

EXACT SCIENCES CORP
Form 424B5
June 07, 2018

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Filed Pursuant to Rule 424(b)(5)
Registration No. 333-218535

The information in this prospectus supplement is not complete and may be changed. This prospectus supplement and the accompanying prospectus are not an offer to sell these securities and are not soliciting an offer to buy these securities in any state where the offer or sale is not permitted.

Subject to Completion
Preliminary Prospectus Supplement dated June 7, 2018

PROSPECTUS SUPPLEMENT

(To prospectus dated June 6, 2017)

\$150,000,000

Exact Sciences Corporation

1.0% Convertible Senior Notes due 2025

We are offering \$150 million aggregate principal amount of 1.0% Convertible Senior Notes due 2025. We will pay interest on the notes on January 15 and July 15 of each year, beginning July 15, 2018. The notes will mature on January 15, 2025, unless earlier repurchased by us or converted.

The notes we are offering will be issued as additional notes under the indenture pursuant to which we previously issued \$690 million aggregate principal amount of our 1.0% Convertible Senior Notes due 2025 in January 2018, which we refer to as the initial notes. The notes will have identical terms as the initial notes, will be treated as a single series of securities with the initial notes under the indenture and will have the same CUSIP number as the initial notes. Holders of the notes and the initial notes will vote as one class under the indenture.

Holders may convert their notes at their option prior to the close of business on the business day immediately preceding July 15, 2024 only under the following circumstances: (1) on any date during any calendar quarter (and only during such calendar quarter) if the closing sale price of our common stock was more than 130% of the applicable conversion price on each applicable trading day for at least 20 trading days (whether or not consecutive) in the period of the 30 consecutive trading days ending on the last trading day of the immediately preceding calendar quarter; (2) during the five business day period following any five consecutive trading day period in which the trading price per \$1,000 principal amount of notes for each trading day during such five trading day period was less than 98% of the product of the closing sale price of our common stock and the applicable conversion rate on each such trading day; or (3) upon the occurrence of specified corporate events. On or after July 15, 2024 until the close of business on the second scheduled trading day immediately preceding the maturity date, holders may convert all or a portion of their notes at any time. Upon conversion, we will pay or deliver, as the case may be, cash, shares of our common stock or a combination of cash and shares of our common stock, at our election, as described in this prospectus supplement.

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The initial conversion rate for the notes will be 13.2569 shares of common stock per \$1,000 principal amount of notes (equivalent to an initial conversion price of approximately \$75.43 per share of common stock). The conversion rate will be subject to adjustment in certain events but will not be adjusted for accrued interest as described herein. Following certain corporate transactions, we will increase the applicable conversion rate for a holder that elects to convert its notes in connection with such corporate transactions by a number of additional shares of our common stock as described in this prospectus supplement.

If we undergo a fundamental change (as defined herein) prior to maturity of the notes, holders will have the right, at their option, to require us to repurchase for cash all or a portion of their notes at a repurchase price equal to 100% of the principal amount of the notes being repurchased, plus accrued and unpaid interest to, but excluding, the fundamental change repurchase date.

We may not redeem the notes prior to the maturity date.

The notes will rank senior in right of payment to all of our future indebtedness that is expressly subordinated in right of payment to the notes; rank equally in right of payment to all of our future liabilities that are not so subordinated; be effectively junior to all of our existing and future secured indebtedness and other secured obligations, to the extent of the value of the assets securing that indebtedness and other secured obligations; and be structurally subordinated to all liabilities (including trade payables) of our subsidiaries.

Our common stock trades on the NASDAQ Capital Market under the symbol "EXAS". On June 6, 2018, the last sale price of the shares as reported on the NASDAQ Capital Market was \$65.36 per share.

Investing in the notes involves risks that are described in the "Risk Factors" section beginning on page S-14 of this prospectus supplement and on page 16 of our Annual Report on Form 10-K for the year ended December 31, 2017.

	Per Note	Total
Public offering price(1)	%	\$
Underwriting discount(2)	%	\$
Proceeds, before expenses, to us	%	\$

(1) The purchase price of the notes in this offering will include accrued interest from, and including, January 17, 2018. The table above does not reflect such accrued interest.

(2) We refer you to the "Underwriting" section of this prospectus supplement for additional information regarding underwriter compensation.

We have granted the underwriters an option to purchase up to an additional \$22,500,000 aggregate principal amount of notes, solely to cover over-allotments, if any, for 30 days after the date of this prospectus supplement.

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

The notes will be ready for delivery in book-entry form only through the facilities of The Depository Trust Company for the accounts of its participants on or about _____, 2018.

Sole Book-Running Manager

BofA Merrill Lynch

The date of this prospectus supplement is June _____, 2018.

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ABOUT THIS PROSPECTUS SUPPLEMENT

This prospectus supplement and the accompanying prospectus form part of a registration statement on Form S-3 that we filed with the Securities and Exchange Commission, or the "SEC," using a "shelf" registration process. This document contains two parts. The first part consists of this prospectus supplement, which provides you with specific information about this offering. The second part, the accompanying prospectus, provides more general information, some of which may not apply to this offering. Generally, when we refer only to the "prospectus," we are referring to both parts combined together with all documents incorporated by reference.

In this prospectus supplement, the "Company," "we," "us" and "our" and similar terms refer to Exact Sciences Corporation and its subsidiaries. References to our "common stock" refer to the common stock of Exact Sciences Corporation.

This prospectus supplement, and the information incorporated herein by reference, may add, update or change information in the accompanying prospectus and in any free writing prospectuses we may provide to you in connection with this offering. You should read both this prospectus supplement and the accompanying prospectus together with additional information described under the headings "Where You Can Find More Information" and "Incorporation of Certain Information by Reference." If there is any inconsistency between the information in this prospectus supplement and the accompanying prospectus, you should rely on the information in this prospectus supplement.

You should rely only on the information contained in or incorporated by reference to this prospectus supplement and the accompanying prospectus. Neither we nor the underwriters have authorized any other person to provide information different from that contained in this prospectus supplement, the accompanying prospectus and the documents incorporated by reference herein and therein. If anyone provides you with different or inconsistent information, you should not rely on it. The information in this prospectus supplement, the accompanying prospectus and in any free writing prospectuses we may provide to you in connection with this offering is accurate only as of their respective dates, regardless of time of delivery. Our business, financial condition, results of operations and prospects may have changed since those dates.

We are offering to sell, and seeking offers to buy, our securities only in jurisdictions where offers and sales are permitted. The distribution of this prospectus supplement and the offering of the securities in certain jurisdictions may be restricted by law. Persons outside the United States who come into possession of this prospectus supplement must inform themselves about, and observe any restrictions relating to, the offering of the securities and the distribution of this prospectus supplement outside the United States. This prospectus supplement does not constitute, and may not be used in connection with, an offer to sell, or a solicitation of an offer to buy, any securities offered by this prospectus supplement by any person in any jurisdiction in which it is unlawful for such person to make such an offer or solicitation.

All references in this prospectus to our consolidated financial statements include, unless the context indicates otherwise, the related notes.

The industry and market data and other statistical information contained in this prospectus supplement, the accompanying prospectus and the documents we incorporate by reference are based on management's own estimates, independent publications, government publications, reports by market research firms or other published independent sources, and, in each case, are believed by management to be reasonable estimates. Although we believe these sources are reliable, we have not independently verified the information. None of the independent industry publications used in this prospectus supplement, the accompanying prospectus or the documents we incorporate by reference were prepared on our or our affiliates' behalf and none of the sources cited by us consented to the inclusion of any data from its reports, nor have we sought their consent.

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This prospectus and the documents incorporated by reference herein include trademarks, tradenames and service marks that are our property or the property of licensors or other third parties. Solely for convenience, such trademarks and tradenames may appear without any " " or "@" symbol. However, failure to include such symbols is not intended to suggest, in any way, that we will not assert our rights or the rights of any applicable licensor or other third party to such trademarks, tradenames and service marks.

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CAUTIONARY STATEMENT ABOUT FORWARD-LOOKING INFORMATION

Certain information set forth in this prospectus supplement, set forth in the accompanying prospectus or incorporated by reference herein or therein, may contain forward-looking statements within the meaning of Section 27A of the Securities Act of 1933, as amended (the "Securities Act"), and Section 21E of the Securities Exchange Act of 1934, as amended (the "Exchange Act"), that are intended to be covered by the "safe harbor" created by those sections. Forward-looking statements, which are based on certain assumptions and describe our future plans, strategies and expectations, can generally be identified by the use of forward-looking terms such as "believe," "expect," "may," "will," "should," "would," "could," "seek," "intend," "plan," "estimate," "goal," "anticipate," "project" or other comparable terms. All statements other than statements of historical facts included in this prospectus supplement regarding our strategies, prospects, financial condition, operations, costs, plans and objectives are forward-looking statements. Examples of forward-looking statements include, among others, statements we make regarding expected future operating results, anticipated results of our sales and marketing efforts, expectations concerning payer reimbursement and the anticipated results of our product development efforts. Forward-looking statements are neither historical facts nor assurances of future performance. Instead, they are based only on our current beliefs, expectations and assumptions regarding the future of our business, future plans and strategies, projections, anticipated events and trends, the economy and other future conditions. Because forward-looking statements relate to the future, they are subject to inherent uncertainties, risks and changes in circumstances that are difficult to predict and many of which are outside of our control. Our actual results and financial condition may differ materially from those indicated in the forward-looking statements. Therefore, you should not rely on any of these forward-looking statements. Important factors that could cause our actual results and financial condition to differ materially from those indicated in the forward-looking statements include, among others, the following: our ability to successfully and profitably market our products and services; the acceptance of our products and services by patients and healthcare providers; our ability to meet demand for our products and services; the willingness of health insurance companies and other payers to cover Cologuard and adequately reimburse us for our performance of the Cologuard test; the amount and nature of competition from other cancer screening and diagnostic products and services; the effects of the adoption, modification or repeal of any healthcare reform law, rule, order, interpretation or policy; the effects of changes in the pricing, coverage and reimbursement for our products and services, including without limitation as a result of the Protecting Access to Medicare Act of 2014; recommendations, guidelines and quality metrics issued by various organizations such as the U.S. Preventive Services Task Force, the American Cancer Society, and the National Committee for Quality Assurance regarding cancer screening or our products and services; our ability to successfully develop new products and services; our success establishing and maintaining collaborative, licensing, supplier and other arrangements; our ability to maintain regulatory approvals and comply with applicable laws, rules and regulations; and the other risks and uncertainties included in this prospectus supplement under the caption "Risk Factors," and those risks and uncertainties described in the documents incorporated by reference into this prospectus supplement and the accompanying prospectus. Therefore, you should not rely on any of these forward-looking statements. We urge you to consider those risks and uncertainties in evaluating our forward-looking statements. All subsequent written and oral forward-looking statements attributable to us or to persons acting on our behalf are expressly qualified in their entirety by the applicable cautionary statements. We further caution readers not to place undue reliance upon any such forward-looking statements, which speak only as of the date made. Except as otherwise required by the federal securities laws, we undertake no obligation to publicly update any forward-looking statement, whether written or oral, that may be made from time to time, whether as a result of new information, future developments or otherwise.

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SUMMARY

The information below is only a summary of more detailed information included elsewhere in or incorporated by reference in this prospectus supplement and the accompanying prospectus. This summary may not contain all the information that is important to you or that you should consider before making an investment decision. Please read this entire prospectus supplement and the accompanying prospectus, including the risk factors, as well as the information incorporated by reference in this prospectus supplement and the accompanying prospectus, carefully.

Our Company

We are a molecular diagnostics company currently focused on the early detection and prevention of some of the deadliest forms of cancer. We have developed an accurate, non-invasive, patient-friendly screening test called Cologuard for the early detection of colorectal cancer and pre-cancer, and we are currently working on the development of additional tests for other types of cancer, with the goal of becoming a leader in cancer diagnostics.

Our Cologuard® Test

Colorectal cancer is the second leading cause of cancer deaths in the United States and the leading cause of cancer deaths in the U.S. among non-smokers. Each year in the U.S. there are approximately:

140,000 new cases of colorectal cancer

51,000 deaths from colorectal cancer

Colorectal cancer treatment represents a significant, growing healthcare cost. As of 2010, \$14 billion was spent annually in the U.S. on colorectal cancer treatment, and the projected annual treatment costs are expected to be \$20 billion in 2020. The incidence of colorectal cancer in Medicare patients is expected to rise from 106,000 cases in 2010 to more than 180,000 cases in 2030.

It is widely accepted that colorectal cancer is among the most preventable, yet least prevented cancers. Colorectal cancer can take up to 10-15 years to progress from a pre-cancerous lesion to metastatic cancer and death. Patients who are diagnosed early in the progression of the disease with pre-cancerous lesions or polyps or early-stage cancer are more likely to have a complete recovery and to be treated less expensively. Accordingly, the American Cancer Society ("ACS") recommends that all people age 50 and older undergo regular colorectal cancer screening. Of the more than 85 million people in the U.S. for whom routine colorectal cancer screening is recommended, 38 percent have not been screened according to current guidelines. Poor compliance with screening guidelines has meant that nearly two-thirds of colorectal cancer diagnoses are made in the disease's late stages. The five-year survival rates for stages 3 and 4 are 70 percent and 13 percent, respectively. We believe the large underserved population of unscreened and inadequately screened patients represents a significant opportunity for a patient-friendly screening test.

Our Cologuard test is a non-invasive stool-based DNA ("sDNA") screening test that utilizes a multi-target approach to detect DNA and hemoglobin biomarkers associated with colorectal cancer and pre-cancer. Eleven biomarkers are targeted that have been shown to be strongly associated with colorectal cancer and pre-cancer. Methylation, mutation, and hemoglobin results are combined in the laboratory analysis through a proprietary algorithm to provide a single positive or negative reportable result.

On August 11, 2014 the U.S. Food and Drug Administration ("FDA") approved Cologuard for use as the first and only sDNA non-invasive colorectal cancer screening test. Our submission to the FDA for Cologuard included the results of our pivotal DeeP-C clinical trial that had over 10,000

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patients enrolled at 90 sites in the U.S. and Canada. The results of our DeeP-C clinical trial for Cologuard were published in the New England Journal of Medicine in April 2014. The peer-reviewed study, "Multi-target Stool DNA Testing for Colorectal-Cancer Screening," highlighted the performance of Cologuard in the trial population:

Cancer Sensitivity: 92%

Stage I and II Cancer Sensitivity: 94%

High-Grade Dysplasia Sensitivity: 69%

Specificity: 87%

The competitive advantages of sDNA screening may provide a significant market opportunity. If the test were used by 40-percent of the eligible U.S. screening population of 85 million people, at a three-year interval, and if average revenue per test was \$500, we estimate that our annual Cologuard revenue would be more than \$5.5 billion, annually.

Our Cologuard Commercialization Strategy

Our commercialization strategy includes three main elements focusing on physicians, patients, and payers.

Physicians and Patients

Our sales team actively engages with physicians and their staffs to emphasize the need for colorectal cancer screening, educate them on the value of Cologuard, and enroll them in our physician ordering system to enable them to prescribe the test. We focus on specific physicians based on a combination of their Cologuard order history and ordering potential and also on physician groups and larger regional and national health systems.

Securing inclusion in guidelines and quality measures is a key part of our physician engagement strategy since many physicians rely on such guidelines and quality measures when making screening recommendations. In June 2016, the US Preventive Services Task Force ("USPSTF") issued an updated recommendation statement for colorectal cancer screening and gave an "A" grade to colorectal cancer screening starting at age 50 and continuing until age 75. The statement specifies seven screening methods, including FIT-DNA (which is Cologuard).

Many professional colorectal cancer screening guidelines in the U.S., including those of the ACS and the National Comprehensive Cancer Network ("NCCN"), recommend regular screening using any of a variety of methods. Since 2008, joint colorectal cancer screening guidelines endorsed by the ACS have included sDNA screening technology as a screening option for the detection of colorectal cancer in average risk, asymptomatic individuals age 50 and older. In October 2014, the ACS updated its colorectal cancer screening guidelines to specifically include Cologuard as a recommended screening test. In June 2016, the NCCN updated its Colorectal Cancer Screening Guidelines to add sDNA screening, at a once-every-three-years interval, to its list of recommended screening tests.

In October 2016, the National Committee for Quality Assurance ("NCQA") included Cologuard testing on a three-year interval in the 2017 Healthcare Effectiveness Data and Information Set ("HEDIS") measures. More than 90 percent of America's health plans measure quality based on HEDIS. In April 2017, the Centers for Medicare & Medicaid Services ("CMS") included Cologuard in its updated 2018 Medicare Advantage Star Ratings program.

A critical part of the value proposition of Cologuard is our compliance program, which involves active engagement with patients and providers. This customer-service-oriented activity is focused on helping patients to complete Cologuard tests that have been ordered for them by their providers.

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After the launch of Cologuard, we initiated a significant public relations effort to engage patients in the United States, and launched demographically-targeted direct-to-patient advertising campaigns in digital, social, print and other channels. In 2016, we began a national television advertising campaign, with a majority of placements in national cable and syndicated programming widely viewed by our target patient demographic. In the second and third quarters of 2017, we extended that campaign with new 30-second commercials intended to increase the cost effectiveness of our broadcast media. During 2018, we plan to maintain our current television advertising efforts, accelerate our investment in digital and social media, and embark upon strategic partnerships designed to increase awareness for Cologuard.

Payers

The cornerstone of our payer-engagement strategy was securing coverage from CMS. Medicare covers approximately 47 percent of patients in the screening population for Cologuard. On October 9, 2014, CMS issued a National Coverage Determination ("NCD") for Cologuard following a parallel review process with the FDA. Cologuard was the first screening test approved by the FDA and covered by CMS through that process. As outlined in the NCD, Medicare Part B covers Cologuard once every three years for beneficiaries who meet all of the following criteria:

age 50 to 85 years,

asymptomatic (no signs or symptoms of colorectal disease including but not limited to lower gastrointestinal pain, blood in stool, positive guaiac fecal occult blood test or fecal immunochemical test), and

at average risk for developing colorectal cancer (e.g., no personal history of adenomatous polyps, colorectal cancer, or inflammatory bowel disease, including Crohn's Disease and ulcerative colitis; no family history of colorectal cancers or adenomatous polyps, familial adenomatous polyposis or hereditary non-polyposis colorectal cancer).

Pursuant to the 2017 Clinical Laboratory Fee Schedule, CMS reimbursed Cologuard at the rate of \$512.43 per test. Under the Protecting Access to Medicare Act of 2014 ("PAMA"), effective January 1, 2018, the CMS reimbursement rate for Cologuard was set at \$508.87, which was the volume-weighted median of private payer rates for Cologuard for the period from January 1, 2016 to June 30, 2016. We expect that the CMS reimbursement rate established for 2018 will remain in place for three years and then be reset based on the volume-weighted median of private payer rates for Cologuard during the data collection period from January 1, 2019 to June 30, 2019. Payments from CMS are currently subject to sequestration.

In addition to Medicare reimbursement, we seek to secure favorable coverage and in-network reimbursement agreements from commercial payers. Most commercial payers have issued positive coverage decisions for Cologuard, and we have entered into contracts with several payers to include Cologuard as an in-network service. In-network agreements with payers have varying terms and conditions, including reimbursement rate, term and termination. From time to time in the ordinary course of our business, we may enter into new agreements, certain existing agreements may expire without renewal and certain other existing agreements may be terminated early by us or the third-party payer. We believe that commercial payers' reimbursement of Cologuard will depend on a number of factors, including payers' determination that it is: sensitive and specific for colorectal cancer; not experimental or investigational; approved or recommended by major organizations' guidelines; reliable, safe and effective; medically necessary; appropriate for the specific patient; and cost-effective. Reimbursement may also be affected by whether Cologuard is in-network for a given payer. Also, some payers may apply various medical management requirements, including a requirement that they give prior authorization for a Cologuard test before they are willing to pay for it. Other payers may perform post-payment reviews or audits, which could lead to payment recoupments. Medical management, such

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as prior authorizations and post-payment review or audits, may require that we, patients, or physicians provide the payer with extensive medical records and other information.

Coverage of Cologuard may also depend, in whole or in part, on whether payers determine, or courts and/or regulatory authorities determine, coverage is required under applicable federal or state laws mandating coverage of certain colorectal cancer screening services. For example, Section 2713 of the Patient Protection and Affordable Care Act ("ACA") mandates that certain health insurers cover evidence-based items or services that have in effect a rating of "A" or "B" in the current recommendations of USPSTF without imposing any patient cost-sharing ("ACA Mandate"). Similarly, federal regulations require that Medicare Advantage plans cover "A" or "B" rated preventive services without patient cost-sharing. Following the June 2016 update to the USPSTF colorectal cancer screening recommendation statement, CMS issued an updated Evidence of Coverage notice for Medicare Advantage plans that affirms such plans must include coverage of Cologuard every three years without patient cost-sharing. While we believe the ACA Mandate will require most health insurers to cover Cologuard without patient cost-sharing (following an initial phase-in period between one and two years from the date of the updated USPSTF recommendation statement depending on the date a given plan year commences), it is possible that certain health insurers will disagree and determine not to cover Cologuard. It may be difficult for us or patients to enforce the ACA Mandate directly, and we may need to rely on states to take enforcement action, which they may choose not to do. It is also possible that the ACA Mandate will be repealed or significantly modified in the future.

We believe quality metrics may influence payers' coverage and contracting decisions, as well as physicians' cancer screening procedures. Some government and private payers are adopting pay-for-performance programs that differentiate payments for healthcare services based on the achievement of documented quality metrics, cost efficiencies or patient outcomes. Payers may look to quality measures such as HEDIS and CMS Star Ratings, to assess quality of care. We believe inclusion in the HEDIS measures and Star Ratings measures may have a positive impact on payers' willingness to reimburse Cologuard, as well as on physicians' willingness to prescribe the test.

Our Clinical Lab Facilities

As part of our commercialization strategy, we established a state-of-the-art, highly automated lab facility that is certified pursuant to federal Clinical Laboratory Improvement Amendments ("CLIA") requirements to process Cologuard tests and provide patient results. Our commercial lab operation is housed in a 50,000 square foot facility in Madison, Wisconsin. At our lab, we currently have the capacity to process approximately two million tests per year. We are expanding our current facility to increase our lab processing capacity to more than three million tests per year around the end of 2018.

During the fourth quarter of 2017, we began construction of a new clinical lab facility in Madison, Wisconsin that is expected to be completed around mid-2019. We expect our total lab capacity at both facilities will be approximately five million tests per year around the end of 2019.

Product Pipeline

We also are developing a pipeline of potential future products and services. We are continuing to collaborate with MAYO Foundation for Medical Education and Research ("MAYO"), our development partner for Cologuard, on developing new tests, with the goal of becoming a leader in the early detection of cancer. We believe our proprietary technology platform provides a strong foundation for the development of additional cancer diagnostic tests. Through our collaboration with MAYO, we have identified proprietary biomarkers for several major cancers, including liver cancer and lung cancer. We have successfully performed validation studies on tissue samples for seven major cancers and on blood samples for four major cancers.

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The ACS estimates that liver cancer will be diagnosed in 42,000 Americans and cause 30,000 deaths in 2018, three-fourths of which will be hepatocellular carcinoma ("HCC"). Incidence and mortality rates are both increasing at approximately 3 percent per year. People who have been diagnosed with cirrhosis of the liver or Hepatitis B are at high risk of developing HCC. Evidence shows that HCC surveillance in these high-risk groups leads to earlier detection and improved outcomes. The NCCN and American Association for the Study of Liver Diseases ("AASLD") guidelines recommend that these two groups be surveilled for HCC every six months using ultrasound and the blood-based biomarker alpha-fetoprotein ("AFP"). However, ultrasound and AFP are documented to have poor sensitivity for early stage cancer, which is the primary target of surveillance. We are currently seeking to develop a blood-based biomarker test to serve as an alternative to ultrasound and AFP for use in HCC surveillance. We published a small case-control study in 2016 showing high accuracy for detecting HCC using a blood-based panel of methylation markers.

The ACS estimates that, in the United States in 2018, lung cancer will be diagnosed in 234,000 people and cause 154,000 deaths. Currently, more than half of lung cancer cases are diagnosed at an advanced stage, after symptoms appear, when the five-year survival rate is in the low single digits. We are currently seeking to develop a blood-based biomarker test to aid in the early detection of lung cancer in individuals with lung nodules discovered through a computerized tomography ("CT") or other scan. Such a test may help reduce the number of unnecessary biopsies and other follow-up procedures, and thereby reduce costs and improve health outcomes.

We also continue to explore opportunities for improving Cologuard, including improvements that could lower our cost of goods or expand the usage of Cologuard to different patient populations.

Corporate Information

Our executive offices are located at 441 Charmany Drive, Madison, Wisconsin 53719. Our telephone number is (608) 284-5700. Our Internet website address is www.exactsciences.com. Our Internet website and the information contained therein or connected thereto are not part of this prospectus supplement or the accompanying prospectus.

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THE OFFERING

The summary below describes the principal terms of the notes. Certain of the terms and conditions described below are subject to important limitations and exceptions. For a more detailed description of the terms and conditions of the notes, see the section entitled "Description of the Notes." With respect to the discussion of the terms of the notes on the cover page, in this section and in the section entitled "Description of the Notes," references to "the Company," "we," "our," and "us" refer solely to Exact Sciences Corporation and not its subsidiaries.

Issuer: Exact Sciences Corporation

Notes Offered: \$150,000,000 aggregate principal amount of 1.0% Convertible Senior Notes due 2025. We have granted the underwriters an option to purchase up to an additional \$22,500,000 aggregate principal amount of notes solely to cover over-allotments. The notes we are offering will be issued as additional notes under the indenture pursuant to which we previously issued \$690,000,000 aggregate principal amount of our 1.0% Convertible Senior Notes due 2025 in January 2018, which we refer to as the initial notes. The notes offered hereby will have identical terms as the initial notes, will be treated as a single series of securities with the initial notes under the indenture and will have the same CUSIP number as the initial notes. Holders of the notes offered hereby and the initial notes will vote as one class under the indenture.

Issue Price: % of the principal amount, plus accrued interest from, and including, January 17, 2018.

Maturity Date: January 15, 2025, unless earlier converted.

Interest and Payment Dates: 1.0% per year, payable semi-annually in arrears on January 15 and July 15 of each year, beginning July 15, 2018. We will pay additional interest, if any, at our election as the sole remedy relating to the failure to comply with our reporting obligations as described under "Description of the Notes Events of Default; Notice and Waiver."

Ranking: The notes will be our senior unsecured obligations that will:

rank senior in right of payment to all of our future indebtedness that is expressly subordinated in right of payment to the notes;

rank equally in right of payment to all of our future liabilities that are not so subordinated;

be effectively junior to all of our existing and future secured indebtedness and other secured obligations, to the extent of the value of the assets securing that indebtedness and other secured obligations; and

be structurally subordinated to all indebtedness and other liabilities (including trade payables) of our subsidiaries.
See "Description of the Notes Ranking."

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As of March 31, 2018, our total consolidated indebtedness (including the outstanding principal amount of the initial notes) was approximately \$694.4 million, \$4.4 million of which was secured. As of March 31, 2018, our subsidiaries had \$8.3 million of indebtedness and other liabilities (including trade payables). After giving effect to the issuance of the notes offered hereby (assuming no exercise of the underwriters' option to purchase additional notes), our total consolidated indebtedness would have been approximately \$844.2 million as of March 31, 2018.

The indenture that will govern the notes does not limit our ability to incur additional indebtedness in the future, including senior secured indebtedness.

Conversion Rights:

Prior to the close of business on the business day immediately preceding July 15, 2024, you may, at your option, convert your notes, in multiples of \$1,000 principal amount, but only under the following circumstances:

on any date during any calendar quarter (and only during such calendar quarter) if the closing sale price (as defined in "Description of the Notes Conversion Rights Conversion Upon Satisfaction of Sale Price Condition") of our common stock was more than 130% of the applicable conversion price on each applicable trading day for at least 20 trading days (whether or not consecutive) in the period of the 30 consecutive trading days ending on the last trading day of the immediately preceding calendar quarter;

during a specified period if we distribute to all or substantially all holders of our common stock any rights, options or warrants (other than pursuant to a stockholders rights plan) entitling them to purchase, for a period of 45 calendar days or less from the announcement date for such distribution, shares of our common stock at a price per share less than the average of the closing sale prices for the ten consecutive trading day period ending on, and including, the trading day immediately preceding the announcement date for such distribution;

during a specified period if we distribute to all or substantially all holders of our common stock cash or other assets, debt securities or rights to purchase our securities (other than pursuant to a stockholders rights plan), which distribution has a per share value exceeding 10% of the closing sale price of our common stock on the trading day immediately preceding the announcement date for such distribution;

during a specified period if a fundamental change occurs or if we engage in certain corporate transactions; or

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during the five business day period following any five consecutive trading day period in which the trading price (as defined in "Description of the Notes Conversion Rights Conversion Upon Satisfaction of Trading Price Condition") per \$1,000 principal amount of notes for each trading day during such five trading day period was less than 98% of the product of the closing sale price of our common stock and the applicable conversion rate on each such trading day.

On or after July 15, 2024 until the close of business on the second scheduled trading day immediately preceding the maturity date, you may, at your option, convert all or any portion of your notes, in multiples of \$1,000 principal amount, at any time, regardless of the foregoing circumstances.

The conversion rate for the notes is initially 13.2569 shares of common stock per \$1,000 principal amount of notes (equivalent to an initial conversion price of approximately \$75.43 per share of common stock), subject to adjustment as described in this prospectus supplement.

Settlement upon Conversion:

Upon conversion, we will pay or deliver, as the case may be, cash, shares of our common stock (and cash in lieu of fractional shares) or a combination of cash and shares of our common stock, at our election. If we satisfy our conversion obligation solely in cash or through payment and delivery, as the case may be, of a combination of cash and shares of our common stock, the amount of cash and shares of common stock, if any, due upon conversion will be based on a daily conversion value or a daily settlement amount, as applicable (as described herein) calculated on a proportionate basis for each VWAP trading day (as defined herein) in a 30 VWAP trading day observation period (as defined herein). You will not receive any separate cash payment or additional shares for interest, if any, accrued and unpaid to the conversion date, except in limited circumstances. Instead, interest will be deemed paid by the cash, shares of our common stock or a combination of cash and shares of our common stock paid or delivered, as the case may be, to you upon conversion of a note. See "Description of the Notes Conversion Rights."

No Redemption:

We may not redeem the notes prior to the maturity date, and no "sinking fund" is provided for the notes, which means we are not required to redeem or retire the notes periodically.

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Adjustment to Conversion Rate Upon a Make-Whole Fundamental Change:

If the effective date (as defined herein) of a make-whole fundamental change (as defined herein) occurs prior to the maturity date of the notes and a holder elects to convert its notes in connection with such make-whole fundamental change, we will increase the conversion rate by a number of additional shares. The number of additional shares will be determined by reference to the table in "Description of the Notes Conversion Rights Adjustment to Conversion Rate Upon a Make-Whole Fundamental Change," based on the effective date and the price paid (or deemed paid) per share of our common stock in such make-whole fundamental change. If holders of our common stock receive only cash in a make-whole fundamental change, the price paid (or deemed paid) per share will be the cash amount paid per share. Otherwise, the price paid (or deemed paid) per share will be equal to the average of the closing sale prices of our common stock over the five trading day period ending on, and including, the trading day immediately preceding the effective date of such make-whole fundamental change.

Fundamental Change Repurchase Right of Holders:

If we undergo a fundamental change (as defined under "Description of the Notes Fundamental Change Put") prior to maturity of the notes, subject to certain conditions, you will have the right, at your option, to require us to repurchase for cash all or a portion of your notes in minimum principal amounts of \$1,000 or whole multiples thereof at a repurchase price equal to 100% of the principal amount of the notes being repurchased, plus accrued and unpaid interest to, but excluding, the fundamental change repurchase date. See "Description of the Notes Fundamental Change Put."

Book-Entry Form:

The notes will be issued in book-entry form and will be represented by permanent global certificates deposited with, or on behalf of, The Depository Trust Company, or DTC, and registered in the name of Cede & Co. as DTC's nominee. Beneficial interests in any of the notes will be shown on, and transfers will be effected only through, records maintained by DTC or its nominee and any such interest may not be exchanged for certificated securities, except in limited circumstances.

Listing:

The initial notes are not listed or quoted, and we do not intend to apply to have the notes listed or quoted on any securities exchange or any automated dealer quotation system. Our common stock is listed on the NASDAQ Capital Market under the symbol "EXAS".

Use of Proceeds:

We estimate that our net proceeds from this offering, after deducting underwriting discounts and estimated offering fees and expenses, will be approximately \$ (or \$ if the underwriters exercise in full their option to purchase \$22,500,000 in additional notes from us).

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Trustee:

We intend to use the net proceeds we receive from this offering for general corporate purposes, including working capital and possible acquisitions. See "Use of Proceeds." U.S. Bank National Association

Public Market for the Notes

We cannot assure you that an active trading market for the notes will be developed or maintained. See "Underwriting New Issue of Notes." We have been advised by the underwriter that it presently intends to make a market in the notes after completion of the offering. However, the underwriter is under no obligation to do so and may discontinue any market-making activities at any time without notice. We do not intend to apply for a listing of the notes on any securities exchange or any automated dealer quotation system.

Risk Factors:

You should read the "Risk Factors" beginning on page S-14 of this prospectus supplement and on page 16 our Annual Report on Form 10-K for the year ended December 31, 2017 and the other information included or incorporated by reference in this prospectus supplement and the accompanying prospectus for a discussion of factors you should consider carefully before deciding to invest in the notes.

U.S. Federal Income Tax Considerations:

For the U.S. federal income tax considerations of the holding, disposition and conversion of the notes, and the holding and disposition of our common stock, see "Material U.S. Federal Income Tax Considerations."

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RISK FACTORS

We operate in a rapidly changing environment that involves a number of risks, some of which are beyond our control. Before deciding whether to invest in the notes offered hereby, you should consider carefully the risk factors related to our indebtedness and the notes described below and the other risk factors related to our business incorporated herein by reference to our Annual Report on Form 10-K for the year ended December 31, 2017. These are the risks and uncertainties we believe are most important for you to consider. We cannot be certain that we will successfully address these risks. If we are unable to address these risks, our business may not grow, our stock price may suffer and we may be unable to stay in business. Additional risks and uncertainties not presently known to us, which we currently deem immaterial or which are similar to those faced by other companies in our industry or business in general, may also impair our business operations.

As used in this section of the prospectus supplement, the term "notes" includes the initial notes, unless the context requires otherwise.

Risks Related to Our Indebtedness and the Notes

The notes are our unsecured senior obligations and will be effectively junior to all of our existing and future secured indebtedness and other secured obligations, to the extent of the value of the assets securing that indebtedness, and be structurally subordinated to all indebtedness and other liabilities (including trade payables) of our subsidiaries.

The notes are our senior unsecured obligations and will effectively rank junior in right of payment to any of our secured indebtedness and other secured obligations to the extent of the value of the assets securing such indebtedness and be structurally subordinated to all indebtedness and other liabilities (including trade payables, but excluding intercompany obligations) of our subsidiaries. In the event of our bankruptcy, liquidation, reorganization or other winding up, our assets that secure indebtedness or other obligations effectively ranking senior in right of payment to the notes will be available to pay obligations on the notes only after the secured indebtedness or other secured obligations has been repaid in full from these assets. There may not be sufficient assets remaining to pay amounts due on any or all of the notes then outstanding. The indenture governing the notes does not prohibit us from incurring additional indebtedness in the future, including senior secured indebtedness, and such indebtedness may be substantial, nor will the indenture prohibit our subsidiaries from incurring additional indebtedness or other liabilities. See "Description of the Notes Ranking."

As of March 31, 2018, our total consolidated indebtedness (including the outstanding principal amount of the initial notes) was approximately \$694.4 million, \$4.4 million of which was secured. As of March 31, 2018, our subsidiaries had approximately \$8.3 million of indebtedness and other liabilities (including trade payables). After giving effect to the issuance of the notes offered hereby (assuming no exercise of the underwriters' option to purchase additional notes), our total consolidated indebtedness would have been approximately \$844.4 million as of March 31, 2018.

Our increased indebtedness could adversely affect our business, financial condition and results of operations and our ability to meet our payment obligations under such indebtedness.

As of March 31, 2018, we had approximately \$694.4 million of total consolidated indebtedness outstanding. After giving effect to this offering of notes, our indebtedness will increase by \$150.0 million (or \$172.5 million if the underwriters exercise their option to purchase additional notes in full) (which amount, with respect to the notes, reflects the face amount of the notes). The indenture that will govern the notes does not restrict our ability to incur additional indebtedness. This level of debt could have significant consequences on our future operations, including:

increasing our vulnerability to adverse economic and industry conditions;

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making it more difficult for us to meet our payment and other obligations under the notes;

making it more difficult to obtain any necessary future financing for working capital, capital expenditures, debt service requirements or other purposes;

requiring the dedication of a substantial portion of any cash flow from operations to service our indebtedness, thereby reducing the amount of cash flow available for other purposes, including capital expenditures;

placing us at a possible competitive disadvantage with competitors that are less leveraged than us or have better access to capital than we have; and

limiting our flexibility in planning for, or reacting to, changes in our business and the markets in which we compete.

Any of the above-listed factors could have an adverse effect on our business, financial condition and results of operations and our ability to meet our payment obligations under the notes.

Our ability to meet our payment and other obligations under the notes depends on our ability to generate significant cash flow in the future. This, to some extent, is subject to general economic, financial, competitive, legislative and regulatory factors as well as other factors that are beyond our control. We cannot assure you that our business will generate cash flow from operations, or that future borrowings will be available to us, in an amount sufficient to enable us to meet our payment obligations under the notes and to fund other liquidity needs. If we are not able to generate sufficient cash flow to service our debt obligations, we may need to refinance or restructure our debt, including the notes, sell assets, reduce or delay capital investments, or seek to raise additional capital. If we are unable to implement one or more of these alternatives, we may not be able to meet our payment obligations under the notes, and this default could cause us to be in default on any other currently existing or future outstanding indebtedness.

We may incur substantially more debt or take other actions which would intensify the risks discussed above.

We and our subsidiaries may be able to incur substantial additional debt in the future, some of which may be secured debt. We and our subsidiaries are not restricted under the terms of the indenture governing the notes from incurring additional debt, securing existing or future debt, recapitalizing our debt or taking a number of other actions that are not limited by the terms of the indenture governing the notes that could have the effect of diminishing our ability to make payments on the notes when due.

Servicing our debt requires a significant amount of cash, and we may not have sufficient cash flow from our business to pay amounts due under our indebtedness, including the notes.

Our ability to make scheduled payments of the principal of, to pay interest on or to refinance our indebtedness, including the notes, depends on our future performance, which is subject to economic, financial, competitive and other factors beyond our control. Our business may not continue to generate cash flow from operations in the future sufficient to service our debt, including the notes, and make necessary capital expenditures. If we are unable to generate such cash flow, we may be required to adopt one or more alternatives, such as selling assets, restructuring debt or obtaining additional equity capital on terms that may be onerous or highly dilutive. Our ability to refinance our indebtedness will depend on the capital markets and our financial condition at such time. We may not be able to engage in any of these activities or engage in these activities on desirable terms, which could result in a default on our debt obligations.

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The terms of the notes will not contain restrictive covenants and provide only limited protection in the event of a change of control.

The indenture under which the notes will be issued does not contain restrictive covenants that would protect you from several kinds of transactions that may adversely affect you. In particular, the indenture does not contain covenants that will limit our ability to pay dividends or make distributions on or redeem our capital stock or limit our ability to incur additional indebtedness and, therefore, may not protect you in the event of a fundamental transaction, a highly leveraged transaction or other similar transaction. The requirement that we offer to repurchase the notes upon a change of control is limited to the transactions specified in the definition of a "fundamental change" under "Description of the Notes Fundamental Change Put." Similarly, the circumstances under which we are required to adjust the conversion rate upon the occurrence of a "make-whole fundamental change" are limited to circumstances where a note is converted in connection with such a transaction as set forth under "Description of the Notes Conversion Rights Adjustment to Conversion Rate Upon a Make-Whole Fundamental Change."

Accordingly, subject to restrictions contained in any future debt agreements, we could enter into certain transactions, such as acquisitions, refinancings or recapitalizations that could affect our capital structure and the value of the notes and common stock but would not constitute a fundamental change under the notes.

Some significant restructuring transactions may not constitute a fundamental change, in which case we would not be obligated to offer to repurchase the notes.

Upon the occurrence of a fundamental change, you have the right to require us to repurchase your notes. However, the fundamental change provisions will not afford protection to holders of the notes in the event of certain transactions. For example, transactions such as leveraged recapitalizations, refinancings, restructurings or acquisitions initiat