# 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): January 16, 2024

# 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)

Che	eck the appropriate box below if the Form 8-K filing is intended to s	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 A	act (17 CFR 240.14a-12)						
	Pre-commencement communications pursuant to Rule 14d-2(b) u	nder the Exchange Act (17 CFR	240.14d-2(b))					
	Pre-commencement communications pursuant to Rule 13e-4(c) un	nder the Exchange Act (17 CFR	240.13e-4(c))					
	Securities registered pursuant to Section 12(b) of the Act:							
		Trading						
	Title of each class	Symbol(s)	Name of each exchange on which registered					
	Common Stock, par value \$0.0001 per share	ACET	The Nasdaq Global Market					
	icate by check mark whether the registrant is an emerging growth courities Exchange Act of 1934 (§ 240.12b-2 of this chapter).	ompany as defined in Rule 405 c	of the Securities Act of 1933 (§ 230.405 of this chapter) or Rule 12b-2 of the					
Eme	erging growth company							
	n emerging growth company, indicate by check mark if the registrate outling standards provided pursuant to Section 13(a) of the Exchange		nded transition period for complying with any new or revised financial					

#### Item 7.01 Regulation FD Disclosure.

On January 16, 2024, Adicet Bio, Inc. (the "Company") posted to the "Presentations & Events" section of the Company's website at investor.adicetbio.com an updated corporate presentation (the "Corporate Presentation"). A copy of the Corporate Presentation is furnished herewith as Exhibit 99.1.

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 9.01 Exhibits.

(d) Exhibits

Exhibit No. Description

99.1 Adicet Bio, Inc. Corporate Presentation, dated January 16, 2024, 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.

January 16, 2024 Date:

By: Name: Title: /s/ Nick Harvey Nick Harvey Chief Financial Officer



## Forward-Looking Statements

This presentation contains "forward-looking statements" of Adicet Bio, Inc. (Adicet or the Company) 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; preclinical and clinical development of Adicet's product candidates, including future plans or expectations for ADI-001 and ADI-270 and the potential safety, durability, tolerability and efficacy of these product candidates; the expected progress, timing and success of the Phase 1 clinical trial of ADI-001, including continued enrollment and expectations around a clinical update in the second half of 2024; the Company's plan to initiate a Phase 1 clinical trial of ADI-001 in lupus nephritis, including the potential for a clinical update in the second half of 2024; the Company's expansion into other autoimmune indications in the future, including IND submissions, acceptances and the initiation of clinical trials; the Company's expectations regarding the submission of an IND for ADI-270 in renal cell carcinoma in the second quarter of 2024 and timing for future clinical updates; and expectations regarding its uses of capital, expenses and financial results, including the 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 global economic conditions and public health emergencies on Adicet's business and financial results, including with respect to disruptions to its preclinical and clinical studies, business operations, and ability to raise additional capital; Adicet's ability to execute on its strategy, including obtaining the requisite regulatory approvals on

#### **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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# Developing Broad Pipeline of Allogeneic $\gamma\delta1$ T Cell Therapies for Al and Cancer

Program	Target	Potential Diseases	Research	IND-Enabling	Clinical	Status			
ADICET WHOLLY OWNED PROGRAMS									
ADI-001	CD20	Autoimmune	-	•	-	IND in LN cleared Dec 2023 Initiate LN Phase 1 2Q 2024 Update planned 2H 2024 Al expansion opportunities			
ADI-001	CD20	NHL		-	_	MCL Phase 1 ongoing* Update planned 2H 2024			
ADI-270	CD70 (TGFβ-DNR)	RCC & Other ST / Heme	_	•	<del>-</del>	IND submission in RCC expected 2Q 2024			
ADI-xxx	PSMA (w/ Armor)	mCRPC		<del></del>	<u> </u>	Preclinical activities			
ADI-925	Tumor stress ligands	Multiple Solid / Heme	-	0	<u> </u>	Preclinical activities			
PARTNERED PROGRAMS									
ADI-002	GPC3	HCC			—	REGENERON			



<sup>\*</sup>Adicet is focused on advancing MCL enrollment in the GLEAN trial and has deprioritized enrolling large B-cell lymphoma patients.

Al= Autoimmune; GPC3= Glypican-3; HCC= Hepatocellular carcinoma; IND= Investigational new drug; LN= Lupus nephritis; mCRPC= Metas

NHL= Non-Hodgkin's lymphoma; PSMA= Prostate specific membrane antigen; RCC= Renal cell carcinoma; ST= Solid tumor

# Adicet Bio Leadership Team



Chen Schor President and CEO



res**TOR**bio



Blake Aftab, Ph.D. Chief Scientific Officer

Chief Medical Officer















Nancy Boman, M.D., Ph.D. Chief Regulatory Officer

Francesco Galimi, M.D., Ph.D.





Don Healey, Ph.D. Chief Technology







Nick Harvey Chief Financial Officer





Amy Locke Head of Human Resources

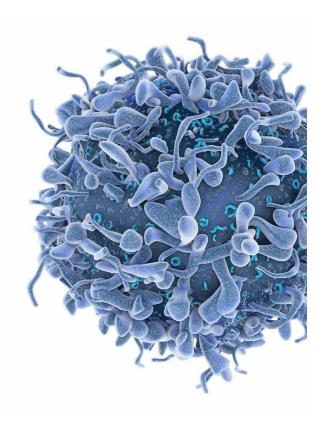








# **ADI-001**Autoimmune Diseases



## Adicet γδ1 CAR T Cell Therapy For Autoimmune Indications

#### ADI-001 Data in NHL Provides Strong Foundation for Future Development in Al

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Exposure Consistent with Approved Autologous CAR T (Cmax, Day 28 Persistence and AUC)

B-Cell Depletion Consistent with Autologous CD19
CAR T in SLE, SSC and IIM

Preferentially Trafficking to Organs/ Tissues

No Significant Risk of CRS, ICANS, or T cell Malignancies
Compared to
Autologous CAR T

Readily Available, "Off-the-Shelf"

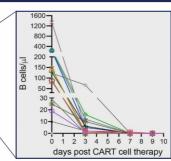
Potential to Dose in Community Setting

AUC= Area Under the Curve; Cmax= Peak plasma concentration; CRS= Cytokine release syndrome; ICANS= Immune effector cell-associated neurotoxicity syndrome; IIM= idiopathic inflammatory myopathy SLE= systemic lupus erythematosus; SSC= systemic scierosis

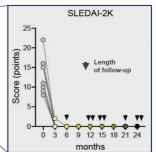


# CAR-T Cell Therapy Depleted B-cells, Drove an Immune Reset, and Achieved Treatment-Free Remissions in Patients with Al Diseases'

- Schett et al. treated 15 patients with AI diseases (SLE, IIM, & SSc) with autologous CD19 CAR T
- Deep B-cell depletion & immune reset observed in all patients



- Lasting treatment-free remissions observed in all SLE patients
- Symptom control with no additional disease-specific treatment achieved in all patients
- Robust efficacy demonstrated in IIM and SSc patients



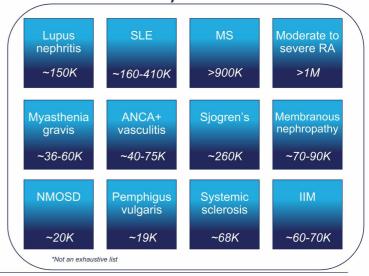
<sup>1</sup> CD19.CAR-T Cell in Refractory Systemic Autoimmune Diseases: A Monocentric Experience from the First 15 Patients. ASH 2023 Note: Third-party data summarized on this slide does not employ ADI-001.



# Opportunity to Address Unmet Needs in a Large Number of B-Cell Mediated Autoimmune Diseases

- POC for CAR-T mediated B-cell depletion demonstrated in multiple AI diseases
  - Lupus and lupus nephritis<sup>1,2,3</sup>
  - Systemic sclerosis<sup>2</sup>
  - Idiopathic inflammatory myopathies<sup>2</sup>
  - Myasthenia gravis<sup>4,5</sup>
- B-cell depletion via CD20 antibodies (i.e., rituximab, obinutuzumab) further validate the therapeutic approach in several of these diseases
- Deep B-cell depletion in the tissues and secondary lymphoid organs highly desirable for therapeutic success

#### Potential autoimmune diseases for development\* & U.S. prevalence

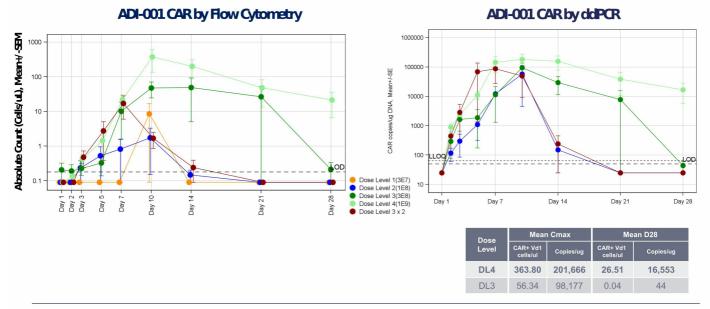


ANCA = Antineutrophilic cytoplasmic antibody; NMOSD= Neuromyelitis optica significantification of the science o



Mackensen A et al. Nature Medicine 2022
 CD19.CAR-T Cell in Refractory Systemic Autoimmune Diseases: A Monocentric Experience from the First Filteen Patients. ASH 2023
 YB323 Poster @ American College of Rheumatology Convergence November 2023

# Cmax, D28 Persistence and AUC Consistent Values Reported for Approved Autologous CD19 CAR T1



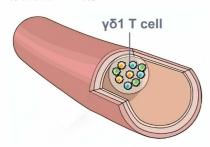
Badbaran, A. Cancers 2020;12, 1970; Locke et al. N Engl J Med 2022; 386:640-654; Neelapu et al. N Engl J Med. 2017;377:2531-2544; Ogasawara et al. Clin Pharmacokinet 60, 1621–1633 (202) D28= Day 28



# $\gamma\delta1$ T Cells Preferentially Home to Tissues

## peripheral blood<sup>10, 11</sup>

% of CD3+: ~1-3%



Ratios empirically calculated or approximated from proportion of %CD3 from literature reports in relative compartment

Images adapted from Hunter et al J Hepatol. 2018 and Ribot et al Nat Rev Immunol. 2021

References:

¹Rancan et al Nat Immunol 2023
²Brauneck et al Front Med 2021
³Davey et al Trends Immunol 2018
²Uger et al Sci Rep 2018
²Wyang 0. et al Exp Ther Med 2020
¹¹Dèchanet et al J Infect Dis 1999

<sup>5</sup>Wu et al Sci Transl Med 2019 <sup>6</sup>Deusch et al Eur J Immunol 1991 <sup>7</sup>Melo et al Clin Immunol 2021 <sup>8</sup>Toulon et al J Exp Med 2009 <sup>9</sup>Wisnewski et al Am J Respir Cell Mol Biol 2000



lymph node<sup>3,4</sup> CD27+ CD62L+

**V**δ1+ ↑↑ Vδ2+ ↓↓



skin<sup>8</sup>

tissue/blood: 8X



breast<sup>5</sup>

tissue/blood: ~15X adipose tissue/blood: 9X



bone marrow<sup>2</sup> tissue/blood: 4X

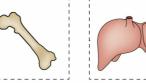


GI<sup>6</sup> tissue/blood: 11X



kidney1

tissue: >3X  $\gamma\delta$  vs  $\alpha\beta$ ~3X more Vδ1 vs Vδ2+



liver<sup>7</sup> tissue/blood: 3X



lung<sup>9</sup>

tissue/blood: 9X

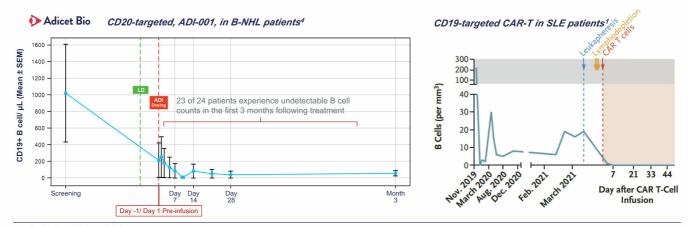
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# ADI-001 in AI Diseases: B-Cell Depletion Consistent with Autologous CD19 CAR T in SLE Academic Studies

#### **B-cell Depletion**

- B-cell depletion data from ADI-001 trial in NHL mirrored experience of autologous CD19 CAR T
- B-cell depletion via CD20 targeting validated by CD20-targeted antibody (obinutuzumab) which demonstrated efficacy on top of SOC in lupus nephritis in a Phase II clinical study<sup>3</sup>



- Mougiakakos MD et al. NEJM 2021 Mackensen A et al. Nature Medicine 2022 Furie RA et al. Ann Rheum Dis. 2022 Adicet internal data

#### ADI-001 in AI Diseases: B-Cell Depletion Consistent with Autologous CD19 CAR T "Off-the-Shelf" with Advantageous Tissue Tropism and Safety Profile

#### **B-cell** depletion

- · B-cell depletion data from ADI-001 trial in NHL mirrors experience of autologous CD19 CAR T in SLE, systemic sclerosis and idiopathic inflammatory myopathy (IIM)<sup>1,2</sup>
- · B-cell depletion via CD20 targeting validated by CD20-targeted antibody (obinutuzumab) which demonstrated efficacy on top of SOC in lupus nephritis in Phase II clinical study3

#### γδ1 T cell homing to tissues of interest

- · Inability to deplete tissue-resident B cells in secondary lymphoid organs or other tissues is a contributing reason for failure of targeted agents in lupus<sup>4,5,6</sup>
- · yδ1 T cells preferentially traffic to organs/tissues<sup>7</sup> and may be ideally suited to deplete B cells in secondary lymphoid organs, kidneys and other organs

#### **Favorable** safety profile "Off-theshelf"

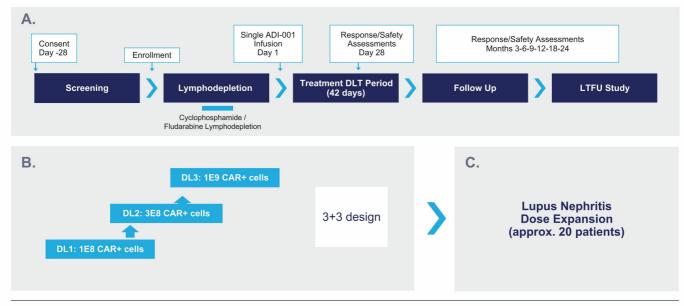
- No significant risk of CRS, ICANS or T cell malignancies compared to autologous CAR T\*
- · ADI-001 is an "off-the-shelf" investigational therapy potentially well suited for autoimmune diseases
  - -Ability to dose in community setting, lower COGs, faster turnaround time

- Mougiakakos MD et al. NEJM 2021 Mackensen A et al. Nature Medicine 2022 Furie RA et al. Ann Rheum Dis. 2022 Kamburova EG et al. American Journal of T
- Reddy VK et al. Rheumatology 2022
   Sadun RE and Foster MH AJKD 2019
   Zhang, PJ Hematol Oncol 2023

\*May 4, 2023 Data-cut date, n=24 evaluable patients



# ADI-001 Phase 1 Study Design: Lupus Nephritis



Clinical protocol enables to de-escalate down to DL-1 of 3E7 CAR+ Cells

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## ADI-001 in Lupus Nephritis: Key Endpoints

# Primary endpoints

#### Part 1

- · DLT incidence and MTD
- · Incidence of treatment-emergent adverse events (TEAEs), including severity, seriousness, and relatedness

#### Part 2

· Safety profile at the MTD/MAD/RP2D of ADI-001 in patients with LN

# Secondary endpoints

- · Levels of ADI-001 cells in peripheral blood
- · Response to treatment:
  - CR or PR on day 28 and month 3, 6, 9, 12, 18, 24 after infusion of ADI-001
- Biomarkers associated with response to treatment:
  - Antibody to dsDNA, antinuclear antibody (ANA) and complement levels

# Exploratory endpoints

• Response per SLEDAI-2K/DORIS criteria



CR= Complete response; DLT= Dose limiting toxicity, dsDNa= Double-strand DNA; MAD= Maximum administered dose; MTD= Maximum tolerated dose; PR= Partial response; RP2D= Recommended Phase 2 dos SLEDAL-X/DDRISE Systemic Jusus enthematiosus disease activity index 2000.

## ADI-001: Lupus Nephritis

#### **Opportunity**

- Type of kidney disease caused by SLE, an AI disease which affects an estimated 325,000 people in the U.S.<sup>1</sup>
- LN is a serious complication of SLE which affects approximately 40% of patients with SLE<sup>2</sup> and occurs when the immune system attacks the kidneys<sup>3</sup>
- Current treatment aims to reduce symptoms, keep the disease from getting worse, and keep the kidneys working well enough not to need dialysis or a kidney transplant

#### Next Steps

- IND cleared for ADI-001 in lupus nephritis in December 2023
- Expect to initiate Phase 1 study 2Q 2024
- Expand to additional Al indications



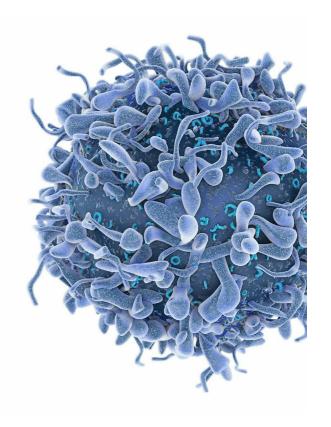
<sup>,</sup> Arthritis Rheum 2008 Jan;58(1):15-25, doi: 10.1002/art.23177.- Estimates of the prevalence of arthritis and other rheumatic conditions in the United States, Part 1

<sup>2</sup> Hoover P.I et a: Kidney Int 2016

Crampton, Steve P. et al. "Skin Malar rash Discoid rash CNS Spleen Splenomegaly Kidney Serum Glomerulonephritis Anti-nuclear antibodies Blood IFN signature Plasma blasts Anemia Thromborytonenia Neurolonical damane Affective disported rung Inflammation Joints Athritis" (2014).



ADI-001 Mantle Cell Lymphoma



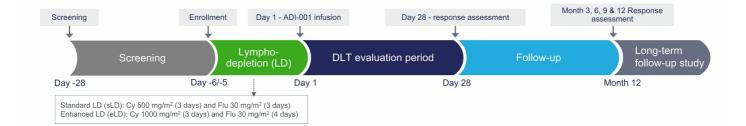
#### ADI-001: Encouraging Early Efficacy Data in Heavily Pretreated NHL Patients

- Demonstrated efficacy and favorable safety in 24 patients with aggressive r/r B-cell NHL
  - Heavily pre-treated patients: median 4 prior lines of therapy with 50% prior CAR T
  - Across All Doses: 71% ORR and 63% CR (LBCL & MCL)
  - RP2D: 75% ORR, 63% CR, 25% 6-months CR rate (LBCL & MCL)
  - MCL (all doses): 80% CR rate, 60% 6-months CR rate
  - No significant incidence of CRS or ICANS or T cell malignancy risk; "off-the-shelf"
  - DL4 (RP2D) Cmax, D28 persistence, AUC consistent with approved CD19 autologous CAR T
- Potential for highly differentiated profile for MCL: High CR Rate, favorable durability, safety, "off-the-shelf" dosing, and ability to dose in community setting
- Evaluating option of advancing to a potentially pivotal study in MCL under an accelerated approval pathway
- Strong foundation for growing pipeline of next-generation engineered γδ1 T cell therapies

May 4, 2023 Data-cut date, n=24 evaluable patients LBCL= Large B-cell lymphoma



## GLEAN: ADI-001 First-in-Human Study (CD20 CAR+ γδ1 T cells)



# ADI-001 Dose (CAR+ Cells) (3 + 3 escalation design)\*

 DL1	DL2	DL3	DL4
3E7	1E8	3E8	1E9

# Primary endpoint:

- Number of DLTs
- Treatment emergent and treatment-related AEs

# Secondary endpoint:

- ORR, DOR, PFS, TTP, and OS
- PK, immunogenicity

#### MCL:

- Enrolling MCL patients 3L+, DL4
- Prior CAR-T allowed

AEs= Adverse events; Cy= Cyclophosphamide; DL= Dose level; DOR= Duration of response; Flu= Fludarabine; GLEAN= Gamma deLta adoptive thErApy for Nhl-1; OS= Overall survival; PFS= Progression-free survival; PK= Pharmacokinetics; RR= Relapsed or refractory; TTP= Time to progression



<sup>\*</sup>Protocol part 1b includes patients receiving single lymphodepletion and two infusions at DL3 (two doses 300 million CAR+ cells, one on day 1 and the second dose on day 7)

#### ADI-001: Patient Characteristics

Patient Characteristics	N (%) (Total N = 24)
Age – median (range)	66.5 (44 - 75)
Sex – number of male	17 (70.8)
B cell malignancy (WHO 2017 classification)	
Large B cell lymphoma (LBCL)	18 (75.0)
· R/R diffuse large B cell lymphoma	13 (54.2)
- R/R high grade B cell lymphoma, double/triple hit	4 (16.7)
· R/R high grade B cell lymphoma, NOS	1 (4.2)
R/R mantle cell lymphoma (MCL)	5 (20.8)
Follicular	1 (4.2)
IPI score (LBCL) - median (range)	2.5 (1 - 4)
Simplified MIPI score-median (range)	5 (4 - 8)
Follicular IPI score-median(range)	2 (2 - 2)
Stage III & IV disease	17 (70.8)
Sum of the product of the diameters at screening - median (range)	3001 (150 - 7919) mm <sup>2</sup>
Prior lines of therapies - median (range)	4 (2 - 9)
Prior anti-CD19 CAR T therapies	12 (50.0)
Prior systemic anti-cancer therapy	
CD20 mAB + anthracycline-based chemo	23 (95.8)
CD20 mAB + non-anthracycline-based chemo	15 (62.5)
Only chemotherapy	1 (4.2)
POLA or POLA-R or POLA-BR	7 (29.2)
BTK inhibitors +/- other drugs (except CD20 combination)	5 (20.8)
CD20 mAB	7 (29.2)
CD19 biologics or combinations	5 (20.8)
Anti-CD19 CAR T	12 (50.0)
Other experimental therapies	2 (8.3)
CD20 +/- Lenalidomide/Bortezomib/Venatoclax/Ibrutinib	2 (8.3)
Refractory status at study entry	
Refractory to first-line therapies	10 (41.7)
Refractory to second-line therapies	12 (50.0)
Refractory to the last course of anti-cancer systemic therapy	17 (70.8)

- 23 patients had aggressive B-cell lymphoma: 18 LBCL and 5 MCL; 1 patient enrolled with follicular lymphoma
- Most patients were heavily pre-treated with median four prior lines of therapy, relatively high tumor burden and poor prognostic outlook
- Twelve patients (50%) progressed following approved autologous anti-CD19 CAR T cell therapy - Yescarta (axi-cel), Breyanzi (liso-cel) or Tecartus (brexu-cel)
  - 8/12 of patients progressed within less than 6 months from date of autologous CAR T administration
- ~70% of patients were refractory to the last course of systemic therapy, and the remaining had relapsed

Aay 4, 2023 Data-cut date, n=24 evaluable patients; IPI= International Prognostic Index; MIPI= Mantle Cell Lymphoma Prognostic Index; WHO= World Health Organization



# ADI-001: Efficacy Summary by Dose Level

	Median No. of Prior Lines	ORR (%)	CR Rate (%)	6-month CR Rate (%)
DL4 (RP2D; LBCL & MCL)	4	6/8 (75.0%)	5/8 (62.5%)	2/8 (25.0%)
DL 4 MCL	4	2/2 (100.0%)	2/2 (100%)	2/2 (100%)
All Doses (LBCL & MCL)	4	17/24 (70.8%)	15/24 (62.5%)	4/24 (16.7%)
All Doses MCL	4	4/5 (80%)	4/5 (80%)	3/5 (60%)

High CR rate and favorable durability in MCL

May 4, 2023 Data-cut date; Data are subject to further review and verification



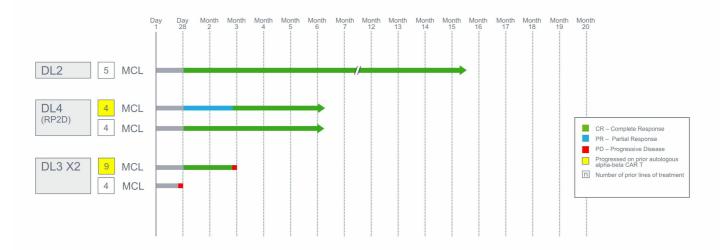
# ADI-001: Safety Data in Efficacy Evaluable Patients

	DL1 (N=3)		DL2 (	N=3)	DL3 (	N=6)	DL3 X2 (N=4) DL4 (N			N=8) Total (N=24)		(N=24)
	Any Grade	Gr>=3	Any Grade	Gr>=3	Any Grade	Gr>=3	Any Grade	Gr>=3	Any Grade	Gr>=3	Any Grade	Gr>=3
CRS	2 (66.7%)	0	0	0	1 (16.7%)	1 (16.7%)	4 (100.0%)	0	4 (50.0%)	0	11 (45.8%)	1 (4.2%)
ICANS	0	0	1 (33.3%)	0	0	0	1 (25.0%)	1 (25.0%)	1 (12.5%)	0	3 (12.5%)	1 (4.2%)
GvHD	0	0	0	0	0	0	0	0	0	0	0	0
DLT	0	0	0	0	0	0	0	0	0	0	0	0
Infection	1 (33.3%)	1 (33.3%)	2 (66.7%)	0	3 (50.0%)	2 (33.3%)	2 (50.0%)	1 (25.0%)	3 (37.5%)	2 (25.0%)	11 (45.8%)	6 (25.0%)
SAE-TEAE	1 (33.3%)	1 (33.3%)	2 (66.7%)	2 (66.7%)	4 (66.7%)	3 (50.0%)	2 (50.0%)	2 (50.0%)	3 (37.5%)	2 (25.0%)	12 (50.0%)	10 (41.7%)
Related SAE- TEAE	1 (33.3%)	0	1 (33.3%)	1 (33.3%)	3 (50.0%)	2 (33.3%)	2 (50.0%)	2 (50.0%)	3 (37.5%)	2 (25.0%)	10 (41.7%)	7 (29.2%)

May 4, 2023 Data-cut date; Data are subject to further review and verification. Safety assessment was performed using the Common Terminology Criteria for Adverse Events (v5) and the American Society for Transplantation and Cellular Therapy criteria. AE= Adverse event; SAE= serious adverse event; TEAE= treatment emergent adverse event



# Strong CR Rate and Durability in 4L+ MCL Patients



ORR: 80%, CR rate: 80%, 6-month CR rate: 60% No significant CRS or ICANS

May 4, 2023 Data-cut date, n=24 evaluable patients; Data are subject to further review and verification.



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## Data Provides Strong Foundation for Future Development in MCL

- √ High CR rate
- √ Favorable durability in late-line patients
- ✓ Superior cell killing potency compared to autologous CAR T¹
- ✓ Cmax, Day 28 persistence and AUC consistent with approved CD19 autologous CAR T
- ✓ Favorable safety profile with no significant risk of CRS, ICANS, or T-cell malignancy
- ✓ Potential to dose in community setting

May 4, 2023 Data-cut date, n=24 evaluable patients

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## Focused Investments in Differentiated Early-Stage Pipeline

# ADI-270: Renal cell carcinoma and other solid tumors

- IND submission in RCC expected 2Q 2024
- Targeting CD70 via CD27-ligand has shown superiority to scFv CARs in preclinical studies
- Innate and adaptive targeting mechanisms associated w/ activity in RCC, AML, and other malignancies
- Armoring via TGFβ dominant-negative receptor addresses suppressive TME and HvG resilience
- Lead CAR demonstrated potency and improved serial killing & resilience against suppressive factors in preclinical models

#### Metastatic castrationresistant prostate cancer program

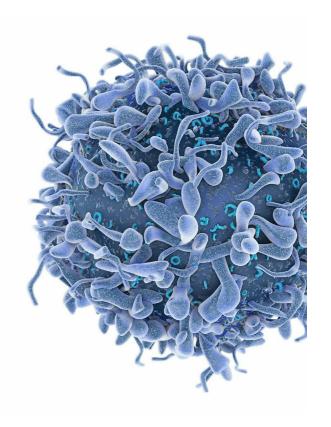
- Adicet lead scFv antibody designed to have similar binding determinants as approved radioligand therapy
- Highly potent cell therapy with armoring intended to address immunosuppressive solid tumor environment & heterogeneous PSMA expression
- Leveraging γδ1 T cell tropism to solid tissues
- Multiple modes of cell killing: CAR mediated, innate and adaptive immunity

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vG=Host vs. Graft; RCC= Renal cell carcinoma; scFv= Single-chain fragment variable; TME= Tumor microenvironment

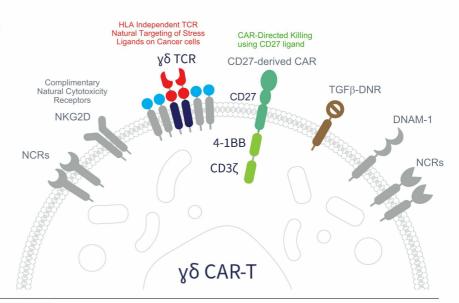


ADI-270 Renal Cell Carcinoma



# ADI-270: Adicet's Armored CD70 CAR γδ1 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 RCC and other indications<sup>2</sup>
- Armoring via dominant negative receptor; addresses TGFβ in TME<sup>3</sup>
- Next-generation CAR format demonstrated potency and improved serial killing, and resilience against suppressive factors and HvG
- Homing and activity of γδ1 T cells demonstrated in RCC<sup>2</sup>

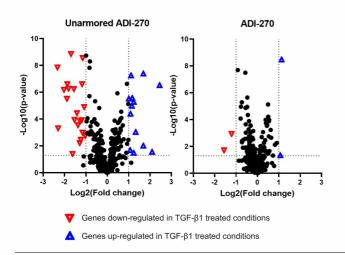


Sauer et al. Blood (2021); <sup>2</sup> Rancan et al. Nat Immunol 2023; <sup>3</sup> Junker et al. Cytokine (2000); HvG=Host vs. Gra

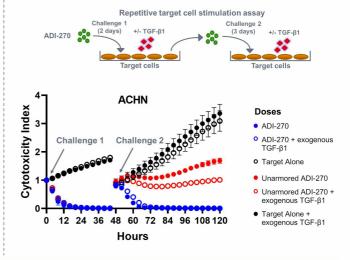
Adicet Bio

# Armored CAR Demonstrated Resilience Against TGFβ and Maintained Potent Cytotoxic Function in Preclinical Models

Armored ADI-270 cells are protected against  $TGF\beta$ -mediated alterations to activation expression profile



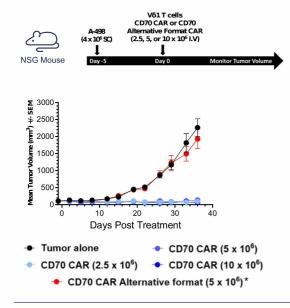
#### Armored CAR demonstrated improved serial killing



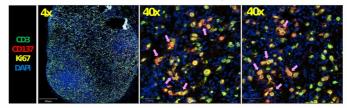
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# CD70 CAR γδ1 T Cells Demonstrated Activity In Vivo and Proliferated Within Tumors



Tumor Infiltration and Proliferation of γδ1 CAR T cells



• Anti-tumor activity associated with CAR  $\gamma\delta 1$  T cell tumor infiltration and proliferation within the tumor bulk as evidenced by areas of marker colocalization noted with pink arrows

Adicet Bio

# Armored CD70 CAR γδ1 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 mAb<sup>4,5</sup>
- Single-digit ORR 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 γδ1 T cells

- Response to low antigen density by design with CD27based CAR (compared to scFvbased CAR)3
- Three mechanisms of action designed to address tumor heterogeneity
- Homing of  $\gamma\delta 1$  T cells reported in RCC
- Next-generation CAR format enhanced durability and improved HvG resilience
- · Inclusion of armoring to address suppressive TME

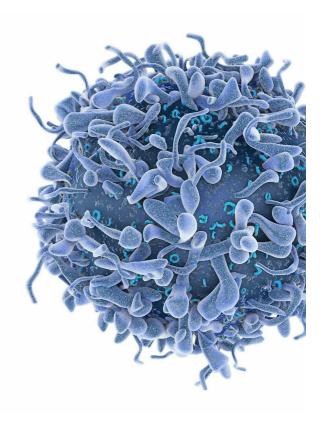
IND filing expected 2Q 2024

- 7. Massard et al. Cancer Chemother Pharmacol (2019) 8. CRISPR Therapeutics Presentation (2022)





# **PSMA Program**



## Armored PSMA CAR γδ1 T Cell Program

#### Program:

- Allogeneic PSMA CAR γδ1 T cell therapy candidate for prostate cancer (PCa)
- Armoring demonstrated with TGFβ-DNR provides functional advantage for treating solid tumors
- Demonstrated penetration & killing in mCRPC primary tumor organoids

#### Historical challenge:

#### Safety issues with PSMA-targeted immunotherapies



- Radioligand therapies are the only approved PSMA-targeted agents and bind functional PSMA homodimer, associated with a favorable and differentiated off-tissue AE profile 1
- Adicet's CAR is designed to have similar binding determinants as radioligand therapies, recognizing a conformational epitope present on the homodimer ( $\uparrow$  selectivity)<sup>1,2</sup>
- Promising safety profile for  $\gamma\delta1$  T cells observed in ADI-001 clinical trial
- Modest efficacy with PSMA-targeted immunotherapies



- Highly potent cell therapy compatible with armoring to address immunosuppressive solid tumor environment & heterogeneous PSMA expression
- Multiple modes of cell killing CAR mediated, innate and adaptive immunity
- Immunosuppressive TME & poor T cell infiltration in PCa



Natural tropism of γδ1 T cells to solid tissues

Adicet approach:

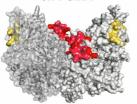
Infiltration of  $\gamma\delta 1$  T cells reported and demonstrated in prostate cancer

Adicet Bio

Deshayes E et al. Cancer (2023)
 Schülke N et al. PNAS (2003)

# Lead PSMA CAR Binding Determinants Designed to be Consistent With Pluvicto, The Only PSMA-Targeted Therapy With Validated Efficacy & Safety Profile

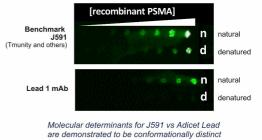
# Binding interface of mAbs on PSMA



Binding interface of Adicet lead binder (red) & benchmark binder J591 (yellow; Tmunity)

# Full-length PSMA Membrane distal Membrane proximal

#### **PSMA Antigen Capture Assay**

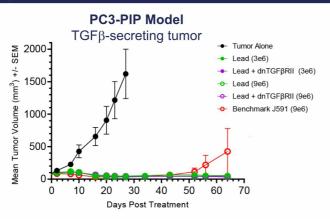


- are demonstrated to be comormationally distinct
- Radioligand therapies bind the functional PSMA homodimer and are associated with a favorable efficacy and off-tissue AE profile, and is
  the only approved PSMA-targeted therapy
- This profile is distinct and differentiated from other approaches for CAR, ADC, and bispecifics that recognize non-conformational, linear, or monomeric epitopes

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# PSMA CAR $\gamma\delta1$ T-Cell Activity Observed in Primary Patient mCRPC Organoids



- Organoid killing

  Too deals

  Organoid killing

  Ond deals

  Ond deal
- Adicet PSMA CAR and armored versions retained Demonstrated infiltration tumor control in TGFβ-secreting prostate cancer model derived tumor organoids
- Adicet's unarmored lead CAR retained maximal control beyond 65 days at 3e6 CAR cell dose
- Benchmark CAR (J591) lost control of tumor despite a significant 9e6 CAR T cell dose
- Demonstrated infiltration and killing of primary patient derived tumor organoids
- V81 T cells alone demonstrated intrinsic infiltration and killing, consistent with reported anti-tumor activity in mCRPC
- Infiltration and killing further enhanced with PSMA CAR

Adicet internal data.

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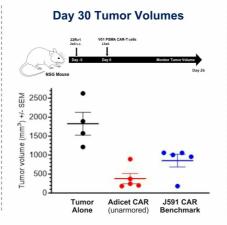


# CAR γδ1 T Cells Highly Active Against 22Rv1 Xenograft Model, A Challenging Model With Intermediate & Heterogenous PSMA Expression

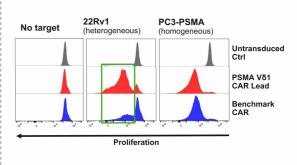
#### 22Rv1 Tumors Express Intermediate and Heterogeneous PSMA



PSMA 2X



#### **Target-Associated Proliferation**



- · Lead PSMA CAR retained robust target-associated proliferation in heterogenous PSMA expressing models
  - Benchmark CAR (J591) did not retain robust proliferation in the context of heterogenous mCRPC
  - Adicet's lead CAR retained superior tumor control versus benchmark CAR

Adicet internal data



# Armored PSMA CAR γδ1 T Cell Opportunity For Differentiation

#### **Target validation**

- PSMA expression is present in >85% of patients with mCRPC with limited expression in normal tissues (100-1,000 times overexpressed)1
- · Clinically validated via multiple modalities:
- PSMA targeted radiotherapy approved for mCRPC<sup>2</sup>
- Immunotherapies (T-cell engaging antibodies and cell therapies) demonstrated PSA, PSMAradiographic, 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 approaches3,6
- · Single mechanism of targeting limits activity in heterogeneous tumors
- Immunosuppressive environment of mCRPC associated with TGFβ7

#### **Opportunity for Adicet and** γδ1 T cells

- Potent CAR construct active against heterogeneous PSMA
- · Three mechanisms of action designed to address tumor heterogeneity
- Homing of γδ1 T cells documented in mCRPC
- · Inclusion of armoring to address suppressive TME
- No significant CRS and ICANS demonstrated with Adicet CAR  $\gamma\delta1$ T cells in clinical trials reported todate; potential to address therapeutic index

- Friedlaender A et al. healthbook TIMES Onco Hema (2023)
   Sartor et al. N Eng J Med (2021)
   Tran et al. Ann Onc. (2020)
   Bendell et al. JCO (2020)

- 5. Slovin et al. JCO (2022) 6. Narayan et al. Nat Med (2022) 7. Mirzaei et al. Int J Biol Macromol



# Potential Near-Term Milestones

ADI-001 Autoimmune Diseases

> ADI-001 MCL

ADI-270 RCC

20	24		2025			
1H	2H		1H		2H	
Initiate LN Phase 1	Phase 1 Da		ary Clinical a in LN  ry Clinical Data in 1-2 additional Al Indications		Clinical Data in 3 Al	
					indications	
Study Enrollment Ongoing	Clinical U	pdate		egulatory Path for y Pivotal Phase 2 Study	Clinical Update	
ADI-270 IND Submission			Potentia	Data RCC and al Expansion to al CD70+ Tumors	Clinical Update	

\$183.3 in cash and cash equivalents as of 9/30/23 Projected cash runway into 2H 2025

Subject to data readouts and regulatory feedback

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