

# 21ST ANNUAL NEEDHAM VIRTUAL HEALTHCARE CONFERENCE

April 13, 2022

### FORWARD-LOOKING STATEMENTS

This presentation contains forward-looking statements within the meaning of, and made pursuant to the safe harbour provisions of, The Private Securities Litigation Reform Act of 1995. All statements contained in this document, other than statements of historical facts or statements that relate to present facts or current conditions, including but not limited to, statements regarding possible or assumed future results of operations, business strategies, research and development plans, regulatory activities, market opportunity, competitive position and potential growth opportunities are forward-looking statements. These statements involve known and unknown risks, uncertainties and other important factors that may cause our actual results, performance or achievements to be materially different from any future results, performance or achievements expressed or implied by the forward-looking statements. In some cases, you can identify forward-looking statements by terms such as "may," "might," "will," "should," "expect," "plan," "aim," "seek," "anticipate," "could," "intend," "target," "project," "contemplate," "believe," "estimate," "predict," "forecast," "potential" or "continue" or the negative of these terms or other similar expressions. The forward-looking statements in this presentation are only predictions. We have based these forward-looking statements largely on our current expectations and projections about future events and financial trends that we believe may affect our business, financial condition and results of operations. These forward-looking statements speak only as of the date of this presentation and are subject to a number of risks, uncertainties and assumptions, some of which cannot be predicted or quantified and some of which are beyond our control, including, among others: our ability to successfully advance our current and future product candidates through development activities, preclinical studies, and clinical trials; our reliance on the maintenance on certain key collaborative relationships for the manufacturing and development of our product candidates; the timing, scope and likelihood of regulatory filings and approvals, including final regulatory approval of our product candidates; the impact of the COVID-19 pandemic on our business and operations, supply chain and labor force; the performance of third parties in connection with the development of our product candidates, including third parties conducting our future clinical trials as well as third-party suppliers and manufacturers; our ability to successfully commercialize our product candidates and develop sales and marketing capabilities, if our product candidates are approved; and our ability to maintain and successfully enforce adequate intellectual property protection. These and other risks and uncertainties are described more fully in the "Risk Factors" section of our most recent filings with the Securities and Exchange Commission and available at www.sec.gov. You should not rely on these forward-looking statements as predictions of future events. The events and circumstances reflected in our forward-looking statements may not be achieved or occur, and actual results could differ materially from those projected in the forward-looking statements. Moreover, we operate in a dynamic industry and economy. New risk factors and uncertainties may emerge from time to time, and it is not possible for management to predict all risk factors and uncertainties that we may face. Except as required by applicable law, we do not plan to publicly update or revise any forward-looking statements contained herein, whether as a result of any new information, future events, changed circumstances or otherwise.





# Future Vision

Establish franchises in B cell malignancies, GBM, solid tumor indications

Century is developing transformative allogeneic cell therapies

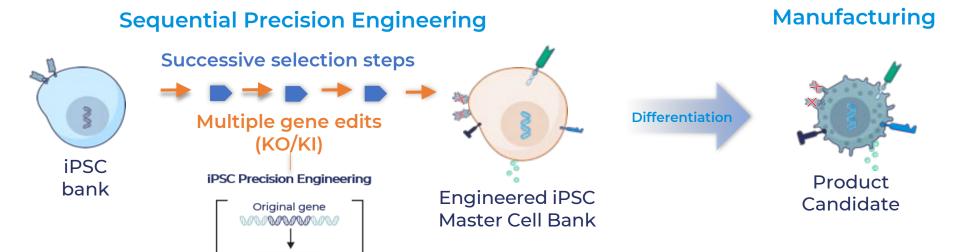
Develop best-in-class iPSC-derived iNK and iT cell platforms

World-class cell engineering capabilities

Financial strength maintained by platform synergies, efficiencies



### PRECISION CRISPR MAD7 GENE EDITING OF IPSC CELLS UNLOCKS TRANSFORMATIONAL POTENTIAL



CRISPR nuclease

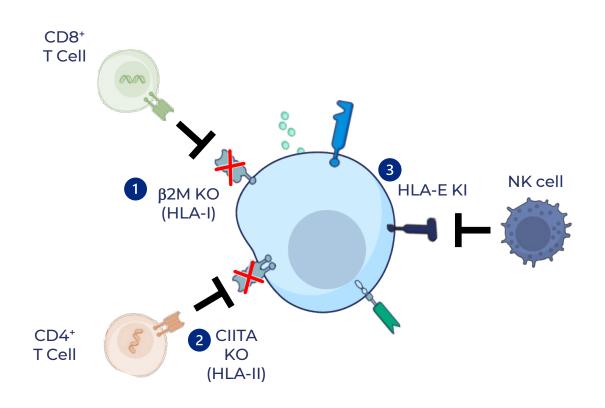
Replacement gene

CRISPR-mediated HDR (MAD7)

- Precise gene editing (MAD 7) enables engineering of candidates with synthetic functionalities
- Quality control by ensuring genomic **integrity** is maintained
- Clonal selection of MCB for homogenous products, scalable process



### ALLO-EVASION<sup>TM</sup> 1.0 DESIGNED TO OVERCOME THREE MAJOR PATHWAYS OF HOST VS GRAFT REJECTION



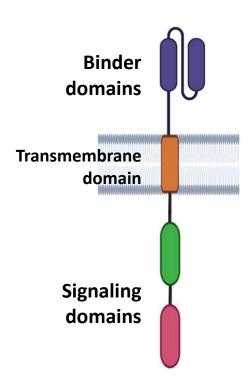
#### Core edits

- 1 Deletion of β2M, a protein required to express HLA-I on the cell surface prevents recognition by CD8 T cells
- 2 Knock out of CIITA eliminates HLA-II expression to escape elimination by CD4 T cells
- 3 Knock-in of HLA-E prevents killing by NK cells



### CENTURY'S PROTEIN SCIENCES ENABLES ALL ASPECTS OF PROTEIN ENGINEERING OF OUR PRODUCT CANDIDATES

#### **CAR Structure**



### **CAR Engineering**

- Enable fit-for-purpose CAR assembly
  - Multiple CAR formats for single- and multi-specific CARs
- CAR Binder Discovery
  - Century Therapeutics' proprietary VHH library
  - Partnerships with Distributed Bio and LakePharma
- Signaling Domains
  - Signaling domains specifically optimized for iNK and iT cells

### **Transgene Engineering**

- Allo-Evasion molecules
- Safety switch
- Cytokines
- · Chimeric proteins

### **CRISPR-MAD7 Protein Engineering**

• Protein optimization, production, and characterization





### Century's multi-suite facility Branchburg, NJ

- cGMP manufacturing facility expected to be operational in 2022
  - 53,000 ft<sup>2</sup> facility
- Capable of production of NK and T cells, as well as other cell types

### Advantages of investing in manufacturing capabilities expertise

- Fully integrated analytical and process development and manufacturing
- More effectively implement process and product changes
- Potentially accelerate clinical development of commercial product
- Potential to achieve economies of scale

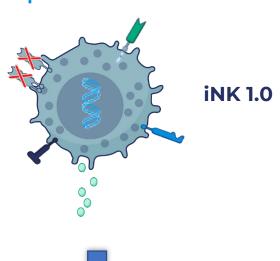


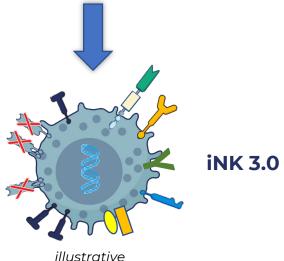
# CELL ENGINEERING ENABLES RAPID PRODUCT ITERATION

Proposed Feature	CNTY-101 iNK 1.0	CNTY-103 iNK 2.0	iNK 3.0 Illustrative
Cytokine support	Soluble IL-15	mbIL-15	mbIL-15
Innate Activity	+	++ (NKG2A KO)	++ (NKG2A KO)
Undisclosed			<b>\</b>
PET tracer			<b>✓</b>
Allo-Evasion			
HLA-1 KO	<b>/</b>	<b>/</b>	<b>/</b>
HLA-II KO	<b>/</b>	<b>/</b>	<b>/</b>
HLA-E KI	<b>/</b>	<b>/</b>	<b>/</b>
Undisclosed			<b>/</b>
# of edits	6	6	12
КО	2	3	4
Site-specific KI	4	3	8

Century's Platform allows for potentially transformative product evolution

#### Fit for purpose tailored functionality



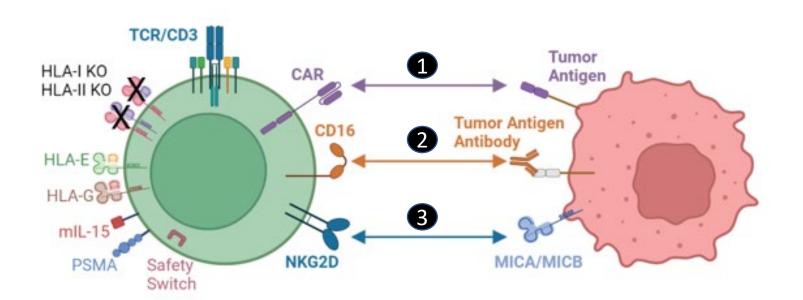




## CENTURY'S FIRST GENERATION γδ CAR-iT CELLS MULTIPLE BUILT-IN PATHWAYS FOR TUMOR KILLING

γδ CAR-iT cell

Tumor cell



### PATHWAYS FOR POTENTIAL TUMOR KILLING

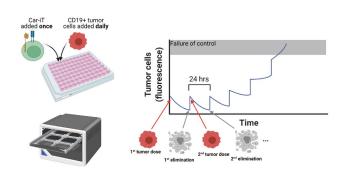
- 1. CAR-mediated killing
- 2. ADCC (Antibody-dependent cellular cytotoxicity)
- 3. Innate-receptor mediated killing (NKG2D, others)

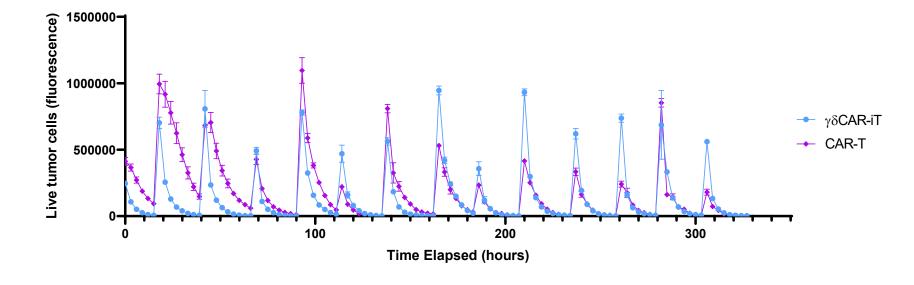


### γδ CAR-iT CELLS KILL LYMPHOMA CELLS THROUGH MULTIPLE ROUNDS OF KILLING WITHOUT REACHING EXHAUSTION

#### **Serial Killing Assay Setup**

#### Serial Killing of CD19<sup>+</sup> Lymphoma Cells by γδ CAR-iT cells







### **CENTURY'S EMERGING FRANCHISES**



- CNTY-101: Lead product candidate, CD19 targeted CAR-iNK
- **CNTY-102**: First  $\gamma\delta$  iT candidate, multi-specific (CD19 + CD79b) CAR-iT

Potentially first product candidate to enter the clinic with edits designed to avoid all major pathways of rejection

CNTY-102 provides optionality to address additional subtypes / use in combination



- CNTY-103: Multi-specific (CD133+ EGFR) CAR iNK for recurrent GBM
- Follow-on candidate

Multi-tumor antigen targeting addresses heterogeneity in GBM tumor cells

Exploring additional tumor antigens and TME modulation features



Solid tumors

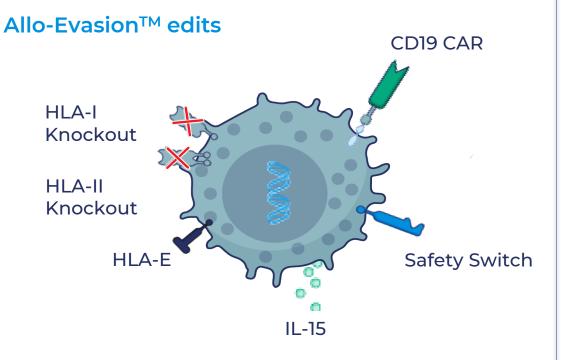
**Future candidate** expected to be announced 4Q 2022

Leverage  $\gamma\delta$  iT platform to target challenging solid tumors



### CNTY-101: NEXT GENERATION CD19 TARGETED PRODUCT

#### HIGHLY DIFFERENTIATED PROFILE



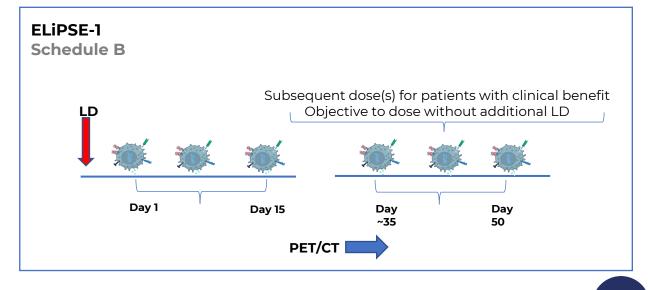
#### **CNTY-101**

First cell product candidate with 6 gene edits introduced with CRISPR-HDR



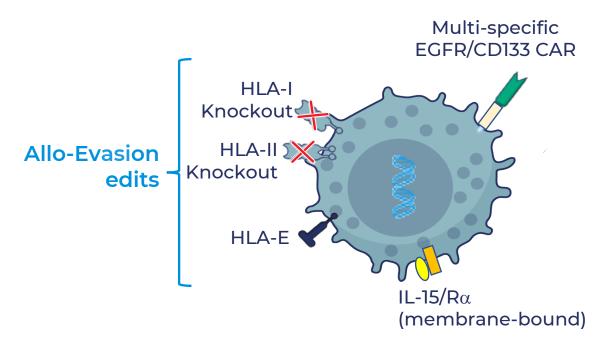
#### **ELIPSE-1: PHASE 1 STUDY DESIGN\***

- Explore 1 to 2 cycles up to 6 doses with single LD conditioning
- Effect of Allo-Evasion on iNK persistence after multiple doses
  - IND expected mid-2022 to advance CNTY-101 into Phase 1 clinical trial



<sup>\*</sup> Subject to FDA approval

### CNTY-103: FIRST PROGRAM IN CNS MALIGNANCIES



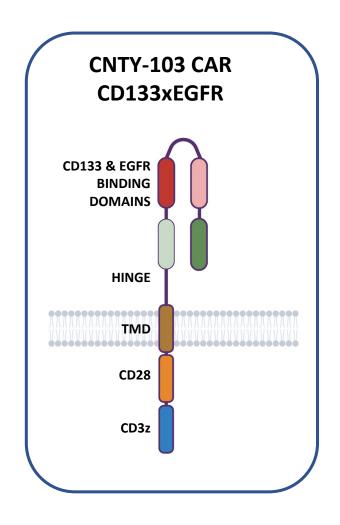
- Targeting EGFR+ GBMs: Provides better coverage intended to ensure maximum tumor clearance
- Targeting CD133+ GBMs: Clears functionally relevant treatment-resistant tumor cells

### UNIQUELY POSITIONED TO ADDRESS CHALLENGES OF GBM

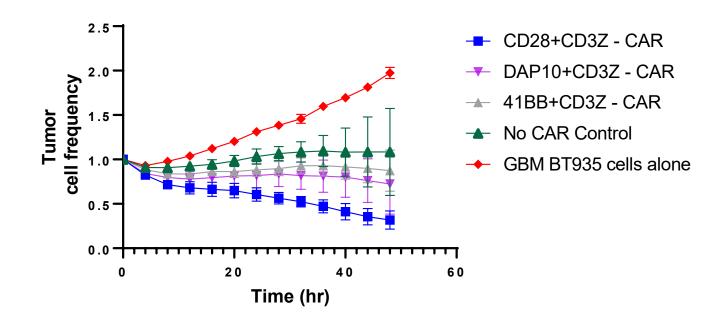
Challenge	Century's Solution			
Trafficking	Local delivery with Ommaya reservoi			
Heterogeneity	Targeting EGFR / EGFRvIII and CD133 (functional marker of treatment-resistant GBM cells)			
Toxicity	Potentially minimize risks like CRS with iNK			
Persistence	Potential to dose as needed			



## MULTIPLE CAR DESIGNS AND SIGNALING DOMAINS WERE INVESTIGATED TO DEVELOP THE CD133xEGFR CAR



Comparison Of The Cytotoxic Activity Of Multiple CD133xEGFR CAR
Constructs Engineered With Different Signaling Domains





Product	iPSC Platform	Targets	Indications	Expected IND Submission	Discovery	Preclinical	Clinical	Collaborator
CNTY-101	iNK	CD19	B-Cell Malignancies	Mid 2022				
CNTY-103	iNK	CD133 + EGFR	Glioblastoma	2023				
CNTY-102	iΤ	CD19 + CD79b	B-Cell Malignancies	2024				
CNTY-104	iNK/iT	Multi- specific	Acute Myeloid Leukemia	2024				ر <sup>ال</sup> ا Bristol Myers Squibb
CNTY-106	iNK/iT	Multi- specific	Multiple Myeloma	2024				الا Bristol Myers Squibb





# CENTURY THERAPEUTICS' UNIQUE INVESTMENT OPPORTUNITY

- Well capitalized with cash runway into 2025
  - Financial strength maintained by operational synergies
- Comprehensive end-to-end allogeneic platform
  - iPSC derived iNK and iT cells, world-class gene editing, protein engineering and manufacturing
- CNTY-101 IND submission on track for mid-2022
  - CD19 targeted cell product designed to avoid 3 pathways of host rejection
- CNTY-103 IND enabling activities to initiate in 2022
  - Century's first solid tumor candidate for GBM
- Experienced partner on board to tackle challenging malignancies
  - Bristol-Myers Squibb brings expertise in oncology and hematology



