers with the expectation that the customers would bill federal programs for the product. In addition, certain marketing practices, including off-label promotion, may also violate false claims laws. To the extent that any product we make is sold in a foreign country, we may be subject to similar foreign laws and regulations.

Further, there has been a recent trend in the increase of federal and state laws and regulations regarding consulting arrangements with physicians. Some states, such as California, Massachusetts and Vermont, mandate that we comply with a state code of conduct, disclose marketing payments made to physicians, and report compliance information to the state authorities. Some states, such as Massachusetts, have created an internet database to provide disclosed information on certain transactions with physicians to the public. The shifting compliance environment and the need to build and maintain robust and expandable systems to comply in multiple jurisdictions with different compliance and reporting requirements increases the possibility that a pharmaceutical company may run afoul of one or more of the requirements.

If our past or present operations, or those of our distributors are found to be in violation of any of the laws described above or any other governmental regulations that apply to us, we may be subject to penalties, including civil and criminal penalties, damages, fines, exclusion from participation in U.S. federal or state health care programs, and the curtailment or restructuring of our operations. Similarly, if the healthcare providers, distributors or other entities with whom we do business are found to be out of compliance with applicable laws and regulations, they may be subject to sanctions, which could also have a negative impact on us. The risk of being found to have violated such laws is increased by the fact that many of them have not been fully interpreted by the regulatory authorities or the courts, and their provisions are open to a variety of interpretations. Any penalties, damages, fines, curtailment or

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restructuring of our operations could materially adversely affect our ability to operate our business and our financial results. Although compliance programs can mitigate the risk of investigation and prosecution for violations of these laws, the risks cannot be entirely eliminated. Any action against us for violation of these laws, even if we successfully defend against it, could cause us to incur significant legal expenses and divert our management's attention from the operation of our business. Moreover, achieving and sustaining compliance with applicable federal and state privacy, security and fraud laws may prove costly.

The design, development, manufacture, supply, and distribution of EXPAREL, DepoCyt(e) and DepoDur is highly regulated and technically complex.

The design, development, manufacture, supply, and distribution of our products EXPAREL, DepoCyt(e) and DepoDur is technically complex and highly regulated. We, along with our third-party providers, must comply with all applicable regulatory requirements of the FDA and foreign authorities. In addition, the facilities used to manufacture, store, and distribute our products are subject to inspection by regulatory authorities at any time to determine compliance with applicable regulations.

The manufacturing techniques and facilities used for the manufacture and supply of our products must be operated in conformity with cGMP. In complying with cGMP requirements, we, along with our suppliers, must continually expend time, money and effort in production, record keeping, and quality assurance and control to ensure that our products meet applicable specifications and other requirements for safety, efficacy and quality. In addition, we, along with our suppliers, are subject to unannounced inspections by the FDA and other regulatory authorities.

Any failure to comply with regulatory and other legal requirements applicable to the manufacture, supply and distribution of our products could lead to remedial action (such as recalls), civil and criminal penalties and delays in manufacture, supply and distribution of our products. For instance, in connection with routine inspections of one of our manufacturing facilities in April and May 2008, the FDA issued a Form 483 Notice of Inspectional Observations identifying certain deficiencies with respect to our laboratory control system for Depocyt(e). As a result, we did not release new lots of Depocyt(e) for a limited time period as we validated a new assay. We also submitted the new assay to the FDA in July 2008 and in August 2008 we began releasing new lots of DepoCyt(e).

If we fail to comply with the extensive regulatory requirements to which we and our products, EXPAREL, DepoCyt(e) and DepoDur, are subject, such products could be subject to restrictions or withdrawal from the market and we could be subject to penalties.

The testing, manufacturing, labeling, safety, advertising, promotion, storage, sales, distribution, export and marketing, among other things, of our products EXPAREL, DepoCyt(e) and DepoDur are subject to extensive regulation by governmental authorities in the United States and elsewhere throughout the world. Quality control and manufacturing procedures regarding EXPAREL, DepoCyt(e) and DepoDur must conform to cGMP. Regulatory authorities, including the FDA, periodically inspect manufacturing facilities to assess compliance with cGMP. Our failure or the failure of our contract manufacturers to comply with the laws administered by the FDA or other governmental authorities could result in, among other things, any of the following:

product recall or seizure;
suspension or withdrawal of an approved product from the market;
interruption of production;
operating restrictions;
warning letters;

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inju	unctions;
fine	es and other monetary penalties;
crin	ninal prosecutions; and
una	nticipated expenditures.

If the government or third-party payers fail to provide coverage and adequate coverage and payment rates for EXPAREL, DepoCyt(e), DepoDur or any future products we may develop, license or acquire, if any, or if hospitals choose to use therapies that are less expensive, our revenue and prospects for profitability will be limited.

In both domestic and foreign markets, sales of our existing products and any future products will depend in part upon the availability of coverage and reimbursement from third-party payers. Such third-party payers include government health programs such as Medicare and Medicaid, managed care providers, private health insurers and other organizations. Coverage decisions may depend upon clinical and economic standards that disfavor new drug products when more established or lower cost therapeutic alternatives are already available or subsequently become available. Assuming coverage is approved, the resulting reimbursement payment rates might not be adequate. In particular, many U.S. hospitals receive a fixed reimbursement amount per procedure for certain surgeries and other treatment therapies they perform. Because this amount may not be based on the actual expenses the hospital incurs, hospitals may choose to use therapies which are less expensive when compared to our product candidates. Accordingly, EXPAREL, DepoCyt(e), DepoDur or any product candidates that we may develop, in-license or acquire, if approved, will face competition from other therapies and drugs for these limited hospital financial resources. We may need to conduct post-marketing studies in order to demonstrate the cost-effectiveness of any future products to the satisfaction of hospitals, other target customers and their third-party payers. Such studies might require us to commit a significant amount of management time and financial and other resources. Our future products might not ultimately be considered cost-effective. Adequate third-party coverage and reimbursement might not be available to enable us to maintain price levels sufficient to realize an appropriate return on investment in product development.

Third party payers, whether foreign or domestic, or governmental or commercial, are developing increasingly sophisticated methods of controlling healthcare costs. In addition, in the United States, no uniform policy of coverage and reimbursement for drug products exists among third-party payers. Therefore, coverage and reimbursement for drug products can differ significantly from payer to payer.

Further, we believe that future coverage and reimbursement will likely be subject to increased restrictions both in the United States and in international markets. Third-party coverage and reimbursement for our products or product candidates for which we receive regulatory approval may not be available or adequate in either the United States or international markets, which could have a negative effect on our business, results of operations, financial condition and prospects.

We are subject to new legislation, regulatory proposals and healthcare payer initiatives that may increase our costs of compliance and adversely affect our ability to market our products, obtain collaborators and raise capital.

In March 2010, the President signed the Patient Protection and Affordable Care Act, as amended by the Health Care and Education Affordability Reconciliation Act, which we refer to collectively as the Health Care Reform Law. The Health Care Reform Law makes extensive changes to the delivery

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of health care in the United States. Among the provisions of the Health Care Reform Law of greatest importance to the pharmaceutical industry are the following:

an annual, nondeductible fee on any entity that manufactures or imports certain branded prescription drugs and biologic agents, apportioned among these entities according to their market share in certain government healthcare programs, beginning in 2011;

an increase in the statutory minimum rebates a manufacturer must pay under the Medicaid Drug Rebate Program, retroactive to January 1, 2010, to 23.1% and 13% of the average manufacturer price for most branded and generic drugs, respectively;

a new Medicare Part D coverage gap discount program, in which manufacturers must agree to offer 50% point-of-sale discounts off negotiated prices of applicable brand drugs to eligible beneficiaries during their coverage gap period, as a condition for the manufacturer's outpatient drugs to be covered under Medicare Part D, beginning in 2011;

extension of manufacturers' Medicaid rebate liability to covered drugs dispensed to individuals who are enrolled in Medicaid managed care organizations, effective March 23, 2010;

expansion of eligibility criteria for Medicaid programs by, among other things, allowing states to offer Medicaid coverage to additional individuals beginning in April 2010 and by adding new mandatory eligibility categories for certain individuals with income at or below 133% of the Federal Poverty Level beginning in 2014, thereby potentially increasing both the volume of sales and manufacturers' Medicaid rebate liability;

expansion of the entities eligible for discounts under the Public Health Service pharmaceutical pricing program, effective in January 2010;

new requirements to report certain financial arrangements with physicians and others, including reporting any "transfer of value" made or distributed to prescribers and other healthcare providers and reporting any investment interests held by physicians and their immediate family members during each calendar year beginning in 2012, with reporting starting in 2013;

a new requirement to annually report drug samples that manufacturers and distributors provide to physicians, effective April 1, 2012;

a licensure framework for follow-on biologic products;

a new Patient-Centered Outcomes Research Institute to oversee, identify priorities in, and conduct comparative clinical effectiveness research, along with funding for such research;

creation of the Independent Payment Advisory Board which, beginning in 2014, will have authority to recommend certain changes to the Medicare program that could result in reduced payments for prescription drugs and those recommendations could have the effect of law even if Congress does not act on the recommendations; and

establishment of a Center for Medicare Innovation at the Centers for Medicare & Medicaid Services to test innovative payment and service delivery models to lower Medicare and Medicaid spending, potentially including prescription drug spending, beginning by January 1, 2011.

These measures could result in decreased net revenues from our pharmaceutical products and decrease potential returns from our development efforts. A number of states have challenged the constitutionality of certain provisions of the Health Care Reform Law, and many of these court challenges are still pending final adjudication. Congress has also proposed a number of legislative initiatives, including possible repeal of the Health Care Reform Law. At this time, it remains unclear whether there will be any changes made to the Health Care Reform Law, whether to certain provisions or its entirety. In addition, some details regarding the implementation of the Health Care Reform Law

are yet to be determined, and at this time, the full effect that the Health Care Reform Law would have on our business remains unclear.

In addition, other legislative changes have been proposed and adopted since the Health Care Reform Law was enacted. Most recently, on August 2, 2011, the President signed into law the Budget Control Act of 2011, which, among other things, creates the Joint Select Committee on Deficit Reduction to recommend proposals in spending reductions to Congress. The Joint Select Committee may consider all elements of discretionary and non-discretionary spending, and its recommendations could result in reduced spending under Medicare and Medicaid for prescription drugs. In the event that the Joint Select Committee is unable to achieve a targeted deficit reduction of at least \$1.2 trillion for the years 2013 through 2021, or Congress does not act on the committee's recommendation, without amendment, by December 23, 2011, an automatic reduction is triggered. These automatic cuts would be made to several government programs and, with respect to Medicare, would include aggregate reductions to Medicare payments to providers of up to 2% per fiscal year, starting in 2013. The full impact on our business of the new law is uncertain. Nor is it clear whether other legislative changes will be adopted, if any, or how such changes would affect the demand for our products.

In addition, there have been a number of other legislative and regulatory proposals aimed at changing the pharmaceutical industry. In particular, California has enacted legislation that requires development of an electronic pedigree to track and trace each prescription drug at the saleable unit level through the distribution system. California's electronic pedigree requirement is scheduled to take effect in January 2015. Compliance with California and future federal or state electronic pedigree requirements may increase our operational expenses and impose significant administrative burdens. As a result of these and other new proposals, we may determine to change our current manner of operation, provide additional benefits or change our contract arrangements, any of which could have a material adverse effect on our business, financial condition and results of operations.

Public concern regarding the safety of drug products such as EXPAREL could result in the inclusion of unfavorable information in our labeling, or require us to undertake other activities that may entail additional costs.

In light of widely publicized events concerning the safety risk of certain drug products, the FDA, members of Congress, the Government Accountability Office, medical professionals and the general public have raised concerns about potential drug safety issues. These events have resulted in the withdrawal of drug products, revisions to drug labeling that further limit use of the drug products and the establishment of risk management programs that may, for example, restrict distribution of drug products after approval. The Food and Drug Administration Amendments Act of 2007, or FDAAA, grants significant expanded authority to the FDA, much of which is aimed at improving the safety of drug products before and after approval. In particular, the FDAAA authorizes the FDA to, among other things, require post-approval studies and clinical trials, mandate changes to drug labeling to reflect new safety information and require risk evaluation and mitigation strategies for certain drugs, including certain currently approved drugs. It also significantly expands the federal government's clinical trial registry and results databank, which we expect will result in significantly increased government oversight of clinical trials. Under the FDAAA, companies that violate these and other provisions of the new law are subject to substantial civil monetary penalties, among other regulatory, civil and criminal penalties. The increased attention to drug safety issues may result in a more cautious approach by the FDA in its review of data from our clinical trials. Data from clinical trials may receive greater scrutiny, particularly with respect to safety, which may make the FDA or other regulatory authorities more likely to require additional preclinical studies or clinical trials. If the FDA requires us to provide additional clinical or preclinical data for EXPAREL, the indications for which this product candidate was approved may be limited or there may be specific warnings or limitations on dosing, and our efforts to commercialize EXPA

Our product, DepoDur, is subject to regulation by the Drug Enforcement Agency and such regulation may affect the sale of DepoDur.

Products used to treat and manage pain, especially in the case of opioids, are from time to time subject to negative publicity, including illegal use, overdoses, abuse, diversion, serious injury and death. These events have led to heightened regulatory scrutiny. Controlled substances are classified by the DEA as Schedule I through V substances, with Schedule I substances being prohibited for sale in the United States, Schedule II substances considered to present the highest risk of abuse and Schedule V substances being considered to present the lowest relative risk of abuse. DepoDur contains morphine, and it is regulated as a Schedule II controlled substance. Despite the strict regulations on the marketing, prescribing and dispensing of such substances, illicit use and abuse of morphine does occur. Thus, the marketing of DepoDur by our partners may generate public controversy that may adversely affect sales of DepoDur and decrease the revenue we receive from the sale of DepoDur.

In addition, we and our contract manufacturers are subject to ongoing DEA regulatory obligations, including, among other things, annual registration renewal, security, recordkeeping, theft and loss reporting, periodic inspection and annual quota allotments for the raw material for commercial production of our products. The DEA, and some states, conduct periodic inspections of registered establishments that handle controlled substances. Facilities that conduct research, manufacture, store, distribute, import or export controlled substances must be registered to perform these activities and have the security, control and inventory mechanisms required by the DEA to prevent drug loss and diversion. Failure to maintain compliance, particularly non-compliance resulting in loss or diversion, can result in regulatory action that could have a material adverse effect on our business, results of operations, financial condition and prospects. The DEA may seek civil penalties, refuse to renew necessary registrations, or initiate proceedings to revoke those registrations. In certain circumstances, violations could lead to criminal proceedings.

Individual states also have controlled substances laws. Though state controlled substances laws often mirror federal law, because the states are separate jurisdictions, they may separately schedule drugs, as well. While some states automatically schedule a drug when the DEA does so, in other states there has to be rulemaking or a legislative action. State scheduling may delay commercial sale of any controlled substance drug product for which we obtain federal regulatory approval and adverse scheduling could have a material adverse effect on the commercial attractiveness of such product. We or our partners must also obtain separate state registrations in order to be able to obtain, handle, and distribute controlled substances for clinical trials or commercial sale, and failure to meet applicable regulatory requirements could lead to enforcement and sanctions from the states in addition to those from the DEA or otherwise arising under federal law.

Risks Related to Intellectual Property

The patents and the patent applications that we have covering our products are limited to specific injectable formulations, processes and uses of drugs encapsulated in our DepoFoam drug delivery technology and our market opportunity for our product candidates may be limited by the lack of patent protection for the active ingredient itself and other formulations and delivery technology and systems that may be developed by competitors.

The active ingredients in EXPAREL, DepoCyt(e) and DepoDur are bupivacaine, cytarabine and morphine, respectively. Patent protection for the bupivacaine, cytarabine and morphine molecules themselves has expired and generic immediate-release products are available. As a result, competitors who obtain the requisite regulatory approval can offer products with the same active ingredients as EXPAREL, DepoCyt(e) and DepoDur so long as the competitors do not infringe any process, use or formulation patents that we have developed for these drugs encapsulated in our DepoFoam drug delivery technology.

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For example, we are aware of at least one long acting injectable bupivacaine product in development which utilizes an alternative delivery system to EXPAREL. Such a product is similar to EXPAREL in that it also extends the duration of effect of bupivacaine, but achieves this clinical outcome using a completely different drug delivery system compared to our DepoFoam drug delivery technology.

The number of patents and patent applications covering products in the same field as EXPAREL indicates that competitors have sought to develop and may seek to market competing formulations that may not be covered by our patents and patent applications. The commercial opportunity for EXPAREL could be significantly harmed if competitors are able to develop and commercialize alternative formulations of bupivacaine that are long acting but outside the scope of our patents.

Now that EXPAREL is approved by the FDA, one or more third parties may challenge the patents covering this product, which could result in the invalidation or unenforceability of some or all of the relevant patent claims. For example, if a third party files an Abbreviated New Drug Application, or ANDA, for a generic drug product containing bupivacaine and relies in whole or in part on studies conducted by or for us, the third party will be required to certify to the FDA that either: (1) there is no patent information listed in the FDA's Orange Book with respect to our NDA for EXPAREL; (2) the patents listed in the Orange Book have expired; (3) the listed patents have not expired, but will expire on a particular date and approval is sought after patent expiration; or (4) the listed patents are invalid or will not be infringed by the manufacture, use or sale of the third-party's generic drug product. A certification that the new product will not infringe the Orange Book-listed patents for EXPAREL, or that such patents are invalid, is called a paragraph IV certification. If the third party submits a paragraph IV certification to the FDA, a notice of the paragraph IV certification must also be sent to us once the third-party's ANDA is accepted for filing by the FDA. We may then initiate a lawsuit to defend the patents identified in the notice. The filing of a patent infringement lawsuit within 45 days of receipt of the notice automatically prevents the FDA from approving the third-party's ANDA until the earliest of 30 months or the date on which the patent expires, the lawsuit is settled, or the court reaches a decision in the infringement lawsuit in favor of the third party. If we do not file a patent infringement lawsuit within the required 45-day period, the third-party's ANDA will not be subject to the 30-month stay. Litigation or other proceedings to enforce or defend intellectual property rights are often very complex in nature, may be very expensive and time-consuming, may divert our management's attention from our core business, and may result in unfavorable results that could adversely impact our ability to prevent third parties from competing with our products.

Because it is difficult and costly to protect our proprietary rights, we may not be able to ensure their protection and all patents will eventually expire.

Our commercial success will depend in part on obtaining and maintaining patent protection and trade secret protection for EXPAREL, DepoCyt(e), DepoDur, DepoFoam and for any product candidates that we may develop, license or acquire and the methods we use to manufacture them, as well as successfully defending these patents and trade secrets against third-party challenges. We will only be able to protect our technologies from unauthorized use by third parties to the extent that valid and enforceable patents or trade secrets cover them.

The patent positions of pharmaceutical and biotechnology companies can be highly uncertain and involve complex legal and factual questions for which important legal principles remain unresolved. No consistent policy regarding the breadth of claims allowed in pharmaceutical or biotechnology patents has emerged to date in the United States. The patent situation outside the United States is even more uncertain. Changes in either the patent laws or in interpretations of patent laws in the United States and other countries may diminish the value of our intellectual property. Accordingly, we cannot predict the breadth of claims that may be allowed or enforced in our patents or in third-party patents.

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The degree of future protection for our proprietary rights is uncertain, because legal means afford only limited protection and may not adequately protect our rights or permit us to gain or keep our competitive advantage. For example:

we may not have been the first to make the inventions covered by each of our pending patent applications and issued patents;

we may not have been the first to file patent applications for these inventions;

others may independently develop similar or alternative technologies or duplicate any of our product candidates or technologies;

it is possible that none of the pending patent applications will result in issued patents;

the issued patents covering our product candidates may not provide a basis for commercially viable active products, may not provide us with any competitive advantages, or may be challenged by third parties;

we may not develop additional proprietary technologies that are patentable; or

patents of others may have an adverse effect on our business.

Patent applications in the United States are maintained in confidence for at least 18 months after their earliest effective filing date. Consequently, we cannot be certain we were the first to invent or the first to file patent applications on EXPAREL, our DepoFoam drug delivery technology or any product candidates that we may develop, license or acquire. In the event that a third party has also filed a U.S. patent application relating to our product candidates or a similar invention, we may have to participate in interference proceedings declared by the U.S. Patent and Trademark Office to determine priority of invention in the United States. The costs of these proceedings could be substantial and it is possible that our efforts would be unsuccessful, resulting in a material adverse effect on our U.S. patent position. Furthermore, we may not have identified all United States and foreign patents or published applications that affect our business either by blocking our ability to commercialize our drugs or by covering similar technologies that affect our drug market.

In addition, some countries, including many in Europe, do not grant patent claims directed to methods of treating humans, and in these countries patent protection may not be available at all to protect our product candidates. Even if patents issue, we cannot guarantee that the claims of those patents will be valid and enforceable or provide us with any significant protection against competitive products, or otherwise be commercially valuable to us.

Some of our older patents have already expired. In the cases of DepoCyt(e) and DepoDur, key patents providing protection in Europe have expired. In the case of EXPAREL, while pending patent applications, if granted, would provide protection for EXPAREL in Europe and the United States through November 2018, an existing formulation patent for EXPAREL will expire in November 2013. Once our patents covering EXPAREL have expired, we are more reliant on trade secrets to protect against generic competition.

We also rely on trade secrets to protect our technology, particularly where we do not believe patent protection is appropriate or obtainable. However, trade secrets are difficult to protect. While we use reasonable efforts to protect our trade secrets, our licensors, employees, consultants, contractors, outside scientific collaborators and other advisors may unintentionally or willfully disclose our information to competitors. Enforcing a claim that a third party illegally obtained and is using our trade secrets is expensive and time consuming, and the outcome is unpredictable. In addition, courts outside the United States are sometimes less willing to protect trade secrets. Moreover, our competitors may independently develop equivalent knowledge, methods and know-how.

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If we fail to obtain or maintain patent protection or trade secret protection for EXPAREL, DepoCyt(e), DepoDur, DepoFoam or any product candidate that we may develop, license or acquire, third parties could use our proprietary information, which could impair our ability to compete in the market and adversely affect our ability to generate revenues and achieve profitability.

If we are sued for infringing intellectual property rights of third parties, it will be costly and time consuming, and an unfavorable outcome in any litigation would harm our business.

Our ability to develop, manufacture, market and sell EXPAREL, our DepoFoam drug delivery technology or any product candidates that we may develop, license or acquire depends upon our ability to avoid infringing the proprietary rights of third parties. Numerous U.S. and foreign issued patents and pending patent applications, which are owned by third parties, exist in the general fields of pain management and cancer treatment and cover the use of numerous compounds and formulations in our targeted markets. Because of the uncertainty inherent in any patent or other litigation involving proprietary rights, we and our licensors may not be successful in defending intellectual property claims by third parties, which could have a material adverse affect on our results of operations. Regardless of the outcome of any litigation, defending the litigation may be expensive, time-consuming and distracting to management. In addition, because patent applications can take many years to issue, there may be currently pending applications, unknown to us, which may later result in issued patents that EXPAREL, DepoCyt(e) or DepoDur may infringe. There could also be existing patents of which we are not aware that EXPAREL, DepoCyt(e) or DepoDur may inadvertently infringe.

There is a substantial amount of litigation involving patent and other intellectual property rights in the biotechnology and biopharmaceutical industries generally. If a third party claims that we infringe on their products or technology, we could face a number of issues, including:

infringement and other intellectual property claims which, with or without merit, can be expensive and time consuming to litigate and can divert management's attention from our core business;

substantial damages for past infringement which we may have to pay if a court decides that our product infringes on a competitor's patent;

a court prohibiting us from selling or licensing our product unless the patent holder licenses the patent to us, which it would not be required to do;

if a license is available from a patent holder, we may have to pay substantial royalties or grant cross licenses to our patents; and

redesigning our processes so they do not infringe, which may not be possible or could require substantial funds and time.

We may be subject to claims that our employees have wrongfully used or disclosed alleged trade secrets of their former employers.

As is common in the biotechnology and pharmaceutical industry, we employ individuals who were previously employed at other biotechnology or pharmaceutical companies, including our competitors or potential competitors. Although no claims against us are currently pending, we may be subject to claims that these employees or we have inadvertently or otherwise used or disclosed trade secrets or other proprietary information of their former employers. Litigation may be necessary to defend against these claims. Even if we are successful in defending against these claims, litigation could result in substantial costs and be a distraction to management.

Risks Related to Our Financial Condition and Capital Requirements

We believe certain matters raise substantial doubt about our ability to continue as a going concern, which may hinder our ability to obtain future financing.

As of September 30, 2011, we believe certain matters raise substantial doubt about our ability to continue as a going concern. Such doubts are based on our recurring losses and our cash used in operating activities. We continue to experience losses. Our ability to continue as a going concern is subject to our ability to generate a profit and/or obtain necessary funding from outside sources, including by the sale of our securities, obtaining loans from financial institutions or other financing arrangements, where possible. Our continued losses increase the difficulty of our meeting such goals and our efforts to continue as a going concern may not prove successful.

We have incurred significant losses since our inception and anticipate that we will incur continued losses for the foreseeable future.

We are an emerging specialty pharmaceutical company with a limited operating history. We have focused primarily on developing and commercializing EXPAREL. We have incurred losses in each year since our inception in December 2006, including net losses of \$27.1 million, \$31.7 million, and \$41.9 million for the years ended December 31, 2010, 2009, and 2008, respectively. As of September 30, 2011, we had an accumulated deficit of \$165.0 million. These losses, among other things, have had, and will continue to have, an adverse effect on our stockholders' equity and working capital. We incurred increased pre-commercialization expenses during 2010 and 2011 as we prepared for the potential commercial launch of EXPAREL, and we expect to incur significant sales, marketing and manufacturing expenses, as well as continued development expenses related to the commercialization of EXPAREL. As a result, we expect to continue to incur significant losses for the foreseeable future. Because of the numerous risks and uncertainties associated with developing pharmaceutical products, we are unable to predict the extent of any future losses or when we will become profitable, if at all.

We may never become profitable.

Our ability to become profitable depends upon our ability to generate revenue from EXPAREL and to continue to generate revenue from DepoCyt(e) and DepoDur. Our ability to generate revenue depends on a number of factors, including, but not limited to, our ability to:

manufacture commercial quantities of EXPAREL, at acceptable cost levels;

continue to manufacture DepoCyt(e) and DepoDur for sale by our commercial partners; and

continue to develop a commercial organization and the supporting infrastructure required to successfully market and sell EXPAREL.

We anticipate incurring significant additional costs associated with the commercialization of EXPAREL. We also do not anticipate that we will achieve profitability for a period of time after generating material revenues, if ever. If we are unable to generate revenues, we will not become profitable and may be unable to continue operations without continued funding.

Under our financing arrangement with Paul Capital, upon the occurrence of certain events, Paul Capital may require us to repurchase the right to receive royalty payments that we assigned to it, or may foreclose on certain assets that secure our obligations to Paul Capital. Any exercise by Paul Capital of its right to cause us to repurchase the assigned right or any foreclosure by Paul Capital would adversely affect our results of operations and our financial condition.

On March 23, 2007, we entered into the Amended and Restated Royalty Interests Assignment Agreement with affiliates of Paul Capital, pursuant to which we assigned to Paul Capital the right to

receive a portion of our royalty payments from DepoCyt(e) and DepoDur. To secure our obligations to Paul Capital, we granted Paul Capital a security interest in collateral which includes the royalty payments we are entitled to receive with respect to sales of DepoCyt(e) and DepoDur, as well as to bank accounts to which such payments are deposited. Under our arrangement with Paul Capital, upon the occurrence of certain events, or the put events, including if we experience a change of control, we or our subsidiary undergo certain bankruptcy events, transfer any or substantially all of our rights in DepoCyt(e) or DepoDur, transfer all or substantially all of our assets, breach certain of the covenants, representations or warranties under the Amended and Restated Royalty Interests Assignment Agreement, or sales of DepoCyt(e) or DepoDur are suspended due to an injunction or if we elect to suspend sales of DepoCyt(e) or DepoDur as a result of a lawsuit filed by certain third parties, Paul Capital may (i) require us to repurchase the rights we assigned to it at a cash price equal to (a) 50% of all cumulative payments made by us to Paul Capital under the Amended and Restated Royalty Interests Assignment Agreement during the preceding 24 months, multiplied by (b) the number of days from the date of Paul Capital's exercise of such option until December 31, 2014, divided by 365. Any exercise by Paul Capital of its right to cause us to repurchase the assigned right or any foreclosure by Paul Capital would adversely affect our results of operations and our financial condition.

Our debt obligations expose us to risks that could adversely affect our business, operating results and financial condition.

We have a substantial level of debt. As of September 30, 2011, we had \$26.25 million in aggregate principal amount of indebtedness outstanding, not including our obligation under the Amended and Restated Royalty Interests Assignment Agreement with Paul Capital. The level and nature of our indebtedness, among other things, could:

make it difficult for us to make payments on our outstanding debt from time to time or to refinance it;

make it difficult for us to obtain any necessary financing in the future for working capital, capital expenditures, debt service, product and company acquisitions or general corporate purposes;

limit our flexibility in planning for or reacting to changes in our business including life cycle management;

reduce funds available for use in our operations;

impair our ability to incur additional debt because of financial and other restrictive covenants;

make us more vulnerable in the event of a downturn in our business;

place us at a possible competitive disadvantage relative to less leveraged competitors and competitors that have better access to capital resources;

restrict the operations of our business as a result of provisions in the Amended and Restated Royalty Interests Assignment Agreement with Paul Capital that restrict our ability to (i) amend, waive any rights under, or terminate any material agreements relating to DepoCyt(e) and DepoDur, (ii) enter into any new agreement or amend or fail to exercise any of our material rights under existing agreements that would materially adversely affect Paul Capital's royalty interest, and (iii) sell any material assets related to DepoCyt(e) or DepoDur; or

impair our ability to merge or otherwise affect the sale of the Company due to the right of the holders of certain of our indebtedness to accelerate the maturity date of the indebtedness in the event of a change of control of the Company.

We will need to raise additional capital to pay our indebtedness as it comes due. If we are unable to obtain funds necessary to make required payments, or if we fail to comply with the various

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requirements of our indebtedness, we would be in default, which would permit the holders of our indebtedness to accelerate the maturity of the indebtedness and could cause defaults under any indebtedness we may incur in the future. Any default under our indebtedness would have a material adverse effect on our business, operating results and financial condition. If we are unable to refinance or repay our indebtedness as it becomes due, we may become insolvent and be unable to continue operations.

For example, our loan and security agreement governing our \$26.25 million credit facility with Hercules Technology Growth Capital, Inc. and Hercules Technology III, L.P., as lenders, or the Hercules Credit Facility, contains a number of affirmative and restrictive covenants, including reporting requirements and other collateral limitations, certain limitations on liens and indebtedness, dispositions, mergers and acquisitions, restricted payments and investments, corporate changes and limitations on waivers and amendments to certain agreements, our organizational documents, and documents relating to debt that is subordinate to our obligations under the Hercules Credit Facility. Our failure to comply with the covenants in the loan and security agreement governing the Hercules Credit Facility could result in an event of default that, if not cured or waived, could result in the acceleration of all or a substantial portion of our debt and potential foreclosure on the assets pledged to secure the debt.

Our short operating history makes it difficult to evaluate our business and prospects.

We were incorporated in December 2006 and have only been conducting operations with respect to EXPAREL since March 2007. Our operations to date have been limited to organizing and staffing our company, conducting product development activities, including clinical trials and manufacturing development activities, for EXPAREL and manufacturing and related activities for DepoCyt(e) and DepoDur. Further, in 2010 and 2011 we began to establish our commercial infrastructure for EXPAREL. Consequently, any predictions about our future performance may not be as accurate as they could be if we had a history of successfully developing and commercializing pharmaceutical products.

We will need additional funding and may be unable to raise capital when needed, which would force us to delay, reduce or eliminate our product development programs or commercialization efforts.

Developing products for use in the hospital setting, conducting clinical trials, establishing outsourced manufacturing relationships and successfully manufacturing and marketing drugs that we may develop is expensive. We will need to raise additional capital to:

fund our operations and continue our efforts to hire, and outsource through our relationship with Quintiles, additional personnel and build a commercial infrastructure to prepare for the commercialization of EXPAREL;

qualify and outsource the commercial-scale manufacturing of our products under cGMP; and

in-license and develop additional product candidates.

We may not have sufficient financial resources to meet all of our objectives, which could require us to postpone, scale back or eliminate some, or all, of these objectives, including our launch activities for EXPAREL. Our future funding requirements will depend on many factors, including, but not limited to:

the costs of establishing a commercial organization to sell, market and distribute EXPAREL;

the success of the commercialization of EXPAREL;

the cost and timing of manufacturing sufficient supplies of EXPAREL in preparation for commercialization;

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the rate of progress and costs of our efforts to prepare for the submission of an NDA for any product candidates that we may in-license or acquire in the future, and the potential that we may need to conduct additional clinical trials to support applications for regulatory approval;

the costs of filing, prosecuting, defending and enforcing any patent claims and other intellectual property rights associated with our product candidates, including any such costs we may be required to expend if our licensors are unwilling or unable to do so;

the effect of competing technological and market developments;

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

the potential that we may be required to file a lawsuit to defend our patent rights or regulatory exclusivities from challenges by companies seeking to market generic versions of extended-release liposome injection of bupivacaine.

Future capital requirements will also depend on the extent to which we acquire or invest in additional complementary businesses, products and technologies.

Until we can generate a sufficient amount of product revenue, if ever, we expect to finance future cash needs through public or private equity offerings, debt financings, product supply revenue and royalties, corporate collaboration and licensing arrangements, as well as through interest income earned on cash and investment balances. We cannot be certain that additional funding will be available on acceptable terms, or at all. If adequate funds are not available, we may be required to delay, reduce the scope of, or eliminate, one or more of our development programs or our commercialization efforts.

Our quarterly operating results may fluctuate significantly.

We expect our operating results to be subject to quarterly fluctuations. Our net loss and other operating results will be affected by numerous factors, including:

our ability to establish the necessary commercial infrastructure to launch EXPAREL without substantial delays, including engaging additional sales and marketing personnel and contracting with third parties for warehousing, distribution, cash collection and related commercial activities;

maintaining our existing manufacturing facilities and expanding our manufacturing capacity, including installing specialized processing equipment for the manufacturing of EXPAREL;

our execution of other collaborative, licensing or similar arrangements and the timing of payments we may make or receive under these arrangements;

variations in the level of expenses related to our future development programs;

any product liability or intellectual property infringement lawsuit in which we may become involved;

regulatory developments affecting EXPAREL or the product candidates of our competitors; and

the level of underlying hospital demand for EXPAREL and wholesaler buying patterns.

If our quarterly or annual operating results fall below the expectations of investors or securities analysts, the price of our common stock could decline substantially. Furthermore, any quarterly or annual fluctuations in our operating results may, in turn, cause the price of our stock to fluctuate substantially. We believe that quarterly comparisons of our financial results are not necessarily meaningful and should not be relied upon as an indication of our future performance.

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Raising additional funds by issuing securities may cause dilution to existing stockholders and raising funds through lending and licensing arrangements may restrict our operations or require us to relinquish proprietary rights.

To the extent that we raise additional capital by issuing equity securities, our existing stockholders' ownership will be diluted. If we raise additional funds through licensing arrangements, it may be necessary to relinquish potentially valuable rights to our potential products or proprietary technologies, or grant licenses on terms that are not favorable to us. Any debt financing we enter into may involve covenants that restrict our operations. These restrictive covenants may include limitations on additional borrowing and specific restrictions on the use of our assets as well as prohibitions on our ability to create liens, pay dividends, redeem our stock or make investments.

We incur significant costs as a result of operating as a public company.

As a public company, we incur significant legal, accounting, insurance and other expenses that we did not incur as a private company, including costs associated with public company reporting requirements. We also have incurred and will incur costs associated with complying with the requirements of the Sarbanes-Oxley Act of 2002 and related rules implemented by the Securities and Exchange Commission and The NASDAQ Global Market. The expenses incurred by public companies generally for reporting and corporate governance purposes have been increasing. We expect these rules and regulations to increase our legal and financial compliance costs and to make some activities more time-consuming and costly, although we are currently unable to estimate these costs with any degree of certainty. These laws and regulations could also make it more difficult or costly for us to obtain or maintain certain types of insurance, including director and officer liability insurance, and we may be forced to accept reduced policy limits and coverage or incur substantially higher costs to obtain the same or similar coverage. These laws and regulations could also make it more difficult for us to attract and retain qualified persons to serve on our board of directors, our board committees or as our executive officers. Furthermore, if we are unable to satisfy our obligations as a public company, we could be subject to delisting of our common stock, fines, sanctions and other regulatory action and potentially civil litigation.

Compliance with Section 404 of the Sarbanes-Oxley Act of 2002 requires our management to devote substantial time to compliance initiatives, and if our independent registered public accounting firm is required to provide an attestation report on our internal controls but is unable to provide an unqualified attestation report, our stock price could be adversely affected.

Pursuant to Section 404 of the Sarbanes-Oxley Act of 2002, or Section 404, we are required to furnish a report by our management on the effectiveness of our internal control over financial reporting. The internal control report must contain (i) a statement of management's responsibility for establishing and maintaining adequate internal control over financial reporting, (ii) a statement identifying the framework used by management to conduct the required evaluation of the effectiveness of our internal control over financial reporting and (iii) management's assessment of the effectiveness of our internal control over financial reporting as of the end of our most recent fiscal year, including a statement as to whether or not internal control over financial reporting is effective.

To achieve compliance with Section 404 within the prescribed period, we will be engaged in a process to document and evaluate our internal control over financial reporting, which is both costly and challenging. In this regard, we will need to continue to dedicate internal resources, hire additional employees for our finance and audit functions, potentially engage outside consultants and adopt a detailed work plan to (i) assess and document the adequacy of internal control over financial reporting, (ii) continue steps to improve control processes where appropriate, (iii) validate through testing that controls are functioning as documented, and (iv) implement a continuous reporting and improvement process for internal control over financial reporting. In addition, in connection with the attestation

process by our independent registered public accounting firm, if required, we may encounter problems or delays in completing the implementation of any requested improvements and receiving a favorable attestation. If we cannot favorably assess the effectiveness of our internal control over financial reporting, or if our independent registered public accounting firm is unable to provide an unqualified attestation report on our internal controls, investors could lose confidence in our financial information and our stock price could decline.

The use of our net operating loss carryforwards and research tax credits may be limited.

We have significant federal and state net operating loss carryforwards and federal and state research and development tax credit carryforwards. Our net operating loss carryforwards and research and development tax credits may expire and not be used. Our net operating loss carryforwards will begin expiring in 2026 for federal purposes and 2016 for state purposes if we have not used them prior to that time, and our federal tax credits will begin expiring in 2027 unless previously used. Our state tax credits carryforward indefinitely. Additionally, our ability to use any net operating loss and credit carryforwards to offset taxable income or tax, respectively, in the future will be limited under Internal Revenue Code Sections 382 and 383 if we have a cumulative change in ownership of more than 50% within a three-year period. The completion of this offering, together with our initial public offering, private placements and other transactions that have occurred, may trigger such an ownership change. In addition, since we will need to raise substantial additional funding to finance our operations, we may undergo further ownership changes in the future. In the event such an ownership change occurs, we will be limited regarding the amount of net operating loss carryforwards and research tax credits that could be utilized annually in the future to offset taxable income or tax, respectively. Any such annual limitation may significantly reduce the utilization of the net operating loss carryforwards and research tax credits before they expire. In addition, California and certain states have suspended use of net operating loss carryforwards for certain taxable years, and other states are considering similar measures. As a result, we may incur higher state income tax expense in the future. Depending on our future tax position, continued suspension of our ability to use net operating loss carryforwards in states in which we are subject to income tax could have an adverse impact on our results of operations and financial condi

Our results of operations and liquidity needs could be materially negatively affected by market fluctuations and economic downturns.

Our results of operations could be materially negatively affected by economic conditions generally, both in the United States and elsewhere around the world. Continuing concerns over inflation, energy costs, geopolitical issues, the availability and cost of credit, the U.S. mortgage market and a declining residential real estate market in the United States have contributed to increased volatility and diminished expectations for the economy and the markets going forward. These factors, combined with volatile oil prices, declining business and consumer confidence and increased unemployment, have precipitated an economic recession and fears of a possible depression. Domestic and international equity markets continue to experience heightened volatility and turmoil. These events and the continuing market upheavals may have an adverse effect on us. In the event of a continuing market downturn, our results of operations could be adversely affected by those factors in many ways, including making it more difficult for us to raise funds if necessary, and our stock price may further decline.

Risks Related to this Offering and Ownership of Our Common Stock

The market price of our common stock is highly volatile, and you may not be able to resell your shares at or above the public offering price.

Our stock price is volatile, and from February 3, 2011, the first day of trading of our common stock, to November 4, 2011, the trading prices of our stock have ranged from \$6.16 to \$15.34 per share.

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Our stock could be subject to wide fluctuations in price in response to various factors, including the following:

the commercial success of EXPAREL; results of clinical trials of our product candidates or those of our competitors; changes or developments in laws or regulations applicable to our product candidates; introduction of competitive products or technologies; failure to meet or exceed financial projections we provide to the public; actual or anticipated variations in quarterly operating results; failure to meet or exceed the estimates and projections of the investment community; the perception of the pharmaceutical industry by the public, legislatures, regulators and the investment community; general economic and market conditions and overall fluctuations in U.S. equity markets; developments concerning our sources of manufacturing supply; disputes or other developments relating to patents or other proprietary rights; additions or departures of key scientific or management personnel; issuances of debt, equity or convertible securities; changes in the market valuations of similar companies; and

In addition, the stock market in general, and the market for small pharmaceutical and biotechnology companies in particular, have experienced extreme price and volume fluctuations that have often been unrelated or disproportionate to the operating performance of these companies. Broad market and industry factors may negatively affect the market price of our common stock, regardless of our actual operating performance.

the other factors described in this "Risk Factors" section.

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

Upon completion of this offering, our executive officers, directors and 5% stockholders and their affiliates will beneficially own approximately 52% of our outstanding voting stock. As a result, these stockholders will have significant influence and may be able to determine all matters requiring stockholder approval. For example, these stockholders may be able to control elections of directors, amendments of our organizational documents, or approval of any merger, sale of assets, or other major corporate transaction. This concentration of ownership could delay or prevent any acquisition of our company on terms that other stockholders may desire.

If you purchase our common stock in this offering, you will incur immediate and substantial dilution in the book value of your shares.

You will suffer immediate and substantial dilution in the net tangible book value of the common stock you purchase in this offering. We expect that the public offering price of our common stock in this offering will be substantially higher than the net tangible book value per share of our outstanding common stock immediately after this offering. Purchasers of common stock in this offering will experience immediate dilution of approximately \$5.57 per share in net tangible book value of the common stock. In the past, we issued options and warrants to acquire common stock at prices

significantly below the public offering price. To the extent these outstanding options and warrants are ultimately exercised, investors purchasing common stock in this offering will sustain further dilution. See "Dilution" for a more detailed description of the dilution to new investors in the offering.

Sales of a substantial number of shares of our common stock in the public market by our existing stockholders could cause our stock price to fall.

Sales of a substantial number of shares of our common stock in the public market or the perception that these sales might occur, could depress the market price of our common stock and could impair our ability to raise adequate capital through the sale of additional equity securities. We are unable to predict the effect that sales may have on the prevailing market price of our common stock.

Substantially all of our existing stockholders are subject to lock-up agreements with the underwriters of this offering that restrict the stockholders' ability to transfer shares of our common stock for at least 90 days from the date of this prospectus, subject to certain exceptions. The lock-up agreements limit the number of shares of common stock that may be sold immediately following the public offering. Subject to certain limitations, 8,902,759 shares will become eligible for sale upon expiration of the lock-up period. In addition, shares issued or issuable upon exercise of options and warrants vested as of the expiration of the lock-up period will be eligible for sale at that time. Sales of stock by these stockholders could have a material adverse effect on the market price of our common stock.

Certain holders of shares of our common stock are entitled to rights with respect to the registration of their shares under the Securities Act of 1933, as amended, or the Securities Act, subject to the 90-day lock-up arrangement described above. Registration of these shares under the Securities Act would result in the shares becoming freely tradable without restriction under the Securities Act, except for shares held by our affiliates as defined in Rule 144 under the Securities Act. Any sales of securities by these stockholders could have a material adverse effect on the trading price of our common stock.

Our management will have broad discretion in the use of the net proceeds from this offering and may not use them effectively.

Our management will have broad discretion in the application of the net proceeds from this offering and our stockholders will not have the opportunity as part of their investment decision to assess whether the net proceeds are being used appropriately. Because of the number and variability of factors that will determine our use of the net proceeds from this offering, their ultimate use may vary substantially from their currently intended use. The failure by our management to apply these funds effectively could harm our business. Pending their use, we may invest the net proceeds from this offering in short-term, investment-grade, interest-bearing instruments and U.S. government securities. These investments may not yield a favorable return to our stockholders.

Because we do not intend to pay dividends on our common stock, your returns will be limited to any increase in the value of our stock.

We have never declared or paid any cash dividends on our capital stock. We currently intend to retain all available funds and any future earnings to support our operations and finance the growth and development of our business and do not anticipate declaring or paying any cash dividends on our common stock for the foreseeable future. Any return to stockholders will therefore be limited to the appreciation of their stock, if any.

Some provisions of our charter documents 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.

Provisions in our restated certificate of incorporation and our bylaws, as well as provisions of the Delaware General Corporation Law, or DGCL, could make it more difficult for a third party to acquire us or increase the cost of acquiring us, even if doing so would benefit our stockholders, including transactions in which stockholders might otherwise receive a premium for their shares. These provisions include:

authorizing the issuance of "blank check" preferred stock, the terms of which may be established and shares of which may be issued without stockholder approval;

prohibiting stockholder action by written consent, thereby requiring all stockholder actions to be taken at a meeting of our stockholders;

eliminating the ability of stockholders to call a special meeting of stockholders; and

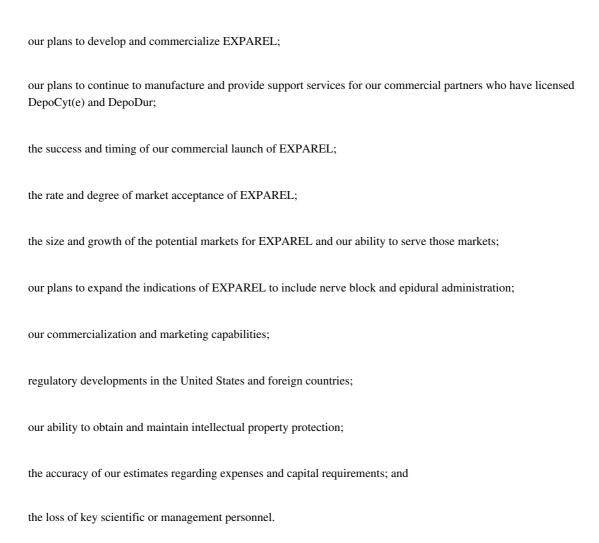
establishing advance notice requirements for nominations for election to the board of directors or for proposing matters that can be acted upon at stockholder meetings.

These provisions may frustrate or prevent any attempts by our stockholders to replace or remove our current management by making it more difficult for stockholders to replace members of our board of directors, which is responsible for appointing the members of our management. In addition, we are subject to Section 203 of the DGCL, which generally prohibits a Delaware corporation from engaging in any of a broad range of business combinations with an interested stockholder for a period of three years following the date on which the stockholder became an interested stockholder, unless such transactions are approved by our board of directors. This provision could have the effect of delaying or preventing a change of control, whether or not it is desired by or beneficial to our stockholders.

CAUTIONARY NOTE REGARDING FORWARD-LOOKING STATEMENTS

This prospectus contains "forward-looking statements" within the meaning of the Private Securities Litigation Reform Act of 1995 (the "PSLRA") and are made pursuant to the safe harbors of the PSLRA. All statements, other than statements of historical facts, included in this prospectus regarding our strategy, future operations, regulatory process, future financial position, future revenue, projected costs, prospects, plans, objectives of management and expected market growth are forward-looking statements. The words "anticipate," "believe," "estimate," "expect," "intend," "may," "plan," "predict," "project," "will," "would" and 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:



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 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 that we may make.

You should read this prospectus and the documents that we have filed as exhibits to the registration statement, of which this prospectus is a part, completely and with the understanding that our actual future results may be materially different from what we expect. We do not assume any obligation to update any forward-looking statements, whether as a result of new information, future events or otherwise, except as required by law.

USE OF PROCEEDS

We estimate that the net proceeds from this offering will be approximately \$43.0 million, or approximately \$49.5 million if the underwriters exercise their option to purchase additional shares from us in full, in each case based on an assumed public offering price of \$7.71 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on November 4, 2011, and after deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us. A \$1.00 increase (decrease) in the assumed public offering price of \$7.71 per share would increase (decrease) our net proceeds from this offering by approximately \$5.6 million, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting the estimated underwriting discounts and commissions and estimated offering expenses.

We intend to use the net proceeds from this offering for the planned manufacture and commercialization of EXPAREL in the United States and for working capital and other general corporate purposes.

The amounts and timing of our actual expenditures may vary significantly depending on numerous factors, including:

the extent to which the FDA may require us to perform additional clinical trials for EXPAREL;

the timing and success of this offering;

the costs of our commercialization activities for EXPAREL;

the cost and timing of expanding our manufacturing facilities and purchasing manufacturing and other capital equipment for EXPAREL and our product candidates;

the scope, progress, results and costs of development for additional indications for EXPAREL and for our product candidates;

the cost, timing and outcome of regulatory review of our product candidates;

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

the extent to which we choose to establish collaboration, co-promotion, distribution or other similar agreements for product candidates;

the costs of preparing, submitting and prosecuting patent applications and maintaining, enforcing and defending intellectual property claims; and

We therefore cannot estimate the amount of net proceeds to be used for all of the purposes described above. We may find it necessary or advisable to use the net proceeds for other purposes, and we will have broad discretion in the application of net proceeds.

any unforeseen or underestimated cash needs.

Following this offering, we believe that our available funds will be sufficient to fund the commercial launch of EXPAREL in the United States. It is possible that we will not achieve the progress that we expect with respect to EXPAREL because the actual costs are difficult to predict and are subject to substantial risks and delays. We have no committed external sources of funds. To the extent that the net proceeds from

this offering and our other capital resources are insufficient to commercially launch EXPAREL, we will need to finance our cash needs through public or private equity offerings, debt financings, corporate collaboration and licensing arrangements or other financing alternatives.

Pending our use of the net proceeds from this offering, we intend to invest the net proceeds in a variety of capital preservation investments, including short-term, investment-grade, interest-bearing instruments and U.S. government securities.

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PRICE RANGE OF COMMON STOCK

Our common stock has been listed on The NASDAQ Global Market under the symbol "PCRX" since our initial public offering on February 3, 2011. Prior to that offering, there was no public market for our common stock. The following table sets forth, for the periods indicated, the high and low intraday sales prices of our common stock as reported by The NASDAQ Global Market:

Year Ended 2011	High	1	Low
Fourth Quarter (through November 4, 2011)	\$ 12.10	\$	6.60
Third Quarter	\$ 12.41	\$	7.06
Second Quarter	\$ 15.34	\$	6.80
First Quarter (beginning February 3, 2011)	\$ 7.60	\$	6.16

On November 4, 2011, the closing price of our common stock as reported on The NASDAQ Global Market was \$7.71 per share. As of the date of this prospectus, we had approximately 37 holders of record of our common stock.

DIVIDEND POLICY

We have never declared or paid any cash dividends on our capital stock. We currently intend to retain earnings, if any, to finance the growth and development of our business. We do not expect to pay any cash dividends on our common stock in the foreseeable future. Payment of future dividends, if any, will be at the discretion of our board of directors and will depend on our financial condition, results of operations, capital requirements, restrictions contained in current or future financing instruments, provisions of applicable law and other factors the board deems relevant. Our ability to pay dividends on our common stock is limited by the covenants of our loan and security agreement governing the Hercules Credit Facility and may be further restricted by the terms of any of our future indebtedness. See "Risk Factors Risks Related to Our Financial Condition and Capital Requirements Our debt obligations expose us to risks that could adversely affect our business, operating results and financial condition."

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CAPITALIZATION

The following table sets forth our cash and cash equivalents, short-term investments and capitalization as of September 30, 2011:

on an actual basis; and

on an as adjusted basis to reflect our issuance and sale of 6,000,000 shares of common stock in this offering at an assumed public offering price of \$7.71 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on November 4, 2011, after deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us.

You should read this table together with our consolidated financial statements and the related notes included elsewhere in this prospectus and the "Management's Discussion and Analysis of Financial Condition and Results of Operations," "Use of Proceeds" and "Selected Consolidated Financial Data."

	As of September 30, 2011				
	Actual (in thousands, exc	As Adjusted ⁽¹⁾ ept share and			
	per share amounts)				
Cash and cash equivalents	\$ 16,402	59,386			
Short-term investments	20,666	20,666			
Total cash, cash equivalents and short-term investments	37,068	80,052			
Long-term debt, including current portion	25,474	25,474			
Royalty interest obligation, including current portion	3,135	3,135			
Total long-term debt and royalty interest obligation	28,609	28,609			
Preferred stock, \$0.001 par value; 5,000,000 shares authorized; no shares issued or outstanding, actual and as adjusted					
Common stock, \$0.001 par value and 250,000,000 authorized: actual, 17,229,892 shares issued and 17,228,827 shares outstanding; as adjusted 23,229,892 shares issued and 23,228,827 shares					
outstanding	17	23			
Additional paid-in capital	178,821	221,799			
Accumulated deficit	(164,956)	(164,956)			
Accumulated other comprehensive loss	(5)	(5)			
Treasury stock, 1,065 shares at cost	(2)	(2)			
Total stockholders' equity	13,875	56,859			
Total capitalization	\$ 42,484	85,468			

A \$1.00 increase (decrease) in the assumed public offering price of \$7.71 per share would increase (decrease) our net proceeds from this offering by approximately \$5.6 million, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting the estimated underwriting discounts and commissions and estimated offering expenses.

The outstanding share information in the table above excludes the following as of September 30, 2011:

2,332,287 shares of common stock issuable upon the exercise of stock options outstanding at a weighted average exercise price of \$3.72 per share;

527,656 shares of common stock issuable upon the exercise of warrants outstanding and exercisable at a weighted average exercise price of \$10.22 per share;

61,242 shares of common stock available for future issuance under our equity compensation plans; and

36,750 shares of common stock issuable upon the exercise of outstanding stock options that we granted after September 30, 2011, at a weighted average exercise price of \$9.74 per share.

DILUTION

If you invest in our common stock, your interest will be diluted immediately to the extent of the difference between the initial public offering price per share you will pay in this offering and the as adjusted net tangible book value per share of our common stock after this offering.

Our historical net tangible book value as of September 30, 2011 was \$6.7 million, or \$0.39 per share of our common stock. Our historical net tangible book value per share represents our total tangible assets less total liabilities, divided by the number of shares of our common stock outstanding on September 30, 2011.

After giving effect to our issuance and sale of 6,000,000 shares of our common stock in this offering at an assumed public offering price of \$7.71 per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on November 4, 2011, after deducting the estimated underwriting discounts and commissions and estimated offering expenses payable by us, the as adjusted net tangible book value as of September 30, 2011 would have been \$49.7 million, or \$2.14 per share. This represents an immediate increase in net tangible book value to existing stockholders of \$1.75 per share. The public offering price per share will significantly exceed the net tangible book value per share. Accordingly, new investors who purchase shares of common stock in this offering will suffer an immediate dilution of their investment of \$5.57 per share. The following table illustrates this per share dilution to the new investors purchasing shares of common stock in this offering without giving effect to the over-allotment option granted to the underwriters:

Assumed public offering price per share		\$ 7.71
Net tangible book value per share as of September 30, 2011	\$ 0.39	
Increase per share attributable to sale of shares of common stock in this offering	1.75	
As adjusted net tangible book value per share after the offering		2.14
Dilution per share to new investors		5.57

A \$1.00 increase (decrease) in the assumed public offering price of \$7.71 per share would increase (decrease) our net tangible book value after this offering by approximately \$5.6 million and dilution per share to new investors by approximately \$0.76, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting the estimated underwriting discounts and commissions and estimated offering expenses.

If the underwriters exercise their over-allotment option in full, the as adjusted net tangible book value will increase to \$2.33 per share, representing an immediate increase to existing stockholders of \$1.94 per share and an immediate dilution of \$5.38 per share to new investors. If any shares are issued upon exercise of outstanding options or warrants, you will experience further dilution.

SELECTED CONSOLIDATED FINANCIAL DATA

The following selected consolidated financial data should be read together with our consolidated financial statements and related notes included elsewhere in this prospectus and the information under "Management's Discussion and Analysis of Financial Condition and Results of Operations". The selected consolidated financial data in this section is not intended to replace our consolidated financial statements and the accompanying notes. The unaudited consolidated financial data include, in the opinion of our management, all adjustments, consisting only of normal recurring adjustments that are necessary for a fair presentation of our financial position and results of operations for these periods. Our historical results for any prior period are not necessarily indicative of results to be expected in any future period, and our results for any interim period are not necessarily indicative of results to be expected for a full fiscal year.

The selected consolidated financial data as of September 30, 2011, and for the nine months ended September 30, 2010 and 2011, have been derived from our unaudited consolidated financial statements included elsewhere in this prospectus.

The selected consolidated financial data as of December 31, 2009 and 2010 and for the years ended December 31, 2008, 2009 and 2010 have been derived from our audited consolidated financial statements included elsewhere in this prospectus.

The selected consolidated financial data as of December 31, 2007 and December 31, 2008 and for the year ended December 31, 2007 have been derived from our consolidated financial statements not included in this prospectus.

The selected consolidated financial data as of December 31, 2006, and for the year December 31, 2006, and for the period from January 1, 2007 through March 23, 2007, have been derived from unaudited consolidated financial statements of the Predecessor, SkyePharma, Inc., not included in this prospectus.

The term Predecessor refers to SkyePharma, Inc. prior to March 24, 2007, or the Acquisition Date, and the term Successor refers to Pacira Pharmaceuticals, Inc. and its consolidated subsidiaries. Our results of operations for the year ended December 31, 2007, while representing a full year for Pacira Pharmaceuticals, Inc., do not reflect the operations of PPI-California until March 24, 2007, after the Acquisition Date. We have presented the Predecessor for the period from January 1, 2007 through March 23, 2007, as we believe it best presents the continuity of operations of the Successor prior to the Acquisition.

		Prede Year		sor anuary 1	Successor												
		Ended		to				Year E	nd	led				Nine Mo	nths	Enc	led
	Dec	ember 31,	, M					Decemb	er					Septe	mbe		
		2006		2007		2007		2008		2009	2010			2010	201	1	
		(unau	dite	ed)				(audi		_			(unaudited)				
Consolidated					((in thousa	nd	s, except sh	ıar	e and per	sha	are data)					
Statement of Operations Data:																	
Revenues:																	
Supply revenue	\$	5,800	\$	684	\$	5,444	\$		\$	6,324	\$	7,640	\$	7,127	\$		4,868
Royalties Collaborative		2,784		500		2,388		3,648		4,044		3,705		2,693			2,743
licensing and development		2.000		-0.4						4.620							• • • •
revenue Revenue from		3,088		204		509		3,425		4,638		3,217		2,551			3,845
SkyePharma PLC		702		39													
Total revenues		12,374		1,427		8,341		13,925		15,006	\$	14,562		12,371]	1,456
Operating expenses: Cost of revenues Research and		15,782		2,825		9,492		17,463		12,301		12,276		10,168		1	0,138
development		16,060		3,251		20,665		33,214		26,233		18,628		14,954		1	2,237
Selling, general and administrative		8,685		2,632		4,170		8,611		5,020		6,030		3,948			13,465
Acquired in-process research and development						12,400											
Total operating expenses		40,527		8,708		46,727		59,288		43,554		36,934		29,070		3	35,840
Loss from operations:		(28,153)		(7,281)		(38,386)		(45,363)		(28,548)		(22,372)		(16,699)		(2	24,384)
Other (expense) income:																	
Loss on early extinguishment of debt												(184)					
Interest income		60		4		491		235		77		146		112			111
Interest (expense) Royalty interest		(11,221)		(2,265)						(1,723)		(3,959)		(2,577)		((4,068)
obligation		4,694		(1,486)		1,686		3,490		(1,880)		(930)		(1,048)			235
Other, net		(2,713)		(13)		16		(224)		367		150		107			61
Total other (expense) income, net		(9,180)		(3,760)		2,193		3,501		(3,159)		(4,777)		(3,406)			(3,661)
Net loss	\$				\$		\$	(41,862)	\$		\$		\$		\$		28,045)
Net loss per share basic and diluted					\$	(77.85)	\$	(79.23)	\$	(55.32)	\$	(47.29)	\$	(35.02)	\$		(1.89)

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Weighted average number of common shares

464,900 528,357 573,118 574,072 574,112 14,826,054

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Predecessor						Successor						
December 31,			December 31,					September 30,				
2006	2	2007	2008 2009				2010		2011			
(unaudited)	naudited) (una		naudited) (audited)		(audited)		(audited)		(unaudited)			
					(i	n thousands	s))				
\$ 627	\$	7,240	\$	12,386	\$	7,077	\$	26,133	\$	16,402		
										20,666		
27,010		2,354		2,341		(1,868)		14,733		25,571		
63,188		39,157		50,541		43,954		66,562		77,609		
21,648		8,241		3,618		25,820		74,660		22,445		
		3		6		6		6				
		1		1		1		1		17		
(319,756)) ((36,193)		(78,055)		(109,762)		(136,911)		(164,956)		
(221,541))	8,937 46		7,490		(22,949)		(48,383)		13,875		
	December 31, 2006 (unaudited) \$ 627 27,010 63,188 21,648	December 31, 2006 (unaudited) (unaudited) (unaudited) \$ 627 \$ 27,010 63,188 21,648	December 31, 2006 2007 (unaudited) (unaudited) \$ 627 \$ 7,240 27,010 2,354 63,188 39,157 21,648 8,241 3 1 (319,756) (36,193) (221,541) 8,937	December 31, 2006 2007 (unaudited) (unaudited) (a \$ 627 \$ 7,240 \$ 27,010 2,354 63,188 39,157 21,648 8,241 3 1 (319,756) (36,193) (221,541) 8,937	December 31, 2006 (unaudited) 2007 (unaudited) December 30 (2008 (audited)) \$ 627 \$ 7,240 \$ 12,386 27,010 2,354 2,341 63,188 39,157 50,541 21,648 8,241 3,618 3 6 1 1 1 (319,756) (36,193) (78,055) (221,541) 8,937 7,490	December 31, December 32008 2006 2007 2008 (unaudited) (unaudited) (audited) (i \$ 627 \$ 7,240 \$ 12,386 \$ 27,010 2,354 2,341 63,188 39,157 50,541 21,648 8,241 3,618 3 6 1 1 1 1 (319,756) (36,193) (78,055) (74,90) (221,541) 8,937 7,490 (36,193) 7,490 (36,193) 7,490 (36,193) 7,490 (36,193) 7,490 (36,193) 7,490 (36,193) 7,490 (36,193) 7,490 (36,193) 7,490 (36,193) 7,490 (36,193) 7,490 (36,193) 7,490 (36,193) 7,490 (36,193) (36,193) 7,490 (36,193) (36,193) (36,193) (36,193) (36,193) (36,193) (36,193) (36,193) (36,193) (36,193) (36,193) (36,193) (36,193) (36,193) (36,193) (36,193)	December 31, 2006 2007 2008 2009 (unaudited) (unaudited) (audited) (in thousands) \$ 627 \$ 7,240 \$ 12,386 \$ 7,077 27,010 2,354 2,341 (1,868) 63,188 39,157 50,541 43,954 21,648 8,241 3,618 25,820 3 6 6 1 1 1 (319,756) (36,193) (78,055) (109,762) (221,541) 8,937 7,490 (22,949)	December 31, 2006 2007 2008 2009 (unaudited) (unaudited) (audited) (audited) (in thousands) \$ 627 \$ 7,240 \$ 12,386 \$ 7,077 \$ 27,010 2,354 2,341 (1,868) 63,188 39,157 50,541 43,954 21,648 8,241 3,618 25,820 3 6 6 6 1 1 1 (319,756) (36,193) (78,055) (109,762) (221,541) 8,937 7,490 (22,949)	December 31, 2006 (unaudited) 2007 (audited) 2008 (audited) 2009 (audited) 2010 (audited) \$ 627 \$ 7,240 \$ 12,386 \$ 7,077 \$ 26,133 27,010 2,354 2,341 (1,868) 14,733 63,188 39,157 50,541 43,954 66,562 21,648 8,241 3,618 25,820 74,660 3 6 6 6 1 1 1 1 1 1 1 1 (319,756) (36,193) (78,055) (109,762) (136,911) (221,541) 8,937 7,490 (22,949) (48,383)	December 31, Sep 2006 2007 2008 2009 2010 (unaudited) (unaudited) (audited) (audited) (audited) (undited) (u		

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

You should read the following discussion and analysis of our financial condition and results of operations together with our financial statements and the related notes appearing at the end of this prospectus. Some of the information contained in this discussion and analysis or set forth elsewhere in this prospectus, including information with respect to our plans and strategy for our business and related financing, includes forward-looking statements that involve risks and uncertainties. You should read the "Risk Factors" section of this prospectus for a discussion of important factors that could cause actual results to differ materially from the results described in or implied by the forward-looking statements contained in the following discussion and analysis.

Overview

We are an emerging specialty pharmaceutical company focused on the development, commercialization and manufacture of proprietary pharmaceutical products, based on our proprietary DepoFoam drug delivery technology, for use in hospitals and ambulatory surgery centers.

On October 28, 2011, the United States Food and Drug Administration, or FDA, approved our New Drug Application, or NDA, for our lead product candidate, EXPAREL, a liposome injection of bupivacaine, an amide-type local anesthetic, indicated for administration into the surgical site to produce postsurgical analgesia. We are developing a sales force entirely dedicated to commercializing EXPAREL comprised of approximately 60 representatives, seven regional managers and a national sales manager. We intend to develop this sales force pursuant to a contract with Quintiles Commercial US, Inc., a division of Quintiles, Inc., or Quintiles, and under the terms of this contract we have the flexibility to hire all or a portion of the sales force dedicated to commercializing EXPAREL as full-time employees of Pacira, upon 60 days notice to Quintiles. We believe that our pre-launch activities including significant personal interactions with our hospital customers, position us for a successful launch of EXPAREL in the first quarter of 2012.

Our two marketed products, DepoCyt(e) and DepoDur, and our proprietary DepoFoam extended release drug delivery technology were acquired as part of the acquisition of our California operating subsidiary, Pacira Pharmaceuticals Inc., or PPI-California, on March 24, 2007, or the Acquisition. DepoCyt(e) is a sustained release liposomal formulation of the chemotherapeutic agent cytarabine and is indicated for the intrathecal treatment of lymphomatous meningitis. DepoCyt(e) was granted accelerated approval by the FDA in 1999 and full approval in 2007. DepoDur is an extended release injectable formulation of morphine indicated for epidural administration for the treatment of pain following major surgery. DepoDur was approved by the FDA in 2004.

We do not expect our currently marketed products, other than EXPAREL, to generate revenue that is sufficient for us to achieve profitability because we expect to continue to incur significant expenses as we commercially launch EXPAREL and advance the development of our product candidates, seek FDA approval for our product candidates that successfully complete clinical trials and develop our sales force and marketing capabilities to prepare for their commercial launch. We also expect to incur additional expenses to add operational, financial and management information systems and personnel, including personnel to support our product development efforts and our obligations as a public reporting company. For us to become and remain profitable, we believe that we must succeed in commercializing EXPAREL or other product candidates with significant market potential.

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Recent Developments

FDA Approval of EXPAREL

On October 28, 2011, our New Drug Application, or NDA for our lead product candidate, EXPAREL, a liposome injection of bupivacaine, an amide-type local anesthetic, indicated for administration into the surgical site to produce postsurgical analgesia was approved by the United States Food and Drug Administration, or FDA.

Hercules Credit Facility

Tranche A of the Hercules Credit Facility is guaranteed by certain of our investors which guarantee is limited to each such investor's pro rata portion of the outstanding principal and accrued and unpaid interest of Tranche A under the Hercules Credit Facility, but in no event exceeding \$11.3 million in the aggregate. The loan and security agreement governing the Hercules Credit Facility provides that, upon the occurrence of certain circumstances and upon our request, the investors' guarantee may be terminated and released. On October 28, 2011, we met the required conditions and requested the release of the guaranty. Upon the release of the investors' guaranty, the interest rate on the Tranche A portion of the term loan will increase to a floating per annum interest rate equal to 11.00% plus the amount, if any, by which the prime rate exceeds 4.00%. In addition, we also elected to extend the interest only period from November 30, 2011 to February 28, 2012.

Novo Milestone

On October 18, 2011, a development milestone was met pursuant to our agreement, or the Novo Agreement, with Novo Nordisk A/S, or Novo, which entitles us to a \$2.0 million cash payment.

Financial Operations Overview

Revenues

Our revenue that is derived from DepoCyt(e) and DepoDur is comprised of two components: supply revenue and royalties. We manufacture these products, which are then sold to our commercial partners. Supply revenue is derived from a contractual supply price paid to us by our commercial partners. Royalties are recognized as the product is sold by our commercial partners and is typically calculated as a percentage of the net selling price, which is net of discounts, returns, and allowances incurred by our commercial partners. Accordingly, the primary factors that determine our revenues derived from DepoCyt(e) and DepoDur are:

the level of orders submitted by our commercial partners;
the timing of running manufacturing lots sold to our commercial partners;
the level of prescription and institutional demand for our products;
unit sales prices;
the amount of gross-to-net sales adjustments realized by our commercial partners; and
exchange rates on European sales, denominated in euros, that are repatriated in dollars.

We also generate collaborative licensing and development revenue from our collaborations with third parties who seek to use our DepoFoam technology to develop extended release formulations of their products and product candidates. For example, in January 2011, we entered into the Novo Agreement, pursuant to which we granted non-exclusive rights to Novo under certain of our patents

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and know-how to develop, manufacture and commercialize formulations of a Novo proprietary drug using our DepoFoam drug delivery technology. Under this agreement, we agreed to undertake specified development and technology transfer activities and to manufacture pre-clinical and certain clinical supplies of such DepoFoam formulated Novo product until the completion of such technology transfer activities. Novo is obligated to pay for all costs we incur in conducting such development, manufacturing and technology transfer activities. We received an upfront license fee of \$1.5 million from Novo. We are also entitled to receive single-digit royalties on sales of such Novo product for up to twelve years following the first commercial sale of such Novo product. In addition, we are entitled to receive up to \$24 million in milestone payments based on achievement of specified development events, and up to an additional \$20 million in milestone payments based on sales of such Novo product exceeding specified amounts.

Cost of Revenues

Cost of revenues consists of the costs associated with producing our products for our commercial partners and providing research and development services to our collaboration partners. In particular, our cost of revenues includes:

manufacturing overhead and fixed costs associated with running two cGMP manufacturing facilities, including salaries and related costs of personnel involved with our manufacturing activities;

allocated overhead, personnel conducting research and development, as well as research and development performed by outside contractors or consultants for our collaborative licensing and development activities;

royalties due to third parties on our revenues;

packaging, testing, freight and shipping;

the cost of active pharmaceutical ingredients; and

overhead costs associated with excess manufacturing capacity, which are charged to cost of revenues as incurred.

Research and Development Expenses

Our research and development expenses consist of expenses incurred in developing, testing, manufacturing and seeking regulatory approval of our product candidates, including:

expenses associated with regulatory submissions, clinical trials and manufacturing, including additional expenses to prepare for the commercial manufacture of EXPAREL, such as the hiring and training of additional personnel;

payments to third-party contract research organizations, contract laboratories and independent contractors;

payments made to consultants who perform research and development on our behalf and assist us in the preparation of regulatory filings;

payments made to third-party investigators who perform research and development on our behalf and clinical sites where such research and development is conducted;

personnel related expenses, such as salaries, benefits, travel and other related expenses, including stock-based compensation; and

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facility, maintenance, and allocated rent, utilities, and depreciation and amortization, and other related expenses.

Clinical trial and manufacturing process development expenses for our product candidates are a significant component of our current research and development expenses. Product candidates in later stage clinical development, such as EXPAREL, generally have higher research and development expenses than those in earlier stages of development, primarily due to the increased size and duration of the clinical trials and manufacturing process development. We coordinate clinical trials through a number of contracted investigational sites and recognize the associated expense based on a number of factors, including actual and estimated subject enrollment and visits, direct pass-through costs and other clinical site fees.

From the Acquisition through September 30, 2011, we incurred research and development expenses of \$111.0 million, of which \$107.2 million is related to the development of EXPAREL. We incurred research and development expenses associated with the development of EXPAREL of \$12.1 million for the nine months ended September 30, 2011 and \$14.2 million for the nine months ended September 30, 2010.

We expect to incur additional research and development expenses as we develop EXPAREL in additional indications. These expenditures are subject to numerous uncertainties regarding timing and cost to completion. Completion of clinical trials may take several years or more and the length of time generally varies according to the type, complexity, novelty and intended use of a product candidate. We are currently unable to determine our future research and development expenses related to EXPAREL because the requirements of any additional clinical trials of EXPAREL for additional indications has yet to be determined. The cost of clinical development may vary significantly due to factors such as the scope, rate of progress, expense and outcome of our clinical trials and other development activities.

Selling, General and Administrative Expenses

Selling, general and administrative expenses consist primarily of salaries, benefits and other related costs, including stock-based compensation, for personnel serving in our executive, finance, accounting, legal, human resource, and sales and marketing functions. Our selling, general and administrative expenses also include facility and related costs not included in research and development expenses and cost of revenues, professional fees for legal, consulting, tax and accounting services, insurance, depreciation and general corporate expenses. We expect that our selling, general and administrative expenses will increase with the continued development and potential commercialization of our product candidates and increased expenses associated with us being a public company. Additionally, we are building a commercial infrastructure in connection with the commercialization of EXPAREL and will outsource most of our sales force through our relationship with Quintiles.

Other Income (Expense)

Other income (expense) consists of interest income, interest expense, and royalty interest obligation. Interest income consists of interest earned on our cash and cash equivalents, and amortization of discount on a note receivable from one of our commercial partners. Interest expense consists primarily of cash and non-cash interest costs related to our credit facility, our secured and unsecured notes issued to certain of our investors that were converted into common stock upon completion of our initial public offering, and negotiated rent deferral payments. Royalty interest obligation consists of our royalty payments made in connection with the amended and restated royalty interests assignment agreement, or the Amended and Restated Royalty Interests Assignment Agreement, with Royalty Securitization Trust I, an affiliate of Paul Capital Advisors, LLC, or Paul Capital.

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We record our royalty interest obligation as a liability in our consolidated balance sheets in accordance with ASC 470-10-25, Sales of Future Revenues. We impute interest expense associated with this liability using the effective interest rate method. The effective interest rate may vary during the term of the agreement depending on a number of factors including the actual sales of DepoCyt(e) and DepoDur and a significant estimation, performed quarterly, of certain of our future cash flows related to these products during the remaining term of the Amended and Restated Royalty Interests Assignment Agreement which terminates on December 31, 2014. The effect of the change in the estimates is reflected in our consolidated statements of operations as royalty interest obligation within other (expense) income. In addition, such cash flows are subject to foreign exchange movements related to sales of DepoCyt(e) and DepoDur denominated in currencies other than U.S. dollars.

Critical Accounting Policies and Use of Estimates

The preparation of our consolidated financial statements requires us to make estimates that affect the reported amounts of assets and liabilities and the disclosure of contingent assets and liabilities at the date of the financial statements as well as the reported revenues and expenses during the reporting periods. On an ongoing basis, we evaluate our estimates and judgments, including those related to clinical trial expenses and stock-based compensation. We base our estimates on historical experience, current business factors and various other assumptions that we believe are necessary to form a basis for making judgements about the carrying values of assets and liabilities and related disclosures. Although we believe our estimates and assumptions are reasonable, actual results may differ significantly from these estimates.

While our significant accounting policies are more fully discussed in Note 2 to our audited consolidated financial statements included in this prospectus, we believe that the following accounting policies are critical to the process of making significant judgments and estimates in the preparation of our financial statements.

Revenue Recognition

We recognize revenue in accordance with SEC Staff Accounting Bulletin, or SAB, No. 104, *Revenue Recognition*, and Statement of Financial Accounting Standards, or ASC 605, *Revenue Recognition*.

Supply revenue

We recognize supply revenue from products manufactured and supplied to our commercial partners, when the following four basic revenue recognition criteria under the related accounting guidance 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. Prior to the shipment of our manufactured products, we conduct initial product release and stability testing in accordance with cGMP. Our commercial partners can return the products within contracted specified timeframes if the products do not meet the applicable inspection tests. We estimate our return reserves based on our experience with historical return rates. Historically, our product returns have not been material.

Royalties

We recognize revenue from royalties based on our commercial partners' net sales of products. Royalties are recognized as earned in accordance with contract terms when they can be reasonably estimated and collectability is reasonably assured. Our commercial partners are obligated to report their net product sales and the resulting royalty due to us within 60 days from the end of each quarter.

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Based on historical product sales, royalty receipts and other relevant information, we accrue royalty revenue each quarter and subsequently true-up when we receive royalty reports from our commercial partners.

Collaborative licensing and development revenue

We recognize revenue from reimbursement received in connection with feasibility studies and development work for third parties who desire to utilize our DepoFoam extended release drug delivery technology for their products, when our contractual services are performed, provided collectability is reasonably assured. Our principal costs under these agreements include our personnel conducting research and development, and our allocated overhead, as well as research and development performed by outside contractors or consultants.

We recognize revenues from non-refundable up-front license fees received under collaboration agreements ratably over the performance period as determined under the collaboration agreement (estimated development period in the case of development agreements, and contract period or longest patent life in the case of supply and distribution agreements). If the estimated performance period is subsequently modified, we will modify the period over which the up-front license fee is recognized accordingly on a prospective basis. Upon termination of a collaboration agreement, any remaining non-refundable license fees received by us, which had been deferred, are generally recognized in full. All such recognized revenues are included in collaborative licensing and development revenue in our consolidated statements of operations.

We recognize revenue from milestone payments received under collaboration agreements when earned, provided that the milestone event is substantive, its achievability was not reasonably assured at the inception of the agreement, we have no further performance obligations relating to the event, and collectability is reasonably assured. If these criteria are not met, we recognize milestone payments ratably over the remaining period of our performance obligations under the collaboration agreement.

Research and Development Expenses

We expense all research and development costs as incurred. We rely on third parties to conduct our preclinical and clinical studies and to provide services, including data management, statistical analysis and electronic compilation for our clinical trials. We track and record information regarding third-party research and development expenses for each study or trial that we conduct and recognize these expenses based on the estimated progress towards completion at the end of each reporting period. Factors we consider in preparing these estimates include the number of subjects enrolled in studies, milestones achieved and other criteria related to the efforts of our vendors. Historically, any adjustments we have made to these assumptions have not been material. Depending on the timing of payments to vendors and estimated services provided, we may record net prepaid or accrued expenses related to these costs.

We expense the manufacturing costs (labor and overhead) of our clinical supplies as incurred. To date, these expenses have not been material. Unused raw material for manufacturing clinical supplies is included in inventory and expensed when used.

Stock-Based Compensation

We have adopted the fair value recognition provisions of Financial Accounting Standards Board Accounting Standards Codification, or ASC, 718 "Accounting for Stock Based Compensation" (formerly Statement of Financial Accounting Standards No. 123(R), Share-Based Payments), which we refer to as ASC 718, using the modified prospective transition method. The modified prospective transition method applies the provisions of ASC 718 to new awards and to awards modified, repurchased or

cancelled after the adoption date. Additionally, compensation cost for the portion of the awards for which the requisite service has not been rendered that are outstanding as of the adoption date is recognized in the Statement of Operations over the remaining service period after the adoption date based on the award's original estimate of fair value. All stock-based awards granted to non-employees are accounted for at their fair value in accordance with ASC 718, and ASC 505, "Accounting for Equity Instruments that are Issued to Other Than Employees for Acquiring, or in Conjunction with Selling, Goods or Services," under which compensation expense is generally recognized over the vesting period of the award. Determining the amount of stock-based compensation to be recorded requires us to develop estimates of fair values of stock options as of the grant date.

For the nine months ended September 30, 2011, we recognized employee stock-based compensation expense of \$2.0 million. The employee stock-based compensation expense recorded during the nine months ended September 30, 2010 and during the year ended December 31, 2010 was not significant. For the years ended December 31, 2009 and 2008, we recognized employee stock-based compensation expense of \$0.5 million and \$0.2 million, respectively.

We account for stock-based compensation by measuring and recognizing compensation expense for all stock-based payments made to employees and directors based on estimated grant date fair values. We use the straight-line method to allocate compensation cost to reporting periods over each optionee's requisite service period, which is generally the vesting period. We estimate the fair value of our stock-based awards to employees and directors using the Black-Scholes option valuation model, or Black-Scholes model. The Black-Scholes model requires the input of subjective assumptions, including the expected stock price volatility, the calculation of expected term and the fair value of the underlying common stock on the date of grant, among other inputs.

The following table summarizes our assumptions used in the Black-Scholes model:

	Nine Months Ended September 30,	Year	Ended Decemb	er 31,
	2011	2010	2009	2008
Expected volatility (weighted average)	77.0%	80.8%	82.0%	78.2%
Expected term (in years)	6.25 years	6.25 years	6.25 years	6.25 years
Risk-free interest rate	1.2-3.7%	1.6-3.4%	2.1-2.7%	1.9-3.8%
Expected dividend yield	None	None	None	None

Expected Volatility The expected volatility rate used to value stock option grants is based on volatilities of a peer group of similar companies whose share prices are publicly available. The peer group was developed based on companies in the pharmaceutical and biotechnology industry in a similar stage of development.

Expected Term We elected to utilize the "simplified" method for "plain vanilla" options to estimate the expected term of stock option grants. Under this approach, the weighted average expected life is presumed to be the average of the vesting term and the contractual term of the option.

Risk-Free Interest Rate The risk-free interest rate assumption was based on zero coupon U.S. Treasury instruments that had terms consistent with the expected term of our stock option grants.

Expected Dividend Yield We have never declared or paid any cash dividends and do not presently plan to pay cash dividends in the foreseeable future.

Results of Operations

(1)

Comparison of Nine Months Ended September 30, 2011 and 2010

	Nine Mon Septem	% Increase/		
	2011	(Decrease)		
	(in thou			
Revenues	\$ 11,456	\$ 12,371	(7)%	
Cost of Revenues	10,138	10,168	(0)%	
Research and development	12,237	14,954	(18)%	
Selling, general and administrative	13,465	3,948	241%	
Other income (expense), net	(3,661)	(3,406)	7%	
Revenues				

The following table sets forth a summary of our supply revenue, royalties and collaborative licensing and development revenue for the nine months ended September 30, 2011 and 2010.

	Nine Mon Septem		% Increase/			
	2011		2010	(Decrease)		
	(in tho	usan	ds)			
DepoCyt(e) ⁽¹⁾						
Supply revenue	\$ 4,868	\$	6,497	(25)%		
Royalties	2,600		2,470	5%		
	7,468		8,967	(17)%		
DepoDur ⁽¹⁾						
Supply revenue			630	(100)%		
Royalties	143		223	(36)%		
regulates	143		853	(83)%		
Total DanaCyt(a) and						
Total DepoCyt(e) and DepoDur revenue ⁽¹⁾	7,611		9,820	(22)0/		
DepoDul Tevenue	7,011		9,620	(22)%		
Collaborative licensing and development revenue	3,845		2,551	51%		
Total revenues	\$ 11,456	\$	12,371	(7)%		

Revenues decreased by \$0.9 million, or 7%, to \$11.5 million in the nine months ended September 30, 2011 as compared to \$12.4 million in the nine months ended September 30, 2010 primarily due to a decrease in supply revenue of \$2.3 million that was partially offset by \$1.3 million increase in collaborative licensing and development revenue. The decrease in supply revenue reflects a lower number of lot sales to our commercial partners. The increase in collaborative licensing and development revenue is primarily attributable to activities performed under the Novo Agreement signed in January 2011.

Total DepoCyt(e) and DepoDur revenue does not include collaborative licensing and development revenue related to DepoCyt(e) and DepoDur.

Cost of Revenues

Cost of revenues remained consistent for the nine months ended September 30, 2011 as compared to the nine months ended September 30, 2010. The cost of revenues for the nine months ended September 30, 2011 reflects a reduction from the nine months ended September 30, 2010 of

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\$1.8 million due to lower lot sales of Depocyt(e) and DepoDur to our commercial partners which was offset by an increase in our excess capacity costs. We h