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# CENTURY THERAPEUTICS - EMERGING LEADER IN IPSC CELL THERAPIES

COMPREHENSIVE
iPSC CELL PLATFORM
FOR IMMUNE
EFFECTOR CELLS

PRODUCT CANDIDATE ENGINE
WITH PIPELINE IN SOLID AND
HEMATOLOGIC MALIGNANCIES

**LEAD PROGRAM**ON TRACK TO FILE IND MID 2022

### **EXPERTISE**

GENETIC & PROTEIN ENGINEERING, PROCESS DEVELOPMENT, AND IMMUNO-ONCOLOGY STATE-OF-THE ART GMP
MANUFACTURING FACILITY EXPECTED
TO BE OPERATIONAL 1Q 2022

# HEADQUARTERED IN PHILADELPHIA

WITH CENTERS OF EXCELLENCE IN SEATTLE AND ONTARIO

\$400.3M

IN CASH, CASH EQUIVALENTS AND MARKETABLE SECURITIES AS OF 9/30/2021 ~150

EMPLOYEES INCLUDING EXPERIENCED LEADERS AND ENTREPRENEURS



### WE ARE BUILDING A NEXT GENERATION CELL THERAPY PLATFORM

- Complex genetic editing
- High replication capacity
- Streamlined manufacturing
- Consistent product
- · Limited gene editing
- Finite replicative capacity
- Lengthy, expensive manufacturing
- Potential weakened donor cells

iPSC derived

Allogeneic (Healthy donor derived)

**Autologous (Patient derived)** 



#### NEXT GENERATION ALLOGENEIC iPSC-BASED PLATFORM

Hypoimmunogenic products generated with Allo-Evasion<sup>TM</sup> technology

#### ALLO-EVASION™ PRODUCES HYPOIMMUNOGENIC CELLS

To potentially prevent graft rejection by patient

#### FIT-FOR-PURPOSE PRODUCTS WITH MULTIPLE GENE EDITS

Cutting edge CRISPR gene editing

#### **MULTISPECIFIC TUMOR TARGETING**

CAR engineering with VHH technology

#### **CAR-INK AND CAR-IT CELL PLATFORMS**

Access to both cell platforms provides optionality



## **PROVEN LEADERSHIP TEAM**



novina MERCK INVENTING FOR LIFE







Osvaldo (Lalo) Flores, CEO

Hy Levitsky, President R&D

Adrienne Farid, COO













**Greg Russotti, CTO** 

Luis Borges, CSO

Michael Diem, CBO



# FOUNDATIONAL INVESTMENTS IN IPSC KNOW-HOW AND MANUFACTURING

# iPSC License and collaboration agreement established in 2018

- Access to clinical grade iPSC lines
- Exclusive IP and know-how to generate immune effector cells using feeder-free methods (NK, T, Mac, DC)
- Dedicated FCDI GMP manufacturing capacity for Century's product candidates
- Leveraging two decades of research & investment at University of Wisconsin and FCDI





# In-House Manufacturing accelerates learnings and enables faster product iteration

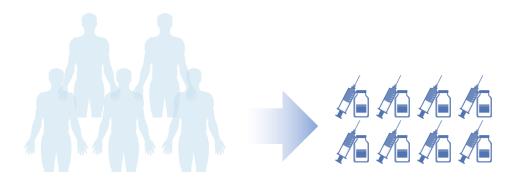
- Century facility expected to be operational by early 2022
  - 53,000 ft<sup>2</sup> facility
  - Designed to produce multiple immune cell types
- Two sites provides optionality and maximizes flexibility





# IPSC TECHNOLOGY CAN OVERCOME LIMITATIONS OF DONOR DERIVED PLATFORMS

## Allogeneic, donor-derived

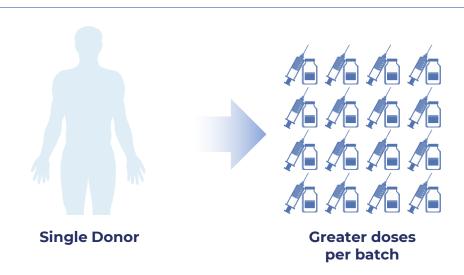


**Multiple donors** 

Fewer doses per batch

- Complex manufacturing, heterogeneous product, limited scale
- Limited genetic engineering options

## Allogeneic, iPSC-derived



- Efficient manufacturing, homogeneous product, greater scale
- Likely unlimited genetic engineering options



### Proprietary CRISPR-MAD7 mediated homology directed repair (HDR)

- MAD7 licensed from Inscripta. Methodology developed at Century
- HDR technology enables precise gene KOs and transgene KIs
- CRISPR protein and guide RNAs delivered using RNP (non-viral)

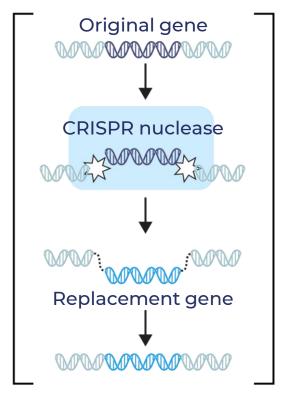
#### Fully characterized, homogeneous drug products

- Sequential gene editing steps allow:
  - Introduction of multiple gene modifications
  - Elimination of clones with chromosomal alterations and structural variants (i.e. translocations, inversions)

#### Master cell banks (MCBs) generated from single-cell clones

- MCBs fully characterized and de-risked genetically
  - Whole genome sequencing
  - Copy number variation (CNV) analysis
  - Transgene copy number by ddPCR

### **iPSC Precision Engineering**

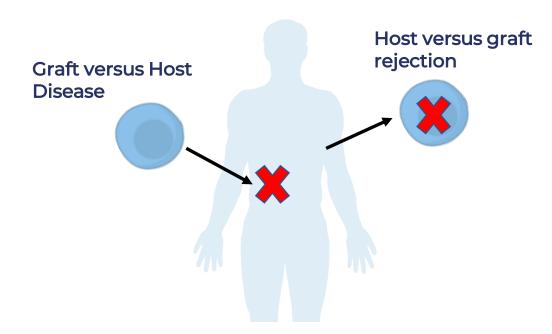


**CRISPR-mediated HDR (MAD7)** 



### OVERCOMING ALLOREACTIVITY CHALLENGES

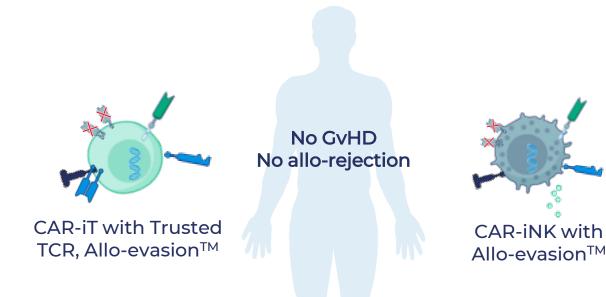
### **Current limitations**



#### Graft versus Host Disease (GvHD)

- Donor T cells damage patient's tissues
   Host versus graft Rejection
  - Patient immune system eliminates allogeneic cells

## Century's solution

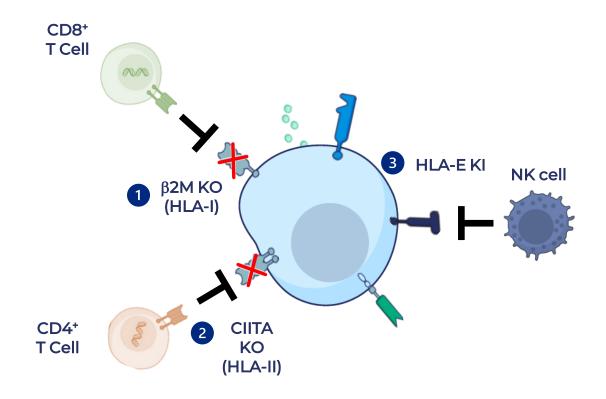


#### Prevention of Graft versus Host Disease

- GvHD is circumvented with Trusted TCR
   Prevention of Allo-Rejection
  - Allo-rejection is circumvented with Allo-evasion™ gene edits



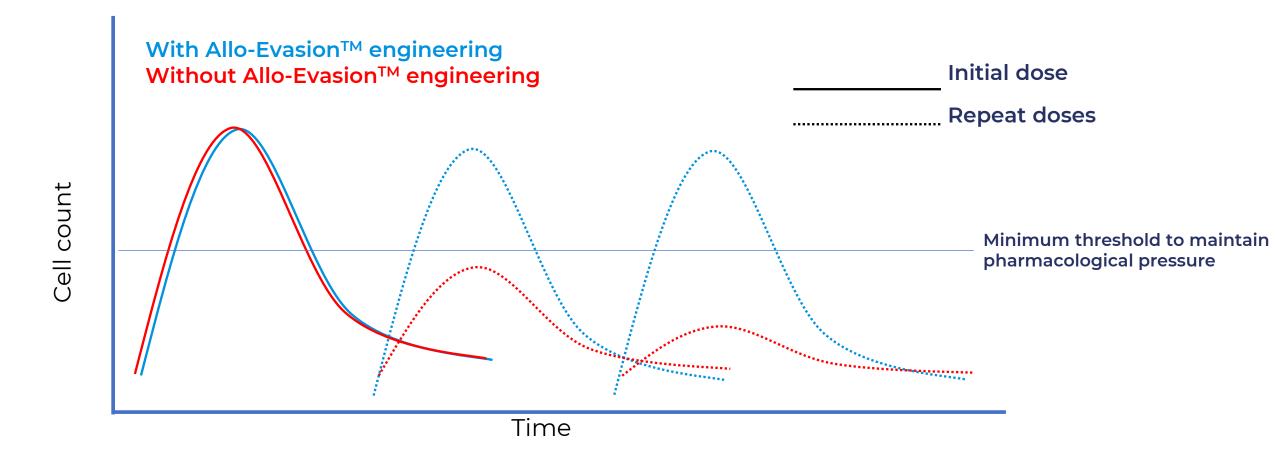
# ALLO-EVASION™ 1.0 DESIGNED TO OVERCOME 3 MAJOR PATHWAYS OF HOST VS GRAFT REJECTION



- Deletion of  $\beta$ 2M, a protein required to express HLA-1 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

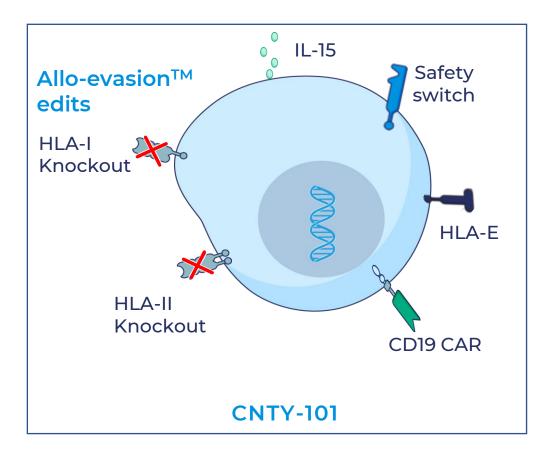


# ILLUSTRATIVE POTENTIAL OF ALLO-EVASION<sup>TM</sup> ON CELLULAR PHARMACOKINETICS AND REPEAT DOSING





# CNTY-101: CAR-INK CANDIDATE IN R/R B-CELL LYMPHOMA



#### CNTY-101 may change the lymphoma treatment paradigm

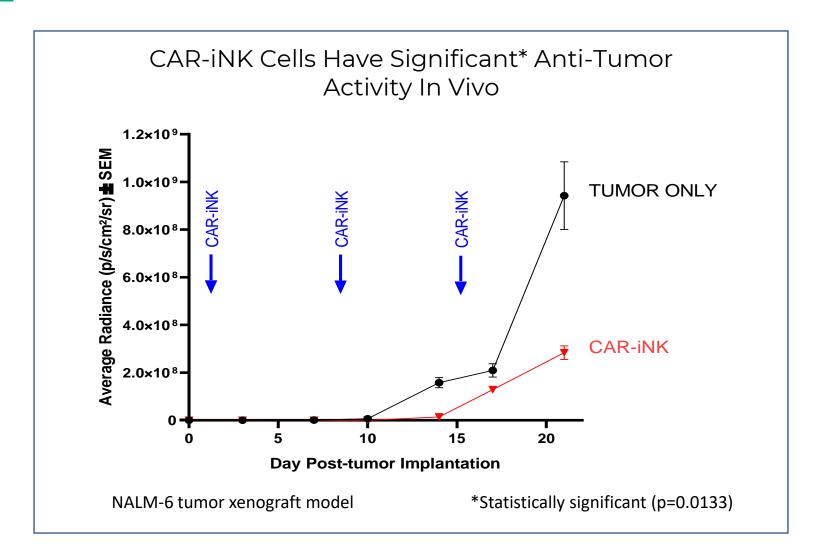
- Has potential to use with milder lymphodepletion regimens
- Potential to re-dose to enhance efficacy
- Designed as an off-the-shelf cell therapy

#### IND filing on track for mid-2022

- P1 in RR NHL set to initiate 2H22
- Generates POC for CAR iNK platform and allo-evasion technology



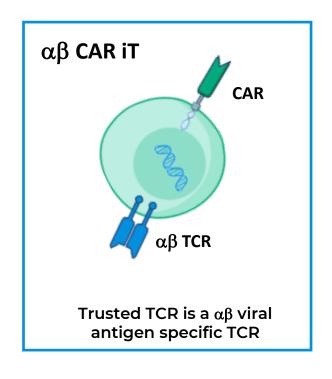
# INK CELL PLATFORM IS OUR MOST ADVANCED PLATFORM

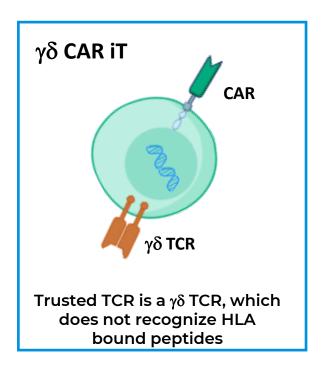




### CENTURY IT CELL PLATFORM: TrueT CELLS WITH TRUSTED TCRs

### Currently Exploring Two Major T Cell Subsets To Develop Century's iT Cell Platform

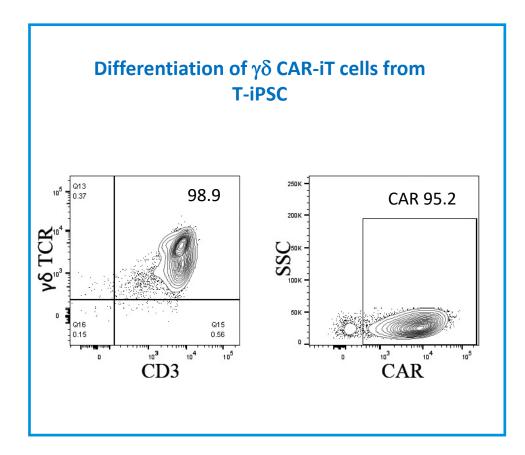




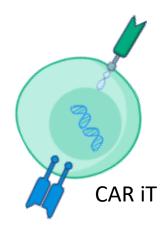
Unique features of Century's iT cell platform:

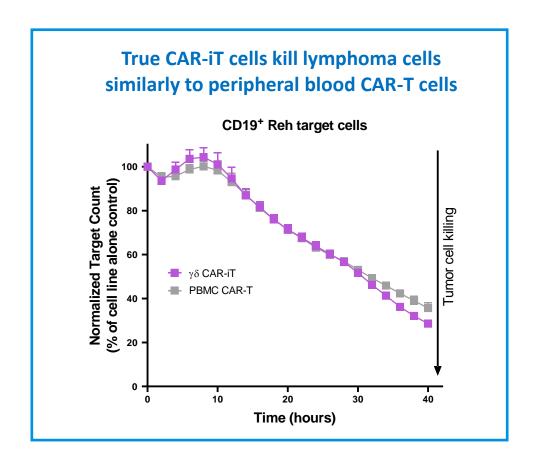
- Retention of a functional TCR intended to improve iT cell differentiation and functionality
- Use of Trusted  $\alpha\beta$  and  $\gamma\delta$  which are not expected to mediate GVHD





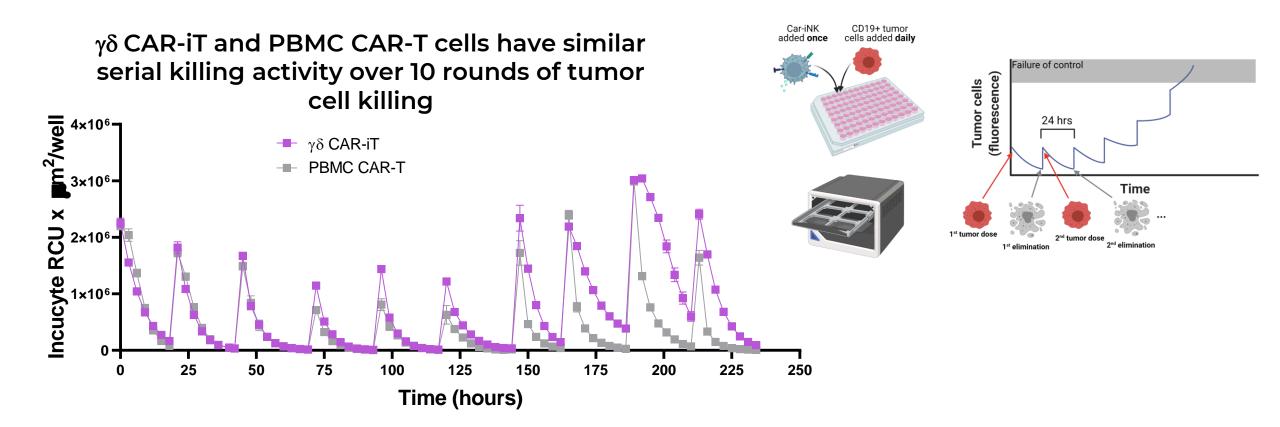
## **TrueT Cells**







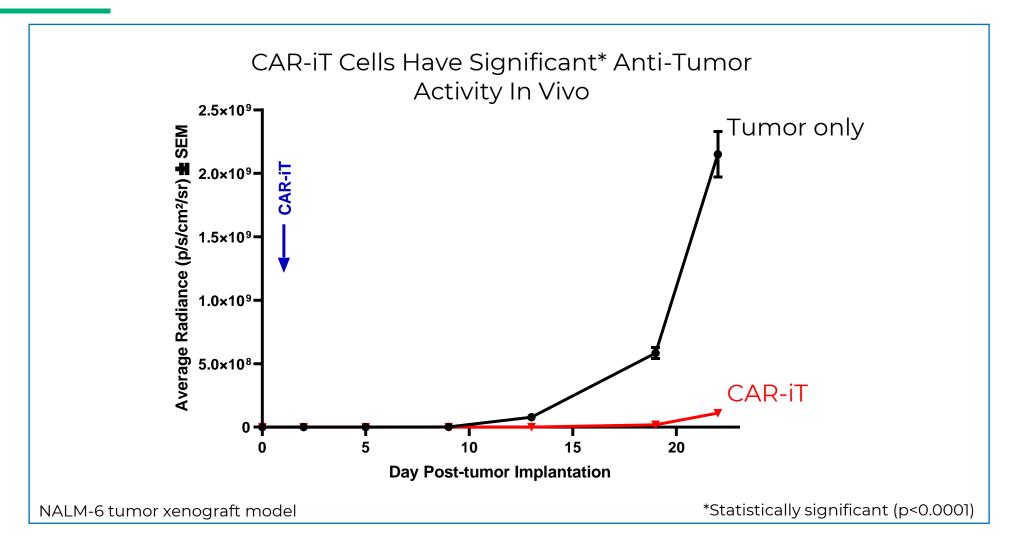
# iPSC-DERIVED $\gamma\delta$ iT CELLS MEDIATE SERIAL KILLING AGAINST LYMPHOMA CELLS





# OUR IT CELL PLATFORM IS CLOSE BEHIND AND MAKING DEMONSTRABLE PROGRESS

Developing  $\alpha\beta$  and  $\gamma\delta$  iT platforms with Trusted TCRs that are not expected to cause GvHD





Product	iPSC Platform	Targets	Indications	Ownership	Expected IND Submission	Discovery Preclinical Phase 1 Phase 2 Phase 3
CNTY- 101	ink	CD19	Lymphoma	CENTURY	Mid 2022	
CNTY- 103	iNK	CD133 + EGFR	Glioblastoma	CENTURY	1H 2023	
CNTY- 102	iT or iNK	CD19 + CD79b	Lymphoma	CENTURY	2H 2023	
CNTY- 104	iT or iNK	Multi- specific	Acute Myeloid Leukemia	CENTURY	1H 2024	





# **CATALYSTS**

Event	Estimated Timing
Close IPO with gross proceeds of \$243M	2Q21 <b>✓</b>
Solicit Pre-IND written feedback for CNTY-101	3Q21 <b>✓</b>
CNTY-101 entered IND enabling studies and manufacturing	4Q21 <b>✓</b>
cGMP manufacturing facility in Branchburg NJ expected to be operational	1Q22
In vivo POC for CNTY-103	1H22
CNTY-101 IND filing	Mid-2022
Initiate CNTY-101 P1 study R/R NHL	2H22
Preliminary safety from CNTY-101 P1	1H23



# CENTURY'S NEXT GENERATION IPSC TECHNOLOGY PLATFORM

