

UNITED STATES  
SECURITIES AND EXCHANGE COMMISSION  
WASHINGTON, D.C. 20549

FORM 10-K

FOR ANNUAL & TRANSITION REPORTS PURSUANT TO  
SECTION 13 OR 15(d) OF THE SECURITIES EXCHANGE ACT OF 1934

(MARK ONE)

Annual Report Pursuant to Section 13 or 15(d) of the Securities Exchange Act of 1934  
For the fiscal year ended December 31, 2013

or

Transition Report Pursuant to Section 13 or 15(d) of the Securities Exchange Act of 1934  
For the transition period from \_\_\_\_\_ to \_\_\_\_\_

Commission File Number: 001-33221

**HERON THERAPEUTICS, INC.**

(Exact name of registrant as specified in its charter)

DELAWARE  
(State or other jurisdiction of incorporation or organization)  
123 SAGINAW DRIVE, REDWOOD CITY, CALIFORNIA  
(Address of principal executive offices)

94-2875566  
(I.R.S. Employer Identification Number)  
94063  
(Zip Code)

Registrant's telephone number, including area code:  
(650) 366-2626

Securities registered pursuant to Section 12(b) of the Act:

COMMON STOCK

Securities registered pursuant to Section 12(g) of the Act:

NONE

Indicate by check mark if the registrant is a well-known seasoned issuer, as defined in Rule 405 of the Securities Exchange Act. Yes  No

Indicate by check mark if the registrant is not required to file reports pursuant to Section 13 or Section 15(d) of the Exchange Act. Yes  No

Indicate by check mark whether the registrant (1) has filed all reports required to be filed by Section 13 or 15(d) of the Securities Exchange Act of 1934 during the preceding 12 months (or for such shorter period that the registrant was required to file such reports), and (2) has been subject to such filing requirements for the past 90 days. Yes  No

Indicate by check mark whether the registrant has submitted electronically and posted on its corporate Web site, if any, every Interactive Data File required to be submitted and posted pursuant to Rule 405 of Regulation S-T (§232.405 of this chapter) during the preceding 12 months (or for such shorter period that the registrant was required to submit and post such files).  Yes  No

Indicate by check mark if disclosure of delinquent filers pursuant to Item 405 of Regulation S-K (§229.405) is not contained herein, and will not be contained, to the best of the registrant's knowledge, in definitive proxy or information statements incorporated by reference in Part III of this Form 10-K or any amendment to this Form 10-K.

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 12b-2 of the Exchange Act. (Check One)

Large accelerated filer  Accelerated filer  Non-accelerated filer  Smaller reporting company

Indicate by check mark whether the registrant is a shell company (as defined in Rule 12b-2 of the Exchange Act). Yes  No

The aggregate market value of the voting and non-voting common equity of the registrant held by non-affiliates of the registrant as of June 30, 2013 was \$73,726,812<sup>(1)</sup> based upon the closing sale price on OTCQB reported for such date.

As of February 28, 2014, 23,723,968 shares of registrant's Common Stock, \$.01 par value, were outstanding.

DOCUMENTS INCORPORATED BY REFERENCE

Certain information required to be disclosed in Part III of this report is incorporated by reference from the registrant's definitive Proxy Statement for the 2014 Annual Meeting of Stockholders, which proxy statement will be filed not later than 120 days after the end of the fiscal

year covered by this report.

- <sup>(1)</sup> Excludes 5,051,049 shares held by directors, officers and stockholders whose ownership exceeds 10% of the outstanding shares at June 30, 2013. Exclusion of such shares should not be construed as indicating that the holders thereof possess the power, directly or indirectly, to direct the management or policies of the registrant, or that such person is controlled by or under common control with the registrant.

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PART I

**Introduction—Forward-Looking Statements**

In this Annual Report on Form 10-K, the “Company,” “Heron Therapeutics,” “we,” “us” and “our” refer to Heron Therapeutics, Inc.

*This Annual Report on Form 10-K contains forward-looking statements within the meaning of Section 21E of the Securities Exchange Act of 1934, as amended, and Section 27A of the Securities Act of 1933, as amended. All statements contained in this Form 10-K, other than statements of historical fact, are forward-looking statements. When used in this report or elsewhere by management from time to time, the words “believe,” “anticipate,” “intend,” “plan,” “estimate,” “expect,” “may,” “will,” “should,” “seeks” and similar expressions are forward-looking statements. Such forward-looking statements are based on current expectations, but the absence of these words does not necessarily mean that a statement is not forward-looking. Forward-looking statements made in this Form 10-K include, but are not limited to, statements about:*

- *the anticipated progress of our research, development and clinical programs and timing of, and prospects for, regulatory approval and commercial introduction of Sustol (also known as APF530) and other future product candidates;*
- *estimates of the timing of our resubmission of the New Drug Application (NDA) for Sustol;*
- *if approved, the timing of market introduction of Sustol or other future product candidates;*
- *our ability to successfully market, commercialize and achieve market acceptance for Sustol or other future product candidates;*
- *our ability to successfully develop other drug candidates utilizing our Biochronomer polymer;*
- *our ability to establish collaborations for our technology, Sustol and other future product candidates;*
- *uncertainties associated with obtaining and enforcing patents;*
- *our estimates for future performance; and*
- *our estimates regarding our capital requirements and our needs for, and ability to obtain, additional financing.*

*Forward-looking statements are not guarantees of future performance and involve risks and uncertainties. Actual events or results may differ materially from those discussed in the forward-looking statements as a result of various factors. For a more detailed discussion of such forward-looking statements and the potential risks and uncertainties that may impact our actual results, see the “Risk Factors” section of this Form 10-K and the other risks and uncertainties described below under the headings: “Our Lead Product Candidate Sustol,” “Development Pipeline,” “Our Technology Platform,” “Our Strategy,” “Manufacturing and Supply,” “Patents and Trade Secrets,” “Competition,” and under “Management’s Discussion and Analysis of Financial Condition and Results of Operations.” These forward-looking statements reflect our view only as of the date of this report. Except as required by law, we undertake no obligations to update any forward looking statements and we disclaim any intent to update forward-looking statements after the date of this report to reflect subsequent developments. Accordingly, you should also carefully consider the factors set forth in other reports or documents that we file from time to time with the Securities and Exchange Commission.*

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**ITEM 1. BUSINESS**

**Company Overview**

Heron Therapeutics, Inc. (formerly A.P. Pharma, Inc.) is a specialty pharmaceutical company developing product candidates using its proprietary Biochronomer™ polymer-based drug delivery platform. This drug delivery platform is designed to improve the therapeutic profile of injectable pharmaceuticals by converting them from products that must be injected once or twice per day to products that need to be injected only once every one or two weeks.

Our lead product candidate, Sustol® (also known as APF530), is being developed for the prevention of acute chemotherapy-induced nausea and vomiting (CINV) for patients undergoing both moderately and highly emetogenic chemotherapy and for the prevention of delayed CINV for patients undergoing moderately emetogenic chemotherapy. One of the most debilitating side effects of cancer chemotherapy, CINV is a leading cause of premature discontinuations of treatment. There is only one injectable 5-HT<sub>3</sub> antagonist approved for the prevention of delayed-onset CINV, so this indication represents an area of particular unmet medical need. Sustol contains the 5-HT<sub>3</sub> antagonist granisetron formulated in the Company's proprietary Biochronomer™ drug delivery system, which allows therapeutic drug levels to be maintained for five days with a single subcutaneous injection. This five-day range is designed to cover the delayed phase of CINV. Granisetron was selected for Sustol because it is widely prescribed by physicians based on a well-established record of safety and efficacy.

In May 2009, we filed an original NDA seeking approval for Sustol with the U.S. Food and Drug Administration (FDA). The FDA issued a Complete Response Letter for the Sustol NDA in March 2010. In September 2012, we resubmitted our NDA for Sustol and, in March 2013, we received a second Complete Response Letter, which identified several remaining issues that need to be addressed prior to approval of the Sustol NDA. We are currently working on addressing these issues and expect to resubmit the Sustol NDA in the second quarter of 2014.

We own the worldwide rights to Sustol and are in the early stages of building the commercial infrastructure necessary to commercialize Sustol in the U.S. on our own, assuming approval by the FDA.

In addition to resubmitting the Sustol NDA, we are seeking to expand the labeled indication for Sustol following its potential approval. In the first half of 2014, we plan to initiate a Phase 3 clinical study with Sustol for the prevention of delayed CINV in patients receiving highly emetogenic chemotherapy, an indication for which no 5-HT<sub>3</sub> antagonist currently is approved. The 1,000-patient study is expected to be completed in late 2014.

Our core Biochronomer technology, on which Sustol and our other product candidates are based, consists of bioerodible polymers designed to release drugs over a defined period of time. We have completed over 100 *in vivo* and *in vitro* studies demonstrating that our Biochronomer technology is potentially applicable to a range of therapeutic areas, including, among others, prevention of CINV, pain management and control of inflammation. We have also completed comprehensive animal and human toxicology studies that have established that our Biochronomer polymers are safe and well tolerated. Furthermore, our Biochronomer technology can be designed to deliver drugs over periods varying from days to potentially multiple weeks. We are exploring the potential use of our Biochronomer polymer with other drugs and intend to pursue the clinical development of one or more other drug candidates based on our proprietary delivery platform.

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In November 2013, we initiated a program to expand our pipeline of sustained-release products, including a new program targeting the relief of post-surgical pain. Preliminary animal studies indicate formulations using our Biochronomer delivery system provided sustained analgesic effects of three to five days. We plan to finalize formulation optimization in 2014 and initiate a Phase 1 study in mid-2014. As of 2012, approximately 25 million procedures<sup>1</sup> associated with post-operative pain were conducted in the U.S. In addition, U.S. post-operative pain market sales were approximately \$3.1 billion<sup>2</sup> in 2012.

We were founded in February 1983 as a California corporation under the name AMCO Polymerics, Inc. (AMCO). AMCO changed its name to Advanced Polymer Systems, Inc. in 1984 and was reincorporated in the state of Delaware in 1987. We changed our name to A.P. Pharma, Inc. in May 2001 to reflect a new pharmaceutical focus.

On January 13, 2014, we amended our Certificate of Incorporation to change our name to Heron Therapeutics, Inc. and to effect a combination of our outstanding common stock at a ratio of one-for-twenty (Reverse Stock Split). All historical share and per share amounts have been adjusted to reflect the Reverse Stock Split. All stock options, convertible notes and warrants outstanding were ratably adjusted to give effect to the Reverse Stock Split.

On January 23, 2014, our common stock was approved for listing and began trading on The NASDAQ Capital Market under the symbol HRTX.

Our principal offices are located at 123 Saginaw Drive, Redwood City, California 94063. Our telephone number is (650) 366-2626. Our website is located at [www.heronrx.com](http://www.heronrx.com). Information contained on, or that can be accessed through, our website is not part of this Annual Report on Form 10-K.

**Our Lead Product Candidate—SUSTOL**

***CINV Background***

Prevention and control of nausea and vomiting, or emesis, are paramount in the treatment of cancer patients. The majority of patients receiving chemotherapy will experience some degree of emesis if not prevented with an antiemetic. Chemotherapy treatments can be classified as moderately emetogenic, meaning that 30–90% of patients would experience CINV, or highly emetogenic, meaning that over 90% of patients would experience CINV, if they were not treated with an antiemetic prior to chemotherapy. Onset of CINV within the first 24 hours is described as “acute,” and CINV that occurs more than 24 hours after treatment is described as “delayed.” Delayed CINV may persist for several days from the time of onset. Prevention of CINV is important because the distress caused by CINV can severely disrupt patient quality of life and can lead some patients to delay or discontinue chemotherapy.

***Current Therapy***

Chemotherapeutic agents activate or destroy cells in the lining of the gut, releasing a neurotransmitter called serotonin. When serotonin binds to 5-hydroxytryptamine type 3 (5-HT<sub>3</sub>) receptors, the patient experiences nausea and vomiting. Granisetron, like other 5-HT<sub>3</sub> antagonists, inhibits the vomiting reflex by preventing serotonin from binding to 5-HT<sub>3</sub> receptors. Physicians may combine 5-HT<sub>3</sub> antagonists with other agents, such as corticosteroids or neurokinin-1 (NK1) antagonists, to better prevent CINV.

<sup>1</sup> Custom Research Project by Design Resources completed in 2013.

<sup>2</sup> Decision Resources, Acute Pain, December 2012.

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Current treatment options for preventing CINV include injectable 5-HT<sub>3</sub> antagonists such as palonosetron (Aloxi®), ondansetron (formerly marketed by GlaxoSmithKline as Zofran®) and granisetron (formerly marketed by Roche as Kytril®). Aprepitant (Emend®), an NK1 antagonist, is also used to prevent CINV and is typically used in combination with an injectable 5-HT<sub>3</sub> antagonist. As shown in the table below, several injectable 5-HT<sub>3</sub> antagonists are approved for the prevention of acute CINV in patients receiving either moderately or highly emetogenic chemotherapy. Within the last several years, generic versions of granisetron and ondansetron have become available. Aloxi is the only injectable 5-HT<sub>3</sub> antagonist approved for the prevention of delayed CINV in patients receiving moderately emetogenic chemotherapy. No injectable 5-HT<sub>3</sub> antagonist is approved for the prevention of delayed CINV in patients receiving highly emetogenic chemotherapy.

**Approved Injectable 5-HT<sub>3</sub> Antagonists**

<b>Chemotherapy Regimen</b>	<b>Acute CINV</b>	<b>Delayed CINV</b>
Moderately Emetogenic	Granisetron (Kytril) Ondansetron (Zofran) Palonosetron (Aloxi)	Palonosetron (Aloxi)
Highly Emetogenic	Granisetron (Kytril) Ondansetron (Zofran) Palonosetron (Aloxi)	None

Despite evidence that delayed CINV affects as many as 50–70% of patients and that more patients experience delayed CINV than acute CINV, oncology nurses and physicians frequently underestimate the magnitude of these problems in the patients for whom they care. This may occur in part since patients often do not report side effects they experience at home following chemotherapy treatments. Even though high percentages of chemotherapy patients experience such delayed nausea and emesis, presently Aloxi is the only injectable 5-HT<sub>3</sub> antagonist approved for the prevention of delayed CINV. We believe that Sustol, if approved, could become the second long-acting product given in a single injection that is capable of addressing this important medical need. Eisai Company, which markets Aloxi in the U.S., reported U.S. Aloxi sales of \$427 million in calendar year 2013.

***Our Solution—SUSTOL***

Our lead product candidate, Sustol, is being developed for the prevention of acute CINV in patients receiving both moderately and highly emetogenic chemotherapy and for the prevention of delayed CINV in patients receiving moderately emetogenic chemotherapy. Sustol is delivered by a single subcutaneous injection and contains the 5-HT<sub>3</sub> antagonist granisetron. Granisetron, for infusion and oral tablets, is approved for the prevention of acute CINV, but not delayed CINV. We selected granisetron for Sustol because it is widely prescribed by physicians based on a well-established record of safety and efficacy and because it became generically available in 2007.

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In our pivotal Phase 3 clinical trial of Sustol, in which we enrolled more than 1,300 patients, we successfully demonstrated that Sustol's efficacy in preventing CINV was statistically non-inferior to that of Aloxi. If we obtain regulatory approval for Sustol, we believe that Sustol will represent an attractive treatment for the many cancer patients that suffer from CINV.

***Phase 2 Clinical Trial***

In September 2005, we completed an open-label Phase 2 clinical trial of Sustol. We evaluated the safety, tolerability and pharmacokinetics of Sustol in 45 cancer patients undergoing either moderately or highly emetogenic chemotherapy. In addition, efficacy endpoints were evaluated relating to emetic events and the use of additional medication for treating CINV. Sustol was well tolerated in this study; there were no serious adverse events attributed to Sustol, and fewer than 10% of participating patients had injection site reactions, all of which were mild.

A substantial proportion of the patients in our Phase 2 clinical trial were complete responders, meaning they experienced no vomiting and received no additional medication for CINV during the observation period. These efficacy results compared favorably to similar data for Aloxi, as reported from its Phase 3 clinical trials. Based on these results, we designed our Phase 3 clinical program to directly compare Sustol to Aloxi in a prospective randomized trial design.

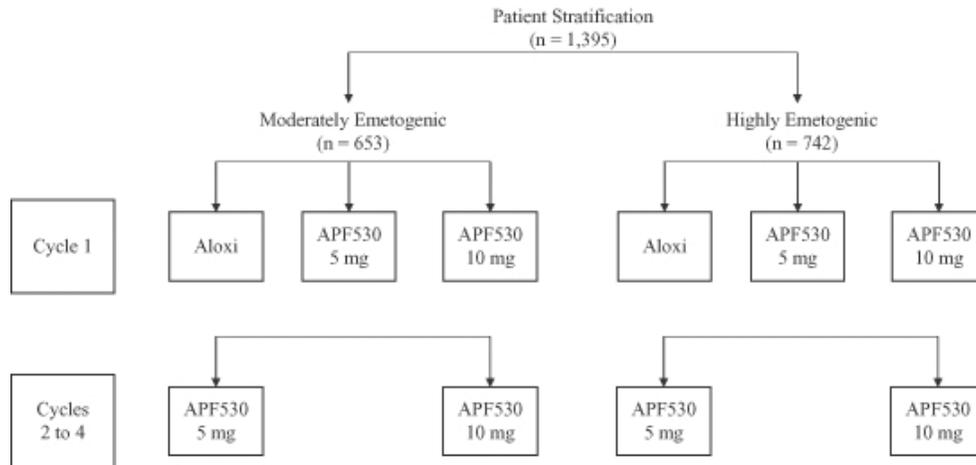
***Pivotal Phase 3 Clinical Trial***

In December 2005, we held our end-of-Phase 2 meeting with the FDA, at which we discussed our registration strategy and our proposed design for the pivotal Phase 3 clinical trial. Following this meeting, we finalized plans for our pivotal Phase 3 clinical trial in accordance with FDA input. The goals of the trial were to demonstrate the safety and efficacy of Sustol in the treatment of CINV following the administration of highly or moderately emetogenic chemotherapy and to establish an effective dose for Sustol. The trial was structured to compare the two Sustol doses (containing 5 mg and 10 mg of granisetron) with the FDA-approved dose of Aloxi across four different primary efficacy endpoints:

- non-inferiority to Aloxi for the prevention of acute CINV in patients receiving moderately emetogenic chemotherapies;
- non-inferiority to Aloxi for the prevention of delayed CINV in patients receiving moderately emetogenic chemotherapies;
- non-inferiority to Aloxi for the prevention of acute CINV in patients receiving highly emetogenic chemotherapies; and
- superiority to Aloxi for the prevention of delayed CINV in patients receiving highly emetogenic chemotherapies (superiority to Aloxi was chosen for this endpoint because Aloxi is not approved for this indication).

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Our pivotal Phase 3 clinical trial was initiated in May 2006 as a multicenter, randomized, observer-blind, actively-controlled, double-dummy, parallel group study that compared the efficacy of Sustol to Aloxi. The trial stratified patients into two groups, one receiving moderately and the other receiving highly emetogenic chemotherapeutic agents in accordance with the Hesketh algorithm, which assigns emetogenic levels based on the chemotherapy agent, drug dosage and combinations employed. In each emetogenic group, patients were randomized during Cycle 1 to receive Sustol high dose (10 mg granisetron), Sustol low dose (5 mg granisetron) or the currently approved dose of Aloxi. For up to three subsequent treatment cycles (Cycles 2–4), the patients were re-randomized to receive either of the two Sustol doses. The diagram below provides further graphical representation of the patient stratification design and target enrollment for patient randomization in our clinical trial. The study completed patient enrollment of 1,395 patients in June 2008, and we announced top-line results on September 30, 2008.



The 10 mg dose of Sustol achieved complete response (CR) rates that were numerically higher than, and statistically non-inferior to, Aloxi across all four assessments. The 5 mg dose of Sustol did not demonstrate non-inferiority to Aloxi for all endpoints. Sustol did not achieve the superiority endpoint for the delayed CINV assessment for highly emetogenic chemotherapies. Aloxi is not FDA approved for the prevention of delayed CINV in patients receiving highly emetogenic chemotherapies; therefore, Sustol needed to be statistically superior to Aloxi for this endpoint to receive FDA approval for this use. CR was defined as the absence of emetic episodes or use of anti-emetic rescue medications during a specified period of time. The time periods studied for CINV onset were 0 to 24 hours after chemotherapy, which is known as acute CINV, and 24 to 120 hours after chemotherapy, which is known as delayed CINV.

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The results summarized below are the primary endpoints from the study, with such data being drawn from the first cycle of treatment:

**Complete Response by Treatment–Cycle 1**

Emetogenicity Level	Treatment Group			Statistics vs. Aloxi (Confidence Interval)	
	SUSTOL (5 mg)	SUSTOL (10 mg)	Aloxi	5 mg	10 mg
Moderately emetogenic	(n=214)	(n=212)	(n=208)		
• Acute CINV	74.8%	76.9%	75.0%	NI (-9.8, 9.3)	NI (-7.5, 11.4)
• Delayed CINV	51.4%	58.5%	57.2%	I (-16.7, 5.1)	NI (-9.5, 12.1)
Highly emetogenic	(n=229)	(n=240)	(n=238)		
• Acute CINV	77.7%	81.3%	80.7%	NI (-12.1, 6.1)	NI (-8.2, 9.3)
• Delayed CINV	62.4%	67.1%	64.3%	NS (-12.6, 8.8)	NS (-7.7, 13.2)

(NI) Non-inferior efficacy was determined using a modified Bonferroni step down procedure. The lower bound of the adjusted Confidence Interval to establish non-inferiority was negative 15%. The Confidence Intervals shown for the moderately emetogenic and highly emetogenic levels are 97.5% and 98.3%, respectively. (NS) = No significant difference. (I) = Inferior efficacy.

Sustol was generally well tolerated, with a side effect profile consistent with previous human use of granisetron and only one serious adverse event reported as possibly attributed to Sustol. In Cycle 1, the data showed a low incidence of patients discontinuing therapy due to any adverse events (related or unrelated to study drugs): 0.5%, 0.9% and 0.9% in the moderately emetogenic patient group, and 2.0%, 3.5% and 1.2% in the highly emetogenic patient group for Sustol 5 mg, Sustol 10 mg and Aloxi, respectively. Further, of the patients completing the first cycle, 1,043 went on to receive a total of 2,374 additional doses of Sustol in Cycles 2 to 4. Of these patients, only 2 (or 0.2%) discontinued therapy due to treatment-related adverse events.

Additional data from the pivotal Phase 3 clinical trial comparing Sustol to Aloxi were released on November 5, 2008 and are reported below. These additional data included predetermined secondary efficacy endpoints and safety data that were not available at the time the top-line data were released. We believe that the overall clinical data package demonstrates the robustness of the Sustol clinical response within and across chemotherapy cycles. Some of the additional key findings follow:

- Collectively, the Phase 3 efficacy and safety data support the conclusion that 10 mg is the most effective dose of Sustol and, therefore, was the selected dose for the NDA.

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- In patients receiving multiple cycles of Sustol, CR rates were observed to generally increase over four cycles of chemotherapy, as shown in the following table:

**Complete Response of SUSTOL 10 mg Dose Over Four Chemotherapy Cycles**

<b>Emetogenicity Level</b>	<b>Cycle 1</b>	<b>Cycle 2</b>	<b>Cycle 3</b>	<b>Cycle 4</b>
<b>Moderately Emetogenic</b>	(n=212)	(n=240)	(n=184)	(n=134)
• Acute (0-24h)	76.9%	77.1%	78.8%	83.6%
• Delayed (24-120h)	59.0%	62.1%	61.4%	66.4%
• Overall (0-120h)	54.2%	58.8%	60.3%	63.4%
<b>Highly Emetogenic</b>	(n=240)	(n=263)	(n=202)	(n=148)
• Acute (0-24h)	81.3%	84.8%	89.6%	87.8%
• Delayed (24-120h)	68.3%	76.0%	81.2%	83.8%
• Overall (0-120h)	64.6%	72.2%	78.7%	79.7%

- The Phase 3 clinical trial protocol predefined multiple primary and secondary endpoints, including complete response, complete control (no emesis, no rescue therapy and no greater-than-mild nausea) and total response (no emesis, no rescue therapy and no nausea) measured over defined time intervals (acute, delayed and overall). Although there were no significant differences between the Sustol 10 mg dose and Aloxi, the response rates for the Sustol 10 mg dose were numerically higher than Aloxi in all nine analyses for moderately emetogenic chemotherapy and in five of nine analyses for highly emetogenic chemotherapy. As noted above, however, Sustol did not achieve the primary endpoint of significant superiority to Aloxi in the highly emetogenic group at any dose.
- The safety profile for Sustol was similar to that for Aloxi; the most notable adverse event was constipation, observed in 15.4% and 13.4% of patients receiving Sustol 10 mg and Aloxi, respectively. Headache was observed in 10.0% and 9.7% of patients receiving the Sustol 10 mg dose and Aloxi, respectively.
- Investigators were required to observe and record all reactions associated with the subcutaneous injection site on days one and five for each treatment cycle. Overall, greater than 90% of the recorded observations were mild in severity, the most common being injection-site redness and bruising. With each additional cycle of treatment, the frequency of injection site reactions decreased, indicating Sustol can safely be administered for multiple cycles.
- During the trial, patients received more than 1,600 separate injections of the Sustol 10 mg dose. Assessment of any injection-site pain was made on days one and five of treatment: on day one, less than 0.1% of injections resulted in any reports of pain; on day five, approximately 4% of injections resulted in a report of pain. All but four of these reports of pain were recorded as mild, with the four recorded as moderate. Additional data from the pivotal Phase 3 clinical trial were presented at the annual meeting of the American Society of Clinical Oncology on June 1, 2009, and are reported below.
- CR rates for the Sustol 10 mg dose were generally higher in patients who had received prior chemotherapy when compared to patients who had not received any previous chemotherapy. Additionally, in all instances, CR rates for Sustol in patients receiving prior chemotherapy were numerically higher than those observed for Aloxi. Based on previous clinical studies, many physicians believe that the risk of CINV increases with each additional cycle of chemotherapy. These new data may suggest potential utility for Sustol in treating patients who have received prior chemotherapy.

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- Of the highly emetogenic chemotherapy regimens, those containing cisplatin are considered to be the most troublesome due to their ability to cause significant delayed CINV. The CR rates for patients receiving cisplatin-based regimens were numerically higher for the Sustol 10 mg dose when compared to Aloxi in both acute and delayed CINV. Specifically, in acute CINV, Sustol had an 81.1% CR rate versus 75.5% for Aloxi, and, in delayed CINV, Sustol had a 66.0% CR rate versus 60.4% for Aloxi. These differences were not statistically significant.
- A pharmacokinetic analysis, conducted in a sub-group of patients, showed that a single Sustol 10 mg dose maintained blood levels of granisetron for the entire five-day period.

***Additional Clinical Studies***

In the first quarter of 2012, at the request of the FDA, the Company completed a thorough QT study of Sustol. The study was conducted to assess the potential for granisetron, the active drug in Sustol, to prolong the QT interval across a wide range of plasma drug concentrations (the QT interval is a measure of the time between the start of the Q wave and the end of the T wave in the heart's electrical cycle, and, in general, the QT interval represents electrical depolarization and repolarization of the left and right ventricles). Prolongation of the QT interval may increase the risk of fatal cardiac tachyarrhythmias. As such, the FDA requires a thorough QT study, which examines a drug's potential to prolong the QT interval, for many drugs in development. Moxifloxacin, a drug known to prolong the QT interval, is a standard positive control used in thorough QT studies. The study met its protocol-specified primary end point and demonstrated that granisetron did not have an effect on cardiac repolarization as measured by prolongation of the QT interval. A pharmacokinetic/pharmacodynamic (PK/PD) analysis demonstrated that there was no relationship between plasma granisetron concentrations and the heart-rate-corrected QT interval (QTc) (slope of zero).

This study was a randomized, double-blind, placebo-controlled, four-way, crossover trial in 56 healthy adults that compared the effects of: (1) Sustol at twice its proposed therapeutic dose; (2) intravenous granisetron at five times its therapeutic dose; (3) oral moxifloxacin (400 mg), a known pro-arrhythmic; and (4) saline placebo, on the surface electrocardiogram with primary focus on the QT interval. The primary end point was to determine that granisetron had no clinically meaningful effect on QTc, defined as the upper bound of the one-sided 95% confidence interval for placebo-adjusted, baseline-subtracted QTc being less than 10 milliseconds at all time points. The primary end point was met irrespective of heart-rate correction methodology (QTcF, QTcI, QTcB). Moxifloxacin, the study's positive control, demonstrated QTc prolongation consistent with previous clinical experience.

Also in the first quarter of 2012, the Company completed a study of the metabolism of Sustol in healthy volunteers. This study was requested by the FDA to corroborate pre-clinical animal metabolism data for the polymer used in Sustol with metabolism data in humans. The study provided quantitative results confirming how the polymer is metabolized by the human body. The results of the study were consistent with preclinical studies, and these results were included in the resubmission of the NDA.

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***New Drug Application***

In May 2009, we filed an NDA for Sustol with the FDA under Section 505(b)(2) of the Federal Food, Drug, and Cosmetic Act (FDCA). In March 2010, we received a Complete Response Letter from the FDA, which stated that the NDA we submitted in May 2009, requesting approval of Sustol, could not be approved as it was initially submitted. The primary points raised in the initial Complete Response Letter were as follows:

*Dosing System*

- The FDA expressed concerns relating to our former two-syringe administration system, including potential issues with the transfer of material from one syringe to the other syringe prior to patient administration, certain components used in the dosing system and the potential risk of improper administration of the drug product.

*Chemistry, Manufacturing and Control*

- The FDA conducted inspections of our facility and several of our contract manufacturing facilities. The FDA identified certain deficiencies during these inspections, and stated that satisfactory resolution of these deficiencies would be required for approval.
- During the NDA review, the FDA asked that we determine if terminal sterilization with gamma irradiation is a feasible approach to enhance the assurance of sterility. We have subsequently demonstrated that terminal sterilization is feasible, and the FDA has requested we change to terminal sterilization prior to approval.
- The FDA requested clarification and revision of certain analytical specifications proposed in our NDA.

*Clinical*

- The FDA did not request additional clinical efficacy studies, although the FDA has asked for the re-presentation and re-analysis of select existing Phase 3 clinical trial data.
- The FDA requested we perform two studies relating to bioavailability and metabolism.
- The FDA did not accept our request to waive the requirement for a thorough QT study.

We met with the FDA in February and March 2011 to clarify the work needed to address the issues identified in the letter and resubmit the NDA. At the February 2011 meeting, we presented information concerning the clinical pharmacology of Sustol and a revised presentation format for certain clinical data from our Phase 3 clinical study. The FDA indicated that we would need to complete a thorough QT study prior to resubmitting its NDA and clarified the requirements for a previously requested metabolism study. The FDA also indicated that the revised presentation format for the clinical data was acceptable for resubmission. The FDA did not request that we conduct any additional efficacy studies. At the March 2011 meeting, the dosing system and the characterization and manufacturing of Sustol were discussed. We also presented the results of the additional analytical work it had completed since receipt of the Complete Response Letter.

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Following these meetings, we performed a number of activities in order to address the items requested by the FDA, including the following:

- We changed our dosing system from the former two-syringe administration system employing a 1-inch needle to a single-syringe system employing a 5/8-inch needle.
- We conducted Human Factors studies to demonstrate that Sustol could be administered safely to patients.
- We modified our proposed Package Insert, product packaging and Instructions for Use to further ensure proper administration of the product.
- We demonstrated that terminal sterilization is feasible and changed our manufacturing process to incorporate terminal sterilization.
- We made other changes to our manufacturing and quality control/quality assurance processes to address concerns raised by the FDA.
- We conducted and completed thorough QT and metabolism studies based on protocols agreed upon with the FDA (see Additional Clinical Studies).

In September 2012, we resubmitted the NDA seeking approval for Sustol. In March 2013, the FDA issued a second Complete Response Letter, which identified several issues that precluded the approval of the Sustol NDA resubmission, including issues relating to:

*Chemistry, Manufacturing and Control*

- The FDA requested resolution of observations made during inspection at two of our contract manufacturing sites. Satisfactory resolution of these issues is required for resubmission and approval.

*Administration*

- The FDA requested that we repeat the Human Factors testing with material made by commercial suppliers.

*Clinical*

- The FDA requested that we provide data sets and algorithms to allow the reanalysis of the clinical data versus the revised American Society of Clinical Oncology (ASCO) 2011 CINV guidelines.

We believe the issues that remain are addressable, and we are working expeditiously to resubmit the Sustol NDA in the second quarter of 2014.

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Section 505(b)(2) of the FDCA permits the FDA, in its review of an NDA, to rely on studies that were not conducted by or for the applicant and to which the applicant has not obtained a right of reference. Such studies can be provided by published literature, or the FDA can rely on previous findings of safety and efficacy for a previously approved drug. Section 505(b)(2) applications may be submitted for drug products that represent a modification (e.g., a new indication or new dosage form) of an eligible approved drug. In such cases, the additional information in 505(b)(2) applications necessary to support the change from the previously approved drug is frequently provided by new studies submitted by the applicant. Our 505(b)(2) application relies on the FDA's previous finding of safety and effectiveness for the active ingredient in Sustol, granisetron, together with the additional clinical and nonclinical data that we submitted to demonstrate the safety and effectiveness of our formulation of the drug product for the indications for which we are seeking approval.

***Development Pipeline***

In addition to resubmitting the Sustol NDA, we are seeking to expand the labeled indication for Sustol following its potential approval. In the first half of 2014, we plan to initiate a Phase 3 clinical study with Sustol for the prevention of delayed CINV in patients receiving highly emetogenic chemotherapy, an indication for which no 5-HT<sub>3</sub> antagonist currently is approved. The 1,000-patient study is expected to be completed in late 2014.

In November 2013, we initiated a program to expand our pipeline of sustained release product candidates, including a new program targeting the relief of post-surgical pain. Preliminary animal studies indicate formulations using our Biochronomer delivery system provided sustained analgesic effects of three to five days. We plan to finalize formulation optimization in 2014 and initiate a Phase 1 study in the second half of 2014. As of 2012, approximately 25 million procedures<sup>3</sup> associated with post-operative pain were conducted in the U.S. In addition, U.S. post-operative pain market sales were approximately \$3.1 billion<sup>4</sup> in 2012.

***Research and Development***

As of December 31, 2013, we had 31 employees engaged in research and development and quality control. Research and development expenses for 2013, 2012 and 2011 were \$32.8 million, \$15.2 million and \$8.2 million, respectively.

***Our Technology Platform***

We have developed a broad family of polymers with unique attributes, known collectively as poly (ortho esters), under the trade name Biochronomer. We have completed over 100 in vivo and in vitro studies demonstrating that our Biochronomer technology is potentially applicable to a range of therapeutic areas, including, among others, pain management, prevention of nausea and control of inflammation. We have also completed comprehensive animal and human toxicology studies that have established that our Biochronomer polymers are safe and well tolerated.

Our Biochronomer technology can provide sustained levels of drugs for prolonged efficacy. The Biochronomer "links," or bonds, are stable at neutral pH conditions. Upon coming into contact with water-containing media, such as internal body fluids, the water reacts with these bonds. This reaction is known as hydrolysis. During the hydrolysis of the Biochronomer links, acidic

<sup>3</sup> Custom Research Project by Design Resources completed in 2013.

<sup>4</sup> Decision Resources, Acute Pain, December 2012.

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elements are produced in a local micro-environment, in a controlled manner, without impacting the overall neutrality of the drug delivery system. These elements assist in the continued, controlled erosion of the polymer with a simultaneous, controlled release of the active drug contained within the Biochronomer. By varying the amount of the acidic elements in the Biochronomer, different rates of hydrolysis may be effectively realized. In this manner, delivery times ranging from days to potentially multiple weeks can be achieved.

Due to the inherent versatility of our Biochronomer technology, products can be designed to deliver drugs at a variety of implantation sites including: under the skin, at the site of a surgical procedure, in joints, in the eye or in muscle tissue. Our Biochronomers can be prepared in a variety of physical forms, ranging from hard, glassy materials to fluids of varying viscosity that are injectable at room temperature, by proper selection of monomers. A significant advantage of our Biochronomer technology is that drugs can be incorporated by standard mixing procedures, allowing the potential production of formulations in the form of injectable gels, microspheres, coatings and strands.

**Our Strategy**

Our primary near-term objective is to obtain FDA approval of the NDA for Sustol. We believe that there is significant market potential for Sustol for the prevention of acute CINV for both moderately and highly emetogenic chemotherapy, and for the prevention of delayed CINV in moderately emetogenic chemotherapy. In addition to resubmitting the Sustol NDA, we plan to initiate a Phase 3 clinical study with Sustol in 2014 for the prevention of delayed CINV in patients receiving highly emetogenic chemotherapy, which represents a possible expanded indication for Sustol, if approved. The 1,000-patient study is expected to be completed in late 2014. We own the worldwide rights to Sustol and are in the early stages of building the commercial infrastructure necessary to commercialize Sustol in the U.S. on our own. Longer term, we intend to become a leading specialty pharmaceutical company focused on improving the effectiveness of existing pharmaceuticals using our proprietary drug delivery technologies.

**Manufacturing and Supply**

We do not currently own or operate manufacturing facilities for the production of clinical or commercial quantities of any of our product candidates. We rely on a small number of third-party manufacturers to produce our compounds and expect to continue to do so to meet the preclinical and clinical requirements of our potential product candidates and for all of our commercial needs. We do not have long-term agreements with any of these third parties. We require in our manufacturing and processing agreements that all third-party contract manufacturers and processors produce active pharmaceutical ingredients (APIs) and finished products in accordance with the FDA's current Good Manufacturing Practices (cGMP) and all other applicable laws and regulations. We maintain confidentiality agreements with potential and existing manufacturers in order to protect our proprietary rights related to our drug candidates.

With regard to our lead product candidate, Sustol, we use third-parties to supply granisetron API, to manufacture our polymer and to formulate, fill and finish our final product. We first source the API, granisetron, from independent suppliers. We then use a different third-party supplier to manufacture our proprietary polymer. Another third-party supplier then receives API granisetron and polymer and formulates them into the bulk drug product and completes the process by filling bulk drug product into syringes. To date, Sustol has been manufactured in small quantities for preclinical studies and clinical trials. If Sustol is approved for commercial sale, we will need to manufacture the product in larger quantities. Significant scale-up of manufactur-

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ing requires additional process development and validation studies, which the FDA must review and approve. We are currently in the process of completing this scale-up and validation work. If approved, the commercial success of Sustol, in the near-term, will be dependent upon the ability of our contract manufacturers to produce a product in commercial quantities at competitive costs of manufacture. If Sustol receives regulatory approval, we plan to scale-up manufacturing through our third-party manufacturers for Sustol with the objective of realizing important economies of scale. These scale-up activities will take time to implement, require additional capital investment, process development and validation studies, and FDA approval. We cannot guarantee that we will be successful in achieving competitive manufacturing costs through such scale-up activities.

**Sales and Marketing**

We own all worldwide rights to Sustol and are in the early stages of building commercial infrastructure necessary to commercialize Sustol in the U.S. on our own. We intend to establish a direct sales force if Sustol is approved.

**Patents and Trade Secrets**

Patents and other proprietary rights are important to our business. We generally seek patent protection for our inventions in key potential markets, and rely upon trade secrets, know-how, continuing technological innovations and licensing opportunities to develop and maintain our competitive position.

As part of our strategy to protect our current product candidates and to provide a foundation for future products, we have filed a number of U.S. patent applications on inventions relating to the composition of a variety of polymers, specific products, product groups and processing technology. As of December 31, 2013, we had a total of 15 issued U.S. patents and an additional 38 issued (or registered) foreign patents. The patents on our bioerodible technologies expire between January 2016 and April 2026. The product Sustol is covered by patents owned by Heron Therapeutics, Inc. that are granted in the United States and in foreign countries. Currently, the product Sustol is covered by six patents issued in the United States and by seven patents issued in foreign countries including Brazil, Canada, European Union (EU), Japan, and Taiwan. U.S. patents covering Sustol have expiration dates ranging from January 2016 to September 2024; foreign patents covering Sustol have expiration dates ranging from January 2017 to September 2025. Granted patents include claims covering the product composition, methods of use and methods of preparation. Our policy is to actively seek patent protection in the United States and to pursue equivalent patent claims in selected foreign countries, thereby seeking patent coverage for novel technologies and compositions of matter that may be commercially important to the development of our business.

Although we believe the bases for these patents and patent applications are sound, they are untested, and there is no assurance that they will not be successfully challenged. There can be no assurance that any patent previously issued will be of commercial value, that any patent applications will result in issued patents of commercial value, or that our technology will not be held to infringe patents held by others.

We also rely on unpatented trade secrets and know-how to protect certain aspects of our production technologies. Our employees, consultants, advisors and corporate partners have entered into confidentiality agreements with us. These agreements, however, may not necessarily provide meaningful protection for our trade secrets or proprietary know-how in the event of unauthorized use or disclosure. In addition, others may obtain access to, or independently develop, these trade secrets or know-how.

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**Competition**

The pharmaceutical industry is highly competitive. Many of our competitors have substantially greater financial, research, development, manufacturing, sales, marketing and distribution resources than we currently do. In addition, they may have significantly more experience in drug development, obtaining regulatory approval and establishing strategic collaborations. We expect any future products we develop to compete on the basis of, among other things, product efficacy and safety, time to market, price, extent of adverse side effects experienced and convenience of administration and drug delivery. We also expect to face competition in our efforts to identify appropriate collaborators or partners to help commercialize our product candidates in our target commercial areas.

Sustol is expected to face significant competition for the prevention of delayed CINV, principally from Eisai's Aloxi (palonosetron). In addition to Aloxi, Sustol will compete with entrenched generic forms of granisetron (formerly marketed by Roche as Kytril) and ondansetron (formerly marketed by GlaxoSmithKline as Zofran). Generic versions of Aloxi may become available after its last scheduled patent expiration date, the longest of which extends to 2024. We are aware, however, of ongoing litigation challenging these patents and, if such challenges are successful, it is possible that generic versions of Aloxi could become available sooner. We are also aware of several companies that have developed, or are developing, both generic and new formulations of granisetron, including transdermal formulations such as ProStrakan's Sancuso® (granisetron transdermal patch), which was approved in 2008.

There are several companies that are developing new formulations of existing drugs using novel drug delivery technologies. The following are some of our major competitors among drug delivery system developers: Alkermes, Inc., Durect Corporation and Pacira Pharmaceuticals, Inc.

**Government Regulation and Product Approvals**

The manufacturing and marketing of our potential products and our ongoing research and development activities are subject to extensive regulation by the FDA and comparable regulatory agencies in state and local jurisdictions and in foreign countries.

***United States Regulation***

Before any of our products can be marketed in the United States, they must be approved by the FDA. To secure approval, any drug we develop must undergo rigorous preclinical testing and clinical trials that demonstrate the product candidate's safety and effectiveness for each chosen indication for use. These extensive regulatory processes control, among other things: the development, testing, manufacture, safety, efficacy, record keeping, labeling, storage, approval, advertising, promotion, sale and distribution of biopharmaceutical products.

In general, the process required by the FDA before investigational drugs may be marketed in the United States involves the following steps:

- preclinical laboratory and animal tests;
- submission of an IND, which must become effective before human clinical trials may begin in the U.S.;

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- adequate and well-controlled human clinical trials to establish the safety and efficacy of the proposed drug for its intended use;
- pre-approval inspection of manufacturing facilities and selected clinical investigators; and
- FDA approval of an NDA or of an NDA supplement (for subsequent indications).

### ***Preclinical Testing***

In the United States, drug candidates are tested in animals until adequate proof-of-safety is established. These preclinical studies generally evaluate the mechanism of action of the product and assess the potential safety and efficacy of the product. Tested compounds must be produced according to applicable cGMP requirements, and preclinical safety tests must be conducted in compliance with FDA and international regulations regarding good laboratory practices (GLP). The results of the preclinical tests, together with manufacturing information and analytical data, are generally submitted to the FDA as part of an IND, which must become effective before human clinical trials may commence. The IND will automatically become effective 30 days after receipt by the FDA, unless before that time the FDA requests an extension or raises concerns about the conduct of the clinical trials as outlined in the application. If the FDA has any concerns, the sponsor of the application and the FDA must resolve the concerns before clinical trials can begin. Submission of an IND may not result in FDA authorization to commence a clinical trial. A separate submission to the existing IND must be made for each successive clinical trial conducted during product development. Regulatory authorities may require additional data before allowing the clinical studies to commence or proceed from one phase to another and could demand that the studies be discontinued or suspended at any time if there are significant safety issues. Furthermore, an independent institutional review board (IRB), for each medical center proposing to participate in the conduct of the clinical trial must review and approve the clinical protocol and patient informed consent before the center commences the study.

### ***Clinical Trials***

Clinical trials for new drug candidates are typically conducted in three sequential phases that may overlap. In Phase 1, the initial introduction of the drug candidate into human volunteers, the emphasis is on testing for safety or adverse effects, dosage, tolerance, metabolism, distribution, excretion and clinical pharmacology. Phase 2 involves studies in a limited patient population to determine the initial efficacy of the drug candidate for specific targeted indications, to determine dosage tolerance and optimal dosage and to identify possible adverse side effects and safety risks. Once a compound shows evidence of effectiveness and is found to have an acceptable safety profile in Phase 2 evaluations, pivotal Phase 3 clinical trials are undertaken to more fully evaluate clinical outcomes and to establish the overall risk/benefit profile of the drug and to provide, if appropriate, an adequate basis for product labeling. During all clinical trials, physicians will monitor patients to determine effectiveness of the drug candidate and to observe and report any reactions or safety risks that may result from use of the drug candidate. The FDA, the IRB (or their foreign equivalents) or the sponsor may suspend a clinical trial at any time on various grounds, including a finding that the subjects are being exposed to an unacceptable health risk.

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***Data Review and Approval***

The data from the clinical trials, together with preclinical data and other supporting information that establishes a drug candidate's safety, are submitted to the FDA in the form of an NDA, or NDA supplement (for approval of a new indication if the product candidate is already approved for another indication). Under applicable laws and FDA regulations, each NDA submitted for FDA approval is usually given an internal administrative review within 60 days following submission of the NDA. If deemed complete, the FDA will "file" the NDA, thereby triggering substantive review of the application.

The FDA can refuse to file any NDA that it deems incomplete or not properly reviewable. The FDA has established internal substantive review goals of six months for priority NDAs (for drugs addressing serious or life threatening conditions for which there is an unmet medical need) and ten months for regular NDAs. The FDA, however, is not legally required to complete its review within these periods, and these performance goals may change over time. Moreover, in many cases, the outcome of the review, even if generally favorable, is not an actual approval, but a "complete response" that describes additional work that must be done before the NDA can be approved. The FDA's review of an NDA may involve review and recommendations by an independent FDA advisory committee. The FDA may deny approval of an NDA, or NDA supplement, if the applicable regulatory criteria are not satisfied, or it may require additional clinical data and/or an additional pivotal Phase 3 clinical trial. Even if such data are submitted, the FDA may ultimately decide that the NDA or NDA supplement does not satisfy the criteria for approval.

Satisfaction of FDA requirements or similar requirements of state, local, and foreign regulatory agencies typically takes several years and requires the expenditure of substantial financial resources. Information generated in this process is susceptible to varying interpretations that could delay, limit or prevent regulatory approval at any stage of the process. Accordingly, the actual time and expense required to bring a product to market may vary substantially. We cannot assure you that we will submit applications for required authorizations to manufacture and/or market potential products or that any such application will be reviewed and approved by the appropriate regulatory authorities in a timely manner, if at all. Data obtained from clinical activities is not always conclusive and may be susceptible to varying interpretations which could delay, limit or prevent regulatory approval. Success in early stage clinical trials does not ensure success in later stage clinical trials. Even if a product candidate receives regulatory approval, the approval may be significantly limited to specific disease states, patient populations and dosages, or have conditions placed on it that restrict the commercial applications, advertising, promotion or distribution of these products.

Once issued, the FDA may withdraw product approval if ongoing regulatory standards are not met or if safety problems occur after the product reaches the market. In addition, the FDA may require testing and surveillance programs to monitor the safety or effectiveness of approved products which have been commercialized, and the FDA has the power to prevent or limit further marketing of a product based on the results of these post-marketing programs. The FDA may also request or require additional clinical trials after a product is approved, which are referred to as Phase 4 clinical studies. The results of Phase 4 clinical studies can confirm the effectiveness of a product candidate and can provide important safety information to augment the FDA's voluntary adverse drug reaction reporting system. Any products manufactured or distributed by us pursuant to FDA approvals would be subject to continuing regulation by the FDA, including record-keeping requirements and reporting of adverse experiences with the drug. Drug manufacturers and their subcontractors are required to register their establishments with the FDA and certain state agencies and are subject to periodic unannounced inspections by the FDA and certain state agencies for compliance with cGMPs, which impose certain procedural and documentation requirements upon us and our third-party manufacturers. We cannot be certain that we, or our present or future suppliers, will be able to comply with the cGMP regulations and other FDA regulatory requirements. If our present or future suppliers are not able to comply with these requirements, the

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FDA may halt our clinical trials, require us to recall a drug from distribution, or withdraw approval of the NDA for that drug. Furthermore, even after regulatory approval is obtained, later discovery of previously unknown problems with a product may result in restrictions on the product or even complete withdrawal of the product from the market.

The FDA closely regulates the marketing and promotion of drugs. Approval may be subject to post-marketing surveillance and other record-keeping and reporting obligations, and involve ongoing requirements. Product approvals may be withdrawn if compliance with regulatory standards is not maintained or if problems occur following initial marketing. A company can make only those claims relating to safety and efficacy that are approved by the FDA. Failure to comply with these requirements can result in adverse publicity, warning letters, corrective advertising and potential civil and criminal penalties. Physicians may prescribe legally available drugs for uses that are not described in the product's labeling and that differ from those tested by us and approved by the FDA. Such off-label uses are common across medical specialties. Physicians may believe that such off-label uses are the best treatment for many patients in varied circumstances. The FDA does not regulate the behavior of physicians in their choice of treatments. The FDA does, however, restrict manufacturers' communications on off-label uses of approved drugs.

***Section 505(b)(2) Applications***

Some of our product candidates may be eligible for submission of applications for approval under the FDA's Section 505(b)(2) approval process, which generally requires less information than the NDAs described above. Section 505(b)(2) was enacted as part of the Drug Price Competition and Patent Term Restoration Act of 1984, also known as the Hatch-Waxman Act, and allows approval of NDAs that rely, at least in part, on studies that were not conducted by or for the applicant and to which the applicant has not obtained a right of reference. Such studies can be provided by published literature, or FDA can rely on previous findings of safety and efficacy for a previously approved drug. Section 505(b)(2) applications may be submitted for drug products that represent a modification (e.g., a new indication or new dosage form) of an eligible approved drug. In such cases, the additional information in 505(b)(2) applications necessary to support the change from the previously approved drug is frequently provided by new studies submitted by the applicant. Because a Section 505(b)(2) application relies in part on previous studies or previous FDA findings of safety and effectiveness, preparing 505(b)(2) applications is generally less costly and time-consuming than preparing an NDA based entirely on new data and information from a full set of clinical trials. The law governing Section 505(b)(2) or FDA's current policies may change in such a way as to adversely affect our applications for approval that seek to utilize the Section 505(b)(2) approach. Such changes could result in additional costs associated with additional studies or clinical trials and delays.

The FDA provides that reviews and/or approvals of applications submitted under Section 505(b)(2) will be delayed in various circumstances. For example, the holder of the NDA for the listed drug may be entitled to a period of market exclusivity during which the FDA will not approve, and may not even review, a Section 505(b)(2) application from other sponsors. If the listed drug is claimed by one or more patents that the NDA holder has listed with the FDA, the Section 505(b)(2) applicant must submit a certification with respect to each such patent. If the 505(b)(2) applicant certifies that a listed patent is invalid, unenforceable or not infringed by the product that is the subject of the Section 505(b)(2) application, it must notify the patent holder and the NDA holder. If, within 45 days of providing this notice, the NDA holder sues the 505(b)(2) applicant for patent infringement, the FDA will not approve the Section 505(b)(2) application until the earlier of a court decision favorable to the Section 505(b)(2) applicant or the expiration of 30 months. The regulations governing marketing exclusivity and patent protection are complex, and it is often unclear how they will be applied in particular circumstances.

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In addition, both before and after approval is sought, we and our collaborators are required to comply with a number of FDA requirements. For example, we are required to report certain adverse reactions and production problems, if any, to the FDA, and to comply with certain limitations and other requirements concerning advertising and promotion for our products. Also, quality control and manufacturing procedures must continue to conform to cGMP after approval, and the FDA periodically inspects manufacturing facilities to assess compliance with continuing cGMP. In addition, discovery of problems, such as safety problems, may result in changes in labeling or restrictions on a product manufacturer or NDA holder, including removal of the product from the market.

***DEA Regulation***

Our research and development processes involve the controlled use of hazardous materials, including chemicals. Some of these hazardous materials are considered to be controlled substances and subject to regulation by the U.S. Drug Enforcement Agency (DEA). Controlled substances are those drugs that appear on one of five schedules promulgated and administered by the DEA under the Controlled Substances Act (CSA). The CSA governs, among other things, the distribution, recordkeeping, handling, security and disposal of controlled substances. We must be registered by the DEA in order to engage in these activities, and are subject to periodic and ongoing inspections by the DEA and similar state drug enforcement authorities to assess ongoing compliance with the DEA's regulations. Any failure to comply with these regulations could lead to a variety of sanctions, including the revocation, or a denial of renewal, of the DEA registration, injunctions or civil or criminal penalties.

***Third-Party Payor Coverage and Reimbursement***

Although none of our current product candidates have been approved or commercialized for any indication, if they are approved for marketing, commercial success of our product candidates will depend, in part, upon the availability of coverage and reimbursement from third-party payors at the federal, state and private levels. Government payor programs, including Medicare and Medicaid, private health care insurance companies and managed care plans have attempted to control costs by limiting coverage and the amount of reimbursement for particular procedures or drug treatments. The U.S. Congress and state legislatures, from time to time, propose and adopt initiatives aimed at cost containment. Ongoing federal and state government initiatives directed at lowering the total cost of health care will likely continue to focus on health care reform, the cost of prescription pharmaceuticals and on the reform of the Medicare and Medicaid payment systems. Examples of how limits on drug coverage and reimbursement in the United States may cause reduced payments for drugs in the future include:

- changing Medicare reimbursement methodologies;
- fluctuating decisions on which drugs to include in formularies;
- revising drug rebate calculations under the Medicaid program or requiring that new or additional rebates be provided to Medicare, Medicaid, other federal or state healthcare programs; and
- reforming drug importation laws.

Some third-party payors also require pre-approval of coverage for new or innovative devices or drug therapies before they will reimburse health care providers that use such therapies. While we cannot predict whether any proposed cost-containment measures will be adopted or otherwise implemented in the future, the announcement or adoption of these proposals could have a material adverse effect on our ability to obtain adequate prices for our product candidates and operate profitably.

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**Foreign Approvals**

In addition to regulations in the United States, we may be subject to a variety of foreign regulations governing clinical trials and commercial sales and distribution of our products. Whether or not we obtain FDA approval for a product, we must obtain approval of a product by the comparable regulatory authorities of foreign countries before we can commence clinical trials or marketing of the product in those countries. The approval process varies from country to country, and the time may be longer or shorter than that required for FDA approval. The requirements governing the conduct of clinical trials, product licensing, pricing and reimbursement vary greatly from country to country.

We have not started the regulatory approval process in any jurisdiction other than the United States, and we are unable to estimate when, if ever, we will commence the regulatory approval process in any foreign jurisdiction. Foreign approvals may not be granted on a timely basis, or at all. Regulatory approval of reimbursement and/or prices is required in most countries other than the United States. The reimbursement and/or prices approved may be too low to generate an acceptable return to us. If we fail to obtain approvals from foreign jurisdictions, the geographic market for our product candidates would be limited.

**Employees**

As of December 31, 2013, we had 47 full-time employees, 5 of whom hold Ph.D. degrees and 1 of whom is an M.D., and approximately 20 full-time equivalent contract workers. There were 31 employees engaged in research and development and quality control, and 16 individuals working in commercial operations, finance, information technology, human resources and administration.

We consider our relations with our employees to be good. None of our employees are covered by a collective bargaining agreement.

**Available Information**

We make available free of charge on or through our Internet website our annual reports on Form 10-K, quarterly reports on Form 10-Q, current reports on Form 8-K and all amendments to those reports as soon as reasonably practicable after they are electronically filed with, or furnished to, the SEC. Our Internet website address is "www.herontx.com." The reference to our Internet website does not constitute incorporation by reference of the information contained on or hyperlinked from our Internet website. We file electronically with the SEC, our annual reports on Form 10-K, quarterly reports on Form 10-Q and current reports on Form 8-K pursuant to Section 13(a) or 15(d) of the Exchange Act. The SEC maintains an Internet site that contains reports, proxy information and information statements, and other information regarding issuers that file electronically with the SEC. The address of that website is <http://www.sec.gov>. The materials are also available at the SEC's Public Reference Room, located at 100 F Street, Washington, D.C. 20549. The public may obtain information through the public reference room by calling the SEC at 1-800-SEC-0330.

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**ITEM 1A. RISK FACTORS**

Our business is subject to various risks, including those described below. You should consider carefully these risk factors and all of the other information included in this Form 10-K. Any of these risk factors could materially adversely affect our business, operating results and financial condition. These risks are not the only ones we face. Additional risks not presently known to us or that we currently believe are immaterial may also significantly impair our business operations. In these circumstances, the market price of our common stock could decline, and you may lose all or part of your investment in our securities. Before you decide whether to purchase any of our common stock, you should carefully consider the risk factors set forth below as may be updated from time to time by our future filings under the Securities Exchange Act of 1934, as amended, or the Exchange Act.

**Risks Related To Our Business**

**We are substantially dependent upon the success of our Sustol product candidate. Clinical trial results and the NDA resubmission for this product may not lead to regulatory approval.**

We have invested a significant portion of our time and financial resources in the development of our most advanced product candidate, Sustol, for which we are initially seeking U.S. Food and Drug Administration (FDA) approval for the prevention of acute chemotherapy-induced nausea and vomiting (CINV) associated with both moderately and highly emetogenic chemotherapy and for the prevention of delayed CINV associated with moderately emetogenic chemotherapy (MEC). We currently also plan to conduct a Phase 3 study which, if successful, may allow us to expand our product label to include delayed CINV associated with highly emetogenic chemotherapy (HEC).

Our near-term ability to generate revenues and our future success, in large part, depends on the approval and successful commercialization of Sustol. We will not be able to commercialize Sustol until we obtain regulatory approval in the United States or foreign countries. In order to satisfy FDA approval standards for the commercial sale of Sustol, we must first successfully resolve the issues identified in the Complete Response Letter received from the FDA in March 2013. This letter identified several issues that precluded the approval of Sustol NDA, including issues relating to: manufacturing of Sustol, the administration of Sustol and our analysis of efficacy data for Sustol under more recent guidelines classifying chemotherapy regimens. Although we are currently working to address these issues and currently expect to resubmit the Sustol NDA in the second quarter of 2014, there can be no assurance that these responses will be sufficient or that we will be able to resubmit within this time period. Further, the FDA's review of our resubmission may not produce positive decisions as to whether:

- Sustol is safe and effective in its proposed use(s) and whether its benefits outweigh the risks;
- the proposed labeling for Sustol has our desired product indication regarding acute and delayed-onset CINV, as well as HEC and MEC regimens; and
- the methods used in manufacturing Sustol and the controls used to maintain its quality are adequate to preserve its identity, strength, quality and purity.

Deficiencies on any of the above, or other factors, could prevent or delay obtaining regulatory approval of Sustol, which would negatively affect our potential revenues, increase our costs and potentially impair our ability to continue as a going concern.

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**We may not obtain regulatory approval for Sustol or any of our product candidates. Regulatory approval may also be delayed or cancelled or may entail limitations on the indicated uses of a proposed product.**

The process for obtaining approval of an NDA is time consuming, subject to unanticipated delays and costs, and requires the commitment of substantial resources. The regulatory process, particularly for pharmaceutical product candidates like ours, is uncertain, can take many years and requires the expenditure of substantial resources. Any product that we or our collaborative partners develop must receive all relevant regulatory agency approvals or clearances, if any, before it may be marketed in the United States or other countries. In particular, human pharmaceutical products are subject to rigorous preclinical and clinical testing and other requirements by the FDA in the United States and similar health authorities in foreign countries. We may not receive necessary regulatory approvals or clearances to market Sustol or any other product candidate. In September 2012, we resubmitted the NDA seeking approval for Sustol with the FDA. In March 2013, we received a second Complete Response Letter, which identified several issues that precluded the approval of the Sustol NDA. We are currently working to address these issues and intend to resubmit the Sustol NDA in the second quarter of 2014. Our NDA resubmission for Sustol may not be approved, or approval may be delayed, as a result of changes in FDA policies for drug approval prior to the FDA's decision on our NDA.

For example, although many drug products have been approved by the FDA in recent years under Section 505(b)(2) of the Federal Food, Drug, and Cosmetic Act, objections have been raised to the FDA's interpretation of Section 505(b)(2). If challenges to the FDA's interpretation of Section 505(b)(2) are successful, the FDA may be required to change its interpretation, which could delay or prevent the approval of our NDA for Sustol. The review of our resubmitted NDA may also be delayed due to the FDA's internal resource constraints. Additionally, data obtained from preclinical and clinical activities are susceptible to varying interpretations that could delay, limit or prevent regulatory agency approvals or clearances. For example, the FDA may require additional clinical data to support approval, such as confirmatory studies and other data or studies to address questions or concerns that may arise during the FDA review process.

Delays in obtaining regulatory approval for Sustol, or the issuance of a third Complete Response Letter, would, among other consequences, delay the launch of Sustol and adversely affect our ability to generate revenue from sales of this product and adversely affect our ability to raise additional capital that would be necessary to sustain our operations. Given the additional delays that we would face prior to obtaining approval for Sustol, if such approval is ever granted, we may need significant additional capital to fund our operations.

Even if granted, regulatory approvals may include significant limitations on the uses for which products may be marketed. Failure to comply with applicable regulatory requirements can, among other things, result in warning letters, imposition of civil penalties or other monetary payments, delay in approving or refusal to approve a product candidate, suspension or withdrawal of regulatory approval, product recall or seizure, operating restrictions, interruption of clinical trials or manufacturing, injunctions and criminal prosecution.

In addition, the marketing and manufacturing of drugs and biological products are subject to continuing FDA review, and later discovery of previously unknown problems with a product, its manufacture or its marketing may result in the FDA requiring further clinical research or restrictions on the product or the manufacturer, including withdrawal of the product from the market.

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If Sustol is approved, but does not attain market acceptance by healthcare professionals and patients, our business prospects and results of operations will suffer.

Even if Sustol receives regulatory approval for commercial sale, the revenue that we may receive from the sale of Sustol may be less than expected and will depend on many factors that are outside of our control. Factors that may affect revenue from Sustol, if approved, include;

- the scope of our approved product label;
- perception of physicians and other members of the health care community of the safety and efficacy relative to that of competing products;
- cost-effectiveness;
- patient and physician satisfaction with the product;
- ability to manufacture commercial product successfully and on a timely basis;
- cost and availability of raw materials;
- market size for the product;
- reimbursement policies of government and third-party payors;
- unfavorable publicity concerning the product or similar drugs;
- the introduction, availability and acceptance of competing treatments;
- adverse event information relating to the product;
- product liability litigation alleging injuries relating to the product;
- product labeling or product insert language required by the FDA or regulatory authorities in other countries;
- the regulatory developments related to the manufacture or continued use of the product;
- extent and effectiveness of sales and marketing and distribution support for the product; and
- our collaborators' decisions as to the timing of product launches, pricing and discounting.

Our product revenue will be adversely affected if, due to these or other factors, the products we or our collaborators are able to commercialize do not gain significant market acceptance.

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**We have a history of losses, we expect to generate losses in the near future, and we may never achieve or maintain profitability.**

We have incurred recurring losses and had an accumulated deficit of \$238.9 million through December 31, 2013. Even if Sustol is approved, we expect to continue to generate substantial losses over at least the next several years as we:

- build a sales force and commence commercialization of Sustol, if approved;
- expand drug product development and commercialization efforts;
- conduct preclinical development and clinical trials; and
- pursue additional applications for our existing delivery technologies.

To achieve and sustain profitability, we must, alone or with others, successfully develop, obtain regulatory approval for, manufacture, market and sell our products. If Sustol is approved for commercialization, we must successfully launch and commercialize the product. If Sustol is not approved, we will likely experience significant delays before we begin to recognize meaningful levels of revenue, if ever. We will incur substantial expenses in our efforts to develop and commercialize products and we may never generate sufficient revenue to become profitable or to sustain profitability.

**Additional capital may be needed to enable us to implement our business plan, and we may be unable to raise capital, which would force us to limit or cease our operations and related product development programs. Raising such capital may have to be accomplished on unfavorable terms, likely causing dilution to our existing stockholders.**

At December 31, 2013, the Company had cash and cash equivalents in the amount of \$72.3 million. We believe that our current working capital balance is sufficient to fund essential operations into 2015; provided, however, that if we pursue additional clinical studies or commercially launch Sustol prior to 2015, we will need to raise additional capital. We are pursuing commercialization of Sustol without a partner for the U.S. market, which will likely require us to obtain additional funding and resources to sustain our operations until we can achieve profitability. The need for and amount of additional funding that we may require depends on various factors, including the results of the on-going regulatory review by the FDA of our Sustol NDA resubmission, the time and costs related to manufacturing of Sustol, if approved, technological and market developments of drugs that may compete with Sustol and the timing of clinical trials to expand our pipeline of sustained-release products. There can be no assurance that Sustol will be approved and, if approved, that we will be successful in obtaining the additional necessary financial resources and expertise, with or without a partner, that will be required to launch Sustol.

We may not be able to raise sufficient additional capital when we need it on favorable or any terms. If we are unable to obtain adequate funds, we may be required to curtail significantly or cease operations.

The timing and degree of any future capital requirements will depend on many factors, including:

- the number and characteristics of product development programs we pursue and the pace of each program;
- the scope, rate of progress, results and costs of preclinical testing and clinical trials;

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- the time, cost and outcome involved in seeking regulatory approvals;
- scientific progress in our research and development programs;
- the magnitude and scope of our research and development programs;
- our ability to establish and maintain strategic collaborations or partnerships for research, development, clinical testing, manufacturing and marketing of our product candidates;
- the cost and timing of establishing sales, marketing and distribution capabilities for a specialty sales force if we commercialize any products independently;
- the cost of establishing clinical and commercial supplies of our product candidates and any products that we may develop; and
- general market conditions.

If we issue additional equity securities or securities convertible into equity securities to raise funds, our stockholders will suffer dilution of their investment, and such issuance may adversely affect the market price of our common stock. Any debt financing we enter into may involve covenants that restrict our operations. These restrictive covenants may include, among other things, limitations on borrowing and specific restrictions on the use of our assets, as well as prohibitions on our ability to create liens, pay dividends, redeem capital stock or make investments. In the event that additional funds are obtained through arrangements with collaborative partners, these arrangements may require us to relinquish rights to some of our technologies, product candidates or products on terms that are not favorable to us or require us to enter into a collaboration arrangement that we would otherwise seek to develop and commercialize ourselves. If adequate funds are not available, we may be required to delay, reduce the scope of, or eliminate one or more of our product development programs and reduce personnel-related and other costs, which will have a material adverse effect on our business.

**The general economic environment in which we operate is experiencing continued weakness and volatility.**

Our ability to secure the additional capital that may be necessary for implementation of our longer-term business plans may be diminished due to the continuing volatile business conditions and financial markets. For example, the difficulty in obtaining additional capital necessary to develop our other product candidates has led us to temporarily suspend certain development programs in recent years. If the economic environment continues its weak recovery and financial markets continue to experience significant volatility, we may have increasing difficulty in raising additional capital when needed.

**We may depend on collaborators as a source of capital and to help us complete the process of developing and testing our products.**

Our strategy for the development, clinical testing and commercialization of our products may require entering into collaborations with corporate partners, licensors, licensees and others. These collaborations may be critical to funding our operations and our success in bringing our products and product candidates to the market and promoting such marketed products

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profitably. We could be dependent upon the subsequent success of these other parties in performing their respective responsibilities and the cooperation of our partners. Our collaborators may not cooperate with us or perform their obligations under our agreements with them. We cannot control the amount and timing of our collaborators' resources that will be devoted to our research activities related to our collaborative agreements with them. Our collaborators may choose to pursue existing or alternative technologies in preference to those being developed in collaboration with us. We may prioritize other programs ahead of collaboration activities such that funding from these other parties could be reduced or deferred. Failure to make or maintain these arrangements, or a delay in a collaborative partner's performance, or factors that may affect our partner's sales may materially adversely affect our business, results of operations and financial condition.

Under agreements with collaborators, we may rely significantly on them, among other activities, to:

- fund research and development activities with us;
- pay us fees upon the achievement of milestones; and
- market for or with us any commercial products that result from our collaborations.

**Clinical trials are expensive and may not result in commercially viable products.**

Conducting clinical trials is a lengthy, time-consuming and expensive process. For example, we have incurred significant expenses in developing Sustol and, even if approved, it may not result in a commercially viable product. We are planning a Phase 3 study of Sustol designed to demonstrate the utility of Sustol in the treatment of delayed-onset CINV in patients receiving HEC regimens. If successful, we intend to submit the results of the study in a post-approval application to expand the label of Sustol to include delayed HEC. There can be no assurance that this study will be successful or that the FDA will grant any such label expansion. Before obtaining regulatory approvals for the commercial sale of any products, we, or our partners, must demonstrate through preclinical testing and clinical trials that our product candidates are safe and effective for their intended uses in humans. We have incurred and will continue to incur substantial expense and devote a significant amount of time to preclinical testing and clinical trials.

Our business, results of operations and financial condition may be materially adversely affected by any delays in, or termination of, our clinical trials. Factors impacting our ability to generate commercially viable products through the conduct of clinical trials include:

- insufficient funds to conduct clinical trials;
- inability to find partners;
- failure of clinical trials to demonstrate the safety and efficacy of our product candidates to the extent necessary to obtain regulatory approvals;
- failure by us or third-party investigators, contract research organizations, or other third parties involved in the research to adhere to regulatory requirements applicable to the conduct of clinical trials;

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- failure of preclinical testing and early clinical trials to predict results of later clinical trials;
- delay in completion of clinical trials, resulting in increased costs; and
- inability to obtain regulatory approval of our product candidates following completion of clinical trials, or delays in obtaining such approvals.

There can be no assurance that if our clinical trials are successfully initiated and completed we will be able to obtain approval by the FDA in the United States or similar regulatory authorities elsewhere in the world in a timely manner, if at all. If we fail to successfully develop and commercialize one or more of our product candidates, we may be unable to generate sufficient revenues to attain profitability and our reputation in the industry and in the investment community would likely be damaged, each of which would cause our stock price to decrease.

**Delays in clinical testing could increase our costs and delay our ability to obtain regulatory approval and commercialize our product candidates.**

Before we, or our collaborators, can receive regulatory approval for the commercial sale of our potential products, the FDA requires extensive preclinical safety testing and clinical trials to demonstrate their safety and efficacy. Significant delays in preclinical and clinical testing could materially impact our product development costs and delay regulatory approval of our product candidates. Completing clinical trials in a timely manner depends on, among other factors:

- obtaining regulatory approval to commence a trial;
- obtaining clinical materials;
- reaching agreement on acceptable clinical study terms with prospective sites and clinical research organizations;
- obtaining institutional review board approval to conduct a study at a prospective site; and
- recruiting patients to participate in a study.

**We rely on third parties to conduct our clinical trials, and their failure to perform their obligations in a timely and competent manner may delay development and commercialization of our product candidates.**

We used clinical research organizations in the United States, Asia and Europe to oversee our clinical trials for Sustol and we expect to use the same or similar organizations for our future clinical trials and pipeline programs. There are numerous alternative sources to provide these services; however, we may face delays outside of our control if these parties do not perform their obligations in a timely or competent fashion, or if we are forced to change service providers. Different cultural and operational issues in foreign countries could cause delays or unexpected problems with patient enrollment or with the data obtained from those locations. If we experience significant delays in the progress of our clinical trials or problems with the quality of data derived from clinical trials, the prospects for approval would decrease.

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**We have yet to demonstrate the full commercial viability of our delivery technology, and we cannot be certain that attainment of such a goal can be accomplished.**

Our bioerodible drug delivery technology is at an early stage of development. We may not be able to substantiate the capability of our drug delivery technology for a variety of reasons, including:

- selection of inappropriate therapeutic compound for delivery;
- selection of inappropriate use or application for the particular product candidate;
- failure to receive regulatory approval on a timely basis or at all; or
- difficulties with manufacturing in commercial quantities at an acceptable cost.

Successful development of delivery technologies requires significant preclinical and clinical testing prior to regulatory approval. Because of these scientific, regulatory and commercial hurdles, any program could be abandoned or otherwise fail, even after significant resources have been expended.

**If our suppliers and contract manufacturers fail to complete pre-commercialization manufacturing development activities for Sustol on a timely basis or fail to comply with stringent regulatory requirements, we will face delays in our ability to obtain regulatory approval for, and to commercialize, Sustol, and our costs will increase.**

We do not manufacture Sustol and do not currently plan to develop any capacity to do so. Instead, we have relied on third parties to manufacture and perform important pre-commercialization manufacturing development activities for Sustol. As part of the process for obtaining regulatory approval, we must demonstrate that the facilities, equipment and processes used to manufacture Sustol are capable of consistently producing a product that meets all applicable quality criteria, and that is comparable to the product that was used in our clinical trials. We must also provide the FDA with information regarding the validation of the manufacturing facilities, equipment and processes of our third-party suppliers and manufacturers, and data supporting the stability of Sustol. If our third-party suppliers and manufacturers are not in compliance with current Good Manufacturing Practice (cGMP) requirements, the approval of our marketing application may be delayed, existing product batches may be compromised, and we may experience delays in the availability of Sustol for commercial distribution.

For example, our most recent Complete Response Letter from the FDA regarding our NDA resubmission for Sustol stated that the NDA could not be approved due to, among other issues, deficiencies observed during an inspection of the facilities used by our third-party suppliers and manufacturers to produce Sustol. If the FDA is not satisfied with our response and any corrective actions taken by these third parties, we may be required to complete additional manufacturing development activities or provide other information to the FDA, which could cause substantial delays in obtaining regulatory approval for Sustol, increase our costs and have a material adverse effect on our business and financial condition.

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**We depend on contract manufacturers and collaborators for manufacturing our products and provide them with technical expertise on the manufacturing process; we also perform quality control testing of the product; if we and our contract manufacturers do not perform as expected, our revenue and customer relations will suffer.**

We do not own or operate manufacturing facilities for the production of clinical or commercial quantities of any product. Our ability to develop and commercialize any products we may develop will depend in part on our ability to manufacture, or arrange for collaborators or other parties to manufacture, our products at a competitive cost, in accordance with regulatory requirements, and in sufficient quantities for clinical testing and eventual commercialization. We do not intend to develop or acquire facilities to manufacture any of our product candidates for clinical trials or commercial purposes in the foreseeable future. We rely on a small number of third-party manufacturers to produce our compounds and expect to continue to do so to meet the preclinical and clinical requirements of our potential products and for all of our commercial needs, some of which are our sole source suppliers at present. We have no long-term agreements with any of these third parties. We may not be able to extend these agreements on satisfactory terms, or at all, and we may not be able to find a replacement contract manufacturer on satisfactory terms or on a timely basis. Additionally, difficult economic conditions may cause operational and financial problems for our third-party suppliers, resulting in their failure and disruption to our operations.

Further, we, along with our contract manufacturers and our collaborators, are required to comply with FDA requirements related to product testing, quality assurance, manufacturing and documentation. Our contract manufacturers, or our collaborators, may not be able to comply with the applicable FDA regulatory requirements. They may be required to pass an FDA pre-approval inspection for conformity with cGMPs before we can obtain approval to manufacture and will be subject to ongoing, periodic, unannounced inspection by the FDA and corresponding state agencies to ensure strict compliance with cGMP, and other applicable government regulations and corresponding foreign standards. If we and our contract manufacturers fail to achieve and maintain high manufacturing standards in compliance with cGMP regulations, we may experience manufacturing errors resulting in patient injury or death, product recalls or withdrawals, delays or interruptions of production or failures in product testing or delivery, delay or prevention of filing or approval of marketing applications for our products, cost overruns or other problems that could seriously harm our business. Not complying with FDA requirements could result in a Warning Letter or an enforcement action such as product seizure, recall, or injunction, prevent commercialization of our product candidates and impair our reputation and results of operations.

Any performance failure on the part of our contract manufacturers or by us could delay clinical development or regulatory approval of product candidates or commercialization of our future products, depriving us of potential product revenue and resulting in additional losses. In addition, our dependence on a third party for manufacturing may adversely affect our future profit margins and limit our ability to commercialize products on a timely and competitive basis. Our ability to replace an existing manufacturer may be difficult because the number of potential manufacturers is limited, and the FDA must approve any replacement manufacturer before we can begin manufacturing Sustol or any of our other product candidates. Such approval would require new testing and compliance inspections. It may be difficult or impossible for us to identify and engage a replacement manufacturer on acceptable terms in a timely manner, or at all.

Sustol or any of our other product candidates may be in competition with other products for access to the facilities of third parties. Consequently, Sustol or any of our other product candidates may be subject to manufacturing delays if collaborators or outside contractors give other companies' products greater priority than our products. For this and other reasons, our collaborators or third-party service providers may not be able to manufacture Sustol or any of our other product candidates in a

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cost-effective or timely manner. If not manufactured in a timely manner, the clinical development of any of our product candidates or their submission for regulatory approval could be delayed, and our ability to deliver products to market on a timely basis could be impaired or precluded.

To date, Sustol has been manufactured in small quantities for preclinical studies and clinical trials. If in the future Sustol or any of our product candidates are approved for commercial sale, we will need to manufacture our products in larger quantities. Significant scale-up of manufacturing may require additional process development and validation studies, which the FDA must review and approve. The commercial success of our products, including Sustol in the near-term, will be dependent upon the ability of our contract manufacturers to produce a product in commercial quantities at competitive costs of manufacture. The ability to do so cannot be presumed. Significant additional development work is required prior to any commercial launch of a product. In the case of Sustol, the high viscosity of the product creates particularly challenging factors relative to attainable production rates and cost of manufacture. If Sustol receives regulatory approval, we plan to scale-up manufacturing for Sustol in order to realize important economies of scale. These scale-up activities would take time to implement, require additional capital investment, process development and validation studies, and FDA approval. We cannot guarantee that we will be successful in achieving competitive manufacturing costs through such scale-up activities.

**If we are unable to establish sales and marketing capabilities or enter into agreements with third parties to sell and market any products we may develop, we may be unable to generate product revenue.**

We do not currently have a sales organization for the sales, marketing and distribution of pharmaceutical products. In order to commercialize any products, we must build our sales, marketing, distribution, managerial and other non-technical capabilities or make arrangements with third parties to perform these services. We have started to establish internal sales and marketing capabilities for Sustol, but may enter into agreements with third parties to sell and market other products we may develop. Although we have hired sales and marketing personnel with prior commercial experience, our company has no direct experience in developing, training or managing a marketing and sales force. The establishment and development of a sales force to market Sustol will be expensive and time consuming and could delay product launch, and we cannot be certain that we will be able to successfully develop this capacity. If we are unable to establish our sales and marketing capability or any other non-technical capabilities necessary to commercialize Sustol, we will need to contract with third parties to market and sell such products we may develop. If we are unable to establish adequate sales, marketing and distribution capabilities, whether independently or with third parties, we may not be able to generate product revenue and may not become profitable.

**If we fail to comply with continuing federal, state and foreign regulations, we could lose our approvals to market drugs, and our business would be seriously harmed.**

Following initial regulatory approval of any drugs we may develop, including Sustol, we will be subject to continuing regulatory review, including review of adverse drug experiences and clinical results that are reported after our drug products become commercially available. This would include results from any post-marketing tests or continued actions required as a condition of approval. The manufacturer and manufacturing facilities we use to make any of our drug candidates will be subject to periodic review and inspection by the FDA. If a previously unknown problem or problems with a product or a manufacturing and laboratory facility used by us is discovered, the FDA or foreign regulatory agency may impose restrictions on that product or on the manufacturing facility, including requiring us to withdraw the product from the market. Any changes to an approved product,

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including the way it is manufactured or promoted, often require FDA approval before the product, as modified, can be marketed. We and our contract manufacturers will be subject to ongoing FDA requirements for submission of safety and other post-market information. If we and our contract manufacturers fail to comply with applicable regulatory requirements, a regulatory agency may:

- issue warning letters;
- impose civil or criminal penalties;
- suspend or withdraw our regulatory approval;
- suspend or terminate any of our ongoing clinical trials;
- refuse to approve pending applications or supplements to approved applications filed by us;
- impose restrictions on our operations;
- close the facilities of our contract manufacturers; or
- seize or detain products or require a product recall.

Additionally, such regulatory review covers a company's activities in the promotion of its drugs, with significant potential penalties and restrictions for promotion of drugs for an unapproved use. Sales and marketing programs are under scrutiny for compliance with various mandated requirements, such as illegal promotions to healthcare professionals. We are establishing a sales force to market Sustol, which may include contracting with third parties to market and sell such products we may develop, and may be unable to ensure that our own employees as well as any third-party employees adhere to legal and regulatory requirements for product advertising and promotion. We are also required to submit information on our open and completed clinical trials to public registries and databases; failure to comply with these requirements could expose us to negative publicity, fines and penalties that could harm our business. If Sustol is approved, we will also be required to comply with the requirements to submit to governmental authorities information on payments to physicians and certain other third parties; failure to comply with these requirements could expose us to negative publicity, fines and penalties that could harm our business.

**If we are unable to recruit and retain skilled employees, we may not be able to achieve our objectives.**

We depend on a small number of key management and technical personnel. Retaining our current employees and recruiting qualified scientific personnel to perform future research and development and commercialization work will be critical to our success. While recent pharmaceutical and biotechnology industry layoffs have somewhat mitigated a usual shortage of skilled personnel in our industry, competition is always present for experienced scientists, and an inability to recruit or retain sufficient skilled personnel could result in delays to product development or approval, loss of sales and diversion of management resources. If we lose members of our senior management team, we may not be able to find suitable replacements and our business may be harmed as a result.

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**We face intense competition from other companies.**

Sustol is expected to face significant competition for the prevention of CINV, principally from Eisai's Aloxi (palonosetron). In addition to Aloxi, Sustol will compete with entrenched generic forms of granisetron (formerly marketed by Roche as Kytril) and ondansetron (formerly marketed by GlaxoSmithKline as Zofran). Generic versions of Aloxi may become available after its scheduled patent expiration date, which was recently extended to 2024. We are aware, however, of ongoing litigation challenging these patents and, if such challenges are successful, it is possible that generic versions of Aloxi could become available sooner. We are also aware of several companies that have developed, or are developing, both generic and new formulations of granisetron, including transdermal formulations such as ProStrakan's Sancuso® (granisetron transdermal patch), which was approved in 2008.

There are several companies that are developing new formulations of existing drugs across various therapeutic areas using novel drug delivery technologies. Many of these companies have substantially greater financial, research and development, manufacturing, sales and marketing and distribution resources and experience than we do. The following are some notable companies among drug delivery system developers: Alkermes, Inc., Durect Corporation, and Pacira Pharmaceuticals, Inc.

Smaller or early stage companies and research institutions may also prove to be significant competitors, particularly through collaborative arrangements with large and established pharmaceutical companies. We will also face competition from these parties in recruiting and retaining qualified scientific and management personnel, establishing clinical trial sites and patient registration for clinical trials, and acquiring and in-licensing technologies and products complementary to our programs or potentially advantageous to our business. If any of our competitors succeed in obtaining approval from the FDA or other regulatory authorities for their products sooner than we do or for products that are more effective or less costly than ours, our commercial opportunity could be significantly reduced. Major technological changes can happen quickly in the biotechnology and pharmaceutical industries, and the development of technologically improved or different products or drug delivery technologies may make our product candidates or platform technologies obsolete or noncompetitive.

**If we cannot establish pricing of our product candidates acceptable to the United States or foreign governments, insurance companies, managed care organizations and other payors, or arrange for favorable reimbursement policies, any product sales will be severely hindered.**

The continuing efforts of the United States and foreign governments, insurance companies, managed care organizations and other payors of health care costs to contain or reduce costs of health care may adversely affect our ability to generate adequate revenues and gross margins to make the products we develop commercially viable. Our ability to commercialize any product candidates successfully will depend in part on the extent to which governmental authorities, private health insurers and other organizations establish appropriate reimbursement levels for the cost of such products and related treatments.

In certain foreign markets, the pricing of prescription pharmaceuticals is subject to government control. In the United States, given recent federal and state government initiatives directed at lowering the total cost of health care, the U.S. Congress and state legislatures will likely continue to focus on health care reform, the cost of prescription pharmaceuticals and on the reform of the Medicare and Medicaid systems. The trend toward managed health care in the United States, which could significantly influence the purchase of health care services and products, as well as legislative proposals to reform health care, control pharmaceutical prices or reduce government insurance programs, may result in lower prices for our product candidates. While

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we cannot predict whether any legislative or regulatory proposals affecting our business will be adopted, the announcement or adoption of these proposals could have a material and adverse effect on our potential revenues and gross margins.

**Our business strategy includes the entry into additional collaborative agreements. We may not be able to enter into additional collaborative agreements or may not be able to negotiate commercially acceptable terms for these agreements.**

Our current business strategy includes the entry into additional collaborative agreements for the development and commercialization of our delivery technologies. The negotiation and consummation of these types of agreements typically involve simultaneous discussions with multiple potential collaborators and require significant time and resources from our officers, business development and research and development staff. In addition, in attracting the attention of pharmaceutical and biotechnology company collaborators, we compete with numerous other third parties with product opportunities as well as the collaborators' own internal product opportunities. We may not be able to consummate additional collaborative agreements, or we may not be able to negotiate commercially acceptable terms for these agreements. If we do not consummate additional collaborative agreements, we may consume money more rapidly on our product development efforts, continue to defer certain development activities or forego the exploitation of certain geographic territories, any of which could have a material adverse effect on our business.

**If we or our collaborators cannot arrange for adequate third-party reimbursement for our products, our future revenue will suffer.**

In both domestic and foreign markets, sales of our potential products, including Sustol, will depend in substantial part on the availability of adequate reimbursement from third-party payors such as government health administration authorities, private health insurers and other organizations. Third-party payors often challenge the price and cost-effectiveness of medical products and services and such pressure may increase in the future. Significant uncertainty exists as to the adequate reimbursement status of newly approved health care products. Any products we are able to successfully develop may not be reimbursable by third-party payors. In addition, our products may not be considered cost-effective and adequate third-party reimbursement may not be available to enable us to maintain price levels sufficient to realize a profit. Legislation and regulations affecting the pricing of pharmaceuticals may change before our products are approved for marketing and any such changes could further limit reimbursement. Reimbursement policies utilized by our collaborators or ourselves may be challenged by regulatory entities, with resultant fines, negative publicity and the need to implement changes that reduce the utilization of our products. If any products we develop do not receive adequate reimbursement, our revenue will be severely limited.

**Our inability to obtain specialized materials could slow down our product development process.**

Some of the critical materials and components used in producing Sustol are sourced from a single supplier. An interruption in supply of a key material could significantly delay our research and development process or increase our expenses.

Specialized materials must often be manufactured for the first time for use in drug delivery technologies, or materials may be used in the technologies in a manner different from their customary commercial uses. The quality of materials can be critical to the performance of a drug delivery technology, so a reliable source of a consistent supply of materials is important. Materials or components needed for our drug delivery technologies may be difficult to obtain on commercially reasonable terms, particularly when relatively small quantities are required or if the materials traditionally have not been used in pharmaceutical products.

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If we are unable to adequately protect or enforce our intellectual property rights or secure rights to third-party patents, we may lose valuable assets, experience reduced market share or incur costly litigation to protect our rights or our third-party collaborators may choose to terminate their agreements with us.

Our success will depend in part on our ability to obtain patents and maintain trade secret protection, as well as successfully defending these patents against challenges, while operating without infringing the proprietary rights of others. We have filed a number of U.S. patent applications on inventions relating to the composition of a variety of polymers, specific products, product groups and processing technology. At December 31, 2013, we had a total of 15 issued U.S. patents and an additional 38 issued (or registered) foreign patents. The patents on the bioerodible technologies expire between January 2016 and April 2026. The product Sustol is covered by patents owned by Heron Therapeutics, Inc. that are granted in the United States and in foreign countries. Currently, the product Sustol is covered by six patents issued in the United States and by seven patents issued in foreign countries including Brazil, Canada, EU, Japan, and Taiwan. U.S. patents covering Sustol have expiration dates ranging from January 2016 to September 2024; foreign patents covering Sustol have expiration dates ranging from January 2017 to September 2025. Granted patents include claims covering the product composition, methods of use and methods of preparation. Our existing patents may not cover future products, additional patents may not be issued, and current patents, or patents issued in the future, may not provide meaningful protection or prove to be of commercial benefit.

The patent positions of pharmaceutical companies, including ours, are uncertain and involve complex legal and factual questions. In addition, the coverage claimed in a patent application can be significantly reduced before the patent is issued. Consequently, our patent applications, or those that are licensed to us, may not issue into patents, and any issued patents may not provide sufficient protection for our product candidates or provide sufficient protection to afford us a commercial advantage against competitive technologies or may be held invalid if challenged or circumvented. Patent applications in the United States are maintained in confidence for at least 18 months after their filing. Consequently, we cannot be certain that the patent applications we are pursuing will lead to the issuance of any patent or be free from infringement or other claims from other parties. Our competitors may also independently develop products similar to ours or design around or otherwise circumvent patents issued to us or licensed by us. In addition, the laws of some foreign countries may not protect our proprietary rights to the same extent as U.S. laws.

We are party to collaborative agreements. These agreements subject us to obligations which must be fulfilled and require us to manage complex relationships with third parties. If we are unable to meet our obligations or manage our relationships with our collaborators under these agreements or enter into additional collaboration agreements or if our existing collaborations are terminated or not extended on terms as beneficial as we anticipate, our revenue may decrease. The loss or diminution of our intellectual property rights could result in a decision by our third-party collaborators to terminate their agreements with us. In addition, these agreements are generally complex and contain provisions that could give rise to legal disputes, including potential disputes concerning ownership of intellectual property and data under collaborations. Such disputes can lead to lengthy, expensive litigation or arbitration, requiring us to divert management time and resources to such dispute.

We also rely upon trade secrets, technical know-how and continuing technological innovation to develop and maintain our competitive position. We require our employees, consultants, advisors and collaborators to execute appropriate confidentiality and assignment-of-inventions agreements with us. These agreements typically provide that all materials and confidential information developed or made known to the individual during the course of the individual's relationship with us is to be kept confidential and not disclosed to third parties except in specific circumstances, and that all inventions arising out of the

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individual's relationship with us shall be our exclusive property. These agreements may be breached, and in some instances, we may not have an appropriate remedy available for breach of the agreements. Furthermore, our competitors may independently develop substantially equivalent proprietary information and techniques, reverse engineer our information and techniques, or otherwise gain access to our proprietary technology. We may be unable to meaningfully protect our rights in trade secrets, technical know-how and other non-patented technology. We may have to resort to litigation to protect our intellectual property rights, or to determine their scope, validity or enforceability. In addition, interference proceedings declared by the U.S. Patent and Trademark Office may be necessary to determine the priority of inventions with respect to our patent applications. Enforcing or defending our proprietary rights is expensive, could cause diversion of our resources and may not prove successful. In addition, courts outside the United States may be less willing to protect trade secrets. Costly and time-consuming litigation could be necessary to seek to enforce and determine the scope of our proprietary rights. Any failure to enforce or protect our rights could cause us to lose the ability to exclude others from using our technology to develop or sell competing products.

**We may infringe on the intellectual property rights of others, and any litigation could force us to stop developing or selling potential products and could be costly, divert management attention and harm our business.**

We must be able to develop products without infringing the proprietary rights of other parties. Because the markets in which we operate involve established competitors with significant patent portfolios, including patents relating to the composition of a variety of polymers, specific products, product groups and processing technology, it could be difficult for us to use our technologies or develop products without infringing the proprietary rights of others. We may not be able to design around the patented technologies or inventions of others and we may not be able to obtain licenses to use patented technologies on acceptable terms, or at all. If we cannot operate without infringing the proprietary rights of others, we will not be able to develop or commercialize some or all of our product candidates, and consequently will not be able to earn product revenue.

If we are required to defend ourselves in a lawsuit, we could incur substantial costs and the lawsuit could divert management attention, regardless of the lawsuit's merit or outcome. These legal actions could seek damages and seek to enjoin testing, manufacturing and marketing of the accused product or process. In addition to potential liability for significant damages, we could be required to obtain a license to continue to manufacture or market the accused product or process and any license required under any such patent may not be made available to us on acceptable terms, if at all.

Periodically, we review publicly available information regarding the development efforts of others in order to determine whether these efforts may violate our proprietary rights. We may determine that litigation is necessary to enforce our proprietary rights against others. Such litigation could result in substantial expense, regardless of its outcome, and may not be resolved in our favor.

**We are exposed to risks and increased expenses as a result of laws requiring filers to evaluate internal controls over financial reporting.**

Section 404 of the Sarbanes-Oxley Act of 2002 (Section 404) requires management to evaluate the effectiveness of our internal controls over financial reporting as of the end of each fiscal year and to include a management report assessing the effectiveness of our internal control over financing reporting in our annual report on Form 10-K for each fiscal year. Our independent registered public accounting firm is required to report on the effectiveness of our internal control over financial

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reporting. We and our independent registered public accounting firm may not be able to conclude on an ongoing basis that we have effective internal controls over financial reporting in accordance with Section 404. We have implemented an ongoing program to perform the system and process evaluation we believe to be necessary to comply with these requirements. However, we cannot assure you that we will be successful in our efforts. We expect to incur increased expense and to devote additional management resources to Section 404 compliance. Any failure to implement required new or improved controls, or difficulties encountered in the implementation or operation of these controls, could harm our operating results and cause us to fail to meet our financial reporting obligations, which could adversely affect our business and reduce our stock price.

Our management, including our Chief Executive Officer and Chief Financial Officer, does not expect that our internal controls over financial reporting will prevent all errors and all fraud. A control system, no matter how well designed and operated, can provide only reasonable, not absolute, assurance that the control system's objectives will be met. We cannot assure you that we or our independent registered public accounting firm will not identify a material weakness in our internal controls in the future, which would require management and our independent registered public accounting firm to evaluate our internal controls as ineffective. If our internal controls over financial reporting are not considered effective, we may experience a loss of public confidence, which could have an adverse effect on our business and on the price of our stock.

**Legislative actions, potential new accounting pronouncements and higher insurance costs are likely to impact our future financial position or results of operations.**

Future changes in financial accounting standards may cause adverse, unexpected fluctuations in the timing of the recognition of revenue or expenses and may affect our financial position or results of operations. New pronouncements and varying interpretations of pronouncements have occurred with frequency and may occur in the future, and we may make changes in our accounting policies in the future. Compliance with changing regulation of corporate governance and public disclosure may result in additional expenses. Changing laws, regulations and standards relating to corporate governance and public disclosure, including the Sarbanes-Oxley Act of 2002, the Dodd-Frank Act, new Securities and Exchange Commission (SEC) regulations and the Public Company Accounting Oversight Board pronouncements, are creating uncertainty for companies such as ours and insurance, accounting and auditing costs are increasing as a result of this uncertainty and other factors. We are committed to maintaining high standards of corporate governance and public disclosure. As a result, we intend to invest all reasonably necessary resources to comply with evolving standards, and this investment may result in increased general and administrative expenses and a diversion of management time and attention from revenue-generating activities to compliance activities.

**We could be exposed to significant product liability claims that could be time consuming and costly to defend, divert management attention and adversely impact our ability to obtain and maintain insurance coverage.**

The testing, manufacture, marketing and sale of our products involve an inherent risk that product liability claims will be asserted against us. Although we are insured against such risks up to an annual aggregate limit in connection with clinical trials and commercial sales of our products, our present product liability insurance may be inadequate and may not fully cover the costs of any claim or any ultimate damages we might be required to pay. Product liability claims or other claims related to our products, regardless of their outcome, could require us to spend significant time and money in litigation or to pay significant damages. Any successful product liability claim may prevent us from obtaining adequate product liability insurance in the future on commercially desirable or reasonable terms. In addition, product liability coverage may cease to be available in sufficient

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amounts or at an acceptable cost. An inability to obtain sufficient insurance coverage at an acceptable cost or otherwise to protect against potential product liability claims could prevent or inhibit the commercialization of our products. A product liability claim could also significantly harm our reputation and delay market acceptance of our products.

**Our use of hazardous materials could subject us to liabilities, fines and sanctions.**

Our laboratory and clinical testing sometimes involve use of hazardous, radioactive or otherwise toxic materials. We are subject to federal, state and local laws and regulations governing how we use, manufacture, handle, store and dispose of these materials. Although we believe that our safety procedures for handling and disposing of such materials comply in all material respects with all federal, state and local regulations and standards, there is always the risk of accidental contamination or injury from these materials. In the event of an accident, we could be held liable for any damages that result and such liability could exceed our financial resources. If we fail to comply with these regulations and standards or with the conditions attached to our operating licenses, the licenses could be revoked, and we could be subjected to criminal sanctions and substantial financial liability or be required to suspend or modify our operations. Compliance with environmental and other laws may be expensive and current or future regulations may impair our development or commercialization efforts.

**Earthquake damage to our facilities could delay our research and development and quality control testing efforts and adversely affect our business.**

Our facility in Redwood City, California, is located in a seismic zone, and there is the possibility of an earthquake, which could be disruptive to our operations and result in delays in our research and development efforts and supplies of Sustol, if approved. In the event of an earthquake, if our facilities or the equipment in our facilities are significantly damaged or destroyed, we may not be able to rebuild or relocate our facility or replace any damaged equipment in a timely manner and our business, financial condition and results of operations could be materially and adversely affected.

**Risks Related To Our Common Stock**

**The price of our common stock has been and may continue to be volatile and our reverse stock split may further increase volatility or cause a decline in value.**

The stock markets, in general, and the markets for drug delivery and pharmaceutical stocks, in particular, have experienced extreme volatility that has often been unrelated to the operating performance of particular companies. These broad market fluctuations may adversely affect the trading price of our common stock. In addition, the limited trading volume of our stock may contribute to its volatility.

Further, our stock price may be subject to additional volatility as a result of our Reverse Stock Split and relisting on the NASDAQ Capital Market. Following reverse splits, the prices of the stocks often trade below the immediate post-split value, resulting in a net loss in value for stockholders.

In the past, following periods of volatility in the market price of a particular company's securities, litigation has often been brought against that company. If litigation of this type is brought against us, it could be extremely expensive and divert management's attention and our company's resources.

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**Our certificate of incorporation, our bylaws and Delaware law contain provisions that could discourage another company from acquiring us and may prevent attempts by our stockholders to replace or remove our current management.**

Provisions of Delaware law, our certificate of incorporation and our bylaws may discourage, delay or prevent a merger or acquisition that stockholders may consider favorable, including transactions in which you might otherwise receive a premium for your shares. In addition, 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 or remove our board of directors. These provisions include authorizing the issuance of "blank check" preferred stock without any need for action by stockholders.

In addition, Section 203 of Delaware General Corporation Law may discourage, delay or prevent a change in control of our company by prohibiting stockholders owning in excess of 15% of our outstanding voting stock from merging or combining with us, unless certain approvals are obtained.

**Further concentration in stockholder ownership could influence strategic actions.**

Our directors, executive officers, principal stockholders and affiliated entities currently beneficially own or control a majority of our outstanding securities. Tang Capital Partners, LP and its affiliates' beneficial ownership in our common stock, as determined in accordance with Rule 13d-3 of the Exchange Act, was approximately 18% as of November 2013, excluding potential further concentration underlying outstanding warrants and our convertible note facility. Kevin C. Tang, the Managing Director of Tang Capital Management, LLC, the general partner of Tang Capital Partners, LP, is also chairman of our board of directors.

Such a concentration of common stock ownership could significantly influence corporate actions on various strategic matters, including, for example, receptivity to collaborations and merger or sale overtures.

**Future sales of our common stock may cause our stock price to decline.**

Our principal stockholders and affiliated entities hold a substantial number of shares of our common stock that they are able to sell in the public market. In addition, they currently own convertible notes and outstanding warrants for additional shares of our common stock. The exercise of these warrants, conversion of the notes or the sale by our current stockholders of a substantial number of shares, or the expectation that such exercises or sales may occur, could significantly reduce the market price of our common stock.

**Future utilization of net operating loss carry-forwards may be impaired due to recent changes in ownership.**

As discussed in Note 12 to the Financial Statements included in Item 8 of this Annual Report on Form 10-K, we believe our net operating losses and tax attributes may be subject to limitation under Section 382 of the Internal Revenue Code of 1986. As a result, our deferred tax assets, and related valuation allowance, have been reduced for the estimated impact of the net operating losses and credits that we currently estimate may expire unused. Utilization of our remaining net operating loss and research and development credit carry-forwards may still be subject to substantial annual limitations due to ownership change limitations provided by the Internal Revenue Code and similar state provisions for ownership changes after December 31, 2013, including those that may come in conjunction with future equity financings or market trades by our stockholders.

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**ITEM 1B. UNRESOLVED STAFF COMMENTS**

None.

**ITEM 2. PROPERTIES**

We lease 26,067 square feet of laboratory, office and warehouse space in Redwood City, California under a lease expiring on November 30, 2016. In addition, we currently lease 2,175 square feet of office space in Stamford, Connecticut on a month-to-month basis and 1,850 square feet of office space in San Diego, California with an initial term that expires in December 2014. The annual rent expense for all properties is approximately \$1.2 million. We believe our facilities are adequate and suitable for current and anticipated needs.

**ITEM 3. LEGAL PROCEEDINGS**

While we are not currently a party to any material pending legal proceedings, from time to time we are named as a party to lawsuits in the normal course of its business. Litigation, in general, and intellectual property litigation, in particular, can be expensive and disruptive to normal business operations. Moreover, the results of legal proceedings are difficult to predict.

**ITEM 4. MINE SAFETY DISCLOSURES**

Not applicable.

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PART II

**ITEM 5. MARKET FOR REGISTRANT'S COMMON STOCK, RELATED STOCKHOLDER MATTERS AND ISSUER PURCHASES OF EQUITY SECURITIES**

Shares of our common stock traded on the OTC Bulletin Board under the symbol APPA.OB until January 10, 2014. Our shares traded under the symbol APPAD following our one-for-twenty reverse stock split (Reverse Stock Split). On January 23, 2014, our common stock was approved for listing and began trading on the NASDAQ Capital Market under the symbol HRTX.

As of February 28, 2014, there were 171 holders of record of our common stock.

We have never paid cash dividends and do not anticipate paying cash dividends in the foreseeable future. The following table shows the range of high and low sales prices for our common stock for the periods indicated as reported by the OTC Bulletin Board, adjusted for the Reverse Stock Split.

<b>2013</b>	<b>High</b>	<b>Low</b>
First Quarter	\$17.80	\$5.60
Second Quarter	9.80	6.00
Third Quarter	10.00	6.40
Fourth Quarter	10.00	6.60
<b>2012</b>	<b>High</b>	<b>Low</b>
First Quarter	\$ 8.60	\$ 3.40
Second Quarter	15.00	7.00
Third Quarter	15.60	11.00
Fourth Quarter	13.40	9.60

On February 28, 2014, the closing sale price of our common stock was \$14.03 per share.

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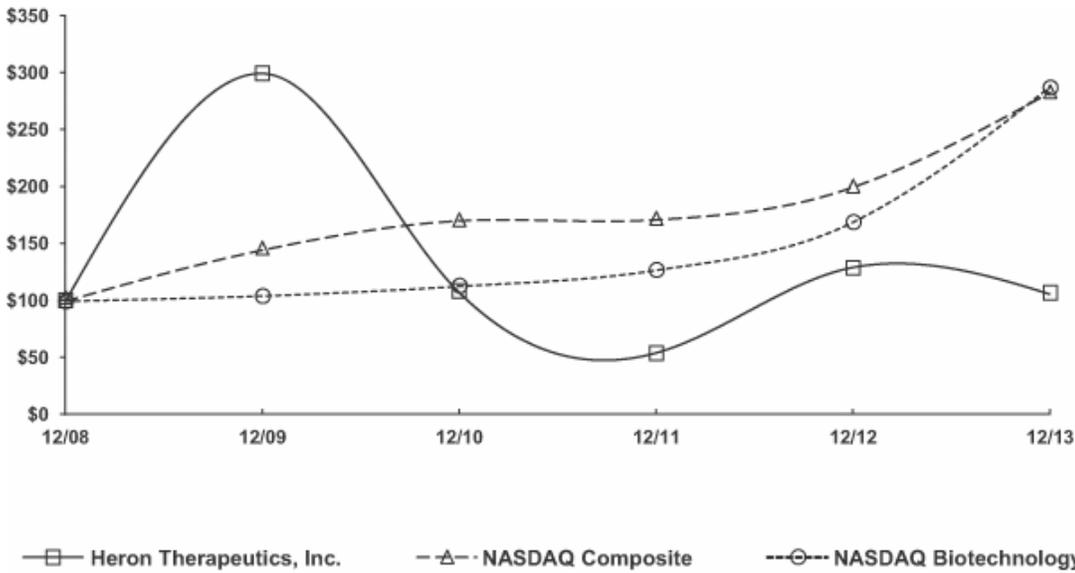
**Stock Performance Graph**

The following is not deemed "filed" with the Securities and Exchange Commission and is not to be incorporated by reference into any filing we make under the Securities Act of 1933, as amended, whether made before or after the date hereof and irrespective of any general incorporation by reference language in such filing.

The following graph shows the value of an investment of \$100 on December 31, 2008 in Heron Therapeutics, Inc. common stock, the NASDAQ Composite Index (U.S.) and the NASDAQ Biotechnology Index. All values assume reinvestment of the pretax value of dividends paid by companies included in these indices and are calculated as of December 31 of each year. Our common stock is traded on the NASDAQ Capital Market. The comparisons shown in the graph are based upon historical data and we caution that the stock price performance shown in the graph is not indicative of, nor intended to forecast, the potential future performance of our stock.

**COMPARISON OF 5 YEAR CUMULATIVE TOTAL RETURN\***

Among Heron Therapeutics, Inc., the NASDAQ Composite Index, and the NASDAQ Biotechnology Index,



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**ITEM 6. SELECTED FINANCIAL DATA**

The following selected financial data should be read in conjunction with the “Management’s Discussion and Analysis of Financial Condition and Results of Operations” and the financial statements and notes thereto, included in Item 8 of this Annual Report on Form 10-K. The financial data does not purport to indicate results of operations as of any future date or for any future period.

For the Years Ended December 31,  
(In thousands, except per share data)

	<b>2013</b>	<b>2012</b>	<b>2011</b>	<b>2010</b>	<b>2009</b>
<b>STATEMENT OF OPERATIONS DATA</b>					
Revenue:					
Contract revenue	\$ —	\$ —	\$ 646	\$ 1,301	\$ 1,261
Operating expenses:					
Research and development	32,780	15,174	8,207	7,264	7,796
General and administrative	21,677	8,657	3,501	3,971	3,707
Operating loss	(54,457)	(23,831)	(11,062)	(9,934)	(10,242)
Gain on sale of interest in royalties	—	—	—	2,500	—
Interest and other income (expense), net	(826)	(599)	(373)	238	24
Loss from continuing operations before income taxes	(55,283)	(24,430)	(11,435)	(7,196)	(10,218)
Benefit from income taxes	—	—	—	—	122
Net loss from continuing operations	(55,283)	(24,430)	(11,435)	(7,196)	(10,096)
Gain (loss) from discontinued operations <sup>(1)</sup>	—	1,082	(379)	(150)	68
Net loss	\$(55,283)	\$(23,348)	\$(11,814)	\$(7,346)	\$(10,028)
Basic and diluted net loss per common share:					
Loss from continuing operations	\$ (3.42)	\$ (2.00)	\$ (1.90)	\$ (3.63)	\$ (6.19)
Net loss	\$ (3.42)	\$ (1.91)	\$ (1.96)	\$ (3.70)	\$ (6.15)
Weighted-average common shares outstanding used to calculate basic and diluted net loss per common share					
	16,163	12,223	6,013	1,984	1,631

<sup>(1)</sup> Loss from discontinued operations represents the loss attributable to our cosmeceutical and toiletries business that was sold to RP Scherer on July 25, 2000. See Note 10 to the Financial Statements included in Item 8 of this Annual Report on Form 10-K.

December 31,	<b>2013</b>	<b>2012</b>	<b>2011</b>	<b>2010</b>	<b>2009</b>
<b>BALANCE SHEET DATA</b>					
Cash, cash equivalents and marketable securities	\$ 72,287	\$ 53,506	\$ 17,974	\$ 2,109	\$ 7,593
Working capital	65,933	49,936	14,547	941	6,426
Total assets	75,937	55,972	19,445	2,911	8,951
Long-term liabilities	—	—	—	35	268
Accumulated deficit	(238,870)	(183,587)	(160,239)	(148,425)	(141,079)
Stockholders’ equity	68,945	51,818	15,752	1,316	6,796

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**ITEM 7. MANAGEMENT'S DISCUSSION AND ANALYSIS OF FINANCIAL CONDITION AND RESULTS OF OPERATIONS**

This Annual Report on Form 10-K contains "forward-looking statements" as defined by the Private Securities Litigation Reform Act of 1995. These forward-looking statements involve risks and uncertainties, including uncertainties associated with: the progress of our research, development and clinical programs; estimates of the timing of our resubmission of our NDA; the timing of regulatory approval and commercial introduction of Sustol and future product candidates; the timing of market introduction of Sustol and future product candidates; our ability to market, commercialize and achieve market acceptance for Sustol and other future product candidates; our ability to establish collaborations for our technology, Sustol and other future product candidates; our ability to develop other drug candidates utilizing our Biochronomer polymer-based drug delivery platform; our estimates for future performance; our estimates regarding our capital requirements and our needs for and ability to obtain additional financing; our ability to protect or enforce our intellectual property rights; volatility in the trading price of our common stock; and other risks and uncertainties identified in our filings with the Securities and Exchange Commission. We caution investors that forward-looking statements reflect our analysis only on their stated date. We do not intend to update them except as required by law.

Unless otherwise noted, all information in this Item 7 regarding share amounts of our common stock, prices per share of our common stock and loss per share has been adjusted to reflect the application of the one-for-twenty reverse stock split of our common stock that we effected on January 13, 2014 (Reverse Stock Split) (see Note 15 to the Financial Statements included in Item 8 of this Annual Report on Form 10-K), on a retroactive basis.

**Overview**

We are a specialty pharmaceutical company developing products using our proprietary Biochronomer™ polymer-based drug delivery platform. This drug delivery platform is designed to improve the therapeutic profile of injectable pharmaceuticals by converting them from products that must be injected once or twice per day to products that need to be injected only once every one or two weeks.

Our lead product candidate, Sustol, is being developed for the prevention of both acute CINV for patients undergoing both moderately or highly emetogenic chemotherapy and for the prevention of delayed CINV for patients undergoing moderately emetogenic chemotherapy. One of the most debilitating side effects of cancer chemotherapy, CINV is a leading cause of premature discontinuation of treatment. There is only one injectable 5-HT<sub>3</sub> antagonist approved for the prevention of delayed-onset CINV, so this indication represents an area of particular unmet medical need. Sustol contains the 5-HT<sub>3</sub> antagonist granisetron formulated in our proprietary Biochronomer drug delivery system, which allows therapeutic drug levels to be maintained for five days with a single subcutaneous injection. This five-day range is designed to cover the delayed phase of CINV. Granisetron was selected for Sustol because it is widely prescribed by physicians based on a well-established record of safety and efficacy.

In May 2009, we filed the original NDA seeking approval for Sustol with the U.S. Food and Drug Administration (FDA). The FDA issued a Complete Response Letter for Sustol in March 2010. We met with the FDA in February and March 2011 to clarify the work needed to address the issues identified in the letter. In September 2012, we resubmitted our NDA for Sustol and,

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in March 2013, we received a second Complete Response Letter, which identified several remaining issues that need to be addressed prior to approval of the Sustol NDA. We are currently working on addressing these issues and expect to resubmit the Sustol NDA in the second quarter of 2014.

We own the worldwide rights to Sustol and are in the early stages of building the commercial infrastructure necessary to commercialize Sustol in the U.S. on our own.

Additionally, we are exploring the potential use of our Biochronomer polymer with other drugs and intend to pursue the clinical development on one or more other drug candidates based on our proprietary delivery platform.

**Critical Accounting Policies and Estimates**

Our accounting policies are more fully described in Note 2 of the Financial Statements included in Item 8 of this Annual Report on Form 10-K. The preparation of financial statements in conformity with U.S. generally accepted accounting principles requires our management to make estimates and assumptions about future events that affect the amounts reported in the financial statements and accompanying notes. Actual results could differ significantly from those estimates. We believe the following policies to be critical to understanding our financial condition, results of operations and expectations for 2014. These policies require management to make significant estimates, assumptions and judgments about matters that are inherently uncertain.

• **Revenue Recognition**

Our revenue arrangements with multiple deliverables are divided into separate units of accounting if certain criteria are met, including whether the delivered item has stand-alone value to the customer and whether there is objective and reliable evidence of the fair value of the undelivered items. The consideration we receive is allocated among the separate units based on their respective fair values, and the applicable revenue recognition criteria are considered separately for each of the separate units. Management's estimate of the selling price is considered when fair value is not determinable. Advance payments received in excess of amounts earned are classified as deferred revenue until earned.

• **Contract Revenue**

Our licensing agreements generally provide for a non-refundable license fee. The license agreements provide for us to earn future revenue through royalty payments. These non-refundable license fees are generally initially reported as deferred revenue and recognized as revenue over an appropriate period, depending on the license. Revenue recognized from deferred license fees is classified as Contract Revenue in our statements of operations.

Contract revenue also relates to research and development arrangements that generally provide for us to invoice research and development fees based on full-time equivalent hours for each project. Revenue from these arrangements is recognized as the related development services are rendered. This revenue approximates the costs incurred.

• **Clinical Trial Accruals**

Our expenses related to clinical trials are based on estimates of the services received and efforts expended pursuant to contracts with multiple research institutions and clinical research organizations that conduct and manage clinical trials on our

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benefit. Since the invoicing related to these services does not always coincide with our financial statement close process, we must estimate the level of services performed and fees incurred in determining the accrued clinical trial costs. The financial terms of these agreements are subject to negotiation and vary from contract to contract, which may result in uneven payment flows. Payments under the contracts depend on factors such as the successful enrollment of patients or achievement of certain events or the completion of portions of the clinical trial or similar conditions. Expenses related to clinical trials generally are accrued based on the level of patient enrollment and services performed by the clinical research organization or related service provider according to the protocol. We monitor patient enrollment levels and related activity to the extent possible and adjust our estimates accordingly. Historically these estimates have been reasonably accurate and no material adjustments have had to be made.

• ***Income Taxes***

We make certain estimates and judgments in determining income tax expense for financial statement purposes. These estimates and judgments occur in the calculation of certain tax assets and liabilities, which arise from differences in the timing of recognition of revenue and expense for tax and financial statement purposes. As part of the process of preparing our financial statements, we are required to estimate our income taxes in each of the jurisdictions in which we operate. This process involves us estimating our current tax exposure under the most recent tax laws and assessing temporary differences resulting from differing treatment of items for tax and financial statement purposes.

We assess the likelihood that we will be able to recover our deferred tax assets. We consider all available evidence, both positive and negative, including our historical levels of income and losses, expectations and risks associated with estimates of future taxable income and ongoing prudent and feasible tax planning strategies in assessing the need for a valuation allowance. If we do not consider it more likely than not that we will recover our deferred tax assets, we will record a valuation allowance against the deferred tax assets that we estimate will not ultimately be recoverable. At December 31, 2013, we believed that the amount of our deferred income taxes would not be ultimately recovered. Accordingly, we recorded a full valuation allowance for deferred tax assets.

Additionally, we believe that our deferred tax assets may have been limited in accordance with a provision of the Internal Revenue Code of 1986, whereby net operating loss and tax credit carryforwards available for use in a given period are limited upon the occurrence of certain events, including a significant change in ownership interests. As a result, our deferred tax assets and related valuation allowance were reduced for the estimated impact of the net operating losses and credits that may expire unused (see Note 12 to the Financial Statements included in Item 8 of this Annual Report on Form 10-K).

Should there be a change in our ability to recover our deferred tax assets, we would recognize a benefit to our tax provision in the period in which we determine that it is more likely than not that we will recover our deferred tax assets.

• ***Stock-Based Compensation***

We account for share-based payment arrangements in accordance with ASC 718, *Compensation – Stock Compensation* and ASC 505-50, *Equity – Equity Based Payments to Non-Employees*, which requires the recognition of compensation expense, using a fair-value based method, for all costs related to share-based payments including stock options, restricted

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stock awards and stock issued under the employee stock purchase plan. These standards require companies to estimate the fair value of share-based payment awards on the date of the grant using an option-pricing model (see Note 8 to the Financial Statements included in Item 8 of this Annual Report on Form 10-K).

**Results of Operations for the years ended December 31, 2013, 2012 and 2011**

The following sets forth the statement of operations data and percentage changes as compared to the prior years (dollar amounts are presented in thousands):

	For the Years Ended December 31,			\$ Change		% Change	
	2013	2012	2011	2013/2012	2012/2011	2013/2012	2012/2011
Total revenue	\$ —	\$ —	\$ 646	\$ —	\$ (646)	nm	(100%)
Research and development	32,780	15,174	8,207	17,606	6,967	116%	85%
General and administrative	21,677	8,657	3,501	13,020	5,156	150%	147%
Interest expense, net	(826)	(599)	(373)	(227)	(226)	38%	61%
Gain (loss) from discontinued operations	—	1,082	(379)	(1,082)	1,461	(100%)	(385%)

**Revenue**

Contract revenue decreased in 2012 by \$0.6 million, or 100%, compared to 2011, as no revenue was earned in 2012. Our contract revenue for 2011 was derived from an agreement with Merial Limited (Merial) that we entered into in September 2009 for a long-acting pain management product for companion animals. In May 2011, we received notice of termination from Merial.

**Research and Development**

Research and development expense in 2013 increased by \$17.6 million, or 116%, compared to 2012. Compared to 2012, headcount-related costs, including stock compensation expense, and project spending for Sustol, including validating our manufacturing processes and increasing the scale of production, were higher in 2013 as we worked to address the issues raised by the FDA in the 2013 Complete Response Letter. Research and development expense for the year 2014 is expected to be higher, as compared to 2013, as we conduct clinical studies on Sustol and continue work on new product development and manufacturing development.

Research and development expense in 2012 increased by \$7.0 million, or 85%, compared to 2011. Compared to 2011, headcount-related costs, including stock compensation expense, and project spending for Sustol were higher in 2012 as we worked to address the issues raised by the FDA in the Complete Response Letter received in 2010.

The scope and magnitude of future research and development expense is difficult to predict given the number of studies that will need to be conducted for any of our potential products. In general, biopharmaceutical development involves a series of steps, beginning with identification of a product candidate and includes proof-of-concept in animals and Phase 1, 2 and 3

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clinical studies in humans. Each step of this process is typically more expensive than the previous one, so success in development results in increasing expenditures. We are exploring the potential use of our Biochronomer polymer with other drugs and intend to pursue the clinical development of one or more other drug candidates based on our proprietary delivery platform.

The major components of research and development expenses were as follows (in thousands):

For the years ended December 31,	2013	2012	2011
Internal research and development costs	\$ 19,403	\$ 6,784	\$ 4,663
External development costs:			
Sustol (CINV product)	12,577	8,030	3,454
External general technology development costs	800	360	90
	<u>\$32,780</u>	<u>\$15,174</u>	<u>\$ 8,207</u>

Internal research and development costs consist of employee salaries and benefits, including stock compensation expense, laboratory supplies, depreciation and allocation of overhead. External development costs include expenditures on polymer development and manufacturing, which are performed on our behalf by third parties.

**General and Administrative**

General and administrative expenses increased in 2013 by \$13.0 million, or 150%, compared to 2012. The increase in 2013 was primarily due to headcount-related costs, including stock compensation expense, consulting costs, professional fees, market research and pre-commercialization activities. General and administrative expense for the year 2014 is expected to be higher as compared to 2013, due to increased costs to support anticipated commercialization activities.

General and administrative expenses increased in 2012 by \$5.2 million, or 147%, compared to 2011. The increase in 2012 was primarily due to headcount-related costs, including stock compensation expense, consulting costs and professional fees related to the NDA resubmission and pre-commercialization activities.

General and administrative expenses consist primarily of salaries and related expenses, professional fees, directors' fees, investor relations costs, pre-commercialization costs, insurance expense and related overhead cost allocation.

**Interest Expense, net**

Interest expense, net of \$0.8 million and \$0.6 million for the years ended December 31, 2013 and 2012 consists primarily of interest expense and amortization of debt discount related to the convertible note financings. Interest expense, net of \$0.4 million for the fiscal year 2011 also included debt issuance costs related to the convertible note financing.

**Discontinued Operations**

On July 25, 2000, we completed the sale of certain technology rights for our cosmeceutical and toiletry business to RP Scherer Corporation (RP Scherer), a subsidiary of Cardinal Health, Inc. Under the terms of the agreement, we guaranteed a minimum

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gross profit percentage on RP Scherer's combined sales of products to Ortho Dermatologics (Ortho) and Dermik Laboratories, Inc. (Dermik) (Gross Profit Guaranty), both of which were acquired by Valeant Pharmaceuticals in July 2011. The Gross Profit Guaranty expense totaled \$944,000 for the first seven guaranty years. Effective March 2007, in conjunction with a sale of assets by RP Scherer's successor company to an Amcol International subsidiary (Amcol), a new agreement was signed between us and Amcol to provide continuity of product supply to Ortho and Dermik. We had previously recorded a liability of \$1.1 million related to the amount that Amcol asserted was due under the Gross Profit Guaranty. In February 2013, an arbitrator ruled that no additional amounts were owed. This event qualified as an adjusting event under ASC 855, Subsequent Event, and in light of the arbitrator's decision in February 2013, which was final and binding, we reversed this accrual as of December 31, 2012. Gain (loss) from discontinued operations represents primarily the gain (loss) attributable to the Gross Profit Guaranty associated with the sale of our cosmeceutical and toiletry business.

***Liquidity and Capital Resources***

We had cash and cash equivalents of \$72.3 million at December 31, 2013. Cash and cash equivalents increased by \$18.8 million at December 31, 2013, as compared to December 31, 2012, primarily due to net cash proceeds received from the November 2013 common stock offering, which was partially offset by cash used in operations and equipment purchases.

Net cash used in operating activities for the year ended December 31, 2013 was \$40.8 million, compared to net cash used in operating activities of \$17.0 million for the year ended December 31, 2012. The \$23.8 million increase in net cash used in operating activities was primarily due to increases in research and development expenses and increased general and administrative expenses to support Sustol.

Net cash used in operating activities for the year ended December 31, 2012 was \$17.0 million, compared to net cash used in operating activities of \$7.7 million for the year ended December 31, 2011. The \$9.3 million increase in net cash used was primarily due to increases in research and development expenses to support Sustol and increased general and administrative expenses.

Net cash used in investing activities was \$1.7 million, \$1.0 million and \$0.5 million for the years ended December 31, 2013, 2012 and 2011, respectively. The increases in net cash used in investing activities were for purchases of equipment.

Our financing activities provided us with \$61.2 million, \$53.5 million and \$24.1 million for the years ended December 31, 2013, 2012 and 2011, respectively. The increase in cash provided by financing activities of \$7.7 million in 2013, as compared to 2012, was primarily due to proceeds received from the November 2013 common stock offering and stock option exercises. The increase in cash provided by financing activities of \$29.4 million in 2012, as compared to 2011, was primarily due to proceeds received from the July 2012 private placement of \$50.5 million and additional proceeds received from the issuance of convertible notes of \$3.0 million, which were partially offset by the April 2011 convertible note financing and the June 2011 private placement.

Historically, we have financed our operations, including technology and product research and development, primarily through sales of our common stock and other securities, royalties received on sales of Retin-A Micro and Carac, the sale of our rights to royalties on sales of Retin-A Micro and Carac, income from collaborative research and development fees, proceeds received from the sales of our Analytical Standards division and our cosmeceutical and toiletry business and interest earned on short-term investments.

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In April 2011, we entered into definitive agreements for a convertible note financing of up to \$4.5 million, which served as a bridge loan to fund the Company's operations until additional financing was secured. The initial funding from the bridge loan was approximately \$1.3 million, net of issuance costs, whereby \$1.5 million aggregate principal amount of convertible notes were issued. In May 2012, the purchasers exercised their rights to purchase the remaining \$3.0 million aggregate principal amount of convertible notes, and we received the additional \$3.0 million of proceeds.

In July 2011, we closed a unit financing where each unit consisted of one share of common stock and a warrant to purchase 0.5 of a share of common stock; whereby the Company received approximately \$22.8 million of proceeds, net of issuance costs.

In July 2012, we closed a common stock financing whereby the Company received approximately \$50.5 million of proceeds, net of issuance costs.

In November 2013, we closed a common stock offering whereby the Company received approximately \$57.8 million of proceeds, net of issuance costs.

We believe that our current cash resources are sufficient to fund essential operations into 2015; provided, however, that if we pursue additional clinical studies or commercially launch Sustol prior to 2015, we will need to raise additional capital. Our capital requirements going forward will depend on numerous factors, including: an approval decision by the FDA with respect to Sustol; the timing of and cost associated with the commercial launch of Sustol, if approved; the degree of commercial success of Sustol; the number and characteristics of product development programs we pursue and the pace of each program including the timing of clinical trials to expand our pipeline of sustained-release products; the scope, rate of progress, results and costs of preclinical testing and clinical trials, including our planned Phase 3 clinical trial of Sustol in the delayed HEC indication; the time, cost and outcome involved in seeking other regulatory approvals; scientific progress in our research and development programs; the magnitude and scope of our research and development programs; our ability to establish and maintain strategic collaborations or partnerships for research, development, clinical testing, manufacturing and marketing of our product candidates; the cost and timing of establishing sales, marketing and distribution capabilities for a specialty sales force if we commercialize other products independently; the cost of establishing clinical and commercial supplies of our product candidates and any products that we may develop; and general market conditions.

We may not be able to raise sufficient additional capital when we need it on favorable terms, or at all. The sale of additional equity in the future may be dilutive to our stockholders. If we are unable to obtain adequate funds on reasonable terms, we may be required to curtail operations significantly or to obtain funds by entering into financing, supply or collaboration agreements on unattractive terms.

**Contractual Obligations**

Below is a summary of fixed payments related to certain contractual obligations (in thousands). This table excludes amounts already recorded on our balance sheet as current liabilities as of December 31, 2013.

	<b>Total</b>	<b>Less than 1 year</b>	<b>2 to 3 years</b>	<b>4 to 5 years</b>	<b>More than 5 years</b>
Other operating leases <sup>(1)</sup>	\$ 2.4	\$ 0.9	\$ 1.5	\$ —	\$ —

<sup>(1)</sup> See Note 7 to the Financial Statements included in Item 8 of this Annual Report on Form 10-K.

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The holders of the convertible notes issued in April 2011 and May 2012 may require prepayment of such notes at any time, at the holders' option. See Note 6 to the Financial Statements included in Item 8 of this Annual Report on Form 10-K.

**Off-Balance-Sheet Arrangements**

As of December 31, 2013, we did not have any off-balance-sheet arrangements, as defined in Item 303(a)(4)(ii) of Regulation S-K.

**Recent Accounting Pronouncements**

Recent accounting pronouncements are disclosed in Note 2 to the Financial Statements included in Item 8 of this Annual Report on Form 10-K.

**ITEM 7A. QUANTITATIVE AND QUALITATIVE DISCLOSURE ABOUT MARKET RISK**

Our exposure to market risk is confined to our cash and cash equivalents. We did not hold any marketable securities at December 31, 2013.

Our debt obligations consist of our convertible debt, which carries a fixed interest rate and, as a result, we are not exposed to interest rate risk on our convertible debt.

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**ITEM 8. FINANCIAL STATEMENTS AND SUPPLEMENTARY DATA**

**Report of Independent Registered Public Accounting Firm**

The Board of Directors and Stockholders of  
Heron Therapeutics, Inc.

We have audited the accompanying balance sheets of Heron Therapeutics, Inc. as of December 31, 2013 and 2012, and the related statements of operations, stockholders' equity and cash flows for each of the three years in the period ended December 31, 2013. These financial statements are the responsibility of the Company's management. Our responsibility is to express an opinion on these financial statements based on our audits.

We conducted our audits in accordance with the standards of the Public Company Accounting Oversight Board (United States). Those standards require that we plan and perform the audit to obtain reasonable assurance about whether the financial statements are free of material misstatement. An audit also includes examining, on a test basis, evidence supporting the amounts and disclosures in the financial statements, assessing the accounting principles used and significant estimates made by management, as well as evaluating the overall presentation of the financial statements. We believe that our audits provide a reasonable basis for our opinion.

In our opinion, the financial statements audited by us present fairly, in all material respects, the financial position of Heron Therapeutics, Inc. at December 31, 2013 and 2012, and the results of its operations and its cash flows for each of the three years in the period ended December 31, 2013, in conformity with U.S. generally accepted accounting principles.

We also have audited, in accordance with the standards of the Public Company Accounting Oversight Board (United States), Heron Therapeutics, Inc.'s internal control over financial reporting as of December 31, 2013, based on criteria established in *Internal Control-Integrated Framework* issued by the Committee of Sponsoring Organizations of the Treadway Commission and our report dated March 4, 2014 expressed an unqualified opinion thereon.

/s/ OUM & Co. LLP

San Francisco, California  
March 4, 2014

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**Report of Independent Registered Public Accounting Firm**

The Board of Directors and Stockholders of  
Heron Therapeutics, Inc.

We have audited Heron Therapeutics, Inc.'s internal control over financial reporting as of December 31, 2013, based on criteria established in *Internal Control – Integrated Framework (1992)* issued by the Committee of Sponsoring Organizations of the Treadway Commission (the COSO criteria). Heron Therapeutics, Inc.'s management is responsible for maintaining effective internal control over financial reporting and for its assessment of the effectiveness of internal control over financial reporting included in the accompanying *Item 9A, Management's Report on Internal Control Over Financial Reporting*. Our responsibility is to express an opinion on the Company's internal control over financial reporting based on our audit.

We conducted our audit in accordance with the standards of the Public Company Accounting Oversight Board (United States). Those standards require that we plan and perform the audit to obtain reasonable assurance about whether effective internal control over financial reporting was maintained in all material respects. Our audit included obtaining an understanding of internal control over financial reporting, assessing the risk that a material weakness exists, and testing and evaluating the design and operating effectiveness of internal control based on the assessed risk. Our audit also included performing such other procedures as we considered necessary in the circumstances. We believe that our audit provides a reasonable basis for our opinion.

A company's internal control over financial reporting is a process designed to provide reasonable assurance regarding the reliability of financial reporting and the preparation of financial statements for external purposes in accordance with generally accepted accounting principles. A company's internal control over financial reporting includes those policies and procedures that (1) pertain to the maintenance of records that, in reasonable detail, accurately and fairly reflect the transactions and dispositions of the assets of the company; (2) provide reasonable assurance that transactions are recorded as necessary to permit preparation of financial statements in accordance with generally accepted accounting principles, and that receipts and expenditures of the company are being made only in accordance with authorizations of management and directors of the company; and (3) provide reasonable assurance regarding prevention or timely detection of unauthorized acquisition, use, or disposition of the company's assets that could have a material effect on the financial statements.

Because of its inherent limitations, internal control over financial reporting may not prevent or detect misstatements. Also, projections of any evaluation of effectiveness to future periods are subject to the risk that controls may become inadequate because of changes in conditions, or that the degree of compliance with the policies or procedures may deteriorate.

In our opinion, Heron Therapeutics, Inc. maintained, in all material respects, effective internal control over financial reporting as of December 31, 2013, based on the COSO criteria.

We also have audited, in accordance with the standards of the Public Company Accounting Oversight Board (United States), the balance sheets of Heron Therapeutics, Inc. as of December 31, 2013 and 2012, and the related statements of operations, stockholders' equity, and cash flows for each of the three years in the period ended December 31, 2013, and our report dated March 4, 2014, expressed an unqualified opinion thereon.

/s/ OUM & Co. LLP

San Francisco, California  
March 4, 2014

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HERON THERAPEUTICS, INC.  
BALANCE SHEETS  
(in thousands, except par value)

December 31,	2013	2012
<b>ASSETS</b>		
Current Assets:		
Cash and cash equivalents	\$ 72,287	\$ 53,506
Prepaid expenses and other current assets	638	584
Total Current Assets	72,925	54,090
Property and equipment, net	2,882	1,752
Other long-term assets	130	130
Total Assets	\$ 75,937	\$ 55,972
<b>LIABILITIES AND STOCKHOLDERS' EQUITY</b>		
Current Liabilities:		
Accounts payable	\$ 1,264	\$ 1,912
Accrued expenses	4,703	1,750
Convertible notes payable to related parties, net of discount	1,025	492
Total Current Liabilities	6,992	4,154
Commitments and Contingencies (Note 7)		
Stockholders' Equity:		
Preferred stock, 2,500 shares authorized; none issued or outstanding at December 31, 2013 and 2012	—	—
Common stock, \$0.01 par value, 75,000 shares authorized; 23,572 shares and 15,111 shares issued and outstanding at December 31, 2013 and 2012, respectively	237	152
Additional paid-in capital	307,578	235,253
Accumulated deficit	(238,870)	(183,587)
Total Stockholders' Equity	68,945	51,818
Total Liabilities and Stockholders' Equity	\$ 75,937	\$ 55,972

See accompanying Notes to Financial Statements.

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HERON THERAPEUTICS, INC.  
STATEMENTS OF OPERATIONS  
(in thousands, except per share amounts)

For the Years Ended December 31,	2013	2012	2011
<b>REVENUE</b>			
Contract revenue	\$ —	\$ —	\$ 646
<b>OPERATING EXPENSES</b>			
Research and development	32,780	15,174	8,207
General and administrative	21,677	8,657	3,501
Total operating expenses	54,457	23,831	11,708
Operating loss	(54,457)	(23,831)	(11,062)
<b>OTHER INCOME (EXPENSES)</b>			
Interest expense, net	(826)	(599)	(373)
Loss from continuing operations	(55,283)	(24,430)	(11,435)
Gain (loss) from discontinued operations	—	1,082	(379)
Net loss	<u>\$(55,283)</u>	<u>\$(23,348)</u>	<u>\$(11,814)</u>
Basic and diluted net loss per share:			
Loss from continuing operations	\$ (3.42)	\$ (2.00)	\$ (1.90)
Net loss	<u>\$ (3.42)</u>	<u>\$ (1.91)</u>	<u>\$ (1.96)</u>
Weighted-average common shares used to compute basic and diluted net loss per share	<u>16,163</u>	<u>12,223</u>	<u>6,013</u>

See accompanying Notes to Financial Statements.

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HERON THERAPEUTICS, INC.  
STATEMENTS OF STOCKHOLDERS' EQUITY  
(in thousands)

	Common Stock		Additional Paid-in Capital	Acc- umulated Deficit	Stock- holders' Equity
	Shares	Amount			
For the Years Ended December 31, 2013, 2012 and 2011					
<b>BALANCE, DECEMBER 31, 2010</b>	2,007	\$ 20	\$ 149,721	\$(148,425)	\$ 1,316
Net loss	—	—	—	(11,814)	(11,814)
Common stock and warrants issued in private placement, net of issuance costs	8,000	80	22,692	—	22,772
Conversion benefit included in convertible notes issued	—	—	1,573	—	1,573
Fair value of stock-based compensation for restricted stock awards issued/(cancelled) to directors	(9)	—	98	—	98
Common stock issued to employees under ESPP	4	—	15	—	15
Stock-based compensation expense related to stock options and ESPP	—	—	1,762	—	1,762
Fair value of warrants issued to non-employee	—	—	30	—	30
<b>BALANCE, DECEMBER 31, 2011</b>	10,002	100	175,891	(160,239)	15,752
Net loss	—	—	—	(23,348)	(23,348)
Common stock issued in private placement, net of issuance costs	5,100	51	50,440	—	50,491
Conversion benefit included in convertible notes issued	—	—	3,169	—	3,169
Common stock issued to employees under ESPP	4	—	22	—	22
Warrant exercise	5	1	(1)	—	—
Stock-based compensation expense related to stock options and ESPP	—	—	5,552	—	5,552
Fair value of warrants issued to non-employee	—	—	180	—	180
<b>BALANCE, DECEMBER 31, 2012</b>	15,111	152	235,253	(183,587)	51,818
Net loss	—	—	—	(55,283)	(55,283)
Common stock issued in private placement, net of issuance costs	7,706	77	57,725	—	57,802
Conversion benefit included in convertible notes issued	—	—	291	—	291
Common stock issued to employees under ESPP	6	1	33	—	34
Common stock issued for stock option exercises	537	5	2,788	—	2,793
Warrant exercise	212	2	598	—	600
Stock-based compensation expense related to stock options and ESPP	—	—	10,890	—	10,890
<b>BALANCE, DECEMBER 31, 2013</b>	23,572	\$ 237	\$ 307,578	\$(238,870)	\$ 68,945

See accompanying Notes to Financial Statements.

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HERON THERAPEUTICS, INC.  
STATEMENTS OF CASH FLOWS  
(in thousands)

For the Years Ended December 31,	2013	2012	2011
<b>CASH FLOWS FROM OPERATING ACTIVITIES:</b>			
Net loss	\$(55,283)	\$ (23,348)	\$(11,814)
Adjustments to reconcile net loss to net cash used in operating activities:			
(Gain) loss from discontinued operations	—	(1,082)	379
Depreciation and amortization	333	197	179
Amortization of debt discount	533	389	103
Stock-based compensation expense	10,890	5,732	1,890
Changes in operating assets and liabilities:			
Accounts receivable	—	—	110
Prepaid expenses and other current assets	(54)	(318)	16
Other long-term assets	—	—	(77)
Accounts payable	(426)	984	467
Accrued expenses	3,244	437	1,298
Deferred revenue	—	—	(272)
Net cash used in operating activities	(40,763)	(17,009)	(7,721)
<b>CASH FLOWS FROM INVESTING ACTIVITIES:</b>			
Purchases of property and equipment	(1,685)	(972)	(513)
Net cash used in investing activities	(1,685)	(972)	(513)
<b>CASH FLOWS FROM FINANCING ACTIVITIES:</b>			
Proceeds from sale of common stock, net of issuance costs	57,802	50,491	—
Proceeds from sale of units of common stock and warrants, net of issuance costs	—	—	22,772
Proceeds from convertible note financing, net of issuance costs	—	3,000	1,312
Proceeds from warrant exercises	600	—	—
Proceeds from the exercise of stock options	2,793	—	—
Proceeds from the issuance of shares under the Employee Stock Purchase Plan	34	22	15
Net cash provided by financing activities	61,229	53,513	24,099
Net increase in cash and cash equivalents	18,781	35,532	15,865
Cash and cash equivalents, beginning of year	53,506	17,974	2,109
Cash and cash equivalents, end of year	\$ 72,287	\$ 53,506	\$ 17,974
Supplemental Cash Flow Data:			
Cash paid for interest	\$ —	\$ —	\$ 2

See accompanying Notes to Financial Statements.

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HERON THERAPEUTICS, INC.  
NOTES TO FINANCIAL STATEMENTS  
DECEMBER 31, 2013, 2012 AND 2011

**NOTE 1 BUSINESS**

Heron Therapeutics, Inc. (formerly A.P. Pharma, Inc.) (the “Company,” “we,” “us” and “our”) is a specialty pharmaceutical company developing products using its proprietary Biochronomer™ polymer-based drug delivery platform. This drug delivery platform is designed to improve the therapeutic profile of injectable pharmaceuticals by converting them from products that must be injected once or twice per day to products that need to be injected only once every one or two weeks.

The Company’s lead product candidate, Sustol, is being developed for the prevention of acute chemotherapy-induced nausea and vomiting (CINV) for patients undergoing both moderately and highly emetogenic chemotherapy and for the prevention of delayed CINV for patients undergoing moderately emetogenic chemotherapy. One of the most debilitating side effects of cancer chemotherapy, CINV is a leading cause of premature discontinuations of treatment. There is only one injectable 5-HT<sub>3</sub> antagonist approved for the prevention of delayed-onset CINV, so this indication represents an area of particular unmet medical need. Sustol contains the 5-HT<sub>3</sub> antagonist granisetron formulated in our proprietary Biochronomer drug delivery system, which allows therapeutic drug levels to be maintained for five days with a single subcutaneous injection. This five-day range is designed to cover the delayed phase of CINV. Granisetron was selected for Sustol because it is widely prescribed by physicians based on a well-established record of safety and efficacy.

In May 2009, we filed the original New Drug Application (NDA) seeking approval for Sustol with the U.S. Food and Drug Administration (FDA). The FDA issued a Complete Response Letter for the Sustol NDA in March 2010. In September 2012, we resubmitted the NDA seeking approval for Sustol with the FDA. In March 2013, we received a second Complete Response Letter, which identifies several issues that preclude approval of the Sustol NDA. We believe the issues that remain are addressable, and we are working expeditiously to resubmit the Sustol NDA in the second quarter of 2014.

We own the worldwide rights to Sustol and are in the early stage of building commercial infrastructure necessary to commercialize Sustol in the U.S. on our own.

Additionally, we are exploring the potential use of our Biochronomer polymer with other drugs and intend to pursue the clinical development of one or more other drug candidates based on our proprietary delivery platform.

In January 2014, we changed our name from A.P. Pharma, Inc. to Heron Therapeutics, Inc. (Name Change). The Name Change is part of our recent corporate restructuring and rebranding of the Company. We believe the Name Change will emphasize our shift from a polymer development company to a commercial phase specialty pharmaceutical company, assuming approval of Sustol. Effective January 13, 2014, we effected a 1-for-20 reverse split of our outstanding common stock (Reverse Stock Split). (See Notes 8 and 15).

All historical share and per share amounts have been adjusted to reflect the Reverse Stock Split. All stock options, convertible notes and warrants outstanding were appropriately adjusted to give effect to the Reverse Stock Split.

**NOTE 2 SUMMARY OF SIGNIFICANT ACCOUNTING POLICIES**

**Basis of Presentation and Use of Estimates**

The preparation of financial statements in conformity with accounting principles generally accepted in the United States of America requires management to make estimates and assumptions that affect the amounts reported in the financial statements

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HERON THERAPEUTICS, INC.  
NOTES TO FINANCIAL STATEMENTS  
DECEMBER 31, 2013, 2012 AND 2011

and accompanying notes. Estimates were made relating to useful lives of fixed assets, valuation allowances, impairment of assets, accruals for research and development expenses and stock-based compensation expenses. Actual results could differ materially from those estimates. Certain amounts previously reported in the financial statements have been reclassified to conform to the current year presentation. Such reclassifications did not affect net loss, stockholders' equity or cash flows.

We have evaluated subsequent events through the date the audited financial statements were issued (See Note 15).

**Liquidity**

We have incurred significant operating losses and negative cash flows from operations and have an accumulated deficit of \$238.9 million as of December 31, 2013. In November 2013, we closed a common stock offering whereby we received approximately \$57.8 million of proceeds, net of issuance costs. During 2011 and 2012, we entered into three financing agreements, which provided us capital to fund operations. In April 2011, we entered into definitive agreements for a convertible note financing of up to \$4.5 million. We received approximately \$1.3 million, net of issuance costs, from the initial closing and an additional \$3.0 million through the issuance of additional convertible notes in May 2012 as a result of the purchasers who participated in the April 2011 convertible note financing fully exercising their rights to purchase additional convertible notes (see Note 6). In June 2011, we entered into definitive agreements for a private placement of units, which were comprised of common stock and warrants (see Note 8). The unit financing, which closed in July 2011, provided us with approximately \$22.8 million of proceeds, net of issuance costs. In July 2012, we closed a common stock financing whereby we received approximately \$50.5 million of proceeds, net of issuance costs (see Note 8). As of December 31, 2013, we had cash and cash equivalents of \$72.3 million.

We believe that our current working capital is sufficient to fund essential operations into 2015; provided, however, that if we pursue additional clinical studies or commercially launch Sustol prior to 2015, we will need to raise additional capital. We may require additional capital to fund our development pipeline programs. If we are unable to obtain sufficient financing on acceptable terms or otherwise, we may be required to reduce or defer our activities.

**Cash Equivalents**

We consider all highly liquid investments with a maturity from the date of purchase of less than three months to be cash equivalents. We have classified all of our investments in certain debt securities as "available-for-sale." As of December 31, 2012, our available-for-sale securities consisted of money market funds primarily containing U.S. government-backed or collateralized overnight securities with original maturities of ninety days or less. The carrying value of our money market funds is included in cash equivalents and approximates their fair value. We have no available-for-sale securities as of December 31, 2013. The Company's bank accounts have been placed under a control agreement in accordance with the April 2011 convertible note financing (see Note 6).

**Property and Equipment**

Property and equipment is stated at cost less accumulated depreciation. Depreciation is computed using the straight-line method over the estimated useful lives of the assets as follows: equipment and machinery, three to five years; furniture and fixtures, five years; and leasehold improvements, over the shorter of the respective lease terms or the respective useful lives of the leasehold improvements.

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HERON THERAPEUTICS, INC.  
NOTES TO FINANCIAL STATEMENTS  
DECEMBER 31, 2013, 2012 AND 2011

**Long-Lived Assets**

As circumstances dictate, we evaluate whether changes have occurred that would require us to consider whether long-lived assets have been impaired. Recoverability of assets to be held and used is determined by comparing the undiscounted net cash flows of long-lived assets to their respective carrying values. If such assets are considered to be impaired, the amount of impairment to be recognized is measured by comparing the projected discounted cash flows using an appropriate discount rate to the asset's carrying value.

**Stock-Based Compensation**

We account for share-based payment arrangements in accordance with ASC 718, *Compensation – Stock Compensation* and ASC 505-50, *Equity – Equity Based Payments to Non-Employees*, which requires the recognition of compensation expense, using a fair-value based method, for all costs related to share-based payments including stock options, restricted stock awards and stock issued under the employee stock purchase plan. These standards require companies to estimate the fair value of share-based payment awards on the date of the grant using an option-pricing model. See Note 8 for further discussion of our stock-based compensation plans.

**Warrants Issued in Connection with Equity Financings**

We generally account for warrants issued in connection with equity financings as a component of equity, unless there is a deemed possibility that we may have to settle warrants in cash.

**Revenue Recognition**

Our revenue arrangements with multiple deliverables are divided into separate units of accounting if certain criteria are met, including whether the delivered item has stand-alone value to the customer and whether there is objective and reliable evidence of the fair value of the undelivered items. The consideration we receive is allocated among the separate units based on their respective fair values, and the applicable revenue recognition criteria are considered separately for each of the separate units. Management's estimate of the selling price is considered when fair value is not determinable. Advance payments received in excess of amounts earned are classified as deferred revenue until earned.

**Contract Revenue**

Our licensing agreements generally provide for a non-refundable license fee. The license agreements provide for us to earn future revenue through royalty payments. These non-refundable license fees are generally initially reported as deferred revenue and recognized as revenue over an appropriate period, depending on the license. Revenue recognized from deferred license fees is classified as Contract Revenue in the accompanying statements of operations.

Contract revenue relates to research and development arrangements that generally provide for us to invoice research and development fees based on full-time equivalent hours for each project. Revenue from these arrangements is recognized as the related development services are rendered. This revenue approximates the costs incurred.

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**Clinical Trial Accruals**

Our expenses related to clinical trials are based on estimates of the services received and efforts expended pursuant to contracts with multiple research institutions and clinical research organizations that conduct and manage clinical trials on our behalf. Since the invoicing related to these services does not always coincide with our financial statement close process, we must estimate the level of services performed and fees incurred in determining the accrued clinical trial costs.

The financial terms of these agreements are subject to negotiation and vary from contract to contract, which may result in uneven payment flows. Payments under the contracts depend on factors such as the successful enrollment of patients, achievement of certain events or the completion of portions of the clinical trial or similar conditions. Expenses related to clinical trials generally are accrued based on the level of patient enrollment and services performed by the clinical research organization or related service provider according to the protocol. We monitor patient enrollment levels and related activity to the extent possible and adjust our estimates accordingly. Historically these estimates have been reasonably accurate and no material adjustments have had to be made.

**Research and Development**

Research and development consists of costs incurred for company-sponsored and collaborative research and development expenses. These costs consist primarily of employee salaries and other personnel-related expenses, facility-related expenses, laboratory consumables, polymer development manufacturing, clinical and pre-clinical related services performed by clinical research organizations, research institutions and other outside service providers.

Research and development expenses under collaborative agreements approximate the revenue recognized, excluding milestone and up-front payments received under such arrangements.

**Net Income (Loss) Per Share**

Basic income (loss) per share is estimated based on the weighted-average number of common shares outstanding. Diluted income (loss) per share is calculated using the weighted-average number of common shares outstanding and other dilutive securities. Dilutive securities are not included in the computation of diluted net loss per share if the inclusion of these potentially dilutive securities is anti-dilutive (see Note 9).

**Concentrations of Credit Risk**

Financial instruments that potentially subject us to concentrations of credit risk consist primarily of cash equivalents. We invest excess cash in a variety of high-grade short-term, interest-bearing securities. This diversification of risk is consistent with our policy to ensure safety of principal and maintain liquidity.

**Segment and Geographic Information**

Our operations are confined to a single business segment, the design and commercialization of polymer technologies for pharmaceutical and other applications.

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**Comprehensive Income (Loss)**

ASC 220, *Comprehensive Income*, requires that an entity's change in equity or net assets during a period from transactions and other events from non-owner sources be reported. Comprehensive loss for the periods reported was comprised solely of our net loss. The comprehensive loss for the years ended December 31, 2013, 2012 and 2011 was \$55.3 million, \$23.3 million and \$11.8 million, respectively. There were no other changes in equity that were excluded from our net loss for all periods.

**Recent Accounting Pronouncements**

There have been no recent accounting pronouncements or changes in accounting pronouncements during the fiscal year ended December 31, 2013 that we believe are of significance, or potential significance, to us.

**NOTE 3 FAIR VALUE MEASUREMENTS**

We follow the provisions of ASC 820-10, Fair Value Measurements and Disclosures, which defines fair value, establishes a framework for measuring fair value in generally accepted accounting principles and requires certain disclosures about fair value measurements. Broadly, the ASC 820-10 framework clarifies that fair value is an exit price, representing the amount that would be received to sell an asset or paid to transfer a liability in an orderly transaction between market participants. As such, fair value is a market-based measurement that should be determined based on assumptions that market participants would use in pricing an asset or liability.

As a basis for considering such assumptions, ASC 820-10 establishes a three tier value hierarchy which prioritizes the inputs used in measuring fair value as follows: Level 1) observable inputs such as quoted prices in active markets; Level 2) inputs other than the quoted prices in active markets that are observable either directly or indirectly; and Level 3) unobservable inputs in which there is little or no market data, which require us to develop our own assumptions. The hierarchy requires us to use observable market data, when available, and to minimize the use of unobservable inputs when determining fair value.

On a recurring basis, we measure our available-for-sale securities at fair value. We used quoted prices in active markets (Level 1) to measure our cash equivalents at fair value on a recurring basis in our balance sheet at December 31, 2012. Cash equivalents consist of highly rated money market funds with maturities of ninety days or less and purchased daily at par value with specified yield rates. Due to the high ratings and short-term nature of these funds, we consider the values of all cash equivalents as Level 1 inputs. We did not hold any cash equivalents at December 31, 2013.

The carrying amounts reported in the balance sheets for accounts payable and accrued expenses approximate fair value because of the short-term nature of these items.

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**NOTE 4 PROPERTY AND EQUIPMENT**

Property and equipment consist of the following (in thousands):

December 31,	2013	2012
Leasehold improvements	\$ 1,360	\$ 1,346
Furniture and equipment	5,795	4,036
Construction-in-progress	—	777
Total property and equipment	7,155	6,159
Accumulated depreciation	(4,273)	(4,407)
Property and equipment, net	\$ 2,882	\$ 1,752

Depreciation expense amounted to \$333,000, \$197,000 and \$179,000 for the years ended December 31, 2013, 2012 and 2011, respectively. At December 31, 2012, construction-in-progress related to equipment purchases not placed in service.

**NOTE 5 ACCRUED EXPENSES**

Accrued expenses consist of the following (in thousands):

December 31,	2013	2012
Research and development costs	\$ 1,795	\$ 651
Accrued compensation	2,566	401
Other	342	698
Total	\$ 4,703	\$ 1,750

**NOTE 6 CONVERTIBLE NOTES PAYABLE TO RELATED PARTIES**

In April 2011, we entered into a Securities Purchase Agreement (Purchase Agreement) with certain institutional investors (Purchasers), including a fund affiliated with Kevin C. Tang, who is the Chairman of our Board of Directors, for a private placement of up to \$4.5 million in Senior Secured Convertible Notes due 2021 (Notes). The Purchase Agreement provided for the Purchasers to purchase \$1.5 million aggregate principal amount of Notes at the initial closing. Pursuant to the Purchase Agreement, the Purchasers had the option to purchase an additional \$3.0 million aggregate principal amount of Notes at any time until May 2, 2013 (Purchase Option). The Notes are convertible into shares of the Company's common stock at a rate of 1,250 shares for every \$1,000 of principal and accrued interest due under the Notes (Conversion Shares).

The cash received from the initial closing of the Note financing, which resulted in the issuance of \$1.5 million aggregate principal amount of Notes, was approximately \$1.3 million, net of issuance costs. In May 2012, the Purchasers exercised their Purchase Option in full, and we received \$3.0 million of cash through the issuance of the remaining \$3.0 million aggregate principal amount of Notes. As a result of the exercise of the Purchase Option, the Purchasers have purchased the full amount of Notes that the Company was obligated to sell under the Purchase Agreement.

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The Notes are secured by substantially all of the assets of the Company, including placing our bank accounts under a control agreement. The Notes initially bore interest at 20% per annum, payable quarterly in cash or in additional principal amount of Notes at the election of the Purchasers. In June 2011, the Notes were amended to reduce the interest rate to 6% per annum effective July 1, 2011. The Notes mature on May 2, 2021; however, the holders of the Notes may require prepayment of the Notes at any time, at each holder's option.

There is no right to convert the Notes to the extent that, after giving effect to such conversion, the holder would beneficially own in excess of 9.99% of the Company's outstanding common stock. Each holder of the Notes can increase or decrease this beneficial ownership conversion limit by written notice to the Company, which will not be effective until 61 days after delivery of the notice.

As of December 31, 2013, the Company was in compliance with all debt-related covenants under the Notes. Upon the occurrence of an event of default under the Notes, the holders of the Notes have the right to require the Company to redeem all or a portion of their Notes.

Pursuant to the Purchase Agreement, the Company filed a registration statement on Form S-1, registering for resale 3.5 million shares (as adjusted for the reverse stock split) underlying the Notes. The registration statement was declared effective on July 29, 2011. The Purchasers have agreed to waive their right to require the Company to maintain the effectiveness of the registration statement and to register the additional shares underlying the Notes until they provide notice otherwise.

The Notes contain an embedded conversion feature that was in-the-money on both issuance dates. Based on an effective fixed conversion rate of 1,250 shares for every \$1,000 of principal and accrued interest due under the Notes, the total conversion benefit at issuance exceeded the loan proceeds. Therefore, a full debt discount was recorded in an amount equal to the face value of the Notes on the issuance dates and the Company began amortizing the resultant debt discount over the respective 10-year term of the Notes. During the year ended December 31, 2013, accrued interest of approximately \$291,000 was paid-in-kind and rolled into the Note principal balance, which resulted in an additional debt discount of approximately \$291,000. For the years ended December 31, 2013, 2012 and 2011, interest expense relating to the stated rate was approximately \$295,000, \$216,000 and \$97,000, respectively, and interest expense relating to the amortization of the debt discount was approximately \$533,000, \$389,000 and \$103,000, respectively.

As of December 31, 2013, the carrying value of the Notes was approximately \$1,025,000, which is comprised of the \$5,033,000 principal amount of the Notes outstanding, less debt discount of \$4,008,000. If the \$5.0 million principal amount of Notes is converted, the Company would issue 6.0 million shares of its common stock. Accrued interest on the principal balance was approximately \$75,000 at December 31, 2013.

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**NOTE 7 COMMITMENTS AND CONTINGENCIES**

Our lease for office, warehouse and laboratory space in Redwood City, California expires in 2016. Our Stamford, Connecticut office space is being leased on a month-to-month basis. Our office space in San Diego, California is being leased for an initial term that expires in December 2014. We also lease certain office equipment under operating lease arrangements. Our future minimum lease payments under these non-cancelable operating leases for facilities and equipment are as follows (in thousands):

For the Years Ended December 31,	<b>Minimum Payments</b>
2014	\$ 922
2015	779
2016	722
Total	<u>\$ 2,423</u>

Total rental expense for facilities and equipment was \$967,000, \$580,000 and \$591,000 for the years ended December 31, 2013, 2012 and 2011, respectively.

As permitted under Delaware law and in accordance with our bylaws, we indemnify our officers and directors for certain events or occurrences while the officer or director is or was serving at our request in such capacity. The term of the indemnification period is for the officer's or director's lifetime. The maximum amount of potential future indemnification is unlimited; however, we have a director or officer insurance policy that limits our exposure and may enable us to recover a portion of any future payments. We believe the fair value of these indemnification agreements is minimal. Accordingly, we have not recorded any liabilities for these agreements as of December 31, 2013.

In the normal course of business, we provide indemnifications of varying scope under our agreements with other companies, typically our clinical research organizations, investigators, clinical sites, suppliers and others. Pursuant to these agreements, we generally indemnify, hold harmless, and agree to reimburse the indemnified parties for losses suffered or incurred by the indemnified parties in connection with use or testing of our products or product candidates or with any U.S. patent or any copyright or other intellectual property infringement claims by any third party with respect to our products. The term of these indemnification agreements is generally perpetual. The potential future payments we could be required to make under these indemnification agreements is unlimited. Historically, costs related to these indemnification provisions have been immaterial. We also maintain various liability insurance policies that limit our exposure. As a result, we believe the fair value of these indemnification agreements is minimal. Accordingly, we have not recorded any liabilities for these agreements as of December 31, 2013.

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**NOTE 8 STOCKHOLDERS' EQUITY**

**Amendments to Certificate of Incorporation**

*Reverse Stock Split*

Effective January 13, 2014, we amended our Certificate of Incorporation to change our name to Heron Therapeutics, Inc. and to effect a 1-for-20 reverse split of our outstanding common stock. The Name Change and Reverse Stock Split were approved by our stockholders on September 19, 2013. As a result of the Reverse Stock Split, the total authorized shares of common stock were reduced from 1,500,000,000 to 75,000,000 shares (See Note 15).

**2009 Private Placement**

In October 2009, in a private placement, we sold 397,727 shares of our common stock at \$17.60 per share and warrants to purchase 198,864 shares of our common stock, exercisable through January 7, 2015, at an exercise price of \$17.60 per share (2009 Private Placement). The purchasers paid \$2.50 per underlying share for the warrants. Additionally, the purchasers had the right to purchase up to an additional 258,264 shares at \$19.40 per share prior to May 14, 2010 and paid \$2.50 per underlying share for the right to purchase such additional shares. These rights expired unexercised on May 14, 2010. Total proceeds from the 2009 Private Placement were approximately \$7.9 million, net of issuance costs. We filed a registration statement on Form S-3 covering 376,631 shares on November 6, 2009, which was declared effective by the SEC on November 17, 2009. On June 30, 2010, we filed a registration statement on Form S-3 covering the remaining 21,096 shares and the 198,864 shares of our common stock underlying the warrants, which was declared effective by the SEC on July 8, 2010. If we fail to keep any registration statements continuously effective, we may be obligated to pay to the holders of the shares and warrants liquidated damages in the amount of 1% per month of the purchase price for the shares and warrants, up to a maximum cap of 8% of such purchase price.

**2011 Private Placement**

In June 2011, we entered into a Securities Purchase Agreement with certain purchasers (Securities Purchase Agreement), pursuant to which we agreed to sell 8.0 million shares of our common stock (Shares) and warrants to purchase 4.0 million shares of our common stock (Warrants) with an exercise price of \$3.60 per share (2011 Private Placement), for an aggregate price of \$24.0 million. The 2011 Private Placement closed on July 1, 2011. For each share purchased, the investors received one Warrant to purchase 0.5 shares of common stock (together with a Share, a Unit), at a purchase price of \$3.00 per Unit. The Warrants were immediately exercisable and expire on the fifth anniversary of the closing date of July 1, 2011. The Warrants may be exercised for cash only or, if a registration statement is not then effective and available for the resale of the shares of common stock issuable upon exercise of the Warrants, by surrender of such Warrant, or a portion of such Warrant, by way of cashless exercise. There is no right to exercise the Warrants to the extent that, after giving effect to such exercise the holder would beneficially own in excess of 9.99% of our outstanding shares of common stock or such other limit as may be designated by any particular purchaser. Each holder of the Warrants can amend or waive the foregoing limitation by written notice to the Company, with such waiver taking effect only upon the expiration of a 61-day notice period.

Under the terms of the Securities Purchase Agreement, on July 29, 2011, the Company filed a registration statement with the SEC to register for resale the Shares and the shares of common stock issuable upon the exercise of the Warrants (the Warrant

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Shares, and collectively with the Shares, the Registrable Securities). The registration statement was declared effective on August 4, 2011. The Company is obligated to maintain the effectiveness of the registration statement with respect to an investor's Registrable Securities until the investor is able to sell the Registrable Securities without limitation or restriction under Rule 144. There is currently only one investor who is an affiliate of the Company and is therefore not able to sell the Registrable Securities without limitation under Rule 144, and that investor has agreed to waive its right to require the Company to maintain the effectiveness of the registration statement until it provides notice otherwise. If the Company fails to keep the registration statement continuously effective for a designated time (with limited exceptions) during the period the Company is obligated to maintain the registration statement, the Company may be obligated to pay to the holders of the Registrable Securities liquidated damages in an amount equal to 1.0% per month of such holder's pro rata interest in the total purchase price of the Private Placement, capped at a total penalty of 6.0%.

The Company received total proceeds of \$22.8 million from the 2011 Private Placement, which was net of issuance costs of approximately \$1.2 million. During the year 2013, the Company received \$0.6 million for an exercise of a Warrant.

**2012 Private Placement**

In July 2012, the Company entered into a securities purchase agreement with certain purchasers, pursuant to which the Company agreed to sell 5.1 million shares of its common stock (2012 Shares) at a purchase price of \$10.50 per share of common stock, for an aggregate price of approximately \$53.6 million (2012 Private Placement). The Company received total proceeds of \$50.5 million from the 2012 Private Placement, which was net of issuance costs of approximately \$3.1 million.

In connection with entering into the securities purchase agreement, the Company also entered into a registration rights agreement. On August 24, 2012, the Company filed a registration statement with the SEC to register the 2012 Shares for resale. The registration statement was declared effective on September 6, 2012. If the Company fails to keep the registration statement continuously effective for a designated time (with limited exceptions), the Company may be obligated to pay to each holder of the 2012 Shares an amount equal to 1.5% per month of the aggregate purchase price of the unregistered 2012 Shares held by such holder, capped at a total penalty of 9.0%.

**2013 Common Stock Offering**

On November 20, 2013, the Company entered into an underwriting agreement (Underwriting Agreement) with Jefferies LLC, as representative of the several underwriters (Underwriters), pursuant to which we agreed to issue and sell an aggregate of 7,500,000 shares of our common stock to the Underwriters (Offering). Under the terms of the Underwriting Agreement, we granted the Underwriters an option for 30 days to purchase up to an additional 1,125,000 shares of our common stock.

The shares in the Offering were sold at a public offering price of \$8.00 per share. The public offering closed on November 25, 2013. We received net proceeds of \$56.3 million, after deducting the Underwriters' discounts and commissions and offering expenses payable by us. On December 5, 2013, the Underwriters exercised their option to purchase 206,500 of the 1,125,000 additional shares of common stock. We received net proceeds of \$1.5 million from issuance of these additional shares.

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The Offering was made pursuant to the Company's effective registration statement on Form S-3 (Registration No. 333-190550), which was previously filed with the Securities and Exchange Commission (SEC) and was declared effective, and a prospectus supplement filed with the SEC. The Offering was not registered under any state blue sky laws and was limited to "Qualified Institutional Buyers" (as defined in Rule 144A under the Securities Act of 1933, as amended) and certain other institutional and accredited investors, as permitted under applicable law. In the Underwriting Agreement, we agreed to indemnify the Underwriters against certain liabilities, including liabilities under the Securities Act of 1933, as amended, or to contribute to payments that the Underwriters may be required to make because of such liabilities.

**Stock-Based Compensation Plans**

We have two types of stock-based compensation plans, which consist of an employee stock purchase plan and three stock option plans.

In 1997, our stockholders approved our 1997 Employee Stock Purchase Plan (the Purchase Plan). In December 2007, May 2009 and June 2011, our stockholders authorized increases in the number of shares reserved for issuance under the Purchase Plan by 5,000, 10,000 and 25,000 shares, respectively, for a total of 50,000 shares reserved at December 31, 2013. Under the terms of the Purchase Plan, employees can elect to have up to a maximum of 10% of their base earnings withheld to purchase our common stock. The purchase price of the stock is 85% of the lower of the closing prices for our common stock on: (i) the first trading day in the enrollment period, as defined in the Purchase Plan, in which the purchase is made, or (ii) the purchase date. The length of the enrollment period may not exceed a maximum of six months. Our compensation committee modified the Purchase Plan such that beginning in May 2008, the length of all offering periods was decreased from 24 months to six months. Enrollment dates are the first business day of May and November and the first enrollment date was April 30, 1997. Approximately 9% of eligible employees participated in the Purchase Plan in 2013. Under the Purchase Plan, we issued 5,634, 4,274 and 3,899 shares in 2013, 2012 and 2011, respectively. The weighted-average fair value per share of purchase rights granted during 2013, 2012 and 2011 was \$7.23, \$5.20 and \$4.00, respectively. The weighted-average exercise price per share of the purchase rights exercised during 2013, 2012 and 2011 was \$6.12, \$5.20 and \$4.00, respectively. We had 16,651, 22,285 and 26,559 shares reserved for issuance under the Purchase Plan at December 31, 2013, 2012 and 2011, respectively.

We currently have one stock option plan from which we can grant options and restricted stock awards to employees, officers, directors and consultants. In December 2007, the stockholders approved our 2007 Equity Incentive Plan (the 2007 Plan). In May 2010 and June 2011, our stockholders approved amendments to our 2007 Equity Incentive Plan to increase the maximum number of shares of common stock available for grant by 100,000 and 4,500,000 shares of common stock, respectively, resulting in an aggregate of 4,750,000 shares of common stock authorized for issuance pursuant to awards granted under our 2007 Equity Incentive Plan. We have also granted stock options and restricted stock awards under the 2002 Stock Incentive Plan (the 2002 Plan) and the Non-Qualified Stock Plan (the NQ Plan) in prior years. We were authorized to issue up to 21,250 shares under the 2002 Plan, which includes an increase of 5,000 shares which were approved by stockholders in May 2006, and 103,125 shares under the NQ Plan, a plan that had not undergone stockholder approval and could only be utilized to grant stock options and restricted stock awards as inducements to attract new employees, to which 50,000 shares were added by the Board of Directors in September 2007, and an additional 50,000 shares were added in July 2008. The remaining shares available in the NQ Plan and 2002 Plan expired in October 2010 and February 2012, respectively. In 2013 and 2012, we

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granted options to certain employees outside of our approved stock option plan. The options to purchase our common stock are granted with an exercise price which equals fair market value of the underlying common stock on the grant dates and expire no later than ten years from the date of grant. The options are exercisable in accordance with vesting schedules that generally provide for them to be fully vested and exercisable four years after the date of grant. Any shares that are issuable upon exercise of options granted that expire or become unexercisable for any reason without having been exercised in full are available for future grant and issuance under the same stock option plan.

As discussed in Note 2, we record stock-based compensation expense based on the fair value of stock options and purchase rights issued to employees in conjunction with our stock option plans or the Purchase Plan on the grant date or purchase date. We also record compensation expense for warrants and stock options issued to non-employees and restricted stock awards to employees and directors.

The fair value of each employee and director grant of options to purchase common stock and purchase rights under the Purchase Plan is estimated on the date of the grant using the Black-Scholes option-pricing model assuming no dividends and the following weighted-average assumptions:

For the Years Ended December 31,	2013	2012	2011
<b>Expected term (years):</b>			
Stock options	6.00	5.51	5.00
Employee Stock Purchase Plan	.49	.49	.50
<b>Risk-free interest rate:</b>			
Stock options	1.5%	0.8%	1.6%
Employee Stock Purchase Plan	0.8%	0.2%	0.8%
<b>Volatility:</b>			
Stock options	105%	106%	106%
Employee Stock Purchase Plan	212%	0.8%	142%

The expected term is based on historical data. The expected term for the Purchase Plan is based on the weighted-average purchase period of the Purchase Plan. The risk-free interest rate is based on the U.S. treasury rate over the expected term of the award. The expected volatility is based on our historical stock prices, and the estimated forfeiture rate of the options is based on historical data.

The Black-Scholes option valuation model requires the input of highly subjective assumptions, including the expected life of the award and stock price volatility. The assumptions listed above represent management's best estimates, but these estimates involve inherent uncertainties and the application of management judgment. As a result, if other assumptions had been used, our recorded stock-based compensation expense could have been materially different.

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Stock-based compensation expense recorded for awards granted under the stock option plans and the Purchase Plan, net of estimated forfeitures, was as follows (in thousands, except per share amounts):

For the Years Ended December 31,	2013	2012	2011
Research and development	\$ 2,562	\$ 1,585	\$ 819
General and administrative	8,328	4,147	1,071
Total	\$10,890	\$5,732	\$1,890
Impact on basic and diluted net loss per common share	\$ 0.67	\$ 0.47	\$ 0.31

We recorded additional stock-based compensation expense in 2013 and 2012 as a result of accelerated vesting of stock options in connection with the resignation of our former chief executive officer and two directors, respectively. No tax benefit was recognized related to stock-based compensation expense since we have incurred operating losses and we have established a full valuation allowance to offset all the potential tax benefits associated with our deferred tax assets.

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The following table summarizes option activity for the years ended December 31, 2013, 2012 and 2011:

	2013			2012		2011		
	Shares	Weighted-Average Exercise Price	Weighted-Average Remaining Contractual Term (Years)	Aggregate Intrinsic Value as of December 31, 2013	Shares	Weighted-Average Exercise Price	Shares	Weighted-Average Exercise Price
Outstanding at beginning of year	4,323,874	\$ 8.46			2,505,283	\$ 6.13	160,833	\$ 29.83
Granted	4,256,971	8.08			1,874,246	11.61	2,403,495	5.19
Exercised	(537,029)	5.20			—	—	—	—
Expired or Forfeited	(1,688,135)	9.45			(55,655)	9.56	(59,045)	32.53
Outstanding at end of year	<u>6,355,681</u>	8.22	8.14	\$ 10,704,578	<u>4,323,874</u>	8.46	<u>2,505,283</u>	6.13
Options exercisable at year end	<u>1,770,597</u>	8.67	5.12	\$ 4,607,452	<u>1,142,099</u>	7.98	<u>337,002</u>	10.68
Options vested or expected to vest	6,292,296	8.22	8.13	\$ 10,614,670	4,283,344	8.45	2,471,725	6.14
Shares available for future grant at year end	1,797,694				1,018,766		2,213,155	
Weighted-average fair value of stock options granted during the year		\$ 8.08				\$ 11.61		\$ 5.19

As of December 31, 2013, there was approximately \$39.1 million of total unrecognized compensation expense related to unvested stock options. This expense is expected to be recognized over a weighted-average period of 2.55 years. Cash received from option exercises for the years ended December 31, 2013, 2012 and 2011 was \$2,793,000, \$0 and \$0, respectively. The total intrinsic value of options exercised in the year ended December 31, 2013, 2012 and 2011 was \$1.7 million, \$0 and \$0, respectively.

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The following table summarizes information about stock options outstanding at December 31, 2013:

Range of Exercise Prices	OPTIONS OUTSTANDING			OPTIONS EXERCISABLE	
	Number Outstanding	Weighted-Average Remaining Contractual Life (Years)	Weighted-Average Exercise Price	Number Exercisable	Weighted-Average Exercise Price
\$3.80 - \$3.80	5,000	7.99	\$ 3.80	2,708	\$ 3.80
\$5.20 - \$5.20	1,072,887	4.02	5.20	997,934	5.20
\$6.40 - \$7.00	386,613	8.60	6.85	80,210	6.66
\$7.20 - \$7.20	3,190,240	9.34	7.20	41,012	7.20
\$7.40 - \$10.60	852,372	9.38	9.89	105,881	10.57
\$11.00 - \$15.00	730,285	7.54	13.15	499,642	12.79
\$15.40 - \$130.40	117,598	6.44	24.14	42,524	39.54
\$139.20 - \$139.20	125	1.08	139.20	125	139.20
\$196.00 - \$196.00	436	0.04	196.00	436	196.00
\$235.12 - \$235.12	125	0.40	235.12	125	235.12
\$3.80 - \$235.12	6,355,681	8.14	\$ 8.22	1,770,597	\$ 8.67

As of December 31, 2013, there were no unvested restricted stock awards granted to employees and directors. The compensation cost that has been expensed in the statements of operations for the restricted stock awards issued to employees and directors was \$0 for 2013 and 2012 and, \$98,000 for 2011, respectively.

The following table summarizes information about the Company's warrants outstanding at December 31, 2013:

	Number of Shares Exercisable	Exercise Price	Expiration Date
Issued to private placement investors in October 2009	198,864	\$17.60	1/7/2015
Issued to private placement investors in July 2011	3,750,000	\$3.60	7/1/2016
Other	20,000	\$13.80	8/1/2015
Total warrants outstanding at December 31, 2013	3,968,864	\$4.35*	

\* Average exercise price

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HERON THERAPEUTICS, INC.  
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**Common Stock Reserved for Future Issuance**

As of December 31, 2013, the Company had reserved shares of common stock for future issuance as follows:

	<b>No. of Shares</b>
Issuance upon exercise of outstanding stock options	6,355,681
Issuance of future grants under stock option plans	1,797,694
Issuance of future grants under employee stock purchase plan	16,651
Issuance of common stock related to convertible notes*	9,884,342
Issuance upon exercise of warrants	3,968,864
Total	<u>22,023,232</u>

\* Assumes all interest payments are paid-in-kind through the maturity date.

**NOTE 9 NET LOSS PER SHARE**

Basic and diluted net loss per share is computed by dividing net loss by the weighted-average number of common shares outstanding for the applicable period. Diluted net loss per share excludes the effect of outstanding potentially dilutive securities because they are anti-dilutive. The following table shows the outstanding potentially dilutive options, warrants and convertible notes as of December 31, 2013, 2012 and 2011 (in thousands):

For the Years Ended December 31,	<b>2013</b>	<b>2012</b>	<b>2011</b>
Number of options outstanding	6,356	4,324	2,505
Number of warrants outstanding	3,969	4,219	4,206
Common stock related to convertible notes outstanding	6,291	5,927	1,967

**NOTE 10 DISCONTINUED OPERATIONS****Cosmeceutical and Toiletry Business**

On July 25, 2000, we completed the sale of certain technology rights for our cosmeceutical and toiletry business to RP Scherer Corporation (RP Scherer), a subsidiary of Cardinal Health, Inc. Under the terms of the agreement, we guaranteed a minimum gross profit percentage on RP Scherer's combined sales of products to Ortho Dermatologics (Ortho) and Dermik Laboratories, Inc. (Dermik) (Gross Profit Guaranty), both of which were acquired by Valeant Pharmaceuticals in July 2011. The Gross Profit Guaranty expense totaled \$944,000 for the first seven guaranty years. Effective March 2007, in conjunction with a sale of assets by RP Scherer's successor company to an Amcol International subsidiary (Amcol), a new agreement was signed between us and Amcol to provide continuity of product supply to Ortho and Dermik. We had previously recorded a liability of \$1.1 million related to the amount that Amcol asserted was due under the Gross Profit Guaranty. In February 2013, an arbitrator ruled that no additional amounts were owed. This event qualified as an adjusting event under ASC 855, Subsequent Event, and in light of the arbitrator's decision in February 2013, which was final and binding, we reversed this accrual as of December 31, 2012.

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HERON THERAPEUTICS, INC.  
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The cosmeceutical and toiletry business is reported as discontinued operations for all periods presented in the accompanying Statements of Operations. Gain (loss) from discontinued operations represents primarily the gain (loss) attributable to changes in estimates of our cosmeceutical and toiletry business that was sold to RP Scherer on July 25, 2000, as follows (in thousands):

For the years ended December 31,	2013	2012	2011
<b>Cosmeceutical and Toiletry Business:</b>			
Change in estimates for guarantees	\$ —	\$1,082	\$(379)

There was no revenue relating to discontinued operations for the years ended December 31, 2013, 2012 and 2011.

Basic and diluted income (loss) per common share from discontinued operations was \$0, \$0.09 and (\$0.06) for the years ended December 31, 2013, 2012 and 2011, respectively.

**NOTE 11 DEFINED CONTRIBUTION PLAN**

We have a defined contribution plan (401k) covering substantially all of our employees. In the past three calendar years, we made matching cash contributions equal to 50% of each participant's contribution during the plan year up to a maximum amount equal to the lesser of 3% of each participant's annual compensation or \$7,650, \$7,500 and \$7,350 for the years 2013, 2012 and 2011, respectively. Such amounts were recorded as expense in the corresponding years. We may also contribute additional discretionary amounts to the defined contribution plan as we may determine. For the years ended December 31, 2013, 2012 and 2011, we contributed to the plan approximately \$134,000, \$71,000 and \$46,000, respectively. No discretionary contributions have been made to the plan since its inception.

**NOTE 12 INCOME TAXES**

There is no provision recorded for the fiscal years ended 2013, 2012 and 2011 because we have incurred operating losses.

Deferred income taxes reflect the net tax effects of net operating loss and tax credit carryovers and temporary differences between the carrying amounts of assets and liabilities for financial reporting purposes and the amounts used for income tax purposes. Significant components of our deferred tax assets are as follows (in thousands):

December 31,	2013	2012	2011
<b>Deferred Tax Assets:</b>			
Net operating loss carryforwards	\$ 31,834	\$ 13,900	\$ 6,700
Research credits	4,792	2,300	2,100
Stock compensation expense	4,124	2,600	600
Other	487	400	900
Total deferred tax assets	41,237	19,200	10,300
Valuation allowance	(41,237)	(19,200)	(10,300)
Net deferred tax assets	\$ —	\$ —	\$ —

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HERON THERAPEUTICS, INC.  
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Realization of our deferred tax assets is dependent upon our future taxable income, if any, the timing and amount of which are uncertain. Accordingly, our deferred tax assets have been fully offset by a valuation allowance. The valuation allowance increased by \$22.0 million and \$8.9 million during 2013 and 2012, respectively.

As of December 31, 2013, we had federal and California net operating loss carryforwards of \$80.1 million and \$79.1 million, respectively, and federal and California research and development tax credit carryforwards of \$2.2 million and \$3.9 million, respectively. Of the carryforwards, federal and California net operating loss carryforwards of \$75.0 million and \$74.0 million, respectively, are subject to annual limitations and will be available from 2014 through 2031, as a result of federal ownership change limitations. The remaining federal and state net operating losses carryforwards expire beginning in 2018 for federal and 2014 for state, through 2033, if not utilized. The federal research credits expire beginning in 2022 and the state research credits have no expiration date.

Federal and state laws limit the use of net operating loss and tax credit carryforwards in certain situations where changes occur in the stock ownership of a company. We conducted an analysis of our stock ownership under Internal Revenue Code Section 382 and have reported our deferred tax assets related to net operating loss and research credit carryforwards after recognizing change of control limitations in 2007, 2011 and 2013. The limitation of our federal and state carryforwards associated with previous net operating loss and research credit carryforwards, and the associated reduction in our deferred tax assets, was offset by a reduction in our valuation allowance. Utilization of our remaining net operating loss and research and development credit carryforwards may still be subject to substantial annual limitations due to ownership change limitations after December 31, 2013. Such an annual limitation could result in the expiration of the net operating loss and research and development credit carryforwards available as of December 31, 2013 before utilization.

The provision for income taxes differs from the amount computed by applying the U.S. federal statutory tax rate (34% in 2013, 2012 and 2011) to income taxes as follows (in thousands):

December 31,	2013	2012	2011
Tax benefit computed at 34%	\$(18,796)	\$(7,937)	\$(4,017)
Stock compensation expense	513	176	148
Other	300	210	70
NOL not benefitted	17,983	7,551	3,799
Tax Provision (Benefit)	\$ —	\$ —	\$ —

We follow the provisions of ASC 740-10-50, Accounting for Uncertainty in Income Tax Provisions. A reconciliation of the beginning and ending amounts of unrecognized tax benefits is as follows (in thousands):

December 31,	2013	2012	2011
Unrecognized tax benefit:			
At the beginning of the period	\$ 120	\$ 120	\$ 120
Gross increases – tax positions in the current period	—	—	—
Gross decreases – tax positions in the current period	—	—	—
At the end of the period	\$120	\$120	\$120

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HERON THERAPEUTICS, INC.  
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The unrecognized tax benefit, if recognized in full, would result in adjustments to deferred taxes and the related valuation allowance. We do not currently anticipate any significant changes to the unrecognized tax benefits in 2014. Our policy is to recognize interest and penalties accrued on any unrecognized tax benefits as a component of tax expense. To date, we have not used the unrecognized tax benefits to reduce any of our past tax obligations. As a result, we had no accrual for the payment of interest and penalties related to the unrecognized tax benefits. As of December 31, 2013, our tax returns were subject to future examination in the U.S. federal and state tax jurisdictions for the tax years 1997 through 2013, due to net operating losses and research credits that are being carried forward.

**NOTE 13 SIGNIFICANT AGREEMENTS****Merial Limited**

In September 2009, we entered into a world-wide license and development agreement with Merial Limited (Merial), a world leading animal health company, for a long acting pain management product for companion animals. We received a nonrefundable upfront license fee and performed reimbursable development services. In May 2011, we received notice of termination from Merial. We recognized \$0.6 million in revenue related to development services to Merial for the year ended 2011.

**NOTE 14 QUARTERLY RESULTS OF OPERATIONS (UNAUDITED)**

The following table presents summarized unaudited results of operations for each of our quarters in the years ended December 31, 2013 and 2012.

**Quarterly Results of Operations**  
**(in thousands, except per share data)**  
**(unaudited)**

	<b>First Quarter</b>	<b>Second Quarter</b>	<b>Third Quarter</b>	<b>Fourth Quarter</b>
<b>Year Ended December 31, 2013</b>				
Total revenue	\$ —	\$ —	\$ —	\$ —
Operating expenses	12,752	15,209	12,664	13,832
Interest and other expense, net	(201)	(204)	(209)	(212)
Net loss	(12,953)	(15,413)	(12,873)	(14,044)
Basic and diluted net loss per share:				
Net loss	\$ (0.85)	\$ (1.01)	\$ (0.84)	\$ (0.75)
<b>Year Ended December 31, 2012</b>				
Total revenue	\$ —	\$ —	\$ —	\$ —
Operating expenses	4,769	4,380	6,054	8,628
Interest and other expense, net	(61)	(146)	(195)	(197)
Loss from continuing operations	(4,830)	(4,526)	(6,249)	(8,825)
Discontinued operations	(91)	(43)	128	1,088
Net loss	(4,921)	(4,569)	(6,121)	(7,737)
Basic and diluted net loss per share:				
Loss from continuing operations	\$ (0.48)	\$ (0.45)	\$ (0.46)	\$ (0.58)
Net loss	(0.49)	(0.46)	(0.45)	(0.51)

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HERON THERAPEUTICS, INC.  
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**NOTE 15 SUBSEQUENT EVENT**

*Reverse Stock Split*

Effective January 13, 2014, we amended our Certificate of Incorporation to change our name to Heron Therapeutics, Inc., and to effect a 1-for-20 reverse split of our outstanding common stock. The Name Change and Reverse Stock Split were approved by our stockholders on September 19, 2013. As a result of the Reverse Stock Split, the total authorized shares of common stock were reduced from 1,500,000,000 to 75,000,000 shares.

All historical share and per share amounts have been adjusted to reflect the Reverse Stock Split. All stock options, convertible notes and warrants outstanding were ratably adjusted to give effect to the Reverse Stock Split.

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**ITEM 9. CHANGES IN AND DISAGREEMENTS WITH ACCOUNTANTS ON ACCOUNTING AND FINANCIAL DISCLOSURE**

None.

**ITEM 9A. CONTROLS AND PROCEDURES**

**Evaluation of Disclosure Controls and Procedures**

Based on an evaluation as of the end of the period covered by this report, our Chief Executive Officer and Chief Financial Officer have concluded that our disclosure controls and procedures (as defined in Rules 13a-15(e) and 15d-15(e) under the Exchange Act) were effective as of the end of the period covered by this report to ensure that information that we are required to disclose in reports that we file or submit under the Exchange Act is recorded, processed, summarized and reported within the time periods specified in SEC rules and forms.

Our disclosure controls and procedures are designed to provide reasonable assurance of achieving their objectives, and our Chief Executive Officer and Chief Financial Officer have concluded that these controls and procedures are effective at the “reasonable assurance” level. We believe that a control system, no matter how well designed and operated, cannot provide absolute assurance that the objectives of the control system are met, and no evaluation of controls can provide absolute assurance that all control issues and instances of fraud, if any, within a company have been detected.

**Management’s Annual Report on Internal Control Over Financial Reporting**

Our management is responsible for establishing and maintaining adequate internal control over financial reporting. A company’s internal control over financial reporting is a process designed to provide reasonable assurance regarding the reliability of financial reporting and the preparation of financial statements for external purposes in accordance with generally accepted accounting principles.

There are inherent limitations in the effectiveness of any system of internal control, including the possibility of human error and the circumvention or overriding of controls. Accordingly, even effective internal controls can provide only reasonable assurances with respect to financial statement preparation. Further, because of changes in conditions, the effectiveness of internal control may vary over time.

Our management assessed the effectiveness of our internal control over financial reporting as of December 31, 2013. In making this assessment, management used the criteria set forth in the report entitled *Internal Control — Integrated Framework (1992)* published by the Committee of Sponsoring Organizations (COSO) of the Treadway Commission. Based on our assessment using the COSO criteria, management concluded that, as of December 31, 2013, our internal control over financial reporting is effective.

Our independent registered public accounting firm, OUM & Co., LLP has audited our Financial Statements included in this Annual Report on Form 10-K and have issued a report on the effectiveness of our internal controls over financial reporting as of December 31, 2013. Their reports appear in Item 8 of this Annual Report on Form 10-K.

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**Changes in Internal Controls Over Financial Reporting**

There were no changes in our internal controls over financial reporting that occurred during our most recent fiscal quarter that have materially affected, or are reasonably likely to materially affect, our internal control over financial reporting.

**ITEM 9B. OTHER INFORMATION**

None.

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PART III

**ITEM 10. DIRECTORS, EXECUTIVE OFFICERS AND CORPORATE GOVERNANCE**

The information required by this item is incorporated by reference into the information set forth under the captions "Election of Directors," "Executive Officers," "Corporate Governance" and "Section 16(a) Beneficial Ownership Reporting Compliance" of our Proxy Statement (the "Proxy Statement") for the 2014 annual meeting of stockholders.

**Code of Ethics**

We have adopted a Code of Ethics that applies to all of our directors, officers and employees. The Code of Ethics is posted on our website at <http://www.herontx.com> under the caption "Investor Relations/ Corporate Governance." If we make any substantive amendments to the code of ethics or grant any waiver, including implicit waiver, from a provision of the code of ethics to our principal executive officer, principal financial officer or principal accounting officer, we will disclose the nature of such amendment or waiver on our website or in a current report on Form 8-K that will be publicly filed.

**ITEM 11. EXECUTIVE COMPENSATION**

The information required by this item is incorporated by reference into the information set forth under the captions "Executive Compensation" and "Director Compensation" of the Proxy Statement.

**ITEM 12. SECURITY OWNERSHIP OF CERTAIN BENEFICIAL OWNERS AND MANAGEMENT AND RELATED STOCKHOLDER MATTERS**

The information required by this item is incorporated by reference into the information set forth under the captions "Common Stock Ownership of Certain Beneficial Owners and Management" and "Equity Compensation Plan Information" of the Proxy Statement.

**ITEM 13. CERTAIN RELATIONSHIPS AND RELATED TRANSACTIONS AND DIRECTOR INDEPENDENCE**

The information required by this item is incorporated by reference into the information set forth under the captions "Related Party Transactions" and "Corporate Governance" of the Proxy Statement.

**ITEM 14. PRINCIPAL ACCOUNTANT FEES AND SERVICES**

The information required by this item is incorporated by reference into the information set forth under the captions "Report of the Audit Committee," "Ratification of Independent Registered Public Accountants" and "Auditors Fees and Services" of the Proxy Statement.

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PART IV

**ITEM 15. EXHIBITS AND FINANCIAL STATEMENT SCHEDULES**

(a) 1. Financial Statements

The financial statements and supplementary data set forth in Part II of the Annual Report on Form 10-K are included herein.

2. Financial Statement Schedules

All schedules have been omitted because the information is not required or is not so material as to require submission of the schedule, or because the information is included in the financial statements or the notes thereto.

3. Exhibits

See Exhibit Index beginning on page 84.

**SIGNATURES**

Pursuant to the requirement of Section 13 or 15(d) of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned, thereunto duly authorized.

**HERON THERAPEUTICS, INC.**

By: /s/ Barry D. Quart  
Barry D. Quart  
Chief Executive Officer  
Date: March 7, 2014

**POWER OF ATTORNEY**

KNOW ALL PERSONS BY THESE PRESENT, that each person whose signature appears below constitutes and appoints Barry D. Quart his true and lawful attorney-in-fact, with the power of substitution, for him in any and all capacities, to sign any amendments to this Annual Report on Form 10-K and to file the same, with exhibits thereto and other documents in connection therewith, with the Securities and Exchange Commission, hereby ratifying and confirming all that said attorney-in-fact, or his substitute or substitutes, may do or cause to be done by virtue hereof.

Pursuant to the requirements of the Securities Exchange Act of 1934, this report has been signed by the following persons in the capacities and on the dates indicated.

<b>Signature</b>	<b>Title</b>	<b>Date</b>
<u>/s/ Barry D. Quart</u> Barry D. Quart	Chief Executive Officer, Director (Principal Executive Officer)	March 7, 2014
<u>/s/ Robert H. Rosen</u> Robert H. Rosen	President, Chief Commercial Officer, Director	March 7, 2014
<u>/s/ Kevin C. Tang</u> Kevin C. Tang	Chairman of the Board of Directors	March 7, 2014
<u>/s/ Stephen R. Davis</u> Stephen R. Davis	Executive Vice-President, Chief Operating Officer, Director	March 7, 2014
<u>/s/ Brian G. Drazba</u> Brian G. Drazba	Vice President, Chief Financial Officer (Principal Financial Officer)	March 7, 2014
<u>/s/ John W. Poyhonen</u> John W. Poyhonen	Director	March 7, 2014
<u>/s/ Craig A. Johnson</u> Craig A. Johnson	Director	March 7, 2014
<u>/s/ Kimberly J. Manhard</u> Kimberly J. Manhard	Director	March 7, 2014

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EXHIBIT INDEX

**FORM 10-K ANNUAL REPORT**

<b>Exhibit Number</b>	<b>Description</b>	<b>Incorporated by Reference Herein</b>	
		<b>Form</b>	<b>Date</b>
2-A	Asset Purchase Agreement between Registrant and R.P. Scherer South, Inc. dated June 21, 2000.	Current Report on Form 8-K, as Exhibit 2.1	August 9, 2000
3-A	Certificate of Incorporation, as amended through July 29, 2009.	Quarterly Report on Form 10-Q for the quarter ended June 30, 2009, as Exhibit 3.1	August 4, 2009
3-B	By laws	Registration Statement on Form S-1 (Registration No. 33-15429) as an Exhibit	
3-C	Amended and Restated Certificate of Designation, Preferences, and Rights of Series A Preferred Stock.	Current Report on Form 8-K, as Exhibit 3.C	December 19, 2006
3-D	Certificate of Amendment of Certificate of Incorporation.	Current Report on Form 8-K, as Exhibit 3.1	June 30, 2011
3-E	Certificate of Amendment of Certificate of Incorporation.	Current Report on Form 8-K, as Exhibit 3.1	January 13, 2014
4-A	Common Stock Certificate.	Registration on Form S-3 (Registration No.333-162968), as Exhibit 4.1	November 6, 2009
4-B	Form of Warrant to Purchase Shares of Common Stock	Current Report on Form 8-K as Exhibit 10.3	October 22, 2009
10-A	Registrant's 1997 Employee Stock Purchase Plan, as amended to date.*	Definitive Proxy on Schedule 14A, as Exhibit B	June 3, 2011
10-B	Lease Agreement between Registrant and Metropolitan Life Insurance Company for lease of Registrant's executive offices in Redwood City dated as of November 17, 1997.	Annual Report on 10-K for the year ended December 31, 1997, as Exhibit 10-E	March 30, 1998
10-C	Registrant's 2002 Equity Incentive Plan dated June 13, 2002.*	Registration on Form S-8 (Registration No.333-90428), as Exhibit No. 99.1	June 13, 2002
10-D	Form of Amended and Restated 2007 Equity Incentive Plan.*	Definitive Proxy on Schedule 14A, as Exhibit A	June 3, 2011

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Exhibit Number	Description	Incorporated by Reference Herein	
		Form	Date
10-E	Form of 2007 Equity Incentive Plan Stock Option Agreement.*	Registration on Form S-8 (Registration No. 333-148660), as Exhibit 4.3	January 14, 2008
10-F	Form of 2007 Equity Incentive Plan Restricted Stock Unit Agreement.*	Registration on Form S-8 (Registration No. 333-148660), as Exhibit 4.4	January 14, 2008
10-G	Form of 2007 Equity Incentive Plan Restricted Stock Award Agreement.*	Annual Report on 10-K for the year ended December 31, 2007, as Exhibit 10-O	March 31, 2008
10-H	Form of 2002 Equity Incentive Plan Stock Option Agreement.*	Annual Report on 10-K for the year ended December 31, 2007, as Exhibit 10-P	March 31, 2008
10-I	Form of 2002 Equity Incentive Plan Restricted Award Agreement.*	Annual Report on 10-K for the year ended December 31, 2007, as Exhibit 10-Q	March 31, 2008
10-J	Amendment to the Registrant's Non-Qualified Stock Plan.*	Quarterly Report on 10-Q for the quarter ended September 30, 2007, as Exhibit 10.16	November 14, 2007
10-K	Form of Indemnification Agreement.*	Annual Report on Form 10-K for the year ended December 31, 2007, as Exhibit 10-S	March 31, 2008
10-L	Registrant's Non-Qualified Stock Plan dated June 13, 2002.*	Registration Statement on Form S-8 (Registration No.333-90428), as Exhibit No. 99.2	June 13, 2002
10-M	Securities Purchase Agreement, dated as of October 19, 2009, by and among the Registrant and the purchasers listed therein.	Current Report on Form 8-K, as Exhibit 10.1	October 22, 2009
10-N	Registration Rights Agreement, dated as of October 22, 2009, by and among the Registrant and the purchasers listed therein.	Current Report on Form 8-K, as Exhibit 10.2	October 22, 2009
10-O	Securities Purchase Agreement, dated as of April 24, 2011, by and among the Company and the purchasers listed therein.	Current Report on Form 8-K, as Exhibit 10.1	April 28, 2011
10-P	Form of Senior Secured Convertible Note due 2021.	Current Report on Form 8-K, as Exhibit 10.2	April 28, 2011

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<u>Exhibit Number</u>	<u>Description</u>	<u>Incorporated by Reference Herein</u>	
		<u>Form</u>	<u>Date</u>
10-Q	Security Agreement, dated as of April 24, 2011, by and between the Company and Tang Capital Partners, LP, as Agent for the Purchasers.	Current Report on Form 8-K, as Exhibit 10.3	April 28, 2011
10-R	Second Amendment to Lease, effective as of April 1, 2011, by and between the Company and Metropolitan Life Insurance Company.	Current Report on Form 8-K, as Exhibit 10.4	April 28, 2011
10-S	Management Retention Agreement, dated as of April 25, 2011, by and between the Company and Michael A. Adam.*	Current Report on Form 8-K, as Exhibit 10.6	April 28, 2011
10-T	Securities Purchase Agreement, dated June 29, 2011, by and between the Company and the purchasers listed on Schedule I thereto.	Current Report on Form 8-K, as Exhibit 10.1	June 30, 2011
10-U	Amendment to Senior Secured Convertible Note Due 2021, dated, June 29, 2011, by and between the Company and the purchasers named in the Securities Purchase Agreement, dated April 24, 2011, by and among the Company and the purchasers listed therein.	Current Report on Form 8-K, as Exhibit 10.2	June 30, 2011
10-V	Third Amendment to Lease, effective as of July 28, 2011, by and between the Company and Metropolitan Life Insurance Company.	Current Report on Form 8-K, as Exhibit 10.1	August 3, 2011
10-W	Securities Purchase Agreement, dated July 25, 2012, by and between the Company and the purchasers named therein.	Current Report on Form 8-K, as Exhibit 10.1	July 25, 2012
10-X	Registration Rights Agreement, dated July 25, 2012, by and between the Company and the purchasers named therein.	Current Report on Form 8-K, as Exhibit 10.2	July 25, 2012

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<u>Exhibit Number</u>	<u>Description</u>	<u>Incorporated by Reference Herein</u>	
		<u>Form</u>	<u>Date</u>
10-Y	Management Retention Agreement as of December 3, 2012, by and between the Company and Mark S. Gelder, M.D.*	Annual Report on Form 10-K for the year ended December 31, 2012, as Exhibit 10-AH	March 1, 2013
10-Z	Executive Employment Agreement, dated May 1, 2013, by and between the Company and Dr. Barry Quart.*	Quarterly Report on Form 10-Q for the quarter ended March 31, 2013, as Exhibit 10-AI	May 10, 2013
10-AA	Executive Employment Agreement, dated May 1, 2013, by and between the Company and Robert Rosen.*	Quarterly Report on Form 10-Q for the quarter ended March 31, 2013, as Exhibit 10-AJ	May 10, 2013
10-AB	Executive Employment Agreement, dated May 1, 2013, by and between the Company and Stephen Davis.*	Quarterly Report on Form 10-Q for the quarter ended March 31, 2013, as Exhibit 10-AK	May 10, 2013
10-AC	Form of Non-Qualified Stock Option Agreement.	Quarterly Report on Form 10-Q for the quarter ended June 30, 2013, as Exhibit 10-AL	August 8, 2013
10-AD	Amendment to Management Retention Agreement, dated as of April 25, 2011, as amended May 29, 2013 (as amended, the "Retention Agreement")*	Quarterly Report on Form 10-Q for the quarter ended June 30, 2013, as Exhibit 10-AM	August 8, 2013
10-AE	Offer Letter dated November 10, 2012 between the Company and Mark S. Gelder, M.D.*	Filed herewith	
10-AF	Offer Letter dated October 16, 2013 between the Company and Brian G. Drazba.*	Filed herewith	
10-AG	Management Retention Agreement as of October 23, 2013, by and between the Company and Brian G. Drazba.*	Filed herewith	
10-AH	Executive Employment Agreement, dated November 1, 2013, by and between the Company and Paul Marshall.*	Filed herewith	
23.1	Consent of Independent Registered Public Accounting Firm.	Filed herewith	
24.1	Power of Attorney	Filed herewith (on signature page)	

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<b>Exhibit Number</b>	<b>Description</b>	<b>Incorporated by Reference Herein</b>	
		<b>Form</b>	<b>Date</b>
31.1	Certification of Chief Executive Officer pursuant to Rules 13A-15(e) Promulgated under the Securities Exchange Act of 1934 as amended.	Filed herewith	
31.2	Certification of Chief Financial Officer pursuant to Rules 13A-15(e) Promulgated under the Securities Exchange Act of 1934 as amended.	Filed herewith	
32.1	Certification of Chief Executive Officer and Chief Financial Officer pursuant to 18 U.S.C. Section 1350, as adopted pursuant to Section 906 of the Sarbanes-Oxley Act of 2002.	Furnished herewith	
101.INS	XBRL Instance Document	Furnished herewith	
101.SCH	XBRL Taxonomy Extension Schema Document	Furnished herewith	
101.CAL	XBRL Taxonomy Extension Calculation Linkbase Document	Furnished herewith	
101.DEF	XBRL Extension Definition	Furnished herewith	
101.LAB	XBRL Taxonomy Extension Label Linkbase Document	Furnished herewith	
101.PRE	XBRL Taxonomy Extension Presentation Linkbase Document	Furnished herewith	

\* Management contract or compensatory plans.



November 10, 2012

Mark Gelder, M.D.

Dear Mark:

It is with great pleasure that we extend an offer of employment with A.P. Pharma, Inc. for the position of Chief Medical Officer of A.P. Pharma, Inc., reporting to the company's Chief Executive Officer. If you accept, it is expected that your first day of employment will be no later than November 26, 2012. Your employment will be "at-will," meaning that either you or the Company can terminate the employment relationship at any time, with or without cause.

You will receive an annual salary of \$350,000.00, payable on a bi-weekly basis in accordance with the Company's normal payroll practices. In addition, following your first day of employment you will receive a \$40,000.00 hiring bonus. Subject to your commencement of employment, you will be granted an option to purchase up to 4,000,000 shares of A.P. Pharma Common Stock, with an exercise price equal to the fair value of our common stock on the date of grant. The option will vest over four years, with 25% cliff vesting at the end of the first year, and then the remaining 75% vesting monthly over the final three years. You will participate in the company's annual management cash bonus program, and will be eligible to receive an annual bonus in a target amount equal to 30% of your annual salary. You will be eligible to participate in all of the company's employee benefit plans and programs and a summary of 2012 benefits accompanies this letter.

If you elect to leave A.P. Pharma prior to your one-year anniversary, you agree to repay your hiring bonus to the Company on your termination.

A Management Retention Agreement also accompanies this letter. This agreement contains certain standard terms and conditions of your employment and severance benefits in certain circumstances.

You will accrue 4.62 hours of vacation per pay period equivalent to 3 weeks every 12 months of employment. In addition, the company typically provides employees with 2 personal days, 9 fixed holidays and 8 days of sick time per year.

Enclosed are copies of our At Will Statement, Confidential Disclosure Agreement, Conflicts of Interest Agreement, and a list of acceptable documents needed to complete an Employment Verification Form I-9 (which will be completed on your first day of employment). All of these forms need to be completed prior to initiating active employment.

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This offer letter, along with the accompanying Management Retention Agreement, constitute our entire agreement and supersede our prior agreements, negotiations and understandings. This agreement may only be amended in writing by mutual consent. This employment agreement will be governed by California laws, without regard to conflict-of-law principles.

This offer will expire as of 5 p.m. (Pacific Time) on Friday, November 16, 2012. If you decide to accept this offer, please scan and e-mail the signed offer letter to my attention at [jwhelan@appharma.com](mailto:jwhelan@appharma.com).

We look forward to having you join A.P. Pharma. If you have any questions regarding any of the information above, please feel free to contact me.

Sincerely,

/s/ John Whelan

John Whelan  
Chief Executive Officer  
[jwhelan@appharm.com](mailto:jwhelan@appharm.com)  
650.366.2626 x223

**The provisions of this offer of employment have been read, are understood, and the offer is herewith accepted.**

/s/ Mark Gelder, M.D.

Mark Gelder, M.D.

Date



October 16, 2013

Brian G. Drazba

Dear Brian:

It is with great pleasure that we extend an offer of employment with A.P. Pharma, Inc. for the position of Chief Financial Officer of A.P. Pharma, Inc., reporting to me the company's Executive Vice President and Chief Operating Officer. Your employment will be "at-will," meaning that either you or the Company can terminate the employment relationship at any time, with or without cause.

You will receive an annual salary of \$285,000.00, payable on a bi-weekly basis in accordance with the Company's normal payroll practices. Subject to your commencement of employment, you will be granted an option to purchase up to 2,000,000 shares of A.P. Pharma Common Stock, with an exercise price equal to the fair value of our common stock on the date of grant. The option will vest over four years, with 25% cliff vesting at the end of the first year, and then the remaining 75% vesting monthly over the final three years. You will participate in the company's annual management cash bonus program, and will be eligible to receive an annual bonus in a target amount equal to 30% of your annual salary. You will be eligible to participate in all of the company's employee benefit plans and programs and a summary of 2013 benefits accompanies this letter.

A Management Retention Agreement also accompanies this letter. This agreement contains certain standard terms and conditions of your employment and severance benefits in certain circumstances.

You will accrue 4.62 hours of vacation per pay period equivalent to 3 weeks every 12 months of employment. In addition, the company typically provides employees with 2 personal days, 9 fixed holidays and 8 days of sick time per year.

Enclosed are copies of our At Will Statement, Confidential Disclosure Agreement, Conflicts of Interest Agreement, and a list of acceptable documents needed to complete an Employment Verification Form I-9 (which will be completed on your first day of employment). All of these forms need to be completed prior to initiating active employment.

This offer letter, along with the accompanying Management Retention Agreement, constitute our entire agreement and supersede our prior agreements, negotiations and understandings. This agreement may only be amended in writing by mutual consent. This employment agreement will be governed by California laws, without regard to conflict-of-law principles.

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This offer will expire as of 5 p.m. (Pacific Time) on Friday, October 25, 2013. If you decide to accept this offer, please scan and e-mail the signed offer letter to my attention at [sdavis@appharma.com](mailto:sdavis@appharma.com).

We look forward to having you join A.P. Pharma. If you have any questions regarding any of the information above, please feel free to contact me.

Sincerely,

/s/ Steve Davis

Steve Davis  
Executive Vice President & COO  
[sdavis@appharm.com](mailto:sdavis@appharm.com)

**The provisions of this offer of employment have been read, are understood, and the offer is herewith accepted.**

/s/ Brian G. Drazba  
Brian G. Drazba

10/21/13  
Date

**A.P. PHARMA, INC.****MANAGEMENT RETENTION AGREEMENT**

This Management Retention Agreement (the "Agreement") is dated as of October 23, 2013, by and between Brian G. Drazba ( "Employee") and A.P. Pharma, Inc., a Delaware corporation (the "Company"). This Agreement is intended to provide Employee with certain benefits described herein upon the occurrence of specific events.

## RECITALS

A. The Company's Board of Directors believes it is in the best interests of the Company and its shareholders to retain Employee and provide incentives to Employee to continue in the service of the Company.

B. The Board of Directors further believes that it is imperative to provide Employee with certain benefits upon Employee's Involuntary Termination or a Change of Control, which benefits are intended to provide Employee with financial security and provide sufficient income and encouragement to Employee to remain employed with the Company, notwithstanding the possibility of a Change of Control.

D. To accomplish the foregoing objectives, the Board of Directors has directed the Company, upon execution of this Agreement by Employee, to agree to the terms provided in this Agreement.

It is therefore agreed as follows:

1. **At-Will Employment.** The Company and Employee acknowledge that Employee's employment is and shall continue to be at-will, as defined under applicable law, and that Employee's employment with the Company may be terminated by either party at any time for any or no reason. If Employee's employment terminates for any reason, Employee shall not be entitled to any payments, benefits, damages, award or compensation other than as provided in this Agreement or otherwise agreed to by the Company. The terms of this Agreement shall terminate upon the earlier of: (i) the date on which Employee ceases to be employed as a corporate officer of the Company, other than as a result of an Involuntary Termination; or (ii) the date that all obligations of the parties hereunder have been satisfied. A termination of the terms of this Agreement pursuant to the preceding sentence shall be effective for all purposes, except that such termination shall not affect the payment or provision of compensation or benefits on account of a termination of employment occurring prior to the termination of the terms of this Agreement. The rights and duties created by this Section 1 are contingent upon Employee's release of claims against the Company (at the time of termination in a form reasonably satisfactory to the Company) and may not be modified in any way except by a written agreement executed by an officer of the Company upon direction from the Board of Directors.

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## 2. **Benefits Upon Termination of Employment.**

(a) **Severance Upon Involuntary Termination.** In the event that Employee suffers an Involuntary Termination at any time under circumstances other than as covered in paragraph 2(b) below, then in addition to all salary and bonuses accrued as of the date of Employee's termination of employment, Employee will be entitled to receive severance benefits as follows: (i) during the period commencing on the date of Employee's termination and ending on the date six (6) months after the effective date of the termination (the "**Severance Period**") the Company shall pay to Employee an amount equal to the monthly base salary which Employee was receiving immediately prior to the Involuntary Termination in accordance with the Company's standard payroll practices; (ii) one-half the average bonus paid by the Company to Employee for services during each of the three 12- month periods (or such shorter period of time during which Employee was eligible for a bonus) prior to the Involuntary Termination date, which payments shall be paid during the Severance Period in accordance with the Company's standard payroll practices; and (iii) reimbursement for or continuation of payment by the Company of its portion of the health insurance benefits provided to Employee immediately prior to the Involuntary Termination pursuant to the terms of the Consolidated Omnibus Budget Reconciliation Act of 1985, as amended ("**COBRA**") or other applicable law through the earlier of the end of the Severance Period or the date upon which Employee is no longer eligible for such COBRA or other benefits under applicable law. In addition, Employee's stock options, restricted stock and other equity awards shall immediately vest, become exercisable and/or the restrictions thereon lapse with respect to that number of shares of Company common stock that otherwise would have vested during the Severance Period had Employee's employment continued. Employee's stock options, restricted stock and other equity awards shall otherwise be subject to the terms of the plan and option or award agreement pursuant to which such options and other equity awards were granted.

(b) **Severance Upon a Change in Control.** In the event that Employee suffers an Involuntary Termination within the twelve (12) month period following the effective date of a Change of Control, then in addition to all salary and bonuses accrued as of the date of Employee's termination of employment, Employee will be entitled to receive severance benefits as follows: (i) during the period commencing on the date of Employee's termination and ending on the date twelve (12) months after the effective date of the termination (the "**Change of Control Severance Period**") the Company shall pay to Employee an amount equal to the greater of (A) the monthly base salary which Employee was receiving immediately prior to the Involuntary Termination or (B) the monthly base salary which Employee was receiving immediately prior to the Change of Control, in each case, in accordance with the Company's standard payroll practices; (ii) the average bonus paid by the Company to Employee for services during each of the three 12- month periods (or such shorter period of time during which Employee was eligible for a bonus) prior to the Involuntary Termination date, which payments shall be paid during the Change of Control Severance Period in accordance with the Company's standard payroll practices; and (iii) reimbursement for or continuation of payment by the Company of its portion of the health insurance benefits provided to Employee immediately prior to the Involuntary Termination pursuant to the terms of COBRA or other applicable law through the earlier of the end of the Change of Control Severance Period or the date upon which Employee is no longer

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eligible for such COBRA or other benefits under applicable law. In addition, Employee's stock options, restricted stock and other equity awards shall immediately vest, become exercisable and/or the restrictions thereon lapse with respect to one hundred percent (100%) of the shares of Company common stock subject thereto. Employee's stock options, restricted stock and other equity awards shall otherwise be subject to the terms of the plan and option or award agreement pursuant to which such options and other equity awards were granted.

(c) **Termination for Cause.** Notwithstanding any other provision of this Agreement, if Employee's employment is terminated for Cause at any time, then Employee shall not be entitled to receive payment of any severance benefits or any continuation or acceleration of stock option vesting or relinquishment of forfeiture and transfer restrictions on restricted stock awards. Employee will receive payment(s) for all salary and bonuses accrued as of the date of Employee's termination of employment.

(d) **Voluntary Resignation.** If Employee voluntarily resigns from the Company under circumstances which do not constitute an Involuntary Termination, then Employee shall not be entitled to receive payment of any severance benefits, or option acceleration, or relinquishment of forfeiture and transfer restrictions. Employee will receive payment(s) for all salary and bonuses accrued as of the date of Employee's termination of employment.

3. **Definition of Terms.** The following terms referred to in this Agreement shall have the following meanings:

(a) **"Cause"** means any of the following: (i) Employee's theft, dishonesty, willful misconduct, breach of fiduciary duty for personal profit, or falsification of any Company or Affiliate documents or records; (ii) Employee's material failure to abide by a Company's or Affiliate's code of conduct or other policies (including without limitation, policies relating to confidentiality and reasonable workplace conduct); (iii) Employee's unauthorized use, misappropriation, destruction or diversion of any tangible or intangible asset or corporate opportunity of the Company or an Affiliate (including, without limitation, Employee's improper use or disclosure of confidential or proprietary information); (iv) any intentional act by Employee which has a material detrimental effect on the Company or an Affiliate's reputation or business; (v) Employee's repeated failure or inability to perform any reasonable assigned duties after written notice from the Company or an Affiliate (including, without limitation, habitual absence from work for reasons other than illness), and a reasonable opportunity to cure, such failure or inability; (vi) any material breach by Employee of any employment or service agreement between Employee and the Company or an Affiliate, which breach is not cured pursuant to the terms of such agreement; or (vii) Employee's conviction (including any plea of guilty or *nolo contendere*) of any criminal act involving fraud, dishonesty, misappropriation or moral turpitude, or which impairs Employee's ability to perform his or her duties with the Company or an Affiliate.

(b) **"Change in Control"** means the occurrence of any of the following:

(i) an Ownership Change Event or a series of related Ownership Change Events (collectively, a **"Transaction"**) in which the stockholders of the Company

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immediately before the Transaction do not retain immediately after the Transaction, in substantially the same proportions as their ownership of shares of the Company's voting stock immediately before the Transaction, direct or indirect beneficial ownership of more than fifty percent (50%) of the total combined voting power of the outstanding voting securities of the Company or such surviving entity immediately outstanding after the Transaction, or, in the case of an Ownership Change Event the entity to which the assets of the Company were transferred (the "**Transferee**"), as the case may be; or

(ii) the liquidation or dissolution of the Company.

For purposes of the preceding sentence, indirect beneficial ownership shall include, without limitation, an interest resulting from ownership of the voting securities of one or more corporations or other business entities which own the Company or the Transferee, as the case may be, either directly or through one or more subsidiary corporations or other business entities. The Board shall have the right to determine whether multiple sales or exchanges of the voting securities in the Company or multiple Ownership Change Events are related, and its determination shall be final, binding and conclusive. The Board may also, but need not, specify that other transactions or events constitute a Change in Control.

(c) "**Involuntary Termination**" shall include any termination by the Company other than for Cause and Employee's voluntary termination within sixty days following the occurrence of any of the following events without Employee's written consent: (i) a material reduction or change in job duties, responsibilities and requirements inconsistent with Employee's position with the Company and Employee's prior duties, responsibilities and requirements or a material negative change in Employee's reporting relationship; (ii) a material reduction of Employee's base compensation (other than in connection with a general decrease in base salaries for most officers of the Company or successor corporation); or (iii) Employee's refusal to relocate to a facility or location more than forty miles from the Company's current location, provided that Employee will not resign due to such change, reduction or relocation without first providing the Company with written notice of the event or events constituting the grounds for his voluntary resignation within thirty days of the initial existence of such grounds and a reasonable cure period of not less than thirty days following the date of such notice.

(d) "**Ownership Change Event**" means the occurrence of any of the following with respect to the Company: (i) the direct or indirect sale or exchange in a single or series of related transactions by the stockholders of the Company of more than fifty percent (50%) of the voting stock of the Company; (ii) a merger or consolidation in which the Company is a party; or (iii) the sale, exchange, or transfer of all or substantially all of the assets of the Company.

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#### 4. Limitation and Conditions on Payments.

(a) **Parachute Payments.** In the event that the severance and other benefits provided for in this Agreement to the Employee: (i) constitute “parachute payments” within the meaning of Section 280G of the Internal Revenue Code of 1986, as amended (the “Code”); and (ii) but for this Section, would be subject to the excise tax imposed by Section 4999 of the Code, then the Employee’s severance benefits under Sections 2(a) and 2(b) shall be payable either:

(i) in full; or

(ii) as to such lesser amount which would result in no portion of such severance benefits being subject to excise tax under Section 4999 of the Code, whichever of the foregoing amounts, taking into account the applicable federal, state and local income taxes and the excise tax imposed by Section 4999, results in the receipt by the Employee on an after-tax basis, of the greatest amount of severance benefits under Section 2(a) and 2(b), notwithstanding that all or some portion of such severance benefits may be taxable under Section 4999 of the Code. Any determination required under this Section 4 shall be made in writing by independent public accountants selected by the Company (the “Accountants”), whose determination shall be conclusive and binding upon the Employee and the Company for all purposes. For purposes of making the calculations required by this Section 4, the Accountants may make reasonable assumptions and approximations concerning applicable taxes and may rely on reasonable, good faith interpretations concerning the application of Section 280G and 4999 of the Code. The Company and the Employee shall furnish to the Accountants such information and documents as the Accountants may reasonably request in order to make a determination under this Section. The Company shall bear all costs the Accountants may reasonably incur in connection with any calculations contemplated by this Section 4. Any reduction in severance benefits required by this Section 4 shall occur in a manner necessary to provide the service provider with the greatest economic benefit. If more than one manner of reduction of severance benefits necessary to arrive at the reduced amount yields the greatest economic benefit to the service provider, the payments and benefits shall be reduced pro rata.

(b) **Release Prior to Receipt of Benefits.** Prior to the receipt of any benefits under this Agreement, Employee shall execute and allow to become effective, a release of claims agreement in a form acceptable to the Company (the “Release”) not later than fifty-two (52) days following Employee’s employment termination in the form provided by the Company. Such Release shall specifically relate to all of Employee’s rights and claims in existence at the time of such execution and shall confirm Employee’s obligations under the Company’s standard form of proprietary information agreement. In no event will severance benefits be provided to Employee until the Release becomes effective. In the event severance payments are delayed because of the effective date of the Release, the Company will pay Employee the severance payments, that Employee would otherwise have received under Section 2(a) on or prior to the effective date of the Release, on the first regular payroll pay day following the effective date of the release, with the balance of the payments being paid as originally scheduled.

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5. **Section 409A.** All severance payments to be made upon a termination of employment under this Agreement may be made only upon a "separation of service" within the meaning of Section 409A of the Code and the Department of Treasury regulations and other guidance promulgated thereunder. Notwithstanding any provision to the contrary in this Agreement, if Employee is deemed by the Company at the time of Employee's separation from service to be a "specified employee" for purposes of Code Section 401A(a)(2)(B)(i), to the extent delayed commencement of any portion of the benefits to which Employee is entitled under this Agreement is required in order to avoid a prohibited distribution under Code Section 409A(a)(2)(B)(i), such portion of Employee's benefits shall not be provided to Employee prior to the earlier of (i) the expiration of the six-month period measured from the date of Employee's "separation of service" with the Company or (ii) the date of Employee's death. Upon the first business day following the expiration of the applicable Code Section 409A(a)(2)(B)(i) period, all payments deferred pursuant to this Section 5 shall be paid in a lump sum to Employee, and any remaining payments due under the Agreement shall be paid as otherwise provided herein. For purposes of Code Section 409A (including, without limitation, for purposes of Treasury Regulation Section 1.409A-2(b)(2)(iii)), Employee's right to receive installment payments under this Agreement shall be treated as a right to receive a series of separate payments and, accordingly, each installment payment hereunder shall at all times be considered a separate and distinct payment. It is intended that none of the severance payments and benefits to be provided hereunder will be subject to Section 409A of the Code and any ambiguities herein will be interpreted to be so exempt. Employee and the Company agree to work together in good faith to consider amendments to this Agreement and to take such reasonable actions which are necessary, appropriate or desirable to avoid imposition of any additional tax or income recognition prior to actual payment to Employee under Section 409A of the Code. Notwithstanding anything to the contrary contained herein, to the extent that any amendment to this Agreement with respect to the payment of any severance payments or benefits would constitute under Code Section 409A a delay in a payment or a change in the form of payment, then such amendment must be done in a manner that complies with Code Section 409A(a)(4)(C).

6. **Conflicts.** Employee represents that Employee's performance of all the terms of this Agreement will not breach any other agreement to which Employee is a party. Employee has not, and will not during the term of this Agreement, enter into any oral or written agreement in conflict with any of the provisions of this Agreement. Employee further represents that Employee is entering into or has entered into an employment relationship with the Company of Employee's own free will and that Employee has not been solicited as an employee in any way by the Company.

7. **Successors.** Any successor to the Company (whether direct or indirect and whether by purchase, lease, merger, consolidation, liquidation or otherwise) to all or substantially all of the Company's business and/or assets shall assume the obligations under this Agreement and agree expressly to perform the obligations under this Agreement in the same manner and to the same extent as the Company would be required to perform such obligations in the absence of a succession. The terms of this Agreement and all of Employee's rights hereunder and thereunder shall inure to the benefit of, and be enforceable by, Employee's personal or legal representatives, executors, administrators, successors, heirs, distributees, devisees and legatees.

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8. **Notice.** Notices and all other communications contemplated by this Agreement shall be in writing and shall be deemed to have been duly given when personally delivered or when mailed by U.S. registered or certified mail, return receipt requested and postage prepaid. Mailed notices to Employee shall be addressed to Employee at the home address which Employee most recently communicated to the Company in writing. In the case of the Company, mailed notices shall be addressed to its corporate headquarters, and all notices shall be directed to the attention of its Secretary.

9. **Miscellaneous Provisions.**

(a) **No Duty to Mitigate.** Employee shall not be required to mitigate the amount of any payment contemplated by this Agreement (whether by seeking new employment or in any other manner), nor shall any such payment be reduced by any earnings that Employee may receive from any other source.

(b) **Waiver.** No provision of this Agreement shall be modified, waived or discharged unless the modification, waiver or discharge is agreed to in writing and signed by Employee and by an authorized officer of the Company (other than Employee). No waiver by either party of any breach of, or of compliance with, any condition or provision of this Agreement by the other party shall be considered a waiver of any other condition or provision or of the same condition or provision at another time.

(c) **Whole Agreement.** No agreements, representations or understandings (whether oral or written and whether express or implied) which are not expressly set forth in this Agreement have been made or entered into by either party with respect to the subject matter hereof. This Agreement supersedes any agreement of the same title and concerning similar subject matter dated prior to the Effective Date, and by execution of this Agreement both parties agree that any such predecessor agreement shall be deemed null and void.

(d) **Choice of Law.** The validity, interpretation, construction and performance of this Agreement shall be governed by the laws of the State of California without reference to conflict of laws provisions.

(e) **Severability.** If any term or provision of this Agreement or the application thereof to any circumstance shall, in any jurisdiction and to any extent, be invalid or unenforceable, such term or provision shall be ineffective as to such jurisdiction to the extent of such invalidity or unenforceability without invalidating or rendering unenforceable the remaining terms and provisions of this Agreement or the application of such terms and provisions to circumstances other than those as to which it is held invalid or unenforceable, and a suitable and equitable term or provision shall be substituted therefore to carry out, insofar as may be valid and enforceable, the intent and purpose of the invalid or unenforceable term or provision.

(f) **Arbitration.** Any dispute or controversy arising under or in connection with this Agreement may be settled at the option of either party by binding arbitration in the County of San Mateo, California, in accordance with the rules of the American Arbitration Association then in effect. Judgment may be entered on the arbitrator's award in any court having jurisdiction. Punitive damages shall not be awarded.

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(g) **Legal Fees and Expenses**. The parties shall each bear their own expenses, legal fees and other fees incurred in connection with this Agreement.

(h) **No Assignment of Benefits**. The rights of any person to payments or benefits under this Agreement shall not be made subject to option or assignment, either by voluntary or involuntary assignment or by operation of law, including (without limitation) bankruptcy, garnishment, attachment or other creditor's process, and any action in violation of this Section 9(h) shall be void.

(i) **Employment Taxes**. All payments made pursuant to this Agreement will be subject to withholding of applicable income and employment taxes.

(j) **Assignment by Company**. The Company may assign its rights under this Agreement to an affiliate, and an affiliate may assign its rights under this Agreement to another affiliate of the Company or to the Company. In the case of any such assignment, the term "Company" when used in a section of this Agreement shall mean the corporation that actually employs the Employee.

(k) **Counterparts**. This Agreement may be executed in counterparts, each of which shall be deemed an original, but all of which together will constitute one and the same instrument.

(l) **Renewal**. This Agreement shall remain in effect until December 31, 2013 and shall be automatically renewed for additional one year periods unless not later than three months prior to December 31<sup>st</sup> of any year either party gives written notice to the other party of the intention to terminate the Agreement effective December 31<sup>st</sup> of that year, provided, that in no event shall this Agreement terminate during the twelve (12) month period commencing upon a Change of Control.

*[Signature page follows]*

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The parties have executed this Agreement on the date first written above.

**A.P. PHARMA, INC.**

By: /s/ Barry D. Quart  
Name: Barry D. Quart, Pharm.D.  
Title: Chief Executive Officer

**EMPLOYEE**

Signature: /s/ Brian G. Drazba  
                  Brian G. Drazba  
Address:

**EXECUTIVE EMPLOYMENT AGREEMENT**

This **EMPLOYMENT AGREEMENT** (the "**Agreement**") is made and entered into effective as of November 1, 2013 (the "**Effective Date**"), by and between A.P. PHARMA, INC. (the "**Company**"), and PAUL MARSHALL (the "**Executive**"). The Company and the Executive are hereinafter collectively referred to as the "**Parties**", and individually referred to as a "**Party**".

**AGREEMENT**

In consideration of the foregoing and the mutual promises and covenants herein contained, and for other good and valuable consideration, the Parties, intending to be legally bound, agree as follows:

**1. EMPLOYMENT.**

**1.1 Title.** The Executive shall initially have the title of Senior Vice President of Technical Operations of the Company and shall serve in such other capacity or capacities as the Company may from time to time prescribe and to which the Executive agrees. The Executive shall initially report to the Chief Executive Officer and the Board of Directors of the Company (the "**Board**").

**1.2 Duties.** The Executive shall do and perform all services, acts or things necessary or advisable to manage and conduct the business of the Company and which are normally associated with the position of Senior Vice President of Technical Operations, consistent with the Bylaws of the Company and as required by the Board.

**1.3 Policies and Practices.** The employment relationship between the Parties shall be governed by the policies and practices established by the Company and the Board. The Executive acknowledges that he has read the Company's employee handbook and other governing policies, which will govern the terms and conditions of his employment with the Company, along with this Agreement. In the event that the terms of this Agreement differ from or are in conflict with the Company's policies or practices or the Company's employee handbook, this Agreement shall control.

**2. LOYAL AND CONSCIENTIOUS PERFORMANCE; NONCOMPETITION.**

**2.1 Loyalty.** During the Executive's employment by the Company, the Executive shall devote the Executive's full business energies, interest, abilities and productive time to the proper and efficient performance of the Executive's duties under this Agreement. Notwithstanding the foregoing, the Executive may provide occasional consulting that are not competitive to the Company.

**2.2 Covenant Not to Compete.** Except with the prior written consent of the Board, which shall not be unreasonably withheld, and except the provisions included in Section 2.1 above, the Executive will not, during his employment by the Company, engage in competition with the Company and/or any of its Affiliates, either directly or indirectly, in any manner or capacity, as adviser, principal, agent, affiliate, promoter, partner, officer, director, employee, stockholder,

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owner, co-owner, consultant, or member of any association or otherwise, in any phase of the business of developing, manufacturing and marketing of products or services which are in the same field of use or which otherwise compete with the products or services or proposed products or services of the Company and/or any of its Affiliates. For purposes of this Agreement, "**Affiliate**" means, with respect to any specific entity, any other entity that, directly or indirectly, through one or more intermediaries, controls, is controlled by or is under common control with such specified entity.

**2.3 Agreement Not to Participate in Company's Competitors** . During the term of this agreement, and except the provisions included in Section 2.1 above, the Executive agrees not to acquire, assume or participate in, directly or indirectly, any position, investment or interest known by the Executive to be adverse or antagonistic to the Company, its business or prospects, financial or otherwise or in any company, person or entity that is, directly or indirectly, in competition with the business of the Company or any of its Affiliates. Ownership by the Executive, as a passive investment, of less than two percent (2%) of the outstanding shares of capital stock of any corporation with one or more classes of its capital stock listed on a national securities exchange or publicly traded on the Nasdaq Stock Market or in the over-the-counter market shall not constitute a breach of this paragraph.

### **3. COMPENSATION OF THE EXECUTIVE.**

**3.1 Base Salary.** The Company shall pay the Executive a base salary of at least \$395,000 per year, less payroll deductions and all required withholdings payable in regular periodic payments in accordance with Company policy. Such base salary shall be prorated for any partial year of employment on the basis of a 365-day fiscal year.

**3.2 Performance Bonus.** In addition to the Executive's base salary, the Executive shall be eligible for a performance bonus based upon the Executive's and the Company's achievement of specified objectives established by the Board during the first quarter of each year after consultation with the Executive, as evaluated by the Board in its discretion. The target bonus for full achievement of all objectives shall be 40% of the Executive's Base Salary.

### **3.3 Equity Incentives.**

(a) As of the Effective Date, Executive will be granted an option to purchase up to 6,000,000 shares of the Company's common stock (the "**Option**"). Subject to the Executive continuing to serve as Senior Vice President of Technical Operations of the Company (except as set forth below), vesting of the Option will be as follows: (i) 4,200,000 shares (the "**Time-Based Shares**") vesting over a four-year period, with 1,050,000 shares vesting on the first anniversary of the date of grant, and then 87,500 shares vesting monthly thereafter over the next three years; (ii) 1,800,000 shares vesting upon receipt by the Company of FDA approval for its investigational new drug APF530 or any other drug product utilizing the Company's Biochronomer technology (either being a "**Qualified Drug**"), (the shares subject to vesting in clause (ii) being the "**Performance-Based Shares**"). If any vesting condition for the Performance-Based Shares is met within 60 days following Executive's termination of service either by the Company without Cause or by the Executive for Good Reason, then the vesting

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condition will be deemed satisfied with respect to that condition (such circumstance being a "**Post-Termination Vesting Event**"). The Option will have a ten-year term and will be treated as an incentive stock option to the maximum extent possible under applicable regulations, with the remainder being non-statutory stock options. The portion of the Option that is vested as of the date of termination of the Executive's service with the Company shall remain exercisable for a period of 90 days following termination. Any portion of the Option that vests as a result of a Post-Termination Vesting Event shall remain exercisable for a period of 90 days following the occurrence of such event.

(b) With respect to all stock options previously granted to Executive by the Company, which options are listed on Schedule A attached hereto (the "**Prior Options**"), the Company and Executive hereby agree that the Prior Options shall be vested with respect to that number of shares reflected under the "Options Considered Vested" column of Schedule A, with the remainder of the Prior Options to be forfeited as of the Effective Date.

**3.4 Changes to Compensation.** The Executive's compensation will be reviewed on a regular basis by the Company and may be changed from time to time as deemed appropriate. For clarity, the provisions of this Section 3.4 are not meant to supersede the right of the Executive to terminate for Good Reason in the event of a material reduction of Executive's Base Salary as provided in Section 4.5.3.

**3.5 Employment Taxes.** All of the Executive's compensation shall be subject to customary withholding taxes and any other employment taxes as are commonly required to be collected or withheld by the Company.

#### **4. TERMINATION.**

**4.1 Termination by the Company.** The Executive's employment by the Company shall be at will. The Executive's employment with the Company may be terminated by the Company at any time and for any reason or no reason, with or without "**Cause**" (as defined below), subject to the provisions of this Section 4.

**4.2 Termination by Mutual Agreement of the Parties.** The Executive's employment pursuant to this Agreement may be terminated at any time upon a mutual agreement in writing of the Parties. Any such termination of employment shall have the consequences specified in such agreement.

**4.3 Termination by the Executive.** The Executive's employment by the Company shall be at will. The Executive shall have the right to resign or terminate the Executive's employment at any time and for any reason, or no reason, with or without "**Good Reason**" (as defined below), subject to the provisions of this Section 4.

#### **4.4 Compensation Upon Termination.**

**4.4.1 With Cause or Without Good Reason.** If the Executive's employment shall be terminated by the Company for Cause, or if the Executive terminates employment hereunder for

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other than Good Reason, the Company shall pay the Executive's base salary and accrued and unused vacation benefits earned through the date of termination at the rate in effect at the time of termination, less standard deductions and withholdings, and the Company shall thereafter have no further obligations to the Executive under this Agreement.

**4.4.2 Without Cause or With Good Reason.** If the Executive's employment shall be terminated by the Company without Cause, or by the Executive for Good Reason, the Executive shall receive the payments specified in Section 4.4.1, and, in addition, within ten days of the Executive's delivery to the Company of a fully effective Release and Waiver in the form attached hereto as **Exhibit A**, within the applicable time period set forth therein, but in no event later than 45 days following termination of the Executive's employment, the Executive shall receive the following: (i) a lump sum payment equal to the sum of (A) the Executive's annual base salary then in effect and (B) the Executive's target performance bonus then in effect, less required deductions and withholdings; (ii) accelerated time-based vesting of shares subject to all stock awards issued by the Company, for the number of shares which would have vested accordingly had the Executive continued employment with the Company for a period of 12 months after termination (for the avoidance of doubt, which shall include partial accelerated vesting of the Time-Based Shares, but not the Performance-Based Shares); and (iii) reimbursement for or continuation of payment by the Company of its portion of the health insurance benefits provided to Executive immediately prior to termination pursuant to the terms of the Consolidated Omnibus Budget Reconciliation Act of 1985, as amended ( "**COBRA** ") or other applicable law for a period of up to 24 months from the date of termination.

**4.4.3 Change in Control.** If the Executive's employment shall be terminated by the Company without Cause, or by the Executive for Good Reason within three months before or within 12 months following a Change in Control, the Executive shall receive the payments specified in Section 4.4.1, and, in addition, within ten days of the Executive's delivery to the Company of a fully effective Release and Waiver in the form attached hereto as **Exhibit A**, within the applicable time period set forth therein, but in no event later than 45 days following termination of the Executive's employment, the Executive shall receive the following: (i) a lump sum payment equal to 150% of the Executive's annual base salary then in effect, less required deductions and withholdings; (ii) the greater of the Executive's target performance bonus then in effect, less required deductions and withholdings, or the Executive's performance bonus paid in the year preceding the year in which termination occurs, less required deductions and withholdings; and (iii) provided that the Executive timely elects continued coverage under COBRA, the COBRA benefit for a period of up to 24 months.

**4.4.4 Parachute Payment.** If any payment or benefit Executive would receive pursuant to a Change in Control or otherwise ("**Payment**") would (i) constitute a "parachute payment" within the meaning of Section 280G of the Code, and (ii) but for this sentence, be subject to the excise tax imposed by Section 4999 of the Code (the "**Excise Tax**"), then such Payment shall be reduced to the Reduced Amount. The "**Reduced Amount**" shall be either (x) the largest portion of the Payment that would result in no portion of the Payment being subject to the Excise Tax or (y) the largest portion, up to and including the total, of the Payment, whichever amount, after taking into account all applicable federal, state and local employment taxes, income taxes, and the Excise Tax (all computed at the highest applicable marginal rate), results in the Executive's

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receipt, on an after-tax basis, of the greater amount of the Payment notwithstanding that all or some portion of the Payment may be subject to the Excise Tax. If a reduction in payments or benefits constituting “parachute payments” is necessary so that the Payment equals the Reduced Amount, reduction shall occur in the following order: reduction of cash payments; cancellation of accelerated vesting of stock awards; reduction of employee benefits. In the event that acceleration of vesting of stock award compensation is to be reduced, such acceleration of vesting shall be cancelled in the reverse order of the date of grant of Executive’s stock awards.

The accounting firm engaged by the Company for general audit purposes as of the day prior to the effective date of the Change in Control shall perform the foregoing calculations. If the accounting firm so engaged by the Company is serving as accountant or auditor for the individual, entity or group effecting the Change in Control, then the Company shall appoint a nationally recognized accounting firm to make the determinations required hereunder. The Company shall bear all expenses with respect to the determinations by such accounting firm required to be made hereunder.

The accounting firm engaged to make the determinations hereunder shall provide its calculations, together with detailed supporting documentation, to the Executive and the Company within 15 calendar days after the date on which the Executive’s right to a Payment is triggered (if requested at that time by the Executive or the Company) or such other time as requested by the Executive or the Company. If the accounting firm determines that no Excise Tax is payable with respect to a Payment, either before or after the application of the Reduced Amount, it shall furnish the Executive and the Company with an opinion reasonably acceptable to the Executive that no Excise Tax will be imposed with respect to such Payment. Any good faith determinations of the accounting firm made hereunder shall be final, binding and conclusive upon the Executive and the Company.

**4.4.5 Application of Section 409A.** Notwithstanding anything to the contrary set forth herein, any payments and benefits provided under this Agreement (the “**Severance Benefits**”) that constitute “deferred compensation” within the meaning of Section 409A of the Internal Revenue Code of 1986, as amended (the “**Code**”) and the regulations and other guidance thereunder and any state law of similar effect (collectively “**Section 409A**”) shall not commence in connection with Executive’s termination of employment unless and until Executive has also incurred a “separation from service” (as such term is defined in Treasury Regulation Section 1.409A-1(h) (“**Separation From Service**”)), unless the Company reasonably determines that such amounts may be provided to Executive without causing Executive to incur the additional 20% tax under Section 409A.

It is intended that each installment of the Severance Benefits payments provided for in this Agreement is a separate “payment” for purposes of Treasury Regulation Section 1.409A- 2(b)(2)(i). For the avoidance of doubt, it is intended that payments of the Severance Benefits set forth in this Agreement satisfy, to the greatest extent possible, the exemptions from the application of Section 409A provided under Treasury Regulation Sections 1.409A-1(b)(4), 1.409A-1(b)(5) and 1.409A-1(b)(9). However, if the Company (or, if applicable, the successor entity thereto) determines that the Severance Benefits constitute “deferred compensation” under Section 409A and Executive is, on the termination of Executive’s service, a “specified employee”

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of the Company or any successor entity thereto, as such term is defined in Section 409A(a)(2)(B)(i) of the Code, then, solely to the extent necessary to avoid the incurrence of the adverse personal tax consequences under Section 409A, the timing of the Severance Benefit payments shall be delayed until the earlier to occur of: (i) the date that is six months and one day after Executive's Separation From Service or (ii) the date of Executive's death (such applicable date, the "**Specified Employee Initial Payment Date**"), and the Company (or the successor entity thereto, as applicable) shall (A) pay to Executive a lump sum amount equal to the sum of the Severance Benefit payments that Executive would otherwise have received through the Specified Employee Initial Payment Date if the commencement of the payment of the Severance Benefits had not been so delayed pursuant to this Section and (B) commence paying the balance of the Severance Benefits in accordance with the applicable payment schedules set forth in this Agreement.

Except to the extent that payments may be delayed until the Specified Employee Initial Payment Date pursuant to the preceding paragraph, on the first regular payroll pay day following the effective date of the Release and Waiver, the Company will pay Executive the Severance Benefits Executive would otherwise have received under the Agreement on or prior to such date but for the delay in payment related to the effectiveness of the Release and Waiver, with the balance of the Severance Benefits being paid as originally scheduled. All amounts payable under the Agreement will be subject to standard payroll taxes and deductions.

#### **4.5 Definitions.**

**4.5.1 Cause.** For purposes of this Agreement, "**Cause**" means that, in the reasonable determination of the Company, the Executive has:

- (i) been indicted for or convicted of or pleaded guilty or no contest to any felony or crime involving dishonesty that is likely to inflict or has inflicted demonstrable and material injury on the business of the Company;
- (ii) participated in any fraud against the Company;
- (iii) willfully and materially breached a Company policy;
- (iv) intentionally damaged any property of the Company thereby causing demonstrable and material injury to the business of the Company; or
- (v) engaged in conduct that, in the reasonable determination of the Company, demonstrates gross unfitness to serve.

Notwithstanding the foregoing, Cause shall not exist based on conduct described in clause (iii) above unless the conduct described in such clause has not been cured within 15 days following the Executive's receipt of written notice from the Company specifying the particulars of the conduct constituting Cause.

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**4.5.2 Change in Control.** For purposes of this Agreement, “**Change in Control**” means the occurrence of any of the following:

- (i) an Ownership Change Event or a series of related Ownership Change Events (collectively, a “**Transaction**”) in which the stockholders of the Company immediately before the Transaction do not retain immediately after the Transaction, in substantially the same proportions as their ownership of shares of the Company’s voting stock immediately before the Transaction, direct or indirect beneficial ownership of more than fifty percent (50%) of the total combined voting power of the outstanding voting securities of the Company or such surviving entity immediately outstanding after the Transaction, or, in the case of an Ownership Change Event the entity to which the assets of the Company were transferred (the “**Transferee**”), as the case may be; or
- (ii) the liquidation or dissolution of the Company.

For purposes of the preceding sentence, indirect beneficial ownership shall include, without limitation, an interest resulting from ownership of the voting securities of one or more corporations or other business entities which own the Company or the Transferee, as the case may be, either directly or through one or more subsidiary corporations or other business entities. The Board shall have the right to determine whether multiple sales or exchanges of the voting securities in the Company or multiple Ownership Change Events are related, and its determination shall be final, binding and conclusive. The Board may also, but need not, specify that other transactions or events constitute a Change in Control.

**4.5.3 Good Reason.** “**Good Reason**” for the Executive to terminate the Executive’s employment hereunder shall mean the occurrence of any of the following events without the Executive’s consent:

- (i) a material reduction (20% or more) by the Company of the Executive’s Base Salary as initially set forth herein or as the same may be increased from time to time;
- (ii) a material reduction by the Company of the Executive’s management responsibilities;
- (iii) a material breach of this Agreement by the Company;

provided however, that any resignation by the Executive due to any of the following conditions shall only be deemed for Good Reason if: (i) the Executive gives the Company written notice of the intent to terminate for Good Reason within 90 days following the first occurrence of the condition(s) that the Executive believes constitutes Good Reason, which notice shall describe such condition(s); (ii) the Company fails to remedy, if remediable, such condition(s) within 15 days following receipt of the written notice (the “**Cure Period**”) of such condition(s) from the Executive; and (iii) Executive actually resigns his employment within the first 15 days after expiration of the Cure Period.

**4.5.4 Ownership Change Event.** For purposes of this Agreement, “**Ownership Change Event**” means the occurrence of any of the following with respect to the Company: (i) the direct

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or indirect sale or exchange in a single or series of related transactions by the stockholders of the Company of more than 50% of the voting stock of the Company; (ii) a merger or consolidation in which the Company is a party; or (iii) the sale, exchange, or transfer of all or substantially all of the assets of the Company.

**5. ASSIGNMENT AND BINDING EFFECT.**

This Agreement shall be binding upon and inure to the benefit of the Executive and the Executive's heirs, executors, personal representatives, assigns, administrators and legal representatives. Because of the unique and personal nature of the Executive's duties under this Agreement, neither this Agreement nor any rights or obligations under this Agreement shall be assignable by the Executive. This Agreement shall be binding upon and inure to the benefit of the Company and its successors, assigns and legal representatives.

**6. CHOICE OF LAW.**

This Agreement is made in California. This Agreement shall be construed and interpreted in accordance with the internal laws of the State of Delaware.

**7. INTEGRATION.**

This Agreement, including **Exhibit A**, contains the complete, final and exclusive agreement of the Parties relating to the terms and conditions of the Executive's employment and the termination of Executive's employment, and supersedes all prior and contemporaneous oral and written employment agreements or arrangements between the Parties. To the extent this Agreement conflicts with the terms of the Company's employee handbook, governing policies, or bylaws, this Agreement controls.

**8. AMENDMENT.**

This Agreement cannot be amended or modified except by a written agreement signed by the Executive and the Company.

**9. WAIVER.**

No term, covenant or condition of this Agreement or any breach thereof shall be deemed waived, except with the written consent of the Party against whom the waiver is claimed, and any waiver or any such term, covenant, condition or breach shall not be deemed to be a waiver of any preceding or succeeding breach of the same or any other term, covenant, condition or breach.

**10. SEVERABILITY.**

The finding by a court of competent jurisdiction of the unenforceability, invalidity or illegality of any provision of this Agreement shall not render any other provision of this Agreement unenforceable, invalid or illegal. Such court shall have the authority to modify or replace the invalid or unenforceable term or provision with a valid and enforceable term or provision which most accurately represents the Parties' intention with respect to the invalid or unenforceable term or provision.

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## **11. INTERPRETATION; CONSTRUCTION.**

The headings set forth in this Agreement are for convenience of reference only and shall not be used in interpreting this Agreement. This Agreement has been drafted by legal counsel representing the Company, but the Executive has been encouraged to consult with, and has consulted with, the Executive's own independent counsel and tax advisors with respect to the terms of this Agreement. The Parties acknowledge that each Party and its counsel has reviewed and revised, or had an opportunity to review and revise, this Agreement, and the normal rule of construction to the effect that any ambiguities are to be resolved against the drafting party shall not be employed in the interpretation of this Agreement.

## **12. REPRESENTATIONS AND WARRANTIES.**

The Executive represents and warrants that the Executive is not restricted or prohibited, contractually or otherwise, from entering into and performing each of the terms and covenants contained in this Agreement, and that the Executive's execution and performance of this Agreement will not violate or breach any other agreements between the Executive and any other person or entity.

## **13. COUNTERPARTS.**

This Agreement may be executed in two counterparts, each of which shall be deemed an original, all of which together shall contribute one and the same instrument.

## **14. ARBITRATION.**

To ensure the rapid and economical resolution of disputes that may arise in connection with the Executive's employment with the Company, the Executive and the Company agree that any and all disputes, claims, or causes of action, in law or equity, arising from or relating to the Executive's employment, or the termination of that employment, will be resolved pursuant to the Federal Arbitration Act and to the fullest extent permitted by law, by final, binding and confidential arbitration conducted by the Judicial Arbitration and Mediation Services ("JAMS"), or its successors, under the then current rules of JAMS for employment disputes; provided that the arbitrator shall: (i) have the authority to compel adequate discovery for the resolution of the dispute and to award such relief as would otherwise be permitted by law; and (ii) issue a written arbitration decision including the arbitrator's essential findings and conclusions and a statement of the award. Both the Executive and the Company shall be entitled to all rights and remedies that either the Executive or the Company would be entitled to pursue in a court of law. The Company shall pay all fees, including the arbitrator's fee. Nothing in this Agreement is intended to prevent either the Executive or the Company from obtaining injunctive relief in court to prevent irreparable harm pending the conclusion of any such arbitration.

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**15. TRADE SECRETS OF OTHERS.**

It is the understanding of both the Company and the Executive that the Executive shall not divulge to the Company and/or its subsidiaries any confidential information or trade secrets belonging to others, including the Executive's former employers, nor shall the Company and/or its Affiliates seek to elicit from the Executive any such information. Consistent with the foregoing, the Executive shall not provide to the Company and/or its Affiliates, and the Company and/or its Affiliates shall not request, any documents or copies of documents containing such information.

**16. ADVERTISING WAIVER.**

The Executive agrees to permit the Company and/or its Affiliates, and persons or other organizations authorized by the Company and/or its Affiliates, to use, publish and distribute advertising or sales promotional literature concerning the products and/or services of the Company and/or its Affiliates, or the machinery and equipment used in the provision thereof, in which the Executive's name and/or pictures of the Executive taken in the course of the Executive's provision of services to the Company and/or its Affiliates, appear. The Executive hereby waives and releases any claim or right the Executive may otherwise have arising out of such use, publication or distribution during the term of this Agreement.

*[Signature Page Follows.]*

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IN WITNESS WHEREOF, the Parties have executed this Agreement as of the date first above written.

**A.P. PHARMA, Inc.**

/s/ Barry Quart

\_\_\_\_\_  
**Barry Quart**  
**Chief Executive Officer**

Dated: 30 Oct 2013

**EXECUTIVE:**

/s/ Paul Marshall

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**PAUL MARSHALL**

Dated: 29 Oct 2013

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**EXHIBIT A**  
**RELEASE AND WAIVER OF CLAIMS**  
**TO BE SIGNED AT TIME OF TERMINATION WITHOUT CAUSE OR**  
**RESIGNATION FOR GOOD REASON**

In consideration of the payments and other benefits set forth in Section 4.4 of the Executive Employment Agreement dated November 1, 2013, to which this form is attached, I, PAUL MARSHALL hereby furnish A.P. PHARMA, INC. (the "**Company**"), with the following release and waiver (the "**Release and Waiver**").

In exchange for the consideration provided to me by the Employment Agreement that I am not otherwise entitled to receive, I hereby generally and completely release the Company and its directors, officers, employees, stockholders, partners, agents, attorneys, predecessors, successors, parent and subsidiary entities, insurers, Affiliates, and assigns from any and all claims, liabilities and obligations, both known and unknown, that arise out of or are in any way related to events, acts, conduct, or omissions occurring prior to my signing this Release and Waiver. This general release includes, but is not limited to: (1) all claims arising out of or in any way related to my employment with the Company or the termination of that employment; (2) all claims related to my compensation or benefits from the Company, including, but not limited to, salary, bonuses, commissions, vacation pay, expense reimbursements, severance pay, fringe benefits, stock, stock options, or any other ownership interests in the Company; (3) all claims for breach of contract, wrongful termination, and breach of the implied covenant of good faith and fair dealing; (4) all tort claims, including, but not limited to, claims for fraud, defamation, emotional distress, and discharge in violation of public policy; and (5) all federal, state, and local statutory claims, including, but not limited to, claims for discrimination, harassment, retaliation, attorneys' fees, or other claims arising under the federal Civil Rights Act of 1964 (as amended), the federal Americans with Disabilities Act of 1990, the federal Age Discrimination in Employment Act of 1967 (as amended) ("**ADEA**"), and the California Fair Employment and Housing Act (as amended).

I also acknowledge that I have read and understand Section 1542 of the California Civil Code which reads as follows: "**A general release does not extend to claims which the creditor does not know or suspect to exist in his favor at the time of executing the release, which if known by him must have materially affected his settlement with the debtor.**" I hereby expressly waive and relinquish all rights and benefits under that section and any law of any jurisdiction of similar effect with respect to any claims I may have against the Company.

I acknowledge that, among other rights, I am waiving and releasing any rights I may have under ADEA, that this Release and Waiver is knowing and voluntary, and that the consideration given for this Release and Waiver is in addition to anything of value to which I was already entitled as an executive of the Company. If I am 40 years of age or older upon execution of this Release and Waiver, I further acknowledge that I have been advised, as required by the Older Workers Benefit Protection Act, that: (a) the release and waiver granted herein does not relate to claims under the ADEA which may arise after this Release and Waiver is executed; (b) I should consult with an attorney prior to executing this Release and Waiver; (c) I have twenty-one (21) days in which to consider this Release and Waiver (although I may

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choose voluntarily to execute this Release and Waiver earlier); (d) I have seven (7) days following the execution of this Release and Waiver to revoke my consent to this Release and Waiver; and (e) this Release and Waiver shall not be effective until the eighth day after I execute this Release and Waiver and the revocation period has expired (the "**Effective Date**").

I understand that among other things, I must not use or disclose any confidential or proprietary information of the Company and I must immediately return all Company property and documents (including all embodiments of proprietary information) and all copies thereof in my possession or control.

This Release and Waiver constitutes the complete, final and exclusive embodiment of the entire agreement between the Company and me with regard to the subject matter hereof. I am not relying on any promise or representation by the Company that is not expressly stated herein. This Release and Waiver may only be modified by a writing signed by both me and a duly authorized officer of the Company.

Date: \_\_\_\_\_

By: \_\_\_\_\_  
**PAUL MARSHALL**

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**Schedule A**

**CONSENT OF INDEPENDENT REGISTERED PUBLIC ACCOUNTING FIRM**

We consent to the incorporation by reference in the Registration Statement on Form S-3 (Nos. 333-162968, 333-167890, 333-183549 and 333-190550) and Form S-8 (Nos. 333-35151, 333-90428, 333-118546, 333-127574, 333-137954, 333-148660, 333-152862, 333-162610, 333-167515, 333-176365, 333-176366 and 333-190549) of our reports dated March 4, 2014 relating to the financial statements and effectiveness of internal control over financial reporting of Heron Therapeutics, Inc., included in the Annual Report on Form 10-K for the year ended December 31, 2013.

/s/ OUM & Co. LLP

San Francisco, California  
March 4, 2014

**CERTIFICATIONS**

I, Barry D. Quart, certify that:

1. I have reviewed this annual report on Form 10-K of Heron Therapeutics, Inc. (the "registrant") ;
2. Based on my knowledge, this report does not contain any untrue statement of a material fact or omit to state a material fact necessary to make the statements made, in light of the circumstances under which such statements were made, not misleading with respect to the period covered by this report;
3. Based on my knowledge, the financial statements, and other financial information included in this report, fairly present in all material respects the financial condition, results of operations and cash flows of the registrant as of, and for, the periods presented in this report;
4. The registrant's other certifying officer and I are responsible for establishing and maintaining disclosure controls and procedures (as defined in Exchange Act Rules 13a-15(e) and 15d-15(e)) and internal control over financial reporting (as defined in Exchange Act Rules 13a-15(f) and 15d-15(f)) for the registrant and have:
  - a) Designed such disclosure controls and procedures, or caused such disclosure controls and procedures to be designed under my supervision, to ensure that material information relating to the registrant, including its consolidated subsidiaries, is made known to us by others within those entities, particularly during the period in which this report is being prepared;
  - b) Designed such internal control over financial reporting, or caused such internal control over financial reporting to be designed under my supervision to provide reasonable assurance regarding the reliability of financial reporting and the preparation of financial statements for external purposes in accordance with generally accepted accounting principles;
  - c) Evaluated the effectiveness of the registrant's disclosure controls and procedures and presented in this report our conclusions about the effectiveness of the disclosure controls and procedures, as of the end of the period covered by this report based on such evaluation; and
  - d) Disclosed in this report any change in the registrant's internal control over financial reporting that occurred during the registrant's most recent fiscal quarter (the registrant's fourth fiscal quarter in the case of an annual report) that has materially affected, or is reasonably likely to materially affect, the registrant's internal control over financial reporting; and
5. The registrant's other certifying officer and I have disclosed, based on our most recent evaluation of internal control over financial reporting, to the registrant's auditors and the audit committee of registrant's board of directors (or persons performing the equivalent functions):
  - a) All significant deficiencies and material weaknesses in the design or operation of internal control over financial reporting which are reasonably likely to adversely affect the registrant's ability to record, process, summarize and report financial information; and
  - b) Any fraud, whether or not material, that involves management or other employees who have a significant role in the registrant's internal control over financial reporting.

Date: March 7, 2014

/s/ Barry D. Quart

Barry D. Quart  
Chief Executive Officer

**CERTIFICATIONS**

I, Brian G. Drazba, certify that:

1. I have reviewed this annual report on Form 10-K of Heron Therapeutics, Inc. (the "registrant") ;
2. Based on my knowledge, this report does not contain any untrue statement of a material fact or omit to state a material fact necessary to make the statements made, in light of the circumstances under which such statements were made, not misleading with respect to the period covered by this report;
3. Based on my knowledge, the financial statements, and other financial information included in this report, fairly present in all material respects the financial condition, results of operations and cash flows of the registrant as of, and for, the periods presented in this report;
4. The registrant's other certifying officer and I are responsible for establishing and maintaining disclosure controls and procedures (as defined in Exchange Act Rules 13a-15(e) and 15d-15(e)) and internal control over financial reporting (as defined in Exchange Act Rules 13a-15(f) and 15d-15(f)) for the registrant and have:
  - a) Designed such disclosure controls and procedures, or caused such disclosure controls and procedures to be designed under my supervision, to ensure that material information relating to the registrant, including its consolidated subsidiaries, is made known to us by others within those entities, particularly during the period in which this report is being prepared;
  - b) Designed such internal control over financial reporting, or caused such internal control over financial reporting to be designed under my supervision to provide reasonable assurance regarding the reliability of financial reporting and the preparation of financial statements for external purposes in accordance with generally accepted accounting principles;
  - c) Evaluated the effectiveness of the registrant's disclosure controls and procedures and presented in this report our conclusions about the effectiveness of the disclosure controls and procedures, as of the end of the period covered by this report based on such evaluation; and
  - d) Disclosed in this report any change in the registrant's internal control over financial reporting that occurred during the registrant's most recent fiscal quarter (the registrant's fourth fiscal quarter in the case of an annual report) that has materially affected, or is reasonably likely to materially affect, the registrant's internal control over financial reporting; and
5. The registrant's other certifying officer and I have disclosed, based on our most recent evaluation of internal control over financial reporting, to the registrant's auditors and the audit committee of registrant's board of directors (or persons performing the equivalent functions):
  - a) All significant deficiencies and material weaknesses in the design or operation of internal control over financial reporting which are reasonably likely to adversely affect the registrant's ability to record, process, summarize and report financial information; and
  - b) Any fraud, whether or not material, that involves management or other employees who have a significant role in the registrant's internal control over financial reporting.

Date: March 7, 2014

/s/ Brian G. Drazba

Brian G. Drazba  
Chief Financial Officer

**CERTIFICATION PURSUANT TO 18 U.S.C. SECTION 1350, AS ADOPTED  
PURSUANT TO SECTION 906 OF THE SARBANES-OXLEY ACT OF 2002**

In connection with the Annual Report of Heron Therapeutics, Inc. (the "Company") on Form 10-K for the year ending December 31, 2013 as filed with the Securities and Exchange Commission on the date hereof (the "Report"), I, Barry D. Quart, Chief Executive Officer of the Company, certify, pursuant to 18 U.S.C. Section 1350, as adopted pursuant to Section 906 of the Sarbanes-Oxley Act of 2002, that:

- (1) The Report fully complies with the requirements of section 13(a) or 15(d) of the Securities Exchange Act of 1934; and
- (2) The information contained in the Report fairly presents, in all material respects, the financial condition and result of operations of the Company.

/s/ Barry D. Quart

Barry D. Quart  
Chief Executive Officer  
March 7, 2014

**CERTIFICATION PURSUANT TO 18 U.S.C. SECTION 1350, AS ADOPTED  
PURSUANT TO SECTION 906 OF THE SARBANES-OXLEY ACT OF 2002**

In connection with the Annual Report of Heron Therapeutics, Inc. (the "Company") on Form 10-K for the year ending December 31, 2013 as filed with the Securities and Exchange Commission on the date hereof (the "Report"), I, Brian G. Drazba, Chief Financial Officer of the Company, certify, pursuant to 18 U.S.C. Section 1350, as adopted pursuant to Section 906 of the Sarbanes-Oxley Act of 2002, that:

- (1) The Report fully complies with the requirements of section 13(a) or 15(d) of the Securities Exchange Act of 1934; and
- (2) The information contained in the Report fairly presents, in all material respects, the financial condition and result of operations of the Company.

/s/ Brian G. Drazba

Brian G. Drazba  
Chief Financial Officer  
March 7, 2014

