



# Spark Therapeutics and Pfizer Announce Longer-term Preliminary Data Showing Consistent and Sustained Factor IX Levels in Hemophilia B at 59th American Society of Hematology (ASH) Annual Meeting and Exposition

Monday, December 11, 2017 - 02:00am

Data include eleventh participant in the trial Annualized bleeding rate (ABR) for all 11 participants was reduced by 97 percent, while annualized infusion rate (AIR) was reduced 99 percent as of Nov. 29, 2017 data cutoff

Spark Therapeutics (NASDAQ:ONCE), a fully integrated gene therapy company dedicated to challenging the inevitability of genetic disease, and Pfizer Inc. (NYSE:PFE), today announced that, with a cumulative follow-up of more than 13 patient years of observation, all 11 participants in the ongoing Phase 1/2 clinical trial of investigational *SPK-9001* for the treatment of patients with hemophilia B had discontinued routine infusions of factor IX concentrates and shown sustained steady-state factor IX activity levels with no serious adverse events, thrombotic events or factor IX inhibitors observed. Based on individual participant history for the year prior to the study, the overall annualized bleeding rate (ABR) was reduced by 97 percent (calculated based on data after week four; 95 percent based on data after infusion) to a mean of 0.3 (0.5) annual bleeds, compared to a mean of 10.5 bleeds annually before *SPK-9001*

administration. Overall annualized infusion rate (AIR) was reduced 99 percent (calculated based on data after week four; 97 percent based on data after infusion) to a mean of 0.8 (1.7) annual infusions, compared to a mean of 62.5 infusions per year before *SPK-9001* administration. Data on all 11 participants were presented today by Lindsey A. George, M.D., attending physician in the Division of Hematology at Children's Hospital of Philadelphia and principal investigator of the trial, at the 59th American Society of Hematology (ASH) Annual Meeting and Exposition in Atlanta.

"We believe these longer-term data are meeting critical goals of our hemophilia programs," said Katherine A. High, M.D., president and head of research and development at Spark Therapeutics. "Now, with four participants followed for more than 18 months, we continue to see consistent levels of factor IX activity, no serious adverse events, as well as a sustained reduction in both the symptoms of hemophilia and the prophylactic and disease management protocols that were used prior to infusion with *SPK-9001*."

As of the Nov. 29, 2017 data cutoff, the mean steady-state factor IX activity level at 12 weeks post-administration for the 11 participants was 36 percent of normal (range as of the data cutoff: 15 to 78 percent). As of the data cutoff, the last participant to be infused, who received *SPK-9001* manufactured using an enhanced process, was out eight months following *SPK-9001* infusion, with a mean factor IX activity level of 60 percent. Spark Therapeutics will enroll up to four additional participants in the current Phase 1/2 clinical trial who will receive *SPK-9001* manufactured using an enhanced process to test its comparability to the *SPK-9001* received by the first 10 participants enrolled in the ongoing trial.

In this open-label, non-randomized and multicenter Phase 1/2 clinical trial, there were no serious adverse events during or following infusion with *SPK-9001*, and no participants experienced thrombotic events or developed factor IX inhibitors. Two participants developed an asymptomatic and transient increase in liver enzymes that resolved with a tapering dose of oral corticosteroids. One participant with severe joint disease administered factor for suspected joint bleeding, while a second participant recorded one spontaneous bleed.

### **About Hemophilia B**

Hemophilia, a rare genetic bleeding disorder that causes the blood to take a long time to clot because of a deficiency in one of several blood clotting factors, is almost exclusively found in males. People with hemophilia are at risk for excessive and recurrent bleeding from modest injuries, which have the potential to be life threatening. People with severe hemophilia often bleed spontaneously into their muscles or joints, or rarely into other

critical closed spaces such as the intracranial space, where bleeding can be fatal. The incidence of hemophilia B is one in 25,000 male births. People with hemophilia B have a deficiency in clotting factor IX, a specific protein in the blood. Hemophilia B also is called congenital factor IX deficiency or Christmas disease. The current standard of care requires recurrent intravenous infusions of either plasma-derived or recombinant factor IX to control and prevent bleeding episodes. There exists a significant need for novel therapeutics to treat people living with hemophilia.

### **About the *SPK-FIX* Program and *SPK-9001***

*SPK-9001* is a novel investigational vector that contains a bio-engineered adeno-associated virus (AAV) capsid and a codon-optimized, high-activity human factor IX gene enabling endogenous production of factor IX.

Spark Therapeutics and Pfizer entered into a collaboration in December 2014 for the *SPK-FIX* program, including *SPK-9001*, under which Spark Therapeutics is responsible for conducting all Phase 1/2 studies for any product candidates, while Pfizer will assume responsibility for pivotal studies, any regulatory activities and potential global commercialization of any products that may result from the collaboration.

### **About Spark Therapeutics**

At Spark Therapeutics, a fully integrated company committed to discovering, developing and delivering gene therapies, we challenge the inevitability of genetic diseases, including blindness, hemophilia and neurodegenerative diseases. We have successfully applied our technology directed to the retina and liver, and currently have four programs in clinical trials or under regulatory review, including the first potential gene therapy for a genetic disease in the United States and product candidates that have shown promising early results in patients with hemophilia. At Spark, we see the path to a world where no life is limited by genetic disease. For more information, visit [www.sparktx.com](http://www.sparktx.com), and follow us on Twitter and LinkedIn.

### **Spark Therapeutics Cautionary note on forward-looking statements**

This release contains "forward-looking statements" within the meaning of the Private Securities Litigation Reform Act of 1995, including statements regarding the company's *SPK-FIX* program. The words "anticipate," "believe," "expect," "intend," "may," "plan," "predict," "will," "would," "could," "should," "continue" and similar expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words. We may not actually achieve the plans, intentions or expectations disclosed in our forward-looking statements, and you should not place undue reliance on our forward-looking statements. Any forward-looking

statements are based on management's current expectations of future events and are subject to a number of risks and uncertainties that could cause actual results to differ materially and adversely from those set forth in, or implied by, such forward-looking statements. These risks and uncertainties include, but are not limited to, the risk that: (i) our lead *SPK-FIX* product candidate, *SPK-9001*, may not produce sufficient data in our Phase 1/2 clinical trial to warrant further development; (ii) our overall collaboration with Pfizer may not be successful; and (iii) any one or more of our product candidates in preclinical or clinical development will not successfully be developed and commercialized. For a discussion of other risks and uncertainties, and other important factors, any of which could cause our actual results to differ from those contained in the forward-looking statements, see the "Risk Factors" section, as well as discussions of potential risks, uncertainties and other important factors, in our Annual Report on Form 10-K, our Quarterly Reports on Form 10-Q and other filings we make with the U.S. Securities and Exchange Commission. All information in this press release is as of the date of the release, and Spark undertakes no duty to update this information unless required by law.

#### **About Pfizer: Working together for a healthier world®**

At Pfizer, we apply science and our global resources to bring therapies to people that extend and significantly improve their lives. We strive to set the standard for quality, safety and value in the discovery, development and manufacture of health care products. Our global portfolio includes medicines and vaccines as well as many of the world's best-known consumer health care products. Every day, Pfizer colleagues work across developed and emerging markets to advance wellness, prevention, treatments and cures that challenge the most feared diseases of our time. Consistent with our responsibility as one of the world's premier innovative biopharmaceutical companies, we collaborate with health care providers, governments and local communities to support and expand access to reliable, affordable health care around the world. For more than 150 years, we have worked to make a difference for all who rely on us. We routinely post information that may be important to investors on our website at [www.pfizer.com](http://www.pfizer.com). In addition, to learn more, please visit us on [www.pfizer.com](http://www.pfizer.com) and follow us on Twitter at @Pfizer and @Pfizer\_News, LinkedIn, YouTube and like us on Facebook at [Facebook.com/Pfizer](https://www.facebook.com/Pfizer).

*Pfizer Disclosure Notice:* The information contained in this release is as of December 11, 2017. Pfizer assumes no obligation to update forward-looking statements contained in this release as the result of new information or future events or developments.

This release contains forward-looking information about *SPK-9001* and the *SPK-FIX* program, including their potential benefits, that involves substantial risks and uncertainties that could cause actual results to differ materially from those expressed or

implied by such statements. Risks and uncertainties include, among other things, the uncertainties inherent in research and development, including the ability to meet anticipated clinical study commencement and completion dates as well as the possibility of unfavorable study results, including unfavorable new clinical data and additional analyses of existing clinical data; risks associated with initial data, including the risk that the final results of the Phase 1/2 study for *SPK-9001* and/or additional clinical trials may be different from (including less favorable than) the initial data results and may not support further clinical development; the risk that clinical trial data are subject to differing interpretations, and, even when we view data as sufficient to support the safety and/or effectiveness of a product candidate, regulatory authorities may not share our views and may require additional data or may deny approval altogether; whether regulatory authorities will be satisfied with the design of and results from our clinical studies; whether and when any applications may be filed with regulatory authorities for *SPK-9001*; whether and when regulatory authorities may approve any such applications, which will depend on the assessment by such regulatory authorities of the benefit-risk profile suggested by the totality of the efficacy and safety information submitted and, if approved, whether *SPK-9001* will be commercially successful; decisions by regulatory authorities regarding labeling and other matters that could affect the availability or commercial potential of *SPK-9001*; and competitive developments.

A further description of risks and uncertainties can be found in Pfizer's Annual Report on Form 10-K for the fiscal year ended December 31, 2016 and in its subsequent reports on Form 10-Q, including in the sections thereof captioned "Risk Factors" and "Forward-Looking Information and Factors That May Affect Future Results," as well as in its subsequent reports on Form 8-K, all of which are filed with the U.S. Securities and Exchange Commission and available at [www.sec.gov](http://www.sec.gov) and [www.pfizer.com](http://www.pfizer.com).

Investor Relations Contact:

Ryan Asay

[ryan.asay@sparktx.com](mailto:ryan.asay@sparktx.com)

(215) 239-6424

Media Contact: Monique da Silva

[communications@sparktx.com](mailto:communications@sparktx.com) (215) 282-7470