A photograph of a woman and a young child embracing and smiling. The woman is on the right, with her arm around the child's shoulder. Both are wearing white shirts. The background is a warm, textured wall with a decorative pattern of golden, leaf-like shapes at the top.

EDG-5506: A Novel Approach to Protect Muscle in Duchenne Muscular Dystrophy

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End Duchenne Tour
Detroit
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Forward-Looking Statements

This presentation contains forward-looking statements that involve substantial risks and uncertainties of Edgewise Therapeutics, Inc. (“Edgewise” or the “Company”). All statements other than statements of historical facts contained in this presentation, including statements regarding our future financial condition, results of operations, business strategy and plans, and objectives of management for future operations, as well as statements regarding industry trends, are forward-looking statements. Such forward-looking statements include, among other things, statements regarding the potential of, and expectations regarding, Edgewise’s drug discovery platform; Edgewise’s product candidates and programs, including EDG-5506; the expected milestones and timing of such milestones for EDG-5506 including the expected timing of reporting of data for EDG-5506 and clinical trials; statements regarding the market opportunity for Edgewise’s product candidates; statements regarding Edgewise’s pipeline of product candidates and programs; and statements regarding Edgewise’s financial position including its liquidity. In some cases, you can identify forward-looking statements by terminology such as “estimate,” “intend,” “may,” “plan,” “potentially” “will” or the negative of these terms or other similar expressions.

We have based these forward-looking statements largely on our current expectations and projections about future events and trends that we believe may affect our financial condition, results of operations, business strategy and financial needs. These forward-looking statements are subject to a number of risks, uncertainties and assumptions, including, among other things: negative impacts of the COVID-19 pandemic on Edgewise’s operations, including clinical trials; risks associated with the process of discovering, developing and commercializing drugs that are safe and effective for use as human therapeutics and operating as an early clinical stage company; Edgewise’s ability to develop, initiate or complete preclinical studies and clinical trials for, obtain approvals for and commercialize any of its product candidates; changes in Edgewise’s plans to develop and commercialize EDG-5506 or any other product candidates; the potential for clinical trials of EDG-5506 or any other product candidates to differ from preclinical,

interim, preliminary, topline or expected results; Edgewise’s ability to enroll patients in its ongoing and future clinical trials; operating results and business generally; Edgewise’s ability to raise funding it will need to continue to pursue its business and product development plans; regulatory developments in the United States and foreign countries; Edgewise’s reliance on third parties, contract manufacturers and contract research organizations; Edgewise’s ability to obtain and maintain intellectual property protection for its product candidates; risks associated with access to capital and credit markets; the loss of key scientific or management personnel; competition in the industry in which Edgewise operates; Edgewise’s ability to develop a proprietary drug discovery platform to build a pipeline of product candidates; general economic and market conditions; and other risks. Information regarding the foregoing and additional risks may be found in the section entitled “Risk Factors” in documents that Edgewise files from time to time with the Securities and Exchange Commission. These risks are not exhaustive. New risk factors emerge from time to time, and it is not possible for our management to predict all risk factors, nor can we assess the impact of all factors on our business or the extent to which any factor, or combination of factors, may cause actual results to differ materially from those contained in, or implied by, any forward-looking statements. You should not rely upon forward-looking statements as predictions of future events. Although we believe that the expectations reflected in the forward-looking statements are reasonable, we cannot guarantee future results, levels of activity, performance or achievements. Except as required by law, we undertake no obligation to update publicly any forward-looking statements for any reason after the date of this presentation.

This presentation concerns product candidates that are under clinical investigation, and which have not yet been approved for marketing by the U.S. Food and Drug Administration (FDA). It is currently limited by federal law to investigational use, and no representation is made as to its safety or effectiveness for the purposes for which it is being investigated.

EDG-5506 is an investigational agent and is not approved in any territory

The Edgewise Approach: *Protect susceptible muscle fibers*



Some muscle fibers are more susceptible to damage due to the lack of functional dystrophin

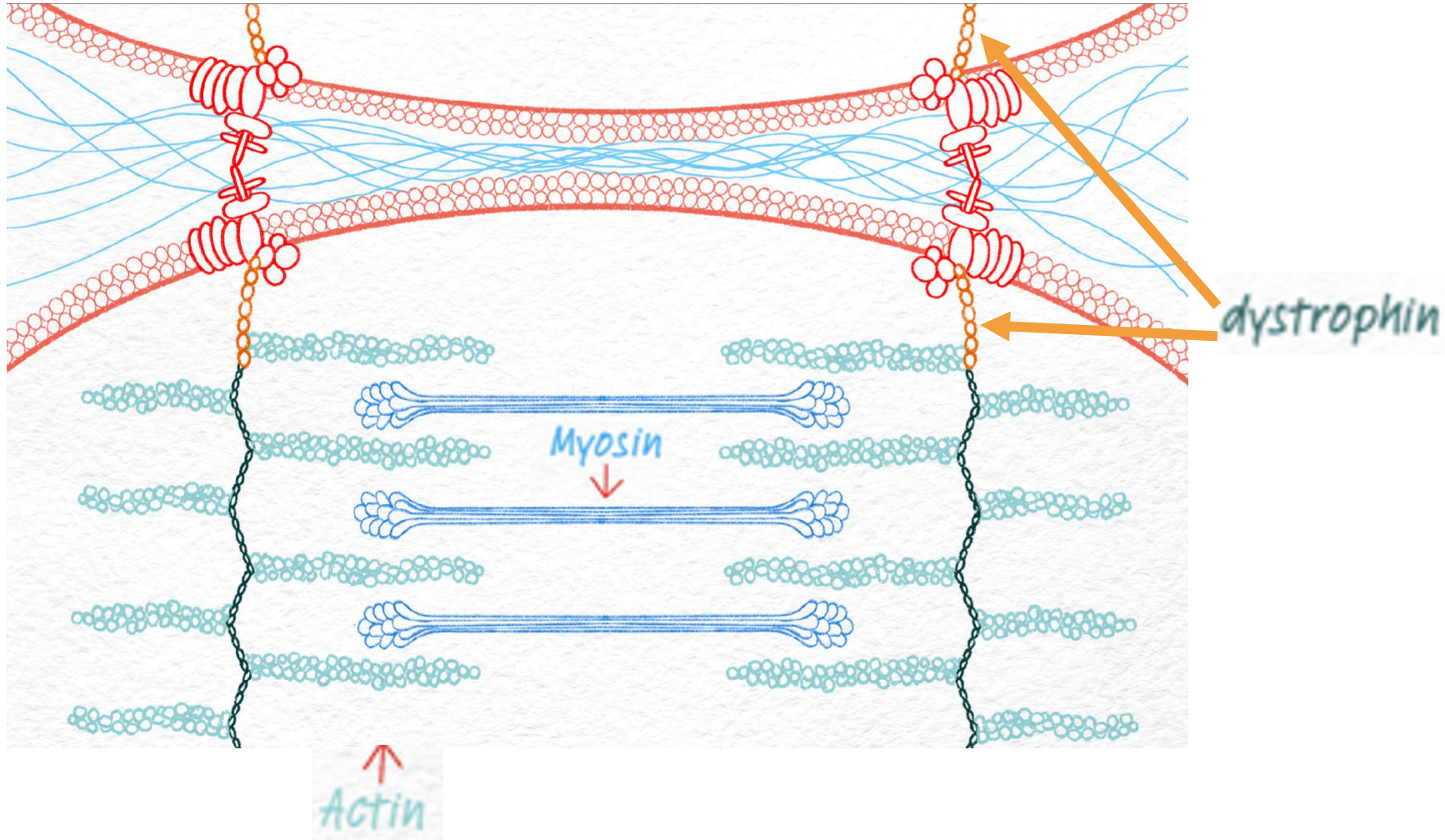


We have designed an investigational therapy, **EDG-5506**, to protect these susceptible muscle fibers from damage



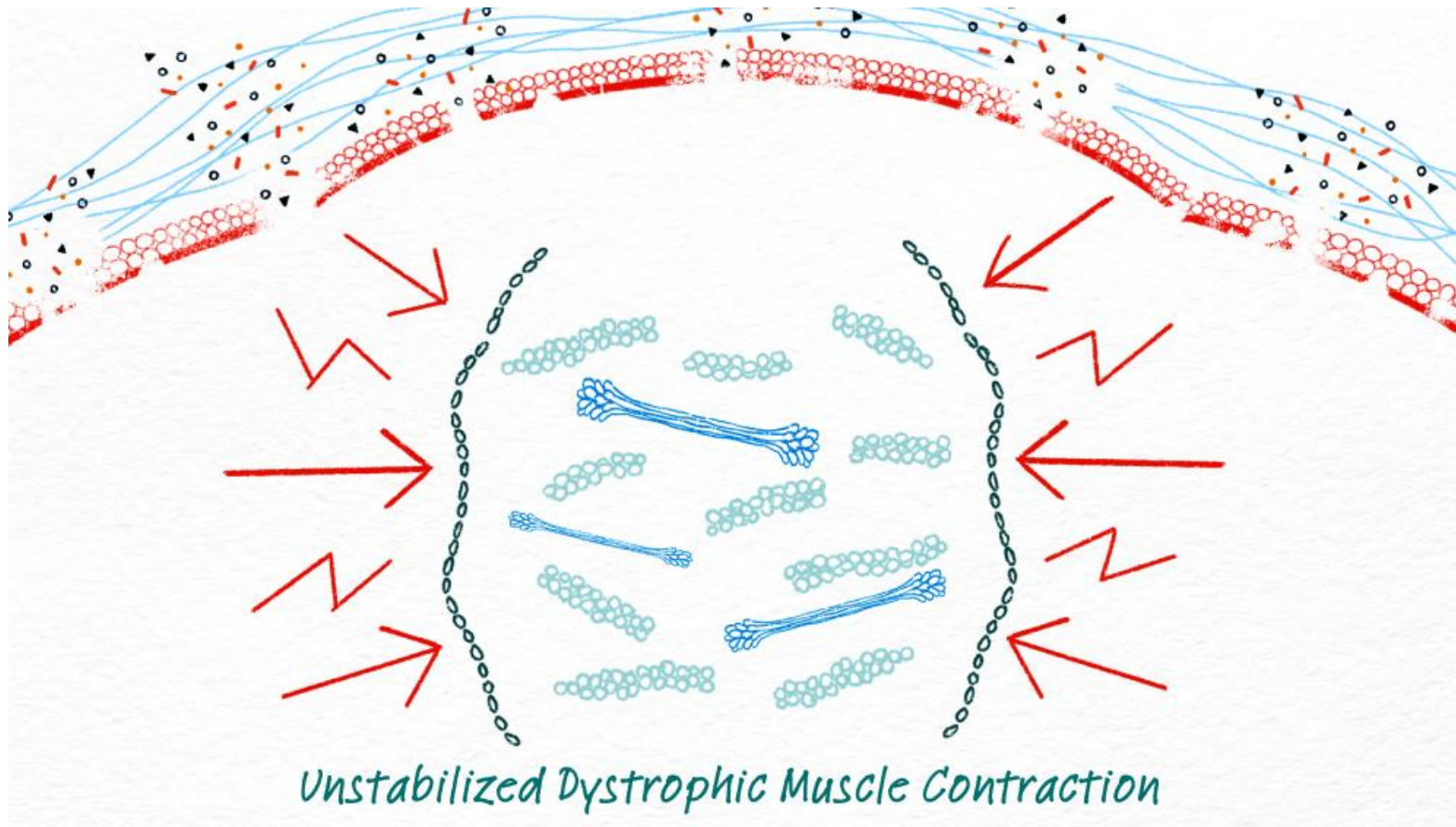
In diseased animal models, **EDG-5506** protected susceptible muscle fibers and prevented long-term development of damage

Dystrophin Protects Muscles from Stress During Contraction Rather than Powering Contraction



The role of dystrophin is to connect parallel muscle fibers to distribute force across muscle during contraction.

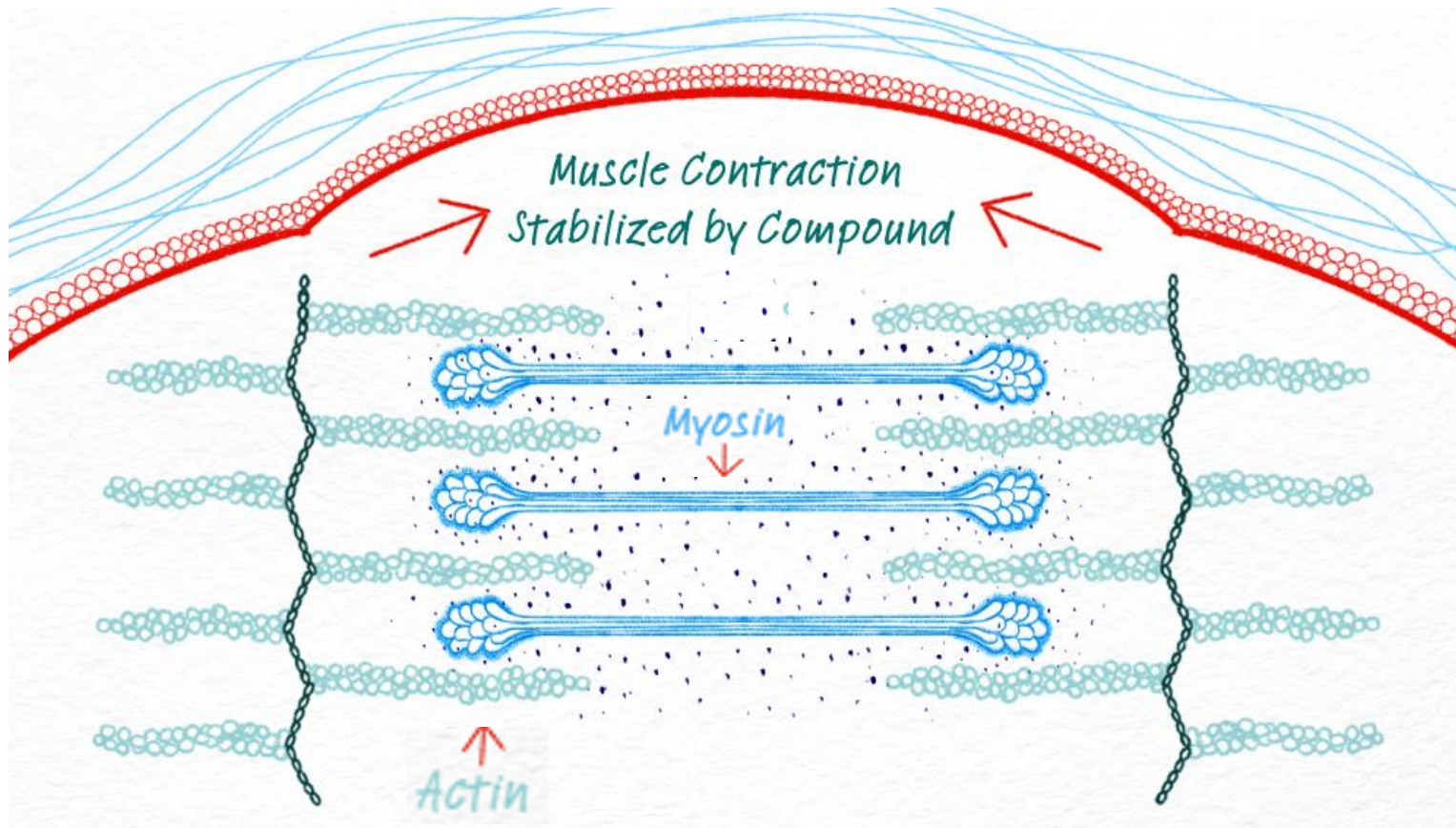
Mechanical Stress Damages Dystrophic Muscle



When dystrophin is lacking, the mechanical stress that occurs during muscle contraction damages the muscle, leading to replacement with fat and fibrotic tissue and ultimately, loss of function.

Unchecked damage leads to fibrosis and functional decline

Myosin Modulation: An Approach to Prevent Mechanical Stress when Dystrophin is Absent



EDG-5506 was designed to protect muscle by modulating myosin, a contractile protein in the fast muscle fibers that are damaged early and to a greater extent in Duchenne and Becker.

Decreased stress allows muscle to function without damage occurring

EDG-5506 Protects Dystrophic Mouse Muscle

Dystrophic mouse muscle is damaged during contraction



Contraction leads to visible changes...

EDG-5506 protects dystrophic mouse muscle during contraction



With EDG-5506, contractions don't cause these changes

EDG-5506 Phase 1 Trial in Unaffected Adults and Those with Becker Muscular Dystrophy



What do we know about side effects?

- When dosed for two weeks, EDG-5506 was well-tolerated
-



How is EDG-5506 given?

- EDG-5506 can be taken as an oral tablet once a day that is absorbed well with or without food
-

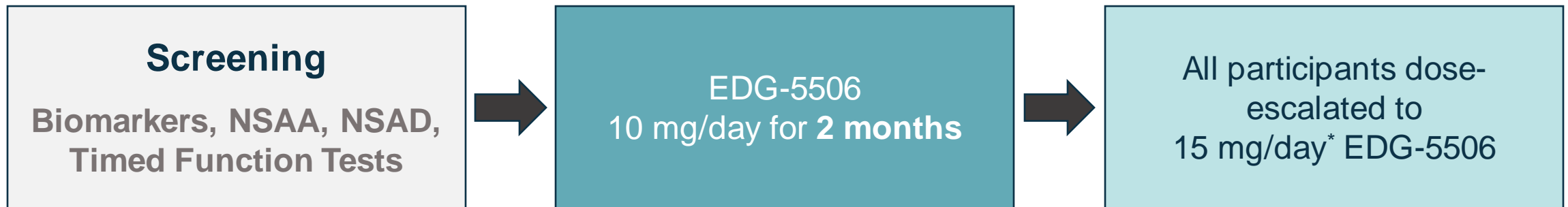
Does EDG-5506 get to the muscle?



- EDG-5506 is highly concentrated in muscle compared to the bloodstream, which tells us that it is getting to where it needs to be to protect the muscle

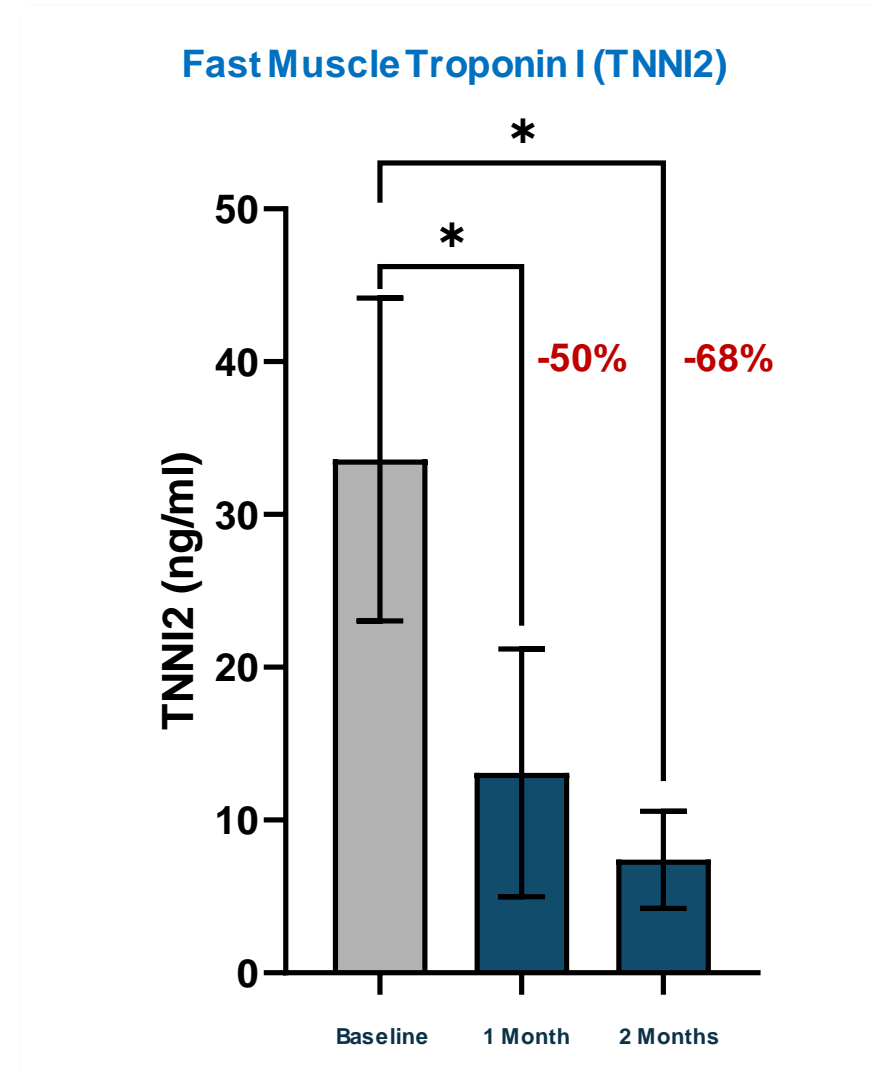
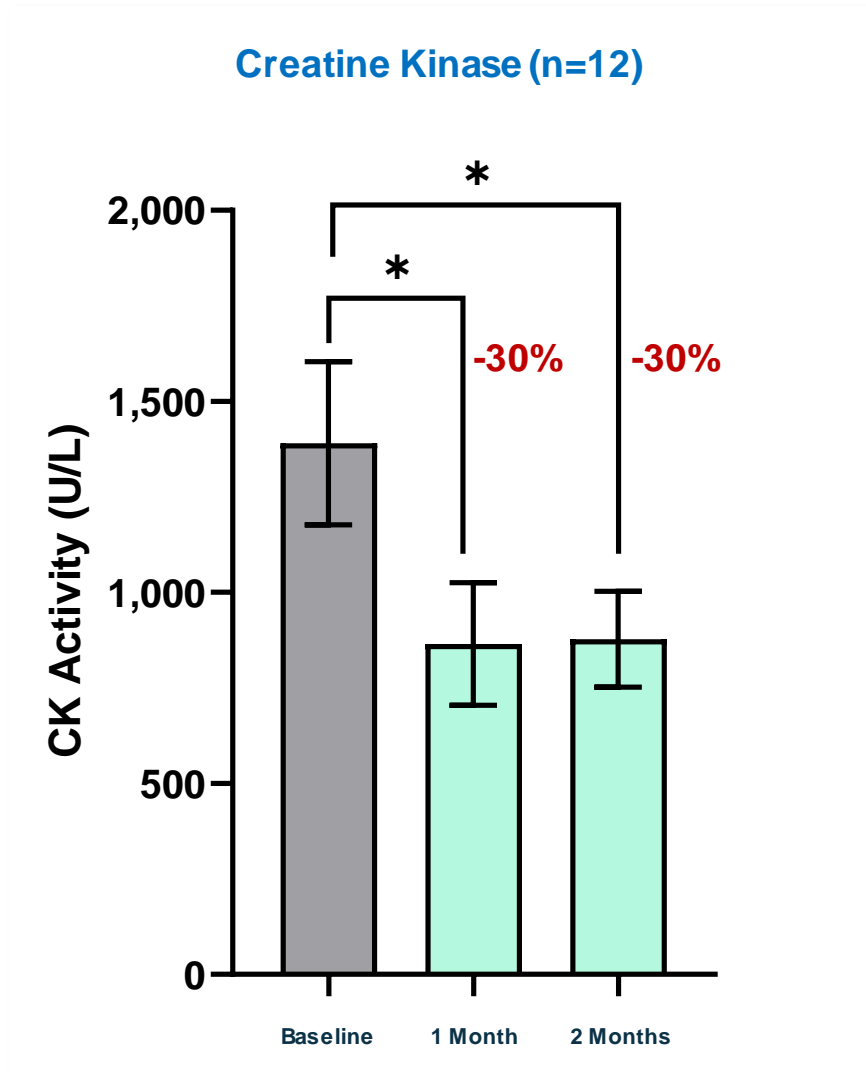
One Year Open Label Study Design

- An open-label, single-center study of EDG-5506 to assess the safety and pharmacokinetics (PK) of EDG-5506 in adults with Becker muscular dystrophy (BMD)
 - 12 patients enrolled, ambulatory with BMD, including all from Phase 1 study



** Protocol allows subsequent dose adjustment after 4 months at 15 mg dose*

2 Months of EDG-5506 Dosing Led to Rapid, Significant and Sustained Decrease in CK and Fast Muscle Troponin in BMD Patients



Source: Data on file; % difference from mean baseline shown

EDG-5506 is Being Developed for Becker and Duchenne Muscular Dystrophy



EDG-5506 is intended to modify disease progression in individuals with Duchenne or Becker with any mutation

EDG-5506 is designed to reduce muscle stress in a type of skeletal muscle fiber that is prone to damage in certain neuromuscular diseases.

Represents a novel mechanism of action

Has the potential to be used alone or in combination with other therapeutic approaches

EDG-5506 Clinical Trials Ongoing and in Planning

- A 12-month open-label study is ongoing, including adults with Becker who were in the Phase 1 trial (NCT05160415)

- A natural history study of those with Becker has begun (ages 8 to 65) (NCT05257473)

- A study in ambulatory adolescent and adult males with Becker is enrolling (ages 12+) (NCT05291091)

- An initial study in ambulatory boys with Duchenne is planned to start in the second half of 2022



THANK YOU!

For questions related to Edgewise clinical trials,
email us at studies@edgewisetx.com