editas

Dima, Tristan, & Stephanie

Corporate Presentation

April 2024

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 and clinical studies and its research and development programs, including initiating the adolescent cohort in the RUBY trial in 2024 and establishing in vivo proof-of-concept for an undisclosed indication in 2024, the timing for the Company's receipt and presentation of data from its clinical trials and preclinical studies, including RUBY clinical updates in mid-2024 and by year-end 2024, the potential of, and expectations for, the Company's product candidates, the timing or likelihood of regulatory filings and approvals, the Company's expectations regarding commercial readiness, and the Company's expectations regarding cash runway. 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 pre-clinical studies and clinical trials, including the RUBY and EdiTHAL trials, and clinical development of the Company's product candidates, including reni-cel; availability and timing of results from pre-clinical studies and clinical trials; whether interim results from a clinical trial will be predictive of the final results of the trial or the results of future trials; expectations for regulatory approvals to conduct trials or to market products and 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 forward-looking statements contained in this presentation represent Company's views only as of the date hereof and should not be relied upon as representing its views as of any subsequent date. Except as required by law, the Company explicitly disclaims any obligation to update any forward-looking statements.



Editas Medicine is a Leader in the CRISPR-based Gene Editing Medicine Field



- Our lead product candidate, reni-cel, is an investigational gene editing medicine that is a potential "best in class" treatment for sickle cell disease and beta thalassemia
- Ongoing RUBY and EdiTHAL clinical trials



- Editas holds an exclusive license to foundational IP for Cas9 and Cas12a for the prevention or treatment of human disease from the Broad Institute and Harvard University
- Source of non-dilutive capital
 - Granted Cas9 sublicenses, including non-exclusive licenses to Vertex Pharmaceuticals and Vor Bio

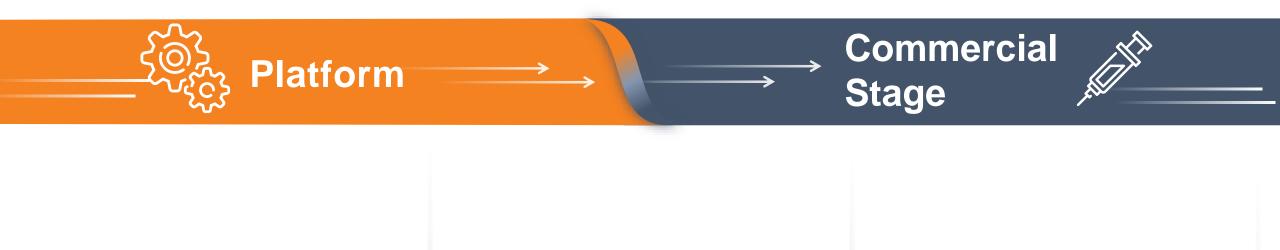


- Proprietary AsCas12a is a high fidelity and high efficiency CRISPR nuclease
- Core expertise in guide RNA design and chemistry for high precision editing
- Longer-term focus on creating important medicines based on *in vivo* gene editing
- Scaled Chemistry, Manufacturing, and Controls (CMC)

 Leadership team with a proven track record of drug development and commercialization



Strategic Framework (From the 2023 J.P. Morgan Healthcare Conference)



Drive reni-cel (EDIT-301) toward BLA and Commercialization Strengthen and Focus Discovery to Build *in vivo* Editing Pipeline Increase Business Development Activities and Monetize IP

Long-Term Vision: A Leader in In Vivo Programmable Gene Editing



Strategic Transformation Toward Long-Term Vision

(From the 2023 J.P. Morgan Healthcare Conference)

Platform

Drive reni-cel (EDIT-301) toward BLA and Commercialization

- Continue ex vivo development of reni-cel (EDIT- 301) for SCD, TDT
 - Enroll 20 patients in RUBY study by year-end
 - Provide RUBY and EdiTHAL data updates by mid-year and year-end
- Divest wholly-owned cell therapy program, continue supporting partnered cell therapy programs
- 7 Terminate AAV IRD programs

Strengthen and Focus Discovery to Build *in vivo* Editing Pipeline

- 🤣 Focus on *in vivo* pipeline build
- Hire new CSO with specific expertise aligned with Editas' vision
- Reset discovery and technology group
- Initiate discovery of *in vivo* editing of HSCs and in other tissues

Increase Business Development Activities and Monetize IP

Commercial

Stage

- Create value through business development to complement core gene editing technology capabilities
- Leverage robust IP portfolio



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Long-Term Vision: A Leader in In Vivo Programmable Gene Editing
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2024 Strategic Objectives

Drive reni-cel (EDIT-301) toward BLA and Commercialization

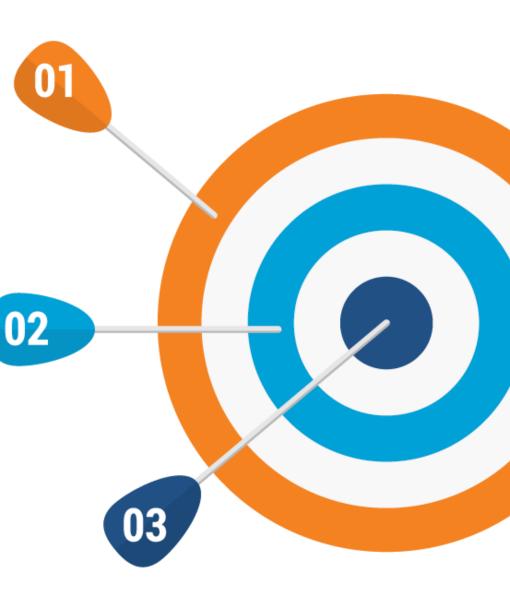
- Continue enrollment and dosing in the RUBY and EdiTHAL trials of reni-cel
- Initiate the adolescent cohort in the RUBY trial
- Present a substantive clinical data set of Sickle cell patients with considerable clinical follow-up in the RUBY study in mid-2024 and by year-end 2024

Strengthen and Focus Discovery to Build *in vivo* Editing Pipeline

- Establish *in vivo* preclinical proof-of-concept for an undisclosed indication
 - Focus on disease targets with high probability of technical, clinical, regulatory, and commercial success
 - Initial focus on hematopoietic stem cells (HSCs)

Increase Business Development Activities and Monetize IP

• Derive revenue from the Company's foundational IP, building on the recently announced license agreements with Vertex Pharmaceuticals and Vor Bio



Sickle Cell Disease (SCD) is an Inherited Life-Threatening Hematological Disorder Manifesting Shortly After Birth

SICKLE CELL DISEASE is a genetic blood disorder caused by mutations in the **HBB gene** that causes sickling of RBCs; this leads to **anemia**, **hemolysis**, **and VOEs**^{1,2}



UPREGULATION OF FETAL HEMOGLOBIN (HbF) is a naturally validated therapeutic strategy to control complications of SCD

SCD AFFECTS^{3,4,5}

PEOPLE IN THE U.S.

~100K

EDITAS EDITS THE HBG1 AND HBG2 PROMOTERS USING AsCAS12a ENZYME, AN APPROACH THAT IS DESIGNED TO:

- Upregulate HbF robustly
- Correct anemia with superior red blood cell production and health vs. BCL11A approach
- Reduce risk of off-target editing with high fidelity and high efficiency proprietary AsCas12a enzyme

Reni-cel is potentially a "best in class" medicine with consistent correction of anemia

HBB, β-globin gene; RBC, red blood cell; SCD, sickle cell disease; VOE, vaso-occlusive event.
1. Kato GJ *et al. Nat Rev Dis Primers* 2018; 4: 18010. 2. Williams TN *et al. Annu Rev Genomics Hum Genet* 2018; 19: 113–147. 3. Sickle Cell Disorders. Available at: https://www.thelancet.com/pb-assets/Lancet/gbd/summaries/diseases/sickle-cell-disorders.pdf. Accessed June 2023. *4.* Wastnedge E *et al. J Glob Health* 2018; 8 (2): 021103. 5. Sickle Cell Disease. Available at: https://www.thelancet.com/pb-assets/Lancet/gbd/summaries/diseases/sickle-cell-disease. Available at: https://www.thelancet.com/pb-assets/Lancet/gbd/summaries/disease. Available at: https://www.thelancet.com/pb-assets/Lancet/gbd/summaries/diseases/sickle-cell-disease. Available at: https://www.thelancet.com/pb-assets/Lancet/gbd/summaries/disease. Available at: https://www.thelancet.com/pb-assets/Lancet/gbd/summaries

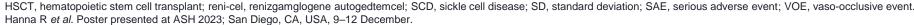
All Treated RUBY Patients Successfully Engrafted, Showed a Favorable *Ruby* Safety Profile

DEMOGRAPHICS	(N=11)	
Genotype, n(%)		
β ^s /β ^s	11 (100)	
Sex, n (%)		
Female	6 (54.5)	
Age, years, mean (SD)	27.6 (4.2)	
Severe VOEs, pre-study annual rate*, mean (SD)	3.9 (1.4)	
	(N=11 ⁺)	
Total reni-cel dose administered, ×10 ⁶ CD34 ⁺ cells/kg, mean (SD)	5.2 (2.5)	
Follow-up duration, months, mean (SD)	6.5 (5.3)	
Time to neutrophil engraftment ^{†, ‡} , days, mean (SD)	23.7 (2.8)	
Time to platelet engraftment ^{†,§} , days, mean (SD)	26.1 (7.7)	

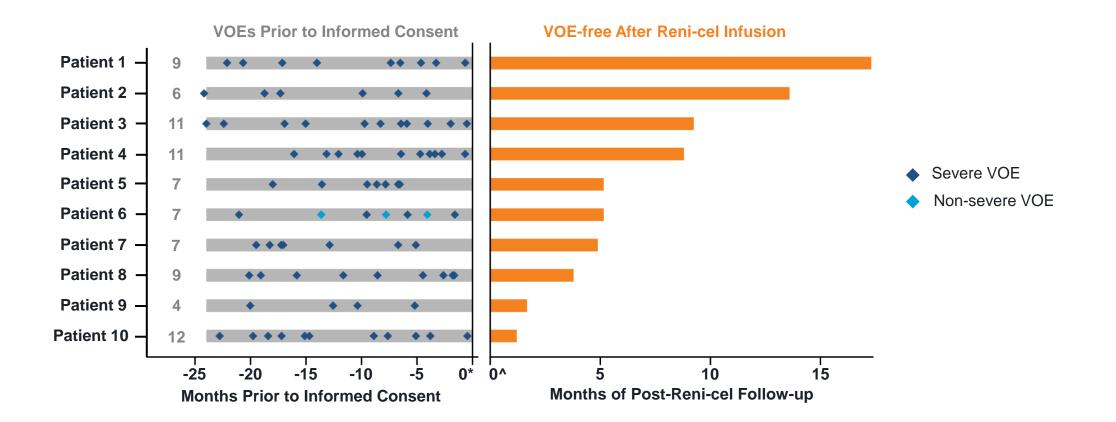
- Safety profile is consistent with myeloablative busulfan conditioning and autologous HSCT
- No serious adverse events (SAEs) related to reni-cel were reported after reni-cel infusion

Data cutoff November 22, 2023.

*The pre-study period is defined as the 2-year period prior to informed consent. [†]One patient had 23 days of follow-up after infusion as of the data cut; neutrophil engraftment and platelet engraftment were not achieved yet; engraftment values are therefore based on n=10. [‡]Three consecutive measurements with absolute neutrophil count (ANC) ≥0.5 × 10⁹/L. [§]Three consecutive measurements with platelet count ≥50 × 10⁹/L starting at least 7 days after the platelet transfusion, and 10 days after thrombopoietin (TPO). No TPO was used for patients after reni-cel infusion.



All Treated RUBY Patients are VOE-free Since Reni-cel Infusion

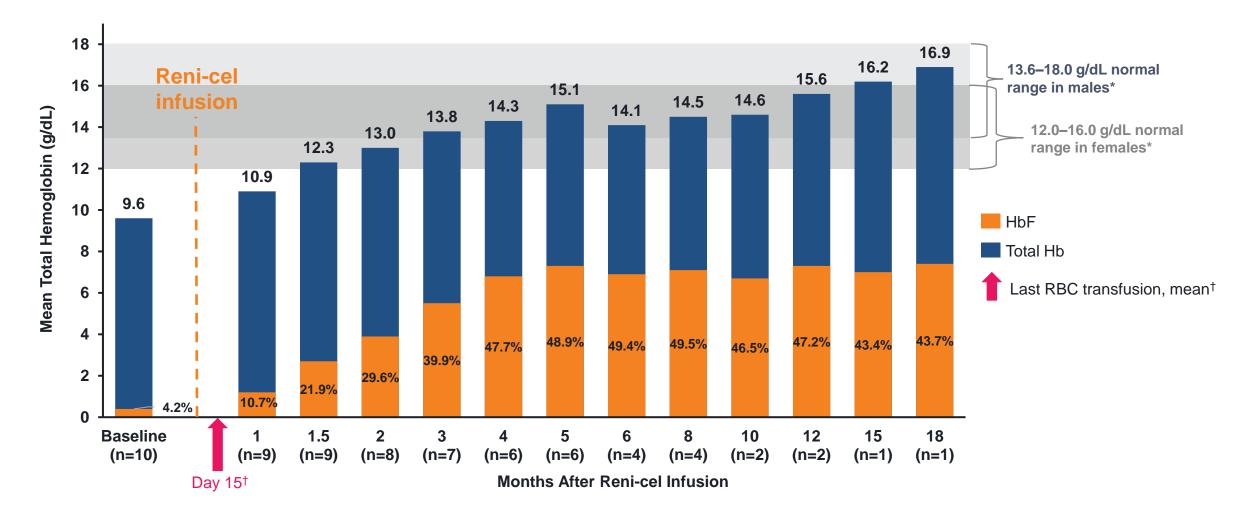


All 10 patients who reached the Month 1 visit have been VOE-free since reni-cel infusion



Data cutoff November 22, 2023. Due to limited follow up period after dosing, Patient 11 was not included. Left panel ends at informed consent date: *Day of informed consent. Right panel starts at infusion date: ^Day reni-cel was infused. reni-cel, renizgamglogene autogedtemcel; VOE, vaso-occlusive event. Hanna R *et al.* Poster presented at ASH 2023; San Diego, CA, USA, 9–12 December.

RUBY Patients Show Total Hb Rapidly Returning to the Normal Range and Clinically Meaningful Improvements in HbF Levels of >40%



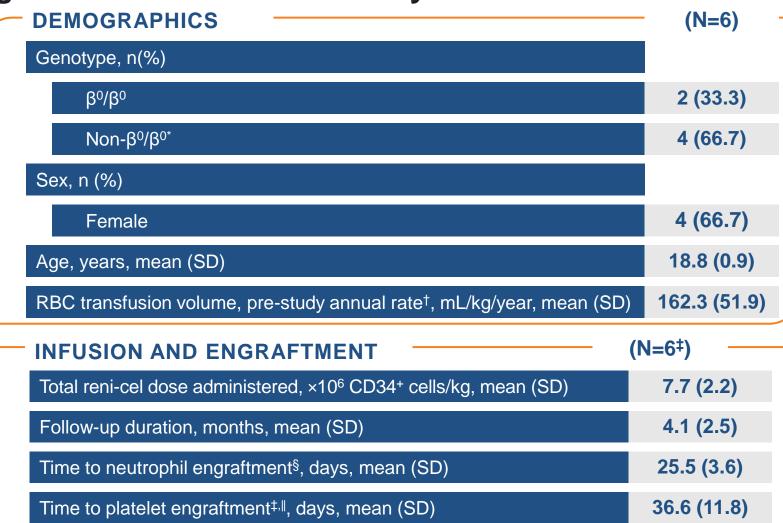
Data cutoff November 22, 2023. Number of male patients = 5; number of female patients = 5. Bars show mean Hb (g/dL). Labels inside / next to the bars indicate mean proportion of HbF as a percentage of total Hb. Mean total Hb concentrations are shown directly above bars.

*Central laboratory reference range. †The last RBC transfusion in patients occurred a mean (SD) of 15.4 (6.0) days after reni-cel infusion (n=10).

Hb, hemoglobin; HbF, fetal hemoglobin; RBC, red blood cell; reni-cel, renizgamglogene autogedtemcel; SCD, sickle cell disease.

Hanna R et al. Poster presented at ASH 2023; San Diego, CA, USA, 9–12 December.

EdiTHAL Patients Successfully Engrafted, Experienced Similar **Engraftment and Similar Safety Profile to RUBY Patients**



Safety profile is consistent with myeloablative busulfan conditioning and autologous HSCT

No serious adverse events (SAEs) related to reni-cel were reported after reni-cel infusion

Data cutoff November 28, 2023.

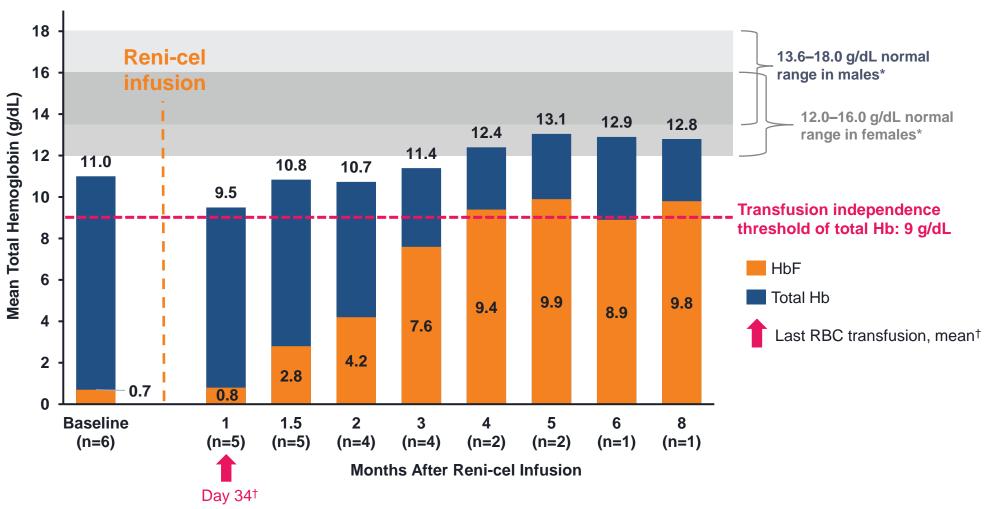


*Non- β^0/β^0 includes β^0/β^+ (n=3) and β^E/β^0 (n=1). [†]The pre-study period is defined as the 2-year period prior to informed consent. [‡]One patient had 36 days of follow-up after infusion as of the data cut; neutrophil was engrafted, but platelet engraftment was not achieved yet; platelet engraftment values are therefore based on n=5. [§]Three consecutive measurements with absolute neutrophil count (ANC) ≥0.5 × 10⁹/L. [¶]Three consecutive measurements with platelet count ≥20 × 10⁹/L starting at least 7 days after the platelet transfusion, and 10 days after thrombopoietin (TPO). No TPO was used for patients after reni-cel infusion. RBC, red blood cell; reni-cel, renizgamglogene autogedtemcel; SD, standard deviation; TDT, transfusion-dependent β-thalassemia; SAE, serious adverse event.

Hanna R et al. Poster presented at ASH 2023; San Diego, CA, USA, 9-12 December.

FDITHAL

EdiTHAL Patients Had Early and Robust Increase of Total Hb Above the Transfusion Independence Threshold



Data cutoff November 28, 2023. Number of male patients = 2; number of female patients = 4. Bars show mean Hb (g/dL). Labels inside / next to the bars indicate mean levels of HbF (g/dL). Mean total Hb concentrations are shown directly above bars.



Hb, hemoglobin; HbF, fetal hemoglobin; reni-cel, renizgamglogene autogedtemcel; TDT, transfusion-dependent β-thalassemia.

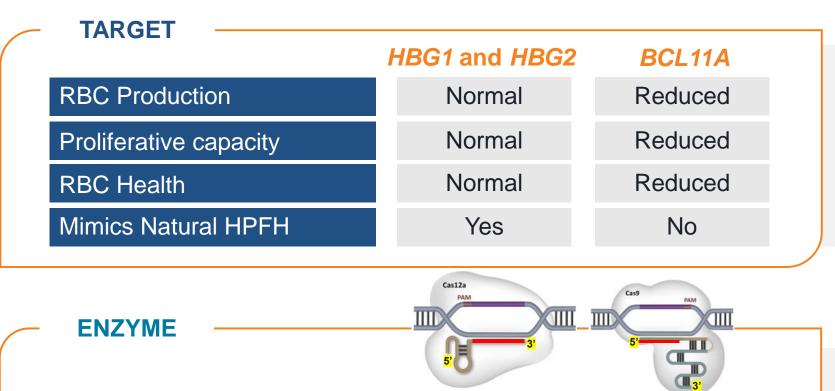
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EDITHAL

Reni-cel's Rational Design: Target Selection AND CRISPR Enzyme Matter in Building a Medicine to Give Best Outcomes to Patients





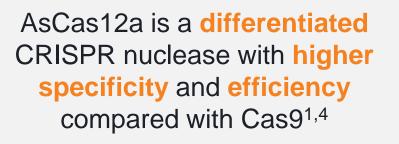


AsCas12a

Higher

Higher

HBG1 and HBG2 promoters are a more appropriate genomic target versus BCL11A for RBC production^{1,2}



Editing Efficiency

Specificity

Images from Moon et al. 2019.3

Cas₉

Lower

Lower

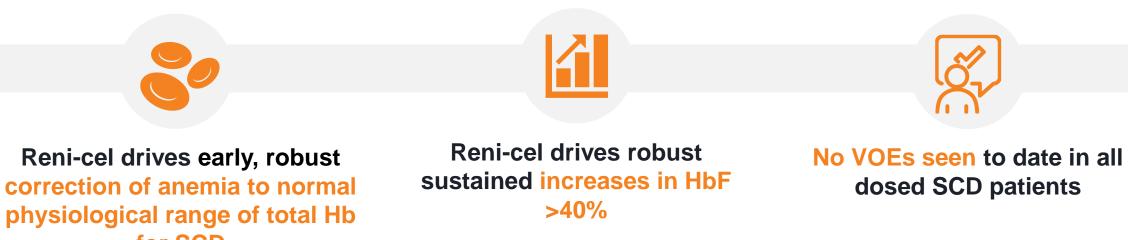
BCL11A, B-cell lymphoma/leukemia 11A gene; Cas9, CRISPR-associated protein 9; AsCas12a, CRISPR-associated protein 12a; CRISPR, clustered regularly interspaced short palindromic repeats; HBG, γglobin gene; HPFH, hereditary persistence of fetal hemoglobin; RBC, red blood cell.



1. Editas Medicine. Data on file. 2. Chang et al. Oral presentation at ASH 2018; San Diego, CA, USA, 2 December 2018. 3. Moon SB et al. Trends in Biotechnology 2019; 37 (8): 870-881. 4. Zhang L et al. Nat Commun. 2021; 12 (1): 3908.

Key Takeaways





for SCD





Reni-cel safety profile consistent with myeloablative busulfan conditioning and autologous HSCT Initial Hb and HbF responses are consistent in SCD and TDT patients at the same follow-up time points



2024 Strategic Objectives

Drive reni-cel (EDIT-301) toward BLA and Commercialization

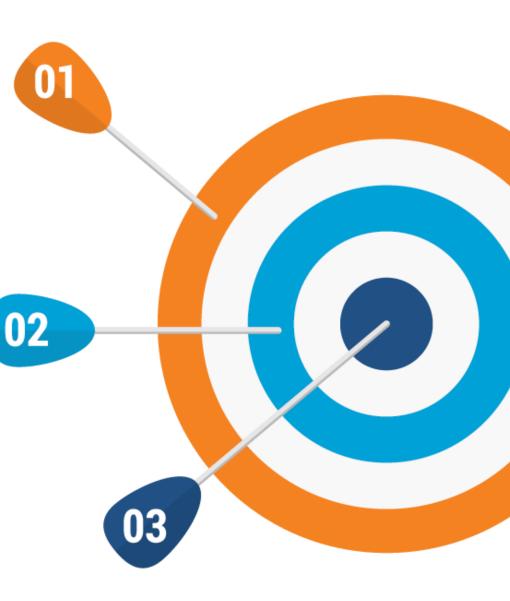
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Investment Highlights

Groundbreaking Science

Strong Leadership Team

Commercial Focus

World leading gene editing platform supported by foundational IP estate.

Shareholder value driven team with a proven track record of drug development & commercialization, strong domain expertise and focus on execution.

Lead asset renizgamglogene autogedtemcel (reni-cel) a potentially differentiated treatment for sickle cell disease and beta thalassemia.

Recent data at the American Society of Hematology (ASH) Annual Meeting supporting differentiation with two additional clinical data updates expected in 2024.

Disciplined, longer-term focus on creating important medicines for people living with serious diseases based on *in vivo* gene editing.

Strong cash position with operational runway into 2026.



Additional Information

Programs Positioned for Clinical Success

	PROGRAM (OR DISEASE CANDIDATE)	PRECLINICAL	IND ENABLING	EARLY-STAGE CLINICAL	LATE-STAGE CLINICAL	DEVELOPMENT & COMMERCIAL PARTNER
HEMOGLOBIN -OPATHIES	renizgamglogene autogedtemcel (reni- cel) : <i>Ex Vivo</i> Autologous Treatment for Sickle Cell Disease (SCD)					
	renizgamglogene autogedtemcel (reni- cel) : <i>Ex Vivo</i> Autologous Treatment for Transfusion-Dependent Beta Thalassemia (TDT)					
	In Vivo HSC Editing					

OTHER ORGANS & TISSUES	Undisclosed Target 1		
	Undisclosed Target 2		

$\alpha\beta$ T Cells (13 total programs)		ر ^{ال} Bristol Myers Squibb [™]
γδ T Cells		Immatics



Experienced Team Focused on Delivering Shareholder Value

