

# **Corporate Overview**

August 2024



## Forward-looking statements

This presentation contains forward-looking statements within the meaning of, and made pursuant to the safe harbor provisions of, The Private Securities Litigation Reform Act of 1995. All statements contained in this presentation, other than statements of historical facts or statements that relate to present facts or current conditions, including but not limited to, statements regarding our clinical development plans and timelines and the initial safety and efficacy profiles of CNTY-101 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 dependence on the success of our lead product candidate, CNTY-101; the ability of CNTY-101 to be administered as part of a multi-dose strategy and to enable responses without lymphodepletion; uncertainties inherent in the results of preliminary data, pre-clinical studies and earlier-stage clinical trials, which may not be predictive of final results or the results of later-stage clinical trials; the timing of

and our ability to successfully enroll the Phase 1 SLE and LN trial; the timing of and our ability to enter dose expansion of the Phase 1 R/R CD19-positive B-cell lymphomas trial; our ability to obtain FDA clearance of our future IND submissions and commence and complete clinical trials on expected timelines, or at all; our reliance on the maintenance of 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 geopolitical issues, banking instability and inflation 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 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; our ability to recruit and maintain key members of management 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.



## Century Therapeutics: Building an industry-leading, nextgeneration allogeneic iPSC-derived cell therapy platform

LIMITLESS POTENTIAL...

PRECISION DESIGN...

**ENDURING IMPACT...** 

Foundational investments in iPSC technology, genetic editing, protein engineering, and manufacturing

Progressing differentiated clinical programs based on Allo-Evasion<sup>™</sup> technology in oncology and autoimmune diseases

Well-capitalized into 2026 to enable delivery on key milestones and clinical data



Overview of Foundational Platform Technologies

#### Century's singular focus:

To deliver best-in-class iPSC-derived cell therapies

# Century platform enables the incorporation of critical features we believe can <u>only</u> be realized via iPSC-derived cell therapies

Infinite replicative capacity at the iPSC stage enables potentially unlimited genomic editing via CRISPR HDR

Single cell cloning of engineered iPSC allows selection of a *fully* characterized clone for a master cell bank, ensuring safety and functional reproducibility of the final drug product

Platform capable of fully **leveraging multiple advances in synthetic biology into a single product** 

Cell expansion during multiple stages of differentiation yields large cell harvests, decreasing risk of cell exhaustion, reducing COGs and providing robust drug inventory that is potentially infinitely replenishable

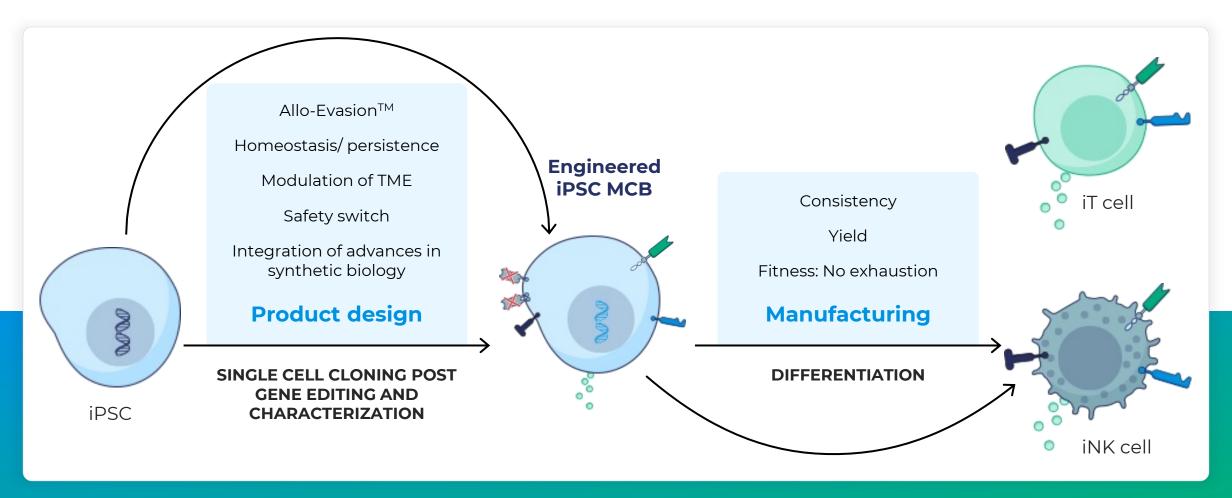
Production from a master cell bank – derived from a single donor – enables *larger* batch sizes and *lower cost of goods than* donor-derived or autologous

Differentiation conditions developed for generating multiple immune effector cells, including NK cells, CD4+ T cells (Th and Treg), CD8+ T cells, monocytes / macrophages



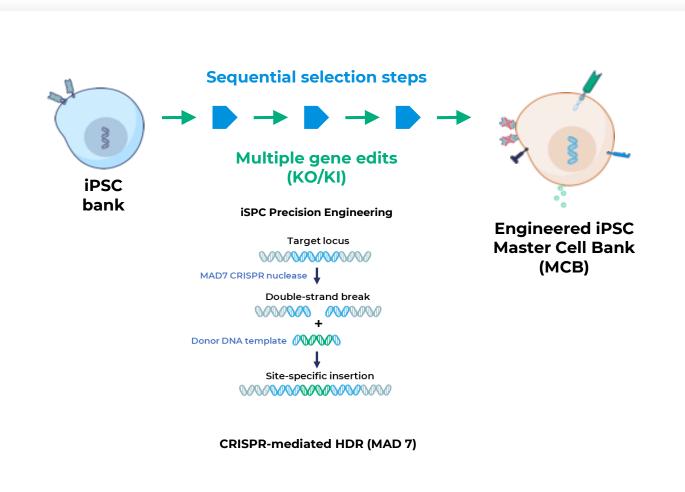
## Century's next-generation allogeneic iPSC technology platform:

Versatility and unprecedented control



Rapid Integration of major advances in product functionality and manufacturability

# Precision CRISPR MAD7 mediated sequential gene editing of iPSC cells generates uniform product candidates



#### **Advantages of Century's Platform**

Precise CRISPR mediated homology directed repair reduces off-target integration

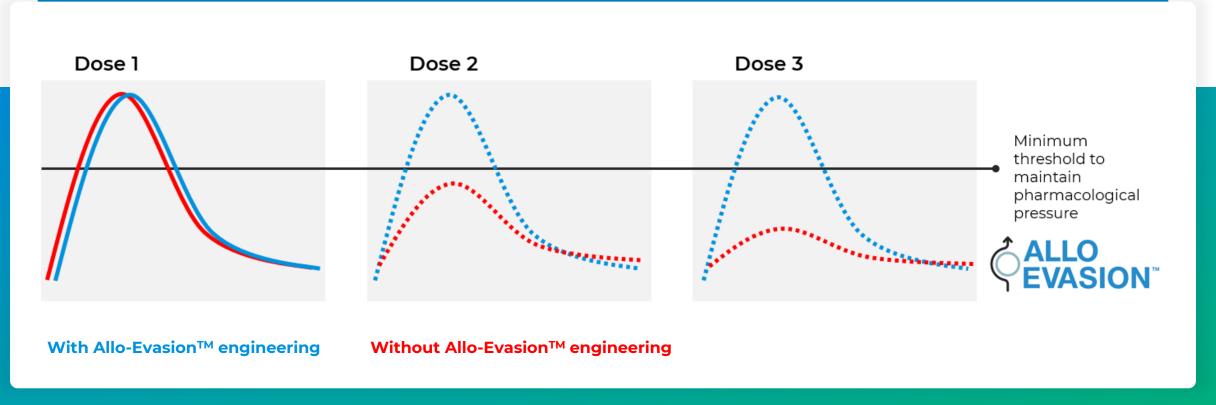
Stepwise and efficient gene editing **avoids risky multiplex modification** and structural variants

**Quality control** through generation of homogenous MCB establishes genomic **product integrity** 

Manufacturing begins at the MCB, confirmed to be **free from genetic aberrations** 

# Potential to drive durable responses with engineering to resist immune rejection

Allo-Evasion<sup>™</sup> edits + repeat dosing = potential greater durability



Next-wave of allogeneic cell therapies must solve for challenge of rejection

# Advancing our leadership in Allo-Evasion<sup>TM</sup> technology

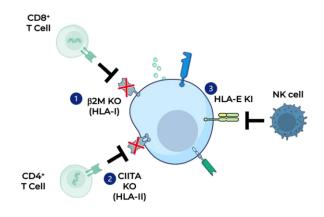
Continuous improvement in holistic immune protection designed to overcome major pathways of host vs. graft rejection

#### Allo-Evasion™ 1.0

#### Allo-Evasion™ 3.0

#### Allo-Evasion™ 5.0

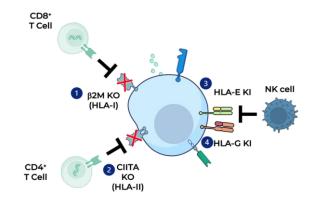
#### Core edits disarm host cells from eliminating therapy



Deletion of \( \beta 2M \), a protein required to express HLA-1 on the cell surface prevents recognition by CD8 T cells

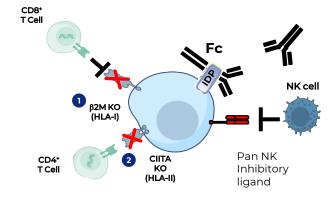
Knock out of CIITA eliminates HLA-II expression to escape elimination by CD4 T cells

Knock-in of HLA-E prevents killing by NK cells



Allo-Evasion™ 1.0 edits plus the incorporation of:

Knock-in of HLA-G improves protection against killing by NK cells



Deletion of 82M. a protein required to express HLA-1 on the cell surface prevents recognition by CD8 T cells

Knock out of CIITA eliminates HLA-II expression to escape elimination by CD4 T cells

Pan-NK inhibitory ligand to provide broader protection against killing by NK cells

IgG degrading protease designed to protect against humoral immunity









# Foundational investments in iPSC manufacturing



#### **Established in-house manufacturing**

- Century 53,000 ft<sup>2</sup> GMP facility
- Designed to produce multiple immune cell types
- Accelerates learnings and enables faster product iteration
- Two sites (FCDI GMP manufacturing, Century in-house manufacturing) provide optionality and maximizes flexibility

#### **Developing fit-for-purpose products**

- Increased process and product consistency
- Scalable platforms and optimized processes to maximize yield, reduce COGs, and meet demand
- Increased cell fitness, as cells do not undergo excessive expansion cycles which often result in cell exhaustion
- Homogeneity of the manufacturing process produces a product candidate that can be readily characterized



### Newly expanded and diversified pipeline

Product candidates spanning cell types and targets in cancer and autoimmune diseases

						Clinical			
Product	iPSC Platform	Targets	Indications	Research	IND-Enabling	P1	P2	P3	Collaborator
Autoimmune Diseases									
CNTY-101	iNK	CD19	B cell-mediated Autoimmune Diseases	CALIPSO-1					
			Autoimmune Diseases						
CNTY-108	iNK/γδ iT	CD19	Autoimmune Diseases						
CLDE-308	αβ iΤ	CD19	Autoimmune Diseases						
CLDE-361	αβ iT	ВСМА	Myasthenia Gravis						
Hematologic a	nd Solid Tumors								
CNTY-101	iNK	CD19	B-Cell Malignancies	E	ELiPSE-1				
CNTY-102	γδ iT	CD19 + CD22	B-Cell Malignancies						
CLDE-308	αβ iΤ	CD19	B-Cell Malignancies						
CNTY-104	iNK/iT	Multi-specific	AML						ullı Bristol Myers Squibb
CNTY-106	iNK/iT	Multi-specific	ММ						ullı Bristol Myers Squibb
CNTY-107	γδ iT	Nectin-4	Solid Tumors						
Research	iT	Not disclosed	Solid Tumors						
Research	iNK/iT	TBD	Hematologic and Solid Tumors						

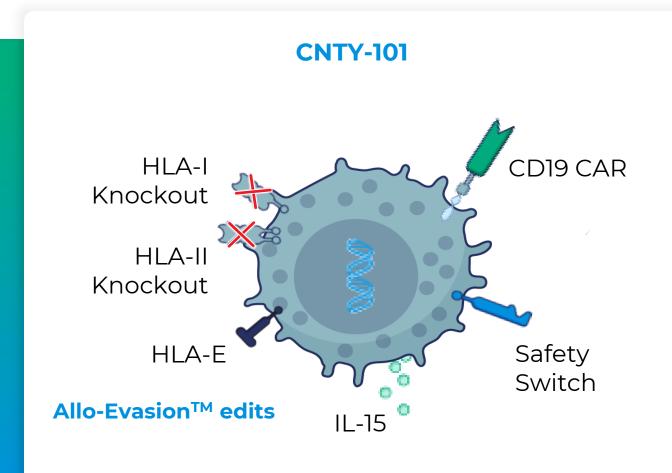




**CNTY-101 Clinical Programs** 

#### CNTY-101: Differentiated next-gen CD19 targeted product

Only cell therapy with six precision gene edits currently in the clinic



# Delivering on our vision to change the cell therapy treatment paradigm

- Goal to improve durability, tolerability and ease of outpatient administration
- Potential to eliminate need for lymphodepletion with subsequent cycles of therapy
- First CD19-targeted agent to test durability benefit of repeat dosing enabled by Allo-Evasion<sup>TM</sup> edits

## CNTY-101 in relapsed/refractory B-cell lymphomas

**Aim**: To deliver durable responses via repeat dosing facilitated by Allo-Evasion™ and extending the period of pharmacologic pressure on tumor cells





#### **Unmet need:**

- Autologous CD19 CAR-T is curative in ~40%¹ of patients
- Autologous CD19 CAR-T access is limited and/or can fail in manufacturing as quality is dependent on patientderived starting material
- Limited options and poor prognosis for patients who fail autologous CAR-T

#### **Potential solution from Century's platform:**

- Off-the-shelf product offers immediate access and consistency
- Multiple doses to increase pharmacological pressure to increase durability
- Host rejection addressed by Allo-Evasion<sup>™</sup> edits



## CNTY-101: ELiPSE-1 (NCT05336409) Phase 1 BOIN design

Patients with CD19+ aggressive and high-risk indolent R/R B-NHL

- DLBCL, HGBL, MCL, PMBCL, FL3B, FL, MZL
- ≥2 prior lines of therapy
- Prior CD19-targeted cell therapy allowed

- Part 1 Dose escalation
  - Schedule A: single dose
  - Schedule B: 1 dose per week x 3 weeks
- Part 2 Dose expansion

# Patient enrollment

LYMPHO-DEPLETION<sup>1</sup>

**Initial Dose** 

#### Schedule A

Dose level 1: 100 million Dose level 2: 300 million

Dose level 3: 1 billion
Dose level 4: 3 billion<sup>3</sup>

DAY 1

CNTY-101

IL-2 x 8 days<sup>3</sup>

#### Schedule B

Dose level 2: 300 million Dose level 3: 1 billion DAY 1 DAY 8 DAY 15

(NTY101)

(IL-2 x 4)

(days)

(IL-2 x 4)

(days)

(IL-2 x 4)

(days)

#### Additional Cycles<sup>2</sup>

First additional cycle: lymphodepletion at investigator's discretion

No lymphodepletion for following cycles

DAY1

CNTY-101

28-DAY DLT PERIOD RESPONSE ASSESSMENT

IL-2 x 8 days<sup>3</sup>

DAY 1 DAY 8 DAY 15

(CNTY-101) (CNTY-101) (CNTY-101)

(IL-2 x 4) (IL-2 x 4) (IL-2 x 4) (days)



ELiPSE-1
enrolled
heavily
pre-treated
R/R B-NHL
patients
across 7 sites

Baseline characteristics	N=12 safety evaluable <sup>1</sup>					
Median age (range, years)	70 (60-76)					
Male, n (%)	9 (75)					
NHL subtype, n (%)						
• DLBCL	7 (58)					
• HRFL	1 (8)					
• MCL	2 (17)					
• MZL	2 (17)					
Prior therapies, median (range)	4 (2-5)					
Response to last line of treatment, n (%)						
• Relapsed	3 (25)					
Refractory	9 (75)					
Received prior autologous* CAR-T, n (%)	3 (25)					
• If no, why						
– Manufacturing fail	1					
– Not eligible	3					
<ul> <li>Not willing to wait</li> </ul>	4 <sup>2</sup>					
<ul> <li>Financial or reimbursement constraints</li> </ul>	1					
*4 subjects received prior CAR T (3 autologous and 1 allogeneic)						

<sup>&</sup>lt;sup>1</sup> As of 27 March 2024 data cutoff, data collection ongoing

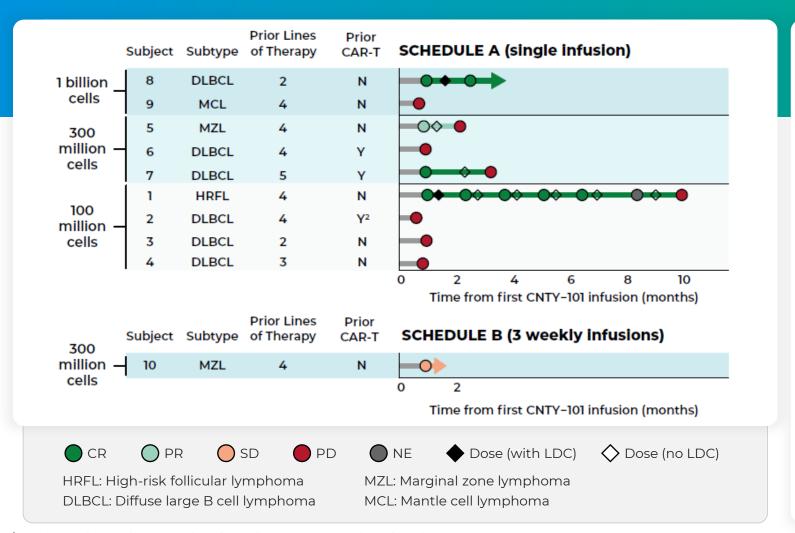


<sup>&</sup>lt;sup>2</sup> One subject received allogeneic CAR-T

HRFL: High-risk follicular lymphoma; DLBCL: Diffuse large B cell lymphoma; MZL: Marginal zone lymphoma; MCL: Mantle cell lymphoma

## CNTY-101 preliminary clinical data

Favorable safety profile and encouraging efficacy across initial dose levels studied



#### Efficacy (n=10)

- 30% CRR and 40% ORR across all dose levels and histologies
- 40% CRR and 60% ORR at highest studied dose levels in Schedule A

#### Safety & Tolerability (n=12)

- No treatment discontinuations due to AES; no GvHD
- CRS: Grade 1 (N=2), Grade 2 (N=2)
  - Hypotension (n=1) and hypoxia
     (n=1) lasted <24hrs</li>
- ICANS: Grade 1 (N=1), resolved in <24hrs</li>



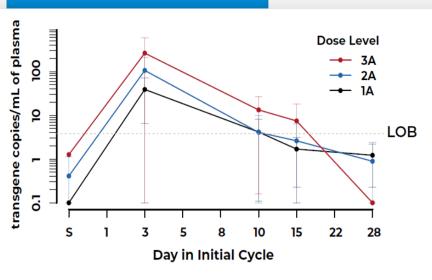
## CNTY-101 emerging pharmacokinetic profile

- Transient detection of CNTY-101 in circulation
- CNTY-101 persistence is detected via a novel cell-free (cf) DNA assay on Day 3 and beyond
- CNTY-101 cfDNA AUC trending to increase with dose
- 3/4 pts who received an additional CNTY-101 cycle without LD had CNTY-101 cfDNA detected at Day 3+

# PBMC genomic DNA Dose Level 3A 2A -0 1A

Day in Initial Cycle





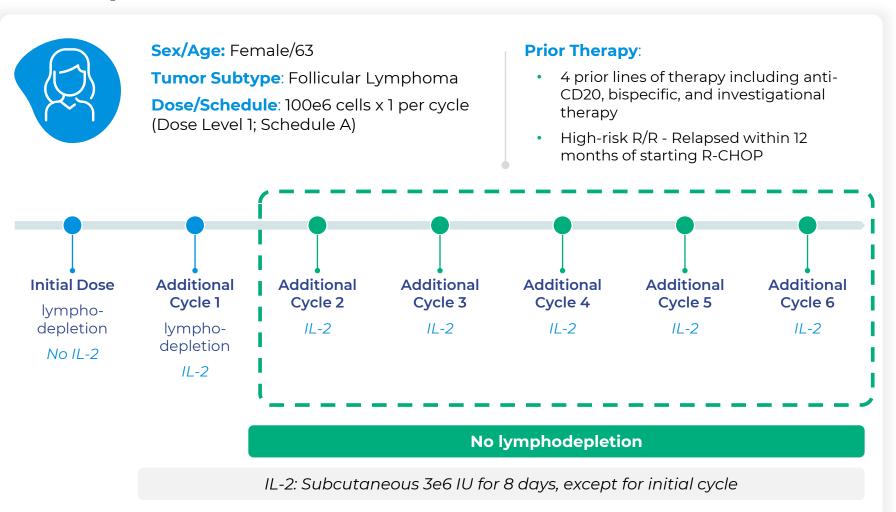
Data is shown as mean ± SD for the initial cycle across subjects at each dose level in Schedule A as of May 1st, 2024 data cutoff date.

# ASH 2023 case study: Dose level 1 patient with 6-month durable complete response

Multiple Doses of CNTY-101, an iPSC-Derived Allogeneic CD19 Targeting CAR-NK Product, are Safe and Result in Tumor Microenvironment Changes Associated with Response: A Case Study

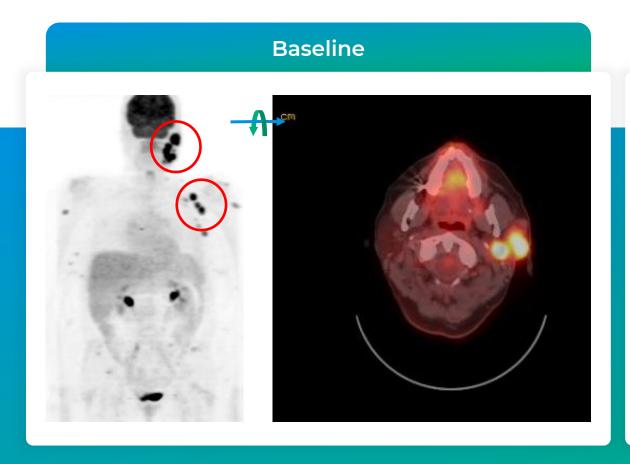
Indu Ramachandran<sup>1</sup>, Sarah Rothman<sup>1</sup>, Mariano Clausi<sup>1</sup>, Kile McFadden<sup>1</sup>, Brenda Salantes<sup>1</sup>, Gloria Jih<sup>1</sup>, Thomas Brigman<sup>1</sup>, Sam Kelly<sup>1</sup>, Matthew S. Hall<sup>1</sup>, Stephanie Yee<sup>1</sup>, Iphigenia Koumenis<sup>1</sup>, Poulomee Das<sup>1</sup>, Jordan Briggs<sup>2</sup>, Tori Braun<sup>2</sup>, Ying Yuan<sup>3</sup>, Elizabeth Devlin<sup>1</sup>, Adrienne Farid<sup>1</sup>, Nikolaus Trede<sup>1</sup>, Tamara K. Moyo<sup>5</sup>, Tahir Latif<sup>4</sup>, Krish Patel<sup>2</sup>

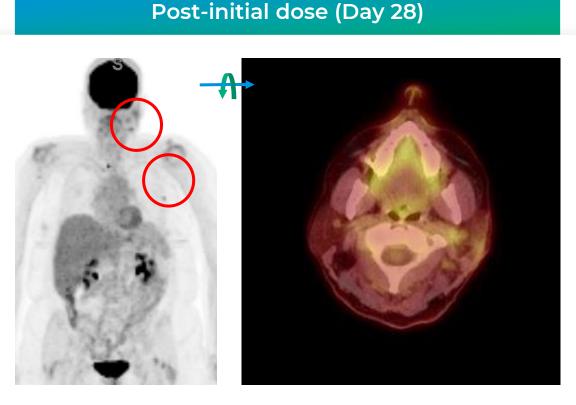
<sup>1</sup>Century Therapeutics, Philadelphia, PA <sup>2</sup>Swedish Cancer Institute, Seattle, WA <sup>3</sup>MD Anderson Cancer Center, Houston, TX <sup>4</sup>Atrium Health Levine Cancer Institute, Charlotte, NC <sup>5</sup>University of Cincinnati Medical Center, Cincinnati, OH



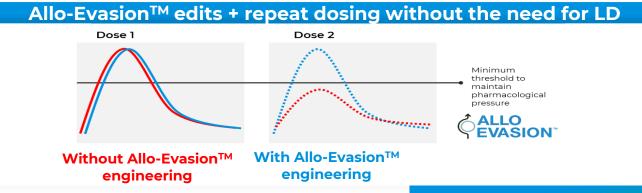


# ASH 2023 case study: Early evidence of anti-lymphoma activity with durable 6-month complete response<sup>^</sup>





# Allo-Evasion<sup>TM</sup> enables repeat dosing without the need for continued lymphodepletion *Initial clinical evidence indicates no sign of allo-rejection for CNTY-101 (ASH case study)*



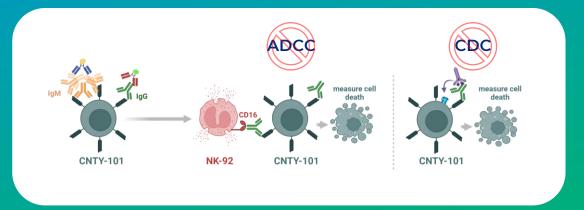
Allo-Evasion<sup>™</sup> provides potential to more tightly control drug exposure to enable sustained pressure on the target

**ELiPSE-1 Clinical Data** 

CNTY-101 cells persist in tissues for at least 3 days as measured by cfDNA; observed with and without LD



Anti-drug antibodies and functional humoral immune response against CNTY-101 are not detected (seven cycles evaluated)





## **ELiPSE-1 initial data: Key takeaways**



Heavily pretreated and refractory patient population treated in first-in-human dose escalation trial



Favorable safety profile; can be delivered in an outpatient setting



Encouraging early efficacy signals at initial 3 dose levels in Schedule A



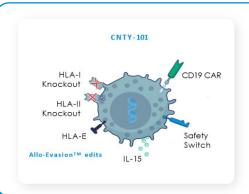
Novel cfDNA assay enables monitoring of CNTY-101 persistence in extravascular space; AUC increase trending with dose



Initial data for CNTY-101 supports the potential for Allo-Evasion™ to enable a multi-dosing regimen without the need for continued lymphodepletion

CNTY-101's manageable initial safety profile, initial response data, and PK/PD supports advancing to higher doses to potentially deepen and prolong clinical response

#### Key differentiators of CNTY-101 in autoimmune disease treatment



# CNTY-101: CD-19 targeted iNK cell therapy with 6 precision gene edits including Allo-Evasion<sup>TM</sup> technology

- Ph1 CALiPSO-1 trial in B cell-mediated autoimmune diseases (Systemic Lupus Erythematosus and Lupus Nephritis) initiated in early 3Q24
- Currently being studied in Ph1 ELiPSE-1 trial in R/R NHL

Key differentiators in AID: (1) Allogeneic (2) NK cells (3) Allo-Evasion™

#### Allogeneic

- Available "off-the-shelf"
- No patient apheresis required
- No manufacturing wait time
- Platform enables lower COGs than donor-derived or autologous

#### **NK cells**

- Killing potency ≥ primary CAR-T
- Trafficking to secondary lymphoid tissues and marrow favors pathogenic B-cell targeting
- Limited *in vivo* expansion

#### Allo-Evasion™

- Avoiding host immune rejection
- Ability to repeat dose without continued lymphodepletion
- Ability to retreat, if needed

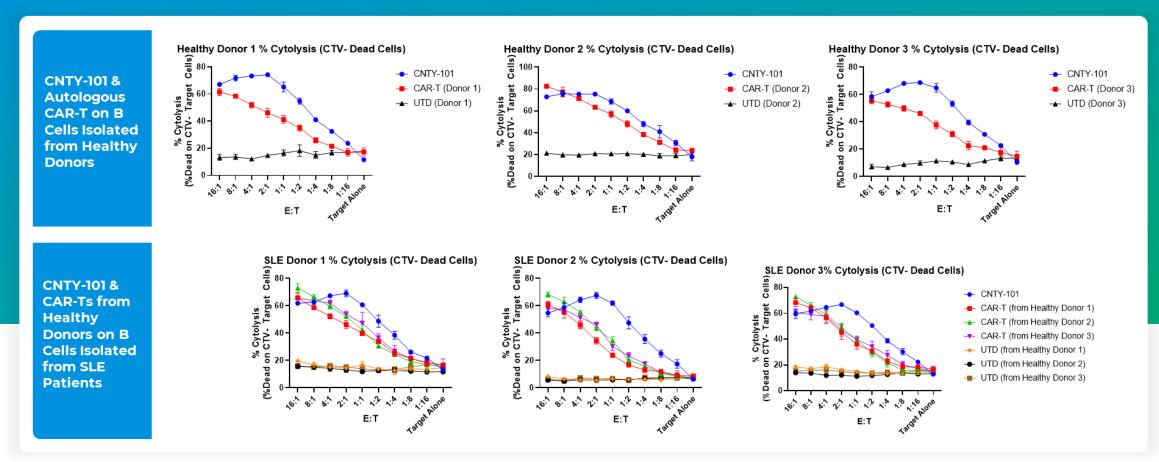
Tighter control over drug exposure:

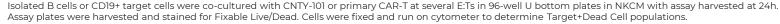
B-cell depletion without prolonged B-cell aplasia

#### CNTY-101: Potential to drive B-cell depletion with tighter control over drug exposure

More potent than primary CAR-T at B-cell killing in preclinical comparison

# CNTY-101 more potent than primary CAR-T cells at B-cell killing at 24 hours in preclinical comparison







# Opportunity in systemic lupus erythematosus and lupus nephritis to improve long-term disease control







## Estimated global prevalence of 3.4 million patients<sup>1</sup>

- Abnormal B cell function and autoantibody production are central to disease pathogenesis
- Major causes of morbidity and mortality involve multiple systems
  - Renal, CNS and cardiovascular involvement are major causes of morbidity and mortality

# Despite approved treatments, significant unmet need remains

- Chronic treatment with broadacting anti-inflammatory and immunosuppressives
- Current treatments fail to significantly impact morbidity in the moderate to severe population
- Treatment toxicity and disease flares remain common

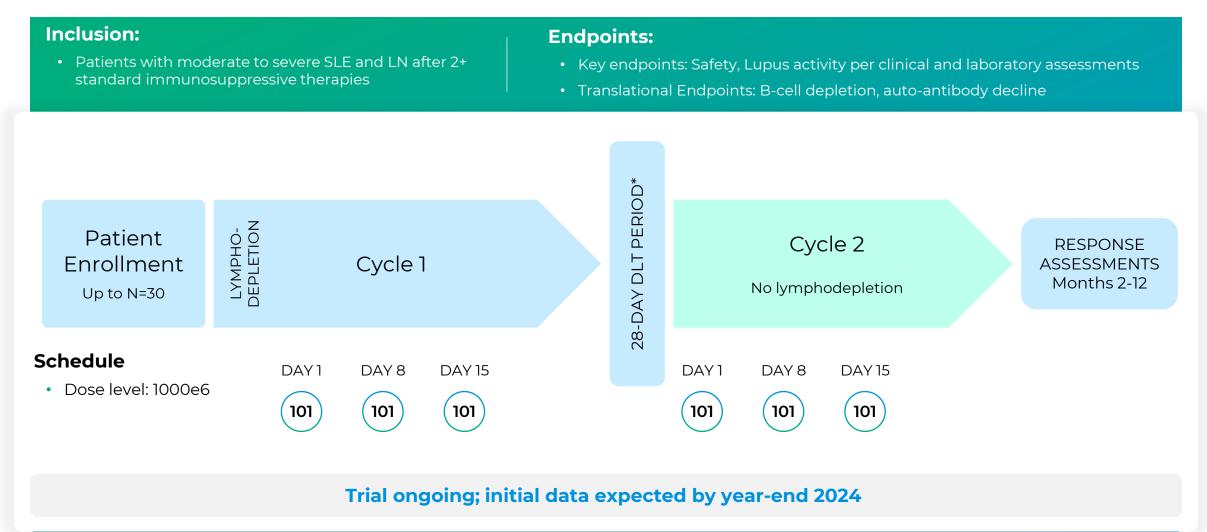
Autologous anti-CD19 CAR T cell therapies have established a promising efficacy proof of concept in SLE/LN

Challenges remain due to potential exposure to CRS and ICANS, product availability, and long-term risks including B-cell aplasia

<sup>1.</sup> Tian J, et al. Ann Rheum Dis 2023;82:351–356 http://dx.doi.org/10.1136/ard-2022-223035

Mackensen A, et al. Nature Medicine 2022 28:10 (2124-2132) https://doi.org/10.1038/s41591-022-02017-5
 Muller, F et al NEJM 2024 390:687 https://www.nejm.org/doi/full/10.1056/NEJMoa2308917
 CNS: Central Nervous System

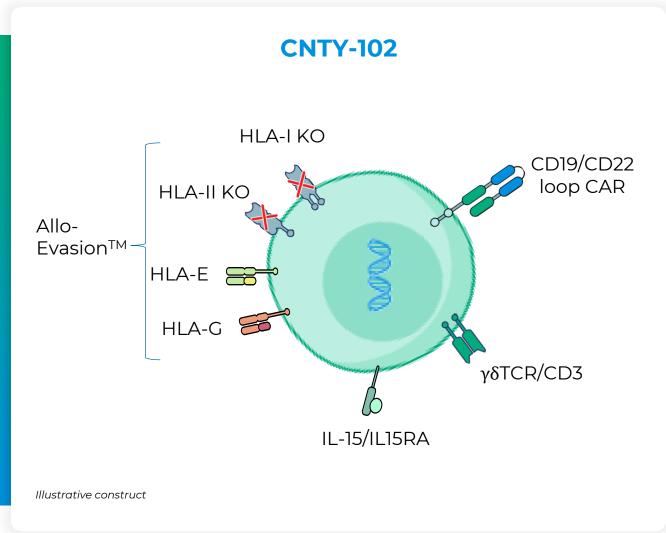
# CNTY-101: CALiPSO-1 (NCT06255028) B cell-mediated autoimmune diseases Phase 1 study





**Discovery Programs** 

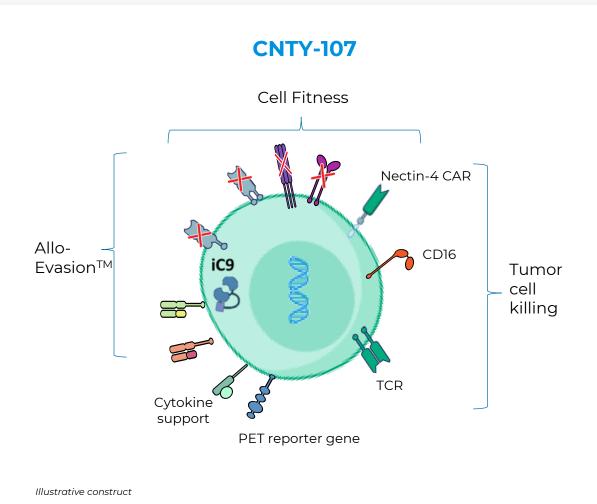
# CNTY-102: Leveraging the next generation $\gamma\delta$ iT cell platform designed to deliver best-in-class potential



# Designed to address factors that limit durability of cell therapy in B-cell malignancies

- γδiT cells have distinct properties that provide optionality in the face of different biological challenges
- Dual targeting designed to counter antigen escape relapse - a major limiting factor for durability of CD19 CAR T therapies
- Armed with Allo-Evasion<sup>™</sup> edits to enable repeat dosing to potentially deliver durable responses

## CNTY-107: First in class Nectin-4 targeted $\gamma\delta$ iT cell therapy



# Leveraging the power of the $\gamma\delta$ iT cell platform for solid tumors

#### **Nectin-4 has been validated by ADC approaches**

- Opportunity to address multiple Nectin-4 positive solid tumors
  - Potential indications include bladder, breast, pancreatic, non-small cell lung cancer, esophageal/gastric, head and neck, and/or ovarian cancers<sup>1</sup>

# $\gamma \delta$ iT allogeneic therapies provide potential to improve upon ADC toxicity profile and efficacy

- Intrinsic homing of  $\gamma\delta$  iT cells to tissues and solid malignancies
- Multi-tumor killing modalities to tackle heterogeneity



# Corporate Position & Upcoming Milestones

# Advancing next-generation iPSC-derived allogeneic NK and T cell therapy candidates for the treatment of cancer and autoimmunity

# Differentiated pipeline based on Allo-Evasion™ technology

✓ Potential to overcome limitations of conventional allogeneic cell therapy

# Encouraging preliminary clinical data from Phase 1 trial of CNTY-101 in R/R B-cell lymphomas

- ✓ Well-tolerated with early evidence of anti-lymphoma activity, and supports the ability to re-dose without lymphodepletion
- ✓ Additional data from EliPSE-1 announced, completed dose escalation

#### **Expansion into additional autoimmune indications**

- CALiPSO-1 trial initiated; amended to include additional cohort of LN patients
- ✓ CNTY-101 has differentiated profile in AID (allogeneic, iNK with Allo-Evasion<sup>™</sup>)
- Clade Therapeutics acquisition further expands and enhances autoimmune opportunities and platform technology

#### In-house manufacturing capabilities

✓ Ability to accelerate learnings and enable faster product iteration

#### **MULTIPLE NEAR-TERM CATALYSTS**

#### Phase 1 ELiPSE-1 trial of CNTY-101 in B-cell malignancies

Progressing into dose expansion in 2H 2024

## Phase 1 CALiPSO-1 trial of CNTY-101 in B-cell mediated autoimmune diseases

Initial clinical data expected by YE 2024

Pursuing additional autoimmune regulatory filings for CNTY-101 in 2H 2024

#### **CASH RESOURCES**

#### Cash runway into 2026

Ended 2Q24 with cash, cash equivalents, and investments of \$269.6M



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**ENDURING IMPACT...** 

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Well-capitalized into 2026 to enable delivery on key milestones and clinical data