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Pioneering the Possible in Gene Editing

January 2025

Forward Looking Statements

This presentation contains forward-looking statements and information within the meaning of The Private Securities Litigation Reform Act of 1995. The words "anticipate," "believe," "continue," "could," "estimate," "expect," "intend," "may," "plan," "potential," "predict," "target," "should," "would," and similar expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words. Forward-looking statements in this presentation include statements regarding the initiation, timing, progress and results of the Company's preclinical studies and its research and development programs, including the Company's expectation to declare two development candidates for its in vivo programs by mid-2025, establish an additional in vivo target cell type/tissue beyond HSCs and the liver by the end of 2025 and achieve in vivo proof of concept by 2027; the timing for the Company's receipt and presentation of data from its preclinical studies, including presenting further in vivo HSC and liver data in 2025; the potential of, and expectations for, the Company's product candidates; the timing or likelihood of regulatory filings and approvals, including the timing of the Company's submission of any IND or CTA and ability to commence clinical trials for its in vivo programs; and the Company's expectations regarding cash runway into the second quarter of 2027. The Company may not actually achieve the plans, intentions, or expectations disclosed in these forward-looking statements, and you should not place undue reliance on these forwardlooking statements. Actual results or events could differ materially from the plans, intentions and expectations disclosed in these forward-looking statements as a result of various important factors, including: uncertainties inherent in the initiation and completion of preclinical studies; availability and timing of results from preclinical studies; expectations for regulatory approvals to conduct trials; and the availability of funding sufficient for the Company's foreseeable and unforeseeable operating expenses and capital expenditure requirements. These and other risks are described in greater detail under the caption "Risk Factors" included in the Company's most recent Annual Report on Form 10-K, which is on file with the Securities and Exchange Commission, as updated by the Company's subsequent filings with the Securities and Exchange Commission, and in other filings that the Company may make with the Securities and Exchange Commission in the future. Any forwardlooking statements contained in this presentation speak only as of the date hereof, and the Company expressly disclaims any obligation to update any forward-looking statements, whether because of new information, future events or otherwise.



In vivo gene editing – a simple IV infusion to cure a disease – will transform human therapeutics globally to the same degree that mobile smart phones have transformed the communications business around the world.





There is a Solution to the Recent Challenges of Gene Therapy and Gene Editing

Problem

- Recent slow launches
- Low total addressable markets (TAMs)
- Reimbursement challenges
- High cost of goods
- Low margins
- Highly complex patient journeys
- Curative business model
- Many recent gene therapy launches pursued competitive indications with already high standards of care

Solution

- Differentiated therapeutic approach to diseases
- Diseases with higher TAMs
- Simple, scalable, lower cost of goods
- High margin *in vivo* delivery methods
- Ability to treat more patients with less burdensome treatment regimen
- Sustainable revenue growth for curative medicines through rapid development of therapies



editas

Has the solution

A different approach focused on functional upregulation treatment strategy (proven by reni-cel).

Solved for delivery using proprietary, targeted LNPs (tLNPs) that allow targeting of multiple tissues, including HSCs, liver, and other tissues using "plug 'n play" process.

Defined path to rapid development of new medicines with "plug 'n play" *in vivo* editing using reprogrammable guide RNA by changing 20 nucleotides to create a new product for a new disease target.

A leading gene editing platform supported by foundational IP estate.

Driven management team with a proven track record of drug development and commercialization, strong domain expertise, and focus on execution.

Strong cash position with operational runway into Q2 2027.



Editas' Differentiated *In Vivo* Gene Editing Upregulation Strategy Designed to Deliver First-to-Market and Best-in-Class Curative Medicines for Genetic Diseases

	editas	Other Approaches		
Therapeutic strategy	Functional upregulation*	Knockdown	Gene correction	
Gene Editing approach	5' Region Or 3' Region	Exon	Exon	
Non gene Editing modality		siRNA, antisense oligos, monoclonal antibody, and small molecule (pill)		
Patient population	All patients (mutation agnostic)	All patients (mutation agnostic)	Subset of patients (single mutation)	
Therapeutic potential	First/best-in class opportunities for loss of function diseases; cannot be addressed via knockdown	Diseases that can be addressed by protein reduction similar to ASO and siRNA	Correction limited to subset of all patients with given disease	



Reni-cel Provides Proof of Concept for Functional Upregulation Strategy and Validates Editing the *HBG* 1/2 Promoter



- TATA
- Reni-cel treatment showed promising results, with robust and clinically meaningful improvements, for gene editing at the HBG1/2 promoters with AsCas12a

- Patients achieved early correction of anemia, durable normalization of total Hb, and sustained increase in HbF ≥40% with pancellular distribution
- Markers of hemolysis improved or normalized by Month 6
- 27 of 28 treated patients were VOE-free post-reni-cel infusion as of the data cutoff date
- Early and sustained meaningful improvements were observed in pain, physical, and social patient-reported outcome domains



 The safety profile was consistent with myeloablative busulfan conditioning and autologous HSCT



Hb, hemoglobin; HbF, fetal hemoglobin; *HBG1/2*, γ-globin genes 1 and 2; HSCT, hematopoietic stem cell treatment; VOE, vaso-occlusive event Data cutoff date of Oct 29, 2024. Presented at the American Society of Hematology (ASH) Annual Meeting and Exposition, December 9, 2024.

Preclinical POC Data in *In Vivo* HSC Editing Leverages Editas' Gene Editing Expertise and Provides Foundation for LNP Platform

Potential for Best-in Class, First-in-Class In Vivo Medicine for Sickle Cell Disease and Beta Thalassemia



Leveraging reni-cel experience with validated target and enzyme that provides for development of a differentiated medicine for sickle cell disease and beta thalassemia



Demonstrated *in vivo* capabilities with **devised novel HSC targeting strategy** and **proprietary LNP** to deliver editing cargo



Produced competitive preclinical data set that outperforms data currently in the public domain



In vivo gene editing medicine for sickle cell disease and beta thalassemia **can expand the total addressable market (TAM)**

Proprietary LNP Platform

- Foundation for an LNP Platform for Delivery to Extrahepatic Tissues
- Ability to deliver gene editing cargo with HSC targeting moiety conjugated to our propriety LNP Platform
- May provide delivery cargo to other tissues and cell types of interest



High Levels of *In Vivo* HBG1/2 Gene Editing Achieved in HSPCs with Optimization of a Proprietary LNP and Novel Targeting Strategy in Mice (



Preclinical PoC achieved for potential treatment of SCD and Beta thalassemia by a clinically validated strategy after a single dose of Editas' proprietary tLNP

In vivo model: NBSGW mouse strain (NOD.Cg-Prkdc^{scid} II2rg^{tm1WjI}/SzJ (NSG) crossed with C57BL/6J-Kit^{W-41J}/J (C57BL/6.Kit^{W41})) engrafted, without irradiation, with human CD34+ cells from peripheral blood after plerixafor mobilization of cells from bone marrow.



HSPC: Hematopoietic stem and progenitor cells; HSPC defined as Lin⁻CD34⁺CD38⁻ cells; HSC defined as Lin⁻CD34⁺CD38⁻CD90⁺CD45RA⁻ cells; % indel indicates level of allelic editing, % HbF+ (Fetal hemoglobin positive) cells by flow cytometry

High Efficiency HSC Delivery Approaching Therapeutic Editing Levels Achieved After a Single Dose of LNP2 in Non-human Primates

GFP mRNA delivery and HBG1/2 editing of HSCs after single dose of LNP2 (moiety 2)



Ongoing evaluation of further optimized formulations, e.g., LNP3 (moiety 2) expected to achieve higher therapeutic editing levels



HSC: Hematopoietic stem cells defined as CD34⁺CD90⁺CD45RA⁻ cells based on Radtke, S. Kiem, HP, et al. (2017). Sci Transl Med 9(414):eaan1145 GFP: Green fluorescent protein

Preclinical NHP PoC Validates High Efficiency Genomic Editing in Liver with First Use of AsCas12a Nuclease Delivery by Lipid Nanoparticle





AsCas12a nuclease and target-specific gRNA delivered using Genevant proprietary LNP

In Vivo Preclinical Proof of Upregulation Strategy Confirmed by Clinically Relevant Target Protein Increase Resulting in Significant Disease Biomarker Reduction in Undisclosed Target 1



In vivo achievement of clinically meaningful level (≥ 2-fold upregulation) of Target 1 protein expression in mice



In vivo PoC for Plug 'n Play Delivery to Extrahepatic Cell Types Achieved with Editas' Proprietary LNP Targeting Platform



In vivo targeting to three extrahepatic cell types at ≥80% efficiency with our plug 'n play platform



A Capital Efficient In Vivo Gene Editing Company

Continued Execution of our Focus to Leverage our Foundational IP Estate for Access to Non-Dilutive Capital

October 03, 2024

Editas Medicine Announces \$50+ Million Monetization Financing with DRI Healthcare Trust

Strengthens balance sheet with non-dilutive capital to enable further pipeline development and related strategic priorities





Editas' Partnerships Validate Science and Value of IP Estate

Editas is well positioned to capture value of IP and leverage non-dilutive financing to focus resources on in vivo pipeline development



Future IP Licensing Opportunities U Multiple clinical-stage programs in development will require an IP License for use of CRISPR Cas9 and Cas12a Technology



2025 Key Anticipated Milestones



Declare Two *in vivo* Development Candidates by Mid-2025

- Candidate in hematopoietic stem cells (HSCs) for the treatment of beta thalassemia and sickle cell disease
- Candidate in liver cells for an undisclosed indication

Establish One Additional Target Tissue

• **Disclose target cell type or tissue by end of 2025**, beyond HSCs and liver cells

Present in vivo preclinical editing data

• In vivo preclinical proof-of-concept in both HSCs and liver cells in large animal models

Derive Revenue from Foundational IP

• Building on the DRI Healthcare monetization, continue to issue sublicenses



2025-2027 Strategic Priorities



Launch Clinical Trials for Multiple in vivo Programs in Multiple Tissues

- Submit at least one IND/CTAs by mid-2026
- Begin at least one human clinical trial by 2H
- Potential for at least one late-stage clinical trial in 2H 2027

Achieve in vivo Human Proof of Concept by year-end 2026

 Achieve human proof-of-concept for at least one indication by year-end 2026

Expand the Range of Diseases Addressable by *in vivo* Gene Upregulation

In vivo proof of concept in at least one tissue beyond HSCs and liver by 2027, demonstrating plug 'n play potential of Editas' proprietary extrahepatic LNP platform





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Additional Information

Programs Positioned for Development

PROGRAM (OR DISEASE CANDIDATE)	PRECLINICAL	IND ENABLING	EARLY-STAGE CLINICAL	LATE-STAGE CLINICAL	DEVELOPMENT & COMMERCIAL PARTNER
HEMOGLOBIN-OPATHIES					
In Vivo HSC Editing – sickle cell disease					
<i>In vivo</i> HSC Editing – beta thalassemia					
OTHER ORGANS & TISSUES					
Liver Upregulation Target 1					
Other Tissue Upregulation Target					
ONCOLOGY					
$\alpha\beta$ T Cells (14 total programs)					ر ^{ال} Bristol Myers Squibb
γδ T Cells					Immatics
iNK Cells					SHORELINE

