

# H.C. WAINWRIGHT GLOBAL INVESTMENT CONFERENCE

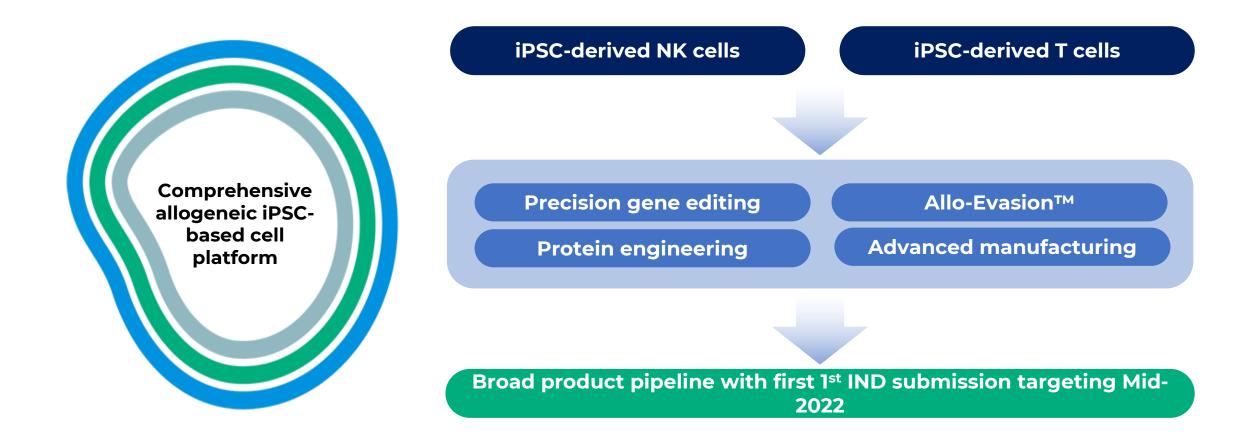
May 24, 2022

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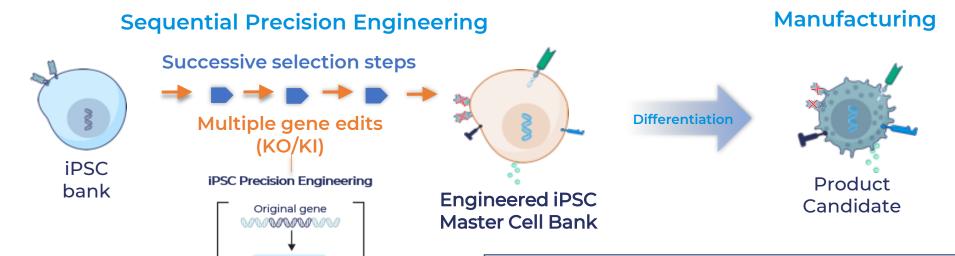


# CENTURY'S NEXT GENERATION IPSC TECHNOLOGY PLATFORM





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



CRISPR nuclease

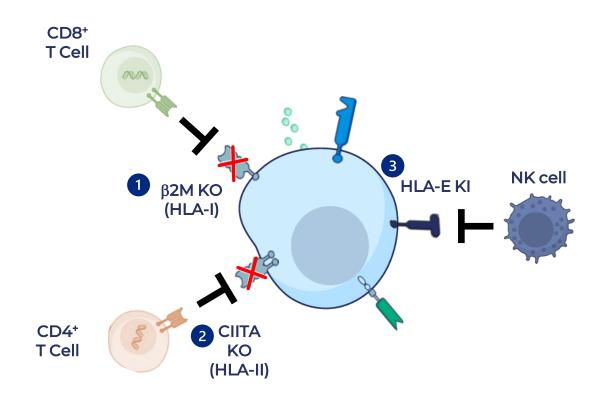
Replacement gene

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™ 1.0 DESIGNED TO OVERCOME THREE MAJOR PATHWAYS OF HOST VS GRAFT REJECTION



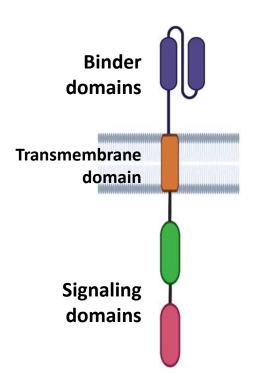
#### **Core edits**

- 1 Deletion of  $\beta$ 2M, 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



# CUTTING EDGE PROTEIN ENGINEERING ENABLES DIFFERENTIATED 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

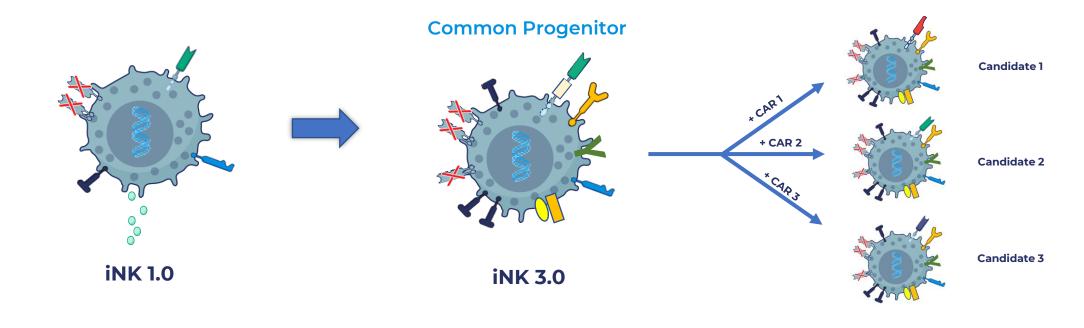
- Fully integrated analytical and process development and manufacturing
- More effectively implement process and product changes
- Potentially accelerate clinical development of commercial product
- Potential to achieve economies of scale



# COMMON PROGENITOR ENABLES EFFICIENT PIPELINE EXPANSION



Additional details to be highlighted at Century's Virtual R&D Day (June 13)



# of edits 6

#### iNK 3.0 common progenitor platform incorporates multiple features:

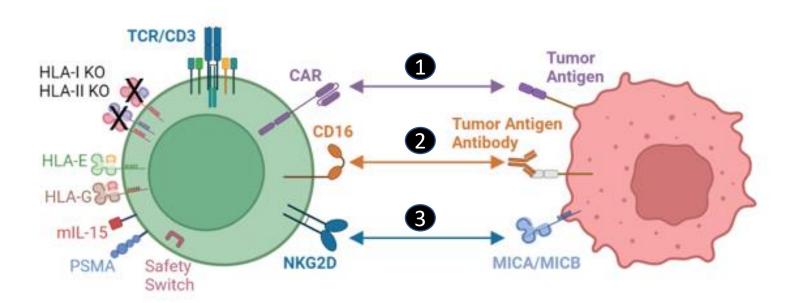
- · Additional Allo-Evasion edits
- Additional innate killing potential
- Additional anti-cancer targeting option
- Cytokine support



# 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

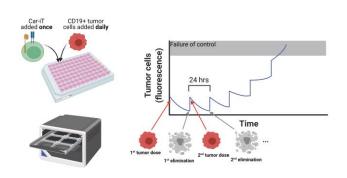
- 1. CAR-mediated killing
- 2. ADCC (Antibody-dependent cellular cytotoxicity)
- 3. Innate-receptor mediated killing (NKG2D, others)

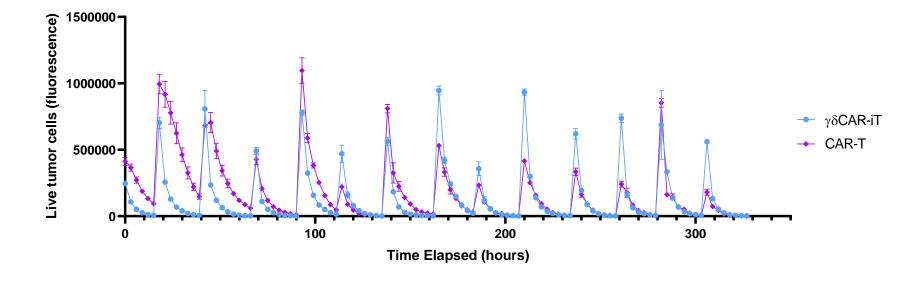


## $\gamma\delta$ CAR-iT CELLS KILL LYMPHOMA CELLS THROUGH MULTIPLE ROUNDS OF KILLING WITHOUT REACHING EXHAUSTION

#### **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

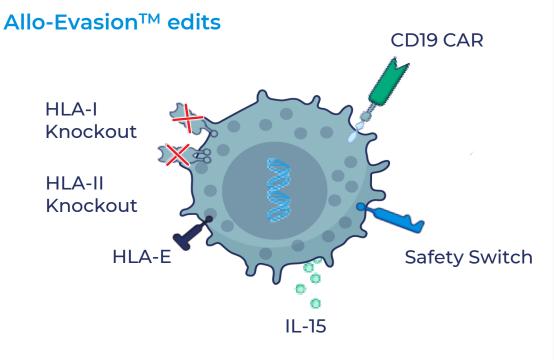
**Future candidate** expected to be announced 4Q 2022

Leverage γδ iT platform to target challenging solid tumors



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

#### HIGHLY DIFFERENTIATED PROFILE

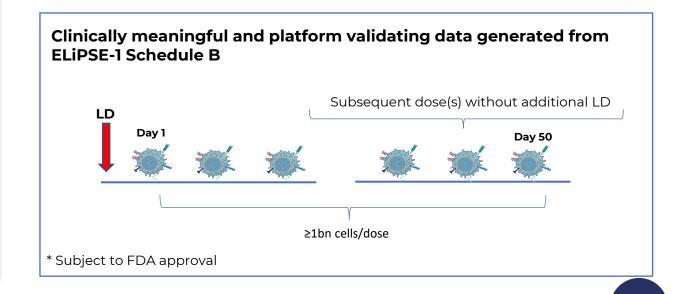


#### **CNTY-101**

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

#### **ELIPSE-1: PHASE 1 STUDY GOALS**

- Dose escalation with multiple doses and single lymphodepletion conditioning regimen
  - IND submission on track for mid-2022 to advance CNTY-101 into Phase 1 clinical trial





### Product candidate pipeline across cell platforms and targets in solid and hematologic cancers

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>اال</sup> Bristol Myers Squibb
CNTY-106	iNK/iT	Multi- specific	Multiple Myeloma	2024				ر <sup>اال</sup> Bristol Myers Squibb
Discovery Research Programs								
	ink/iT	TBD	Solid Tumors	TBD				
	iNK/iT	TBD	Hematological Malignancies	TBD				

**Solid Tumors** 



# 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
- Poised to become clinical stage company with CNTY-101 IND submission mid-2022
  - Product engine enables delivery of 5 INDs over next 3 years



