

NanoString Technologies Inc  
Form S-1  
January 13, 2014  
**Table of Contents**

As filed with the Securities and Exchange Commission on January 13, 2014

Registration No. 333-

**UNITED STATES**  
**SECURITIES AND EXCHANGE COMMISSION**  
**Washington, D.C. 20549**

**FORM S-1**  
**REGISTRATION STATEMENT Under The Securities Act of 1933**

**NANOSTRING TECHNOLOGIES, INC.**

*(Exact name of registrant as specified in its charter)*

**Delaware**  
*(State or other jurisdiction of  
incorporation or organization)*

**3826**  
*(Primary Standard Industrial  
Classification Code Number)*  
**530 Fairview Avenue, N., Suite 2000**

**20-0094687**  
*(I.R.S. Employer  
Identification Number)*

**Seattle, Washington 98109**

**(206) 378-6266**

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

Edgar Filing: NanoString Technologies Inc - Form S-1

**R. Bradley Gray**

**President and Chief Executive Officer**

**530 Fairview Avenue, N., Suite 2000**

**Seattle, Washington 98109**

**(206) 378-6266**

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

**Copies to:**

**Patrick J. Schultheis**

**Kathryn Surace-Smith**

**Alan F. Denenberg**

**Wilson Sonsini Goodrich & Rosati,**

**Vice President, General Counsel**

**Davis Polk & Wardwell LLP**

**Professional Corporation**

**530 Fairview Avenue, N., Suite 2000**

**1600 El Camino Real**

**701 Fifth Avenue, Suite 5100**

**Seattle, Washington 98109**

**Menlo Park, California 94025**

**Seattle, Washington 98104**

**(206) 378-6266**

**(650) 752-2000**

**(206) 883-2500**

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

If any of the securities being registered on this Form are to be offered on a delayed or continuous basis pursuant to Rule 415 under the Securities Act, as amended, check the following box. "

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

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

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

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    
 (Do not check if a smaller reporting company)

Smaller reporting company "

**CALCULATION OF REGISTRATION FEE**

Title of each class of securities to be registered	Proposed maximum aggregate offering price(1)	Amount of registration fee
Common Stock, par value \$0.0001 per share	\$ 63,250,000	\$ 8,147

(1) Estimated solely for the purpose of calculating the amount of the registration fee pursuant to Rule 457(o) under the Securities Act of 1933, as amended. Includes the aggregate offering price of additional shares that the underwriters have the option to purchase to cover overallotments, if any.

**The registrant hereby amends this Registration Statement on such date or dates as may be necessary to delay its effective date until the registrant shall file a further amendment that specifically states that this Registration Statement shall thereafter become effective in accordance with Section 8(a) of the Securities Act of 1933, as amended, or until the Registration Statement shall become effective on such date as the Securities and Exchange Commission, acting pursuant to such Section 8(a), may determine.**

**Table of Contents**

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

*PROSPECTUS (Subject to Completion)*

Issued January 13, 2014

**\$55,000,000**

**COMMON STOCK**

NanoString Technologies, Inc. is offering shares of its common stock with an aggregate public offering price of approximately \$55,000,000.

Our common stock is listed on The NASDAQ Global Market under the symbol NSTG. The last reported sale price of our common stock on The NASDAQ Global Market on January 10, 2014 was \$19.51 per share.

We are an emerging growth company as defined under the federal securities laws and, as such, have elected to comply with certain reduced public company reporting requirements.

	Per Share	Total
Public offering price	\$	\$

Edgar Filing: NanoString Technologies Inc - Form S-1

Underwriting discounts and commissions	\$	\$
Proceeds to NanoString Technologies, Inc. before expenses <sup>(1)</sup>	\$	\$

(1) See Underwriting.

We expect to grant the underwriters an option to purchase additional shares of common stock to cover overallocments up to an aggregate public offering price of approximately \$8,250,000.

*Investing in our common stock involves a high degree of risk. See Risk Factors beginning on page 11.*

Neither the Securities and Exchange Commission nor any state securities commission has approved or disapproved of these securities or determined if this prospectus is truthful or complete. Any representation to the contrary is a criminal offense.

The underwriters expect to deliver the shares to purchasers on or about \_\_\_\_\_, 2014.

**J.P. Morgan**

Leerink Partners

, 2014

**Morgan Stanley**

Baird

**Table of Contents****TABLE OF CONTENTS**

<u>Prospectus Summary</u>	1
<u>Risk Factors</u>	11
<u>Special Note Regarding Forward-Looking Statements</u>	36
<u>Use of Proceeds</u>	37
<u>Market Price of Common Stock</u>	38
<u>Dividend Policy</u>	39
<u>Capitalization</u>	40
<u>Dilution</u>	41
<u>Selected Consolidated Financial Data</u>	43
<u>Management's Discussion and Analysis of Financial Condition and Results of Operations</u>	45
<u>Business</u>	67
<u>Management</u>	107
<u>Executive Compensation</u>	119
<u>Certain Relationships and Related Party Transactions</u>	132
<u>Principal Stockholders</u>	138
<u>Description of Capital Stock</u>	143
<u>Material U.S. Federal Tax Consequences to Non-U.S. Holders of Common Stock</u>	148
<u>Underwriting</u>	151
<u>Legal Matters</u>	158
<u>Experts</u>	158
<u>Where You Can Find Additional Information</u>	158
<u>Index to Consolidated Financial Statements</u>	F-1

We and the underwriters have not authorized anyone to provide any information other than that contained in this prospectus or in any free writing prospectus prepared by or on behalf of us or to which we have referred you. We take no responsibility for, and can provide no assurance as to the reliability of, any other information that others may give you. We and the underwriters are not making an offer to sell these securities in any jurisdiction where the offer or sale is not permitted. You should assume that the information appearing in this prospectus is accurate only as of the date on the front cover of this prospectus. Our business, financial condition, results of operations and prospects may have changed since that date.

For investors outside of the United States: We have not and the underwriters have not done anything that would permit this offering or possession or distribution of this prospectus in any jurisdiction where action for that purpose is required, other than the United States. Persons outside of the United States who come into possession of this prospectus must inform themselves about, and observe any restrictions relating to, the offering of the shares of common stock and the distribution of this prospectus outside of the United States.

**Table of Contents**

**PROSPECTUS SUMMARY**

*This summary highlights selected information contained elsewhere in this prospectus and is qualified in its entirety by the more detailed information and financial statements included elsewhere in this prospectus. It does not contain all of the information that may be important to you and your investment decision. You should carefully read this entire prospectus, including the matters set forth under Risk Factors, Management's Discussion and Analysis of Financial Condition and Results of Operations and our financial statements and related notes.*

**NanoString Technologies, Inc.**

**Overview**

We develop, manufacture and sell robust, intuitive products that unlock scientifically valuable and clinically actionable genomic information from minute amounts of tissue. Our nCounter Analysis System directly profiles hundreds of molecules simultaneously using a novel barcoding technology that is powerful enough for use in research, yet simple enough for use in clinical laboratories worldwide. We market systems and related consumables to researchers in academic, government, and biopharmaceutical laboratories for use in understanding fundamental biology and the molecular basis of disease and to clinical laboratories and medical centers for diagnostic use. We have an installed base of more than 180 systems, which our customers have used to publish more than 360 peer-reviewed papers. As researchers discover how genomic information can be used to improve clinical decision-making, these discoveries can be translated and validated as diagnostic tests based on our nCounter Elements reagents. In certain situations, we intend to translate their discoveries into *in vitro* diagnostic assays. In September 2013, we received 510(k) clearance from the U.S. Food and Drug Administration, or FDA, to market in the United States a version of our first molecular diagnostic product, the Prosigna Breast Cancer Assay, or Prosigna, an assay providing an assessment of a patient's risk of recurrence for breast cancer. In November 2013, we commercially launched the nCounter diagnostic system, or nCounter Dx Analysis System, in the United States, including a dual-mode configuration that can be used both for Prosigna and for research applications. In December 2013, we commercially launched Prosigna in the United States and announced that national diagnostic laboratories ARUP Laboratories, Laboratory Corporation of America Holdings and Quest Diagnostics have chosen to add Prosigna to their suites of breast cancer diagnostic tests. We expect Prosigna testing services to become available in the United States in the first quarter of 2014. In September 2012, we received European Union regulatory clearance for Prosigna providing an assessment of a patient's risk of recurrence for breast cancer and the intrinsic subtype of the patient's tumor. In February 2013, we commercially launched Prosigna in Europe and Israel.

The role of genomic information in research and medical practice is evolving rapidly. The advent of new technologies that sequence and digitally count discrete nucleic acids, commonly referred to as next generation sequencing, or NGS, is accelerating the discovery of the relationships between the genome and human disease. Researchers are applying this wealth of new information to identify biological pathways, which are networks of tens or hundreds of genes that act in concert to produce biological functions. Researchers then seek to translate this understanding of the genomic basis of disease into the development of diagnostic tools that can be used to profile an individual patient's biological pathways as well as develop targeted drug therapies. Precise, simple and robust profiling of biological pathways presents both an analytical challenge for researchers and an opportunity to improve patient outcomes in the future.

Our nCounter Analysis System enables genomic analysis on a scale appropriate for pathway-based biology by digitally quantifying the activity of up to 800 genes simultaneously in a single minute tissue sample. The sensitivity and precision of our novel barcoding chemistry allows the measurement of subtle changes in genomic activity efficiently, which is essential in both research and diagnostics because tissue samples are often available only in very small quantities. This problem is especially acute in cancer research, which is typically conducted using biopsies that are often stored in a format known as formalin-fixed paraffin embedded, or FFPE, which

## Table of Contents

complicates subsequent analysis of genetic material. The nCounter Analysis System is an easy-to-use and flexible solution that allows researchers to efficiently test hypotheses across thousands of different samples. As a result, the nCounter Analysis System is particularly useful for discovering and validating networks of genes that characterize and help predict disease states, enabling the development of diagnostics and medicines designed specifically for treating patients with certain genomic profiles. Researchers may use nCounter to develop their own diagnostic tests based on our nCounter Elements reagents or we may selectively partner with them to translate their discoveries into *in vitro* diagnostic assays.

Prosigna, our first molecular diagnostic test, is based on a collection of 50 genes known as the PAM50 gene signature, which was discovered by several of our life sciences customers. We secured an exclusive worldwide license to the PAM50 gene signature in 2010. Prosigna can provide a breast cancer patient and her physician with a subtype classification based on the fundamental biology of the patient's tumor, as well as a prognostic score that predicts the probability of cancer recurrence over 10 years. Our goal is for physicians to use Prosigna to guide therapeutic decisions so that patients receive only therapeutic interventions from which they are likely to benefit. In 2011, we conducted a clinical study, which we refer to as our TransATAC study, based on material extracted from tumor samples of more than 1,000 evaluable patients from the Arimidex, Tamoxifen, Alone or in Combination, or ATAC, study. In our study, investigators performed Prosigna on these samples that had been previously analyzed using Genomic Health's *Oncotype DX*, the historical market leader in breast cancer recurrence testing. The results of our TransATAC study demonstrated the ability of Prosigna to indicate risk of recurrence in postmenopausal women with hormone receptor-positive early stage breast cancer treated with endocrine therapy alone. In comparing the risk estimate provided by Prosigna to the risk estimate previously generated using *Oncotype DX*, investigators concluded that Prosigna is capable of providing more prognostic information than *Oncotype DX*. Based on the results of this study and multi-site analytical validation studies, we received European Union, or EU, regulatory clearance for Prosigna, known as a CE mark. As part of our preparation for regulatory submission in the United States, we conducted a second clinical study, which we refer to as our ABCSG8 study, based on tumor samples of more than 1,400 evaluable patients from the Austrian Breast & Colorectal Cancer Study Group 8. Our ABCSG8 study confirmed the conclusion that Prosigna can indicate risk of recurrence as previously demonstrated in our TransATAC study. In September 2013, we received 510(k) clearance from the FDA to market in the United States a version of Prosigna providing an assessment of a patient's risk of recurrence for breast cancer. In November 2013, we commercially launched the nCounter Dx Analysis System in the United States. In December 2013, we commercially launched Prosigna in the United States. National diagnostic laboratories ARUP Laboratories, Laboratory Corporation of America Holdings and Quest Diagnostics have chosen to add Prosigna to their suites of breast cancer diagnostic tests, and the laboratories at the University of Alabama at Birmingham Comprehensive Cancer Center and University of North Carolina Lineberger Comprehensive Cancer Center will be among the initial facilities to offer the Prosigna assay in the United States, with the earliest testing beginning during the first quarter of 2014. These laboratories collectively serve the pathology testing needs of a substantial portion of breast cancer patients throughout the United States. We expect additional clinical laboratories to adopt Prosigna in the future. We intend to conduct future clinical studies to evaluate Prosigna's ability to guide physicians and patients in making additional treatment decisions, which may include the selection of the appropriate chemotherapy regimen, the duration of adjuvant endocrine therapy, and whether to use adjuvant radiation therapy, and, if such studies are successfully completed, to seek 510(k) clearance or PMA approval in the United States for such indications in the future.

In November 2013, we began offering a version of the nCounter Dx Analysis System to high-complexity, CLIA-certified laboratories for research and diagnostics purposes. This FLEX configuration of the nCounter Dx Analysis System provides clinical laboratories a single platform with the flexibility to support both clinical testing, by running Prosigna, and research, by processing translational research experiments using our custom CodeSets and panels. The nCounter Elements General Purpose Reagents, or GPRs, provide further flexibility by allowing laboratories to develop their own Laboratory Developed Tests for gene expression, copy number variation and gene fusion signatures, which can be performed by a laboratory and may include genetic tests and other tests for rare conditions. GPRs are chemical reagents that have general laboratory application, and are not labeled or otherwise intended for a specific diagnostic application.



---

## **Table of Contents**

Prosigna is regulated as an *in vitro* diagnostic test and we distribute it as a kit for use on our nCounter Analysis System in clinical laboratories. We expect that our future *in vitro* diagnostic products will be regulated and distributed in a similar manner. This is in contrast to most complex genomic tests, which are currently regulated as services and are usually offered only by a limited number of specialized laboratories. The current centralized laboratory model for complex genomic testing can result in complicated logistics for the treating physician, including slower test result turnaround times and limited international access to tests as compared to local testing. In addition, most clinical laboratories cannot currently share in the revenue associated with offering patients complex genomic tests. We believe that our decentralized model will transform the current paradigm of complex genomic testing by allowing physicians worldwide to provide more comprehensive personalized diagnoses, broadening patient access, and increasing the degree to which clinical laboratories can profit by providing molecular diagnostic testing services.

We generated revenue of \$11.7 million, \$17.8 million and \$23.0 million in 2010, 2011 and 2012, respectively, and \$21.3 million in the nine months ended September 30, 2013, while incurring net losses of \$12.8 million, \$10.9 million and \$17.7 million in 2010, 2011 and 2012, respectively, and \$20.5 million in the nine months ended September 30, 2013.

## **Investment Highlights**

*Highly synergistic life sciences tools and molecular diagnostics business model.* Our nCounter Analysis System's key attributes appeal specifically to life sciences researchers focused on pathway-based biology who wish to translate and validate their discoveries into diagnostic tests. When these researchers identify clinically valuable genomic assays, they can pursue diagnostic applications through Laboratory Developed Tests enabled by our nCounter Elements reagents. In certain situations, we may in-license their discoveries and translate them into molecular *in vitro* diagnostic assays, such as Prosigna.

*Platform optimized for pathway-based biology.* Our system's ability to precisely measure subtle changes in the activity of hundreds of genes simultaneously within precious tissue samples is a significant advantage over traditional tools. While powerful enough for advanced research applications, our system's reliability and simplified workflow enables use in clinical laboratories worldwide. Innovations to improve the cost, performance, and footprint of our system will expand the range of customers that can benefit from using our platform in research and diagnostic applications.

*Recurring sales of proprietary consumables create a predictable revenue stream.* Because we are the exclusive provider of proprietary reagents for the nCounter Analysis System, the growth of our installed instrument base should drive an increasingly predictable stream of recurring consumable revenue. In 2010, 2011, 2012 and 2013, our average consumable revenue per installed instrument exceeded \$100,000 per year.

*Decentralized approach to complex genomic testing.* We believe that offering molecular diagnostics as Laboratory Developed Tests or *in vitro* diagnostic kits for use in local clinical laboratories will improve patient care by reducing turnaround times and allowing physicians worldwide, many of whom do not currently have access to these tests, to provide more comprehensive personalized diagnoses. In addition to broadening patient access, our decentralized business model will allow hospitals and pathology laboratories to profit by in-sourcing their molecular diagnostic testing services.

*Clinically validated assay targeting the significant and growing breast cancer diagnostics market.* In September 2012, we received an EU regulatory clearance for Prosigna, an assay providing an assessment of a patient's risk of recurrence for breast cancer and the intrinsic subtype of the patient's tumor, and in February 2013 we commercially launched Prosigna in Europe and Israel. In September 2013, we received 510(k) clearance from the FDA to market in the United States a version of Prosigna

## **Table of Contents**

providing an assessment of a patient's risk of recurrence for breast cancer. In November 2013, we commercially launched the nCounter Dx Analysis System in the United States, including a dual-mode configuration that can be used both for Prosigna and for research applications. In December 2013, we commercially launched Prosigna in the United States, and we expect Prosigna testing services to become available in the first quarter of 2014. Our TransATAC clinical study of material extracted from tumor samples from more than 1,000 evaluable patients that had been previously analyzed using Genomic Health's *Oncotype DX*, the historical market leader, provided evidence of clinical validity of Prosigna in predicting the risk of distant recurrence of breast cancer, and the investigators concluded that Prosigna is capable of providing more prognostic information than *Oncotype DX*. Our ABCSG8 clinical study based on tumor samples of more than 1,400 evaluable patients confirmed the conclusion that Prosigna can indicate risk of recurrence as previously demonstrated in our TransATAC study. We intend to pursue further clinical studies evaluating our test's ability to inform treatment decisions for which no genomic diagnostic tests are currently available.

*Capital efficient diagnostics business model.* We believe that our diagnostics business model is more capital efficient than the clinical laboratory services model and has the potential to become profitable on a small revenue base. Our diagnostics business leverages many of the capabilities of our life sciences business, including our technology platform and sales, product development, manufacturing, and administrative functions. Because we provide GPRs for Laboratory Developed Tests or *in vitro* diagnostics kits, rather than clinical laboratory services, we do not incur the costs of clinical laboratory infrastructure, sample logistics, or contracting with and billing managed care organizations. We believe that our customers will be motivated by the potential to improve patient care, broaden patient access and profit from testing services based on Prosigna and other potential nCounter-based diagnostics, which will encourage market adoption and potentially reduce sales and marketing expenditures relative to a centralized laboratory model.

## **Our Target Markets**

Over the last decade, methods of measuring genomic information have advanced substantially. However pathway-based research and the development of diagnostic tests require analysis of multiple genes and sensitivity to small changes in expression, which can be challenging for traditional genomic tools. In general, DNA microarrays and tube-based PCR methods require complex, time-consuming workflows and relatively large amounts of sample tissue to accurately characterize biological pathway activation. In both life sciences research and clinical medicine, there is a growing need for improved technologies that can precisely and rapidly measure the activation state of hundreds of genes simultaneously across a large number of precious samples, thereby providing a simple and reliable means to characterize biological pathways within minute tissue specimens.

## ***Life Sciences Research***

According to Strategic Directions International, Inc., life sciences researchers spent approximately \$28 billion on tools and related consumables in 2011. In the decade since the completion of the Human Genome Project, improvements in NGS technology have greatly reduced the cost of sequencing a human genome and increased throughput and precision, which has led to an abundance of new biological information. In order to gather insights from this information, researchers must first distill and then efficiently analyze large pools of data. Gene expression analysis has emerged as a primary tool that researchers use to extract meaningful insights from networks of genes, which enables them to validate and then translate their findings into the development of diagnostics and medicines. According to Percepta Associates, a provider of consulting services to bioscience companies, the 2012 global market for gene expression profiling products is estimated to be \$1.2 billion.

## **Table of Contents**

### ***Molecular Diagnostics***

According to Frost and Sullivan, the molecular diagnostics market totaled approximately \$4.1 billion in 2010 and is expected to reach \$6.2 billion by 2014. Growth in the molecular diagnostics market has been driven by technological innovations that have increased sensitivity, decreased turnaround times, simplified workflow, and lowered costs when compared to other techniques. In addition, the medical community has seen a trend in favor of decentralized diagnostic testing as a result of the convenience of local testing, hospitals and medical centers increasingly viewing their laboratories as profit centers and a need to increase access to tests for patients outside of the United States. We believe that there is an opportunity to improve the quality of diagnosis and treatment of diseases by developing and commercializing comprehensive, simple and widely available diagnostic products based on gene expression analysis. Molecular diagnostics have had a significant impact on the treatment of breast cancer, which had a worldwide incidence of 1.4 million per year in 2008 according to the World Health Organization. Over the last decade, genomic tests for breast cancer have improved the accuracy of prognosis and efficacy of treatment by assessing the risk of cancer recurrence for individual patients.

### **Our Solution**

Our nCounter Analysis System is an automated, multi-application, digital detection and counting system which directly profiles hundreds of molecules simultaneously using a novel barcoding technology that is powerful enough for use in research, yet simple enough for use in clinical laboratories worldwide. Our nCounter Analysis System consists of two automated instruments that prepare and analyze tissue samples using proprietary reagents, which can only be obtained from us. Our life sciences research customers purchase instruments from us and then purchase our panels, custom CodeSets, nCounter Elements reagents and related consumables for the specific experiment or assay they wish to conduct. Our diagnostics customers will either purchase or lease instruments from us and also generally purchase nCounter Elements reagents or our diagnostic kits for tests that they intend to run. Our nCounter Analysis System offers a number of compelling advantages, including:

*Optimized for Pathway-Based Biology.* The nCounter Analysis System can profile up to 800 molecules in a single test tube, which allows customers to analyze interactions among hundreds of genes that mediate biological pathways.

*Digital Precision.* Our molecular barcodes hybridize directly to the target molecules in a sample allowing them to be counted. This generates digital data (1 molecule = 1 count) of excellent quality over a wide dynamic range of measurements. Our nCounter Analysis System provides excellent reproducibility and avoids the potential bias that may be introduced by the sample division and extended amplification that are generally required for qPCR-based techniques.

*Simple Workflow.* The nCounter Analysis System's minimal sample preparation and automated workflow enable the performance of gene expression analysis across hundreds of genes simultaneously, with approximately 24 hours between the time a sample is loaded into the system and results are obtained. Our nCounter Analysis System generates data that customers can evaluate without the use of complex bioinformatics.

*Flexible Sample Requirements.* The nCounter Analysis System is able to unlock genomic information from minute amounts of a variety of challenging tissue samples, including FFPE samples, cell lysates, and single cells.

*Versatility.* The FLEX configuration of the nCounter Dx Analysis System provides clinical laboratories a single platform with the flexibility to support both clinical testing, by running Prosigna, and research, by processing translational research experiments and multiplexed assays using our custom CodeSets and panels. The nCounter Elements GPRs provide further flexibility by enabling laboratories to develop their own Laboratory Developed Tests for gene expression, copy number variation and gene fusion signatures.

## **Table of Contents**

### **Our Strategy**

Our goal is to provide products that empower scientists to understand the molecular basis of disease and empower physicians to put genomic medicine into practice. To accomplish this goal, we intend to continue providing technologies that are powerful enough for research, yet simple and robust enough for use in clinical laboratories worldwide.

Our strategy includes the following key elements:

Establish the nCounter Analysis System as the global standard for gene expression analysis.

Expand the installed base of the nCounter Analysis System in biopharmaceutical and academic research.

Broaden the addressable market of the nCounter Analysis System through continued innovation.

Build a menu of diagnostic content in collaboration with researchers comprising both proprietary *in vitro* diagnostic kits and Laboratory Developed Tests based on nCounter Elements reagents.

Execute high quality clinical studies to support regulatory authorizations, market adoption and reimbursement of diagnostic products.

Enable clinical laboratories worldwide to provide complex genomic testing using our *in vitro* diagnostic products.

Drive physician demand for nCounter Analysis System-based diagnostic products.

Capture capital efficiencies stemming from our diagnostics business model.

### **Risks Associated With Our Business**

Our business is subject to numerous risks, including:

We have incurred losses since we were formed and expect to incur losses in the future. We cannot be certain that we will achieve or sustain profitability.

Our financial results may vary significantly from quarter to quarter which may adversely affect our stock price.

Commercialization of Prosigna in Europe and the United States and development and commercialization of other diagnostic products worldwide are key elements of our strategy. If we fail to successfully commercialize Prosigna or such other products, we may never receive a return on the significant investments in sales and marketing, regulatory, manufacturing and quality assurance personnel we have made, and further investments we intend to make, which would adversely affect our growth prospects, operating results and financial condition.

## Edgar Filing: NanoString Technologies Inc - Form S-1

Further approval and/or clearance by the FDA and by foreign regulatory authorities for our diagnostic tests will take significant time and require significant research, development and clinical study expenditures, may result in a clearance that does not allow us to differentiate our diagnostic tests, including Prosigna, from alternatives and ultimately may not succeed.

The life sciences research and diagnostics markets are highly competitive. If we fail to compete effectively, our business and operating results will suffer.

If Medicare and other third-party payors in the United States and foreign countries do not approve reimbursement for diagnostic tests enabled by our technology, the commercial success of our diagnostic products would be compromised.

---

## **Table of Contents**

If we are unable to protect our intellectual property effectively, our business would be harmed. For additional information about the risks we face, please see the section of this prospectus captioned Risk Factors.

### **Recent Developments**

We anticipate that our fourth quarter 2013 total revenue will be approximately \$10.1 million and full year 2013 total revenue will be approximately \$31.4 million, representing approximately 55% and 37% year-over-year growth, respectively.

We anticipate instrument revenue of approximately \$5.3 million for the fourth quarter of 2013, representing approximately 112% growth compared to \$2.5 million for the fourth quarter of 2012. Our installed base of nCounter Analysis Systems stood at more than 180 systems as of the end of 2013. Consumables revenue is anticipated to be approximately \$4.4 million in the fourth quarter of 2013, representing approximately 19% growth over the \$3.7 million reported for the fourth quarter of 2012.

We are in the process of finalizing our financial results for 2013, and therefore our finalized and audited results are not yet available. The preliminary expectations regarding 2013 instrument, consumable and total revenue are the responsibility of management, are subject to management's review and actual results could differ from management's expectations. The actual results are also subject to audit by our independent registered public accounting firm and no assurance is given by our independent registered public accounting firm on such preliminary expectations. In addition, although we expect to experience an operating and net loss for 2013, we are not able to provide an estimate of such results at the time of this prospectus. However, we expect our operating loss and net loss for the fourth quarter of 2013 to increase compared to the fourth quarter of 2012. Accordingly, no conclusions should be drawn as to the size of our loss based on the foregoing revenue estimates. Our expectations regarding 2013 revenue are not necessarily indicative of results expected in future periods.

In January 2014, we entered into a non-binding letter of intent for a credit agreement with a lender which would allow us to refinance our existing credit facility and potentially incur up to an aggregate of \$45 million in term loan borrowings or up to an aggregate of approximately \$52 million if we elect to exercise in full an option to pay in kind a portion of the interest that would accrue on the borrowings under the credit agreement. We expect this credit agreement to contain customary conditions to borrowings, events of default and negative covenants, including covenants that could limit our ability to, among other things, incur additional indebtedness, liens or other encumbrances, make dividends or other distributions, and sell or transfer assets. We also expect that the credit agreement will include certain customary financial covenants. Our obligations under the credit agreement would be secured by substantially all of our assets. However, there can be no assurance that we will successfully enter into this credit agreement.

### **Corporate History and Information**

We were incorporated in Delaware in June 2003. Our principal executive offices are located at 530 Fairview Avenue, N., Suite 2000, Seattle, Washington 98109. Our telephone number is (206) 378-6266. Our website address is [www.nanostring.com](http://www.nanostring.com). Information contained on the website is not incorporated by reference into this prospectus, and should not be considered to be part of this prospectus.

Unless the context indicates otherwise, as used in this prospectus, the terms NanoString, we, us and our refer to NanoString Technologies, Inc. and its subsidiaries, NanoString Technologies Europe Limited, NanoString Technologies SAS, NanoString Technologies Asia Pacific Limited, NanoString Technologies International, Inc., NanoString Technologies GmbH and NanoString Technologies Singapore Pte Limited. We use NanoString®, NanoString Technologies®, nCounter®, Molecules that Count®, Prosig™, nCounter Elements™ and other marks as trademarks in the United States and other countries. This prospectus contains references to our trademarks as well as third-party trademarks. Solely for convenience, trademarks and trade names referred to in this prospectus, including logos, artwork and other visual displays, may appear without the ® or ™ symbols, but such references are not intended to indicate in any way that we will not assert, to the fullest extent under applicable law, our rights or the rights of the applicable licensor to these trademarks and trade names. We do not intend our use of third-party trade names or trademarks to imply a relationship with, or endorsement or sponsorship of us by, any other entity.

**Table of Contents**

**The Offering**

Common stock offered by us	shares	
Common stock to be outstanding after this offering	shares (or full)	if the underwriters exercise their overallotment option in
Overallotment option	shares	

Use of proceeds

We intend to use the net proceeds from this offering: (1) to further commercialize Prosigna, including establishing a dedicated oncology sales force; (2) to expand the clinical utility of Prosigna and develop other potential diagnostic product opportunities; (3) to expand life sciences commercial operations to grow and support the installed base of our nCounter Analysis Systems among life sciences research customers in the United States and internationally; (4) to develop new life sciences applications, chemistry and instrumentation for our nCounter technology platform; and (5) for working capital and other general corporate purposes. We may also use a portion of the net proceeds to acquire, license and invest in complementary products, technologies or businesses; however, we currently have no agreements or commitments to complete any such transaction. See Use of Proceeds.

NASDAQ trading symbol NSTG

The number of shares of common stock to be outstanding following this offering is based on 14,616,871 shares of common stock outstanding as of September 30, 2013, and excludes:

1,873,491 shares of common stock issuable upon exercise of options outstanding as of September 30, 2013, at a weighted-average exercise price of \$3.41 per share;

1,891,069 shares of common stock reserved for future issuance under stock-based compensation plans, including 1,609,819 shares of common stock reserved for issuance under our 2013 Equity Incentive Plan, and any future automatic increase in shares reserved for issuance under such plan, and 281,250 shares of common stock reserved for issuance under our 2013 Employee Stock Purchase Plan, and any future automatic increase in shares reserved for issuance under such plan; and

617,605 shares of common stock issuable upon the exercise of warrants outstanding as of September 30, 2013, at a weighted-average exercise price of \$8.78 per share.

Unless otherwise indicated, this prospectus reflects and assumes the following:

no exercise of outstanding options to purchase common stock or warrants to purchase common stock after September 30, 2013; and

no exercise by the underwriters of their overallotment option.





**Table of Contents****Summary Consolidated Financial Data**

We have derived the following summary of statements of operations data for the years ended December 31, 2010, 2011 and 2012 from audited consolidated financial statements appearing elsewhere in this prospectus. We derived the following statements of operations for the nine months ended September 30, 2012 and 2013 and the balance sheet data as of September 30, 2013 from unaudited interim financial statements included elsewhere in this prospectus. In the opinion of management, the unaudited interim financial statements reflect all adjustments, which include normal recurring adjustments, necessary for a fair presentation of the financial statements. Historical results are not necessarily indicative of the results that may be expected in the future and the results for the nine months ended September 30, 2013 are not necessarily indicative of the results that may be expected for the full year or any other period. The summary consolidated financial data set forth below should be read together with the financial statements and the related notes to those statements, as well as the sections of this prospectus captioned "Selected Consolidated Financial Data" and "Management's Discussion and Analysis of Financial Condition and Results of Operations."

	Year Ended December 31,			Nine Months Ended	
	2010	2011	2012	2012	2013
	(In thousands, except per share data)				
<b>Consolidated Statements of Operations:</b>					
<b>Revenue</b>	\$ 11,730	\$ 17,800	\$ 22,973	\$ 16,480	\$ 21,283
<b>Costs and expenses:</b>					
Cost of revenue	9,128	9,777	12,361	9,076	10,188
Research and development	7,547	8,990	11,635	8,253	10,469
Selling, general and administrative	8,027	9,529	15,486	10,588	20,822
Total costs and expenses	24,702	28,296	39,482	27,917	41,479
Loss from operations	(12,972)	(10,496)	(16,509)	(11,437)	(20,196)
<b>Other income (expense):</b>					
Interest income	29	10	21	17	28
Interest expense	(94)	(599)	(804)	(551)	(1,412)
Other income (expense)	254	80	(29)	(26)	(30)
Revaluation of preferred stock warrant liability	15	73	(387)	150	1,156
Total other income (expense)	204	(436)	(1,199)	(410)	(258)
Net loss	(12,768)	(10,932)	(17,708)	(11,847)	(20,454)
Accretion of mandatorily redeemable convertible preferred stock	(4,351)	(5,251)	(7,533)	(5,515)	(4,653)
Net loss attributable to common stockholders	\$ (17,119)	\$ (16,183)	\$ (25,241)	\$ (17,362)	\$ (25,107)
Net loss per share - basic and diluted	\$ (54.17)	\$ (50.10)	\$ (71.10)	\$ (51.06)	\$ (4.74)
Weighted-average shares used in computing basic and diluted net loss per share	316	323	355	340	5,292

**Table of Contents**

	As of September 30, 2013	
	Actual	As Adjusted <sup>(1)</sup>
	(In thousands)	
<b>Consolidated Balance Sheet Data:</b>		
Cash, cash equivalents and short-term investments	\$ 52,214	\$
Working capital	52,054	
Total assets	71,293	
Total long-term debt	18,213	
Total stockholders' equity	39,891	

- (1) Gives effect to the sale and issuance by us of \_\_\_\_\_ shares of common stock in this offering based upon an assumed public offering price of \$ \_\_\_\_\_ per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on January \_\_\_\_\_, 2014, after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us.

---

**Table of Contents**

**RISK FACTORS**

*Investing in our common stock involves a high degree of risk. Before making an investment decision, you should carefully consider the risks and uncertainties described below, which we believe are the material risks associated with our business and this offering. Our business, financial condition, operating results or growth prospects could be harmed by any of these risks. In that event, the trading price of our common stock could decline due to any of these risks, and you may lose all or part of your investment. In assessing these risks, you should also refer to all of the other information contained in this prospectus, including our financial statements and related notes.*

**Risks Related to our Business and Strategy**

*We have incurred losses since we were formed and expect to incur losses in the future. We cannot be certain that we will achieve or sustain profitability.*

We have incurred losses since we were formed and expect to incur losses in the future. We incurred net losses of \$20.5 million and \$11.8 million for the nine months ended September 30, 2013 and 2012, respectively. As of September 30, 2013, we had an accumulated deficit of \$118.0 million. We expect that our losses will continue for at least the next several years as we will be required to invest significant additional funds toward development and commercialization of our technology. We also expect that our selling, general and administrative expenses will continue to increase due to the additional costs associated with establishing a dedicated oncology diagnostics sales force and the increased administrative costs associated with being a public company. Our ability to achieve or sustain profitability is based on numerous factors, many of which are beyond our control, including the market acceptance of our products, future product development and our market penetration and margins. We may never be able to generate sufficient revenue to achieve or sustain profitability.

*Our financial results may vary significantly from quarter to quarter which may adversely affect our stock price.*

Investors should consider our business and prospects in light of the risks and difficulties we expect to encounter in the new, uncertain and rapidly evolving markets in which we compete. Because these markets are new and evolving, predicting their future growth and size is difficult. We expect that our visibility into future sales of our products, including volumes, prices and product mix between instruments and consumables, will continue to be limited and could result in unexpected fluctuations in our quarterly and annual operating results.

Numerous other factors, many of which are outside our control, may cause or contribute to significant fluctuations in our quarterly and annual operating results. These fluctuations may make financial planning and forecasting difficult. In addition, these fluctuations may result in unanticipated decreases in our available cash, which could negatively affect our business and prospects. Factors that may contribute to fluctuations in our operating results include many of the risks described in this section. In addition, one or more of such factors may cause our revenue or operating expenses in one period to be disproportionately higher or lower relative to the others. Our products involve a significant capital commitment by our customers and accordingly involve a lengthy sales cycle. We may expend significant effort in attempting to make a particular sale, which may be deferred by the customer or never occur. Accordingly, comparing our operating results on a period-to-period basis may not be meaningful, and investors should not rely on our past results as an indication of our future performance. If such fluctuations occur or if our operating results deviate from our expectations or the expectations of securities analysts, our stock price may be adversely affected.

*If we do not achieve, sustain or successfully manage our anticipated growth, our business and growth prospects will be harmed.*

We have experienced significant revenue growth in a short period of time. We may not achieve similar growth rates in future periods. Investors should not rely on our operating results for any prior periods as an indication of our future operating performance. If we are unable to maintain adequate revenue growth, our

## **Table of Contents**

financial results could suffer and our stock price could decline. Furthermore, growth will place significant strains on our management and our operational and financial systems and processes. For example, commercialization of the Prosigna Breast Cancer Assay, or Prosigna, in Europe and the United States and development and commercialization of this test and other diagnostic products worldwide are key elements of our growth strategy and will require us to hire and retain additional sales and marketing, regulatory, manufacturing and quality assurance personnel. If we do not successfully forecast the timing of regulatory clearance or approval for product marketing in additional jurisdictions and subsequent demand for our diagnostic products or manage our anticipated expenses accordingly, our operating results will be harmed.

***If Prosigna fails to achieve and sustain sufficient market acceptance, we will not generate expected revenue, and our prospects may be harmed.***

Commercialization of Prosigna in Europe, the United States and the other jurisdictions in which we intend to pursue regulatory approval is a key element of our strategy. Currently, most oncologists seeking sophisticated gene expression analysis for diagnosing and profiling breast cancer in their patients ship tissue samples to a limited number of centralized laboratories typically located in the United States. We may experience reluctance, or refusal, on the part of physicians to order, and third-party payors to pay for, Prosigna if the results of our research and clinical studies, and our sales and marketing activities relating to communication of these results, do not convey to physicians, third-party payors and patients that Prosigna provides equivalent or better prognostic information. In addition, breast cancer treatment guidelines recommend that chemotherapy be considered in many cases, in combination with other patient factors. Accordingly, physicians may be reluctant to order a test, such as Prosigna, that may suggest recommending against chemotherapy. Furthermore, our diagnostic tests would be performed by pathologists in local laboratories, rather than by a vendor in a remote centralized laboratory, which requires us to educate pathologists regarding the benefits of this business model and oncologists regarding the reliability and consistency of results generated locally.

These hurdles may make it difficult to convince health care providers that tests using our technologies are appropriate options for cancer diagnostics, may be equivalent or superior to available tests, and may be at least as cost effective as alternative technologies. Furthermore, we may encounter significant difficulty in gaining inclusion in breast cancer treatment guidelines, obtaining patient reimbursement from public and private payors, and gaining broad market acceptance of Prosigna. If we fail to successfully commercialize Prosigna, we may never receive a return on the significant investments in sales and marketing, regulatory, manufacturing and quality assurance personnel we have made, and further investments we intend to make, which would adversely affect our growth prospects, operating results and financial condition.

***Our future success is dependent upon our ability to expand our customer base and introduce new applications.***

Our current customer base is primarily composed of academic institutions, government laboratories and biopharmaceutical companies that perform analyses using our nCounter Analysis System for research use only. Our success will depend, in part, upon our ability to increase our market penetration among these customers and to expand our market by developing and marketing new life sciences applications, developing a lower cost instrument that would be attractive to more researchers, and introducing diagnostic products into clinical laboratories after obtaining regulatory authorization. For example, we must convince physicians and third-party payors that our diagnostic products, such as Prosigna, are cost effective in obtaining prognostic information that can inform treatment decisions and that our nCounter Analysis System could enable an equivalent or superior approach that lessens reliance on centralized laboratories. Furthermore, we expect that increasing the installed base of our nCounter Analysis Systems will drive demand for our relatively high margin consumable products. If we are not able to successfully increase our installed base of nCounter Analysis Systems, sales of our consumable products and our margins may not meet expectations. Attracting new customers and introducing new applications requires substantial time and expense. Any failure to expand our existing customer base, or launch new applications, would adversely affect our ability to improve our operating results.

## **Table of Contents**

***Our life sciences research business depends on levels of research and development spending by academic and governmental research institutions and biopharmaceutical companies, a reduction in which could limit demand for our products and adversely affect our business and operating results.***

In the near term, we expect that our revenue will be derived primarily from sales of our nCounter Analysis Systems to academic institutions, governmental laboratories and biopharmaceutical companies worldwide for research applications. The demand for our products will depend in part upon the research and development budgets of these customers, which are impacted by factors beyond our control, such as:

changes in government programs that provide funding to research institutions and companies;

macroeconomic conditions and the political climate;

changes in the regulatory environment;

differences in budgetary cycles;

market-driven pressures to consolidate operations and reduce costs; and

market acceptance of relatively new technologies, such as ours.

For example, in the United States, automatic across-the-board cuts in government spending, or sequestration, took effect on March 1, 2013. These cuts impacted the budgets of government agencies, such as the National Institutes of Health, which provide significant funding for cancer research and other diseases, however, as of the date of this prospectus the full impact of the cuts is unknown. We believe that the uncertainty regarding the availability of research funding, including the impact of sequestration, has adversely affected our historical operating results and any continuing uncertainty may adversely affect sales to customers or potential customers that rely on government funding. In addition, academic, governmental and other research institutions that fund research and development activities may be subject to stringent budgetary constraints that could result in spending reductions, reduced allocations or budget cutbacks, which could jeopardize the ability of these customers to purchase our products.

Our operating results may fluctuate substantially due to reductions and delays in research and development expenditures by these customers. Any decrease in our customers' budgets or expenditures, or in the size, scope or frequency of capital or operating expenditures, could materially and adversely affect our business, operating results and financial condition.

***Our sales cycle is lengthy and variable, which makes it difficult for us to forecast revenue and other operating results.***

Our sales process involves numerous interactions with multiple individuals within an organization, and often includes in-depth analysis by potential customers of our products, performance of proof-of-principle studies, preparation of extensive documentation and a lengthy review process. As a result of these factors, the large capital investment required in purchasing our instruments and the budget cycles of our customers, the time from initial contact with a customer to our receipt of a purchase order can vary significantly and be up to 12 months or longer. Given the length and uncertainty of our sales cycle, we have in the past experienced, and likely will in the future experience, fluctuations in our instrument sales on a period-to-period basis. In addition, any failure to meet customer expectations could result in customers choosing to retain their existing systems or to purchase systems other than ours.

***Our reliance on distributors for sales of our products outside of the United States could limit or prevent us from selling our products in foreign markets and impact our revenue.***

We have established exclusive distribution agreements for our nCounter Analysis System and related consumable products within parts of Europe, the Middle East, Asia Pacific and South America. We intend to continue to grow our business internationally, and to do so we must

attract additional distributors and retain

## **Table of Contents**

existing distributors to maximize the commercial opportunity for our products. There is no guarantee that we will be successful in attracting or retaining desirable sales and distribution partners or that we will be able to enter into such arrangements on favorable terms. Distributors may not commit the necessary resources to market and sell our products to the level of our expectations or may choose to favor marketing the products of our competitors. If current or future distributors do not perform adequately, or we are unable to enter into effective arrangements with distributors in particular geographic areas, we may not realize long-term international revenue growth.

***If we do not obtain additional regulatory clearances or approvals to market products other than Prosigna for diagnostic purposes, we will be limited to marketing such products for research use only. In addition, if we are unable to obtain additional regulatory clearances or approvals to market Prosigna in additional countries or if regulatory limitations are placed on our diagnostic products our business and growth will be harmed.***

We have received regulatory clearance in the United States under a 510(k) for a version of our first diagnostic product, Prosigna, providing an assessment of a patient's risk of recurrence for breast cancer, and we have obtained a CE mark for Prosigna which permits us to market that assay for diagnostic purposes in Europe. We do not have regulatory clearance or approval to market any other product for diagnostic purposes or to market Prosigna for diagnostic purposes in any other market, other than Israel. Other than with respect to Prosigna in such jurisdictions, we are limited to marketing our products for research use only, which means that we cannot make any diagnostic or clinical claims. We intend to seek regulatory authorizations in other jurisdictions to market Prosigna for diagnostic purposes; however, we cannot assure investors that we will be successful in doing so. Similarly, if we do not obtain additional regulatory clearances or approvals to market future products or future indications for diagnostic purposes, if unexpected regulatory limitations are placed on our products or if we fail to successfully commercialize such products, the market potential for our diagnostic products would be constrained, and our business and growth prospects would be adversely affected.

***As part of our current business model, we will seek to enter into strategic collaborations and licensing arrangements with third parties to develop diagnostic tests.***

We have relied, and expect to continue to rely, on strategic collaborations and licensing agreements with third parties for discoveries based on which we develop diagnostic tests. For example, we licensed the rights to intellectual property that forms the basis of Prosigna from Bioclassifier, LLC, which was founded by several of our life sciences research customers engaged in translational research. In addition, in February 2013, we secured an option from The Broad Institute, a leading non-profit molecular medicine institute in Cambridge, Massachusetts, to acquire an exclusive worldwide license for a gene signature that could be used, after further development, as a Laboratory Developed Test, or, after appropriate regulatory authorization, for a second molecular diagnostic product focused on hepatocellular carcinoma, or HCC. We intend to enter into more such arrangements with our life sciences customers and other researchers for future diagnostic products. However, there is no assurance that we will be successful in doing so. In particular, our life sciences research customers are not obligated to collaborate with us or license technology to us, and they may choose to develop diagnostic products themselves or collaborate with our competitors. Establishing collaborations and licensing arrangements is difficult and time-consuming. Discussions may not lead to collaborations or licenses on favorable terms, if at all. To the extent we agree to work exclusively with a party in a given area, our opportunities to collaborate with others could be limited. Potential collaborators or licensors may elect not to work with us based upon their assessment of our financial, regulatory or intellectual property position. Even if we establish new relationships, they may never result in the successful development or commercialization of future tests.

***New diagnostic product development involves a lengthy and complex process, and we may be unable to commercialize on a timely basis, or at all, any of the tests we develop.***

Few research and development projects result in successful commercial products, and success in early clinical studies often is not replicated in later studies. For example, even though the results of our clinical studies that used samples from the Arimidex, Tamoxifen, Alone or in Combination, or ATAC, study and the Austrian

---

## **Table of Contents**

Breast & Colorectal Cancer Study Group 8, or ABCSG8, study of postmenopausal women with HR+ early stage breast cancer were favorable, there is no guarantee that any future studies will be successful. At any point, we may abandon development of a product candidate or we may be required to expend considerable resources repeating clinical studies, which would adversely impact potential revenue and our expenses. In addition, any delay in product development would provide others with additional time to commercialize competing products before we do, which in turn may adversely affect our growth prospects and operating results.

***Our research and development efforts will be hindered if we are not able to contract with third parties for access to archival tissue samples.***

Under standard clinical practice, tumor biopsies removed from patients are preserved and stored in formalin-fixed paraffin embedded, or FFPE, format. We rely on our ability to secure access to these archived FFPE tumor biopsy samples, as well as information pertaining to the clinical outcomes of the patients from which they were derived for our clinical development activities. Others compete with us for access to these samples. Additionally, the process of negotiating access to archived samples is lengthy because it typically involves numerous parties and approval levels to resolve complex issues such as usage rights, institutional review board approval, privacy rights, publication rights, intellectual property ownership and research parameters. If we are not able to negotiate access to archived tumor tissue samples with hospitals, clinical partners, pharmaceutical companies, or companies developing therapeutics on a timely basis, or at all, or if other laboratories or our competitors secure access to these samples before us, our ability to research, develop and commercialize future products will be limited or delayed.

***The life sciences research and diagnostic markets are highly competitive. If we fail to compete effectively, our business and operating results will suffer.***

We face significant competition in the life sciences research and diagnostics markets. We currently compete with both established and early stage life sciences research companies that design, manufacture and market instruments and consumables for gene expression analysis, single-cell analysis, polymerase chain reaction, or PCR, digital PCR, other nucleic acid detection and additional applications. These companies use well established laboratory techniques such as microarrays or quantitative PCR, or qPCR, as well as newer technologies such as next generation sequencing. We believe our principal competitors in the life sciences research market are Affymetrix, Agilent Technologies, Bio-Rad, Exiqon, Fluidigm, High Throughput Genomics, Illumina, Life Technologies, Luminex, Perkin Elmer, Qiagen and Roche Applied Science. In addition, there are a number of new market entrants in the process of developing novel technologies for the life sciences market, including companies such as RainDance Technologies and Wafergen Bio-Systems.

We also compete with commercial diagnostics companies. We believe our principal competitor in the breast cancer diagnostics market is Genomic Health, which provides gene expression analysis at its central laboratory in Redwood City, California and currently commands a substantial majority of the market. We also face competition from companies such as Agendia, Clariant (a GE Healthcare company), Genoptix (a division of Novartis) and bioMeri  ux, which also offer services by means of centralized laboratories that profile gene or protein expression in breast cancer. In Europe, we also face regional competition from smaller companies such as Sividon Diagnostics, maker of EndoPredict, a distributed test for breast cancer recurrence, and other independent laboratories.

Most of our current competitors are either publicly traded, or are divisions of publicly-traded companies, and enjoy a number of competitive advantages over us, including:

greater name and brand recognition, financial and human resources;

broader product lines;

larger sales forces and more established distributor networks;

substantial intellectual property portfolios;



**Table of Contents**

larger and more established customer bases and relationships; and

better established, larger scale, and lower cost manufacturing capabilities.

We believe that the principal competitive factors in all of our target markets include:

cost of capital equipment;

cost of consumables and supplies;

reputation among customers;

innovation in product offerings;

flexibility and ease-of-use;

accuracy and reproducibility of results; and

compatibility with existing laboratory processes, tools and methods.

We believe that additional competitive factors specific to the diagnostics market include:

breadth of clinical decisions that can be influenced by information generated by tests;

volume, quality, and strength of clinical and analytical validation data;

availability of reimbursement for testing services; and

economic benefit accrued to customers based on testing services enabled by products.

We cannot assure investors that our products will compete favorably or that we will be successful in the face of increasing competition from new products and technologies introduced by our existing competitors or new companies entering our markets. In addition, we cannot assure investors that our competitors do not have or will not develop products or technologies that currently or in the future will enable them to produce competitive products with greater capabilities or at lower costs than ours. Any failure to compete effectively could materially and adversely affect our business, financial condition and operating results.

***We have limited experience in marketing and selling our products, and if we are unable to successfully commercialize our products, our business may be adversely affected.***

We have limited experience marketing and selling our products. Our nCounter Analysis System was introduced for sale in the life sciences research market in 2008, and was introduced for sale in the diagnostics market in Europe and Israel in February 2013, and in the United States in

## Edgar Filing: NanoString Technologies Inc - Form S-1

November 2013. We sell our products through our own sales force in North America and through a combination of our own sales force and distributors in Europe, Middle East, Asia Pacific and South America. In the future, we intend to establish distributor relationships in other parts of the world; however, we may not be able to market and sell our products effectively.

Our future sales of diagnostic products, including Prosigna, will depend in large part on our ability to successfully establish an oncology diagnostics sales force and to increase the scope of our marketing efforts. Because we have limited experience in marketing and selling our products in the diagnostics market, our ability to forecast demand, the infrastructure required to support such demand and the sales cycle to diagnostics customers is unproven. If we do not build an efficient and effective sales force targeting this market, our business and operating results will be adversely affected.

***We may not be able to develop new products or enhance the capabilities of our systems to keep pace with rapidly changing technology and customer requirements, which could have a material adverse effect on our business and operating results.***

Our success depends on our ability to develop new products and applications for our technology in existing and new markets, while improving the performance and cost-effectiveness of our systems. New technologies,

## **Table of Contents**

techniques or products could emerge that might offer better combinations of price and performance than our current or future products and systems. Existing markets for our products, including gene expression analysis, single-cell analysis and copy number variation, as well as potential markets for our diagnostic product candidates, are characterized by rapid technological change and innovation. It is critical to our success that we anticipate changes in technology and customer requirements and to successfully introduce new, enhanced and competitive technologies to meet our customers' and prospective customers' needs on a timely and cost-effective basis. At the same time, however, we must carefully manage the introduction by us of new products. If customers believe that such products will offer enhanced features or be sold for a more attractive price, they may delay purchases until such products are available. We may also have excess or obsolete inventory of older products as we transition to new products and our experience in managing product transitions is very limited. If we do not successfully innovate and introduce new technology into our product lines or manage the transitions to new product offerings, our revenues, results of operations and business will be adversely impacted.

Competitors may be able to respond more quickly and effectively than we can to new or changing opportunities, technologies, standards or customer requirements. We anticipate that we will face increased competition in the future as existing companies and competitors develop new or improved products and as new companies enter the market with new technologies.

*New market opportunities may not develop as quickly as we expect, limiting our ability to successfully market and sell our products.*

The market for our products is new and evolving. Accordingly, we expect the application of our technologies to emerging opportunities will take several years to develop and mature and we cannot be certain that these market opportunities will develop as we expect. For example, in September 2012, we launched a single cell gene expression application for our nCounter Analysis System, which applies our technology to, amongst other things, improve single cell analytic workflow for gene expression analysis, and in July 2013, we launched nCounter Elements, a new digital molecular barcoding chemistry that allows users to design their own customized assays using standard sets of barcodes provided by us. The future growth of the market for these products depends on many factors beyond our control, including recognition and acceptance of our applications by the scientific community and the growth, prevalence and costs of competing methods of genomic analysis. If the markets for nCounter Elements, single cell analysis or others do not develop as we expect, our business may be adversely affected. In addition, we commercially launched Prosigna in Europe and Israel in February 2013 and we intend to offer Prosigna in other countries outside of the United States. Genomic testing for breast cancer is not widely available outside of the United States and the market for such tests is new. The future growth of the market for genomic breast cancer testing will depend on physicians' acceptance of such testing and the availability of reimbursement for such tests. Our success in these new markets will depend to a large extent on our ability to successfully market, sell and establish reimbursement for products using our technologies. If we are not able to successfully market and sell our products or to achieve the revenue or margins we expect, our operating results may be harmed and we may not recover our product development and marketing expenditures.

*We are dependent on single source suppliers for some of the components and materials used in our products, and the loss of any of these suppliers could harm our business.*

We rely on Precision System Science, Co., Ltd of Chiba, Japan, to build our nCounter Prep Station and Korvis LLC of Corvallis, Oregon, to build our nCounter Digital Analyzer. Each of these contract manufacturers are sole suppliers. Since our contracts with these instrument suppliers do not commit them to carry inventory or make available any particular quantities, they may give other customers' needs higher priority than ours, and we may not be able to obtain adequate supplies in a timely manner or on commercially reasonable terms. We also rely on sole suppliers for various components we use to manufacture our consumable products. We periodically forecast our needs for such components and enter into standard purchase orders with them. If we were to lose such suppliers, there can be no assurance that we will be able to identify or enter into agreements with alternative suppliers on a timely basis on acceptable terms, if at all. If we should encounter delays or difficulties in securing

## **Table of Contents**

the quality and quantity of materials we require for our products our supply chain would be interrupted which would adversely affect sales. If any of these events occur, our business and operating results could be harmed.

### ***We may experience manufacturing problems or delays that could limit our growth or adversely affect our operating results***

Our consumable products are manufactured at our Seattle facility using complex processes, sophisticated equipment and strict adherence to specifications and quality systems procedures. Any unforeseen manufacturing problems, such as contamination of our facility, equipment malfunction, or failure to strictly follow procedures or meet specifications, could result in delays or shortfalls in production of our consumable products. Identifying and resolving the cause of any such manufacturing issues could require substantial time and resources. If we are unable to keep up with demand for our products by successfully manufacturing and shipping our products in a timely manner, our revenue could be impaired, market acceptance for our products could be adversely affected and our customers might instead purchase our competitors' products.

In addition, the introduction of new products may require the development of new manufacturing processes and procedures. While all of our codesets are produced using the same basic processes, significant variations may be required to meet product specifications. Developing such a process can be very time consuming, and any unexpected difficulty in doing so could delay the introduction of a product.

### ***Our future capital needs are uncertain and we may need to raise additional funds in the future.***

We believe that our existing cash and cash equivalents, including the funds raised in this offering, will be sufficient to meet our anticipated cash requirements for at least the next 12 months. However, we may need to raise substantial additional capital to:

expand the commercialization of our products;

fund our operations; and

further our research and development.

Our future funding requirements will depend on many factors, including:

market acceptance of our products;

the cost and timing of establishing additional sales, marketing and distribution capabilities;

the cost of our research and development activities;

the cost and timing of regulatory clearances or approvals;

the effect of competing technological and market developments; and

the extent to which we acquire or invest in businesses, products and technologies, including new licensing arrangements for new products, although we currently have no commitments or agreements to complete any such transactions.

We cannot assure you that we will be able to obtain additional funds on acceptable terms, or at all. If we raise additional funds by issuing equity or equity-linked securities, our stockholders may experience dilution. Debt financing, if available, may involve covenants restricting our

## Edgar Filing: NanoString Technologies Inc - Form S-1

operations or our ability to incur additional debt. Any debt or additional equity financing that we raise may contain terms that are not favorable to us or our stockholders. If we raise additional funds through collaboration and licensing arrangements with third parties, it may be necessary to relinquish some rights to our technologies or our products, or grant licenses on terms that are not favorable to us. If we are unable to raise adequate funds, we may have to liquidate some or all of our assets, or delay, reduce the scope of or eliminate some or all of our development programs.

If we do not have, or are not able to obtain, sufficient funds, we may have to delay development or commercialization of our products or license to third parties the rights to commercialize products or technologies

## **Table of Contents**

that we would otherwise seek to commercialize. We also may have to reduce marketing, customer support or other resources devoted to our products or cease operations. Any of these factors could harm our operating results.

***If our Seattle facility becomes unavailable or inoperable, we will be unable to continue manufacturing our consumables or process sales orders, and our business will be harmed.***

We manufacture our consumable products in our facility in Seattle, Washington. In addition, our Seattle facility is the center for order processing, receipt of our prep station and digital analyzer manufactured by third-party contract manufacturers and shipping products to customers. Our facility and the equipment we use to manufacture our consumable products would be costly, and would require substantial lead time, to repair or replace. Seattle is situated near active earthquake fault lines. The facility may be harmed or rendered inoperable by natural or man-made disasters, including earthquakes and power outages, which may render it difficult or impossible for us to produce our tests for some period of time. The inability to manufacture consumables or to ship products to customers for even a short period of time may result in the loss of customers or harm our reputation, and we may be unable to regain those customers in the future. Although we possess insurance for damage to our property and the disruption of our business, this insurance, and in particular earthquake insurance, which is limited, may not be sufficient to cover all of our potential losses and may not continue to be available to us on acceptable terms, if at all.

***We expect to generate a substantial portion of our revenue internationally and are subject to various risks relating to our international activities which could adversely affect our operating results.***

During 2012 and the nine months ended September 30, 2013, approximately 31% and 29%, respectively, of our revenue was generated from sales to customers located outside of North America. We believe that a significant percentage of our future revenue will come from international sources as we expand our overseas operations and develop opportunities in additional areas. Engaging in international business involves a number of difficulties and risks, including:

required compliance with existing and changing foreign regulatory requirements and laws;

required compliance with anti-bribery laws, such as the U.S. Foreign Corrupt Practices Act and U.K. Bribery Act, data privacy requirements, labor laws and anti-competition regulations;

export or import restrictions;

various reimbursement and insurance regimes;

laws and business practices favoring local companies;

longer payment cycles and difficulties in enforcing agreements and collecting receivables through certain foreign legal systems;

political and economic instability;

potentially adverse tax consequences, tariffs, customs charges, bureaucratic requirements and other trade barriers;

difficulties and costs of staffing and managing foreign operations; and

## Edgar Filing: NanoString Technologies Inc - Form S-1

difficulties protecting or procuring intellectual property rights.

As we expand internationally our results of operations and cash flows will become increasingly subject to fluctuations due to changes in foreign currency exchange rates. Historically, most of our revenue has been denominated in U.S. dollars, although we have sold our products and services in local currency outside of the United States, principally the Euro. Our expenses are generally denominated in the currencies in which our operations are located, which is primarily in the United States. As our operations in countries outside of the United States grow, our results of operations and cash flows will be subject to fluctuations due to changes in

## **Table of Contents**

foreign currency exchange rates, which could harm our business in the future. For example, if the value of the U.S. dollar increases relative to foreign currencies, in the absence of a corresponding change in local currency prices, our revenue could be adversely affected as we convert revenue from local currencies to U.S. dollars.

If we dedicate significant resources to our international operations and are unable to manage these risks effectively, our business, operating results and prospects will suffer.

***The enactment of legislation implementing changes in the U.S. taxation of international business activities or the adoption of other tax reform policies could materially impact our future financial position and results of operations.***

Recent changes to U.S. tax laws, including limitations on the ability of taxpayers to claim and utilize foreign tax credits and the deferral of certain tax deductions until earnings outside of the United States are repatriated to the United States, as well as changes to U.S. tax laws that may be enacted in the future, could impact the tax treatment of future foreign earnings. Should the scale of our international business activities expand, any changes in the U.S. taxation of such activities could increase our worldwide effective tax rate and harm our future financial position and results of operations.

***Our ability to use net operating losses to offset future taxable income may be subject to certain limitations.***

As of December 31, 2012, we had federal net operating loss carryforwards, or NOLs, to offset future taxable income of approximately \$62.2 million, which expire in various years beginning in 2023, if not utilized. A lack of future taxable income would adversely affect our ability to utilize these NOLs. In addition, under Section 382 of the Internal Revenue Code, a corporation that undergoes an ownership change is subject to limitations on its ability to utilize its NOLs to offset future taxable income. We may have already experienced one or more ownership changes. Depending on the timing of any future utilization of our carryforwards, we may be limited as to the amount that can be utilized each year as a result of such previous ownership changes. In addition, future changes in our stock ownership, including this or future offerings, as well as other changes that may be outside of our control, could result in additional ownership changes under Section 382 of the Internal Revenue Code. Our NOLs may also be impaired under similar provisions of state law. We have recorded a full valuation allowance related to our NOLs and other deferred tax assets due to the uncertainty of the ultimate realization of the future benefits of those assets.

***Provisions of our debt instruments may restrict our ability to pursue our business strategies.***

Our credit facility requires us, and any debt instruments we may enter into in the future may require us, to comply with various covenants that limit our ability to, among other things:

dispose of assets;

complete mergers or acquisitions;

incur indebtedness;

encumber assets;

pay dividends or make other distributions to holders of our capital stock;

make specified investments;



## Edgar Filing: NanoString Technologies Inc - Form S-1

change certain key management personnel; and

engage in transactions with our affiliates.

These restrictions could inhibit our ability to pursue our business strategies. In addition, we are subject to a financial covenant based on life sciences revenue. If we default under our credit facility, and such event of

-20-

## **Table of Contents**

default was not cured or waived, the lenders could terminate commitments to lend and cause all amounts outstanding with respect to the debt to be due and payable immediately, which in turn could result in cross defaults under other debt instruments. Our assets and cash flow may not be sufficient to fully repay borrowings under all of our outstanding debt instruments if some or all of these instruments are accelerated upon a default.

We may incur additional indebtedness in the future. For example, in January 2014, we entered into a non-binding letter of intent for a credit agreement with a lender which would allow us to refinance our existing credit facility and potentially incur up to an aggregate of \$45 million in term loan borrowings or up to an aggregate of approximately \$52 million if we elect to exercise in full an option to pay in kind a portion of the interest that would accrue on the borrowings under the credit agreement. The debt instruments governing such indebtedness could contain provisions that are as, or more, restrictive than our existing debt instruments. If we are unable to repay, refinance or restructure our indebtedness when payment is due, the lenders could proceed against the collateral granted to them to secure such indebtedness or force us into bankruptcy or liquidation.

### ***Acquisitions or joint ventures could disrupt our business, cause dilution to our stockholders and otherwise harm our business.***

We may acquire other businesses, products or technologies as well as pursue strategic alliances, joint ventures, technology licenses or investments in complementary businesses. We have not made any acquisitions to date, and our ability to do so successfully is unproven. Any of these transactions could be material to our financial condition and operating results and expose us to many risks, including:

disruption in our relationships with customers, distributors or suppliers as a result of such a transaction;

unanticipated liabilities related to acquired companies;

difficulties integrating acquired personnel, technologies and operations into our existing business;

diversion of management time and focus from operating our business to acquisition integration challenges;

increases in our expenses and reductions in our cash available for operations and other uses; and

possible write-offs or impairment charges relating to acquired businesses.

Foreign acquisitions involve unique risks in addition to those mentioned above, including those related to integration of operations across different cultures and languages, currency risks and the particular economic, political and regulatory risks associated with specific countries.

Also, the anticipated benefit of any acquisition may not materialize. Future acquisitions or dispositions could result in potentially dilutive issuances of our equity securities, the incurrence of debt, contingent liabilities or amortization expenses or write-offs of goodwill, any of which could harm our financial condition. We cannot predict the number, timing or size of future joint ventures or acquisitions, or the effect that any such transactions might have on our operating results.

### ***If we are unable to recruit, train and retain key personnel, we may not achieve our goals.***

Our future success depends on our ability to recruit, train, retain and motivate key personnel, including our senior management, research and development, manufacturing and sales and marketing personnel. Competition for qualified personnel is intense, particularly in the Seattle, Washington area. Our growth depends, in particular, on attracting, retaining and motivating highly-trained sales personnel with the necessary scientific background and ability to understand our systems at a technical level to effectively identify and sell to potential new customers. In particular, the commercial launch of Prosigna requires us to establish a dedicated oncology diagnostics sales force to fully optimize the breast cancer diagnostic market opportunity. We do not maintain fixed term employment contracts or key man life insurance with any of our employees. Because of the complex and technical nature of our products and the dynamic market in which we compete, any failure to attract,

train, retain and motivate qualified personnel could materially harm our operating results and growth prospects.

## **Table of Contents**

### ***Undetected errors or defects in our products could harm our reputation, decrease market acceptance of our products or expose us to product liability claims.***

Our products may contain undetected errors or defects when first introduced or as new versions are released. Disruptions or other performance problems with our products may damage our customers' business and could harm our reputation. If that occurs, we may incur significant costs, the attention of our key personnel could be diverted, or other significant customer relations problems may arise. We may also be subject to warranty and liability claims for damages related to errors or defects in our products. A material liability claim or other occurrence that harms our reputation or decreases market acceptance of our products could harm our business and operating results.

The sale and use of products or services based on our technologies, or activities related to our research and clinical studies, could lead to the filing of product liability claims if someone were to allege that one of our products contained a design or manufacturing defect which resulted in the failure to adequately perform the analysis for which it was designed. A product liability claim could result in substantial damages and be costly and time consuming to defend, either of which could materially harm our business or financial condition. We cannot assure investors that our product liability insurance would adequately protect our assets from the financial impact of defending a product liability claim. Any product liability claim brought against us, with or without merit, could increase our product liability insurance rates or prevent us from securing insurance coverage in the future.

### ***We face risks related to handling of hazardous materials and other regulations governing environmental safety.***

Our operations are subject to complex and stringent environmental, health, safety and other governmental laws and regulations that both public officials and private individuals may seek to enforce. Our activities that are subject to these regulations include, among other things, our use of hazardous materials and the generation, transportation and storage of waste. We could discover that we or an acquired business is not in material compliance with these regulations. Existing laws and regulations may also be revised or reinterpreted, or new laws and regulations may become applicable to us, whether retroactively or prospectively, that may have a negative effect on our business and results of operations. It is also impossible to eliminate completely the risk of accidental environmental contamination or injury to individuals. In such an event, we could be liable for any damages that result, which could adversely affect our business.

### **Risks Related to Government Regulation and Diagnostic Product Reimbursement**

#### ***Our research use only products for the life sciences market could become subject to regulation as medical devices by the FDA or other regulatory agencies in the future which could increase our costs and delay our commercialization efforts, thereby materially and adversely affecting our life sciences business and results of operations.***

In the United States, most of our products are currently labeled and sold for research use only, or RUO, and not for the diagnosis or treatment of disease, and are sold to pharmaceutical and biotechnology companies, academic institutions and life sciences laboratories. Because such products are not intended for use in clinical practice in diagnostics, and the products cannot include clinical or diagnostic claims, they are not subject to regulation by the FDA as medical devices. In particular, while the FDA regulations require that RUO products be labeled, "For Research Use Only. Not for use in diagnostic procedures," the regulations do not subject such products to the FDA's pre- and post-market controls for medical devices. In November 2013, the FDA issued a final guidance on RUO products, which, among other things, reaffirmed that a company may not make clinical or diagnostic claims about an RUO product. Although not suggested in the final RUO guidance, if in the future the FDA modifies its approach to regulating our products labeled for research use only, it could reduce our revenue or increase our costs and adversely affect our business, prospects, results of operations or financial condition. In the event that the FDA requires marketing authorization of our RUO products in the future, there can be no assurance that the FDA will ultimately grant any clearance or approval requested by us in a timely manner, or at all.

---

## **Table of Contents**

In addition, we sell dual-use instruments with software that have both FDA-cleared functions and research functions, for which FDA approval or clearance is not required. Dual-use instruments are subject to FDA regulation since they are intended, at least in part, for use by customers performing clinical diagnostic testing. There is a risk that the FDA could take enforcement action against a manufacturer for distributing dual-use instruments if the FDA determines that approval or clearance was required for those functions for which FDA approval or clearance has not been obtained, and the instruments are being sold off-label. There is also a risk that the FDA could broaden its current regulatory enforcement of dual-use instruments through additional FDA oversight.

***Our GPRs may be used by clinical laboratories to create Laboratory Developed Tests, which could in the future be subject to regulation as medical devices, which could materially and adversely affect our life sciences business and results of operations.***

Recently, we launched nCounter Elements, a new digital molecular barcoding chemistry that allows users to design their own customized assays using standard sets of barcodes provided by us with the laboratories' choice of oligonucleotide probes. nCounter Elements are considered GPRs by the FDA, that are Class I medical devices, and we listed nCounter Elements with the FDA as GPRs in July 2013.

A clinical laboratory can use nCounter Elements to create what is called a Laboratory Developed Test. Laboratory Developed Tests are diagnostic tests that are developed and performed by a laboratory and include genetic tests and other tests for rare conditions. In June 2013, the Commissioner of the FDA stated that the FDA intends to further regulate Laboratory Developed Tests; however, it is unclear whether, when and to what extent the FDA will do so. Restrictions on Laboratory Developed Tests by the FDA could restrict the demand for our products, including nCounter Elements. Additionally, compliance with additional regulatory burdens could be time consuming and costly. If the FDA regulates Laboratory Developed Tests, such regulation could adversely affect our prospects, results of operations and financial condition.

***Approval and/or clearance by the FDA and foreign regulatory authorities for our diagnostic tests will take significant time and require significant research, development and clinical study expenditures and ultimately may not succeed.***

Before we begin to label and market our products for use as clinical diagnostics in the United States, thereby subjecting them to FDA regulation as medical devices, unless an exemption applies, we are required to obtain either prior 510(k) clearance or prior pre-market approval, or PMA, from the FDA. In September 2013, we received FDA 510(k) clearance for Prosigna as a prognostic indicator for distant recurrence-free survival at 10 years in post-menopausal women with Stage I/II lymph node-negative or Stage II lymph node-positive (1-3 positive nodes) hormone receptor-positive breast cancer who have undergone surgery in conjunction with locoregional treatment and consistent with standard of care. In the future we plan to submit a separate application for approval of Prosigna to report intrinsic subtype and we expect that this application will require a PMA supported by additional clinical studies. We intend to pursue additional intended uses for Prosigna, which may require more burdensome regulatory processes than the 510(k) clearance process, including PMAs. Even if granted, a 510(k) clearance or PMA approval for any future product would likely place substantial restrictions on how our device is marketed or sold, and the FDA will continue to place considerable restrictions on our products, including, but not limited to, quality system regulations, or QSR, registering manufacturing facilities, listing the products with the FDA, and complying with labeling, marketing, complaint handling, adverse event and medical device reporting requirements and corrections and removals. Obtaining FDA clearance or approval for diagnostics can be expensive and uncertain, and generally takes from several months to several years, and generally requires detailed and comprehensive scientific and clinical data. Notwithstanding the expense, these efforts may never result in FDA approval or clearance. Even if we were to obtain regulatory approval or clearance, it may not be for the uses we believe are important or commercially attractive, in which case we would not be permitted to market our product for those uses.

## **Table of Contents**

Sales of our diagnostic products outside the United States are subject to foreign regulatory requirements governing clinical studies, vigilance reporting, marketing approval, manufacturing, product licensing, pricing and reimbursement. These regulatory requirements vary greatly from country to country. As a result, the time required to obtain approvals outside the United States may differ from that required to obtain FDA approval, and we may not be able to obtain foreign regulatory approvals on a timely basis or at all. Approval by the FDA does not ensure approval by regulatory authorities in other countries, and approval by one foreign regulatory authority does not ensure approval by regulatory authorities in other countries or by the FDA, and foreign regulatory authorities could require additional testing. In addition, FDA regulates exports of medical devices. Failure to comply with these regulatory requirements or to obtain required approvals could impair our ability to commercialize our diagnostic products outside of the United States.

***We expect to rely on third parties to conduct any future studies of our diagnostic products that may be required by the FDA or other regulatory authorities, and those third parties may not perform satisfactorily.***

We do not have the ability to independently conduct the clinical studies or other studies that may be required to obtain FDA and other regulatory clearance or approval for our diagnostic products, including Prosigna. Accordingly, we expect to rely on third parties, such as medical institutions and clinical investigators, to conduct such studies. Our reliance on these third parties for clinical development activities will reduce our control over these activities. These third-party contractors may not complete activities on schedule or conduct studies in accordance with regulatory requirements or our study design. Our reliance on third parties that we do not control will not relieve us of any applicable requirement to prepare, and ensure compliance with, various procedures required under good clinical practices. If these third parties do not successfully carry out their contractual duties or regulatory obligations or meet expected deadlines, if the third parties need to be replaced or if the quality or accuracy of the data they obtain is compromised due to their failure to adhere to our clinical protocols or regulatory requirements or for other reasons, our studies may be extended, delayed, suspended or terminated, and we may not be able to obtain regulatory approval for our diagnostic products.

***We are subject to ongoing and extensive regulatory requirements, and our failure to comply with these requirements could substantially harm our business.***

Following our obtaining CE Mark in the EU and receipt of FDA 510(k) clearance in September 2013 for Prosigna, we are subject to ongoing ISO and FDA obligations and continued regulatory oversight and review, including routine inspections by EU Notified Bodies and by the FDA of our manufacturing facilities and compliance with requirements such as ISO 13485 and quality system regulations, or QSRs, which establish extensive requirements for quality assurance and control as well as manufacturing procedures; requirements pertaining to the registration of our manufacturing facilities and the listing of our devices with the FDA; continued complaint, adverse event and malfunction reporting; corrections and removals reporting; and labeling and promotional requirements. The promotional claims we can make for Prosigna are limited to the cleared indication. For instance, in the United States the following special conditions for use are listed in the intended use: Prosigna is not intended for diagnosis, to predict or detect response to therapy or to help select the optimal therapy for patients. We may also be subject to additional FDA post-marketing obligations. If we are not able to maintain regulatory compliance, we may not be permitted to market our diagnostic products and/or may be subject to enforcement by EU Competent Authorities and the FDA such as the issuance of warning or untitled letters, fines, injunctions, and civil penalties; recall or seizure of products; operating restrictions; and criminal prosecution. In addition, we may be subject to similar regulatory regimes of foreign jurisdictions as we continue to commercialize our products in new markets. Adverse Notified Body, EU Competent Authority or FDA action in any of these areas could significantly increase our expenses and limit our revenue and profitability.

***If Medicare and other third-party payors in the United States and foreign countries do not approve reimbursement for diagnostic tests enabled by our technology, the commercial success of our diagnostic products would be compromised.***

Successful commercialization of our diagnostic products depends, in large part, on the availability of adequate reimbursement for testing services that our diagnostic products enable from government insurance

## **Table of Contents**

plans, managed care organizations and private insurance plans. There is significant uncertainty surrounding third-party reimbursement for the use of tests that incorporate new technology, such as Prosigna. If we are unable to obtain positive policy decisions from third-party payors approving reimbursement for our tests at adequate levels, the commercial success of our products would be compromised and our revenue would be significantly limited. Even if we do obtain reimbursement for our tests, Medicare, Medicaid and private and other payors may withdraw their coverage policies, cancel their contracts with us at any time, review and adjust the rate of reimbursement, require co-payments from patients or stop paying for our tests, which would reduce revenue for testing services based on our technology, and indirectly, demand for diagnostic products. In addition, insurers, including managed care organizations as well as government payors such as Medicare and Medicaid, have increased their efforts to control the cost, utilization and delivery of healthcare services, which may include decreased coverage or reduced reimbursement. From time to time, Congress has considered and implemented changes to the Medicare fee schedules in conjunction with budgetary legislation, and pricing and payment terms, including the possible requirement of a patient co-payment for Medicare beneficiaries for tests covered by Medicare, and are subject to change at any time. Reductions in the reimbursement rate of third-party payors have occurred and may occur in the future. Reductions in the prices at which testing services based on our technology are reimbursed could have a negative impact on our revenue.

In many countries outside of the United States, various coverage, pricing and reimbursement approvals are required. We expect that it will take several years to establish broad coverage and reimbursement for testing services based on our products with payors in countries outside of the United States, and our efforts may not be successful.

***We may be subject, directly or indirectly, to federal and state healthcare fraud and abuse laws and other federal and state laws applicable to our marketing practices. If we are unable to comply, or have not complied, with such laws, we could face substantial penalties.***

As we begin commercializing Prosigna and any other potential diagnostic products in the United States, our operations will be directly, or indirectly through our customers, subject to various federal and state fraud and abuse laws, including, without limitation, the federal and state anti-kickback statutes and state and federal marketing compliance laws and gift bans. These laws may impact, among other things, our proposed sales and marketing and education programs and require us to implement additional internal systems for tracking certain marketing expenditures and reporting them to government authorities. In addition, we may be subject to patient privacy regulation by both the federal government and the states in which we conduct our business. The laws that may affect our ability to operate include:

the federal Anti-kickback Law and state anti-kickback prohibitions;

the federal physician self-referral prohibition, commonly known as the Stark Law, and the state equivalents;

the federal Health Insurance Portability and Accountability Act of 1996, as amended;

the Medicare civil money penalty and exclusion requirements;

the federal False Claims Act civil and criminal penalties and state equivalents; and

state physician gift bans and marketing expenditure laws.

If our operations are found to be in violation of any of the laws described above or any other governmental regulations that apply to us, we may be subject to penalties, including civil and criminal penalties, damages, fines and the curtailment or restructuring of our operations, any of which could adversely affect our ability to operate our business and our results of operations.

***Healthcare policy changes, including legislation reforming the United States healthcare system, may have a material adverse effect on our financial condition and results of operations.***

## Edgar Filing: NanoString Technologies Inc - Form S-1

The Patient Protection and Affordable Care Act, as amended by the Health Care and Education Affordability Reconciliation Act, collectively, the PPACA, enacted in March 2010, makes changes that are expected to significantly impact the pharmaceutical and medical device industries and clinical laboratories.

-25-



---

## **Table of Contents**

Beginning in 2013, each medical device manufacturer will have to pay a sales tax in an amount equal to 2.3% of the price for which such manufacturer sells its medical devices. We expect that the new tax will apply to some or all of our diagnostic products. The PPACA also mandates a reduction in payments for clinical laboratory services paid under the Medicare Clinical Laboratory Fee Schedule of 1.75% for the years 2011 through 2015 and a productivity adjustment to the Clinical Laboratory Fee Schedule. These or any future proposed or mandated reductions in payments may apply to some or all of the clinical laboratory tests that our diagnostics customers use our technology to deliver to Medicare beneficiaries, and may indirectly reduce demand for our diagnostic products.

Other significant measures contained in the PPACA include coordination and promotion of research on comparative clinical effectiveness of different technologies and procedures, initiatives to revise Medicare payment methodologies, such as bundling of payments across the continuum of care by providers and physicians, and initiatives to promote quality indicators in payment methodologies. The PPACA also includes significant new fraud and abuse measures, including required disclosures of financial arrangements with physician customers, lower thresholds for violations and increasing potential penalties for such violations. In addition, the PPACA establishes an Independent Payment Advisory Board, or IPAB, to reduce the per capita rate of growth in Medicare spending. The IPAB has broad discretion to propose policies to reduce health care expenditures, which may have a negative impact on payment rates for services, including our tests. The IPAB proposals may impact payments for clinical laboratory services that our diagnostics customers use our technology to deliver beginning in 2016 and for hospital services beginning in 2020, and may indirectly reduce demand for our diagnostic products.

In addition to the PPACA, the effect of which cannot presently be quantified, various healthcare reform proposals have also emerged from federal and state governments. Changes in healthcare policy, such as the creation of broad test utilization limits for diagnostic products in general or requirements that Medicare patients pay for portions of clinical laboratory tests or services received, could substantially impact the sales of our tests, increase costs and divert management's attention from our business. Such co-payments by Medicare beneficiaries for laboratory services were discussed as possible cost savings for the Medicare program as part of the debt ceiling budget discussions in mid-2011 and may be enacted in the future. In addition, sales of our tests outside of the United States will subject us to foreign regulatory requirements, which may also change over time.

We cannot predict whether future healthcare initiatives will be implemented at the federal or state level or in countries outside of the United States in which we may do business, or the effect any future legislation or regulation will have on us. The taxes imposed by the new federal legislation and the expansion in government's effect on the United States healthcare industry may result in decreased profits to us, lower reimbursements by payors for our products or reduced medical procedure volumes, all of which may adversely affect our business, financial condition and results of operations.

### **Risks Related to Intellectual Property**

*If we are unable to protect our intellectual property effectively, our business would be harmed.*

We rely on patent protection as well as trademark, copyright, trade secret and other intellectual property rights protection and contractual restrictions to protect our proprietary technologies, all of which provide limited protection and may not adequately protect our rights or permit us to gain or keep any competitive advantage. As of December 31, 2013, we owned or exclusively licensed seven issued U.S. patents and approximately 23 pending U.S. patent applications, including provisional and non-provisional filings. We also owned or licensed approximately 73 pending and granted counterpart applications worldwide, including 22 country-specific validations of four European patents. If we fail to protect our intellectual property, third parties may be able to compete more effectively against us and we may incur substantial litigation costs in our attempts to recover or restrict use of our intellectual property.

We cannot assure investors that any of our currently pending or future patent applications will result in issued patents, and we cannot predict how long it will take for such patents to be issued. Further, we cannot

## **Table of Contents**

assure investors that other parties will not challenge any patents issued to us or that courts or regulatory agencies will hold our patents to be valid or enforceable. We cannot guarantee investors that we will be successful in defending challenges made against our patents and patent applications. Any successful third-party challenge to our patents could result in the third party or the unenforceability or invalidity of such patents.

The patent positions of life sciences companies can be highly uncertain and involve complex legal and factual questions for which important legal principles remain unresolved. No consistent policy regarding the breadth of claims allowed in such companies' patents has emerged to date in the United States. Furthermore, in the biotechnology field, courts frequently render opinions that may affect the patentability of certain inventions or discoveries, including opinions that may affect the patentability of methods for analyzing or comparing DNA.

In particular, the patent positions of companies engaged in development and commercialization of genomic diagnostic tests, like Prosigna, are particularly uncertain. Various courts, including the U.S. Supreme Court, have recently rendered decisions that impact the scope of patentability of certain inventions or discoveries relating to genomic diagnostics. Specifically these decisions stand for the proposition that patent claims that recite laws of nature (for example, the relationships between gene expression levels and the likelihood of risk of recurrence of cancer) are not themselves patentable unless those patent claims have sufficient additional features that provide practical assurance that the processes are genuine inventive applications of those laws rather than patent drafting efforts designed to monopolize the law of nature itself. What constitutes a sufficient additional feature is uncertain. Accordingly, this evolving case law in the United States may adversely impact our ability to obtain new patents and may facilitate third-party challenges to our existing owned and licensed patents. One of our main areas of intellectual property, namely patents we license directed to the use of gene expression markers as part of genomic diagnostic tests, may be affected by these decisions.

The laws of some non-U.S. countries do not protect intellectual property rights to the same extent as the laws of the United States, and many companies have encountered significant problems in protecting and defending such rights in foreign jurisdictions. The legal systems of certain countries, particularly certain developing countries, do not favor the enforcement of patents and other intellectual property protection, particularly those relating to biotechnology, which could make it difficult for us to stop the infringement of our patents. Proceedings to enforce our patent rights in foreign jurisdictions could result in substantial cost and divert our efforts and attention from other aspects of our business.

Changes in either the patent laws or in interpretations of patent laws in the United States or other countries may diminish the value of our intellectual property. We cannot predict the breadth of claims that may be allowed or enforced in our patents or in third-party patents. For example:

We might not have been the first to make the inventions covered by each of our pending patent applications.

We might not have been the first to file patent applications for these inventions.

Others may independently develop similar or alternative products and technologies or duplicate any of our products and technologies.

It is possible that our pending patent applications will not result in issued patents, and even if they issue as patents, they may not provide a basis for commercially viable products, may not provide us with any competitive advantages, or may be challenged and invalidated by third parties.

We may not develop additional proprietary products and technologies that are patentable.

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

## Edgar Filing: NanoString Technologies Inc - Form S-1

We apply for patents covering our products and technologies and uses thereof, as we deem appropriate. However, we may fail to apply for patents on important products and technologies in a timely fashion or at all.

In addition to pursuing patents on our technology, we take steps to protect our intellectual property and proprietary technology by entering into confidentiality agreements and intellectual property assignment

-27-

## Table of Contents

agreements with our employees, consultants, corporate partners and, when needed, our advisors. Such agreements may not be enforceable or may not provide meaningful protection for our trade secrets or other proprietary information in the event of unauthorized use or disclosure or other breaches of the agreements, and we may not be able to prevent such unauthorized disclosure. Monitoring unauthorized disclosure is difficult, and we do not know whether the steps we have taken to prevent such disclosure are, or will be, adequate. If we were to enforce a claim that a third party had illegally obtained and was using our trade secrets, it would be expensive and time consuming, and the outcome would be unpredictable. In addition, courts outside the United States may be less willing to protect trade secrets.

In addition, competitors could purchase our products and attempt to replicate some or all of the competitive advantages we derive from our development efforts, willfully infringe our intellectual property rights, design around our protected technology or develop their own competitive technologies that fall outside of our intellectual property rights. If our intellectual property is not adequately protected so as to protect our market against competitors' products and methods, our competitive position could be adversely affected, as could our business.

We have not yet registered certain of our trademarks, including Prosigna, in all of our potential markets. If we apply to register these trademarks, our applications may not be allowed for registration, and our registered trademarks may not be maintained or enforced. In addition, opposition or cancellation proceedings may be filed against our trademark applications and registrations, and our trademarks may not survive such proceedings. If we do not secure registrations for our trademarks, we may encounter more difficulty in enforcing them against third parties than we otherwise would.

To the extent our intellectual property, including licensed intellectual property, offers inadequate protection, or is found to be invalid or unenforceable, we would be exposed to a greater risk of direct competition. If our intellectual property does not provide adequate protection against our competitors' products, our competitive position could be adversely affected, as could our business. Both the patent application process and the process of managing patent disputes can be time consuming and expensive.

***We depend on certain technologies that are licensed to us. We do not control these technologies and any loss of our rights to them could prevent us from selling our products.***

We rely on licenses in order to be able to use various proprietary technologies that are material to our business, including our core digital molecular barcoding technology licensed from the Institute for Systems Biology and technology relating to Prosigna licensed from Bioclassifier, LLC. We do not own the patents that underlie these licenses. Our rights to use these technologies and employ the inventions claimed in the licensed patents are subject to the continuation of and compliance with the terms of those licenses.

In some cases, we do not control the prosecution, maintenance, or filing of the patents to which we hold licenses, or the enforcement of these patents against third parties. Some of our patents and patent applications were either acquired from another company who acquired those patents and patent applications from yet another company, or are licensed from a third party. Thus, these patents and patent applications are not written by us or our attorneys, and we did not have control over the drafting and prosecution. The former patent owners and our licensors might not have given the same attention to the drafting and prosecution of these patents and applications as we would have if we had been the owners of the patents and applications and had control over the drafting and prosecution. We cannot be certain that drafting or prosecution of the licensed patents and patent applications by the licensors have been or will be conducted in compliance with applicable laws and regulations or will result in valid and enforceable patents and other intellectual property rights.

Enforcement of our licensed patents or defense of any claims asserting the invalidity of these patents is often subject to the control or cooperation of our licensors. Certain of our licenses contain provisions that allow the licensor to terminate the license upon specific conditions. Our rights under the licenses are subject to our continued compliance with the terms of the license, including the payment of royalties due under the license. Because of the complexity of our products and the patents we have licensed, determining the scope of the license

## **Table of Contents**

and related royalty obligation can be difficult and can lead to disputes between us and the licensor. An unfavorable resolution of such a dispute could lead to an increase in the royalties payable pursuant to the license or termination of the license. If a licensor believed we were not paying the royalties due under the license or were otherwise not in compliance with the terms of the license, the licensor might attempt to revoke the license. If such an attempt were successful, we might be barred from producing and selling some or all of our products.

In addition, certain of the patents we have licensed relate to technology that was developed with U.S. government grants. Federal regulations impose certain domestic manufacturing requirements with respect to some of our products embodying these patents.

***We may be involved in lawsuits to protect or enforce our patents and proprietary rights, to determine the scope, coverage and validity of others' proprietary rights, or to defend against third-party claims of intellectual property infringement, any of which could be time-intensive and costly and may adversely impact our business or stock price.***

We have received notices of claims of infringement and misappropriation or misuse of other parties' proprietary rights in the past and may from time to time receive additional notices. Some of these claims may lead to litigation. We cannot assure investors that we will prevail in such actions, or that other actions alleging misappropriation or misuse by us of third-party trade secrets, infringement by us of third-party patents and trademarks or other rights, or the validity of our patents, trademarks or other rights, will not be asserted or prosecuted against us.

Litigation may be necessary for us to enforce our patent and proprietary rights or to determine the scope, coverage and validity of the proprietary rights of others. Litigation could result in substantial legal fees and could adversely affect the scope of our patent protection. The outcome of any litigation or other proceeding is inherently uncertain and might not be favorable to us, and we might not be able to obtain licenses to technology that we require. Even if such licenses are obtainable, they may not be available at a reasonable cost. We could therefore incur substantial costs related to royalty payments for licenses obtained from third parties, which could negatively affect our gross margins. Further, we could encounter delays in product introductions, or interruptions in product sales, as we develop alternative methods or products. In addition, if we resort to legal proceedings to enforce our intellectual property rights or to determine the validity, scope and coverage of the intellectual property or other proprietary rights of others, the proceedings could be burdensome and expensive, even if we were to prevail. Any litigation that may be necessary in the future could result in substantial costs and diversion of resources and could have a material adverse effect on our business, operating results or financial condition.

As we move into new markets and applications for our products, incumbent participants in such markets may assert their patents and other proprietary rights against us as a means of slowing our entry into such markets or as a means to extract substantial license and royalty payments from us. Our competitors and others may now and in the future have significantly larger and more mature patent portfolios than we currently have. In addition, future litigation may involve patent holding companies or other adverse patent owners who have no relevant product revenue and against whom our own patents may provide little or no deterrence or protection. Therefore, our commercial success may depend in part on our non-infringement of the patents or proprietary rights of third parties. We are aware of a third party, Genomic Health, Inc., that has issued patents and pending patent applications in the United States, Europe and other jurisdictions that claim methods of using certain genes that are included in Prosigna. We believe that Prosigna does not infringe any valid issued claim. Numerous significant intellectual property issues have been litigated, and will likely continue to be litigated, between existing and new participants in our existing and targeted markets and competitors may assert that our products infringe their intellectual property rights as part of a business strategy to impede our successful entry into those markets. Third parties may assert that we are employing their proprietary technology without authorization. In addition, our competitors and others may have patents or may in the future obtain patents and claim that use of our products infringes these patents. We could incur substantial costs and divert the attention of our management and technical personnel in defending against any of these claims. Parties making claims against us may be able to obtain injunctive or other relief, which could block our ability to develop, commercialize and sell products, and

## **Table of Contents**

could result in the award of substantial damages against us. In the event of a successful claim of infringement against us, we may be required to pay damages and obtain one or more licenses from third parties, or be prohibited from selling certain products. We may not be able to obtain these licenses at a reasonable cost, if at all. We could therefore incur substantial costs related to royalty payments for licenses obtained from third parties, which could negatively affect our gross margins. In addition, we could encounter delays in product introductions while we attempt to develop alternative methods or products to avoid infringing third-party patents or proprietary rights. Defense of any lawsuit or failure to obtain any of these licenses on favorable terms could prevent us from commercializing products, and the prohibition of sale of any of our products could materially affect our ability to grow and gain market acceptance for our products.

Furthermore, because of the substantial amount of discovery required in connection with intellectual property litigation, there is a risk that some of our confidential information could be compromised by disclosure during this type of litigation. In addition, during the course of this kind of litigation, there could be public announcements of the results of hearings, motions or other interim proceedings or developments. If securities analysts or investors perceive these results to be negative, it could have a substantial adverse effect on the price of our common stock.

In addition, our agreements with some of our suppliers, distributors, customers and other entities with whom we do business require us to defend or indemnify these parties to the extent they become involved in infringement claims against us, including the claims described above. We could also voluntarily agree to defend or indemnify third parties in instances where we are not obligated to do so if we determine it would be important to our business relationships. If we are required or agree to defend or indemnify any of these third parties in connection with any infringement claims, we could incur significant costs and expenses that could adversely affect our business, operating results, or financial condition.

***We may be subject to damages resulting from claims that we or our employees have wrongfully used or disclosed alleged trade secrets of our employees former employers.***

Many of our employees were previously employed at universities or other life sciences companies, including our competitors or potential competitors. Although no claims against us are currently pending, we may be subject to claims that these employees or we have inadvertently or otherwise used or disclosed trade secrets or other proprietary information of their former employers. Litigation may be necessary to defend against these claims. If we fail in defending such claims, in addition to paying monetary damages, we may lose valuable intellectual property rights. A loss of key research personnel work product could hamper or prevent our ability to commercialize certain potential products, which could severely harm our business. Even if we are successful in defending against these claims, litigation could result in substantial costs and be a distraction to management.

***Our products contain third-party open source software components, and failure to comply with the terms of the underlying open source software licenses could restrict our ability to sell our products.***

Our products contain software tools licensed by third-party authors under open source licenses. Use and distribution of open source software may entail greater risks than use of third-party commercial software, as open source licensors generally do not provide warranties or other contractual protections regarding infringement claims or the quality of the code. Some open source licenses contain requirements that we make available source code for modifications or derivative works we create based upon the type of open source software we use. If we combine our proprietary software with open source software in a certain manner, we could, under certain open source licenses, be required to release the source code of our proprietary software to the public. This would allow our competitors to create similar products with less development effort and time and ultimately could result in a loss of product sales.

Although we monitor our use of open source software to avoid subjecting our products to conditions we do not intend, the terms of many open source licenses have not been interpreted by U.S. courts, and there is a risk that these licenses could be construed in a way that could impose unanticipated conditions or restrictions on our ability to commercialize our products. Moreover, we cannot assure investors that our processes for controlling

## **Table of Contents**

our use of open source software in our products will be effective. If we are held to have breached the terms of an open source software license, we could be required to seek licenses from third parties to continue offering our products on terms that are not economically feasible, to re-engineer our products, to discontinue the sale of our products if re-engineering could not be accomplished on a timely basis, or to make generally available, in source code form, our proprietary code, any of which could adversely affect our business, operating results, and financial condition.

*We use third-party software that may be difficult to replace or cause errors or failures of our products that could lead to lost customers or harm to our reputation.*

We use software licensed from third parties in our products. In the future, this software may not be available to us on commercially reasonable terms, or at all. Any loss of the right to use any of this software could result in delays in the production of our products until equivalent technology is either developed by us, or, if available, is identified, obtained and integrated, which could harm our business. In addition, any errors or defects in third-party software, or other third-party software failures could result in errors, defects or cause our products to fail, which could harm our business and be costly to correct. Many of these providers attempt to impose limitations on their liability for such errors, defects or failures, and if enforceable, we may have additional liability to our customers or third-party providers that could harm our reputation and increase our operating costs.

We will need to maintain our relationships with third-party software providers and to obtain software from such providers that does not contain any errors or defects. Any failure to do so could adversely impact our ability to deliver reliable products to our customers and could harm our results of operations.

### **Risks Related to Our Common Stock and this Offering**

*The price of our common stock may be volatile, and you could lose all or part of your investment.*

The trading price of our common stock has fluctuated and may continue to fluctuate substantially. Since shares of our common stock were sold in our initial public offering in June 2013 at a price of \$10.00 per share, the reported high and low sales prices of our common stock ranged from \$20.79 to \$7.01 through January 10, 2014. The trading price of our common stock depends on a number of factors, including those described in this Risk Factors section, many of which are beyond our control and may not be related to our operating performance. These fluctuations could cause you to lose all or part of your investment in our common stock since you might be unable to sell your shares at or above the price you paid in this offering. Factors that could cause fluctuations in the trading price of our common stock include the following:

actual or anticipated quarterly variation in our results of operations or the results of our competitors;

announcements by us or our competitors of new products, significant contracts, commercial relationships or capital commitments;

failure to obtain or delays in obtaining product approvals or clearances from the FDA or foreign regulators;

adverse regulatory or reimbursement announcements;

issuance of new or changed securities analysts' reports or recommendations for our stock;

developments or disputes concerning our intellectual property or other proprietary rights;

commencement of, or our involvement in, litigation;

## Edgar Filing: NanoString Technologies Inc - Form S-1

market conditions in the life sciences research and molecular diagnostics markets;

manufacturing disruptions;

any future sales of our common stock or other securities;

any change to the composition of the board of directors or key personnel;

-31-



## **Table of Contents**

expiration of contractual lock-up agreements with our executive officers, directors and security holders;

announcements by us or our competitors of significant acquisitions, strategic partnerships, joint ventures or capital commitments;

general economic conditions and slow or negative growth of our markets; and

the other factors described in this Risk Factors section.

The stock market in general, and market prices for the securities of life sciences and diagnostic companies like ours in particular, have from time to time experienced volatility that often has been unrelated to the operating performance of the underlying companies. These broad market and industry fluctuations may adversely affect the market price of our common stock, regardless of our operating performance. In several recent situations where the market price of a stock has been volatile, holders of that stock have instituted securities class action litigation against the company that issued the stock. If any of our stockholders were to bring a lawsuit against us, the defense and disposition of the lawsuit could be costly and divert the time and attention of our management and harm our operating results.

### ***An active trading market for our common stock may not be sustained.***

Until recently, there has been no public market for our common stock. Although our common stock is listed on The NASDAQ Global Market, the market for our shares has demonstrated varying levels of trading activity. Furthermore, the current level of trading may not be sustained in the future. The lack of an active market for our common stock may impair investors' ability to sell their shares at the time they wish to sell them or at a price that they consider reasonable, may reduce the fair market value of their shares and may impair our ability to raise capital.

### ***If securities or industry analysts do not publish research reports about our business, or if they issue an adverse opinion about our business, our stock price and trading volume could decline.***

The trading market for our common stock will be influenced by the research and reports that industry or securities analysts publish about us or our business. If one or more of the analysts who cover us issues an adverse opinion about our company, our stock price would likely decline. If one or more of these analysts ceases coverage of us or fails to regularly publish reports on us, we could lose visibility in the financial markets, which in turn could cause our stock price or trading volume to decline.

### ***Future sales of our common stock in the public market could cause our stock price to fall.***

Our stock price could decline as a result of sales of a large number of shares of our common stock after this offering or the perception that these sales could occur. These sales, or the possibility that these sales may occur, also might make it more difficult for us to sell equity securities in the future at a time and at a price that we deem appropriate.

Holders of approximately 7.4 million shares (including shares underlying outstanding warrants described in the section of this prospectus captioned Description of Capital Stock Warrants ), or approximately 50%, of our outstanding shares, have rights, subject to some conditions, to require us to file registration statements covering the sale of their shares or to include their shares in registration statements that we may file for ourselves or other stockholders. We have also registered the offer and sale of all shares of common stock that we may issue under our equity compensation plans.

In addition, in the future, we may issue additional shares of common stock or other equity or debt securities convertible into common stock in connection with a financing, acquisition, litigation settlement, employee arrangements or otherwise. Any such issuance could result in substantial dilution to our existing stockholders and could cause our stock price to decline.

---

**Table of Contents**

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

Our executive officers, directors and principal stockholders, together with their respective affiliates, beneficially owned approximately 58.6% of our outstanding common stock as of December 31, 2013, and we expect that upon completion of this offering, that same group will beneficially own approximately % of our outstanding common stock (assuming no exercise of the underwriters overallotment option). Accordingly, after this offering, our executive officers, directors and principal stockholders will continue to be able to determine the composition of the board of directors, retain the voting power to approve all matters requiring stockholder approval, including mergers and other business combinations, and continue to have significant influence over our operations. This concentration of ownership could have the effect of delaying or preventing a change in our control or otherwise discouraging a potential acquirer from attempting to obtain control of us, which in turn could have a material adverse effect on our stock price and may prevent attempts by our stockholders to replace or remove the board of directors or management.

***Our management team has broad discretion to use the net proceeds from this offering and its investment of these proceeds may not yield a favorable return. We may invest the proceeds of this offering in ways with which investors disagree.***

We have broad discretion as to how to spend and invest the proceeds from this offering, and we may spend or invest these proceeds in a way with which our stockholders disagree. Accordingly, investors will need to rely on our judgment with respect to the use of these proceeds. We intend to use the proceeds from this offering: (1) to further commercialize Prosigna, including establishing a dedicated oncology sales force; (2) to expand the clinical utility of Prosigna and develop other potential diagnostic product opportunities; (3) to expand life sciences commercial operations to grow and support the installed base of our nCounter Analysis Systems among life sciences research customers in the United States and internationally; (4) to develop new life sciences applications, chemistry and instrumentation for our nCounter technology platform; and (5) for working capital and other general corporate purposes. We may also use a portion of the net proceeds to acquire, license and invest in complementary products, technologies or businesses; however, we currently have no agreements or commitments to complete any such transaction. These uses may not yield a favorable return to our stockholders.

We cannot specify with certainty all of the particular uses for the net proceeds to be received upon the closing of this offering. In addition, the amount, allocation and timing of our actual expenditures will depend upon numerous factors, including the revenue generated from the sale of our products. Accordingly, we will have broad discretion in using these proceeds. In addition, until the net proceeds are used, they may be placed in investments that do not produce significant income or that may lose value.

***Anti-takeover provisions in our charter documents and under Delaware or Washington law could make an acquisition of us difficult, limit attempts by our stockholders to replace or remove our current management and limit our stock price.***

Provisions of our certificate of incorporation and bylaws may delay or discourage transactions involving an actual or potential change in our control or change in our management, including transactions in which stockholders might otherwise receive a premium for their shares, or transactions that our stockholders might otherwise deem to be in their best interests. Therefore, these provisions could adversely affect the price of our stock. Among other things, the certificate of incorporation and bylaws:

permit the board of directors to issue up to 15,000,000 shares of preferred stock, with any rights, preferences and privileges as they may designate;

provide that the authorized number of directors may be changed only by resolution of the board of directors;

provide that all vacancies, including newly-created directorships, may, except as otherwise required by law, be filled by the affirmative vote of a majority of directors then in office, even if less than a quorum;

**Table of Contents**

divide the board of directors into three classes;

provide that a director may only be removed from the board of directors by the stockholders for cause;

require that any action to be taken by our stockholders must be effected at a duly called annual or special meeting of stockholders and may not be taken by written consent;

provide that stockholders seeking to present proposals before a meeting of stockholders or to nominate candidates for election as directors at a meeting of stockholders must provide notice in writing in a timely manner, and meet specific requirements as to the form and content of a stockholder's notice;

prevent cumulative voting rights (therefore allowing the holders of a plurality of the shares of common stock entitled to vote in any election of directors to elect all of the directors standing for election, if they should so choose);

provide that special meetings of our stockholders may be called only by the chairman of the board, our chief executive officer or by the board of directors; and

provide that stockholders are permitted to amend the bylaws only upon receiving at least two-thirds of the total votes entitled to be cast by holders of all outstanding shares then entitled to vote generally in the election of directors, voting together as a single class.

In addition, because we are incorporated in Delaware, we are governed by the provisions of Section 203 of the Delaware General Corporation Law, which generally prohibits a Delaware corporation from engaging in any of a broad range of business combinations with any interested stockholder for a period of three years following the date on which the stockholder became an interested stockholder. Likewise, because our principal executive offices are located in Washington, the anti-takeover provisions of the Washington Business Corporation Act may apply to us under certain circumstances now or in the future. These provisions prohibit a target corporation from engaging in any of a broad range of business combinations with any stockholder constituting an acquiring person for a period of five years following the date on which the stockholder became an acquiring person. See the section of this prospectus captioned Description of Capital Stock Anti-Takeover Effects of Delaware and Washington Law and Our Certificate of Incorporation and Bylaws.

***We are an emerging growth company, and any decision on our part to comply only with certain reduced reporting and disclosure requirements applicable to emerging growth companies could make our common stock less attractive to investors.***

We are an emerging growth company, as defined in the Jumpstart Our Business Startups Act, or the JOBS Act, enacted in April 2012, and, for as long as we continue to be an emerging growth company, we may choose to take advantage of exemptions from various reporting requirements applicable to other public companies but not to emerging growth companies, including, but not limited to, not being required to have our independent registered public accounting firm audit our internal control over financial reporting under Section 404, reduced disclosure obligations regarding executive compensation in our periodic reports and proxy statements, and exemptions from the requirements of holding a nonbinding advisory vote on executive compensation and stockholder approval of any golden parachute payments not previously approved. We could be an emerging growth company until December 31, 2018, although, if we have more than \$1.0 billion in annual revenue, if the market value of our common stock that is held by non-affiliates exceeds \$700 million as of June 30 of any year, or we issue more than \$1.0 billion of non-convertible debt over a three-year period before the end of that five-year period, we would cease to be an emerging growth company as of the following December 31. We cannot predict if investors will find our common stock less attractive if we choose to rely on these exemptions. If some investors find our common stock less attractive as a result of any choices to reduce future disclosure, there may be a less active trading market for our common stock and our stock price may be more volatile.

As an emerging growth company the JOBS Act allows us to delay adoption of new or revised accounting pronouncements applicable to public companies until such pronouncements are made applicable to private companies. We have elected to use this extended transition period under the JOBS Act. As a result, our financial



## **Table of Contents**

statements may not be comparable to the financial statements of issuers who are required to comply with the effective dates for new or revised accounting standards that are applicable to public companies, which may make our common stock less attractive to investors.

***Complying with the laws and regulations affecting public companies will increase our costs and the demands on management and could harm our operating results.***

As a public company, and particularly after we cease to be an emerging growth company, we will incur significant legal, accounting and other expenses that we did not incur as a private company. In addition, the Sarbanes-Oxley Act and rules subsequently implemented by the SEC and The NASDAQ Global Market impose numerous requirements on public companies, including requiring changes in corporate governance practices. Also, the Exchange Act requires, among other things, that we file annual, quarterly and current reports with respect to our business and operating results. Our management and other personnel will need to devote a substantial amount of time to compliance with these laws and regulations. These burdens may increase as new legislation is passed and implemented, including any new requirements that the Dodd-Frank Wall Street Reform and Consumer Protection Act of 2010 may impose on public companies. These requirements have increased and will continue to increase our legal, accounting, and financial compliance costs and have made and will continue to make some activities more time consuming and costly. For example, we expect these rules and regulations to make it more difficult and more expensive for us to obtain director and officer liability insurance, and in the future we may be required to accept reduced policy limits and coverage or to incur substantial costs to maintain the same or similar coverage. These rules and regulations could also make it more difficult for us to attract and retain qualified persons to serve on our board of directors or our board committees or as executive officers.

The Sarbanes-Oxley Act requires, among other things, that we assess the effectiveness of our internal control over financial reporting annually and the effectiveness of our disclosure controls and procedures quarterly. In particular, beginning January 1, 2014, Section 404 of the Sarbanes-Oxley Act, or Section 404, requires us to perform system and process evaluation and testing of our internal control over financial reporting to allow management to report on, and our independent registered public accounting firm potentially to attest to, the effectiveness of our internal control over financial reporting. As an emerging growth company, we expect to avail ourselves of the exemption from the requirement that our independent registered public accounting firm attest to the effectiveness of our internal control over financial reporting under Section 404. However, we may no longer avail ourselves of this exemption when we cease to be an emerging growth company. When our independent registered public accounting firm is required to undertake an assessment of our internal control over financial reporting, the cost of our compliance with Section 404 will correspondingly increase. Our compliance with applicable provisions of Section 404 will require that we incur substantial accounting expense and expend significant management time on compliance-related issues as we implement additional corporate governance practices and comply with reporting requirements. Moreover, if we are not able to comply with the requirements of Section 404 applicable to us in a timely manner, or if we or our independent registered public accounting firm identifies deficiencies in our internal control over financial reporting that are deemed to be material weaknesses, the market price of our stock could decline and we could be subject to sanctions or investigations by the SEC or other regulatory authorities, which would require additional financial and management resources.

Furthermore, investor perceptions of our company may suffer if deficiencies are found, and this could cause a decline in the market price of our stock. Irrespective of compliance with Section 404, any failure of our internal control over financial reporting could have a material adverse effect on our stated operating results and harm our reputation. If we are unable to implement these requirements effectively or efficiently, it could harm our operations, financial reporting, or financial results and could result in an adverse opinion on our internal control over financial reporting from our independent registered public accounting firm.

**Table of Contents**

**SPECIAL NOTE REGARDING FORWARD-LOOKING STATEMENTS**

This prospectus contains forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995 that are based on management's beliefs and assumptions and on information currently available to management. Some of the statements under Prospectus Summary, Risk Factors, Management's Discussion and Analysis of Financial Condition and Results of Operations and Business and elsewhere in this prospectus contain forward-looking statements. In some cases, you can identify forward-looking statements by the following words: may, will, could, would, should, expect, intend, plan, anticipate, believe, estimate, predict, project, potential, continue, and other similar terms or these terms or other comparable terminology, although not all forward-looking statements contain these words.

These statements involve risks, uncertainties and other factors that may cause actual results, levels of activity, performance or achievements to be materially different from the information expressed or implied by these forward-looking statements. Although we believe that we have a reasonable basis for each forward-looking statement contained in this prospectus, we caution you that these statements are based on a combination of facts and factors currently known by us and our projections of the future, about which we cannot be certain. Forward-looking statements in this prospectus include, but are not limited to, statements about:

our expectations regarding our 2013 operating results, including our expectations regarding instrument, consumable and total revenue and operating and net loss;

our ability to successfully commercialize Prosigna, our first product for which we have obtained a CE mark in the European Union and, in September 2013, received 510(k) clearance from the U.S. Food and Drug Administration, or FDA;

the implementation of our business model and strategic plans for our business;

the regulatory regime and our ability to secure regulatory clearance or approval for the clinical use of our products, domestically and internationally;

our strategic relationships, including with patent holders of our technologies, manufacturers and distributors of our products, and third parties who conduct our clinical studies;

our intellectual property position;

our expected use of proceeds;

our expectations regarding the market size and growth potential for our life sciences and diagnostic businesses;

any estimates regarding expenses, future revenues, capital requirements, and stock performance; and

our ability to sustain and manage growth, including our ability to develop new products and enter new markets.

In addition, you should refer to the Risk Factors section of this prospectus for a discussion of other important factors that may cause actual results to differ materially from those expressed or implied by the forward-looking statements. As a result of these factors, we cannot assure you that the forward-looking statements in this prospectus will prove to be accurate. Furthermore, if the forward-looking statements prove to be inaccurate, the inaccuracy may be material. In light of the significant uncertainties in these forward-looking statements, you should not regard

## Edgar Filing: NanoString Technologies Inc - Form S-1

these statements as a representation or warranty by us or any other person that we will achieve our objectives and plans in any specified time frame, or at all. We undertake no obligation to publicly update any forward-looking statements, whether as a result of new information, future events or otherwise, except as required by law.

This prospectus contains market data and industry forecasts that were obtained from industry publications. These data involve a number of assumptions and limitations, and you are cautioned not to give undue weight to such estimates. We have not independently verified any third-party information. While we believe the market position, market opportunity and market size information included in this prospectus is generally reliable, such information is inherently imprecise.

**Table of Contents**

**USE OF PROCEEDS**

The net proceeds to us from the sale of shares of our common stock that we are selling in this offering will be approximately \$            million, based upon an assumed public offering price of \$            per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on           , 2014, after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us. If the underwriters' overallotment option is exercised in full, our net proceeds would be approximately \$            million, after deducting estimated underwriting discounts and commissions and estimated offering expenses payable by us.

We currently expect to use the net proceeds from this offering: (1) to further commercialize Prosigna, including establishing a dedicated oncology sales force; (2) to expand the clinical utility of Prosigna and develop other potential diagnostic product opportunities; (3) to expand life sciences commercial operations to grow and support the installed base of our nCounter Analysis Systems among life sciences research customers in the United States and internationally; (4) to develop new life sciences applications, chemistry and instrumentation for our nCounter technology platform; and (5) for working capital and other general corporate purposes.

We may also use a portion of the net proceeds to acquire, license and invest in complementary products, technologies or businesses; however, we currently have no agreements or commitments to complete any such transaction.

We cannot specify with certainty all of the particular uses for the net proceeds to be received upon the closing of this offering. In addition, the amount, allocation and timing of our actual expenditures will depend upon numerous factors, including the revenue generated from the sale of our products to life sciences customers and the sale of Prosigna. Accordingly, we will have broad discretion in using these proceeds. Pending their uses, we plan to invest the net proceeds of this offering in short-term, interest-bearing, investment-grade instruments, certificates of deposit or direct or guaranteed obligations of the U.S. government.



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

Our common stock has been listed on The NASDAQ Global Market under the symbol `NSTG` since June 26, 2013. Prior to that date, there was no public trading market for our common stock. The following table sets forth for the periods indicated the high and low sales prices per share of our common stock as reported on The NASDAQ Global Market:

	<b>High</b>	<b>Low</b>
<b>Year Ended December 31, 2013:</b>		
Second Quarter (from June 26, 2013)	\$ 9.90	\$ 7.81
Third Quarter	14.10	7.01
Fourth Quarter	18.09	8.64
<b>Year Ended December 31, 2014:</b>		
First Quarter (through January 10, 2014)	\$ 20.79	\$ 16.28

On January 10, 2014, the last reported sale price of our common stock on The NASDAQ Global Market was \$19.51 per share. As of December 31, 2013, we had approximately 92 holders of record of our common stock. The actual number of stockholders is greater than this number of record holders and includes stockholders who are beneficial owners but whose shares are held in street name by brokers and other nominees. This number of holders of record also does not include stockholders whose shares may be held in trust by other entities.

**Table of Contents**

**DIVIDEND POLICY**

We have never declared or paid any cash dividends on our common stock or any other securities. We anticipate that we will retain all available funds and any future earnings, if any, for use in the operation of our business and do not anticipate paying cash dividends in the foreseeable future. In addition, our credit facility materially restricts, and future debt instruments we issue may materially restrict, our ability to pay dividends on our common stock. Payment of future cash dividends, if any, will be at the discretion of the board of directors after taking into account various factors, including our financial condition, operating results, current and anticipated cash needs, the requirements of current or then-existing debt instruments and other factors the board of directors deems relevant.

**Table of Contents****CAPITALIZATION**

The following table summarizes our capitalization as of September 30, 2013:

on an actual basis; and

on an as adjusted basis, giving effect to the sale and issuance by us of \_\_\_\_\_ shares of common stock in this offering, at an assumed public offering price of \$ \_\_\_\_\_ per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on \_\_\_\_\_, 2014, after deducting underwriting discounts and commissions and estimated offering expenses payable by us.

Investors should read the information in this table together with the financial statements and related notes to those statements, as well as the sections of this prospectus captioned "Selected Consolidated Financial Data" and "Management's Discussion and Analysis of Financial Condition and Results of Operations."

	As of September 30, 2013	
	Actual	As Adjusted
	(In thousands, except per share amounts)	
Total long-term debt (including current portion)	\$ 18,213	\$
Stockholders' equity (deficit):		
Preferred stock, \$0.0001 par value per share; 15,000 shares authorized, no shares issued or outstanding, actual and as adjusted		
Common stock, \$0.0001 par value per share, 150,000 shares authorized, 14,616 shares issued and outstanding, actual; shares issued and outstanding, as adjusted	1	
Additional paid-in capital	157,890	
Other comprehensive income	5	
Accumulated deficit	(118,005)	
Total stockholders' equity	39,891	
Total capitalization	\$ 58,104	\$

The number of shares of common stock to be outstanding following this offering is based on 14,616,871 shares of common stock outstanding as of September 30, 2013, and excludes:

1,873,491 shares of common stock issuable upon exercise of options outstanding as of September 30, 2013, at a weighted-average exercise price of \$3.41 per share;

1,891,069 shares of common stock reserved for future issuance under stock-based compensation plans, including 1,609,819 shares of common stock reserved for issuance under our 2013 Equity Incentive Plan, and any future automatic increase in shares reserved for issuance under such plan, and 281,250 shares of common stock reserved for issuance under our 2013 Employee Stock Purchase Plan, and any future automatic increase in shares reserved for issuance under such plan; and

617,605 shares of common stock issuable upon the exercise of warrants outstanding as of September 30, 2013, at a weighted-average exercise price of \$8.78 per share.



**Table of Contents**

**DILUTION**

Purchasers of common stock offered by this prospectus will suffer immediate and substantial dilution in the net tangible book value per share of common stock. Our net tangible book value as of September 30, 2013 was approximately \$39.9 million, or approximately \$2.73 per share of common stock. Net tangible book value per share represents the amount of total tangible assets less total liabilities, divided by the number of shares of our common stock outstanding as of September 30, 2013.

Dilution in net tangible book value per share represents the difference between the amount per share paid by purchasers in this offering and the net tangible book value per share of our common stock immediately after this offering. After giving effect to an assumed sale of shares of common stock in this offering at an assumed public offering price of \$ \_\_\_\_\_ per share, which was the last reported sales price of our common stock on The NASDAQ Global Market on January \_\_\_\_\_, 2014, and after deducting the underwriting discounts and commissions and estimated offering expenses payable by us, our as adjusted net tangible book value as of September 30, 2013 would have been approximately \$ \_\_\_\_\_, or \$ \_\_\_\_\_ per share of common stock. This represents an immediate increase in net tangible book value of \$ \_\_\_\_\_ per share of common stock to our existing stockholders and an immediate dilution in net tangible book value of \$ \_\_\_\_\_ per share of common stock to investors participating in this offering.

The following table illustrates this dilution on a per share basis to new investors:

Assumed public offering price per share	\$
Historical net tangible book value per share as of September 30, 2013	\$ 2.73
Increase attributable to this offering	

As adjusted net tangible book value per share as of September 30, 2013 after giving effect to this offering	
Dilution per share to investors participating in this offering	\$

Each \$1.00 increase or decrease in the assumed public offering price of \$ \_\_\_\_\_ per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on January \_\_\_\_\_, 2014, would increase or decrease, as applicable, our pro forma net tangible book value per share to new investors by \$ \_\_\_\_\_, and would increase or decrease, as applicable, dilution per share to new investors in this offering by \$ \_\_\_\_\_, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting estimated underwriting discounts and commissions payable by us. In addition, to the extent any outstanding options or warrants to purchase common stock are exercised, new investors would experience further dilution. If the underwriters exercise their over-allotment option in full, the pro forma net tangible book value per share of our common stock immediately after this offering would be \$ \_\_\_\_\_ per share, and the dilution in pro forma net tangible book value per share to new investors in this offering would be \$ \_\_\_\_\_ per share.

The following table summarizes, on a pro forma basis as of September 30, 2013, the differences between the existing stockholders and the new investors purchasing shares of our common stock in this offering with respect to the number of shares purchased from us, the total consideration paid or to be paid to us, which includes net proceeds received from the issuance of common stock, cash received from the exercise of stock options, and the average price per share paid or to be paid to us at the public offering price of \$ \_\_\_\_\_ per share, before deducting underwriting discounts and commissions and estimated offering expenses payable by us (in thousands, except per share amounts):

	Shares Purchased		Total Consideration		Weighted Average Price Per Share
	Number	Percent	Amount	Percent	
Existing stockholders		%	\$	%	\$
New investors					
<b>Total</b>		<b>100.0%</b>	<b>\$</b>	<b>100.0%</b>	



**Table of Contents**

Each \$1.00 increase or decrease in the assumed public offering price of \$ \_\_\_\_\_ per share, which was the last reported sale price of our common stock on The NASDAQ Global Market on January \_\_\_\_\_, 2014, would increase or decrease, as applicable, the total consideration paid by new investors and total consideration paid by all stockholders by approximately \$ \_\_\_\_\_, assuming that the number of shares offered by us, as set forth on the cover page of this prospectus, remains the same and after deducting estimated underwriting discounts and commissions payable by us. In addition, to the extent any outstanding options or warrants to purchase common stock are exercised, new investors will experience further dilution

The number of shares of common stock to be outstanding following this offering is based on 14,616,871 shares of common stock outstanding as of September 30, 2013, and excludes:

1,873,491 shares of common stock issuable upon exercise of options outstanding as of September 30, 2013, at a weighted-average exercise price of \$3.41 per share;

1,891,069 shares of common stock reserved for future issuance under stock-based compensation plans, including 1,609,819 shares of common stock reserved for issuance under the 2013 Equity Incentive Plan, and any future automatic increase in shares reserved for issuance under such plan, and 281,250 shares of common stock reserved for issuance under the 2013 Employee Stock Purchase Plan, and any future automatic increase in shares reserved for issuance under such plan; and

617,605 shares of common stock issuable upon the exercise of warrants outstanding as of September 30, 2013, at a weighted-average exercise price of \$8.78 per share.

Share reserves for the equity incentive plans will also be subject to automatic annual increases in accordance with the terms of the plans. To the extent that new options are issued under the equity benefit plans or we issue additional shares of common stock in the future, there will be further dilution to investors participating in this offering.

**Table of Contents****SELECTED CONSOLIDATED FINANCIAL DATA**

The following selected statement of operations data for the years ended December 31, 2010, 2011 and 2012 and the balance sheet data as of December 31, 2011 and 2012 have been derived from audited consolidated financial statements included elsewhere in this prospectus. The selected statements of operations for the nine months ended September 30, 2012 and 2013 and the balance sheet data as of September 30, 2013 have been derived from unaudited interim financial statements included elsewhere in this prospectus. The selected statements of operations for the years ended December 31, 2008 and 2009 and the balance sheet data as of December 31, 2008, 2009 and 2010 have been derived from audited consolidated financial statements which are not included in this prospectus. In the opinion of management, the unaudited interim financial statements reflect all adjustments, which include only normal recurring adjustments, necessary for a fair presentation of the financial statements. Historical results are not necessarily indicative of the results that may be expected in the future and the results for the nine months ended September 30, 2013 are not necessarily indicative of results to be expected for the full year or any other period. You should read the following selected financial and other data below in conjunction with the financial statements and related notes included elsewhere in this prospectus and the sections of this prospectus captioned Management's Discussion and Analysis of Financial Condition and Results of Operations.

	2008	Year Ended December 31,			Nine Months		
	2008	2009	2010	2011	2012	Ended September 30,	
						2012	
						2013	
	(In thousands, except per share amounts)						
<b>Consolidated Statements of Operations:</b>							
<b>Revenue</b>	\$ 1,613	\$ 7,288	\$ 11,730	\$ 17,800	\$ 22,973	\$ 16,480	\$ 21,283
<b>Costs and expenses:</b>							
Cost of revenue	1,450	5,874	9,128	9,777	12,361	9,076	10,188
Research and development	4,428	4,550	7,547	8,990	11,635	8,253	10,469
Selling, general and administrative	4,513	5,464	8,027	9,529	15,486	10,588	20,822
Total costs and expenses	10,391	15,888	24,702	28,296	39,482	27,917	41,479
Loss from operations	(8,778)	(8,600)	(12,972)	(10,496)	(16,509)	(11,437)	(20,196)
<b>Other income (expense):</b>							
Interest income	51	64	29	10	21	17	28
Interest expense	(193)	(320)	(94)	(599)	(804)	(551)	(1,412)
Other income (expense)			254	80	(29)	(26)	(30)
Revaluation of preferred stock warrant liability	35	19	15	73	(387)	150	1,156
Total other income (expense)	(107)	(237)	204	(436)	(1,199)	(410)	(258)
Net loss	(8,885)	(8,837)	(12,768)	(10,932)	(17,708)	(11,847)	(20,454)
Accretion of mandatorily redeemable convertible preferred stock	(1,708)	(2,551)	(4,351)	(5,251)	(7,533)	(5,515)	(4,653)
Net loss attributable to common stockholders	\$ (10,593)	\$ (11,388)	\$ (17,119)	\$ (16,183)	\$ (25,241)	\$ (17,362)	\$ (25,107)
Net loss per share - basic and diluted	\$ (34.50)	\$ (36.62)	\$ (54.17)	\$ (50.10)	\$ (71.10)	\$ (51.06)	\$ (4.74)
Weighted-average shares used in computing basic and diluted net loss per share	307	311	316	323	355	340	5,292



**Table of Contents**

	As of December 31,					As of
	2008	2009	2010	2011	2012	September 30, 2013
	(In thousands)					
<b>Consolidated Balance Sheet Data:</b>						
Cash, cash equivalents and short-term investments	\$ 3,508	\$ 1,739	\$ 4,366	\$ 10,868	\$ 21,692	\$ 52,214
Working capital	(4,224)	1,385	2,944	12,236	19,937	52,054
Total assets	9,564	9,367	13,275	24,584	37,406	71,293
Total long-term debt	8,878	1,274	1,829	1,887	12,759	18,213
Mandatorily redeemable convertible preferred stock	21,276	38,551	57,887	80,957	103,622	
Total stockholders' equity (deficit)	(25,194)	(36,565)	(53,517)	(69,451)	(93,760)	39,891

-44-

---

**Table of Contents**

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

*You should read the following discussion and analysis together with the financial statements and the related notes to those statements included elsewhere in this prospectus. This discussion contains forward-looking statements that involve risks and uncertainties. As a result of many factors, such as those set forth in the section of the prospectus captioned "Risk Factors" and elsewhere in this prospectus, our actual results may differ materially from those anticipated in these forward-looking statements.*

**Overview**

We develop, manufacture and sell robust, intuitive products that unlock scientifically valuable and clinically actionable genomic information from minute amounts of tissue. Our nCounter Analysis System directly profiles hundreds of molecules simultaneously using a novel barcoding technology that is powerful enough for use in research, yet simple enough for use in clinical laboratories worldwide. We market systems and related consumables to researchers in academic, government, and biopharmaceutical laboratories for use in understanding fundamental biology and the molecular basis of disease and to clinical laboratories and medical centers for diagnostic use. We have an installed base of more than 180 systems, which our customers have used to publish more than 360 peer-reviewed papers. As researchers discover how genomic information can be used to improve clinical decision-making, these discoveries can be translated and validated as diagnostic tests based on our nCounter Elements reagents. In certain situations, we intend to translate their discoveries into *in vitro* diagnostic assays. In September 2013, we received 510(k) clearance from the FDA to market in the United States a version of Prosigna providing an assessment of a patient's risk of recurrence for breast cancer. In November 2013, we commercially launched the nCounter Dx Analysis System in the United States. In December 2013, we commercially launched Prosigna in the United States and announced that national diagnostic laboratories ARUP Laboratories, Laboratory Corporation of America Holdings and Quest Diagnostics have chosen to add Prosigna to their suites of breast cancer diagnostic tests. We expect Prosigna testing services to become available in the United States in the first quarter of 2014. In September 2012, we received European Union regulatory clearance for our first molecular diagnostic product, the Prosigna Breast Cancer Assay, or Prosigna, an assay providing an assessment of a patient's risk of recurrence for breast cancer and the intrinsic subtype of the patient's tumor. In February 2013, we commercially launched Prosigna in Europe and Israel.

We derive a substantial majority of our revenue from the sale of our products, which consist of our nCounter instruments and related proprietary consumables, which we call CodeSets, nCounter Elements reagents and Master Kits. We sell two types of CodeSets: custom orders and standard sets, which we call panels. We also derive revenue from processing fees related to proof-of-principle studies we conduct for potential customers and extended service contracts for our nCounter Analysis Systems.

Until recently, we have sold our products for research use only. After buying an nCounter Analysis System, life sciences customers purchase consumables from us for use in their experiments. Our instruments are designed to work only with our consumable products. Accordingly, as the installed base of our instruments grows, we expect recurring revenue from consumable sales to become an increasingly important driver of our operating results.

We have begun to offer instruments and consumables for use in diagnostic testing. In September 2012, we obtained a CE mark for Prosigna, our first diagnostic product, and, in early 2013 we commercially launched Prosigna in Europe, including in France, Germany, Greece, Italy, Spain, Turkey and the United Kingdom, and Israel. In September 2013, we received 510(k) clearance from the FDA to market in the United States a version of Prosigna providing an assessment of a patient's risk of recurrence for breast cancer. In November 2013, we commercially launched the nCounter Dx Analysis System in the United States, including a dual-mode configuration that can be used both for Prosigna and for research applications. In December 2013, we commercially launched Prosigna in the United States. National diagnostic laboratories ARUP Laboratories, Laboratory Corporation of America Holdings and Quest Diagnostics have chosen to add Prosigna to their suites of breast cancer diagnostic tests, and the laboratories at the University of Alabama at Birmingham Comprehensive Cancer Center and

## **Table of Contents**

University of North Carolina Lineberger Comprehensive Cancer Center will be among the initial facilities to offer the Prosigna assay in the United States, with the earliest testing beginning during the first quarter of 2014. These laboratories collectively serve the pathology testing needs of a substantial portion of breast cancer patients throughout the United States.

In November 2013, we began offering a version of the nCounter Dx Analysis System to high-complexity, CLIA-certified laboratories for research and diagnostics purposes. This FLEX configuration of the nCounter Dx Analysis System provides clinical laboratories a single platform with the flexibility to support both clinical testing, by running Prosigna, and research, by processing translational research experiments using our custom CodeSets and panels. The nCounter Elements GPRs provide further flexibility by allowing laboratories to develop their own Laboratory Developed Tests for gene expression, copy number variation and gene fusion signatures, which can be performed by a laboratory and may include genetic tests and other tests for rare conditions.

To support the commercial launch of Prosigna, we are establishing a dedicated oncology diagnostics sales force. As a result, we expect sales and marketing expenses and operating losses to increase as we market the product. In addition, we expect sales to grow gradually as more systems are installed, Prosigna gains inclusion in important breast cancer treatment guidelines and to the extent reimbursement by third-party payors becomes more broadly available.

We use third-party contract manufacturers to produce the two instruments comprising the nCounter Analysis System. We manufacture consumables at our Seattle, Washington facility. This operating model is designed to be capital efficient and to scale efficiently as our product volumes grow. We focus a substantial portion of our resources on developing new products and solutions. We invested \$7.5 million, \$9.0 million and \$11.6 million in 2010, 2011 and 2012, respectively, and \$8.3 million and \$10.5 million in the nine months ended September 30, 2012 and 2013, respectively, in research and development and intend to continue to make significant investments in research and development.

Our total revenue increased to \$23.0 million in 2012 from \$17.8 million in 2011 and \$11.7 million in 2010, and to \$21.3 million for the nine months ended September 30, 2013 from \$16.5 million for the nine months ended September 30, 2012, which was driven by the sale of additional nCounter Analysis Systems and consumables for use on our growing installed base of instruments. Historically, we have generated a substantial majority of our revenue from sales to customers in North America; however, we expect sales in other regions to increase over time. We have never been profitable and had net losses of \$12.8 million, \$10.9 million, \$17.7 million, \$20.5 million and \$11.8 million in 2010, 2011 and 2012, and for the nine months ended September 30, 2013 and 2012, respectively. As of September 30, 2013, our accumulated deficit was \$118.0 million.

## **Key Financial Metrics**

We are organized as, and operate in, two reportable segments: our life sciences business and our diagnostics business. Our life sciences business provides instruments, consumables and services for efficiently profiling the activity of hundreds of genes simultaneously from a single tissue sample. Our diagnostics business provides molecular diagnostic kits to pathology labs enabling complex molecular testing on a decentralized basis.

Our chief operating decision maker is the chief executive officer. The chief operating decision maker reviews financial information presented on a total company basis, accompanied by information about segment revenue and certain direct sales and marketing expenses by segment. Our chief operating decision maker evaluates performance based on these two measures. The chief operating decision maker does not review segment information related to cost of revenue, research and development or other selling, general and administrative expenses.

## **Revenue**

We generate revenue from the sale of our products and related services. We are organized as, and operate in, two reportable segments: life sciences and diagnostics. For a description of our revenue recognition policies, see the section of this prospectus captioned "Critical Accounting Policies and Significant Estimates Revenue Recognition."

## **Table of Contents**

### *Product Revenue*

Our products consist of our nCounter Analysis System and related consumables. Our nCounter Analysis System typically consists of one nCounter Digital Analyzer and one nCounter Prep Station. The U.S. list price of one research use only nCounter Analysis System is \$235,000. Outside the United States, depending on the country, the list price is generally higher. Systems are sold to distributors at a discount to list price. For our life sciences customers, related consumables include (1) panels, which are standard pre-manufactured CodeSets, (2) custom CodeSets, which we manufacture to the specific requirements of an individual researcher, (3) nCounter Elements reagents, and (4) Master Kits, which are ancillary reagents, cartridges, tips and reagent plates required to setup and process samples in our instruments. Product revenue also includes payments for instrument installation. Currently, our customer base is primarily composed of academic institutions, government laboratories, and biopharmaceutical companies that perform analyses using our nCounter Analysis System and purchase consumables for research use only. In 2010, 2011, 2012 and 2013, our average life sciences consumable revenue per system exceeded \$100,000 per year.

For our diagnostics customers, we sell our nCounter Dx Analysis Systems to customers or offer to lease them under reagent rental arrangements where an instrument is placed at a customer location at minimal direct cost and the customer commits to purchase a minimum volume of consumable diagnostic kits or nCounter Elements reagents over a period of time. Beginning in November 2013, we have offered the FLEX configuration of our nCounter Dx Analysis System that can be used for Prosigna and research applications. The list price for our nCounter Dx Analysis System is higher than the research use only version. To date, all diagnostics customers have elected to purchase instruments; however, we expect that in the future, certain customers will elect to lease them. The revenue derived from the sale of diagnostic kits and nCounter Elements reagents will be driven by a combination of the number of tests performed by our customers as well as the price of each kit. The list price of a Prosigna test in the United States and Europe is \$2,080 and 1,550 per patient, respectively. Although the price of Prosigna and our additional future diagnostic products will depend on many factors, including whether and how much third-party payors will reimburse laboratories for conducting such tests, we expect that the gross margin for our diagnostic kits will be higher than for our life sciences research consumables. We plan to sell Prosigna kits to our lab customers, who will be responsible for providing the testing service and contracting and billing payors. We also plan to sell Prosigna kits to clinical laboratories on a fixed dollars-per-kit basis, which would not expose us to direct third-party payor reimbursement risk. However, we anticipate providing customary volume discounts, and in some cases, introductory pricing during the period in which third-party payor reimbursement is being established. As a result, we expect the average selling price per Prosigna test to be between \$1,500 and \$2,000 in the United States.

### *Service Revenue*

Service revenue consists of fees associated with extended service contracts and conducting proof-of-principle studies. We include a one-year warranty with the sale of our instruments and offer extended service contracts, which are purchased by a majority of our customers. We selectively provide proof-of-principle studies to prospective customers in order to help them better understand the benefits of the nCounter Analysis System.

### *Revenue by Geography*

We sell our life sciences products through our own sales force in the United States, Canada, Singapore and certain European countries. We sell through distributors in other parts of the world. As we have expanded our European life sciences direct sales force and entered into agreements with distributors of our life sciences products in Europe, the Middle East, Asia Pacific and South America, the amount of revenue generated outside of North America has generally increased, although there have been significant quarter-to-quarter fluctuations. In the future, we intend to expand our sales force and establish additional distributor relationships outside the United States to better access international markets.

**Table of Contents**

The following table reflects product revenue by geography and as a percentage of total product revenue, based on the billing address of our customers. North America consists of the United States, Canada and Mexico; and Asia Pacific includes Japan, China, South Korea, Singapore, Malaysia and Australia.

	Year Ended December 31,						Nine Months Ended September 30,			
	2010		2011		2012		2012		2013	
	(Dollars in thousands)									
North America	\$ 10,643	91%	\$ 14,044	79%	\$ 15,906	69%	\$ 11,809	72%	\$ 15,028	71%
Europe & Middle East	909	8	2,918	16	4,167	18	2,863	17	3,702	17
Asia Pacific	178	1	838	5	2,900	13	1,808	11	2,553	12
Total	\$ 11,730	100%	\$ 17,800	100%	\$ 22,973	100%	\$ 16,480	100%	\$ 21,283	100%

Most of our revenue is denominated in U.S. dollars. Our expenses are generally denominated in the currencies in which our operations are located, which is primarily in the United States. Changes in foreign currency exchange rates have not materially affected us to date; however, they may become material to us in the future as our operations outside of the United States expand.

**Cost of Revenue**

Cost of revenue consists primarily of costs incurred in the production process, including costs of purchasing instruments from third-party contract manufacturers, consumable component materials and assembly labor and overhead, installation, warranty, service and packaging and delivery costs. In addition, cost of revenue includes royalty costs for licensed technologies included in our products, provisions for slow-moving and obsolete inventory and stock-based compensation expense. We provide a one-year warranty on each nCounter Analysis System sold and establish a reserve for warranty repairs based on historical warranty repair costs incurred.

We expect the average unit costs of our instruments to decline in future periods as a result of our ongoing efforts to develop a lower-cost nCounter Analysis System to expand our market opportunity among smaller laboratories. We expect the unit costs of consumable products to decline as a result of our ongoing efforts to improve our manufacturing processes and expected increases in production volume and yields. Although the unit costs of our custom CodeSets vary, they are generally higher as a percentage of the related revenue than our panels, *in vitro* diagnostic kits and nCounter Elements GPRs.

**Operating Expenses****Research and Development**

Research and development expenses consist primarily of salaries and benefits, occupancy, laboratory supplies, consulting fees and related costs, costs associated with licensing molecular diagnostics rights and clinical study expenses (including the cost of tissue samples) to support the regulatory approval or clearance of diagnostic products. We have made substantial investments in research and development since our inception. Our research and development efforts have focused primarily on the tasks required to enhance our technologies and to support development and commercialization of new and existing products and applications for both our life sciences and diagnostics businesses. We believe that our continued investment in research and development is essential to our long-term competitive position and expect these expenses to increase in future periods.

Given the relatively small size of our research and development staff and the limited number of active projects at any given time, we have found that, to date, it has been effective for us to manage our research and development activities on a departmental basis. Accordingly, we do not require employees to report their time by project nor do we allocate our research and development costs to individual projects. The following table shows the composition of total research and development expense by functional area for the periods indicated. Prior to

**Table of Contents**

2012, research and development expense related to our core nCounter platform technology and diagnostic product development were combined.

	Year Ended December 31,			Nine Months Ended	
	2010	2011	2012	September 30, 2012	2013
	(In thousands)				
Core nCounter platform technology and diagnostic product development	\$ 3,649	\$ 4,359	\$	\$	\$
Core nCounter platform technology			1,537	997	2,639
Manufacturing process development	878	969	1,183	932	1,102
Life sciences products and applications	1,848	1,875	2,183	1,543	2,135
Diagnostic product development			4,783	3,326	3,384
Facility allocation	1,173	1,787	1,949	1,455	1,209
<b>Total</b>	<b>\$ 7,548</b>	<b>\$ 8,990</b>	<b>\$ 11,635</b>	<b>\$ 8,253</b>	<b>\$ 10,469</b>

Our clinical studies employ a retrospective / prospective design, which means that we use samples that were previously collected from patients and for which the treatment regimen and ultimate patient outcome is known. Such studies are capital efficient as they do not require recruiting new patients and running prospective trials and they can be completed much more quickly than typical prospective clinical trials. We intend to use a similar approach whenever possible for the additional clinical studies we intend to conduct in support of our future regulatory submissions to expand the indications for Prosigna and for future diagnostic products.

We expect to license additional molecular diagnostic rights as part of our strategy to develop additional diagnostic products. For example, in February 2013 we secured an option from a customer to acquire an exclusive worldwide license for a gene signature that could be used, after further development, as a Laboratory Developed Test, or, after appropriate regulatory authorization, for a second molecular diagnostic product to identify patients with cirrhosis who are at highest risk of developing the most common type of liver cancer, HCC, and to determine whether a patient who has been diagnosed with HCC is likely to have a recurrence. The related option fee was expensed in the first quarter of 2013. Such arrangements may include upfront, milestone or annual cash payments and revenue-based royalties. We believe that our continued investment in research and development is essential to our long-term competitive position and expect these expenses to increase in future periods.

*Selling, General and Administrative*

Selling, general and administrative expenses consist primarily of costs for our sales and marketing, finance, human resources, information technology, business development and general management functions, as well as professional services, such as legal, consulting and accounting services. We expect selling, general and administrative expenses to increase in future periods as the number of sales, technical support and marketing and administrative personnel grows and we continue to introduce new products, broaden our customer base and grow our business. In particular, the continued commercialization of Prosigna requires us to establish a dedicated oncology diagnostics sales force which will increase selling and marketing expenses significantly. Our legal, accounting and compliance costs have also increased as a result of our becoming a public company, and we expect them to continue to increase.

**Factors Affecting Our Performance***Instrument Installed Base*

Our future financial performance will be driven in large part by the rate of sales of our nCounter Analysis Systems, which typically consist of one nCounter Digital Analyzer and one nCounter Prep Station. In some cases, our customers increase the throughput of their nCounter Analysis System by purchasing up to three nCounter Prep Stations per nCounter Digital Analyzer. We plan to grow our system sales in the coming years.

## **Table of Contents**

through multiple strategies, including expanding our sales efforts outside of the United States and continuing to enhance the underlying technology and applications for both life sciences research and diagnostics use. As part of this strategy, we have increased our life sciences sales and marketing headcount by over 30% in 2013 in an effort to increase the rate of sales of our nCounter Analysis Systems. Similarly, since January 2013, we have contracted with nine additional distributors bringing our total to 15. As our installed base of instruments grows, we solicit feedback from our customers and focus our research and development efforts on enabling the nCounter Analysis System for additional applications, which in turn helps to drive additional sales of our instruments and consumables. We are developing a new generation of the nCounter Analysis System that we believe will increase our addressable market and simplify the procurement processes of our potential customers. The new generation system will be a single instrument with a reduced footprint that combines the prep station and the digital analyzer. We plan to reduce the cost of the new generation system through the adoption of new, less expensive technologies. We are targeting release of the new generation system in 2014.

Our sales process involves numerous interactions with multiple individuals within an organization, and often includes in-depth analysis by potential customers of our products, performance of proof-of-principle studies, preparation of extensive documentation and a lengthy review process. As a result of these factors, the large capital investment required in purchasing our instruments and the budget cycles of our customers, the time from initial contact with a customer to our receipt of a purchase order can vary significantly and be up to 12 months or longer. Given the length and uncertainty of our sales cycle, we have in the past experienced, and likely will in the future experience, fluctuations in our instrument sales on a period-to-period basis. We are developing an nCounter Analysis System that we intend to offer at a lower price, which we believe will simplify the procurement processes of our potential customers as well as increase our addressable market.

We have sold more than 180 nCounter Analysis Systems, which we count based on the number of nCounter Digital Analyzers sold given that a system may couple an analyzer with multiple nCounter Prep Stations. Management focuses on instrument unit sales as a primary indicator of current business success and a leading indicator of likely future sales of consumables.

### ***Recurring Consumable Revenue***

Our instruments are designed to be used only with our consumables. This closed system model generates recurring revenue from each instrument we sell. Management focuses on recurring consumable revenue per system as an indicator of the continuing value generated by each system. We calculate recurring consumable revenue per system quarterly by dividing consumable revenue recognized in a particular quarter (other than consumable revenue related to proof-of-principle studies) by the total number of nCounter Analysis Systems installed as of the last day in the immediately preceding quarter. We believe that our recurring consumable revenue is driven by our customers' ability to extract value from up to 800 data points per sample and to process hundreds of samples in a relatively short period of time with little hands-on preparation using our nCounter Analysis System, enabling them to process more units of consumables per unit of time. In 2010, 2011, 2012 and 2013, our average consumable revenue per system exceeded \$100,000 per year.

As the installed base of the nCounter Analysis Systems expands, consumables revenue is expected to increase and over time should be an increasingly important contributor to our total revenue. Over time, we believe that consumables revenue should be subject to less period-to-period fluctuation than our instrument sales revenue.

### ***Revenue Mix and Gross Margin***

Our product revenue is derived from sales of the nCounter Analysis System and related consumables. Generally, our consumables have higher gross margins than our instruments. There will be fluctuations in mix between instruments and consumables from period to period. Although results may vary period to period, over time, as our installed base of systems grows, consumables should constitute a larger percentage of total revenue, which would increase our gross margins. In addition, we expect both the average selling price and the manufacturing cost of our instruments to decrease following the introduction of future generations of our

**Table of Contents**

nCounter Analysis System. Future instrument selling prices and gross margins may fluctuate as we introduce new products and reduce our product costs and from variability in the timing of new product introductions.

We derive service revenue from extended service contracts, which are purchased by a majority of our customers. Additionally, we selectively provide proof-of-principle studies in connection with prospective sales to customers to demonstrate the performance of our nCounter Analysis System.

The following table reflects instrument, consumable and service revenue in absolute dollars and as percentage of total revenue.

	Year Ended December 31,						Nine Months Ended September 30,			
	2010		2011		2012		2012		2013	
	(Dollars in thousands)									
Product revenue:										
Instruments	\$ 6,472	55%	\$ 7,112	40%	\$ 8,786	38%	\$ 6,262	38%	\$ 7,714	36%
Consumables	5,034	43	9,997	56	13,036	57	9,325	57	12,421	58
Service revenue	224	2	691	4	1,151	5	893	5	1,148	6
Total	\$ 11,730	100%	\$ 17,800	100%	\$ 22,973	100%	\$ 16,480	100%	\$ 21,283	100%

**Impact of Our Diagnostic Products Strategy**

We have only recently commercially launched the diagnostic version of our nCounter System and Prosigna. Over time, we intend to build a menu of additional diagnostic tests that can be run on our nCounter Analysis System. As researchers discover how genomic information can be used to improve clinical decision-making, these discoveries can be translated and validated as diagnostic tests based on our nCounter Elements reagents. In certain situations, we intend to translate their discoveries into *in vitro* diagnostic assays. We in-licensed the rights to intellectual property that forms the basis of Prosigna from Bioclassifier, LLC, which was founded by several of our life sciences research customers. We intend to enter into similar arrangements with our life sciences research customers and other researchers for future diagnostic gene signatures. Our strategy is to target intellectual property rights to potential diagnostic methods that are well understood, have the potential to facilitate changes in treatment with a major impact on outcome and cost, have the potential to support value-based pricing, and for which tissue samples for clinical validation are readily available. For example, in February 2013 we secured an option from a customer to acquire an exclusive worldwide license for a gene signature that could be used, after further development, as a Laboratory Developed Test, or, after appropriate regulatory authorization, for a second molecular diagnostic to identify patients with cirrhosis who are at highest risk of developing HCC and to determine whether a patient who has been diagnosed with HCC is likely to have a recurrence. This disciplined approach is designed to efficiently focus our research and development investment on development of potential products, rather than discovery of new gene signatures. Licenses may include upfront, milestone and/or annual cash payments and revenue-based royalties. The number and amount of such payments and royalty rates are expected to vary depending on the level of development and commercial potential of in-license opportunities.

We believe that our *in vitro* diagnostics business model is more capital efficient than the clinical laboratory services model and has the potential to become profitable on a relatively small revenue base. Our diagnostics business leverages many of the capabilities of our life sciences business, including our technology platform and instrument sales, product development, manufacturing, and administrative functions. Because we provide *in vitro* diagnostics kits rather than clinical laboratory services, we do not incur the costs of clinical laboratory infrastructure, sample logistics, or contracting with and billing managed care organizations. We believe that our customers will be motivated by the potential to improve patient care, broaden patient access and profit from testing services based on Prosigna and other potential nCounter-based diagnostics, which will encourage market adoption and potentially reduce sales and marketing expenditures relative to a centralized laboratory model.



**Table of Contents****Results of Operations****Comparison of Nine Months Ended September 30, 2012 and 2013***Revenue; Cost of Revenue; Gross Profit*

	Nine Months Ended September 30,		Change 2012 v. 2013	
	2012	2013	Dollars	Percentage
	(Dollars in thousands)			
Product revenue:				
Instruments:				
Life Sciences	\$ 6,262	\$ 7,226	\$ 964	15%
Diagnostics		488	488	
Total Instrument revenue	6,262	7,714	1,452	23
Consumables:				
Life Sciences	9,325	12,380	3,055	33
Diagnostics		41	41	
Total Consumable revenue	9,325	12,421	3,096	33
Service Revenue	893	1,148	255	29
Total revenue	16,480	21,283	4,803	29
Cost of revenue	9,076	10,188	1,112	12
Gross profit	\$ 7,404	\$ 11,095	\$ 3,691	50

## Gross margin

45%

52%

Instrument revenue increased significantly for the nine month period ended September 30, 2013 due to an increase in the number of instruments sold. Instrument revenue included approximately \$0.5 million from our diagnostics segment for the nine month period ended September 30, 2013. The increase in consumable revenue was driven by growth in our installed base of instruments. The increase in service revenue was primarily related to an increase in the number of instruments covered by service contracts.

The increase in cost of revenue was related to the increased volume of both instruments and consumables sold. Gross margin improved due to cost efficiencies associated with increased consumables production volume and several large custom consumable orders with unusually low per unit manufacturing costs. These improvements were partially offset by a shift in product mix toward instruments.

*Research and Development Expense*

	Nine Months Ended September 30,		Change 2012 v. 2013	
	2012	2013	Dollars	Percentage
	(Dollars in thousands)			
Research and development expense	\$ 8,253	\$ 10,469	\$ 2,216	27%

The increase reflected a \$2.2 million increase in personnel-related expenses to support the advancement of our nCounter technology and clinical development of Prosigna. The increase in the period included \$0.6 million of increased engineering costs for development of the next generation of our nCounter system. Decreases in Prosigna clinical study costs of \$0.7 million, after completion of the ABCSG8 study in late 2012, partially offset the increases.

*Selling, General and Administrative Expense*

Edgar Filing: NanoString Technologies Inc - Form S-1

	<b>Nine Months Ended September 30,</b>		<b>Change 2012 v. 2013</b>	
	<b>2012</b>	<b>2013</b>	<b>Dollars</b>	<b>Percentage</b>
	<b>(Dollars in thousands)</b>			
Selling, general and administrative expense	\$ 10,588	\$ 20,822	\$ 10,234	97%

-52-

**Table of Contents**

The increase for the nine month period was primarily attributable to \$3.7 million of increased staffing and personnel-related costs to support sales and marketing and administration; \$2.1 million of increased external marketing and other consulting costs related to the commercial launch of Prosigna; \$2.6 million of increased legal costs and \$0.8 million of increased corporate professional fees and other public company costs.

*Other Income (Expense), Net*

	Nine Months Ended September 30,		Change 2012 v. 2013	
	2012	2013	Dollars (Dollars in thousands)	Percentage
Interest income	\$ 17	\$ 28	\$ 11	65%
Interest expense	(551)	(1,412)	(861)	156
Other income (expense)	(26)	(30)	(4)	15
Revaluation of preferred stock warrant liability	150	1,156	1,006	671
<b>Total other income (expense), net</b>	<b>\$ (410)</b>	<b>\$ (258)</b>	<b>\$ 152</b>	<b>(37)</b>

The increase in interest expense was driven by increased borrowing under our credit facility during 2013, from \$7.5 million as of September 30, 2012 to \$18.0 million as of September 30, 2013.

The increase in other income from the revaluation of the preferred stock warrant liability resulted from a re-measurement of the fair value of preferred stock warrants using the Black-Scholes option pricing model, which was primarily impacted by a decrease in the valuation of the underlying stock. Upon closing of our initial public offering in July 2013, all outstanding preferred stock was automatically converted into common stock, and the warrants to purchase preferred stock converted into warrants to purchase common stock. As a result, the preferred stock warrant liability was reclassified to stockholders' equity.

*Comparison of Years Ended December 31, 2011 and 2012**Revenue; Cost of Revenue; Gross Profit*

	Year Ended December 31,		Change 2011 v. 2012	
	2011	2012	Dollars (Dollars in thousands)	Percentage
<b>Revenue:</b>				
<b>Product revenue:</b>				
Instruments	\$ 7,112	\$ 8,786	\$ 1,674	24%
Consumables	9,997	13,036	3,039	30
Service revenue	691	1,151	460	67
<b>Total revenue</b>	<b>17,800</b>	<b>22,973</b>	<b>5,173</b>	<b>29</b>
<b>Cost of revenue</b>	<b>9,777</b>	<b>12,361</b>	<b>2,584</b>	<b>26</b>
<b>Gross profit</b>	<b>\$ 8,023</b>	<b>\$ 10,612</b>	<b>\$ 2,589</b>	<b>32</b>
<b>Gross margin</b>	<b>45%</b>	<b>46%</b>		

The increase in instrument revenue was attributable to an increase in the number of systems sold, primarily related to an increase in sales outside of the United States. The net selling price of our instruments was relatively flat. The increase in consumable revenue was related to our increased instrument installed base. Overall, we derived \$3.3 million in incremental revenue from customers outside of North America as a result of the expansion of our overseas sales and marketing efforts.

## Edgar Filing: NanoString Technologies Inc - Form S-1

The increase in cost of revenue was attributable to an increase in the number of systems sold, as well as the increased costs associated with higher volumes of consumables sold. Gross margin was relatively flat, consistent with the relatively constant product mix in the two years.

**Table of Contents***Research and Development Expense*

	Year Ended December 31,		Change 2011 v. 2012	
	2011	2012	Dollars	Percentage
			(Dollars in thousands)	
Research and development expense	\$ 8,990	\$ 11,635	\$ 2,645	29%

The increase was primarily attributable to a \$2.2 million increase in clinical study and sample acquisition costs and an \$0.8 million increase in facility-related costs due to the expansion of our facility. The increases were offset in part by the absence of \$0.5 million in third-party in-license fees incurred in 2011.

*Selling, General and Administrative Expense*

	Year Ended December 31,		Change 2011 v. 2012	
	2011	2012	Dollars	Percentage
			(Dollars in thousands)	
Selling, general and administrative expense	\$ 9,529	\$ 15,486	\$ 5,957	63%

The increase was primarily attributable to a \$2.5 million increase in personnel-related expenses as a result of increased sales and administrative headcount to support the growth of our business, \$2.0 million in marketing consulting costs in preparation for the commercial launch of Prosigna, and \$0.9 million in corporate and intellectual property-related legal costs.

*Other Income (Expense), Net*

	Year Ended December 31,		Change 2011 v. 2012	
	2011	2012	Dollars	Percentage
			(Dollars in thousands)	
Interest income	\$ 10	\$ 21	\$ 11	110%
Interest expense	(599)	(804)	(205)	34
Other income (expense)	80	(29)	(109)	(136)
Revaluation of preferred stock warrant liability	73	(387)	(460)	(630)
Total other income (expense), net	\$ (436)	\$ (1,199)	\$ (763)	175

The increase in interest expense was driven by the increase in borrowings under our existing credit facility compared to the prior period level of borrowings under our 2010 loan and security agreement and convertible subordinated notes.

The increase in expense from the revaluation of the preferred stock warrant liability was driven by an increase in the valuation of our stock.

**Table of Contents****Comparison of Years Ended December 31, 2010 and 2011***Revenue; Cost of Revenue; Gross Profit*

	Year Ended December 31,		Change 2010 v. 2011	
	2010	2011	Dollars (Dollars in thousands)	Percentage
Revenue:				
Product revenue:				
Instruments	\$ 6,472	\$ 7,112	\$ 640	10%
Consumables	5,034	9,997	4,963	99
Service revenue	224	691	467	208
Total revenue	11,730	17,800	6,070	52
Cost of revenue	9,128	9,777	649	7
Gross profit	\$ 2,602	\$ 8,023	\$ 5,421	208
Gross margin	22%	45%		

The increase in instrument revenue was attributable to an increase in the number of systems sold, which was primarily related to an increase outside of the United States. The net selling price of our instruments was relatively flat. The increase in consumable revenue was related to our increased instrument installed base as well as the introduction of new applications and panel products. Overall, we derived \$2.7 million in incremental revenue from customers outside of North America as a result of the expansion of our overseas sales and marketing efforts.

The increase in cost of revenue was attributable to an increase in the number of systems sold, as well as the increased costs associated with higher volumes of consumables sold. This increase was largely offset by a decrease in manufacturing costs for certain consumable products. Gross margin increased primarily due to manufacturing process improvements, raw material costs reductions, increased manufacturing volumes, and a shift in product mix toward consumables, all of which lowered costs as a percentage of revenue.

*Research and Development Expense*

	Year Ended December 31,		Change 2010 v. 2011	
	2010	2011	Dollars (Dollars in thousands)	Percentage
Research and development expense	\$ 7,547	\$ 8,990	\$ 1,443	19%

The increase was primarily attributable to a \$0.9 million increase in personnel related expenses as a result of increased headcount, a \$0.7 million increase in facility costs as we leased additional space, and \$0.5 million in third-party in-license fees incurred in 2011. Partially offsetting the increase was the absence of \$0.7 million in development costs incurred in 2010 for a second generation of our instruments launched in 2011.

*Selling, General and Administrative Expense*

	Year Ended December 31,		Change 2010 v. 2011	
	2010	2011	Dollars (Dollars in thousands)	Percentage
Selling, general and administrative expense	\$ 8,027	\$ 9,529	\$ 1,502	19%

The increase was primarily attributable to a \$1.6 million increase in personnel-related expenses, largely as a result of increased sales and administrative headcount, and to a lesser extent, an increase in facility costs as we leased additional space. Partially offsetting the increase was

the reduction of \$0.5 million in consulting fees related to the evaluation of the Prosigna market opportunity.

**Table of Contents***Other Income (Expense), Net*

	Year Ended December 31,		Change 2010 v. 2011	
	2010	2011	Dollars (Dollars in thousands)	Percentage
Interest income	\$ 29	\$ 10	\$ (19)	(66%)
Interest expense	(94)	(599)	(505)	537
Other income (expense)	254	80	(174)	(69)
Revaluation of preferred stock warrant liability	15	73	58	387
<b>Total other income (expense), net</b>	<b>\$ 204</b>	<b>\$ (436)</b>	<b>\$ (640)</b>	<b>(314)</b>

The increase in interest expense was driven by a \$5.0 million increase in borrowings under our 2010 loan and security agreement in November 2010 and the issuance of an aggregate of \$5.0 million in convertible promissory notes in June and September 2011. The decrease in other income was attributable to a one-time payment of \$0.2 million received in 2010 under the Qualifying Therapeutic Discovery Project Program.

**Quarterly Results of Operations**

The following tables set forth selected unaudited quarterly statements of operations data for the last eleven fiscal quarters. The unaudited interim financial statements for each of these quarters have been prepared on the same basis as the audited consolidated financial statements included elsewhere in this prospectus and, in the opinion of management, reflect all adjustments, which include only normal recurring adjustments, necessary to a fair statement of our results of operations and financial position for these periods. These data should be read in conjunction with the audited consolidated financial statements and accompanying notes included elsewhere in this prospectus. These quarterly operating results are not necessarily indicative of our operating results for any future period.

	Three Months Ended										
	March 31, 2011	June 30, 2011	September 30, 2011	December 31, 2011	March 31, 2012	June 30, 2012	September 30, 2012	December 31, 2012	March 31, 2013	June 30, 2013	September 30, 2013
(Dollars in thousands)											
<b>Product Revenue:</b>											
Instruments											
Life Sciences	\$ 2,060	\$ 1,898	\$ 1,078	\$ 2,076	\$ 1,500	\$ 2,579	\$ 2,183	\$ 2,524	\$ 1,639	\$ 2,320	\$ 3,267
Diagnostics										203	285
<b>Total Instrument revenue</b>	<b>2,060</b>	<b>1,898</b>	<b>1,078</b>	<b>2,076</b>	<b>1,500</b>	<b>2,579</b>	<b>2,183</b>	<b>2,524</b>	<b>1,639</b>	<b>2,523</b>	<b>3,552</b>
Consumables											
Life Sciences	2,147	2,843	2,634	2,373	2,703	3,054	3,568	3,711	3,699	4,305	4,376
Diagnostics											41
<b>Total Consumable Revenue</b>	<b>2,147</b>	<b>2,843</b>	<b>2,634</b>	<b>2,373</b>	<b>2,703</b>	<b>3,054</b>	<b>3,568</b>	<b>3,711</b>	<b>3,699</b>	<b>4,305</b>	<b>4,417</b>
Service Revenue	137	138	181	235	299	310	284	258	338	390	420
<b>Total revenue</b>	<b>4,344</b>	<b>4,879</b>	<b>3,893</b>	<b>4,684</b>	<b>4,502</b>	<b>5,943</b>	<b>6,035</b>	<b>6,493</b>	<b>5,676</b>	<b>7,218</b>	<b>8,389</b>
Costs and expenses:											
Cost of revenue	2,532	2,477	2,189	2,579	2,656	3,334	3,086	3,285	2,882	3,522	3,784
Research and development	2,620	2,229	1,884	2,257	2,197	2,971	3,085	3,382	3,059	3,626	3,784
Selling, general and administrative	2,327	2,473	2,110	2,619	3,167	3,251	4,170	4,898	6,126	6,708	7,988
Other (income) expenses	19	67	282	68	92	(42)	360	789	868	(1,143)	533
<b>Total costs and expenses</b>	<b>7,498</b>	<b>7,246</b>	<b>6,465</b>	<b>7,523</b>	<b>8,112</b>	<b>9,514</b>	<b>10,701</b>	<b>12,354</b>	<b>12,935</b>	<b>12,713</b>	<b>16,089</b>



Edgar Filing: NanoString Technologies Inc - Form S-1

Net loss	\$ (3,154)	\$ (2,367)	\$ (2,572)	\$ (2,839)	\$ (3,610)	\$ (3,571)	\$ (4,666)	\$ (5,861)	\$ (7,259)	\$ (5,495)	\$ (7,700)
----------	------------	------------	------------	------------	------------	------------	------------	------------	------------	------------	------------

-56-

## **Table of Contents**

Consistent with others in our industry, we have experienced variations in revenue related to the ordering patterns of our customers. The third calendar quarter has tended to be relatively weak for instruments and consumables due to summer holidays and vacations of potential academic customers. In addition, we believe that instrument sales will typically be stronger in the fourth quarter of each calendar year due to the availability of residual funding for capital expenditures prior to the end of many customers' fiscal year and less strong in the first quarter of each calendar year as a result of such expenditures. As a result of such factors, we expect to continue to see seasonality and quarter-to-quarter variations in our revenue, especially related to instruments.

Cost of revenue tends to be higher on a percentage basis in the first quarter of each year due to lower revenue over which we must spread our fixed manufacturing and overhead costs. Product mix from quarter-to-quarter also impacts cost of revenue on a percentage basis.

Research and development expense has varied from quarter-to-quarter due to the timing of license payments and clinical study activity. For example, we incurred third-party clinical study costs of \$1.0 million, \$0.5 million and \$0.7 million in the second, third and fourth quarter of 2012, respectively. We believe that our continued investment in research and development is essential to our long-term competitive position, and we expect these expenses to continue to increase in future periods.

Selling, general and administrative expense increased each quarter in 2012 primarily due to increased sales and marketing headcount and marketing consulting costs in preparation for the commercial launch of Prosigna. Similarly, selling, general and administrative expense increased in each of the first three quarters of 2013, primarily attributable to increased staffing and personnel-related costs to support sales, marketing and administration, increased legal costs, the majority of which related to a law suit settled in September 2013, and increased corporate professional fees and other costs of being a public company.

## **Liquidity and Capital Resources**

As of September 30, 2013, we had cash, cash equivalents and short-term investments of \$52.2 million. Since inception, we have financed our operations primarily through the sale of equity securities and, to a lesser extent, from borrowings. Our principal uses of cash are funding our operations, debt service payments as described below, and capital expenditures.

### ***Sources of Funds***

Our cash used in operations for the year ended December 31, 2012 and the nine months ended September 30, 2013 was \$14.8 million and \$21.5 million, respectively. During 2012, we incurred \$13.0 million in term loan borrowings under our credit facility and amended the credit facility to allow for the incurrence of up to an additional \$10.0 million in term loan borrowings. Also in 2012, we issued and sold shares of Series E preferred stock which generated proceeds after offering expenses of \$15.1 million. In April 2013, we incurred \$5.0 million of the remaining term loan borrowings under our credit facility. In July 2013, we raised \$50.2 million, before offering expenses, in our initial public offering. Our cash, cash equivalents and short-term investments are sufficient to meet our anticipated cash needs for at least the next 12 months, when taken with the net proceeds of this offering. However, we may need to raise additional capital to expand the commercialization of our products, fund our operations and further our research and development activities. Our future funding requirements will depend on many factors, including: market acceptance of our products; the cost and timing of establishing additional sales, marketing and distribution capabilities; the cost of our research and development activities; the cost and timing of regulatory clearances or approvals; the effect of competing technological and market developments; and the extent to which we acquire or invest in businesses, products and technologies, including new licensing arrangements for new products, although we currently have no commitments or agreements to complete any such transactions.

From time to time, we may explore additional financing sources and means to lower our cost of capital, which could include equity, equity-linked and debt financing. There can be no assurance that any additional financing will be available to us on acceptable terms. If we raise additional funds by issuing equity or equity-linked securities, our stockholders may experience dilution. Debt financing, if available, may involve

---

**Table of Contents**

covenants restricting our operations or our ability to incur additional debt. Any debt or additional equity financing that we raise may contain terms that are not favorable to us or our stockholders. If we raise additional funds through collaboration and licensing arrangements with third parties, it may be necessary to relinquish some rights to our technologies or our products, or grant licenses on terms that are not favorable to us. If we are unable to raise adequate funds, we may have to liquidate some or all of our assets, or delay, reduce the scope of or eliminate some or all of our development programs. If we do not have, or are not able to obtain, sufficient funds, we may have to delay development or commercialization of our products or license to third parties the rights to commercialize products or technologies that we would otherwise seek to commercialize. We also may have to reduce marketing, customer support or other resources devoted to our products or cease operations.

In January 2014, we entered into a non-binding letter of intent for a credit agreement with a lender which would allow us to refinance our existing credit facility and potentially incur up to an aggregate of \$45 million in term loan borrowings or up to an aggregate of approximately \$52 million if we elect to exercise in full an option to pay in kind a portion of the interest that would accrue on the borrowings under the credit agreement. We expect this credit agreement will contain customary conditions to borrowings, events of default and negative covenants, including covenants that could limit our ability to, among other things, incur additional indebtedness, liens or other encumbrances, make dividends or other distributions, and sell or transfer assets. We also expect that the credit agreement will include certain customary financial covenants. Our obligations under the credit agreement will be secured by substantially all of our assets. However, there can be no assurance that we will successfully enter into this credit agreement.

*Credit Facility*

In March 2012, we entered into a loan and security agreement, which we refer to as our credit facility, pursuant to which we incurred \$7.5 million in term loan borrowings, which we refer to as Term A borrowings. Pursuant to the credit facility, we incurred an additional \$5.5 million of term loan borrowings in December 2012, which we refer to as Term B borrowings. Also in December 2012 and April 2013, we amended the credit facility to allow for the incurrence on or prior to April 30, 2013 of up to an additional \$10.0 million in term loan borrowings, of which we incurred \$5.0 million and refer to as Term C borrowings. Interest on term loan borrowings is determined at the time of borrowing and accrues at a fixed rate equal to the three month LIBOR plus 8.39% (subject to a LIBOR floor of 0.50%). Through January 2014, we are required to only pay interest on outstanding term borrowings on a monthly basis. Following the expiration of the interest only payment period, we are required to pay principal and interest in 30 equal monthly payments, plus an end of term payment equal to 5.5% of the amount borrowed. We may at our option prepay all of the term loan borrowings by paying the lender, among other things, all principal and accrued interest, the end of term payment plus a make-whole premium. Pursuant to the credit facility, from time to time we can also incur revolver borrowings of up to the lesser of \$2.0 million and a borrowing base tied to the amount of eligible accounts receivable. Interest on revolver borrowings accrues at a floating rate equal to the prime rate plus 3.70% (subject to a floor of 6.95%) and is payable monthly. We are also required to pay a fee of 0.075% per month on the unused portion of the revolver borrowings.

The credit facility contains customary conditions to borrowing, events of default and covenants, including covenants that restrict our ability to dispose of assets, merge with or acquire other entities, incur indebtedness, incur encumbrances, make distributions to holders of our capital stock, make investments or engage in transactions with our affiliates. In addition, we must comply with a financial covenant based on life sciences revenue. This financial covenant is measured monthly on a trailing three month basis. We were in compliance with all covenants as of December 31, 2012 and September 30, 2013. Our obligations under the credit facility are secured by substantially all of our assets other than intellectual property.

As of September 30, 2013, we had \$7.5 million of Term A borrowings outstanding, \$5.5 million of Term B borrowings and \$5.0 million of Term C borrowings, which each accrue interest at 8.89%.

We issued the lenders warrants to purchase an aggregate of 44,389 shares of our Series D preferred stock in connection with entering into the credit facility and warrants to purchase an additional 32,551 shares of Series D

---

## **Table of Contents**

preferred stock in connection with the incurrence of the Term B borrowings. Such warrants have ten year terms and an exercise price of \$8.45 per share. In December 2012 we issued the lenders warrants to purchase an aggregate 20,837 shares of our Series E preferred stock at an exercise price of \$14.40 per share in connection with the amendment of the credit facility. In addition, in April 2013 we issued the lenders warrants to purchase an aggregate of 10,418 shares of our Series E preferred stock at an exercise price of \$14.40 per share in connection with the incurrence of Term C borrowings. In connection with our initial public offering, these warrants became exercisable for shares of our common stock.

### *Convertible Promissory Notes*

In June 2011 and September 2011, we issued approximately \$5.0 million aggregate principal amount of our subordinated convertible promissory notes to existing investors. Interest on the notes accrued on the unpaid principal balance at 8.0% per year. The principal amount of and accrued interest on the subordinated convertible notes converted into an aggregate of 602,172 shares of our Series D preferred stock in November 2011. In addition, we issued warrants to purchase an aggregate of 118,368 shares of our Series D preferred stock at an exercise price of \$8.45 per share to the holders of the subordinated convertible notes. These warrants will expire upon the earliest of (1) November 1, 2018, (2) a change in control of our company and (3) the sale of all or substantially all of our assets. Following our initial public offering, such warrants became exercisable for an aggregate of 617,605 shares of our common stock.

### *2010 Loan and Security Agreement*

In November 2010, we amended our then-existing loan and security agreement to provide for up to \$2.0 million of equipment term borrowings and, subject to certain conditions, up to \$3.0 million in revolver borrowings. Under the agreement, we incurred \$1.9 million of equipment term borrowings, which were payable over periods of up to 36 months in equal monthly installments of principal and interest. In addition, we incurred maximum borrowings under the revolver of \$2.7 million. As amended, interest on borrowings under the 2010 loan and security agreement accrued at a floating rate equal to the bank's prime reference rate (subject to a floor of LIBOR plus 2.5%, or, if LIBOR cannot be determined, then 2.5%), plus up to 1.5% depending on the facility. All of the indebtedness incurred under the 2010 loan and security agreement was repaid in 2012 in connection with the entry into our existing credit facility.

We also issued the lender under the 2010 loan and security agreement warrants to purchase 4,691 shares of our Series B preferred stock at an exercise price of \$17.47 per share. After giving effect to the anti-dilution provisions of such warrant, in connection with our initial public offering, these warrants became exercisable for 7,315 shares of our common stock. These warrants will expire in October 2014.

### *Use of Funds*

Our principal uses of cash are funding our operations, satisfaction of our obligations under our debt instruments, and other working capital requirements. Over the past several years, our revenue has increased significantly from year to year and, as a result, our cash flows from customer collections have increased. However, our operating expenses have also increased as we have invested in growing our existing life sciences business and in developing Prosigna and preparing it for commercialization. As a result, our cash used in operating activities has either remained relatively constant or increased. We expect our operating cash requirements to increase in the future as we (1) increase sales and marketing activities to expand the installed base of our nCounter Analysis Systems among life sciences research customers and clinical laboratories, (2) commercialize, and conduct studies to expand the clinical utility of, Prosigna, and (3) develop new applications, chemistry and instruments for our nCounter platform.

We will need to raise additional funds to support our operations, and such funding may not be available to us on acceptable terms, or at all. If we are unable to raise additional funds when needed, our operations and ability to execute our business strategy could be adversely affected. We may seek to raise additional funds through equity, equity-linked or debt financings. If we raise additional funds through the incurrence of

**Table of Contents**

indebtedness, such indebtedness would have rights that are senior to holders of our equity securities and could contain covenants that restrict our operations. Any additional equity financing may be dilutive to our stockholders.

**Historical Cash Flow Trends**

The following table shows a summary of our cash flows for the periods indicated:

	Years Ended December 31,			Nine Months Ended September 30,	
	2010	2011	2012 (In thousands)	2012	2013
Cash used in operating activities	\$ (10,965)	\$ (10,692)	\$ (14,808)	\$ (11,140)	\$ (21,483)
Cash used in investing activities	(1,932)	(2,800)	(428)	(312)	(33,506)
Cash provided by financing activities	15,524	19,994	26,060	5,945	52,589

**Operating Cash Flows**

We derive operating cash flows from cash collected from the sale of our products and services. These cash flows received are outweighed by our use of cash for operating expenses to support the growth of our business. As a result, we have historically experienced negative cash flows from operating activities as we have expanded our business in the United States and other markets and this will likely continue for the foreseeable future.

Net cash used in operating activities for the nine months ended September 30, 2013 consisted of our net loss of \$20.5 million and \$2.4 million of cash used for working capital purposes. These uses were partially offset by \$1.4 million of net non-cash income and expense items, such as depreciation and amortization, stock-based compensation and change in the fair value of preferred stock warrants.

Net cash used in operating activities for the nine months ended September 30, 2012 consisted of our net loss of \$11.8 million and \$1.4 million of cash used for working capital purposes. These uses were partially offset by \$2.1 million of non-cash expense items, such as depreciation and amortization and stock-based compensation.

Net cash used in operating activities for 2012 consisted of our net loss of \$17.7 million and changes in our operating assets and liabilities of \$0.4 million, which were partially offset by non-cash expense items such as depreciation and amortization of our equipment and leasehold improvements of \$1.9 million, stock-based compensation of \$0.7 million, revaluation of preferred stock warrant liability of \$0.4 million and amortization of debt discounts and issuance cost of \$0.1 million.

Net cash used in operating activities for 2011 consisted of our net loss of \$10.9 million and changes in our operating assets and liabilities of \$1.8 million, which were partially offset by non-cash expense items including depreciation and amortization of our equipment and leasehold improvements of \$1.5 million, amortization of debt discounts and issuance costs of \$0.3 million and stock-based compensation of \$0.2 million.

Net cash used in operating activities for 2010 consisted of our net loss of \$12.8 million, offset in part by a \$0.6 million change in operating assets and liabilities and non-cash expense items such as depreciation and amortization of our equipment and leasehold improvements of \$1.0 million and stock-based compensation of \$0.1 million.

**Investing Cash Flows**

Net cash used in investing activities for each of the periods presented was primarily for the purchase of laboratory, manufacturing and computer equipment and software to support our expanding infrastructure. In 2011, we leased additional laboratory and office space and incurred \$1.8 million in expenses related to leasehold improvements and our restricted cash related to this leased space increased by \$0.1 million. In 2012 and 2013, we purchased lesser amounts of property and equipment required to support the growth and expansion of our operations.

## Table of Contents

Our most significant cash flows used in investing activities for the nine months ended September 30, 2013 are for the purchase of short-term investments. These amounts primarily relate to shifts between cash and cash equivalents and short-term investments. Because we manage our cash usage with respect to our total cash, cash equivalents and short-term investments, we do not consider these cash flows to be important to an understanding of our liquidity and capital resources. Also included in investing activities for each of the periods presented are the purchases of laboratory, manufacturing and computer equipment and software to support our expanding infrastructure. We made no major capital expenditures for the remainder of 2013.

### *Financing Cash Flows*

Historically, we have funded our operations through the issuance of equity securities and the incurrence of indebtedness.

Net cash provided by financing activities for the nine months ended September 30, 2013 consisted of net proceeds of \$47.4 million from our initial public offering, proceeds from term loan borrowings of \$5.0 million and proceeds from exercise of stock options of \$0.4 million. These proceeds were partially offset by repayments of borrowings of \$0.2 million.

Net cash provided by financing activities for the nine months ended September 30, 2012 consisted of \$7.5 million of borrowing under our existing credit facility, which was partially offset by repayments of borrowings under a previous loan agreement of \$1.7 million.

For 2012, net cash provided by financing activities consisted of \$13.0 million of borrowing under our credit facility and \$15.1 million from the issuance of Series E preferred stock. This was partially offset by repayments of borrowings under our 2010 loan and security agreement of \$1.7 million and payments related to deferred offering costs of \$0.6 million.

For 2011, net cash provided by financing activities consisted of the issuance of Series D preferred stock which generated proceeds of \$14.9 million, issuance of our subordinated convertible notes which generated proceeds of \$5.0 million and incurrence of an aggregate of \$5.0 million of borrowings under our 2010 loan and security agreement, which were offset in part by repayment of \$4.9 million of such borrowings.

For 2010, net cash provided by financing activities consisted of the issuance of \$15.0 million of Series C preferred stock and net incurrence of indebtedness under our 2010 loan and security agreement of \$0.5 million.

### **Contractual Obligations**

The following table reflects a summary of our contractual obligations as of December 31, 2012.

Contractual Obligations <sup>(1)</sup>	Total	Payments due by period			
		Less than 1 Year	1-3 Years (In thousands)	3-5 Years	More than 5 Years
Operating lease obligations <sup>(2)</sup>	\$ 7,295	\$ 2,009	\$ 3,950	\$ 1,336	\$
Long-term debt obligations <sup>(3)</sup>	13,993	3,009	10,984		
Inventory purchase obligations <sup>(4)</sup>	2,604	2,604			
Total	\$ 23,892	\$ 7,622	\$ 14,934	\$ 1,336	\$

(1) Excludes royalty obligations based on net sales of products, including royalties payable to the Institute for Systems Biology, as any such amounts are not currently determinable.

(2) Operating lease costs are primarily for office, laboratory and manufacturing space at our headquarters.

## Edgar Filing: NanoString Technologies Inc - Form S-1

- (3) Includes principal and interest on long-term debt obligations other than our Term C borrowings.
  
- (4) Purchase obligations consist of contractual and legally binding commitments under outstanding purchase orders to purchase long lead time inventory items.

-61-

---

## **Table of Contents**

### **Critical Accounting Policies and Significant Estimates**

Our discussion and analysis of our financial condition and results of operations are based upon our financial statements which have been prepared in accordance with U.S. generally accepted accounting principles, or GAAP. The preparation of these financial statements requires us to make estimates and judgments that affect the reported amounts of assets and liabilities and related disclosure of contingent assets and liabilities, revenue and expenses at the date of the financial statements. Generally, we base our estimates on historical experience and on various other assumptions in accordance with GAAP that we believe to be reasonable under the circumstances. Actual results may differ from these estimates.

Critical accounting policies and estimates are those that we consider the most important to the portrayal of our financial condition and results of operations because they require our most difficult, subjective or complex judgments, often as a result of the need to make estimates about the effect of matters that are inherently uncertain. Our critical accounting policies and estimates include those related to:

revenue recognition;

stock-based compensation;

inventory valuation;

fair value measurements; and

income taxes.

### ***Revenue Recognition***

We recognize revenue when: (1) persuasive evidence of an arrangement exists; (2) delivery has occurred or services have been rendered; (3) the price to the customer is fixed or determinable; and (4) collectability is reasonably assured. We generate revenue from the sale of products and services. Our products consist of our proprietary nCounter Analysis System and related consumables. Services consist of extended warranties and service fees for assay processing. A delivered product or service is considered to be a separate unit of accounting when it has value to the customer on a stand-alone basis. Products or services have value on a stand-alone basis if they are sold separately by any vendor or if the customer could resell the delivered product.

Systems product revenue is recognized upon installation and calibration in geographic regions where such services are only available from our specialized technicians. In these regions, systems and related installation and calibration are considered to be one unit of accounting, as systems are required to be professionally installed and calibrated before use. In certain geographic regions, installation and calibration services are available from other vendors, and in such regions they are considered separate revenue elements. Consumables are considered to be separate units of accounting as they are sold separately. Consumables product revenue is recognized upon shipment.

Service revenue is recognized when earned, which is generally upon the rendering of the related services. Extended warranties and service fees for assay processing are each considered separate units of accounting as they are sold separately. We offer extended warranties on our nCounter Analysis System for periods ranging from 12 to 36 months after the end of the standard 12-month warranty period. Extended warranties are generally separately priced. Revenue from extended warranties are deferred and recognized in income on a straight-line basis over the warranty period.

We allocate the contract consideration at the inception of the contract to the deliverables based upon their relative selling prices. To date, selling prices have been established by reference to vendor specific objective evidence based on stand-alone sales transactions for each deliverable. Vendor specific objective evidence is considered to have been established when a substantial majority of individual sales transactions within the previous 12 month period fall within a reasonably narrow range, which we have defined to be plus or minus 15% of the median sales price of actual stand-alone sales transactions. We use our best estimate of selling price for individual deliverables when vendor specific objective evidence or third-party evidence is unavailable. Allocated revenue is only recognized for each deliverable when the revenue recognition criteria



have been met.

## **Table of Contents**

### ***Stock-based Compensation***

Prior to the closing of our initial public offering, we granted stock options at exercise prices believed to be equal to the fair value of the common stock underlying such options as determined by the board of directors, with input from management, on the date of grant. Because such grants occurred prior to the public trading of our common stock, the board of directors exercised significant judgment in determining the fair market value of our common stock. The valuations were consistent with the guidance and methods outlined in the AICPA Practice Aid Valuation of Privately-Held-Company Equity Securities Issued as Compensation, or AICPA Practice Aid, for all option grant dates. After the closing of the initial public offering, we granted stock options with exercise prices based on market prices.

We account for stock-based compensation at fair value. Stock-based compensation costs are recognized based on their grant date fair value estimated using the Black-Scholes option pricing model. Stock-based compensation expense recognized in the statement of comprehensive income (loss) is based on options ultimately expected to vest and has been reduced by an estimated forfeiture rate based on our historical and expected forfeiture patterns. We use the straight-line method of allocating compensation cost over the requisite service period of the related award.

Determining the fair value of stock-based awards at the grant date under the Black-Scholes option pricing model requires judgment, including estimating the value per share of our common stock, risk-free interest rate, expected term and dividend yield and volatility. The assumptions used in calculating the fair value of stock-based awards represent our best estimates based on management judgment and subjective future expectations. These estimates involve inherent uncertainties. If any of the assumptions used in the Black-Scholes option pricing model significantly change, stock-based compensation for future awards may differ materially from the awards granted previously.

The expected term of options granted is based on historical experience of similar awards and expectations of future employee behavior. The risk-free interest rate for the expected term of the option is based on the U.S. Treasury yield curve in effect at the time of grant. We have not paid and do not anticipate paying cash dividends on our common stock; therefore, the expected dividend yield is assumed to be zero. We based our estimate of volatility on the estimated volatility of similar companies whose share prices are publicly available.

### ***Inventory Valuation***

Inventory consists of raw materials, certain component parts to be used in manufacturing our products and finished goods. Inventory is stated at the lower of cost or market. Cost is determined using a standard cost system, whereby the standard costs are updated periodically to reflect current costs and market represents the lower of replacement cost or estimated net realizable value. We record adjustments to inventory for potentially excess, obsolete, slow-moving or impaired items. The business environment in which we operate is subject to rapid changes in technology and customer demand. We regularly review inventory for excess and obsolete products and components, taking into account product life cycle and development plans, product expiration and quality issues, historical experience and our current inventory levels. If actual market conditions are less favorable than anticipated, additional inventory adjustments could be required.

### ***Fair Value Measurements***

We establish the fair value of our assets and liabilities using the price that would be received to sell an asset or paid to transfer a financial liability in an orderly transaction between market participants at the measurement date. A fair value hierarchy is used to measure fair value. The three levels of the fair value hierarchy are as follows:

Level 1 Quoted prices in active markets for identical assets and liabilities.

Level 2 Quoted prices for similar instruments in active markets; quoted prices for identical or similar instruments in markets that are not active; and model-derived valuations in which all significant inputs and significant value drivers are observable in active markets.

---

**Table of Contents**

Level 3 Valuations derived from valuation techniques in which one or more significant inputs or significant value drivers are unobservable.

Prior to the closing of our initial public offering, we recorded preferred stock warrant liability at fair value. Preferred stock warrant liability was categorized as Level 3 because it was valued based on unobservable inputs and our judgment due to the absence of quoted market prices, inherent lack of liquidity and the long-term nature of such financial instruments. We performed a fair value assessment of the preferred stock warrant inputs on a quarterly basis using the Black-Scholes option pricing model. The assumptions used in the Black-Scholes option pricing model are inherently subjective and involve significant judgment. Changes in our judgments could have had a material impact on our results of operations and financial position. Any change in fair value was recognized as a component of other income (expense) on the statements of comprehensive income (loss).

***Income Taxes***

We use the liability method of accounting for income taxes. Deferred tax assets and liabilities are recognized for the future tax consequences attributable to the differences between the financial statement carrying amounts of existing assets and liabilities and their respective tax basis. Deferred tax assets and liabilities are measured using enacted tax rates expected to be in effect when such assets and liabilities are recovered or settled. The effect on deferred tax assets and liabilities of a change in tax rates is recognized in the year that includes the enactment date. We determine deferred tax assets including net operating losses and liabilities, based on temporary differences between the book and tax bases of assets and liabilities. We believe that it is currently more likely than not that our deferred tax assets will not be realized, and as such, a full valuation allowance is required.

We utilize a two-step approach for evaluating uncertain tax positions. Step one, recognition, requires us to determine if the weight of available evidence indicates that a tax position is more likely than not to be sustained upon audit, including resolution of related appeals or litigation processes, if any. If a tax position is not considered more likely than not to be sustained, no benefits of the position are recognized. If we determine that a position is more likely than not to be sustained, then we proceed to step two, measurement, which is based on the largest amount of benefit which is more likely than not to be realized on effective settlement. This process involves estimating our actual current tax exposure, including assessing the risks associated with tax audits, together with assessing temporary differences resulting from the different treatment of items for tax and financial reporting purposes. If actual results differ from our estimates, our net operating loss and credit carryforwards could be materially impacted.

At December 31, 2012, we had federal net operating loss carryforwards, or NOLs, of approximately \$62.2 million and federal research and experimentation credit carryforwards of approximately \$1.0 million, which may be used to reduce future taxable income or offset income taxes due. These NOLs and credit carryforwards expire beginning in 2023 through 2032.

Our realization of the benefits of the NOLs and credit carryforwards is dependent on sufficient taxable income in future fiscal years. We have established a valuation allowance against the carrying value of our deferred tax assets, as it is not currently more likely than not that we will be able to realize these deferred tax assets. In addition, utilization of NOLs and credits to offset future income subject to taxes may be subject to substantial annual limitations due to the change in ownership provisions of the Internal Revenue Code of 1986, or the Code, and similar state provisions. We may have already experienced one or more ownership changes. Depending on the timing of any future utilization of our carryforwards, we may be limited as to the amount that can be utilized each year as a result of such previous ownership changes. Future changes in our stock ownership, including this offering or future offerings, as well as other changes that may be outside our control could potentially result in further limitations on our ability to utilize our net operating loss and tax credit carryforwards.

## **Table of Contents**

We do not anticipate that the amount of our existing unrecognized tax benefits will significantly increase or decrease within the next 12 months. Due to the presence of NOLs in most jurisdictions, our tax years remain open for examination by taxing authorities back to the inception of the company.

### **Recent Accounting Pronouncements**

We have reviewed recent accounting pronouncements and concluded that they are either not applicable to our business, or that no material effect is expected on the consolidated financial statements as a result of future adoption.

As an emerging growth company the JOBS Act allows us to delay adoption of new or revised accounting pronouncements applicable to public companies until such pronouncements are made applicable to private companies. As a result, our financial statements may not be comparable to the financial statements of issuers who are required to comply with the effective dates for new or revised accounting standards that are applicable to public companies.

### **Off-Balance Sheet Arrangements**

We do not have any relationships with unconsolidated entities or financial partnerships, such as entities often referred to as structured finance or special purpose entities, which would have been established for the purpose of facilitating off-balance sheet arrangements or for any other contractually narrow or limited purpose.

### **Inflation**

We do not believe that inflation has had a material effect on our business, financial condition or results of operations. If our costs were to become subject to significant inflationary pressures, we may not be able to fully offset such higher costs through price increases. Our inability or failure to do so could adversely affect our business, financial condition and results of operations.

### **Quantitative and Qualitative Disclosures about Market Risk**

We are exposed to various market risks, including changes in commodity prices and interest rates. Market risk is the potential loss arising from adverse changes in market rates and prices. Prices for our products are largely denominated in U.S. dollars and, as a result, we do not face significant risk with respect to foreign currency exchange rates.

#### ***Interest Rate Risk***

Generally, our exposure to market risk has been primarily limited to interest income sensitivity, which is affected by changes in the general level of U.S. interest rates, particularly because the majority of our investments are in short-term debt securities. The primary objective of our investment activities is to preserve principal while at the same time maximizing the income we receive without significantly increasing risk. To minimize risk, we maintain our portfolio of cash, cash equivalents and short-term investments in a variety of interest-bearing instruments, which have included U.S. government and agency securities, high-grade U.S. corporate bonds and money market funds. Declines in interest rates, however, would reduce future investment income. A 1% decline in interest rates, occurring on October 1, 2013 and sustained throughout the period ended September 30, 2014, would not be material.

As of September 30, 2013, the principal and accrued interest outstanding under our term borrowings was \$18.3 million. The interest rates on our term borrowings under our credit facility are fixed. If overall interest rates had increased by 10% during the periods presented, our interest expense would not have been affected.

**Table of Contents**

***Foreign Currency Exchange Risk***

As we expand internationally our results of operations and cash flows will become increasingly subject to fluctuations due to changes in foreign currency exchange rates. Historically, a majority of our revenue has been denominated in U.S. dollars, although we sell our products and services in local currency outside of the United States, principally the Euro. Our expenses are generally denominated in the currencies in which our operations are located, which is primarily in the United States. The effect of a 10% adverse change in exchange rates on foreign denominated cash, receivables and payables would not have been material for the periods presented. As our operations in countries outside of the United States grow, our results of operations and cash flows will be subject to fluctuations due to changes in foreign currency exchange rates, which could harm our business in the future. To date, we have not entered into any material foreign currency hedging contracts although we may do so in the future.

---

**Table of Contents**

**BUSINESS**

**Overview**

We develop, manufacture and sell robust, intuitive products that unlock scientifically valuable and clinically actionable genomic information from minute amounts of tissue. Our nCounter Analysis System directly profiles hundreds of molecules simultaneously using a novel barcoding technology that is powerful enough for use in research, yet simple enough for use in clinical laboratories worldwide. We market systems and related consumables to researchers in academic, government, and biopharmaceutical laboratories for use in understanding fundamental biology and the molecular basis of disease and to clinical laboratories and medical centers for diagnostic use. We have an installed base of more than 180 systems, which our customers have used to publish more than 360 peer-reviewed papers. As researchers discover how genomic information can be used to improve clinical decision-making, these discoveries can be translated and validated as diagnostic tests based on our nCounter Elements reagents. In certain situations, we intend to translate their discoveries into *in vitro* diagnostic assays. In September 2013, we received 510(k) clearance from the FDA to market in the United States a version of Prosigna providing an assessment of a patient's risk of recurrence for breast cancer. In November 2013, we commercially launched the nCounter diagnostic system in the United States, including a dual-mode configuration that can be used both for Prosigna and for research applications. In December 2013, we commercially launched Prosigna in the United States and announced that national diagnostic laboratories ARUP Laboratories, Laboratory Corporation of America Holdings and Quest Diagnostics have chosen to add Prosigna to their suites of breast cancer diagnostic tests. We expect Prosigna testing services to become available in the United States in the first quarter of 2014. In September 2012, we obtained CE Mark in the European Union for our first molecular diagnostic product, the Prosigna Breast Cancer Assay, or Prosigna, an assay providing an assessment of a patient's risk of recurrence for breast cancer and the intrinsic subtype of the patient's tumor. In February 2013, we commercially launched Prosigna in Europe and Israel.

The role of genomic information in research and medical practice is evolving rapidly. The advent of new technologies that sequence and digitally count discrete nucleic acids, commonly referred to as next generation sequencing, or NGS, is accelerating the discovery of the relationships between the genome and human disease. Researchers are applying this wealth of new information to identify biological pathways, which are networks of tens or hundreds of genes that act in concert to produce biological functions. Researchers then seek to translate this understanding of the genomic basis of disease into the development of diagnostic tools that can be used to profile an individual patient's biological pathways as well as develop targeted drug therapies. Precise, simple and robust profiling of biological pathways presents both an analytical challenge for researchers and an opportunity to improve patient outcomes in the future.

Our nCounter Analysis System enables genomic analysis on a scale appropriate for pathway-based biology by digitally quantifying the activity of up to 800 genes simultaneously in a single minute tissue sample. The sensitivity and precision of our novel barcoding chemistry allows the measurement of subtle changes in genomic activity efficiently, which is essential in both research and diagnostics because tissue samples are often available only in very small quantities. This problem is especially acute in cancer research, which is typically conducted using biopsies that are often stored in a format known as formalin-fixed paraffin embedded, or FFPE, which complicates subsequent analysis of genetic material. The nCounter Analysis System is an easy-to-use and flexible solution that allows researchers to efficiently test hypotheses across thousands of different samples. As a result, the nCounter Analysis System is particularly useful for discovering and validating networks of genes that characterize and help predict disease states, enabling the development of diagnostics and medicines designed specifically for treating patients with certain genomic profiles. Researchers may use nCounter to develop their own diagnostic tests based on our nCounter Elements reagents or we may selectively partner with them to translate their discoveries into *in vitro* diagnostic assays.

Prosigna, our first molecular diagnostic test, is based on a collection of 50 genes known as the PAM50 gene signature, which was discovered by several of our life sciences customers. We secured an exclusive worldwide license to the PAM50 gene signature in 2010. Prosigna can provide a breast cancer patient and her physician with a subtype classification based on the fundamental biology of the patient's tumor, as well as a prognostic score that predicts the probability of cancer recurrence over 10 years. Our goal is for physicians to use Prosigna to guide therapeutic decisions

---

**Table of Contents**

so that patients receive only therapeutic interventions from which they are likely to benefit. In 2011, we conducted a clinical study, which we refer to as our TransATAC study, based on material extracted from tumor samples of more than 1,000 evaluable patients from the ATAC study. In our study, investigators performed Prosigna on these samples that had been previously analyzed using Genomic Health's *Oncotype DX*, the historical market leader in breast cancer recurrence testing. The results of our TransATAC study demonstrated the ability of Prosigna to indicate risk of recurrence in postmenopausal women with hormone receptor-positive early stage breast cancer treated with endocrine therapy alone. In comparing the risk estimate provided by Prosigna to the risk estimate previously generated using *Oncotype DX*, investigators concluded that Prosigna is capable of providing more prognostic information than *Oncotype DX*. Based on the results of this study and multi-site analytical validation studies, we received European Union, or EU, regulatory clearance for Prosigna, known as a CE mark. As part of our preparation for regulatory submission in the United States, we conducted a second clinical study, which we refer to as our ABCSG8 study, based on tumor samples of more than 1,400 evaluable patients from the Austrian Breast & Colorectal Cancer Study Group 8. Our ABCSG8 study confirmed the conclusion that Prosigna can indicate risk of recurrence as previously demonstrated in our TransATAC study. In September 2013, we received 510(k) clearance from the FDA to market in the United States a version of Prosigna providing an assessment of a patient's risk of recurrence for breast cancer. In November 2013, we commercially launched the nCounter Dx Analysis System in the United States. In December 2013, we commercially launched Prosigna in the United States. National diagnostic laboratories ARUP Laboratories, Laboratory Corporation of America Holdings and Quest Diagnostics have chosen to add Prosigna to their suites of breast cancer diagnostic tests, and the laboratories at the University of Alabama at Birmingham Comprehensive Cancer Center and University of North Carolina Lineberger Comprehensive Cancer Center will be among the initial facilities to offer the Prosigna assay in the United States, with the earliest testing beginning during the first quarter of 2014. These laboratories collectively serve the pathology testing needs of a substantial portion of breast cancer patients throughout the United States. We expect additional clinical laboratories to adopt Prosigna in the future. We intend to conduct future clinical studies to evaluate Prosigna's ability to guide physicians and patients in making additional treatment decisions, which may include the selection of the appropriate chemotherapy regimen, the duration of adjuvant endocrine therapy, and whether to use adjuvant radiation therapy, and, if such studies are successfully completed, to seek 510(k) clearance or PMA approval in the United States for such indications in the future.

In November 2013, we began offering a version of the nCounter Dx Analysis System to high-complexity, CLIA-certified laboratories for research and diagnostics purposes. This FLEX configuration of the nCounter Dx Analysis System provides clinical laboratories a single platform with the flexibility to support both clinical testing, by running Prosigna, and research, by processing translational research experiments using our custom CodeSets and panels. The nCounter Elements GPRs provide further flexibility by allowing laboratories to develop their own Laboratory Developed Tests for gene expression, copy number variation and gene fusion signatures, which can be performed by a laboratory and may include genetic tests and other tests for rare conditions.

Prosigna is regulated as an *in vitro* diagnostic test and we distribute it as a kit for use on our nCounter Analysis System in clinical laboratories. We expect that our future *in vitro* diagnostic products will be regulated and distributed in a similar manner. This is in contrast to most complex genomic tests, which are currently regulated as services and are usually offered only by a limited number of specialized laboratories. The current centralized laboratory model for complex genomic testing can result in complicated logistics for the treating physician, including slower test result turnaround times and limited international access to tests as compared to local testing. In addition, most clinical laboratories cannot currently share in the revenue associated with offering patients complex genomic tests. We believe that our decentralized model will transform the current paradigm of complex genomic testing by allowing physicians worldwide to provide more comprehensive personalized diagnoses, broadening patient access, and increasing the degree to which clinical laboratories can profit by providing molecular diagnostic testing services.

We generated revenue of \$11.7 million, \$17.8 million and \$23.0 million in 2010, 2011 and 2012, respectively, and \$21.3 million in the nine months ended September 30, 2013, while incurring net losses of \$12.8 million, \$10.9 million and \$17.7 million in 2010, 2011 and 2012, respectively, and \$20.5 million in the nine months ended September 30, 2013.

## **Table of Contents**

### **Our Market Opportunity**

Every living organism has a genome that contains the full set of biological instructions required to build and maintain life. By analyzing the variations in genomes, genes and gene activity in and between organisms, researchers can determine their functions and roles in health and disease. An improved understanding of the genome and its functions allows researchers to drive advancements in scientific discovery. As they make scientific discoveries, researchers have been able to translate some of these findings into clinical applications that improve patient care.

A gene is a specific set of instructions embedded in the DNA of a cell. For a gene to be turned on, or expressed, the cell must first transcribe a copy of its DNA sequence into molecules of messenger RNA. Then, the cell translates the expressed information contained in the RNA into proteins that control most biological processes. In addition to the translated RNAs, there are many types of non-coding RNAs that are involved in many cellular processes and the control of gene expression, including microRNA, or miRNA, and long noncoding RNA, or lncRNA.

Biological pathways are the networks of tens or hundreds of genes that work in concert to produce a biological function. Understanding the activation state of pathways and disruptions in individual elements of these pathways provides significant insight into the fundamental basis of disease and facilitates data driven treatment decisions. Therapeutic interventions, such as drugs, can be used to treat disease by activating or inactivating biological pathways that are relevant to disease. As a result, pathway-based biology has become a widely adopted paradigm that researchers use to understand biological processes and has assisted them in the development of diagnostics and drugs to treat disease. To be successful in their research, these scientists need the ability to precisely and simultaneously measure the activation state of the tens or hundreds of genes that comprise biological pathways.

Over the last decade, methods of measuring genomic information have advanced substantially. However pathway-based research and the development of diagnostic tests require analysis of multiple genes and sensitivity to small changes in expression, which can be challenging for traditional genomic tools. In general DNA microarrays and tube-based qPCR methods, require complex, time consuming workflows and relatively large amounts of sample tissue to accurately characterize biological pathway activation. In both life sciences research and clinical medicine, there is a growing need for improved technologies that can precisely and rapidly measure the activation state of hundreds of genes simultaneously across a large number of precious samples, thereby providing a simple and reliable means to characterize biological pathways within minute tissue specimens.

### ***Existing Technologies***

#### ***Microarrays***

Microarrays are tools used to measure gene expression based on the relative brightness of a fluorescent dot on a glass, silicon or other semiconductor surface. Microarrays are sometimes referred to as an analog technology because an indirect measurement of the brightness level, rather than a direct measurement of the molecule, is used to infer the level of gene expression. Traditional microarrays are capable of cost effectively analyzing a large number of targets on a small number of samples, however they have limited sensitivity, precision and dynamic range. In addition, the relatively low throughput and complex workflows of microarrays makes them challenging to use in diagnostic applications.

#### ***Next Generation Sequencing***

Next generation sequencing, or NGS, is a technology that rapidly sequences nucleic acids, and then uses a computationally intense process to count discrete copies of each nucleic acid. While the primary application of this technology is to decode the sequence of DNA, researchers can also use NGS to quantify gene expression using an application called RNA sequencing, or RNA-Seq, which counts the number of times a specific gene sequence is detected in a sample. This is referred to as a digital technology because it relies on a direct count of molecules, rather than an indirect analog measurement of the level of gene expression. This digital gene expression analysis technology can provide higher sensitivity, precision, and dynamic range than traditional



## **Table of Contents**

analog gene expression tools, such as microarrays. As a result, RNA-Seq is replacing microarrays as the platform of choice for genome-wide expression analysis in research. In recent years, many innovations have been made to improve the precision, workflow and automation of these systems. However, RNA-Seq is not well suited for diagnostic use in most clinical laboratories due to its complexity, computational intensity, limited throughput and need for expert technicians. In addition, it is challenging to perform RNA-Seq analyses with small amounts of tissue, especially FFPE samples, which is a limiting factor in large scale studies and many clinical applications.

### *Real-time or Quantitative Polymerase Chain Reaction*

Quantitative polymerase chain reaction, or qPCR, is a technique that can measure gene expression usually on a single target in a particular cell or tissue. First, an enzyme is used to create a complementary DNA, or cDNA, of the RNA target to be measured. Then polymerase chain reaction, or PCR, is used to amplify the cDNA copy through the use of enzymes and repeated heating and cooling cycles, with fluorescent dyes being incorporated during each amplification cycle. Finally, the expression of the gene of interest is inferred based on the number of amplification cycles required for the fluorescent amplified target to become detectable. qPCR is sometimes referred to as an analog technology because the number of cycles of amplification, rather than a direct measure, is used to infer the level of gene expression. The wide availability of qPCR chemistry makes it a popular approach for measuring the expression of a single gene but expanding this method to the analysis of multiple hundreds of genes requires complex liquid handling automation and process optimization. Recently, researchers have used microfluidic approaches that divide samples across multiple parallel qPCR reactions, each measuring a single gene, in order to profile multiple genes in parallel. This approach requires additional liquid-handling steps and can result in less efficient use of tissue samples. Finally, the use of enzymes in numerous cycles of amplification can introduce distortion and bias into the data, depending on the process controls and expertise of the person conducting the experiment.

### *Life Sciences Research*

According to Strategic Directions International, Inc., life sciences researchers spent approximately \$28 billion on tools and related consumables in 2011. In the decade since the completion of the Human Genome Project, improvements in NGS technology have greatly reduced the cost of sequencing a human genome and increased throughput and precision, which has led to an abundance of new biological information. In order to gather insights from this information, researchers must first distill and then efficiently analyze large pools of data. Gene expression analysis has emerged as a primary tool that researchers use to extract meaningful insights from networks of genes, which enables them to validate and then translate their findings into the development of diagnostics and medicines. According to Percepta Associates, a provider of consulting services to bioscience companies, the 2012 global market for gene expression profiling products is estimated to be \$1.2 billion.

Academic, government, and biopharmaceutical researchers engaged in gene expression analysis typically focus on making biological discoveries that may lead to the development of relevant medical products and better informed treatment decisions for physicians and patients. They have traditionally performed these experiments using microarrays or qPCR. Recently, RNA-Seq has dramatically enhanced researchers' ability to discover patterns of gene expression that have biological meaning. However, researchers are increasingly performing analyses on a larger number of genes and samples and are seeking new methods of interrogation that would allow them to:

increase the number of genes that can be analyzed simultaneously in order to understand the complete biological pathway involving multiple genes;

improve the overall efficiency of their laboratories by simplifying workflow and accelerating the rate of successfully completing their research;

provide more reliable, precise and reproducible data about targeted genes and biological pathways;

maximize the amount of genomic information extracted from precious tissue samples;

minimize the computational intensity of complex genomic analysis;

process difficult-to-work-with specimens, such as tumor biopsies stored in FFPE format; and

## **Table of Contents**

create more systematic and reliable ways to help transition their research discoveries into future clinical products.

We believe that the above items create an opportunity for technologies that are optimized for pathway-based biology. Based on 2011 market data regarding the installed base of microarray systems at that technology's peak, we estimate that the potential market opportunity of our current generation of nCounter Analysis System is approximately 3,000 systems.

### ***Molecular Diagnostics***

As researchers have discovered links between specific genomic variants and diseases, their discoveries have led to the development of genome-based diagnostics that have the potential to inform therapeutic interventions, predict the risk or onset of disease and detect residual disease on an individualized basis. As a result, these molecular diagnostic tests have made individualized treatment possible and have an increasingly important influence on decisions that physicians and patients make regarding treatment and care.

According to Frost and Sullivan, the molecular diagnostics market totaled approximately \$4.1 billion in 2010 and is expected to reach \$6.2 billion by 2014. Growth in the molecular diagnostics market has been driven by technological innovations that have increased sensitivity, decreased turnaround times, simplified workflow, and lowered costs when compared to other techniques. In addition, the medical community has seen a trend in favor of decentralized diagnostic testing as a result of the convenience of local testing, hospitals and medical centers increasingly viewing their laboratories as profit centers and a need to increase access to tests for patients outside of the United States. We believe that there is an opportunity to improve the quality of diagnosis and treatment of diseases by developing and commercializing comprehensive, simple and widely available diagnostic products based on gene expression analysis.

Cancer is a disease generally caused by genetic mutations in cells. The behavior of cancer cells is extremely complex, depending on many different genes and the interactions of those genes. It is often impossible for researchers to identify a single gene that adequately signals a more aggressive or less aggressive type of cancer. However, in some cases, researchers have been able to identify more aggressive or less aggressive types of cancer through gene expression analysis of biological pathways. Multi-gene expression analysis has the potential to considerably improve the decisions of oncologists as they care for their patients. Based on the pattern of gene expression, oncologists can determine which specific treatments are most likely to be effective for an individual patient, monitor a patient's response to those treatments, and determine the likelihood of recurrence.

Enhanced understanding of the genomic basis of disease has driven the development of molecular diagnostics for a number of indications. This trend is exemplified by the development of multi-gene tests for breast cancer. Over the last decade, genomic tests for breast cancer have improved the accuracy of prognosis and efficacy of treatment by providing risk assessments particular to individual patients. As a result of individualized risk profiling, thousands of patients have been spared unnecessary treatment while many others have been placed on more appropriate treatment regimens.

These multi-gene breast cancer tests are widely available in the United States, but are not generally available in other countries, despite the fact that the United States accounts for less than 15% of the global annual

## Table of Contents

breast cancer incidence. The following provides a summary of the incidence of breast cancer worldwide in 2008 based on data published by the World Health Organization:

<sup>(1)</sup>Ferlay J, Shin HR, Bray F, Forman D, Mathers C and Parkin DM. GLOBOCAN 2008 v2.0, Cancer Incidence and Mortality Worldwide: IARC CancerBase No. 10 [Internet]. Lyon, France: International Agency for Research on Cancer; 2010. Available from: <http://globocan.iarc.fr>, accessed on 03/11/2012.

Multi-gene molecular diagnostic tests for breast cancer are provided by several companies today, including Genomic Health, Agendia and Clariant (a GE Healthcare company). These tests are offered as services with the analysis conducted at company-owned centralized laboratories. When a physician orders the test, the pathologist sends a tumor block or thin sections from the biopsy specimen to the centralized laboratory for analysis. The lab operator then uses a multi-gene panel to determine a risk category for each patient, which predicts that individual's likelihood of recurrence. The lab operator typically analyzes the tumor tissue and delivers results to the treating physician within 10 to 14 days of receipt of the tumor sample.

Despite their positive impact on patient care, existing molecular diagnostic tests for breast cancer available prior to Prosigna, which we call first-generation tests, have several limitations, including:

*High Frequency of Intermediate Risk Results.* The first-generation tests generally sort patients into two or three risk categories, such as high/intermediate/low risk. In clinical practice, patients who are low risk are generally spared chemotherapy and patients who are high risk generally receive chemotherapy. In contrast, there is no standard of care for how intermediate risk patients should be treated. The ambiguity of treatment and outcomes associated with an intermediate risk score is challenging for physicians and unsettling for patients.

*Available Exclusively from Specialized Laboratories.* The first-generation tests are based on techniques such as qPCR, microarrays or multiplexed immunohistochemistry and are generally administered by highly-skilled technicians usually working in centralized laboratories. This centralized model may result in complicated logistics for the treating physician, including slow test result turnaround times, added shipping and logistics costs and limited international access to tests. In addition, the economic value of providing the centralized genomic tests that improve clinical decision-making has been captured by a small number of specialized laboratories. Most clinical laboratories are unable to perform or profit from these tests. In addition, because most of the specialized laboratories are in the United States, patients living outside of the United States may be challenged to gain access to these genomic tests.

*Unclear Utility in Other Treatment Decisions.* The first-generation tests were specifically designed to provide a physician with an understanding of the residual risk of recurrence for patients receiving

## **Table of Contents**

hormonal therapy, thereby informing the physician's decision whether to also administer adjuvant chemotherapy. The genes included in the first-generation tests were selected primarily, and in many cases exclusively, based on the correlation of their expression with the risk of cancer recurrence in such patients. It remains unclear as to whether the specific genes included in these first-generation tests will be able to inform other important treatment decisions, such as the selection of specific adjuvant chemotherapy regimens, the duration of adjuvant endocrine therapy, or the decision of whether to administer adjuvant radiation therapy.

In contrast to the central laboratory-based first-generation molecular diagnostic test for breast cancer, the medical community has seen a trend in favor of decentralized diagnostic testing. Tests for HIV, Hepatitis C, Influenza and MRSA, which were once centralized, are now often conducted in hospital laboratories or at the point of care. We believe that this trend of decentralized testing will continue as a result of many factors, including:

*Convenience.* We believe that physicians would prefer that molecular diagnostic tests be performed at a local level and in the same laboratory that performs other tests that the physicians may order. Local molecular diagnostic testing could provide physicians the same rapid turnaround of test results that they have learned to expect for other types of tests.

*Economic Advantages.* We believe that hospitals and medical centers desire to make their clinical laboratories profit centers by performing tests and billing third-party payors. As diagnostic technologies become less complicated to administer, hospitals and medical centers tend to favor in-sourcing tests.

*International Availability.* There is a critical need to increase access to molecular diagnostic tests for patients that live outside the United States. Currently, patients living outside the United States may be challenged to gain access to tests that are provided only by specialized laboratories located within the United States. We believe genomic testing will become more available to patients throughout the world when it can be provided by their local clinical laboratories.

We believe that the market for complex molecular diagnostics will require increased precision, increased breadth of decision making information, and a decentralized approach that is in line with other applications of diagnostic testing.

## **Our Solution**

Our nCounter Analysis System is an automated, multi-application, digital detection and counting system which directly profiles hundreds of molecules simultaneously using a novel barcoding technology that is powerful enough for use in research, yet simple enough for use in clinical laboratories worldwide. Our nCounter Analysis System consists of two automated instruments that prepare and analyze tissue samples using proprietary reagents, which can only be obtained from us. Our life sciences research customers purchase instruments from us and then purchase our panels, custom CodeSets, nCounter Elements reagents and related consumables for the specific experiment or assay they wish to conduct. Our diagnostics customers will either purchase or lease

## **Table of Contents**

instruments from us and also generally purchase nCounter Elements reagents or our diagnostic kits for tests that they intend to run.

Our nCounter Analysis System offers a number of compelling advantages, including:

*Optimized for Pathway-Based Biology.* The nCounter Analysis System can profile up to 800 molecules in a single test tube, which allows customers to analyze interactions among hundreds of genes that mediate biological pathways.

*Digital Precision.* Our molecular barcodes hybridize directly to the target molecules in a sample allowing them to be counted. This generates digital data (1 molecule = 1 count) of excellent quality over a wide dynamic range of measurements and provides excellent reproducibility.

*Simple Workflow.* The nCounter Analysis System's minimal sample preparation and automated workflow enable the performance of gene expression analysis across hundreds of genes simultaneously in approximately 24 hours between the time a sample is loaded into the system and results are obtained. Our nCounter Analysis System generates data that customers can evaluate without the use of complex bioinformatics.

*Flexible Sample Requirements.* The nCounter Analysis System is able to unlock genomic information from minute amounts of a variety of challenging tissue samples, including FFPE samples, cell lysates and single cells.

*Versatility.* The FLEX configuration of the nCounter Dx Analysis System provides clinical laboratories a single platform with the flexibility to support both clinical testing, by running Prosigna, and research, by processing translational research experiments and multiplexed assays using our custom CodeSets and panels. The nCounter Elements GPRs provide further flexibility by enabling laboratories to develop their own Laboratory Developed Tests for gene expression, copy number variation and gene fusion signatures.

Our nCounter Analysis System enables research from basic discovery to the development and commercialization of molecular diagnostic tests on a single platform. We believe that our nCounter Analysis System is complementary to and synergistic with digital gene expression on next generation sequencers (using RNA-Seq).

Many of our research customers are transitioning their efforts to discover important patterns of gene expression away from the analog technology of microarrays to the digital technology of NGS, using the application RNA-Seq.

## **Table of Contents**

Often, these customers perform follow-up experiments to validate the results of their RNA-Seq discovery experiments by testing hundreds or thousands of samples. Increasingly, researchers are performing these validation experiments using our nCounter Analysis System, which combines the precision of digital gene expression with the high throughput and simple workflow required to efficiently process large numbers of samples. Because our nCounter Analysis System can directly measure RNA without the use of enzymes or amplification (except for single-cell applications), we believe that researchers view the nCounter Analysis System as providing an independent means of validating that observations made initially using RNA-Seq are real rather than artifacts associated with the complex series of steps necessary for sequencing. In addition, because the operation of our nCounter Analysis System is simple and intuitive, it provides a practical technology enabling the translation of potential biomarkers into diagnostic tests that can be performed in the clinical laboratory, after appropriate regulatory authorization.

The figure below illustrates the current and future uses of and opportunities for digital technology in translational genomics.

### ***Life Sciences Research***

The nCounter Analysis System enables our life sciences research customers to conduct research on a scale that is well suited for pathway-based biology. The precision, ease of use and flexibility of our nCounter Analysis System allows researchers to efficiently test their hypotheses in thousands of different samples and is particularly useful for identifying networks of genes that characterize and predict disease.

## **Table of Contents**

### *Life Sciences Applications*

Our nCounter Analysis System is capable of supporting a number of life sciences applications based upon the measurement of the concentration or amount of a target nucleic acid. Key applications currently supported include:

*Gene Expression.* Researchers use the nCounter Analysis System to measure the degree to which individual genes in pathways are turned on or off by simultaneously quantifying the amount of messenger RNA, or mRNA, associated with each of up to 800 genes.

*Single Cell Gene Expression.* Historically, most gene-expression profiling has been performed on populations of cells where observed expression levels represent an average of the unique expression states of each cell within the population. The nCounter Analysis System is capable of measuring gene expression of 20 to 800 genes from a single cell, thereby elucidating previously hidden relationships between individual cells within a population.

*miRNA Expression.* Researchers can use the nCounter Analysis System to measure the simultaneous expression levels of up to 800 different miRNAs. The nCounter Analysis System is capable of highly multiplexed, direct digital detection and counting of miRNAs in a single reaction without amplification, thereby delivering high levels of sensitivity, specificity, precision, and linearity. We currently enable miRNA experiments for use in tissue from humans, mice, rats, and fruit flies.

*Copy Number Variation.* Researchers can use the nCounter Analysis System to probe for structural variations that result in cells having an abnormal number of copies of one or more sections of the DNA. Researchers are able to conduct large-scale, statistically-powered studies of these copy number variations, or CNVs, by leveraging the nCounter Analysis System's multiplexing capacity to assay up to 800 DNA regions in a single tube, with as little as 300 ng of DNA.

We also support research directed toward particular gene fusions, gene-expression regulatory elements called long non-coding RNA, or lncRNA, and experiments based on a technique used to investigate the DNA targets of transcription factors called chromatin immunoprecipitation, or ChIP.

Our customers have used the nCounter Analysis System to publish more than 360 peer-reviewed papers. In 2013 alone, our customers published more than 180 peer-reviewed papers incorporating data generated using the nCounter Analysis System. The most frequent topic of nCounter-based peer-reviewed publications is cancer research, including biomarker discovery and validation. Other frequent topics include immunology and inflammation, infectious disease and developmental and cell biology.

### *Consumables*

Following their purchase of our nCounter Analysis System, our life sciences research customers purchase consumables from us consisting of CodeSets and other consumables that are designed for the specific experiment that they intend to run. Our instruments are designed to be used only with our consumables. This closed system model generates recurring revenue from each instrument we sell. We believe that our recurring consumable revenue is driven by our customers' ability to extract value from up to 800 data points per sample and to process hundreds of samples in a relatively short period of time with little hands-on preparation using our nCounter Analysis System, enabling them to process more units of consumables per unit of time.

### *Molecular Diagnostics*

We believe that the attributes that make the nCounter Analysis System attractive to researchers also have the potential to make the system attractive to hospitals and clinical laboratories that desire to conduct molecular diagnostic tests. The precision, ease of use and flexibility of the nCounter Analysis System will allow medical technicians in pathology labs to conduct complex molecular diagnostic tests with minimal training. We expect these tests to encompass both Laboratory Developed Tests based on our nCounter Elements reagents and *in vitro* diagnostic kits, initially Prosigna.





---

## **Table of Contents**

Prosigna is designed to address the limitations of first-generation tests, including:

*Fewer Intermediate Risk Patients in Node-Negative Disease.* In our TransATAC study, Prosigna was performed on material extracted from tumor samples from more than 1,000 evaluable patients that were previously analyzed using Genomic Health's Oncotype DX, a widely-used first-generation test that is offered as a laboratory-developed test. Each patient in the study was assigned to a risk group based on risk estimates generated separately by Prosigna and Oncotype DX, and using prospectively defined risk cutoffs. Cutoffs for low, intermediate and high were <10%, 10% to 20% and >20% estimated risk of recurrence, respectively. In a comparison of the sizes of the risk groups in patients with node-negative disease, Prosigna assigned 26% fewer patients to an intermediate score than Oncotype DX. The reduction in the size of the intermediate risk group in node-negative patients is a result primarily of Prosigna's ability to reclassify patients that are classified by Oncotype DX as intermediate risk to high risk. We believe this study provides evidence of Prosigna's potential ability to clarify treatment decisions.

*Available on a Decentralized Basis.* Prosigna will be available for use on the nCounter Analysis System on a distributed basis in the clinical laboratories of hospitals and medical centers worldwide, which aligns Prosigna with the evolving trend towards decentralized testing. We believe that by distributing molecular diagnostics to local labs, we will provide faster turnaround for patients and enable succinct, comprehensive reports from pathologists, resulting in enhanced patient care. We also believe that this model will increase the degree to which clinical laboratories can profit by providing molecular diagnostic testing services. In addition, our decentralized model can help address the needs of patients outside of the United States by enabling local laboratories to provide testing.

*Potentially More Treatment Decisions.* Prosigna measures the expression of up to 50 genes, providing a more detailed profile of the biology of a patient's tumor than the first-generation tests. In addition, Prosigna utilizes the concept of intrinsic subtypes, a fundamental method of classifying breast tumors into the four distinct subtypes of that disease. By providing a more detailed view of tumor biology and determining the intrinsic subtype of breast cancer patients, Prosigna has the potential to inform not only the decision of whether to administer adjuvant chemotherapy, but also potentially inform other important treatment decisions. These decisions may include the selection of specific adjuvant chemotherapy for individual patients, the duration of adjuvant endocrine therapy, and the use of adjuvant radiation therapy. We intend to perform clinical studies validating Prosigna's ability to inform additional treatment decisions, and to seek a PMA to enable Prosigna to report these intrinsic subtypes for use in informing clinical decisions within the United States.

In November 2013, we began offering a version of the nCounter Dx Analysis System to high-complexity, CLIA-certified laboratories for research and diagnostics purposes. This FLEX configuration of the nCounter Dx Analysis System provides clinical laboratories a single platform with the flexibility to support both clinical testing, by running Prosigna, and research, by processing translational research experiments using our custom CodeSets and panels. The nCounter Elements GPRs provide further flexibility by allowing laboratories to develop their own Laboratory Developed Tests for gene expression, copy number variation and gene fusion signatures, which can be performed by a laboratory and may include genetic tests and other tests for rare conditions.

We believe that the strengths of our diagnostics platform, which will enable us to commercialize Prosigna on a decentralized basis, could be applied to *in vitro* diagnostic kits for other cancers after securing the requisite regulatory authorizations. Over time, we intend to identify other tests and develop them for use on our nCounter Analysis System.

### **Our Strategy**

Our goal is to provide products that empower scientists to understand the molecular basis of disease and empower physicians to put genomic medicine into practice. To accomplish this goal, we intend to continue

## **Table of Contents**

providing technologies that are powerful enough for research, yet simple and robust enough for use in clinical laboratories worldwide.

Our strategy includes the following key elements:

*Establish the nCounter Analysis System as the Global Standard for Gene Expression Analysis.* We will promote the power of pathway-based biology and the benefits of digitally-quantified gene expression. One element of this strategy is to recognize and publicize the breadth of scientific achievements and peer-reviewed publications in top-tier journals based on our nCounter Analysis System. Since 2009, more than 360 peer-reviewed papers have been published by researchers using the nCounter Analysis System. We plan to also highlight successful examples of using the nCounter Analysis System to translate research discoveries into diagnostic tests, beginning with Prosigna as well as future examples of Laboratory Developed Tests and additional *in vitro* diagnostic kits as they emerge.

*Expand the Installed Base of the nCounter Analysis Systems in Biopharmaceutical and Academic Research.* We will continue to engage with researchers through direct sales efforts in North America and Europe, increasing the installed base of our nCounter Analysis Systems. We will target translational researchers in genome centers, academic medical centers and biopharmaceutical companies. We will continue to focus primarily on cancer researchers, because these researchers recognize the significant value of our technology when analyzing small biopsy samples stored in challenging formats such as FFPE. We also intend to expand our existing geographic reach, both directly and through distributors.

*Broaden the Addressable Market of the nCounter Analysis System through Continued Innovation.* We will continue to invest in product development efforts that increase the capabilities of our nCounter Analysis System and expand the universe of potential customers. We expect that these investments will lead to additional protocols, enhanced chemistries, and new generations of instruments and software. We will prioritize innovations that increase the flexibility of the nCounter Analysis System to process small and degraded samples, and increase the ease and speed with which users can select target genes and design CodeSets for their particular experimental needs. We will also prioritize innovations that are expected to reduce the cost and footprint of the nCounter Analysis System, which will help us to target a broader range of customers.

*Build a Menu of Diagnostic Content in Collaboration with Researchers Comprising Both Proprietary In Vitro Diagnostic Kits and Laboratory Developed Tests Based on nCounter Elements Reagents.* We intend to continue cultivating relationships with leading researchers who are working to establish the connection between genomics and clinical decision-making, many of whom are already using the nCounter Analysis System today in their translational research. When these researchers invent new diagnostic methods, we intend to enable development of Laboratory Developed Tests based on our nCounter Elements reagents. In certain situations, we may gain exclusive access to these clinically valuable gene signatures through licensing and collaboration arrangements. For example, the intellectual property that forms the basis of Prosigna was in-licensed from Bioclassifier, LLC, which was founded by several of our life sciences research customers. In addition, in February 2013, we secured an option from The Broad Institute to acquire an exclusive worldwide license for a gene signature that could be used, after further development, as a Laboratory Developed Test, or, after appropriate regulatory authorization, for a second molecular diagnostic product focused on hepatocellular carcinoma, or HCC. We will seek to enter into similar arrangements with our life sciences research customers and other researchers for future diagnostic gene signatures. Our strategy is to target intellectual property rights to potential diagnostic methods that are well understood, have the potential to facilitate changes in treatment with a major impact on outcome and cost, have the potential to support value-based pricing, and for which tissue samples for clinical validation are readily available. This disciplined approach is designed to efficiently focus our research and development investment on the development of potential products, rather than discovery of new gene signatures. Our initial focus will be on cancer gene signatures with the goal to individualize major treatment decisions so that patients are more likely to receive only those interventions from which they are likely to benefit.

*Execute High Quality Clinical Studies to Support Regulatory Authorizations, Market Adoption and Reimbursement of Diagnostic Products.* For each diagnostic product we intend to develop, we plan to design and execute clinical programs that achieve high standards of clinical evidence. Whenever possible, these programs

## **Table of Contents**

will be designed to achieve Level 1 clinical evidence, the standard generally required for inclusion in clinical practice guidelines. We will take advantage of tissue samples and outcomes from past controlled randomized clinical studies wherever possible. This strategy has been demonstrated in the first two studies for Prosigna where we were able to access the tissue and outcome information from breast cancer drug trials that totaled more than 2,400 evaluable patients.

*Enable Clinical Laboratories Worldwide to Provide Complex Genomic Testing Using Our In Vitro Diagnostic Products.* We intend to enable diagnostic tests in clinical laboratories worldwide by facilitating development of Laboratory Developed Tests based on nCounter Elements reagents and distributing nCounter Analysis System-based *in vitro* diagnostic kits. Our strategy is to drive improved and more efficient clinical decisions while allowing laboratories to profit from the valuable testing services enabled by our technology. By providing the nCounter Analysis Systems to labs that already conduct most of the diagnostic testing and control access to tissue samples, we intend to enable pathologists and labs to provide advanced genomic testing that previously required the shipment of tissue samples to a centralized lab.

*Drive Physician Demand for nCounter Analysis System-Based Diagnostic Products.* While our existing life science focused commercial team will continue to focus on building an installed base of nCounter Analysis Systems in research and clinical laboratories in regions where we have secured the necessary regulatory authorizations, we will continue to establish a commercial organization focused on driving orders for Prosigna tests toward clinical laboratories. This Prosigna sales force will directly target its efforts on the oncologists and pathologists, and will adopt promotional and marketing practices that have been employed successfully by the biopharmaceutical industry to educate treating physicians on the ability of our molecular tests to inform treatment decisions in accordance with the product labeling. We will communicate with patients, oncologists, pathologists, hospital administrators and other key stakeholders to outline the clinical and economic advantages of bringing complex genomic tests in-house.

*Capture Capital Efficiencies Stemming from our Diagnostics Business Model.* We plan to leverage other capabilities we have built to support our life sciences business, including our technology platform and product development, manufacturing, and administrative functions, to build our diagnostics business at minimal incremental cost. We will rely on the clinical laboratory infrastructure, sample logistics, managed care contracting and billing operations of our laboratory customers, further reducing our capital requirement. We intend to coordinate commercial efforts with the sales and marketing personnel of the clinical laboratories offering clinical testing services based on our diagnostic products. We believe that this approach will yield a diagnostics business model that is more capital efficient than a clinical laboratory services model and has the potential to become profitable on a relatively small revenue base.

## **Our Products and Technology**

The fundamental technology employed in our nCounter Analysis System was conceived at the Institute for Systems Biology in the laboratories of Dr. Leroy Hood, a renowned pioneer in genomics and personalized medicine. Our life sciences research customers purchase instruments from us and then purchase our panels or custom CodeSets and related consumables for the specific experiment they wish to conduct. Our diagnostics customers will either purchase or lease instruments from us and also generally purchase nCounter Elements reagents or our diagnostic kits for tests that they intend to run.

### ***nCounter Analysis System***

The nCounter Analysis System is an automated, multi-application, digital detection and counting system consisting of one or more nCounter Prep Stations and one nCounter Digital Analyzer. Since 2008, we have marketed a research use only version of the system, and in 2013 we introduced the nCounter Dx Analysis System to be marketed to clinical laboratories. The nCounter Dx Analysis System comes in two configurations, one that only runs Prosigna and one that is called the FLEX Configuration, a dual-mode system that runs Prosigna in one mode and our life science applications, including nCounter Elements, in the other.

## **Table of Contents**

### *Instruments and Software*

The nCounter Prep Station is the automated liquid handling component of the nCounter Analysis System that processes samples after they are hybridized and prepares the samples for data collection on the nCounter Digital Analyzer. The nCounter Digital Analyzer collects data from samples by taking images of the immobilized fluorescent reporters in the sample cartridge and processing the data into output files, which include the target identifier and related count numbers along with a broad set of internal controls that validate the precision of each assay. The currently available nCounter Prep Station and nCounter Digital Analyzer were designed and are manufactured under ISO 13485:2003, the quality standard for *in vitro* diagnostic platforms and medical devices. We also provide our research customers with the nSolver Analysis Software, a data analysis program that offers researchers the ability to quickly and easily quality check, normalize, and analyze their data without having to use any additional software for data analysis. The diagnostic version of our nCounter Analysis System includes the software that runs Prosigna and generates individualized patient reports.

### *Simple and Rapid NanoString Workflow*

The nCounter Analysis System's simple three step workflow takes approximately 24 hours and requires approximately 15 minutes of hands-on time by the user:

during step 1, up to 12 targeted research tissue samples (or up to 48 samples with sample multiplexing) or up to 10 breast cancer tissue samples in the case of Prosigna, and our reagents or diagnostic kits are injected into strip tubes that we provide and allowed to hybridize overnight;

in step 2, the strip tube is loaded into our nCounter Prep Station, which purifies the mixtures and moves them onto a cartridge with 12 flow-cells where the fluorescent barcodes are captured and affixed onto a glass surface of the cartridge and oriented in one direction; and

in step 3, the cartridge is placed into our nCounter Digital Analyzer, which uses fluorescent microscopy and image analysis software to automatically count the barcodes and provide the level of expression of each target in the sample.

When the nCounter Analysis System is run in life science mode, a user can process up to approximately 36 samples per day by installing one Prep Station with a single Digital Analyzer. One can increase the number of samples analyzed to 108 samples per day on a single Digital Analyzer if it is coupled with three Prep Stations. This throughput can be quadrupled using sample multiplexing for experiments targeting 200 genes or fewer. For Prosigna, a clinical laboratory can process up to 30 samples per day on an nCounter Dx Analysis System.

### ***Life Sciences Research***

Following purchase of our nCounter Analysis System, life sciences research customers purchase panels, custom CodeSets targeted to a specific experiment or nCounter Elements reagents.

#### *Panels*

We offer more than 20 panels that are pre-manufactured and targeted to a specific experiment, including the following:

*Gene Expression Panels.* Preassembled CodeSets that include all of the consumables required to perform the assay on the nCounter Analysis System. We offer nCounter Gene Expression Panels to conduct a wide variety of gene analysis, including analysis of kinase genes, cancer-related human genes, immunology-related genes, and inflammation-related genes.

*miRNA Expression Assay Kits.* A family of panels that provide a cost-effective profiling solution capable of highly multiplexed, direct digital detection and counting of up to 800 miRNAs in a single reaction without amplification. These provide our customers

## Edgar Filing: NanoString Technologies Inc - Form S-1

with high levels of sensitivity, specificity, precision, and linearity. Separate panels are available for use with samples from humans, mice, rats, and fruit flies.

-80-

## **Table of Contents**

*Cancer Copy Number Variation Panel.* Allows researchers to conduct large-scale, copy number variation projects by leveraging the nCounter Analysis System's multiplexing capacity to assay up to 800 regions in a single tube, with as little as 300 ng of starting material.

*nCounter Leukemia Fusion Gene Expression Assay Kit.* A panel that allows researchers to profile a broad set of fusion genes which result from balanced translocations in different leukemia subtypes. In addition to fusion genes, the kit includes probes for 11 wild-type genes involved in translocations and 12 leukemia-related biomarkers.

*Human Karyotype Panel.* Allows for the simultaneous measurement of all 23 pairs of human chromosomes, including 338 individual regions.

### *Custom CodeSets*

We also work with our customers to develop custom CodeSets to enable them to evaluate specific genes that are the subject of their study. Our customers provide us a list of targets for which we subsequently build a unique CodeSet. Our design process leverages full length sequences for the DNA or RNA molecules that our customers are interested in detecting and prevents cross hybridization to non-target molecules in the sample. The custom CodeSet design process occurs in four distinct steps:

### *nCounter Elements*

In July 2013, we launched nCounter Elements, a new digital molecular barcoding chemistry that allows users to design their own customized assays using standard sets of barcodes provided by us with the laboratories' choice of oligonucleotide probes that they can purchase independently from an oligonucleotide manufacturer. Clinical laboratories can use nCounter Elements to create Laboratory Developed Tests, which are diagnostic tests that are developed and performed by a laboratory and include genetic tests and other tests for rare conditions. In addition, the highly flexible architecture of nCounter Elements enables a broad range of basic research studies where iterative design and refinement of assays are important.

nCounter Elements reagents have been registered with the FDA as GPRs and are available for use in developing Laboratory Developed Tests, pursuant to a licensing arrangement.

### *Master Kits*

Our nCounter Master Kit includes all of the ancillary reagents and plasticware required for our customers to be able to setup and process samples in the nCounter Prep Station and nCounter Digital Analyzer. The components of the Master Kit include the sample cartridge, strip tubes, tips, buffers, and reagent plates.

---

## **Table of Contents**

### ***Molecular Diagnostics***

Our nCounter Analysis System's ability to simultaneously quantify gene expression on tens or hundreds of genes from minimal amounts of FFPE tissue make it well suited for profiling pathway activation in tumor samples. In addition, the nCounter Analysis System has the precision, reproducibility, and simple workflow required of technologies used in clinical laboratories.

Following purchase or lease of our nCounter Dx Analysis System, we expect that our clinical laboratory customers will build menus of diagnostic tests comprised of Laboratory Developed Tests, Prosigna and other *in vitro* diagnostic kits we develop and sell in the future. These customers will use the nCounter Dx Analysis System, nCounter Elements reagents and *in vitro* diagnostic kits to provide clinical diagnostic services. Currently, Prosigna is the only *in vitro* diagnostic kit available for use on our nCounter Dx Analysis System. Over time, we intend to develop, obtain regulatory authorization for, and sell additional *in vitro* diagnostic kits, each of which will enable a unique diagnostic test.

### ***Informing Breast Cancer Treatment using the PAM50 Gene Signature***

In 2009, leading cancer researchers first described a new gene expression signature, called PAM50, based on the expression of 50 genes in tumor tissue. PAM50 provides two types of information regarding a breast cancer patient's tumor. First, PAM50 assigns each patient's tumor to one of four intrinsic subtypes, a classification based on the fundamental biology of that individual's breast tumor. Second, PAM50 provides a prognostic risk of recurrence, or ROR, score that assesses the probability that a cancer will recur in the future in patients who will be treated with hormonal therapy.

The intrinsic subtypes of breast cancer were first described in 2000 and have been repeatedly observed across multiple studies and technology platforms. Each patient's breast cancer can be classified into one of four intrinsic subtypes (Luminal A, Luminal B, HER2-enriched, and Basal-like) that describe the fundamental biology of the tumor, conveying valuable information about an individual patient's prognosis and likelihood of response to specific therapies. In June 2011, the widely-recognized St. Gallen International Breast Cancer Treatment Guidelines adopted the intrinsic subtypes as a standard approach to classifying early stage breast cancer, and in general, the basis for systemic therapy recommendations. The PAM50 gene signature represents a molecular approach to intrinsic subtyping and has been described in multiple peer-reviewed publications.

In September 2012 the online edition of the journal Nature published a study of the molecular biology of breast cancer, using the intrinsic subtypes as defined by the PAM50 gene signature as an organizing framework for analyzing genomic and proteomic aberrations. This study, which was an outcome of The Cancer Genome Atlas Initiative and was titled "Comprehensive molecular portraits of human breast tumours," represents a thorough description of breast cancer genomics. The study involved the analysis of tissue from 800 breast cancer tumors by a total of six technology platforms, covering genomics, epigenetics, and proteomics. The research concluded that diverse genetic and epigenetic alterations converge phenotypically into the four main breast cancer subtypes defined by PAM50.

Studies using PAM50 and other methods for assigning intrinsic subtype have suggested that PAM50 may be useful in improving several treatments decisions in breast cancer by:

providing prognostic information that may help physicians and patients decide whether the addition of adjuvant chemotherapy to hormonal therapy is appropriate;

providing prognostic information that may help physicians and patients decide whether extended endocrine therapy is appropriate;

providing information that may help physicians choose which adjuvant chemotherapy regimen to select for an individual patient; and

providing information that may help physicians and patients decide whether adjuvant radiation therapy is appropriate.



**Table of Contents***Development and Validation of the Prosigna Breast Cancer Assay*

In 2010, we acquired an exclusive worldwide license to develop *in vitro* diagnostic and research products for breast cancer based on the PAM50 gene signature. In 2010, we began developing an *in vitro* diagnostic kit based on the PAM50 gene signature, simplifying and optimizing the test for use on the nCounter Analysis System. In 2011, we performed the first in a series of clinical studies designed to validate the test's ability to provide prognostic information for postmenopausal women with HR+ early stage breast cancer treated with endocrine therapy alone using material extracted from tumor samples from 1,017 patients from the TransATAC population of which 1,007 samples passed prespecified criteria and yielded evaluable results. TransATAC is a translational study group that has used the tumor tissue and data from a subset of the 9,366 women enrolled (1996-2000) in the ATAC (Arimidex, Tamoxifen, Alone or in Combination) trial to study the molecular characteristics of tumors in postmenopausal women with HR+ early stage breast cancer of pathological grade 1, 2 or 3. Access to the samples is controlled by a steering committee of the TransATAC study group. In 2010, we submitted a proposal to the steering committee seeking access to the samples and data to validate Prosigna, and our proposal was approved. In 2011, we jointly designed the study with a member of the TransATAC study group who subsequently served as the lead investigator in our study. Following the steering committee's approval of our proposed study, we agreed to cover the costs of the study. The TransATAC population had been previously used in 2008 to clinically validate the current market leader in breast cancer prognosis and prediction, Genomic Health's Oncotype DX, which is a laboratory-developed test that is administered using a centralized laboratory service model. Our study used RNA that had been extracted by Genomic Health from 1,017 tumor samples from patients with postmenopausal HR+ early stage breast cancer during the 2008 study. Because both the Oncotype DX results and the outcomes of the patients associated with each RNA sample were known, the study provided an opportunity to measure both the ability of PAM50 to provide prognostic information, and how the prognostic information provided by Prosigna compares to data previously collected using Oncotype DX. Results of our TransATAC study were presented in December 2011 at the CTRC-AACR San Antonio Breast Cancer Symposium. In July 2013 the TransATAC results were published in the Journal of Clinical Oncology.

In 2012, we performed a second clinical validation study to test the ability of Prosigna to estimate the prognosis of postmenopausal women with HR+ early stage breast cancer treated with endocrine therapy alone that evaluated tumor samples from 1,620 patients enrolled in the Austrian Breast & Colorectal Cancer Study Group 8, or ABCSG8, trial, of which 1,478 samples passed prespecified criteria and yielded evaluable results. The ABCSG8 trial enrolled 3,714 women (1996-2003) to compare the safety and efficacy of tamoxifen alone to sequential treatment with tamoxifen followed by anastrozole in postmenopausal women with HR+ early stage breast cancer of pathological grade 1 or 2. Access to the ABCSG8 samples and data is controlled by the chairman of the ABCSG, who, in 2010, agreed to collaborate with us as the lead investigator on a study to validate Prosigna. In 2011, we jointly designed the study with him and agreed to cover the costs of the study. Investigators at the British Columbia Cancer Agency, or BCCA, performed the Prosigna test using the nCounter Analysis System installed in BCCA's Center for Translational and Applied Genomics on tumor samples which had been stored in FFPE format from participants in the original ABCSG8 study. Results of our ABCSG8 study were presented in December 2012 at the CTRC-AACR San Antonio Breast Cancer Symposium. In December 2013, the results of the ABCSG8 study were published in the Annals of Oncology, the Journal of the European Society of Medical Oncology.

Beginning in 2012, we planned and executed a series of prospectively defined analyses of the data sets from the ATAC and ABCSG8 trials designed to clinically validate additional features and benefits of Prosigna. Results from three of these analyses have been presented publicly at medical meetings. At the European Society of Medical Oncology meeting in September 2012, the investigators of our TransATAC study presented results indicating that the ROR score provided by Prosigna adds significant prognostic information to clinical-pathology variables for recurrence between five and 10 years after diagnosis, which is often referred to as late recurrence. In September 2013, these results were published in the Journal of the National Cancer Institute. At the IMPAKT Breast Cancer Conference in May 2013, the investigators of our TransATAC study and our ABCSG8 study presented additional analyses providing further evidence that Prosigna provides valuable information that could assist with treatment decisions by helping to identify patients at highest risk of this late recurrence. In addition, the results of an analysis of the combined data set of the ABCSG8 and ATAC studies was presented during the

## **Table of Contents**

2013 Annual American Society of Clinical Oncology, or ASCO, Meeting in June 2013 and demonstrated that Prosigna can identify a clinically significant number of low risk patients with one or two positive nodes. Several additional analyses of the ABCSG8 and ATAC studies are planned or ongoing.

In 2012, we performed a series of multi-site analytical validation studies intended to show that Prosigna provides consistent and reliable results, independent of the specific instrument, laboratory or operator performing the testing. We presented results from these analytical validation studies in March 2013 at the United States & Canadian Academy of Pathology annual meeting.

### *Clinical Validation of Prosigna for Indicating Prognosis in Postmenopausal HR+ Early Stage Breast Cancer Patients*

Our TransATAC and ABCSG8 studies were performed using similar statistical analysis plans, allowing results on the prognostic performance of Prosigna in each study to be compared. Both studies met their primary and secondary objectives, and the data support the following conclusions:

in satisfaction of the primary objective of both studies, the ROR score was significantly related to outcome, and added significant prognostic information about 10 year distant recurrence risk to standard clinical-pathological variables in the study populations as a whole. In satisfaction of a secondary objective of both studies, similar results were achieved in all three prospectively defined clinically important subsets of patients: node-negative, node-positive and HER2-negative;

in satisfaction of the primary objective of our ABCSG8 study, the low, intermediate, and high risk patient groups as defined by Prosigna had different distant recurrence free survival rates at 10 years in the study population as a whole, showing that Prosigna can accurately categorize patients based on prognosis; and

in satisfaction of a secondary objective of both studies, patients with different intrinsic subtypes as reported by Prosigna had significantly different outcomes when treated with endocrine therapy alone, reinforcing the power of intrinsic subtyping as a descriptor of breast cancer tumor biology.

When taken together, we believe that our TransATAC and ABCSG8 studies provide strong evidence for Prosigna's clinical validity. The following tables and figures summarize the prognostic performance of Prosigna.

*Prosigna's ROR score is significantly related to outcome in both node-negative and node-positive breast cancer patients (TransATAC study)*

**Table of Contents**

These curves illustrate the relationship of the ROR score to 10 year risk of distant recurrence in node-negative patients, patients with one to three positive nodes and patients with four or more positive nodes in the TransATAC population. In each subset of patients, as the ROR score increases, so too does 10 year risk of distant recurrence.

*Prosigna adds statistically significant prognostic information beyond standard clinical-pathological variables in both studies*

Patient population	Number of patients	TransATAC		Number of patients	ABCSG8	
		DLR-X <sup>2</sup>	p-value		DLR-X <sup>2</sup>	p-value
All evaluable	1,007	34.2	<0.0001	1,478	53.5	<0.0001
Node-negative	739	25.0	<0.0001	1,047	25.6	<0.0001
Node-positive	268	9.3	0.0023	382 <sup>(1)</sup>	26.0	<0.0001
Her2-negative	888	28.9	<0.0001	1,397	47.5	<0.0001

(1) Includes patients with only one to three positive nodes.

The above table shows the prognostic information provided by the ROR score in the TransATAC and ABCSG8 study populations. The statistic DLR-X2 measures the amount of prognostic information which the ROR score provides beyond the standard clinical-pathological variables. Since statistical significance is defined as p<0.05, these studies showed that the ROR score is significantly related to outcome and adds statistically significant prognostic information about 10 year distant recurrence risk to standard clinical-pathological variables in the study population as a whole and in all three prospectively defined clinically important subsets of patients in both studies.

*Prosigna's risk groups have statistically significant different outcomes in the study populations as a whole*

Risk Group	Number of Patients (%)	TransATAC Estimated DRFS at 10 years		Number of Patients (%)	ABCSG8 Estimated DRFS at 10 years	
		Percent [95% CI] <sup>(1)</sup>	p-value <sup>(2)</sup>		Percent [95% CI] <sup>(1)</sup>	p-value <sup>(2)</sup>
Low	437 (43%)	96% [94%-98%]	<0.0001	502 (34%)	97% [95%-98%]	0.0093
Intermediate	254 (25%)	86% [81%-90%]	NA	478 (32%)	91% [88%-94%]	NA
High	316 (31%)	63% [57%-69%]	<0.0001	498 (34%)	80% [76%-83%]	0.0004
Total	1,007 (100%)			1,478 (100%)		

(1) DRFS = Distant Recurrence Free Survival; CI = Confidence Interval.

(2) P-value calculated based on comparison to intermediate risk group.

For the CE-marked version of Prosigna, the above table illustrates the result that in both studies the Prosigna-defined risk groups in the study populations as a whole have different 10 year distant recurrence free survival rates. In both studies, the low and high risk groups showed distant recurrence free survivals with statistically significant differences from the intermediate risk group.

**Table of Contents**

Patients with node-negative disease categorized as Luminal A subtype had higher distant recurrence free survival than those categorized as Luminal B subtype

Subtype	TransATAC			ABCSG8		
	Number of Patients (%)	Percent [95% CI] <sup>(1)</sup>	p-value <sup>(2)</sup>	Number of Patients (%)	Percent [95% CI] <sup>(1)</sup>	p-value <sup>(2)</sup>
Luminal A	529 (72%)	94% [92%-96%]	NA	725 (69%)	95% [93%-96%]	NA
Luminal B	176 (24%)	74% [68%-81%]	<0.0001	284 (27%)	87% [83%-90%]	0.0019
Total <sup>(3)</sup>	705 (95%)			1,009 (96%)		

(1) DRFS = Distant Recurrence Free Survival; CI = Confidence Interval.

(2) P-value calculated based on comparison to Luminal A group.

(3) Total number of patients is less than 100% of evaluable patients because a small number of patients were categorized as Basal-like or HER2-enriched subtypes.

The above table illustrates the result that in both studies intrinsic subtypes as defined by Prosigna in the node-negative patients have different 10 year distant recurrence free survivals. In each study, patients categorized as Luminal A have a statistically significantly higher estimated distant recurrence free survival at 10 years than patients categorized as Luminal B.

#### Comparison of Prosigna and Oncotype DX Performance in Our TransATAC Study

The TransATAC population had been previously used in 2008 to clinically validate the current market leader in breast cancer prognosis and prediction, Genomic Health's *Oncotype DX*, which is a laboratory-developed test that uses a centralized laboratory service model. The PAM50 study used RNA that had been extracted by Genomic Health from 1,017 tumor samples from patients with postmenopausal HR+ early stage breast cancer during the 2008 study. Because both the *Oncotype DX* results and the outcomes of the patients associated with each RNA sample were known, the study provided an opportunity to measure how the prognostic information provided by Prosigna compares to that provided by *Oncotype DX* in this study population.

In order to compare how the two tests separated patients according to risk in this study, risk groups were defined based on each test's estimate of the risk of distant recurrence at 10 years within the TransATAC population. Risk score thresholds to define the risk groups were chosen for each test based on the results of our TransATAC study in order to define risk groups that contain patients with the same risk. In order to achieve these comparable risk groups, the cut points used for *Oncotype DX* were different than those used by Genomic Health.

For each test, the low risk group was prospectively defined as patients with less than a 10% estimated risk of recurrence. For each test, the intermediate risk group was prospectively defined as patients with between a 10% and 20% estimated risk of recurrence. For each test, the high risk group was prospectively defined as patients with greater than a 20% estimated risk of recurrence. The figure below summarizes the sizes and outcomes of the risk groups defined by each test in patients with node-negative disease.

## **Table of Contents**

*Prosigna's ROR score identified more high risk patients and fewer intermediate risk patients than Oncotype DX's RS score in this study in patients with node-negative disease*

This figure illustrates the result that, in patients with node-negative disease, Prosigna assigned 26% fewer patients to the intermediate risk group than did Oncotype DX (180 patients vs. 243 patients) in this study. In addition, in patients with node-negative disease, Prosigna assigned more patients to the high risk group than did Oncotype DX; however, the low risk and high risk groups defined by each test have similar outcomes as illustrated by the overlapping Kaplan-Meier curves. This observation led the independent investigators of our TransATAC study to conclude that, in patients with node-negative disease, Prosigna assigned fewer patients to the intermediate risk group than Oncotype DX RS, with equivalent or higher separation between the low and high risk groups.

*Clinical validation of Prosigna ability to help identify node-positive early-stage breast cancer patients at low risk of recurrence*

In June 2013, we presented results from the combined data analysis of the ATAC and ABCSG8 studies at the Annual American Society of Clinical Oncology, or ASCO, meeting. These results were derived from the analysis of the combined data set and long term follow-up from 2,485 patients in the ABCSG8 and ATAC studies. Node-positive patients in the combined data set were grouped into one of three categories based on the number of positive nodes.

According to current treatment guidelines by the National Comprehensive Cancer Network, or NCCN, postmenopausal women with node-positive, HR+ early-stage breast cancer should be considered high risk and should receive adjuvant chemotherapy in addition to five years of endocrine therapy. The results of both randomized clinical studies and meta-analyses suggest that a substantial portion of women with node-positive disease may be adequately treated with adjuvant endocrine therapy alone. However, identifying these low risk node-positive patients has been challenging due to the heterogeneity of the node-positive patient population, which includes patients with different numbers of positive lymph nodes and diverse tumor genomics.

The objective of the new study was to determine whether the risk score provided by Prosigna provides additional prognostic information for risk of metastasis over and above standard clinical variables alone in patients with either one positive lymph node, or two to three positive lymph nodes. Of all the patients in the study with one positive node, 40% were categorized as low-risk based on their risk score and experienced an absolute 10-year risk of distant recurrence rate of 6.6%, while 71% were categorized as Luminal A subtype and experienced an absolute 10-year risk of distant recurrence of 8.4%. Separately, the analysis also demonstrated that patients with the Luminal A subtype have statistically significant different risk of metastasis than Luminal B.

The investigators in this study concluded that Prosigna helped identify a subset of postmenopausal women with node-positive HR+ early stage breast cancer, including patients with one positive node, as well as some with two positive nodes that had a low risk of recurrence. The authors concluded that identifying this subset of patients may help physicians assess treatment options, including whether the patients might be adequately treated with adjuvant endocrine therapy alone.

---

## **Table of Contents**

### *Clinical Validation of Prosigna for Indicating Risk of Late Recurrence in Postmenopausal HR+ Early Stage Breast Cancer*

Despite recent improvements in breast cancer treatment, some women with HR+ early-stage breast cancer remain at risk of disease recurrence after remaining recurrence-free for the first five years following diagnosis. Identifying newly diagnosed women with HR+ breast cancer who are at highest risk of having their cancer recur between five and 10 years after diagnosis is a priority for oncologists seeking to help breast cancer patients make more informed treatment decisions. These patients may benefit by extending the duration of their adjuvant endocrine therapy beyond five years.

The ability of Prosigna to estimate risk of recurrence between five and 10 years after diagnosis in postmenopausal women with HR+, node-positive and node-negative early-stage breast cancer has been demonstrated in two independent studies. Results from both of these studies were presented at the IMPAKT Breast Cancer Conference in May 2013.

In one study, the investigators of our TransATAC study assessed and compared the value of five different prognostic scores for indicating risk of distant recurrence in the first five years after diagnosis, and between five and 10 years after diagnosis, for all patients in the ATAC trial. In a multivariate analysis, the ROR score was one of only two genomic prognostic scores that provided additional prognostic information regarding risk of distant recurrence between five and 10 years. The results of the TransATAC late recurrence study were published in the Journal of the National Cancer Institute in September 2013.

In a second study, the investigators of our ABCSG8 study found that the ROR score provided by Prosigna added prognostic information about the risk of late recurrence of breast cancer to the standard pathological variables in a study population including 1,478 postmenopausal women with hormone receptor positive, node-positive and node-negative early-stage breast cancer who participated in the ABCSG8 trial ( $p < 0.0001$ ). Patients with no recurrence by year five and categorized as low risk based on the ROR score had Distant Recurrence Free Survival, or DRFS, of 98.7% at year 10, while patients with no recurrence by five years and categorized as high risk based on the ROR score had DRFS of 91.5% at year 10. These investigators concluded that, in combination with the late recurrence results from the TransATAC study, Prosigna's ability to indicate risk of late recurrence in postmenopausal HR+ early stage breast cancer has achieved Level 1 clinical evidence, the standard generally required for inclusion in clinical practice guidelines.

### *Multi-site analytical validation of Prosigna.*

Regulatory clearance or approval for *in vitro* diagnostic kits generally requires studies that demonstrate that a diagnostic test can be reliably run in multiple qualified laboratories with adequate precision and reproducibility. In 2012, we performed a series of multi-site analytical validation studies intended to show that Prosigna provides consistent and reliable results, independent of the specific instrument, laboratory or operator performing the testing. The results of these studies were presented publicly at the United States & Canadian Academy of Pathology annual meeting in March 2013. These results support the ability of the Prosigna assay to generate consistent results when used in qualified clinical labs in different cities and countries.

The objective of these studies was to assess the analytical robustness of Prosigna when used in qualified clinical laboratories. Prosigna assays were run independently at three different testing sites by a total of six different operators using three different reagent lots. Reproducibility was assessed by testing multiple tissue sections from each of 43 FFPE breast tumor blocks across the three sites following review of hematoxylin and eosin stained slides by an independent pathologist at each site. The magnitude of different sources of analytical variation in the assay were further characterized by testing five pooled breast tumor RNA samples more than 100 times each.

When starting with FFPE tissue blocks, these studies showed that Prosigna's reproducibility, including all analytical and pre-analytical variables, was characterized by a total standard deviation of just 2.9 ROR units on a zero-to-100 scale. In addition, there was an average site-to-site concordance of 97% in reporting of intrinsic subtype. When starting from pooled RNA, Prosigna's precision was characterized by a total standard deviation of less than 1 ROR unit. There was no statistically significant bias in results between sites or operators.

---

## **Table of Contents**

### *Prosigna in the European Union and Other Countries that Recognize the CE Mark.*

In September 2012, we obtained CE mark designation for Prosigna for use as a semi-quantitative *in vitro* diagnostic assay using the gene expression profile of cells found in FFPE breast tumor tissue to assess the 10 year risk of distant recurrence in postmenopausal women with HR+ early stage breast cancer treated with endocrine therapy alone. This CE-marked product is indicated for use in patients with either node-negative or node-positive disease, and provides physicians and their patients with the intrinsic subtype of a patient's breast cancer tumor, ROR score, and risk category (high/intermediate/low). In early 2013, we began marketing this test in Europe, including in France, Germany, Greece, Italy, Spain, Turkey and the United Kingdom, and Israel. In April 2013, we installed the first diagnostic systems in Europe, which will initially be used for clinical studies of Prosigna's impact on adjuvant treatment decisions in early stage breast cancer called decision impact studies. The list price of Prosigna kits in Europe is the equivalent of \$1,550 per patient.

### *Prosigna in the United States.*

In September 2013, we received 510(k) clearance from the FDA to market in the United States a version of Prosigna providing a prognostic indicator for distant recurrence-free survival at 10 years, and is indicated for postmenopausal women with Stage I/II lymph node-negative or Stage II lymph node-positive (one to three positive nodes) hormone receptor-positive breast cancer who have undergone surgery in conjunction with locoregional treatment consistent with standard of care. For each patient, the Prosigna Assay reports the Prosigna Score, which is referred to as the ROR Score in the scientific literature and outside the United States, and a risk category based on both the Prosigna Score and nodal status. Node-negative patients are classified as low, intermediate or high risk, while node-positive patients are classified as low or high risk. Prosigna is not intended for diagnosis, to predict or detect response to therapy, or to help select the optimal therapy for patients. We expect Prosigna to be competitive with other products that are currently available in the United States given the advantages demonstrated by our TransATAC and ABCSG8 clinical studies. In the future, we plan to submit a separate application for approval to report intrinsic subtype. If we obtain approval to report intrinsic subtyping from the FDA, we expect our competitive position in the United States will be enhanced. We expect that this future application will require a PMA supported by additional clinical studies.

The U.S. list price for a Prosigna kit is expected to be \$2,080 per test (which is comparable to the pricing in jurisdictions accepting the CE-marked version of Prosigna). We plan to sell Prosigna kits to our lab customers, who will be responsible for providing the testing service and contracting and billing payors. We also plan to sell Prosigna kits to clinical laboratories on a fixed dollars-per-kit basis, which would not expose us to direct third-party payor reimbursement risk. However, we anticipate providing customary volume discounts, and in some cases, introductory pricing during the period in which third-party payor reimbursement is being established. As a result, we expect the average selling price per Prosigna test to be between \$1,500 and \$2,000 in the United States.

## **Intellectual Property**

We must develop and maintain protection on the proprietary aspects of our technologies in order to remain competitive. We rely on a combination of patents, copyrights, trademarks, trade secret and other intellectual property laws and confidentiality, material transfer agreements, licenses, invention assignment agreements and other contracts to protect our intellectual property rights.

As of December 31, 2013, we owned or exclusively licensed seven issued U.S. patents and approximately 23 pending U.S. patent applications, including provisional and non-provisional filings. We also owned or licensed approximately 73 pending and granted counterpart applications worldwide, including 22 country-specific validations of four European patents. The issued U.S. patents that we own or exclusively license are expected to expire between July 3, 2021 and March 28, 2029. We have either sole or joint ownership positions in all of our pending U.S. patent applications. Where we jointly own cases, we have negotiated license or assignment provisions for exclusive rights. For our material nCounter Analysis System and Prosigna product rights, we are the exclusive licensee. We also generally protect our newly developed intellectual property by

**Table of Contents**

entering into confidentiality agreements that include intellectual property assignment clauses with our employees, consultants and collaborators.

Our patent applications relate to the following three main areas:

our nCounter Analysis System biology, chemistry, software and hardware;

specific applications for our nCounter Analysis System technology; and

our gene expression markers, methods and algorithms for recurrence and drug response in certain forms of cancer.

The following patents and patent applications (including expected 20 year expiration dates) relate to our nCounter Analysis System:

<b>Patent and Patent Application Numbers</b>	<b>Form of Ownership</b>	<b>Expected Expiration Date</b>	<b>Description</b>
US 7,473,767, US 7,919,237, US 8,148,512, EP Patent No. 1448581, AU Patent No. 2002327202, CA Patent No. 2452712, JP Patent No. 4343682, USSN 13/794,299, US 8,492,094 and foreign applications in certain jurisdictions claiming priority to PCT/US2002/021278	In-licensed from the Institute for Systems Biology	7/3/2021	Directed to compositions and methods of immobilization and detection
EP Patent No. 1963531, AU Patent No. 2006330830, USSN 13/794,424 and foreign applications in certain jurisdictions claiming priority to PCT/US2006/049274	Co-owned with the Institute for Systems Biology	12/22/2026	Directed to compositions and methods of immobilization and detection
EP Patent No. 1963500, USSN 11/645,270 and foreign applications in certain jurisdictions claiming priority to PCT/US2006/049279	Owned	12/22/2026	Directed to methods of immobilization and detection
US 7,941,279, AU Patent No. 2007268027, CA Patent No. 2653095, JP Patent No. 5081232 and foreign applications in certain jurisdictions claiming priority to PCT/US2007/012130	Owned	5/21/2027	Directed to compositions
US 8,415,102, USSN 13/788,133 and foreign applications in certain jurisdictions claiming priority to PCT/US2008/059959	Owned	4/10/2028	Directed to methods of manufacture
US 8,519,115, USSN 13/957,029 and foreign applications in certain jurisdictions claiming priority to PCT/US2009/053790	Owned	8/13/2029	Directed to compositions and methods of detections



**Table of Contents**

The following patent applications (including expected 20 year expiration dates) relate to specific applications for our nCounter Analysis System:

<b>Patent Application Numbers</b>	<b>Form of Ownership</b>	<b>Expected Expiration Date</b>	<b>Description</b>
USSN 12/904,078 and foreign applications in certain jurisdictions claiming priority to PCT/US2010/052556	Owned	10/13/2030	Directed to compositions and methods of detection
USSN 13/025,458 and foreign applications in certain jurisdictions claiming priority to PCT/US2011/024519	Owned	2/11/2031	Directed to compositions and methods of detection
USSN 13/049,682 and foreign applications in certain jurisdictions claiming priority to PCT/US2011/028657	Owned	3/16/2031	Directed to methods of detection
USSN 14/007,586	Owned	3/28/2032	Directed to compositions and methods of diagnosis
USSN 13/530,848 and foreign applications in certain jurisdictions claiming priority to PCT/US2012/043799	Owned	6/22/2032	Directed to compositions and methods of detection
USSN 14/078,009 and PCT/US2013/069665	Owned	11/12/2033	Directed to compositions and methods of diagnosis
Additional pending provisional patent applications	Owned	2033	Directed to nCounter Analysis System methods of use

**Table of Contents**

The following patent applications (including expected 20 year expiration dates) relate to our gene expression markers:

<b>Patent Application Numbers</b>	<b>Form of Ownership</b>	<b>Expected Expiration Date</b>	<b>Description</b>
USSN 13/959,575 and foreign applications in certain jurisdictions claiming priority to PCT/US2006/044737	In-licensed from Bioclassifier, LLC	11/17/2026	Directed to methods of prognosis
EP Patent No. 2297359, USSN 12/995,450 and foreign applications in certain jurisdictions claiming priority to PCT/ US2009/045820	In-licensed from Bioclassifier, LLC	6/1/2029	Directed to methods of prognosis
USSN 13/421,367 and foreign applications in certain jurisdictions claiming priority to PCT/ US2012/029226	In-licensed from Bioclassifier, LLC	3/15/2032	Directed to methods of treatment and determining drug response
USSN 13/690,891 and PCT/ US2012/067317	In-licensed from Bioclassifier, LLC	11/30/2032	Directed to methods of treatment and determining drug response
USSN 13/899,656 and PCT/US2013/042157	Owned	5/22/2033	Directed to compositions and methods of using gene expression markers
USSN 13/930,249 and PCT/US2013/048551	In-licensed from Bioclassifier, LLC	6/28/2033	Directed to methods of treatment and determining drug response
Additional pending provisional patent applications	Owned or In-licensed from Bioclassifier, LLC	2033	Directed to compositions and methods of using gene expression markers, some of which are owned by NanoString Technologies, Inc. and some of which are encompassed by our license agreement with Bioclassifier, LLC

We intend to file additional patent applications in the United States and abroad to strengthen our intellectual property rights; however, our patent applications (including the patent applications listed above) may not result in issued patents, and we cannot assure investors that any patents that have issued or might issue will protect our technology. We have received notices of claims of potential infringement from third parties and may receive additional notices in the future. When appropriate, we have taken a license to the intellectual property rights from such third parties. For additional information, see the section of this prospectus captioned **Risk Factors** **Risks Related to Intellectual Property**.

We own a number of trademarks and develop names for our new products and as appropriate secure trademark protection for them, including domain name registration, in relevant jurisdictions.

**Collaborations; License Agreements**

We have relied, and expect to continue to rely, on strategic collaborations and licensing agreements with third parties. For example, our base molecular barcoding technology is in-licensed from the Institute for Systems Biology and the intellectual property that forms the basis of Prosigna is in-licensed from Bioclassifier, LLC. In addition to the licenses with the Institute for Systems Biology and Bioclassifier, we rely on other license and supply arrangements for proprietary components which require us to pay royalties on the sale of our products.

---

**Table of Contents**

Other life sciences research customers are using our nCounter Analysis System to discover gene expression signatures that we believe could form the basis of future diagnostic products. Currently, we are considering several of these gene signatures for in-licensing. For example, in February 2013 we secured an option from a customer to acquire an exclusive worldwide license for a gene signature that could be used, after further development, as a Laboratory Developed Test, or, after appropriate regulatory authorization, for a second molecular diagnostic product to identify patients with cirrhosis who are at highest risk of developing HCC and to determine whether a patient who has been diagnosed with HCC is likely to have a recurrence. Our licensing arrangements with the Institute for Systems Biology and Bioclassifier are discussed below in greater detail.

***Institute for Systems Biology***

In 2004, we entered into an agreement with the Institute for Systems Biology pursuant to which the Institute granted to us an exclusive, subject to certain government rights, worldwide license, including the right to sublicense, to the digital molecular barcoding technology on which our nCounter Analysis System is based, including 13 patents and patent applications. We issued 15,625 shares of our common stock to the Institute for Systems Biology as partial consideration for entry into the license agreement. Pursuant to the terms of the amended license agreement, we are required to pay the Institute for Systems Biology royalties on net sales of products sold by us, or our sublicensees, at a low single digit percentage rate. Royalties owed to the Institute for Systems Biology had been subject to annual minimums, which have expired. Through December 31, 2013, we have paid aggregate royalties of \$1.7 million to the Institute for Systems Biology. Unless earlier terminated in accordance with the terms of the amended license agreement, the agreement will terminate upon the expiration of the last to expire patent licensed to us. The Institute for Systems Biology has the right to terminate the agreement under certain situations, including our failure to meet certain diligence requirements or our uncured material breach of the agreement.

***Bioclassifier, LLC***

In July 2010, we entered into an exclusive license agreement with Bioclassifier, LLC, pursuant to which Bioclassifier granted to us an exclusive, subject to certain government rights, worldwide license, with the right to sublicense, to certain intellectual property rights and technology, including intellectual property rights that comprise eight non-provisional patent applications as of December 31, 2013, in the field of research products and prognostic and/or diagnostic tests for cancer, including Prosigna. Bioclassifier has licensed these rights from the academic institutions that employed the cancer researchers that discovered or were involved in the initial development of PAM50. This license agreement was amended and restated in February 2012, with the changes retroactively effective to the July 2010 date of the original agreement. Pursuant to the terms of the amended and restated license agreement, we are required to pay Bioclassifier the greater of certain minimum royalty amounts and mid-single digit to low double digit percentage royalties on net sales of products and/or methods sold by us that are covered by patent rights or include, use or are technology licensed to us. Our obligation to pay royalties to Bioclassifier expires on a country-by-country basis upon the expiration of the last patent licensed or, if a product or method includes, uses or is technology licensed to us but is not covered by a patent licensed to us, ten years after the first commercial sale of the product or method in such country. We are also required to pay Bioclassifier low to mid double digit percentage of any income received by us from the grant of a sublicense by use to the patents or technology licensed us under the agreement. We are also required to meet certain development and commercialization milestones extending to 2015. Through December 31, 2013, we have paid Bioclassifier \$365,000 of which \$175,000 will be credited against future royalties owed.

Additionally, we are obligated to pay certain fees to Bioclassifier if we do not meet certain milestones within predetermined time periods. The agreement specifies that we will control and be responsible for the costs of prosecuting and enforcing the intellectual property licensed in certain major market countries. The agreement also includes customary rights of termination for Bioclassifier, including for our uncured material breach or our bankruptcy.

## **Table of Contents**

### **Research and Development**

We have committed, and expect to continue to commit, significant resources to developing new technologies and products, improving product performance and reliability and reducing costs. We have assembled experienced research and development teams at our Seattle, Washington location with the scientific, engineering, software and process talent that we believe is required to successfully grow our business. As of December 31, 2013, we had approximately 38 employees in research and development, of which 17 hold a Ph.D. degree and five hold a M.S. degree. We are currently focused on several products and enhancements in both our future diagnostic products and current life sciences research offerings. Our research and development expenses for the years ended December 31, 2010, 2011 and 2012 and the nine months ended September 30, 2013 were \$7.5 million, \$9.0 million, \$11.6 million and \$10.5 million, respectively.

#### ***nCounter Technology***

We are continuously seeking to improve the nCounter Analysis System, including improvements to the technology and accessibility. As we make improvements, we anticipate that we will make available new and improved generations of the nCounter Analysis System to be used in both our life sciences research and diagnostics businesses.

Our technology development efforts are focused on:

*Applications.* We plan to develop additional application areas to enable researchers to apply the nCounter Analysis System to new experimental paradigms. Currently, we are updating our panel product line with panels focused on cancer pathways and the immune response to cancer. We are also focused on improving and expanding the ability of our technology to detect gene fusions. Finally, we are exploring the application of the nCounter Analysis System for the digital multiplexed quantitation of proteins, which may allow researchers to measure multiple nucleic acids and proteins with a single instrument using small amounts of precious sample.

*Instruments.* We are developing a new generation of the nCounter Analysis System that we believe will increase our addressable market and simplify the procurement processes of our potential customers. The new generation system will be a single instrument with a reduced footprint that combines the prep station and the digital analyzer. We plan to reduce the cost of the new generation system through the adoption of new, less expensive technologies. We are targeting release of the new generation system in 2014.

#### ***Expanding Clinical Utility of the Prosigna Breast Cancer Assay***

We plan to extend the clinical utility of Prosigna to inform other major treatment decisions in breast cancer, after appropriate regulatory authorization. The decisions about receiving extended adjuvant endocrine therapy, adjuvant chemotherapy or adjuvant radiation therapy have significant objective quality of life implications because of the acute and long term risk of side effects, some of them severe (including death), that are caused by these treatments. In addition, there are significant health economic consequences to decisions regarding these therapies based both on the cost of the treatments themselves and of treating their side effects. Therefore, a pressing issue is to identify the individual patients who need or are likely to benefit from extended adjuvant endocrine therapy, adjuvant radiation therapy and adjuvant chemotherapy so that the rest of the patients can be spared these treatments without affecting their long term outcome.

Our efforts to expand the clinical utility of Prosigna are focused on:

*Duration of Endocrine Therapy.* For postmenopausal women with early stage breast cancer, the standard of care is to treat with five years of endocrine therapy after diagnosis and initial treatment. However, the results of meta-analyses demonstrate that even with the standard adjuvant endocrine treatment, a significant percentage of women continue to have recurrences of breast cancer after five years from initial diagnosis. Recently reported randomized trials comparing five years of adjuvant

---

**Table of Contents**

endocrine therapy with extending adjuvant endocrine therapy beyond five years have shown that extended endocrine therapy reduces both breast cancer recurrence and breast cancer mortality. Three clinical studies have demonstrated that the Prosigna score adds prognostic information to the standard clinical parameters for determining the risk of distant recurrence after five years and that Prosigna-defined risk groups identify patients with different risks of distant recurrence after five years. We intend to conduct clinical studies to validate the ability of Prosigna-defined risk groups to identify postmenopausal women with early stage breast cancer who are likely to receive little or no benefit from treatment with extended endocrine therapy.

*Chemotherapy Selection.* The first-generation genomic tests for breast cancer have improved physicians' ability to determine which individual patients can be safely spared adjuvant chemotherapy. However, for those patients who will go on to receive adjuvant chemotherapy, physicians may select from several commonly used adjuvant chemotherapy regimens, including CMF (cyclophosphamide/ methotrexate/5-flouracil), anthracycline-containing regimens or taxane-containing regimens. The first-generation genomic breast cancer tests do not inform the selection of specific adjuvant chemotherapy regimen. Over the past several years, studies have been presented and published that suggest that the PAM50 gene signature, which is the basis of Prosigna, can inform the selection of adjuvant chemotherapy regimen. In 2012, U.S.-based researchers published a study indicating that a qPCR-based version of PAM50 predicted which breast cancer patients benefit from anthracycline-based chemotherapy regimens. In 2013, Danish researchers published a study indicating that Prosigna predicted which breast cancer patients benefit from gemcitabine. In 2013, Spanish researchers published a study demonstrating that the proliferation score of a qPCR-based version of PAM50 predicted which breast cancer patients benefit from weekly paclitaxel. In the future, we intend to perform clinical studies designed to demonstrate that Prosigna can aid in the selection of chemotherapy regimen in breast cancer patients. We have secured access to tissue samples and outcomes from two randomized, controlled clinical studies that may be used in an effort to clinically validate this intended use.

*Radiation Therapy in Early Stage Breast Cancer.* Recently presented research suggests that by determining a patient's intrinsic subtype, physicians could identify those patients who do not benefit from adjuvant radiation therapy. In a previously conducted clinical trial that enrolled postmenopausal women with T1/T2, node-negative early stage breast cancer, patients with Luminal A tumors (approximately 50% of the patients enrolled in the trial) received little or no benefit from adjuvant radiation therapy, whereas patients with tumors of other subtypes received significant benefit. We intend to conduct clinical studies to validate the ability of intrinsic subtype as determined by Prosigna to identify postmenopausal women with early stage breast cancer who are likely to receive little or no benefit from treatment with adjuvant radiation therapy. We have secured access to tissue samples and outcomes from one randomized, controlled clinical study that may be used for clinically validating this intended use.

*Ductal Carcinoma in situ Treatment.* Ductal Carcinoma in situ, or DCIS, is characterized by a clonal proliferation of epithelial cells confined within the lumen of the mammary duct. DCIS is usually asymptomatic; however, screening mammography programs have led to a substantial increase in the incidence of DCIS in the past two decades so that it represented 20% of breast cancers diagnosed in the United States in 2004, according to the National Cancer Institute's Surveillance, Epidemiology and End Results Program. The major clinical risk in DCIS is its progression to invasive carcinoma. Since DCIS has a variable natural history, the major treatment decision relates to how aggressively to treat DCIS when it is diagnosed. Mastectomy is a highly effective, although radical, treatment for DCIS as it is curative in 98 to 99% of patients with either gross or mammographically detected DCIS. A recent study has demonstrated that a multi-gene assay containing a subset of the genes used in the *Oncotype DX* test could identify a group of patients with such a low risk of recurrent DCIS that they are unlikely to benefit from treatment beyond limited surgical resection. These data suggest other genomic tests, including Prosigna, may also be able to identify low risk patients who may be spared aggressive treatment. We intend to conduct clinical studies to validate the ability of Prosigna to identify DCIS.

**Table of Contents**

patients who may be spared aggressive treatment. We have applied for access to a cohort of DCIS tissue samples from patients entered into a randomized, controlled clinical trial with long-term follow-up, and have received approval from the relevant committee within the institution controlling these samples.

Our clinical studies to date have employed a retrospective / prospective design, which means that we use samples that were previously collected from patients and for which the treatment regimen and ultimate outcome of each patient are known. Such studies are capital efficient as they do not require recruiting new patients and running prospective trials and they can be completed much more quickly than typical prospective clinical trials. We intend to use a similar approach whenever possible for the additional clinical studies we intend to conduct in support of our future regulatory submissions seeking to expand the indications for Prosigna and for future diagnostic products.

In the future, we do intend to participate in prospective clinical studies that require recruiting new patients. Thus far, we have accepted invitations to participate in two such prospective studies that are being organized and sponsored by cooperative groups. We are not and do not expect to be financially responsible for conducting either trial; however, we may provide in-kind support through the contribution of Prosigna *in vitro* diagnostic kits or sale of kits at a discounted price. These studies are:

*RxSPONDER trial (SWOG 1007).* The RxSPONDER trial is a Phase III clinical trial organized by the Southwest Oncology Group and sponsored by the National Cancer Institute. The primary objective of this trial is to determine the effect of endocrine therapy with versus without chemotherapy in patients with node-positive breast cancer who do not have high Recurrence Scores (RS) by *Oncotype DX*. The trial also has several secondary objectives related to other breast cancer assays, including PAM50. One secondary objective is to perform other assays or tests (in particular the ROR score) as they are developed and validated that measure potential benefit of chemotherapy, and to compare them to *Oncotype DX*. Another secondary objective is to determine the role of other assays, including PAM50, as indicators of Disease Free Survival, Distant Disease Free Survival, and Local Disease Free Interval of patients randomized to chemotherapy versus no chemotherapy.

*Optimal Personalised Treatment of early breast cancer using Multi-parameter Analysis (OPTIMA) trial.* The OPTIMA trial is a multi-center partially blind randomized clinical trial of early stage breast cancer patients in the United Kingdom. The OPTIMA trial seeks to advance the development of personalized medicine in breast cancer by using multi-parameter tests to help identify those women who are likely to benefit from chemotherapy and helping spare those who are unlikely to benefit from an unnecessary and unpleasant treatment. In the United Kingdom, the OPTIMA study population would ordinarily be treated with a combination of chemotherapy and endocrine therapy. The OPTIMA trial compares the management of patients using test-directed assignment to chemotherapy with standard management (i.e., chemotherapy) in a non-inferiority design. OPTIMA prelim is the preliminary phase of the study sponsored by the U.K. Health Technology Assessment of the National Institute of Health Research which will evaluate the performance and health-economics of alternative multi-parameter tests to determine which technology should be evaluated in the main trial. This decision will be informed by a combined primary outcome measure including concordance of test results, cost-effectiveness and deliverability of pathology services. All patients in the OPTIMA prelim trial will be tested with the *Oncotype DX* test as well as additional biomarkers and tests, including Prosigna. In addition, we are exploring potential clinical studies with the goal of investigating Prosigna's ability to provide other clinically useful information to physicians treating women with HR+ early stage breast cancer, including identifying which patients are likely to benefit from adjuvant chemotherapy.

***Future Molecular Diagnostics***

In addition to the development of Prosigna, we are currently evaluating several molecular signatures which have the potential to create additional diagnostic products or enable Laboratory Developed Tests based on nCounter Elements. We intend to license rights to molecular diagnostic intellectual property as part of our

## **Table of Contents**

strategy to develop additional diagnostic products and enable Laboratory Developed Tests, with a particular focus on licensing rights from our life sciences research customers who are seeking to translate their research into clinical products or services after the necessary regulatory authorizations are secured. We intend to target intellectual property rights that are well understood, have the potential to facilitate changes in treatment with a major impact on outcome and cost, have the potential to support value-based pricing and with respect to which tissue samples for clinical validation are readily available.

In February 2013, we secured an option to acquire an exclusive worldwide license for a 186 gene signature that could be used, after further development, as a Laboratory Developed Test, or after appropriate regulatory authorization, for a second molecular diagnostic product to determine the prognosis of patients diagnosed with the most common type of liver cancer, HCC, or with hepatitis C-related early-stage cirrhosis. We secured the option from The Broad Institute acting on behalf of the inventors' institutions. During the period in which the option can be exercised, we plan to assess the feasibility of developing an *in vitro* diagnostic assay based on the HCC gene signature for use on the nCounter Analysis System.

HCC, a form of liver cancer, is an increasingly prevalent clinical diagnosis and is the third most common cause of cancer-related death globally. While incidence rates of HCC have been lower in the United States than in many countries historically, domestic age-adjusted HCC incidence rates have doubled in recent decades. In fact, primary liver cancer mortality rates have increased faster than mortality rates for any other leading cause of cancer in the United States. HCC develops from advanced fibrosis of the liver, or cirrhosis, which is estimated to affect one to two percent of the world's population. The prognosis for patients with advanced HCC is poor, with a reported five-year survival rate of approximately 10%, thus it is important that patients be diagnosed with HCC when it is at an early stage and treatable with surgery. Since there is a high rate of recurrence of HCC after the treatment of the primary tumor, it is important to identify those patients with a high risk of recurrence so that these recurrences can be treated before advanced disease develops.

A paper in the New England Journal of Medicine in 2008 by Hoshida, et al, described the HCC gene signature in connection with a method for conducting gene expression analysis on RNA extracted from liver tissue adjacent to HCC tumors. Using this method, the authors discovered a 186-gene signature which identifies those HCC patients who have a poor prognosis because of a high rate of recurrence after primary treatment. This gene signature was highly correlated with survival in a training set of 82 Japanese patients and was validated in an independent set of 225 patients from the United States and Europe. A paper published online in January 2013 in the journal Gastroenterology demonstrated that this same 186-gene signature also identifies those patients with hepatitis C-related early-stage cirrhosis who have a poor prognosis because of their high rate of developing HCC.

## **Sales and Marketing**

We began selling nCounter Analysis Systems to life sciences researchers in 2008 and began sales efforts in the diagnostics market in Europe, including in France, Germany, Greece, Italy, Spain, Turkey and the United Kingdom, and Israel in early 2013, and in the United States in November 2013. We sell our instruments and life sciences research products primarily through our own sales force in North America and through a combination of direct and distributor channels in Europe, the Middle East, Asia Pacific and South America. We have agreements with 15 distributors, each of which is exclusive within a certain territory. In the event the distributor does not meet minimum performance requirements, we may terminate the distribution agreement or convert from an exclusive to non-exclusive arrangement within the territory, allowing us to enter into arrangements with other distributors for the territory. None of our customers represented more than 10% of our revenue for the years ended December 31, 2010, 2011 or 2012 or the nine months ended September 30, 2013.

## ***Instrumentation and Life Sciences Research***

Our sales and marketing efforts for instrumentation and in the life sciences market are targeted at department heads, research or clinical laboratory directors, principal investigators, core facility directors, and research scientists and pathologists at leading academic institutions, biopharmaceutical companies, publicly and

## **Table of Contents**

privately-funded research institutions and contract research organizations. We seek to increase awareness of our products among our target customers through direct sales calls, trade shows, seminars, academic conferences, web presence and other forms of internet marketing.

Our nCounter Analysis Systems are relatively new to the life sciences and diagnostic market place and our instruments require a significant capital investment or commitment to a reagent rental agreement. Our sales process involves numerous interactions with multiple people within an organization, and often includes in-depth analysis by potential customers of our products, proof-of-principle studies, preparation of extensive documentation and a lengthy review process. As a result of these factors, the large capital investment required in purchasing our instruments and the budget cycles of our customers, the time from initial contact with a customer to our receipt of a purchase order can vary significantly and be up to 12 months or longer. Given the length and uncertainty of our sales cycle, we have in the past experienced, and likely will in the future experience, fluctuations in our instrument sales on a period-to-period basis. We are developing a research use nCounter Analysis System that we intend to offer at a lower price, which we believe will simplify the procurement processes of our potential life science research customers as well as increase our addressable market. We also continue to develop enhancements to both the chemistries and assays that are run on the nCounter Analysis System, which may drive further adoption.

### ***Molecular Diagnostics***

We intend to sell Prosigna kits via a three-pronged effort. First, we will seek to establish third-party reimbursement and patient access for clinical testing services that our diagnostics customers will provide based upon our products by educating third-party payors regarding the clinical utility and health economic value of the clinical tests enabled by our technology. Second, we will seek to establish an installed base of nCounter Analysis Systems by selling or leasing instruments to select clinical laboratories, with initial sales efforts directed at large commercial laboratories and academic medical centers that treat a high volume of breast cancer patients. In December 2013, we announced that national diagnostic laboratories ARUP Laboratories, Laboratory Corporation of America Holdings and Quest Diagnostics have chosen to add Prosigna to their suites of breast cancer diagnostic tests, and the laboratories at the University of Alabama at Birmingham Comprehensive Cancer Center and University of North Carolina Lineberger Comprehensive Cancer Center will be among the initial facilities to offer the Prosigna assay in the United States, with the earliest testing beginning during the first quarter of 2014. Third, we will drive physician demand for clinical testing services enabled by our diagnostic products, and direct test orders toward those laboratories which have adopted our technology.

We intend to have a direct sales model in the United States and most of Europe. In other countries, we intend to have either direct sales, distributor relationships or a mix of both. Because oncology and pathology are relatively concentrated medical specialties, we believe that a focused marketing organization and specialized sales force with regional and local experience can effectively build interest within clinical laboratories and generate physician demand for Prosigna. Where appropriate, we intend to coordinate commercial efforts with the sales and marketing personnel of the clinical laboratories offering clinical testing services based on our diagnostic products. We believe that these clinical laboratories will be motivated to coordinate commercial efforts by the potential to improve patient care, broaden patient access and profit from testing services based on Prosigna and other potential nCounter-based diagnostics. We believe this direct sales approach, coupled with our multiple publications of clinical data in peer-reviewed journals, provides the best opportunity to increase patient and physician demand.

In connection with the U.S. launch of Prosigna, we have been actively recruiting sales professionals to build a dedicated sales force to educate medical oncologists about Prosigna. We intend to use a phased approach to build our sales force, initially hiring approximately 15 field based oncology-focused sales representatives during the first quarter of 2014, with the expectation that the sales force will grow once internal milestones related to treatment guideline inclusion and third-party payor reimbursement have been achieved. We also intend to build a small team of medical science liaisons to complement the sales force and expect to continue to rely on our existing sales professionals to place nCounter Dx Analysis Systems in clinical labs.



**Table of Contents**

**Manufacturing; Suppliers**

We use third-party contract manufacturers to produce our instruments and raw materials for our consumables, and we build the CodeSets and reagent packages at our Seattle, Washington facility.

***Instruments***

We outsource manufacturing of our nCounter Prep Stations and nCounter Digital Analyzers. Precision System Science, Co., Ltd. of Chiba, Japan, or PSS, is our sole source supplier for the nCounter Prep Station.

Korvis Automation Inc., or Korvis, is our sole source supplier for our nCounter Digital Analyzers at its facility in Corvallis, Oregon. The facilities at which our instruments are built have been certified to ISO 13485:2003 standards. Our contracts with these instrument suppliers do not commit them to carry inventory or make available any particular quantities. Under the terms of the two instrument supply agreements, we are required to place binding purchase orders for instruments that will be delivered to us by the supplier three to six months from the date of placement of the purchase order. Although qualifying alternative third-party manufac