

# **Corporate Presentation**

## A New Class of Cell Therapies & Genetic Medicines with the Capacity to Cure



AUGUST 2024

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## On a mission to advance a new class of cell therapies & genetic medicines



UNMATCHED PLATFORM

Innovating with powerful, proprietary, and differentiated genetic engineering technologies

# Proprietary validated, non-viral platform positions Poseida for leadership in allogeneic cell therapy and genetic medicines

## Unique T<sub>SCM</sub>-based Allo CAR-T Therapies

- Proven results & 3 clinical stage therapies
- Differentiated CAR-T product profiles
- ✓ In house GMP manufacturing
- Further opportunities with preclinical pipeline
- ✓ Well-suited for oncology & beyond

## Differentiated Genetic Medicines

- Stable integration for durability of response
- ✓ Can re-dose, titrate
- Suitable for patients of all ages
- Nonviral approach for lower cost and better safety
- Broad potential across genetic diseases

## Effective Corporate Stewardship

- Strong, select partnerships with Roche and Astellas
  - Validate technology
  - Fund program activities
  - Provide nondilutive financing
- Disciplined and seasoned management team



# Our robust pipeline spans allogeneic CAR-T and non-viral genetic medicines



\*Solid tumor targets identified in conjunction with the research collaboration and license agreement with Astellas are yet to be disclosed





# Allogeneic CAR-T

The Future of Cell Therapy is Allo



# Poseida is emerging as an industry leader in allogeneic CAR-T

POSEIDA'S VISION: Our T<sub>SCM</sub>-rich allogeneic CAR-T will enable all patients who can benefit from transformational cell therapy to do so

"Built in" product differentiation through unique T<sub>SCM</sub>-rich CAR-T approach

Fully proprietary genetic engineering toolkit designed for T<sub>SCM</sub>-rich allo CAR-T

Clinical proof-of-concept delivered through lead P-BCMA-ALLO1 program

Manufacturing platform advancing in lockstep with clinical development

Robust and growing multi-asset pipeline

Allo CAR-T company of choice for top pharma (Roche, Astellas)

Holistic systems engineering approach to allogeneic cell therapy



# Stem cell memory T cells ( $T_{SCM}$ ) are the ideal cell type for CAR-T and have always been our focus and key source of differentiation





The Poseida difference: Our proprietary tools uniquely provide the capabilities required to produce T<sub>SCM</sub>-rich allogeneic CAR-T



# PiggyBac is an effective, non-viral system that inserts one or more genes in a single step to deliver a $T_{SCM}$ -rich cell therapy product

### PIGGYBAC FOR CAR-T DRUG DEVELOPMENT

- Non-viral, transposon-based system that avoids viral vector issues such as cost, quality/safety, and limited cargo capacity
- Preferentially transposes naïve and stem cell memory T cells resulting in T<sub>SCM</sub> rich product
- Works well in resting T cells so T<sub>SCM</sub> phenotype can be preserved
- Large cargo capacity can deliver one or more CARs, inducible safety switch, and selectable marker in single step
- Achieves stable DNA integration and works in a wide variety of cell types





# PiggyBac enables Poseida to build tremendous functionality into its allogeneic CAR-T cell therapies





Poseida's Cas-CLOVER gene editing system addresses graft vs. host and host vs. graft alloreactivity in allogeneic CAR-T while preserving product stemness



#### HIGH-FIDELITY, DESIGNED FOR LITTLE TO NO OFF-TARGET CUTTING

- Utilizes deactivated Cas9 (dCas9) as a binding protein with gRNA
- dCas9 guides a fused nuclease domain from the Clo051 enzyme, which only cuts DNA when bound to its matching pair
- Edits in current clinical-stage CAR-T include TCR and B2M (MHC I) knockouts



- Unlike many other approaches to gene editing, effective in resting T cells
- Fidelity possibly up to 25-fold greater than CRISPR-Cas9
- High editing efficiency in resting T cells results in high levels (50–70%) of T<sub>SCM</sub>
- Preserves stemness because T cell activation, which stimulates differentiation, is avoided
- Multiplexing potential for multiple edits in a single efficient step

 Madison et al., Mol Ther Nucleic Acids. 2022; 2. Alvarez et al., Mol Ther. 31(4), Supp. 1, S1-794. 2023. 3. Data on file, Manuscript in preparation (Poseida Therapeutics) 4. Gilmore et al., NEJM 2021; 5. Longhurst et al., NEJM 2024; 6. Ren et al., Clin Cancer Res., 2017; 7. Antoniani et al., Blood. 2018; 8. Georgiadis et al., Mol Ther. 2018; 9. Webber et al., Nature Comm., 2019; 10. Fix et al., J Immunother Cancer. 2022; 11. Ottaviano et al. Sci. Trans. Med., 2022; 12. Zhang et al., Nature., 2022; 13. Cancellieri et al., Nature Genetics 2023; 14. Poseida R&D Day, April 17, 2024; 15. Alvarez et al., Mol Ther. 2023.



## P-BCMA-ALLO1 represents a new category of allogeneic CAR-T with a compelling value proposition for patients with multiple myeloma



# P-BCMA-ALLO1

- Lead program in heme malignancies partnered with Roche
- Targeting B cell maturation antigen (BCMA)
- Early data (ASH 2023) is pace-setting for allo CAR-T in MM

## Ongoing clinical development with additional data updates in 2H2024

Target product value proposition		P-BCMA-ALLO1 early product profile	
☆	<ul> <li>Effective myeloma control</li> <li>High response rates</li> <li>Deep responses including in hard to treat patients</li> </ul>	<ul> <li>✓ 82% ORR<sup>1</sup></li> <li>✓ 100% ORR in BCMA bispecific-naïve</li> <li>✓ sCR, MRD<sup>-</sup> responses</li> <li>✓ Prior CAR-T, high-risk patient responses</li> </ul>	ots ses
☆	Favorable emerging safety profile and well-tolerated	<ul> <li>✓ No GvHD, DLT</li> <li>✓ Low rates, CRS, neurotox all Gr ≤2</li> <li>✓ Non-viral approach with built in safe</li> </ul>	ty switch
☆	Avoid unnecessary burden	<ul> <li>No invasive patient apheresis</li> <li>No anti-myeloma bridging therapy</li> <li>Low CRS, neurotox limits adjunctive use for side effects</li> </ul>	herapy
*	Reliable quality	<ul> <li>Treatment of all enrolled patients, w in-spec product</li> </ul>	ith
☆	Convenient, rapid, and accessible for patients	<ul> <li>Shipping from inventory</li> <li>Outpatient usage</li> <li>Treatment in 1 week</li> <li>Low manufacturing cost</li> </ul>	

#### 1. In P1/P2 cohorts

ORR = overall response rate; sCR = stringent complete response; MRD = minimal residual disease; GvHD = graft vs. host disease; DLT = dose limiting toxicities; CRS = cytokine release syndrome



# ASH 2023: Rapid and convenient CAR-T administration for entire intent-to-treat (ITT) population without need for apheresis



- 100% of ITT population underwent LD and received
   P-BCMA-ALLO1
   (1 patient had not begun LD by data cutoff date)
- No patient required bridging therapy
- Median time from enrollment to:
  - Start of LD was 1 day<sup>2</sup>
  - P-BCMA-ALLO1 infusion was 7 days<sup>2</sup>
- Patients were heavily pretreated with median 7 lines of therapy; ~40% previous BCMA therapy and 30% high-risk cytogenetics

1. Interim safety analysis on patients (n = 33) given an infusion of P-BCMA-ALLO1 (including cyclic arm patient) and with a minimum of 4 weeks

follow-up. Data cutoff for safety and efficacy analysis was Oct. 23rd, 2023

N=33, analysis excludes patient retreated with P-BCMA-ALLO1
 ITT = intent-to-treat defined at enrollment; LD = lymphodepletion



# ASH 2023: Deep responses and a high response rate in BCMA naïve and prior BCMA therapy exposed RRMM patients receiving adequate lymphodepletion



#### ORR by treatment arm

- 82% overall ORR in pooled P1/P2 cohorts
- 100% ORR in all P1/P2 patients who had received any prior anti-myeloma treatment, including auto BCMA CAR-T, other than teclistamab
- sCR rate 40% (2/5 pts) in P2 cohort responders
- Some P1, P2 patients achieving MRD<sup>-</sup> status, including patients those with highrisk cytogenetics, extra medullary disease
- At data cut off, 8 of 9 responders in P1 &
   P2 arms still in response

Not included in total: one Arm S patient non-evaluable for response assessment and one Arm S patient that was retreated.
 Two P2 patients who had received prior BCMA auto CAR-T (idecabtagene vicleucel and P-BCMA-101), both achieved VGPR.
 ORR = overall response rate; sCR = stringent complete response; MRD = minimal residual disease



## ASH 2023: P-BCMA-ALLO1 is well tolerated in RRMM patients

### SAFETY SUMMARY

- Dose-levels through 6 × 10<sup>6</sup> cells/kg cleared with no DLTs
- No GvHD observed at any dose
- Low CRS incidence (21%), Grade ≤ 2 in severity
- Neurotoxicity (Grade ≤ 2) observed in 2 patients (6%)
  - No Parkinson's-like symptoms
- Serious infections were uncommon even in the higher LD arms
- Grade ≥ 3 TEAEs were associated mainly with LD and myeloma



#### All treatment-emergent adverse events grade $\geq 3$



# ASH 2023: $T_{SCM}$ -rich P-BCMA-ALLO1 cells traffic to tumor site, differentiate to an active oncolytic cell type, and persist

### DATA FROM A PATIENT ACHIEVING AN MRD-NEGATIVE, STRINGENT COMPLETE RESPONSE

- T<sub>SCM</sub>-rich CAR-T can be thought of as prodrugs, meaning they can expand and differentiate into T effector cells after infusion into a patient
- Patient data obtained >6 weeks after treatment supports this premise
  - CAR-T drug product infused was rich in T<sub>SCM</sub> and central memory (T<sub>SCM</sub>) T cells carrying the CAR
  - In bone marrow, where myeloma is found, a tissue sample collected >6 weeks after infusion showed cells, especially CD8+ "killer" T cells became oncolytic effector (T<sub>EFF</sub>) CAR-T cells
  - CAR-T cells were plentiful in the bone marrow, making up ~14% of all marrow lymphocytes at this timepoint, and ~70-fold more concentrated than in peripheral blood
- First known clinical evidence supporting T<sub>SCM</sub>-rich CAR-T hypothesis for an allogeneic CAR-T



Final drug product (FP) phenotype for patient #23 in comparison to CAR-T phenotype by %  $T_{SCM}$ ,  $T_{CM}$ ,  $T_{EM}$  and  $T_{EFF}$  of CD4 and CD8 CAR-T cells



# Summary: P-BCMA-ALLO1 is a promising "off-the-shelf" T<sub>SCM</sub>-rich allogeneic CAR-T therapy based upon preliminary phase 1 results

- Rapid, accessible treatment to meet urgent patient needs
  - 100% treatment of the ITT population with in-spec product and no bridging therapy
  - Median "brain-to-vein" time (enrollment to infusion) of 7 days, including lymphodepletion
- Favorable emerging safety profile
  - No GvHD or DLT and low rates of CRS, neurotoxicity all  $Gr \le 2$
- Deep clinical responses in very heavily pretreated patients receiving adequate lymphodepletion
  - 82% overall ORR in pooled P1/P2 cohorts
  - 100% ORR in all P1/P2 patients who had received any prior anti-myeloma treatment, including auto BCMA CAR-T, other than teclistamab
  - sCR rate 40% (2/5 pts) in P2 cohort responders
  - Some P1, P2 patients achieving MRD- status, including patients with high-risk cytogenetics, extra-medullary disease
- Prodrug-like P-BCMA-ALLO1 cells demonstrate expansion, trafficking to site of malignancy, differentiation, and persistence
- Further clinical development of P-BCMA-ALLO1 is ongoing, with data updates in 2024 (coordinated with Roche)



# Poseida's manufacturing platform, used across all products, delivers $T_{SCM}$ -rich product with high purity





# P-CD19CD20-ALLO1 – Poseida's first dual CAR-T is partnered with Roche



#### Motivation

- Post CAR-T relapse remains a common problem in B-cell malignancies
- Early autologous CD19/CD20 CAR-T data suggests dual targeting can be effective
- Allogeneic approach would be commercially attractive

### **Clinical Trial**

- Phase 1 study enrolling B-cell malignancies (NCT04960579)
- 3x3 design, with flexibility for expansion
- Poseida produced GMP product

#### Status

- Lymphodepletion learnings from other programs incorporated
- Initiated Phase 1 clinical trial

## P-CD19CD20-ALLO1



- Differentiated, carrying 2 full length CARs and other Poseida platform elements<sup>1</sup>
- First known allogeneic dual CD19+CD20 targeting CAR-T

## Data update planned for 2H24\*

\*Subject to coordination with Roche



## P-MUC1C-ALLO1 is Poseida's lead solid tumor allogeneic CAR-T program

#### Motivation

- High unmet medical need in many epithelial cell-derived tumors
- Poseida's autologous PSMA program showed clinical effect of T<sub>SCM</sub>-rich CAR-T in a solid tumor

### **Clinical Trial**

- Phase 1 basket study enrolling treatment resistant breast, ovarian, pancreatic and other tumors (NCT05239143)
- Flexible 3x3 design; Poseida produced GMP product

#### Status

- Favorable early data (ESMO-IO 2022)
  - No DLT, CRS, GvHD or neurotoxicity
  - PR in breast cancer
- Ongoing exploration of dosing regimen
  - Cell dose, increasing lymphodepletion, dose administration

### P-MUC1C-ALLO1



- Unique approach to targeting MUC1C protein at tumor specific moiety
- Also carries Poseida's platform<sup>1</sup> elements

## Data update planned for 2H24

1. Safety switch, selectable marker, TCR KO,  $\beta$ 2M KO DLT=dose limiting toxicities; CRS= cytokine release syndrome; GvHD= graft vs. host disease



## The power of partnerships: Roche collaboration validates allogeneic platform

Meaningful clinical progress for partnered programs and ongoing milestone achievement

### OVERVIEW

- P-BCMA-ALLO1 licensed
- P-CD19CD20-ALLO1 licensed
- P-BCMACD19-ALLO1 option program
- P-CD70-ALLO1 option program
- Research collaboration with 6 additional Heme Targets



#### ECONOMIC SUPPORT

- \$110 million in upfront payments (August 2022)
- Based on progress made, recent milestone acceleration extended cash runway by ~6 months, runway now into the second half of 2025
- Research, development, launch, and net sales milestones and other payments potentially up to \$6 billion in aggregate value, plus royalties
- Significant direct and indirect support from Roche and additional upside

Excitement Around Heme CAR-T

Continued Validation of Platform

Supporting Financial Position



In April 2024, Poseida entered into research collaboration and licensing agreement with Astellas to develop novel allogeneic cell therapies targeting solid tumors



- Astellas to leverage Poseida's allogeneic CAR-T platform to develop innovative programs targeting solid tumors in combination with Astellas' *convertible*CAR technology
- Research collaboration and license agreement for two solid tumor product candidates based on one Poseida-developed CAR-T construct with first target identified
- Astellas to reimburse Poseida for research costs and will be responsible for development and commercialization of collaboration products
- Poseida received \$50 million upfront plus potential developmental and sales milestones and contingency payments of up to \$550 million, in addition to low double digit tiered royalties as a percentage of net sales

Collaboration enables development in areas beyond Poseida's core pipeline focus and expands strategic investment Astellas announced in August 2023 to support the advancement of Poseida's breakthrough research



# Poseida's consistent platform reflects a holistic systems engineering approach to CAR-T set to deliver product after product



Poseida has taken a deliberate platform approach to develop a pipeline of allogeneic CAR-T programs that use the same proprietary technologies, the same manufacturing platform and reflect similar production methods



## Platform enables delivery of a continuous pipeline of products







# **Genetic Medicines**

Non-viral Delivery for Gene Insertion and Gene Editing to Enable Access for All Patients



# Poseida's vision for genetic medicine

Effective - capacity to cure\*

Safe – non-viral, low immunogenicity lipid nanoparticles

Provide patients with corrective, transformational therapeutic benefit through medicines that insert, delete, or modify genes Durable - stable genome editing/insertion

Patient-friendly - single or short course of treatment

Scalable - can be produced at scale and cost-effectively

Broad applicability - treat patients of all types & ages

Versatile – insert genes of any size, remove genes or signals, across cell types



## Our non-viral delivery technology is poised to unlock the field of genetic medicine





## This product vision requires an entirely new suite of technologies



- Integrated, stable expression
- Large cargo capacity for whole genes
- Safe harbor insertion, including in non-dividing cells
- Re-dosable, reversible and scarless

High fidelity gene editing

RNA-guided DNA nuclease

Exceptional fidelity

- Efficient
- Applicable to different cell types
- Multiplexing potential



- ✓ Low immunogenicity
- Titrate-to-efficacy dosing
- ✓ Scalable
- Favorable cost of goods

Our technologies could be used individually or together to deliver transformational therapies



## Versatility in developing products tailored to therapeutic need

Potential to add proprietary innovation elements onto core technology components





# DNA insertion technology enables whole gene functional correction

Key advantages of our gene insertion approach over Cas9 knock-ins and episomal strategies



Poseida is progressing *Site-Specific Super piggyBac (ssSPG)*, our next generation gene editing/insertion technology with the potential for unprecedented on-target efficiency



# Cas-CLOVER: Potentially the cleanest gene editing

High-fidelity system offers potential as highly differentiated system with low to no off-target editing



Cas-CLOVER gene editing system yields **20-40x higher fidelity** than Cas9 High on-target performance

Differentiated system with low to no off-target editing across multiple cells/targets

 Madison et al., Mol Ther Nucleic Acids. 2022; 2. Alvarez et al., Mol Ther. 31(4), Supp. 1, S1-794. 2023. 3. Data on file, Manuscript in preparation (Poseida Therapeutics) 4. Gilmore et al., NEJM 2021; 5. Longhurst et al., NEJM 2024; 6. Ren et al., Clin Cancer Res., 2017; 7. Antoniani et al., Blood. 2018; 8. Georgiadis et al., Mol Ther. 2018; 9. Webber et al., Nature Comm., 2019; 10. Fix et al., J Immunother Cancer. 2022; 11. Ottaviano et al. Sci. Trans. Med., 2022; 12. Zhang et al., Nature., 2022; 13. Cancellieri et al., Nature Genetics 2023.



## Delivery: Non-viral LNP technology enables broad hepatocyte DNA delivery

Technology advancements enabling Poseida as a leader in non-viral gene delivery



Juvenile immunocompetent mice single dose of LNP intravenously; immunostaining for transgene protein (pink).

Poseida LNP can deliver both DNA and RNA



## Focused development of key programs within areas of significant opportunity





# P-KLKB1-101: HAE patients have an unmet need for a safe therapy with durable efficacy



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 Ren et al., Clin Cancer Res., 2017; 2. Antoniani et al., Blood. 2018; 3. Georgiadis et al., Mol Ther. 2018; 4. Webber et al., Nature Comm., 2019; 5. Gilmore et al., NEJM 2021; 6. Fix et al., J Immunother Cancer. 2022; 7. Ottaviano et al. Sci. Trans. Med., 2022; 8. Zhang et al., Nature., 2022; 9. Cancellieri et al., Nature Genetics 2023; 10. Longhurst et al., NEJM 2024. 11. Madison et al., Mol Ther Nucleic Acids. 2022; 12. Data on file, Manuscript in preparation (Poseida Therapeutics)



# Stable targeted reduction of HAE biomarker with KLKB1 gene editing



Interim non-human primate (NHP) data demonstrate favorable tolerability & liver editing approaching desired therapeutic range



## Poseida's non-viral system has potential to address unmet needs for Hemophilia A patients

- Non-viral lipid nanoparticle (LNP) delivery less immunogenic
- Greater access without concerns of prior viral exposure
- Titrate-to efficacy, or re-dosing, for a personalized therapy

- Large transposon cargo capacity enables whole gene restoration
- Optimally suited for both FVIII gene along with key *cis*-regulatory elements



- Transposition in hepatocytes for potential long-term durability
- 13 months of FVIII expression with potential for longer
- Key advantages in adolescents, for early intervention

- Flexibility: modulate through an inducible off-switch
- Titrate down, switch off, or swap out therapies

Durability, pediatric ineligibility, non-viral, and personalized dosing are significant limitations associated with gene therapies available to Hemophilia A patients today



# P-FVIII-101: Non-viral delivery of Factor VIII via piggyBac for Hemophilia A

Hemophilia A opportunity remains wide-open for a better, more durable gene therapy approach

- Caused by deficiency in functional coagulation Factor VIII
- ~1 in 5,000 male births with ~60% of patients suffering from severe form<sup>1</sup>
- Disease managed with lifelong, repeated IV or subcutaneous administrations





#### Nanoparticle delivery of piggyBac enables:

- Permanent disease correction: DNA is stably integrated into liver hepatocytes, durability observed for 13 months
- Large size of F8 easily accommodated with all key regulatory elements; favorable genotoxicity profile
- Biodegradable Lipid Nanoparticle avoids AAV toxicity and enables redosing and treatment all ages



# Poseida has a strong track record for execution with multiple potential milestones on the horizon

#### CAR-T CLINICAL AND MANUFACTURING PROGRESS

- P-BCMA-ALLO1 Phase 1 data (AACR '24, ASH '23, ESMO IO '22)
- P-MUC1C-ALLO1 cell expansion improvement
- Internal GMP manufacturing, with improvements

#### ADVANCING GENETIC MEDICINES TOWARDS CLINIC

- ✓ Genetic medicines **R&D day** (Apr '24)
- Preclinical data and technologies (ASGCT '24, ISTH '23, ASGCT '23, ASH '22)
- Upcoming INTERACT for FVIII; one or more INDs anticipated in '25

#### NON-DILUTIVE FINANCING THROUGH PARTNERSHIPS

- Astellas strategic investment (Aug '23)
- Astellas solid tumor research collaboration (May '24)
- Highly productive Roche collaboration (initiated Aug '22) with acceleration and increased probability of payments

#### **UPCOMING 2024 MILESTONES**

- P-BCMA-ALLO1 in relapsed/refractory multiple myeloma (RRMM):
  - New data anticipated for presentation at the International Myeloma Society 21<sup>st</sup> Annual Meeting, being held September 25-28, 2024, in Rio de Janeiro
  - Additional clinical updates are planned for the second half of 2024, subject to coordination with Roche

#### P-MUC1C-ALLO1 in solid tumors:

Clinical update planned for the second half of 2024

#### P-CD19CD20-ALLO1 in B-cell malignancies:

 Interim data update anticipated in the second half of 2024, subject to coordination with Roche

### P-KLKB1-101 for HAE & P-FVIII-101 for Hemophilia A:

- Data updates anticipated in the fourth quarter 2024
- Poseida Cell Therapy R&D Day November 14



## Strong financial position bolstered by strategic investments

Well capitalized into the second half of 2025, enabling time to key inflection points with upcoming data readouts (P-BCMA-ALLO1, P-MUC1C-ALLO1 and P-CD19CD20-ALLO1)

### Cash Runway Into Second Half 2025

- **\$238M** in cash, cash equivalents and short-term investments as of June 30, 2024
- Well capitalized into the second half of 2025 based on existing cash and expected baseline nearterm payments from Roche

Key Relationships Validate Long-term Vision

- Roche collaboration: Accelerated and increased certainty of achieving upcoming milestones and payments in November 2023
- Astellas research collaboration and license agreement to develop novel allogeneic solid tumor targeted cell therapies for \$50M upfront plus potential developmental and sales milestones of up to \$550M in total

### Potential for Additional Value Creation

- Significant upside potential through achievement of additional milestones and other payments under the Roche collaboration, not already included in current cash runway
- Business development opportunities have potential to further extend cash runway with non-dilutive capital



## Passionate and experienced leadership team driven to unleash value





**Mark Gergen** Executive Chairman

POSE



## **Thank You**

## A New Class of Cell Therapies & Genetic Medicines with the Capacity to Cure

