### UNITED STATES SECURITIES AND EXCHANGE COMMISSION

WASHINGTON, D.C. 20549

### FORM 8-K

### **CURRENT REPORT**

Pursuant to Section 13 or 15(d) of the Securities Exchange Act of 1934

Date of Report (Date of earliest event reported): November 10, 2022

### Adicet Bio, Inc.

(Exact name of Registrant as Specified in Its Charter)

**Delaware** (State or Other Jurisdiction of Incorporation) 001-38359 (Commission File Number)

81-3305277 (IRS Employer Identification No.)

200 Berkeley Street, 19th Floor Boston, Massachusetts (Address of Principal Executive Offices)

02116 (Zip Code)

Registrant's Telephone Number, Including Area Code: (650) 503-9095

Not applicable (Former Name or Former Address, if Changed Since Last Report)

	(rotinet Name of Former Address, it Changed Since Last Report)						
Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions:							
Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)							
Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)							
☐ Pre-commencement communications	Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))						
☐ Pre-commencement communications	Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))						
Securities registered pursuant to Section 12(b) of the Act:							
Title of each clas	s	Trading Symbol(s)	Name of each exchange on which registered				
Common Stock, par value \$0.0	0001 per share	ACET	The Nasdaq Global Market				
Indicate by check mark whether the registrant is an emerging growth company as defined in Rule 405 of the Securities Act of 1933 (§ 230.405 of this chapter) or Rule 12b-2 of the Securities Exchange Act of 1934 (§ 240.12b-2 of this chapter).							
Emerging growth company ⊠							
If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act.							

#### Item 7.01 Regulation FD Disclosure.

On November 10, 2022, Adicet Bio, Inc. ("Adicet") held its Virtual Research and Development ("R&D") Day. In connection with the R&D Day, Adicet has made its presentation from the R&D Day available on its website at investor.adicetbio.com/events-and-presentations/events. A copy of the presentation is also furnished herewith as Exhibit 99.1 to this Current Report on Form 8-K.

The information in this Item 7.01, including Exhibit 99.1 attached hereto, is intended to be furnished and shall not be deemed "filed" for purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the "Exchange Act"), or otherwise subject to the liabilities of that section, nor shall it be deemed incorporated by reference in any filing under the Securities Act of 1933, as amended, or the Exchange Act, except as expressly set forth by specific reference in such filing.

#### Item 8.01 Other Events.

Adicet's ongoing research efforts are focused on "off-the-shelf" gamma delta T cells, engineered with chimeric antigen receptors (CARs) and adaptors (CAds), to enhance selective tumor targeting and facilitate innate and adaptive anti-tumor immune response for durable activity in patients.

On November 10, 2022, Adicet provided the below updates regarding its product candidates and R&D programs:

- Adicet's pipeline is led by ADI-001, a first-in-class allogeneic gamma delta T cell therapy expressing a CAR targeting CD20, is in an ongoing Phase 1 study for the treatment of relapsed or refractory B-cell non-Hodgkin's lymphoma (NHL).
- Adicet's pipeline also includes ADI-925, ADI-925 is an engineered Chimeric Adapter (CAd) γδ1 T cell product candidate targeting stress ligands, including MICA/MICB & ULBP1-6, expressed on malignant cells.
- Adicet announced its newly created approach for targeting CD70 and noted preliminary preclinical data supporting the target in both AML and RCC.
- Adicet announced its program targeting PSMA and noted that its lead PSMA CAR construct demonstrated improved cytotoxicity and targeting compared to a reference benchmark.
- · Adicet announced an early program targeting B7-H6, a member of the B7 family that is well characterized for its role in innate immunosurveillance of tumors.

#### Item 9.01 Exhibits.

(d) Exhibits

#### Exhibit No. Description

99.1 <u>R&D Da</u>

R&D Day Presentation by Adicet Bio, Inc. on November 10, 2022, furnished herewith.

104 Cover Page Interactive Data File (embedded within the Inline XBRL document)

#### **SIGNATURES**

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

### ADICET BIO, INC.

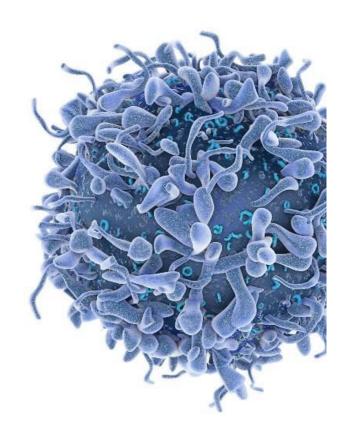
By: Name: Title: Date: November 10, 2022

/s/ Nick Harvey Nick Harvey Chief Financial Officer



# Virtual R&D Event

November 10, 2022



### Forward-Looking Statements

This presentation contains "forward-looking statements" of Adicet Bio, Inc. (Adicet) within the meaning of the Private Securities Litigation Reform Act of 1995 relating to business and operations of Adicet. These forward-looking statements include, but are not limited to, express or implied statements regarding the potential safety, durability, and therapeutic effects of ADI-001 and preclinical programs, including the implementation, timing and success of ADI-001, ADI-925 and the additional preclinical programs in Adicet's pipeline; plans and timing for the release of additional clinical data from Adicet's Phase 1 trial of ADI-001 in relapsed/refractory NHL patients; future progress of the Phase 1 trial of ADI-001, including ongoing patient enrollment; expectations regarding future regulatory filings for product candidates in the Company's pipeline, including the planned IND submission for ADI-925 in the second half of 2023; timing of a dose selection for the Phase 2 trial by the end of 2022 and initiation of a potentially pivotal program in the first half of 2023, expectations for the in-house manufacturing capabilities in Adicet's Redwood City facility, and Adicet's growth as a company and expectations regarding the advancement of its product candidates and expected cash runway. Any forward-looking statements in this presentation are based on management's current expectations and beliefs of future events, and are subject to a number of risks and uncertainties that could cause actual results to differ materially and adversely from those set forth in or implied by such forward-looking statements, including without limitation, the effect of COVID-19 on Adicet's business and financial results, including with respect to disruptions to its clinical trials, business operations, and ability to raise additional capital; Adicet's ability to execute on its strategy, including obtaining the requisite regulatory approvals on the expected timeline if at all; that positive results, including interim results, from a clinical

#### Industry and Market Information

Information regarding market share, market position and industry data pertaining to Adicet's business contained in this presentation consists of estimates based on data and reports compiled by industry professional organizations and analysts and Adicet's knowledge of their industry. Although Adicet believes the industry and market data to be reliable, this information could prove to be inaccurate. You should carefully consider the inherent risks and uncertainties associated with the market and other industry data contained in this presentation. Forward-looking information obtained from third-party sources is subject to the same qualifications and the additional uncertainties as the other forward-looking statements in this presentation.



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# Agenda

Introduction	Chen Schor		
Preclinical Program Overview	Blake Aftab, Ph.D.		
Perspectives	Marco Davila, M.D., Ph.D.		
Closing Remarks	Chen Schor		
Q&A	Chen Schor Blake Aftab, Ph.D. Marco Davila, M.D., Ph.D. Francesco Galimi, M.D., Ph.D. Nick Harvey		



Chen Schor President and CEO



Blake Aftab, Ph.D. Chief Scientific Officer



Marco Davila, M.D., Ph.D. Roswell Park Comprehensive Cancer Center



Francesco Galimi, Mick Harvey M.D., Ph.D. Chief Financial Officer Chief Financial Officer





Leaders in Developing Allogeneic CAR and CAd γδ T Cell Therapies to Fight Cancer

Chen Schor, President and CEO

# Adicet Highlights: Leading The Way With γδ T Cell Therapies To Fight Cancer

#### 2015

 Adicet Bio Founded

Dec 2021



50% CR rate

with ADI-001 in

aggressive NHL

#### June 2022

- 75% CR rate, encouraging preliminary durability, no significant CRS, ICANS or Gr3+ infection rate in
  - Potential for outpatient dosing

aggressive NHL



### 2015 - 2021

- Developed fundamental understanding of γδ T cell biology
- Optimized robust manufacturing process for allogeneic off-the-shelf γδ1 T cell therapy
- · Built foundational IP portfolio

#### November 2022

- Four new highly differentiated γδ T cell therapy pipeline programs
- Building on years of expertise, IP and know-how in γδ T cell biology



#### Now & Future





- Initiate at least one potentially pivotal program in H1/2023
- IND submission for ADI-925 expected in H2/2023

CR: Complete response; CRS: Cytokine release syndrome; y6: Garmia delta; Gr3: Grade tirree; IDANS; Immune affector call-associated neurotoxicity syndrome: IP: Intellactual property: NHL: Non-Hodoxin's.



# Adicet CAR $\gamma\delta$ T Cell Platform Potential Advantages: Engineered to Address Activity, Tumor Homing, Safety, and COGs Limitations

		CAR yo T Cells	Key Supporting Data	
Adaptive a  *Active tum  Predomina  Preclinical	Innate anti-tumor response	~	PRE-CLINICAL:	
	Adaptive anti-tumor response	./	Nishimoto et. al. Clinical & Translational Immunology 2022; Makkouk et. al. JITC 2021; Azameera et. al. ISCT 2022	
	Adaptive anti-tumor response	<b>V</b>	Single dose protects from repeat tumor challenge (Romero et al. ASGCT 2019)	
	Active tumor homing	<b>/</b>	<ol> <li>Gamma delta 1 CAR T cells expansion capacity is better than CAR NK cells and comparable or better then alpha-beta CAR T cells (Nishimoto et al)</li> </ol>	
	Dradominantly activating recentor averaging	~	<ul> <li>4) Predominantly activating receptors (Nishimoto, Makkouk, and Azameera et. al. publications)</li> </ul>	
	Predominantly activating receptor expression		CLINICAL:	
	Preclinical persistence by repeat tumor challenge	<b>/</b>	<ol> <li>CRs demonstrated with ADI001 starting at 30M CAR+ cells (flat dose) in bulks tumors &gt; 6,000 mm (ASCO 2022 presentation)</li> </ol>	
		~	<ul> <li>2) Active dose of ADI-001 ~ 1% of NK total dose per lympho-depletion cycle (ASCO 2022 presentation)</li> </ul>	
	Prognostic value of tumor infiltration		3) Gentles et. Al. Nat Med. 2015	
Safety*	Low GvHD risk			
			CLINICAL:	
	Low risk of cytokine release syndrome ≥ grade 3 risk	<b>/</b>	No GvHD and no ≥ grade 3 GvHD cases with ADI-001 (ASCO 2022)	
SDOO	No gene editing required (may affect efficacy)	1	PRE-CLINICAL:	
	The game canning requires (may amost emeasy)		- (1) No gene editing with ADI-001	
	Scalable manufacturing	<b>/</b>	(2) Manufacturing for pivotal and commercial with CRO	



### Adicet's Pipeline Strategy

- · Unmet medical need
- · Current standard of care
- · Competitive landscape

γδ T cells innate and adaptive anti-tumor activity

Engineer and enhance targeting with CAR or CAd Follow γδ1 T cell tissue residence and infiltration in solid tumors

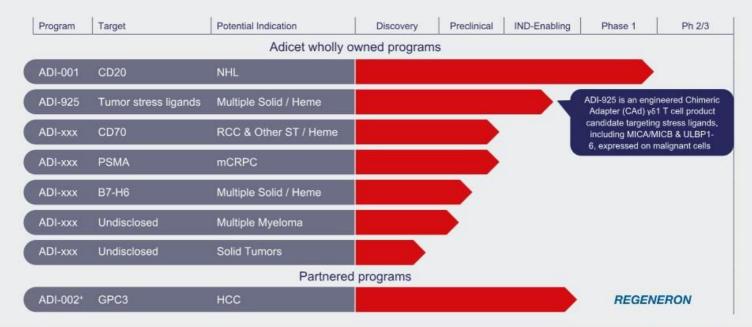
Armor to preserve activity in immunosuppressive tumor microenvironment



- 1) Hematologic malignancies where Adicet established clinical POC, or
- 2) Indications where infiltrating γδ T cells correlated with OS, or
- 3) Indications where donor lymphocyte infusions, enriched with  $\gamma\delta$  T cells, have shown clinical benefit

Opportunity for highly differentiated clinical benefits

# Building a Broad Pipeline of First-in-Class CAR and CAd $\gamma\delta$ T Cell Product Candidates



<sup>\*</sup>Rependent exercised its notion to license the exclusive worldwide rights to ADL002



HCC: Hepstocellular carcinoms; mCRPC: Metastatic castration-resistant prostate cancer; MICA/MICE; Major histocompatibility complex (MHC) Class i chain-related protein A/B; NH Non-Hoddsir/s lumphorum; PSMA: Prestate-search; membrans artistem; RCC: Renat call carcinoma. ST: Solid tumpo; ULBP: ULB fabriding protein

# ADI-925: First-in-Class Enhanced $\gamma\delta$ 1 T Cell Product Candidate for Multiple Heme and Solid Tumors

We engineered our  $\gamma\delta$  1 T cells to significantly enhance their intrinsic innate and adaptive anti-tumor activity by rewiring signaling pathways in the cell

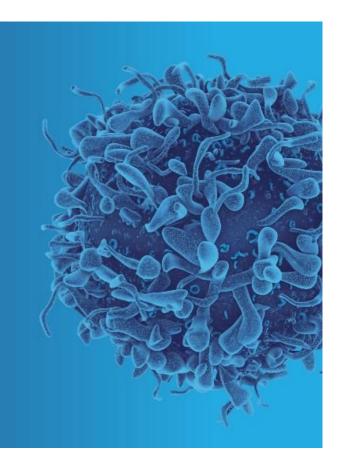
This means that one cell therapy product may demonstrate anti-tumor activity in multiple hematologic malignancies and solid tumors without a CAR.

ADI-925 may provide these benefits to cancer patients



# Preclinical Pipeline Overview

Blake Aftab, Ph.D. Chief Scientific Officer



# Building a Broad Pipeline of First-in-Class CAR and CAd $\gamma\delta$ T Cell Product Candidates

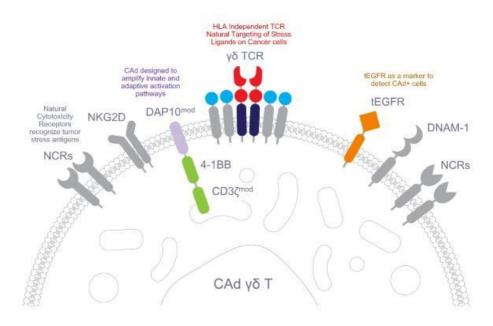


Adicet Bio

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### ADI-925: Engineered γδ1 Chimeric Adaptor T Cell Product Candidate

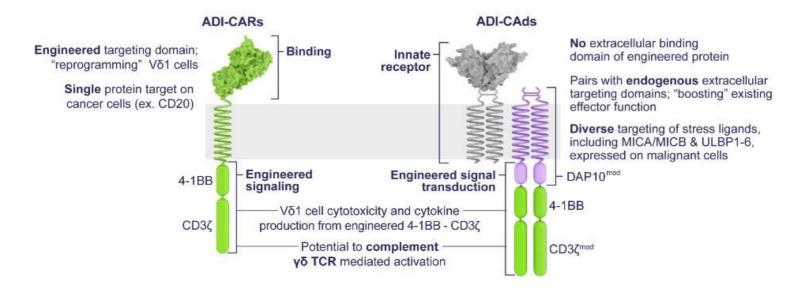
- ADI-925 is designed to enhance the innate and adaptive anti-tumor activity of Vδ1 T cell
- ADI-925 is an engineered Chimeric Adapter (CAd) Vδ1 T cell therapy candidate targeting stress ligands, including MICA/MICB & ULBP1-6 expressed on malignant cells
- ADI-925 has demonstrated increased anti-tumor activity at lower concentrations of Vδ1 T cells
- Developed in-house with broad IP on file



DAP10: Hematopoletic cell signal transducer, DNAM: DNAX accessory molecule; MIC: Major histocompatibility complex class I chain-related protein mod: Modified domain[s]; NCR: natural cytotoxicity receptor; NKG2D: Killer cell lectin like Receptor K1; TCR: T cell receptor; EGFR: Truncated epidermal growth factor receptor.



### First-in-class CAd Enhancement of Intrinsic $\gamma\delta$ T Cell Activity

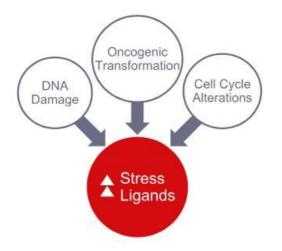


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ADI: Adicet Bio, ULBP: UL16-binding protein

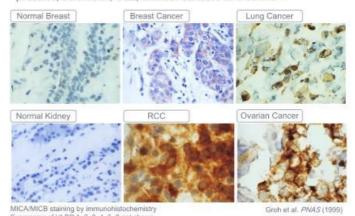
# Tumor Stress Ligands Targeted by ADI-925 Are Ubiquitously Expressed Across a Broad Range of Indications

Expression of stress ligands across indications is coupled to drivers of tumor formation<sup>1</sup>



# Expression of stress ligands present in broad range of primary tumor specimens

AML, MM, melanoma, MCC, TNBC, HCC, Cervical, NSCLC, RCC, prostate, colorectal, CLL, bladder cancers and others<sup>2,3</sup>



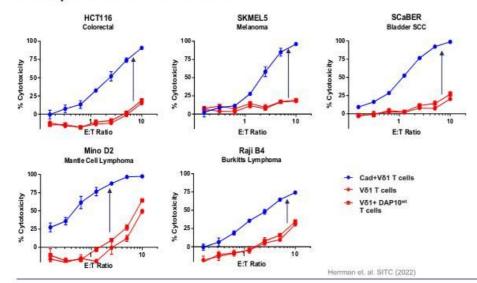
Adicet Bio

Jones et al. Cancers (2022)
 Zhao et al. Oncofarger (2017)
 Groh et al. PNAS (1999)

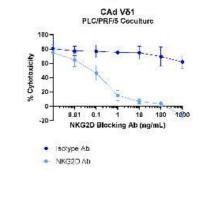
# ADI-925 Targets a Broad Panel of Malignancies



# ADI-925 effectively enhanced cytotoxic potency of $\gamma\delta$ T cells in broad panel of cancer cell lines



# Enhancement was primarily driven via endogenous NKG2D



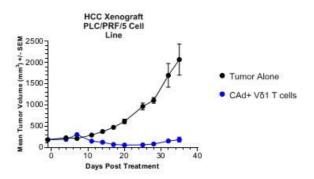
# ADI-925 Demonstrated Enhanced Potency of $\gamma\delta$ T Cells



### Potent Killing with CAd Engineered γδ T Cells

# HCC Cell Line HepG2 Tumor Alone Tumor Alone Positive Control-CAR Võ1 T cells CAd+ Võ1 T cells Co-culture time (hrs)

### Potent Activity in Solid Tumor Models



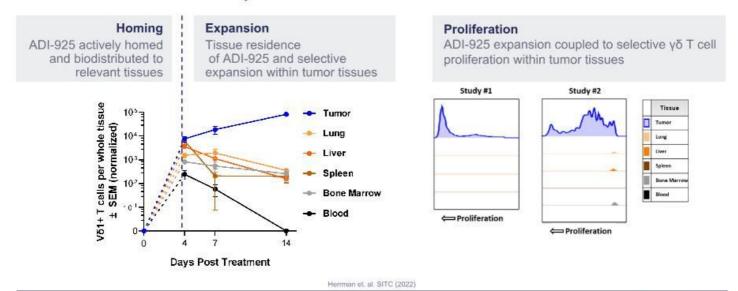
Herman et. al. SITC (2022



# ADI-925 Illustrated Preferential Homing and Expansion Within Tumors



### **Hepatocellular Carcinoma Tumor Model**



# ADI-925: Opportunity For Differentiation

### **Target validation**

- Presence of γδ T cells in tumors correlates with OS3,4,5,6
- · Many stress antigens selected by evolution to mark malignant cells
- Unmodified allogeneic γδ T cell therapy shows encouraging clinical signal in AML1
- · Orthogonal NKG2D CARs have demonstrated clinical POC7

### Key challenges

- · Potency of non-engineered cell monotherapy may be limited
- · Lack of approaches to enhance intrinsic γδ T cell activity beyond that of correlation
- · Solid tumors may require engineered effector targeting coupled to tumor and tissue specific homing

### Opportunity for ADI-925 to address broad landscape

- · Enhanced natural cytotoxic effector function
- · Targeting multiple stress antigens addressing tumor heterogeneity
- · Broad, clinically relevant homing in solid tumors
- · Prominent cell expansion capacity within tumor

IND filing expected H2/2023

7. NCT04823944; Ph1 update

OS: Overall survival



# Building a Broad Pipeline of First-in-Class CAR and CAd $\gamma\delta$ T Cell Product Candidates



ADI-925 is an engineered Chimeric Adapter (CAd) y61 T cell product candidate targeting stress ligands, including MICA/MICB & ULBP1-6, expressed on malignant cell



### Armored CD70 CAR γδ T Cell Opportunity For Differentiation

### Target validation

- · CD70 expression is present in majority of patients with RCC (80%)1 & AML (>96%)2
- · Including, expression on both leukemic blasts and leukemic stem
- · Preliminary clinical validation of target in both AML and RCC:
- · Clinical activity observed in AML with CD70-targeted mAb4.5
- · Single-digit OR and double-digit SD rates with ADCs in RCC (& AML), limited by payload-driven toxicities<sup>6,7,8</sup>
- · Disease control seen with unarmored allogeneic αβ T-cell therapy (incl. one CR in advanced RCC patient)8

### Key challenges

- Modest responses rates with CD70-targeted agents to-date
- · Agents with limited mechanisms of action do not address tumor heterogeneity
- · No tissue-specific mechanisms for tropism with any agents (ADCs, mAbs, αβ T-cell therapy)
- · Payload-driven toxicities with **ADCs**
- · Immunosuppressive environment of RCC and other solid tumors

### Opportunity for Adicet and yo T cells

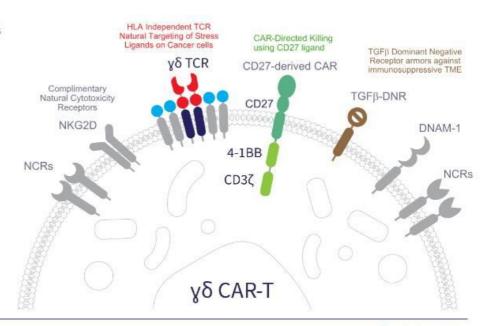
- · Response to low antigen density by design with CD27based CAR (compared to scFv-based CAR)3
- · Three mechanisms of action designed to address tumor heterogeneity
- Homing of γδ T cells reported in RCC
- · Inclusion of armoring to address suppressive TME

Massard et al. Cancer Chemother Pharmacol (2019)
 R. CRISPR Therapeutics Presentation (2022)



### Adicet's Armored CD70 CAR $\gamma\delta$ T Cell

- Targeting CD70 via CD27-ligand has shown superiority to scFv CARs<sup>1</sup>
- Innate and adaptive targeting mechanisms associated with activity in AML and RCC indications
- Armoring via dominant negative receptor; addresses TGFβ in TME<sup>2</sup>
- Lead CAR demonstrated potency and improved serial killing, and resilience against suppressive factors
- Supports functional enhancement illustrated in preclinical models



Sauer et al. Blood (2021)
 Junker et al. Cylokine (2000)

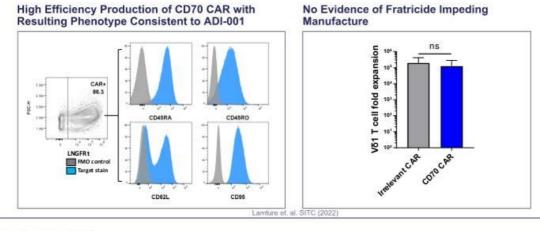


### Armored CD70 CAR $\gamma\delta$ T Cells Generated with High Efficiency





- · CD70 CAR γδ T cells produced with high transduction efficiency
- Phenotype similar to that of Adicet's established ADI-001 process<sup>1</sup>
- Production of CD70 CAR γδ T cells does not indicate fratricide



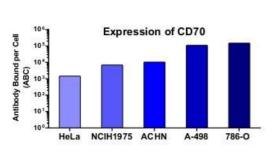
## CD70 CAR γδ T Cells Retained Cytotoxicity Across Range PRESENTED AT SITC 2022 of Target Expression



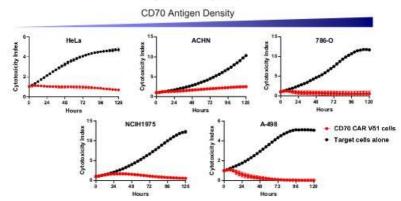


- · CD27-derived targeting of CD70 resulted in activation and anti-tumor activity across range of target antigen densities
- Target recognition applicable across range of indications and cell lines
  - Lung Adenocarcinoma, Renal Cell Carcinoma (x3), and Cervical Carcinoma shown

### Anti-Tumor Cytotoxicity at Low E:T Ratio



Lamture et. al. SITC (2022)





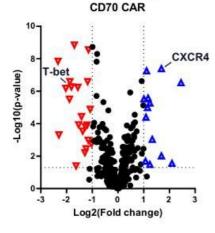
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### TGFβ Plays a Key Immunosuppressive Factor in TME









What is the purpose for armoring cells?

- High levels of TGFβ in tumor microenvironment resulted in differentially downregulated (red) or upregulated (blue) genes
- Alterations propagate immunosuppressive effects<sup>1</sup> and may blunt efficacy of immunotherapies<sup>2</sup> and CAR T<sup>3</sup>
- Armoring aims to preserve T cell function in the presence of immunosuppressive factors like TGFB

▲ Genes up-regulated in TGF-β1 treated conditions

Genes down-regulated in TGF-β1 treated conditions

Lamture et. al. SITC (2022)

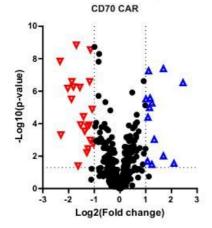
Batille & Massague. *Immunity (2*019)
 Silk et al. *Immunology* (2022)
 Zhang et al. *Front Cell Dev Biol* (2021)



# Armoring CD70 CAR $\gamma\delta$ T Cell Improved Resilience







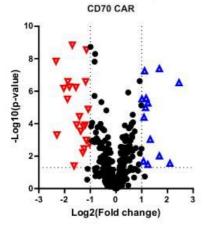
Add TGFβ-DNR Armor

Lamture et. al. SITC (2022

# Armoring CD70 CAR γδ T Cell Improved Resilience

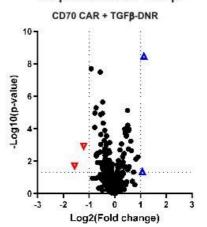








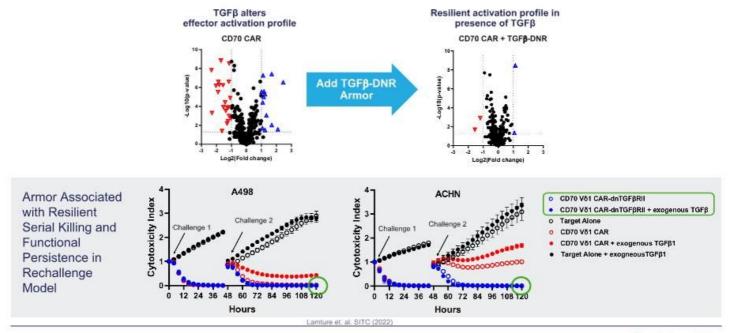
# Resilient activation profile in presence of TGFβ



Lamture et. al. SITC (2022)

# Armoring CD70 CAR γδ T Cell Improved Resilience





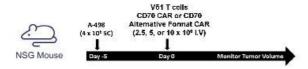
Adicet Bio

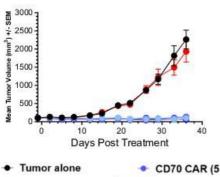
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# CD70 CAR $\gamma\delta$ T Cells Demonstrated Activity In Vivo and Proliferated Within Tumors





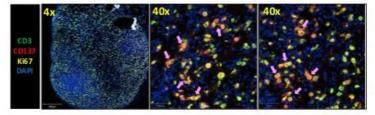




Tumor alone
 CD70 CAR (5 x 10<sup>6</sup>)
 CD70 CAR (10 x 10<sup>6</sup>)
 CD70 CAR (10 x 10<sup>6</sup>)

CD70 CAR Alternative format (5 x 10<sup>6</sup>)

Tumor Infiltration and Proliferation of γδ CAR T cells



- CD70 CAR γδ T cells demonstrated robust tumor growth inhibition
- Anti-tumor activity associated with CAR γδ T cell tumor infiltration and proliferation within the tumor bulk

Lamture et. al. SITC (2022)



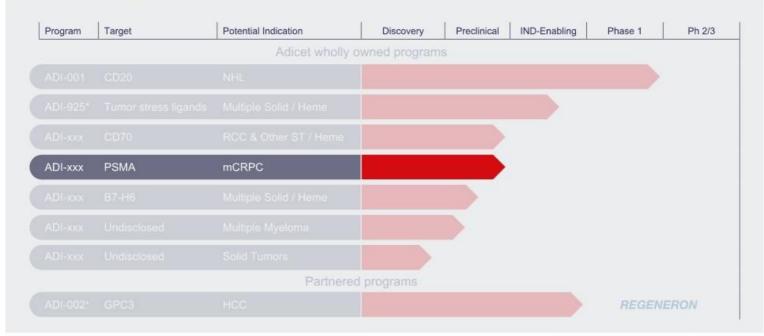
### Armored CD70 CAR γδ T Cells: Summary and Next Steps

- Armored CD70 CAR  $\gamma\delta$  T cell program produced a compelling lead that warrants further development
- Program on-track for achieving differentiated target profile and proceeding to IND-candidate nomination
- Additional data expected to be presented in 2023

# Opportunity for Adicet and γδ T cells

- Response to low antigen density by design with CD27based CAR (compared to scFv-based CAR)<sup>3</sup>
- Three mechanisms of action designed to address tumor heterogeneity
- Homing of γδ T cells documented in RCC and improved AML OS
- Inclusion of armoring to address suppressive TME

# Building a Broad Pipeline of First-in-Class CAR and CAd $\gamma\delta$ T Cell Product Candidates



ADI-925 is an engineered Chimeric Adapter (CAd) y61 T cell product candidate targeting stress ligands, including MICA/MICB & ULBP1-6, expressed on malignant cell



### Armored PSMA CAR γδ T Cell Opportunity For Differentiation

### Target validation

- · PSMA expression is present in >85% of patients with mCRPC1 with limited expression in normal tissues (100-1,000 times overexpressed)
- · Clinically validated via multiple modalities:
- · PSMA targeted radiotherapy approved for mCRPC2
- · Immunotherapies (T-cell engaging antibodies and cell therapies) demonstrated PSA, PSMA-radiographic, and **RECIST responses** in early clinical studies3,4,5

### Key challenges

- · Limited therapeutic index due to CRS, ICANS, and macrophage activation syndrome with PSMA targeted T cell engagers and alpha-beta CAR T cell approaches4,6
- · Single mechanism of targeting limits activity in heterogeneous tumors
- · Immunosuppressive environment of mCRPC associated with TGFB7

### Opportunity for Adicet and yo T cells

- · Potent CAR construct active against heterogeneous PSMA
- · Three mechanisms of action designed to address tumor heterogeneity
- · Homing of γδ T cells documented in mCRPC
- · Inclusion of armoring to address suppressive TME
- · No significant CRS and ICANS demonstrated with Adicet CAR yo T cells in clinical trials reported to-date; potential to address therapeutic index

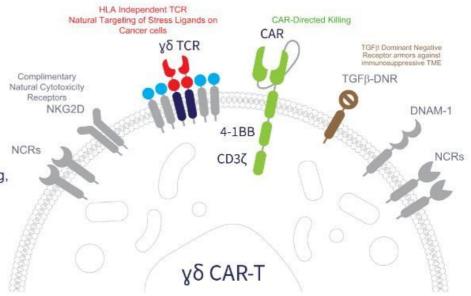
- Train et al. Ann Onc. (2020)
   Nirazaei et al. Int J Biol Macromol (2022)
   Slovin et al. JCO supp (2022)
   Natayan et al. Nat Med (2022)

mCRPC: metastatic castrate-resistant prostate cancer; PSMA: Prostate-specific membrane antigen; RECIST; Response Evaluation Criteria in Solid Tumors



### Armored PSMA CAR γδ T Cell Program

- Lead candidate targeting PSMA demonstrated improved characteristics versus benchmark<sup>1</sup>
- Heterogeneous PSMA
- Armoring technology via TGFβ-DNR improved activity, serial killing, and functional resilience



1. Liu et al. Cancer Res. (1997)

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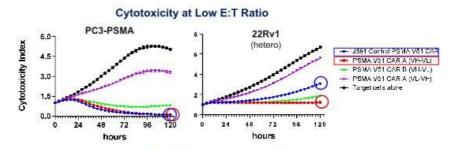


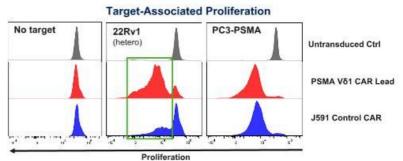
# Adicet's Lead PSMA CAR Construct Demonstrated Improved Cytotoxicity and Targeting Over Benchmark



#### Lead CAR Construct demonstrated:

- Efficient activation of NFAT signaling
- Recognition of intermediate/ heterogeneous PSMA expression
- Broad target-associated proliferation
- Improved cytotoxicity over benchmark1
- Improved in vivo activity over benchmark1





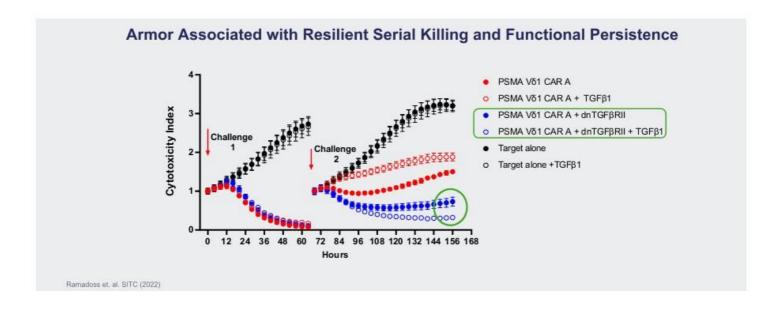
Ramadoss et al. SITC 2022

1. Liu et al. Cancer Res. (1997)



# Armored PSMA CAR $\gamma\delta$ T Cell Program Improved Resilience with TGF $\beta$ -DNR



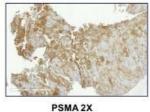


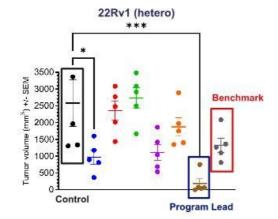
Adicet Bio

# Armored PSMA CAR γδ T Cell Program In Vivo Activity and Next Steps









- Armored PSMA CAR γδ T cell program demonstrated significant antitumor activity across mCRPC models
- Program lead is progressing through efficacy and manufacturing assessment for IND-candidate confirmation

Ramadoss et. al, SITC (2022)

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# Building a Broad Pipeline of First-in-Class CAR and CAd $\gamma\delta$ T Cell Product Candidates



'ADI-925 is an engineered Chimeric Adapter (CAd) y61 T cell product candidate targeting stress ligands, including MICA/MICB & ULBP1-6, expressed on malignant cell



### Armored B7-H6 CAR γδ T Cell Opportunity For Differentiation

### **Target validation**

- · B7-H6 is expressed in >95% of CRC cases, and other gastrointestinal cancers, with limited expression in normal tissues1
- · Well established biological role in natural tumor surveillance and active clinical development
- · Preclinical activity with bispecific T-cell engaging antibodies2 and NKp30 related CAR T3
- · In vivo activity with B7-H6 targeted therapies against both hematological and solid tumor indications<sup>2,3,4</sup>

#### Key challenges

- · Susceptibility to inhibition by shed
- · Selective and specific antigen targeting
- · Single MOA overcome by heterogeneous antigen expression in solid tumors
- · Limited intratumoral homing and proliferation
- · Immunosuppressive TME

### Opportunity for Adicet and yδ T cells

- · Potential to be first-in-class B7-H6 targeted cell therapy product into clinic
- · Designed to retain activity in presence of shed antigen and armored against TME
- High level of homing for Vδ1 T cells in GI and colorectal tissues
- · Three mechanisms for antitumor activity (CAR, innate, and adaptive immunity) designed to address tumor heterogeneity

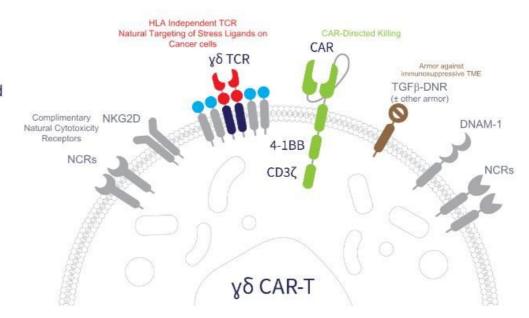
1. NCT04752215 2. Zhang et al. Clin Can Res (2022)

CRC: Colorectal cancer; GI: Gastrointestinal; MOA: Mechanism of action



### Armored B7-H6 CAR γδ T Cell Program

- Library of de novo CAR constructs screened for binding characteristics and improved performance versus benchmark<sup>1</sup>
- CAR optimization underway to define lead construct in 2023
- Iterative and novel armoring enhancements being explored



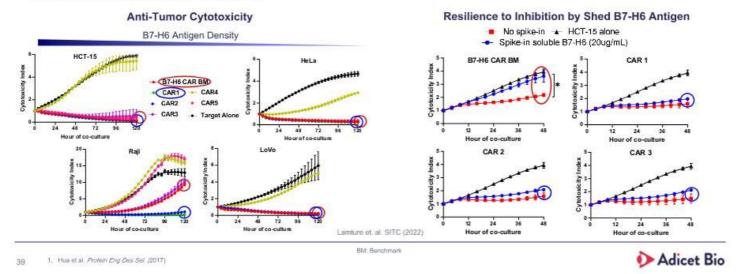
Adicet Bio

1. Hus et al. Protein Eng Des Sel. (2017)

# B7-H6 CAR $\gamma\delta$ T Cells Applicable Across Range of Indications



- B7-H6 CAR γδ T cells demonstrated anti-tumor activity across range of target antigen densities and indications
- Adicet's tailored scFv screening strategy has yielded CAR leads resilient to inhibition by shed antigen compared to benchmark<sup>1</sup>



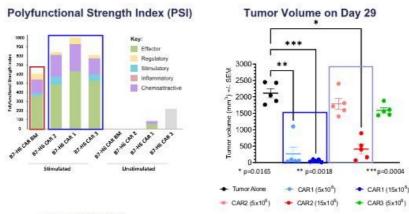
## B7-H6 CAR γδ T Cells Demonstrated Anti-Tumor Activity and Improved Polyfunctional Strength







- B7-H6 CAR γδ T cells demonstrated improved polyfunctional strength compared to benchmark CAR<sup>1</sup>
- Efficacy studies support potent anti tumor activity
- Lead discovery proceeding with additional armoring technologies



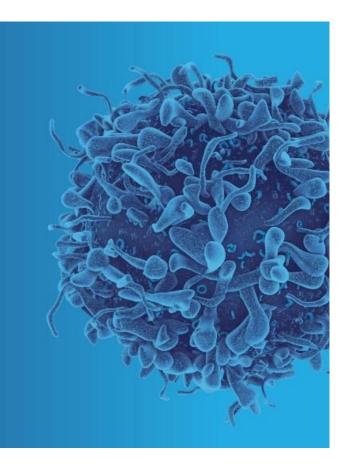
Lamture et. al. SITC (2022)



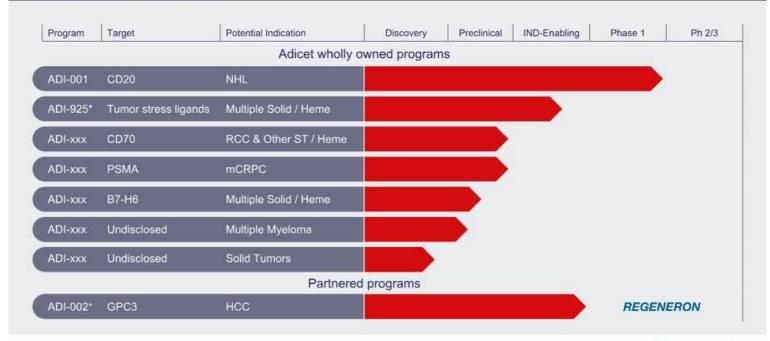


# Summary of Pipeline and 2022 SITC Data

Blake Aftab, Ph.D. Chief Scientific Officer



# Building a Broad Pipeline of First-in-Class CAR and CAd $\gamma\delta$ T Cell Product Candidates



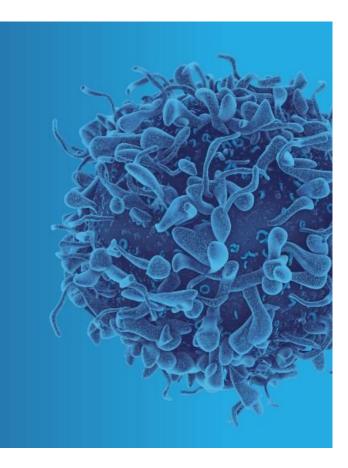
'ADI-925 is an engineered Chimeric Adapter (CAd) y61 T cell product candidate targeting stress ligands, including MICA/MICB & ULBP1-6, expressed on malignant cells





### Perspectives: Adicet γδ T Cell Pipeline

Marco Davila, M.D., Ph.D.
Roswell Park Comprehensive Cancer Center



### Perspectives



Marco Davila M.D., Ph.D. Associate Director and SVP Translational Research Roswell Park Comprehensive Cancer Center Adicet Bio Scientific Advisory Board Member **Question 1**: Can you share your preliminary thoughts on the pipeline we presented today? What do these engineered  $\gamma\delta 1$  T cell programs offer?



### Perspectives



Marco Davila M.D., Ph.D. Associate Director and SVP Translational Research Roswell Park Comprehensive Cancer Center Adicet Bio Scientific Advisory Board Member **Question 1:** Can you share your preliminary thoughts on the pipeline we have compiled here? What do these engineered  $\gamma\delta 1$  T cell programs offer?

**Question 2:** Can you share your thoughts on ADI-925 and the chimeric adaptor technology?



### Perspectives



Ph.D.
Associate Director and SVP
Translational Research
Roswell Park Comprehensive
Cancer Center
Adicet Bio Scientific Advisory
Board Member

**Question 1:** Can you share your preliminary thoughts on the pipeline we have compiled here? What do these engineered  $\gamma\delta 1$  T cell programs offer?

Question 2: Can you share your thoughts on ADI-925 and the chimeric adaptor technology?

**Question 3:** With CD70 and PSMA, can you provide your perspectives in the context of previous approaches to these targets?

Adicet Bio



Chen Schor, President and CEO



Sattva Neelapu, M.D.
Department of
Lymphoma-Myeloma
Division of Cancer
Medicine
The University of
Texas, MD Anderson
Cancer Center

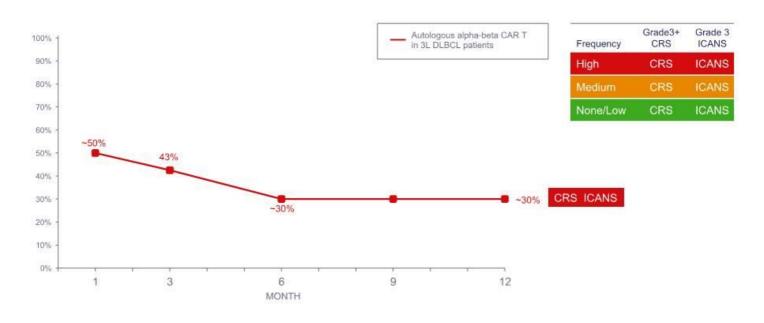


Adicet Bio to discuss recent data from ongoing Phase 1 clinical study in R/R aggressive B-cell NHL

When: Sunday, December 11 at 8:00 a.m. CT/ 9:00 a.m. ET



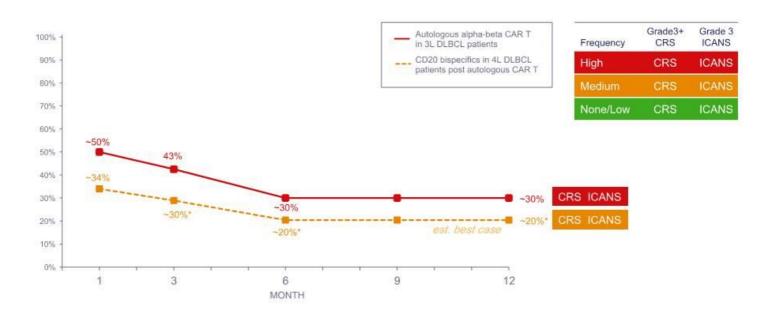
### Advanced Therapies CR Rate Over Time in 3L and 4L Aggressive NHL



Autologous alpha-beta CAR T in 3L DLBCL patients N Engl J Med. (2017) December 28; 377(26): 2531–2544 J Clin Oncol 38:3095-3106. CD20 bispecifics in 4L DLBCL patients post autologous CAR T Gloftsmab ASCO (2022). Epocritamab EHA (2022), GMAB PR June 11 (2022)
\*\*CR rate durability for post-CAR T bispecifics modeled as durability of 3L+ autologous CAR-T (for modeling purpose only) 3L= Third line, 4L= Fourth line



### Advanced Therapies CR Rate Over Time in 3L and 4L Aggressive NHL



Autologous alpha-beta CAR T in 3L DLBCL patients N Engl J Med. (2017) December 28; 377(26): 2531–2544 J Clin Oncol 38: 3095-3106. CD20 bispecifics in 4L DLBCL patients post autologous CAR T
Glofitamab ASCO (2022), Escoritamab EHA (2022), GMAB PR June 11 (2022)
\*CR rate durability for post-CAR T bispecifics modeled as durability of 3L+ autologous CAR-T (for modeling purpose only)



### **Anticipated Near-Term Milestones**

### ADI-001 Phase 1 Study in R/R NHL

- Complete dose escalation through DL4; backfill enrollment to DL3
- · ASH clinical update Dec. 11 at 8:00 a.m. CT
- Establish recommended Phase 2 dose by end of 2022

### Pipeline and Manufacturing

- ADI-925: IND submission expected H2/2023
- One new IND planned every 12-18 months
- Leverage in-house GMP manufacturing to support expanding clinical pipeline

### ADI-001 - Expansion

- Discuss with the FDA and EMA the design of two potentially pivotal studies and a path to support BLA and MAA submissions
- Initiate a potentially pivotal program in H1/2023
- · Initiate additional expansion cohorts in 2023

### Corporate

 Well financed into H1/2025 with \$282M cash and cash equivalents (as of 9/30/22)

Adicet Bio

BLA: Biologics License Application; DL: Dose level; EMA: European Medicines Agency; FDA; U.S. Food and Drug Administration; MAA: Marketing Authorization Application; R/R= Relapsed/refractory



