



# Leaders in Developing Allogeneic CAR $\gamma\delta$ 1 T Cell Therapies to Fight Autoimmune Diseases and Cancer



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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 our preclinical and clinical studies, business operations, employee hiring and retention, 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 preclinical or clinical study may not necessarily be predictive of the results of future or ongoing studies; clinical studies may fail to demonstrate adequate safety and efficacy of Adicet's product candidates, which would prevent, delay, or limit the scope of regulatory approval and commercialization; and regulatory approval processes of the FDA and comparable foreign regulatory authorities are lengthy, time-consuming, and inherently unpredictable; and Adicet's ability to meet production and product release expectations. For a discussion of these and other risks and uncertainties, and other important factors, any of which could cause Adicet's actual results to differ from those contained in the forward-looking statements, see the section entitled "Risk Factors" in Adicet's most recent annual report on Form 10-K and our periodic reports on Form 10-Q and Form 8-K filed with the U.S. Securities and Exchange Commission (SEC), as well as discussions of potential risks, uncertainties, and other important factors in Adicet's other filings with the SEC.

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# Developing Broad Pipeline of Allogeneic $\gamma\delta 1$ T Cell Therapies for Autoimmune Diseases and Cancer

Program	Target	Potential Diseases	Research	IND-Enabling	Clinical	Status
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## ADICET WHOLLY OWNED PROGRAMS

<b>ADI-001</b>	CD20	Autoimmune	●	●	●	IND in LN cleared Dec 2023 Initiate LN Phase 1 2Q 2024 Update planned 2H 2024 AI expansion opportunities
<b>ADI-001</b>	CD20	NHL	●	●	●	MCL Phase 1 ongoing* Update planned 2H 2024
<b>ADI-270</b>	CD70 (TGF $\beta$ -DNR)	RCC & Other ST / Heme	●	●	○	IND submission in RCC expected 2Q 2024
<b>ADI-xxx</b>	PSMA (w/ Armor)	mCRPC	●	○	○	Preclinical activities
<b>ADI-925</b>	Tumor stress ligands	Multiple Solid / Heme	●	○	○	Preclinical activities

## PARTNERED PROGRAMS

<b>ADI-002</b>	GPC3	HCC	●	●	○	<b>REGENERON</b>
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\*Adicet is focused on advancing MCL enrollment in the GLEAN trial and has deprioritized enrolling large B-cell lymphoma patients.

AI=Autoimmune; GPC3= Glypican-3; HCC= Hepatocellular carcinoma; IND= Investigational new drug; LN= Lupus nephritis; mCRPC= Metastatic castration-resistant prostate cancer; MCL= Mantle cell lymphoma; NHL= Non-Hodgkin's lymphoma; PSMA= Prostate specific membrane antigen; RCC= Renal cell carcinoma; ST= Solid tumor

# Adicet Bio Leadership Team



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Chief Technology  
Officer



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Chief Scientific Officer



Nick Harvey  
Chief Financial Officer



Francesco Galimi, M.D., Ph.D.  
Chief Medical Officer



Amy Locke  
Head of Human Resources



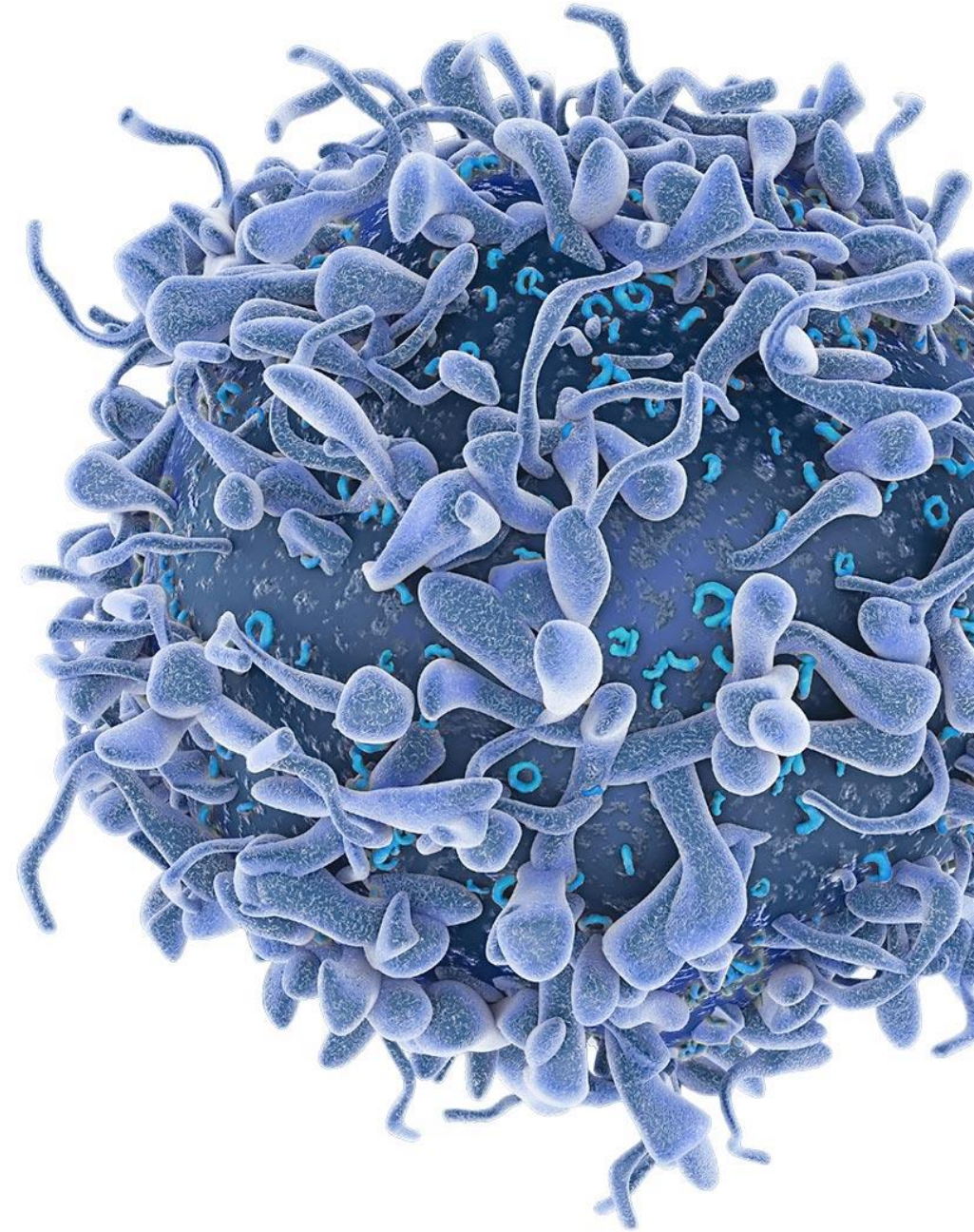
Nancy Boman, M.D., Ph.D.  
Chief Regulatory Officer





# ADI-001

## Autoimmune Diseases



# Adicet $\gamma\delta 1$ CAR T Cell Therapy For Autoimmune Indications

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

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

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

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

Readily Available, "Off-the-Shelf"

Preferentially Trafficking to Organs/ Tissues

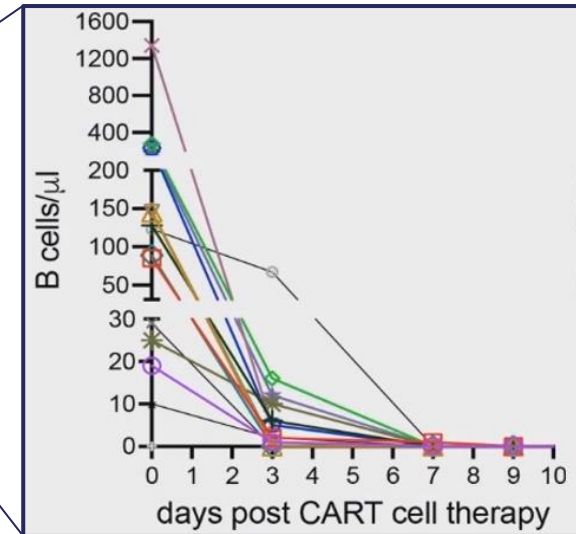
Potential to Dose in Community Setting



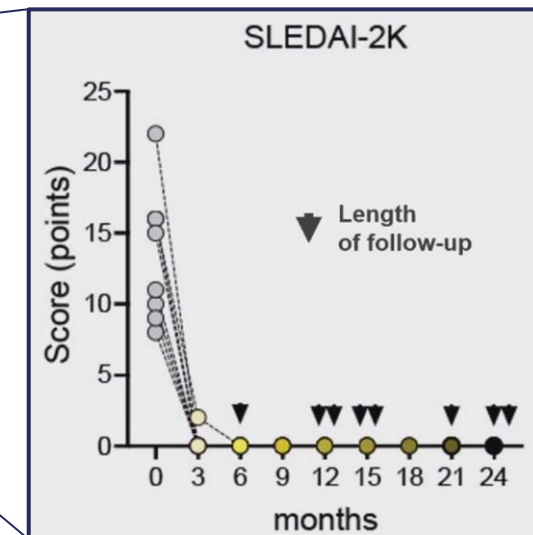
AUC= Area Under the Curve; Cmax= Peak plasma concentration; CRS= Cytokine release syndrome; ICANS= Immune effector cell-associated neurotoxicity syndrome; IIM= idiopathic inflammatory myopathy; SLE= systemic lupus erythematosus; SSC= systemic sclerosis

# CAR-T Therapy Depleted B-cells, Drove an Immune Reset, and Achieved Treatment-Free Remissions in Patients with Autoimmune Diseases<sup>1</sup>

- Schett et al. treated 15 patients with autoimmune 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 autoimmune 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

1. Mackensen A et al. Nature Medicine 2022  
 2. CD19.CAR-T Cell in Refractory Systemic Autoimmune Diseases: A Monocentric Experience from the First Fifteen Patients. ASH 2023  
 3. YTB323 Poster @ American College of Rheumatology Convergence November 2023  
 4. Haghikia A et al. Lancet Neurology 2023  
 5. Granit V et al. Lancet Neurology (2023)

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

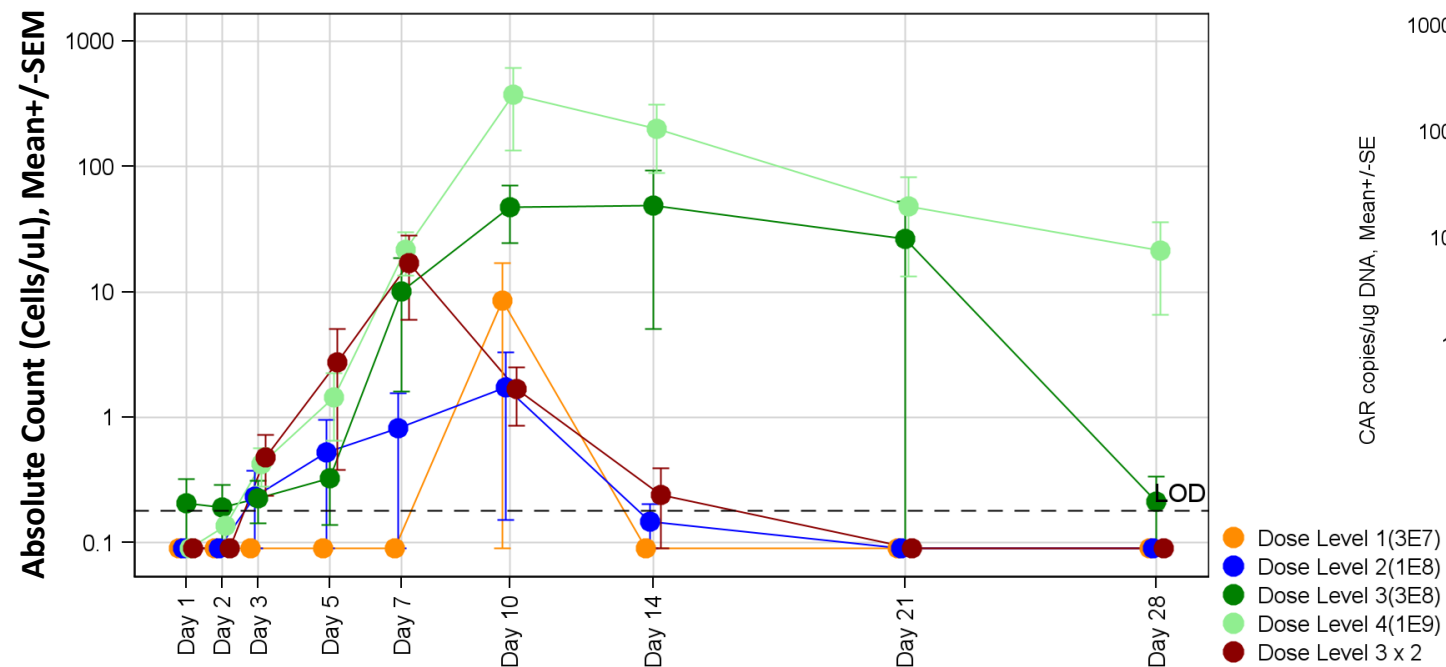
Lupus nephritis ~150K	SLE ~160-410K	MS >900K	Moderate to severe RA >1M
Myasthenia gravis ~36-60K	ANCA+ vasculitis ~40-75K	Sjogren's ~260K	Membranous nephropathy ~70-90K
NMOSD ~20K	Pemphigus vulgaris ~19K	Systemic sclerosis ~68K	IIM ~60-70K

\*Not an exhaustive list

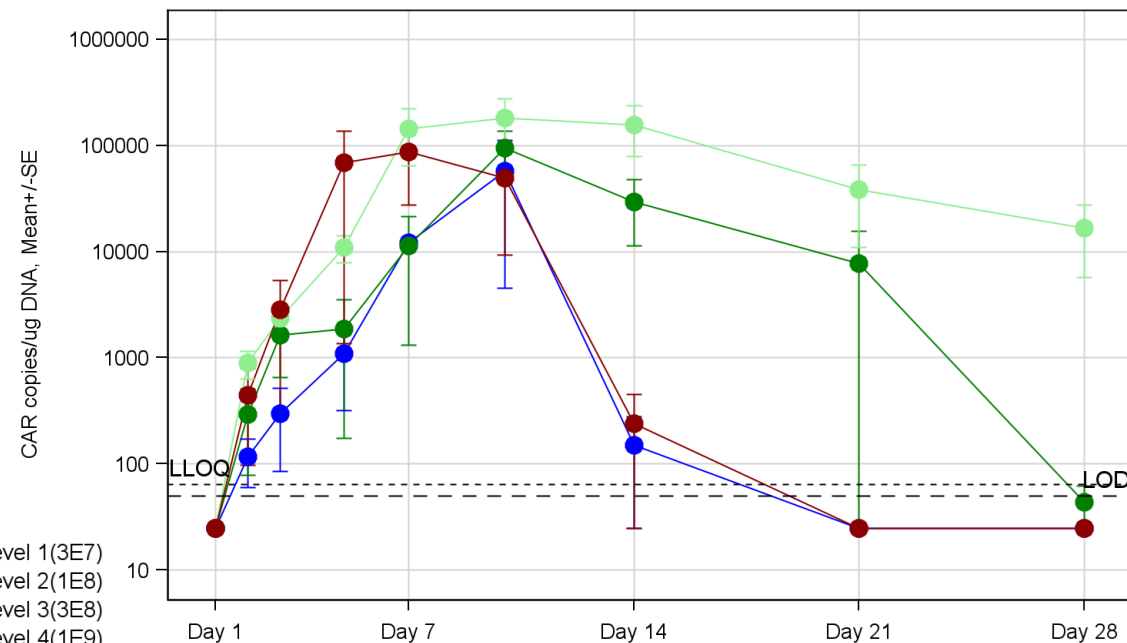


# Cmax, D28 Persistence and AUC Consistent Values Reported for Approved Autologous CD19 CAR T<sup>1</sup>

## ADI-001 CAR by Flow Cytometry

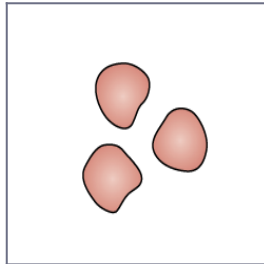


## ADI-001 CAR by ddPCR



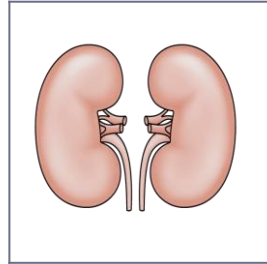
Dose Level	Mean Cmax		Mean D28	
	CAR+ Vd1 cells/ul	Copies/ug	CAR+ Vd1 cells/ul	Copies/ug
DL4	363.80	201,666	26.51	16,553
DL3	56.34	98,177	0.04	44

# GD1 T Cells Preferentially Traffick to Organs & Tissues: Attacking a Source of Resistance to Antibody Therapies



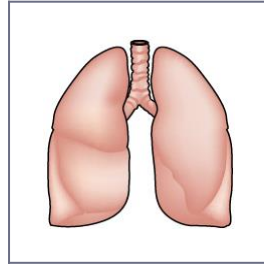
**lymph node**<sup>3,4</sup>

CD27+  
CD62L+  
Vδ1+ ↑↑  
Vδ2+ ↓↓



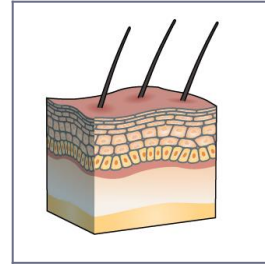
**kidney**<sup>1</sup>

tissue: **>3X**  
γδ vs αβ  
~**3X** more  
Vδ1 vs  
Vδ2+



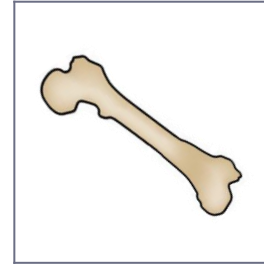
**lung**<sup>9</sup>

issue/blood:  
**9X**



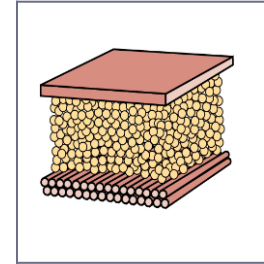
**skin**<sup>8</sup>

tissue/blood:  
**8X**



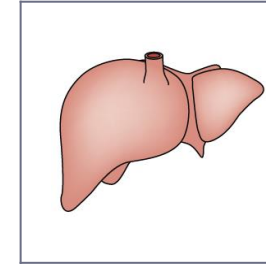
**bone marrow**<sup>2</sup>

tissue/blood:  
**4X**



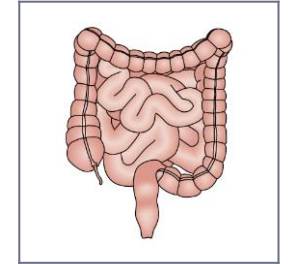
**Breast**<sup>5</sup>

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



**liver**<sup>7</sup>

tissue/blood:  
**3X**



**GI**<sup>6</sup>

tissue/blood:  
**11X**

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:

<sup>1</sup>Rancan *et al Nat Immunol* 2023

<sup>2</sup>Brauneck *et al Front Med* 2021

<sup>3</sup>Davey *et al Trends Immunol* 2018

<sup>4</sup>Uger *et al Sci Rep* 2018

<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

<sup>10</sup>Wang Q. *et al Exp Ther Med* 2020

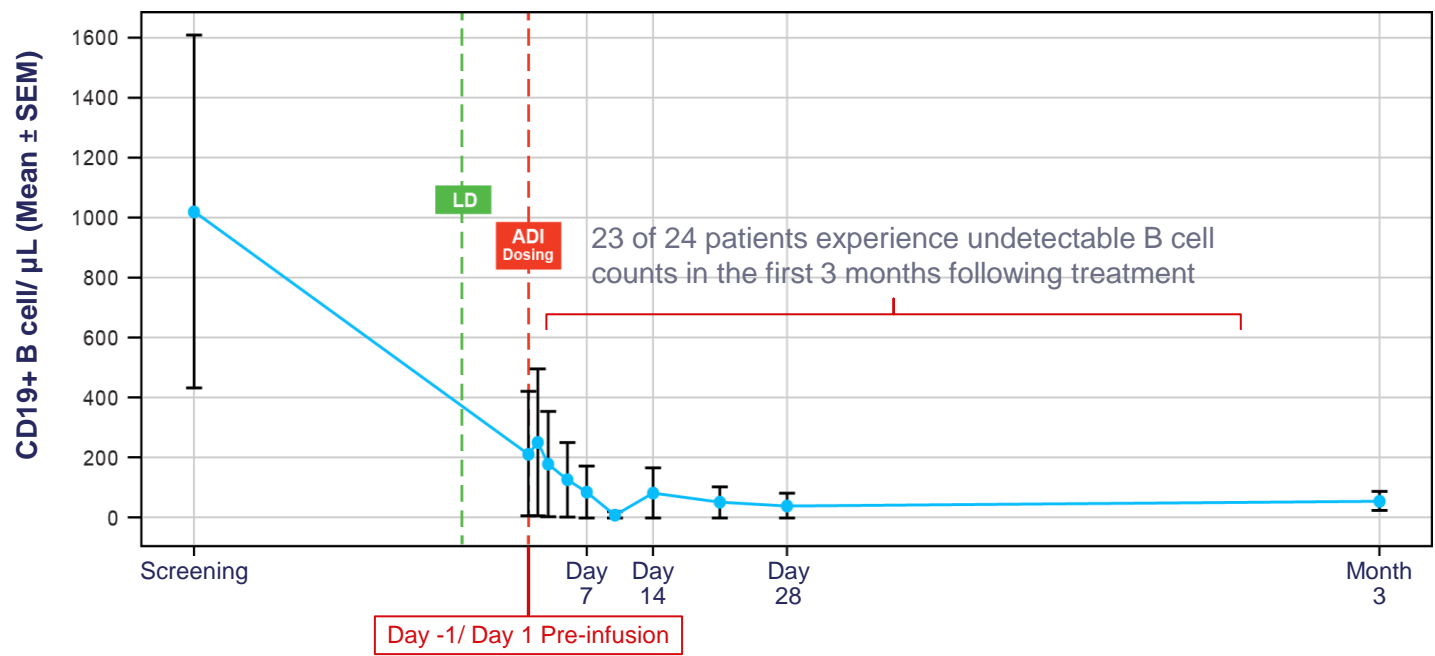
<sup>11</sup>Déchanet *et al J Infect Dis* 1999

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

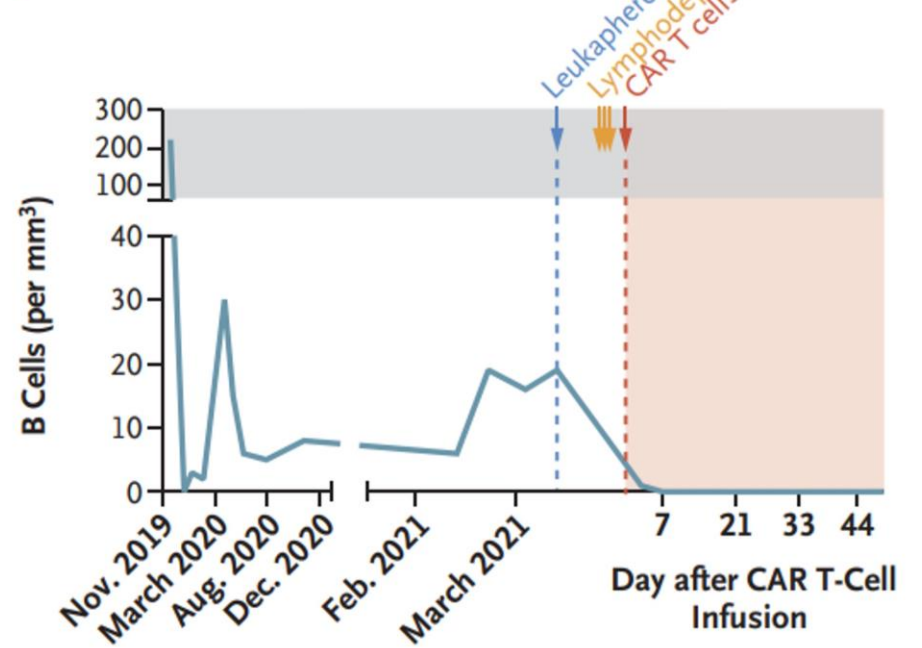
## B-cell Depletion

- B-cell depletion data from ADI-001 trial in NHL mirrored experience of autologous CD19 CAR T in SLE<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 a Phase II clinical study<sup>3</sup>

**Adicet Bio** CD20-targeted, ADI-001, in B-NHL patients<sup>4</sup>



CD19-targeted CAR-T in SLE patients<sup>1</sup>



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

SOC= Standard of care

# ADI-001 in Autoimmune 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 study<sup>3</sup>

### $\gamma\delta$ 1 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>
- $\gamma\delta$ 1 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-the-shelf”

- 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

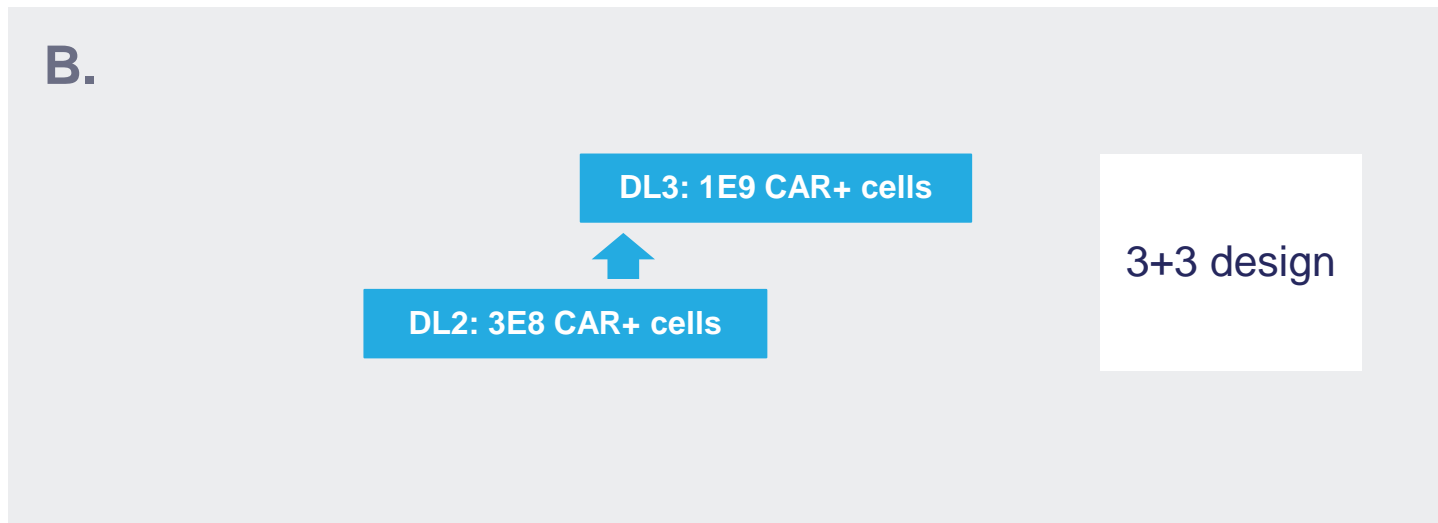
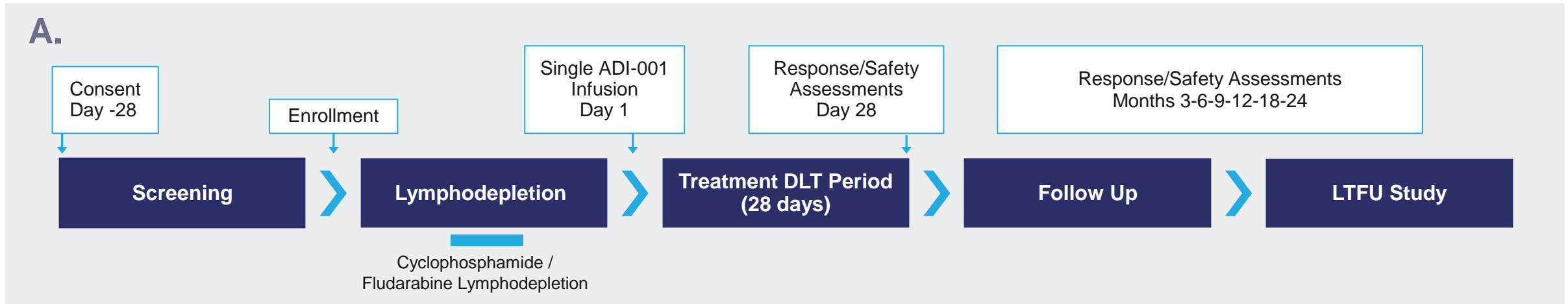
1. Mouggiakakos MD et al. NEJM 2021  
2. Mackensen A et al. Nature Medicine 2022  
3. Furie RA et al. Ann Rheum Dis. 2022  
4. Kamburova EG et al. American Journal of Transplantation 2013

5. Reddy VK et al. Rheumatology 2022  
6. Sadun RE and Foster MH AJKD 2019  
7. Zhang, PJ Hematol Oncol 2023

COGs= Cost of goods

\* ADI-001 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

# 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

# ADI-001: Lupus Nephritis

## Opportunity

- Type of kidney disease caused by SLE, an autoimmune 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 AI indications

1. 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.

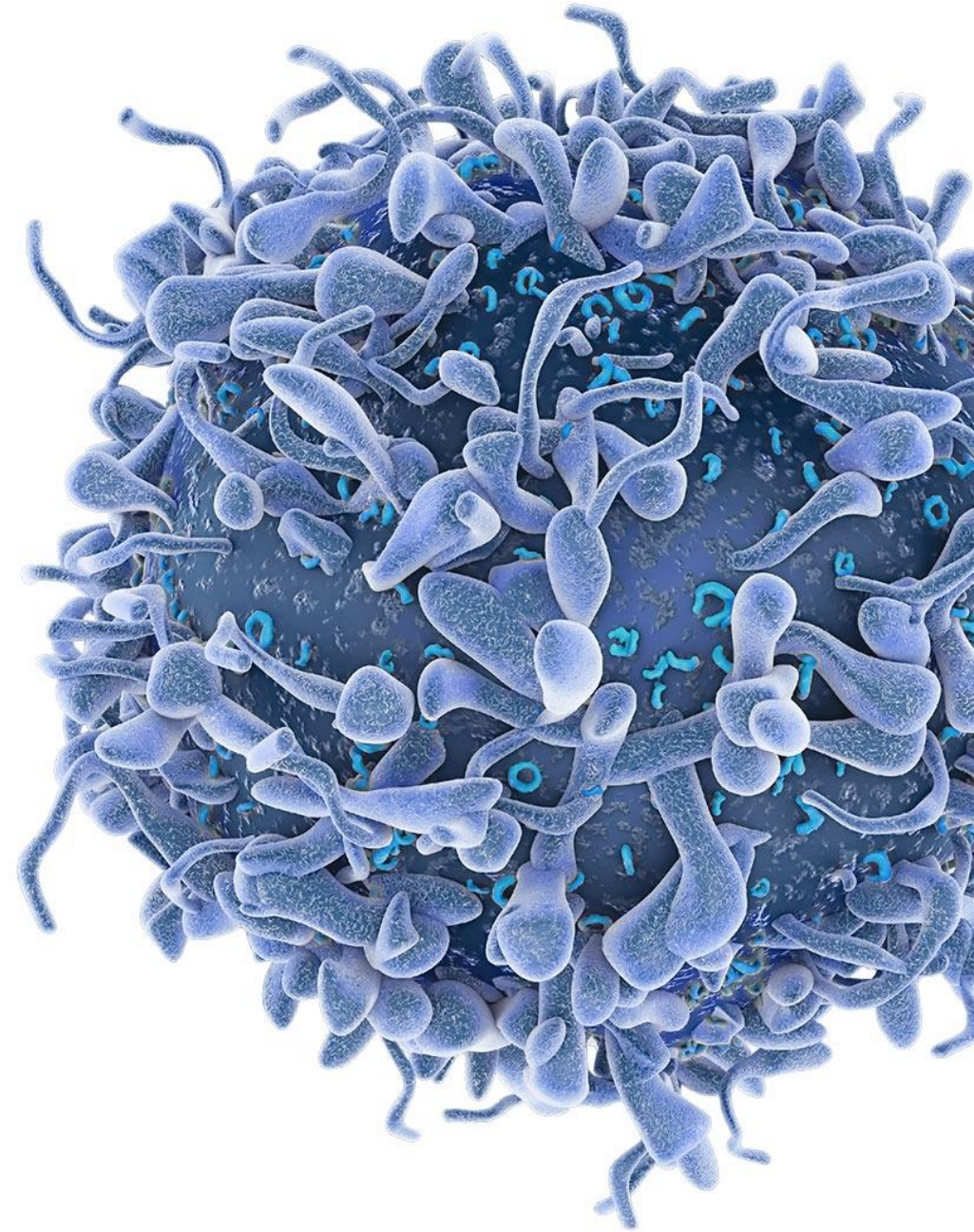
2. Hoover PJ et al.; Kidney Int 2016

3. 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 Thrombocytopenia Neurological damage Affective disorder Lung Inflammation Joints Arthritis." (2014).



# ADI-001

## Mantle Cell Lymphoma

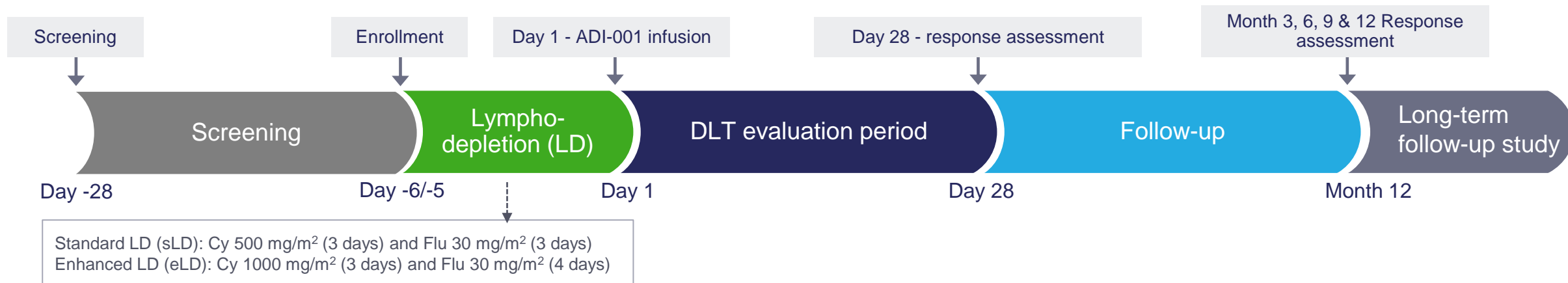




# ADI-001: Encouraging Early Efficacy Data in Heavily Pretreated NHL Patients

- 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  $\gamma\delta 1$  T cell therapies

# GLEAN: ADI-001 First-in-Human Study (CD20 CAR+ $\gamma\delta$ 1 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

\*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)
<b>Age – median (range)</b>	66.5 (44 - 75)
Sex – number of male	17 (70.8)
<b>B cell malignancy (WHO 2017 classification)</b>	
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)
<b>IPI score (LBCL) - median (range)</b>	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)
<b>Sum of the product of the diameters at screening - median (range)</b>	3001 (150 - 7919) mm <sup>2</sup>
Prior lines of therapies - median (range)	4 (2 - 9)
<b>Prior anti-CD19 CAR T therapies</b>	12 (50.0)
<b>Prior systemic anti-cancer therapy</b>	
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)
<b>Refractory status at study entry</b>	
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	13 (54.2)

- 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

# ADI-001: Efficacy Summary by Dose Level

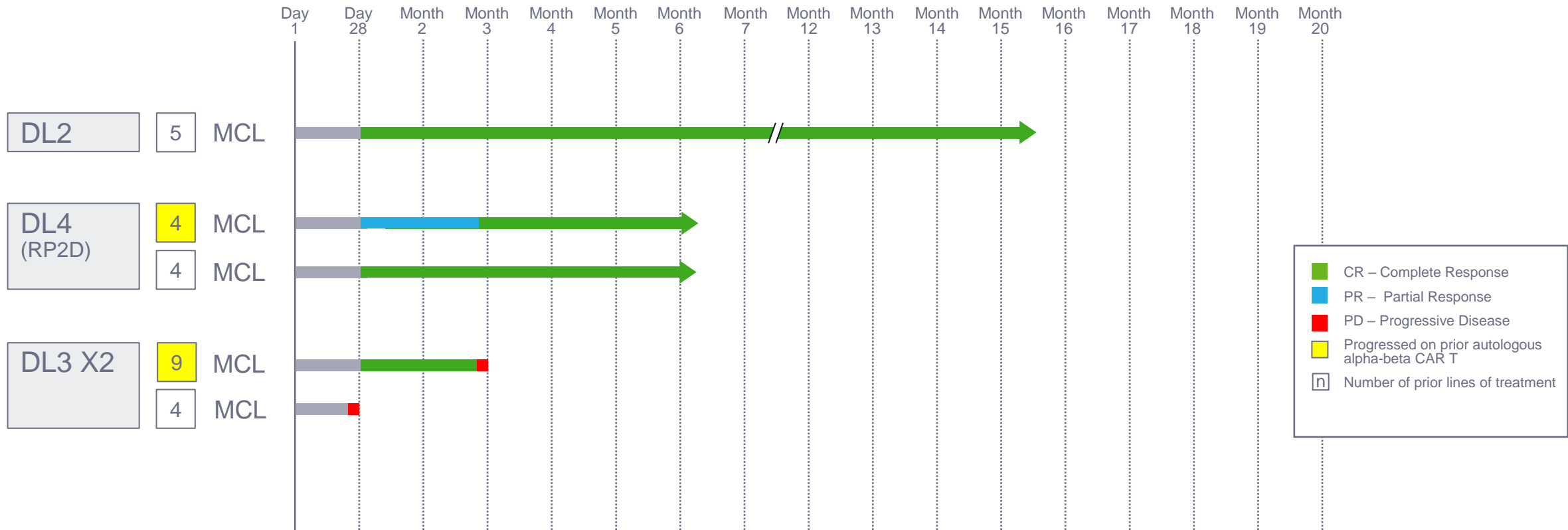
	Median No. of Prior Lines	ORR (%)	CR Rate (%)	6-month CR Rate (%)
DL4 (RP2D; LBCL & MCL)	4	6/8 (75.0%)	5/8 (62.5%)	2/8 (25.0%)
<b>DL 4 MCL</b>	<b>4</b>	<b>2/2 (100.0%)</b>	<b>2/2 (100%)</b>	<b>2/2 (100%)</b>
All Doses (LBCL & MCL)	4	17/24 (70.8%)	15/24 (62.5%)	4/24 (16.7%)
<b>All Doses MCL</b>	<b>4</b>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

**High CR rate and favorable durability in MCL**

# ADI-001: Safety Data in Efficacy Evaluable Patients

	DL1 (N=3)		DL2 (N=3)		DL3 (N=6)		DL3 X2 (N=4)		DL4 (N=8)		Total (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
<b>CRS</b>	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%)
<b>ICANS</b>	0	0	1 (33.3%)	0	0	0	1 (25.0%)	1 (25.0%)	1 (12.5%)	0	3 (12.5%)	1 (4.2%)
<b>GvHD</b>	0	0	0	0	0	0	0	0	0	0	0	0
<b>DLT</b>	0	0	0	0	0	0	0	0	0	0	0	0
<b>Infection</b>	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%)
<b>SAE-TEAE</b>	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%)
<b>Related SAE-TEAE</b>	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%)

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



**ORR: 80%, CR rate: 80%, 6-month CR rate: 60%**

**No significant CRS or ICANS**

# 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 T<sup>1</sup>
- ✓ C<sub>max</sub>, 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

# 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 TGF $\beta$  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 castration-resistant prostate cancer program

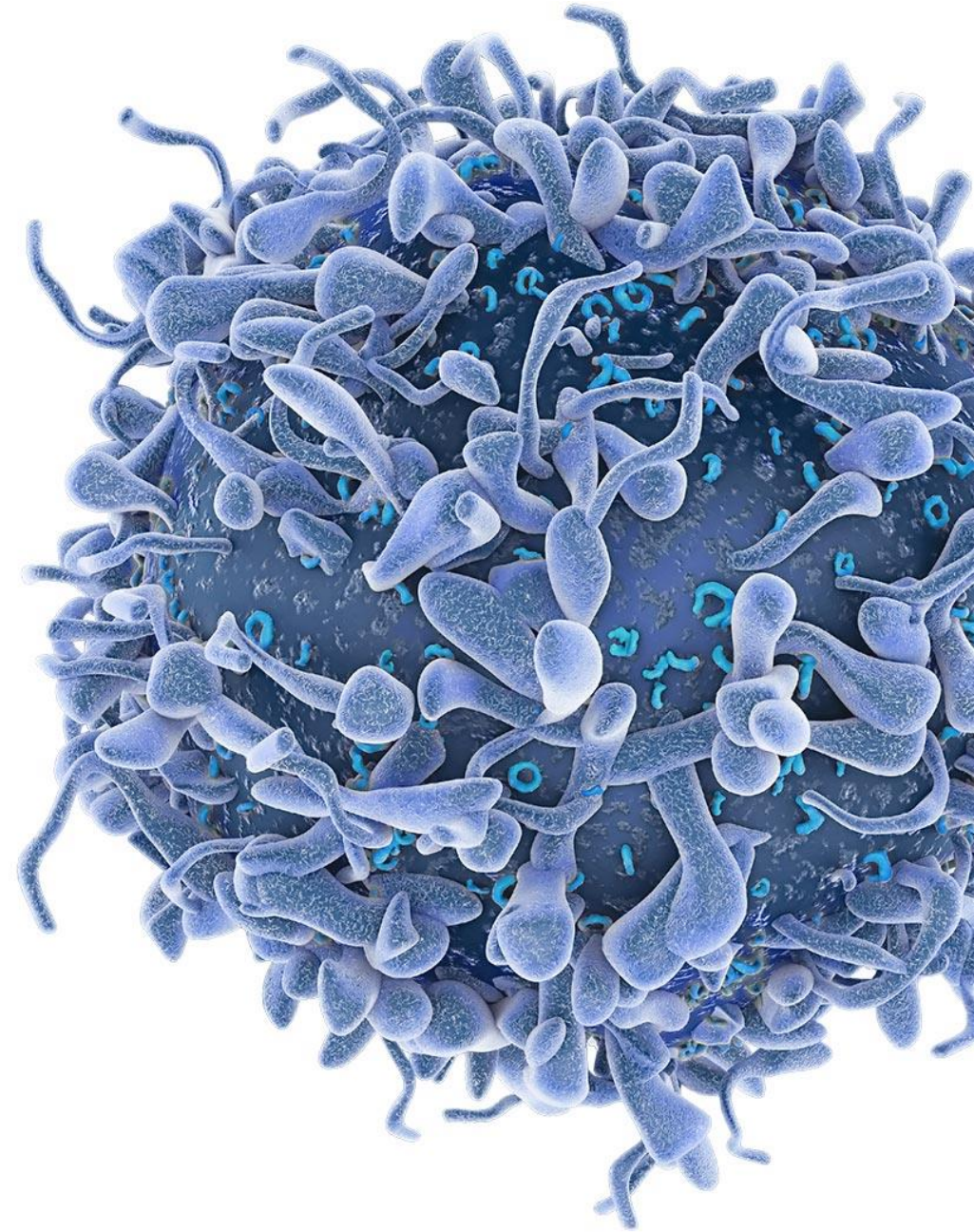
- Adicet lead scFv antibody designed to have similar binding determinants as approved radioligand therapy
- Highly potent cell therapy with armoring intended to address immunosuppressive solid tumor environment & heterogeneous PSMA expression
- Leveraging  $\gamma\delta$ 1 T cell tropism to solid tissues
- Multiple modes of cell killing: CAR mediated, innate and adaptive immunity



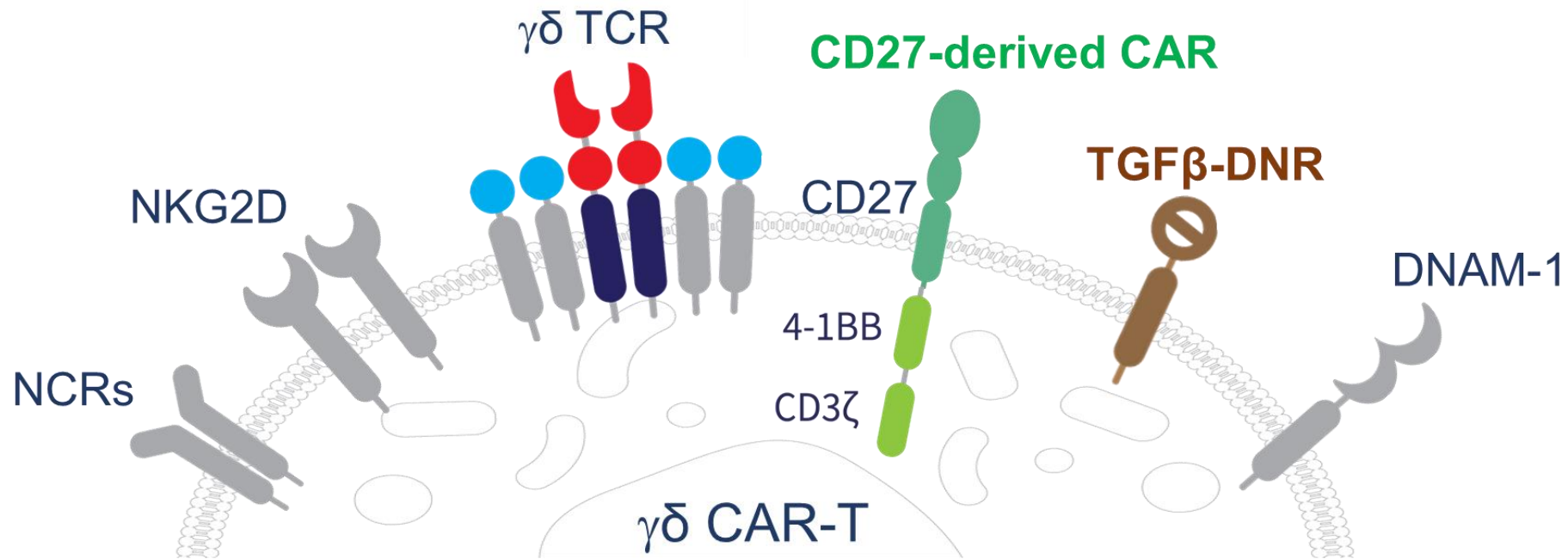


# ADI-270

## Renal Cell Carcinoma & Other CD70+ Diseases



# ADI-270: Designed to address multiple refractory cancers

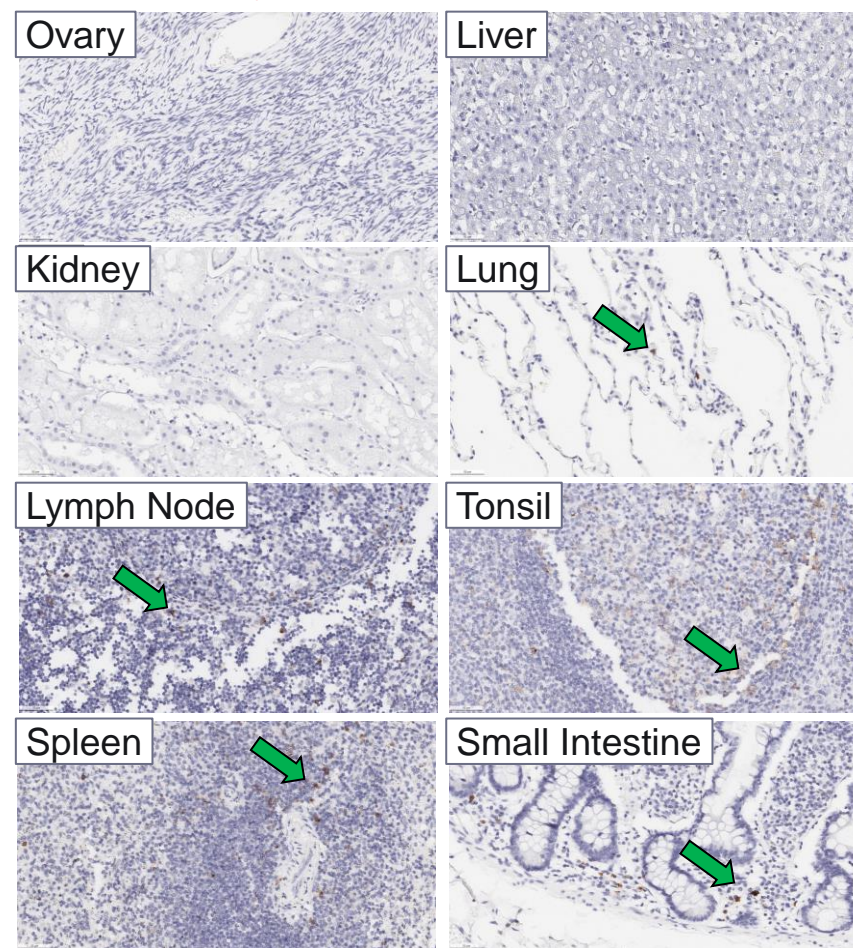


- CAR utilizes CD27 as the binding domain and contains CD27 and 4-1BB costimulatory domains plus CD3 $\zeta$  (3<sup>rd</sup> gen)
- Inactive form of TGF $\beta$  receptor II to mitigate the immunosuppressive effects of TGF $\beta$  within the tumor microenvironment
- Host vs graft armoring against alloreactive activated CD70+ T cells to increase persistence
- Combines endogenous  $\gamma\delta$  innate and adaptive mechanisms to recognize and kill malignant cells

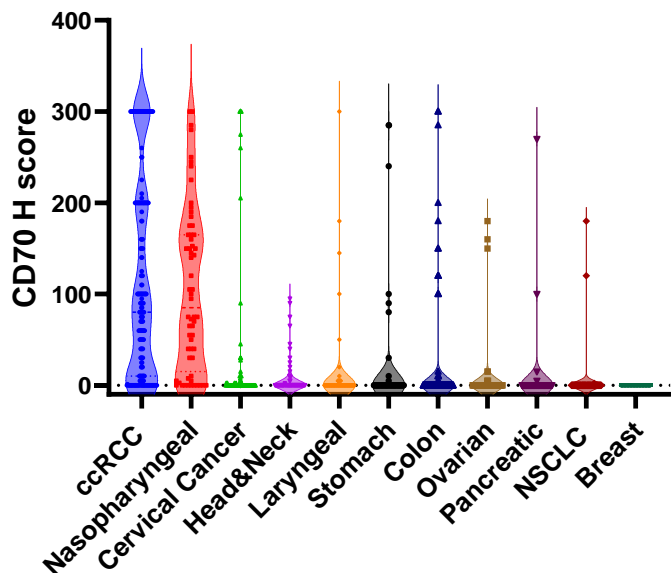
# CD70 is expressed on multiple solid and hematological cancers with limited expression in normal tissues

- High expression in multiple solid and heme malignancies
  - Beyond ccRCC and NPC, multiple solid tumors are of interest when paired with CD70 screening
- Minimal expression on normal tissues (activated lymphocytes)
- Target has clinical safety experience

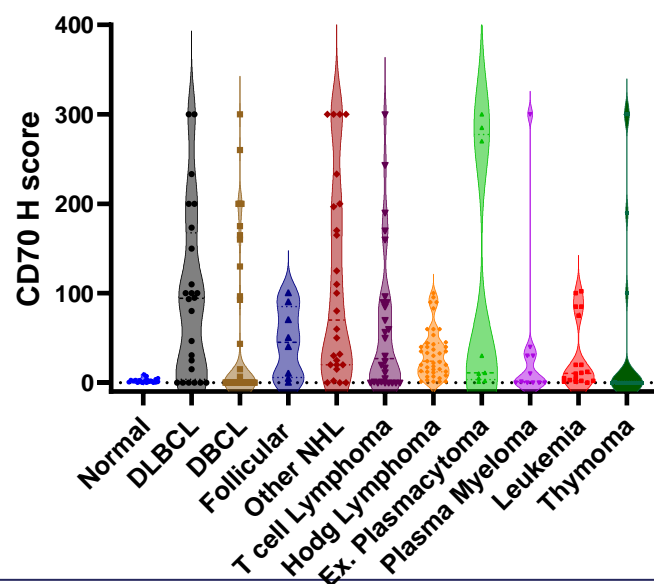
Representative images from a normal tissue array stained for CD70



Solid tumors

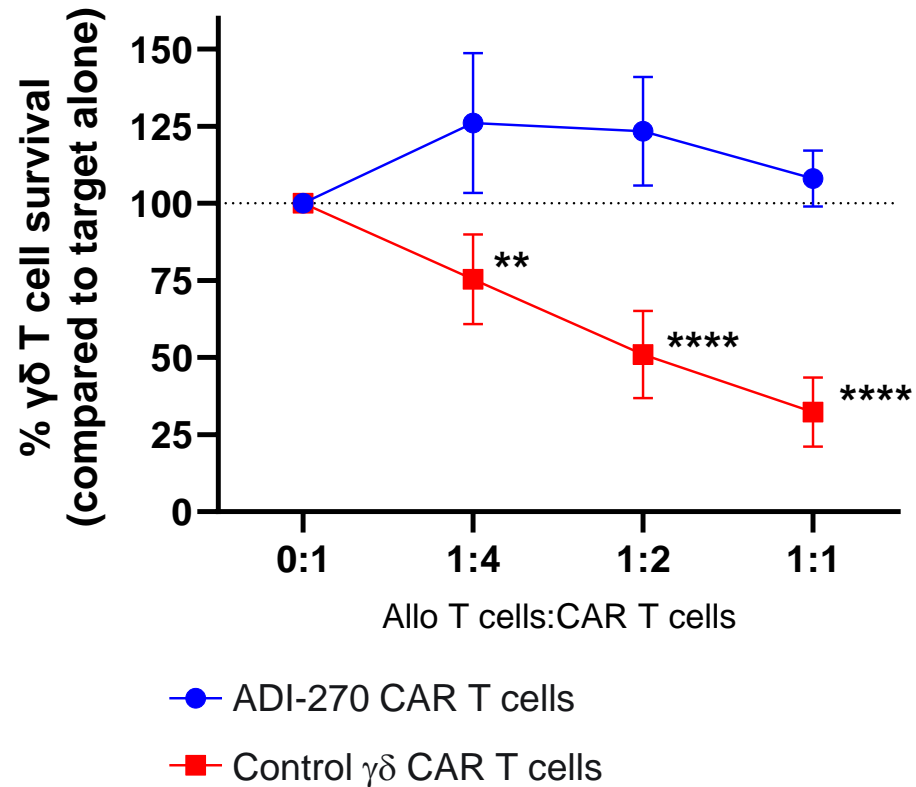


Hematological malignancies

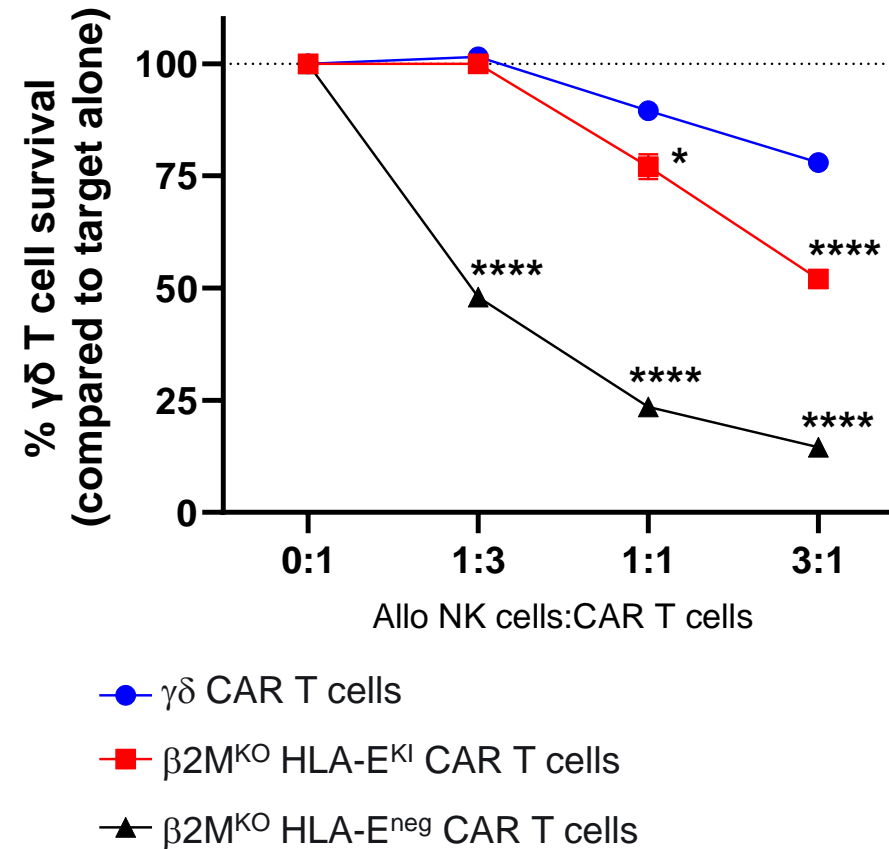


# ADI-270 may be less susceptible to T and NK rejection by host

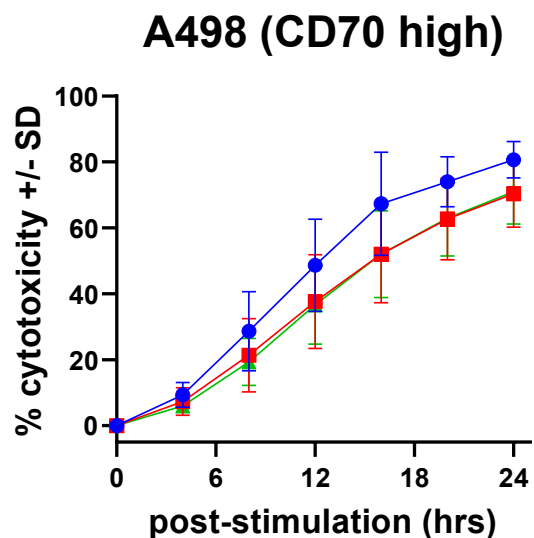
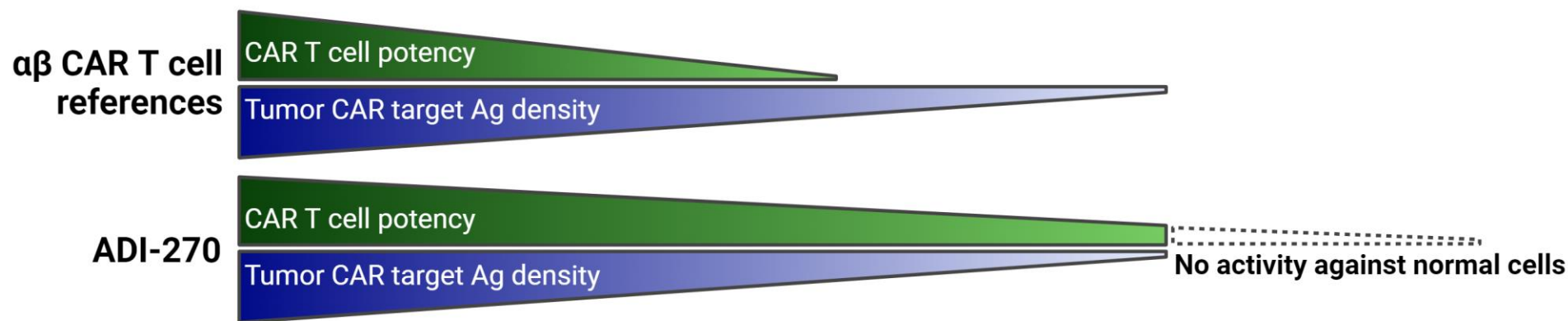
## CD70 targeting less susceptible to T cell rejection



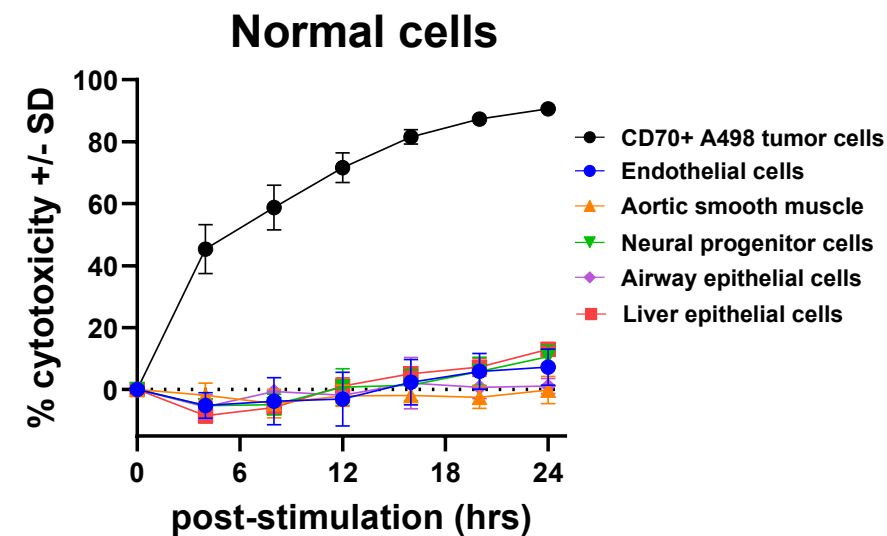
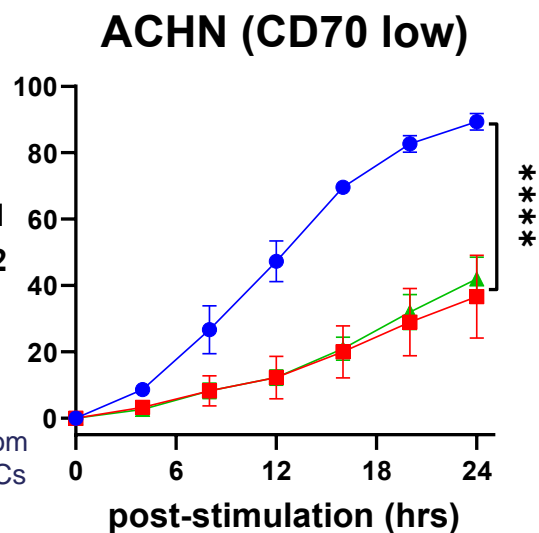
## $\gamma\delta$ CAR T cells less susceptible to NK rejection



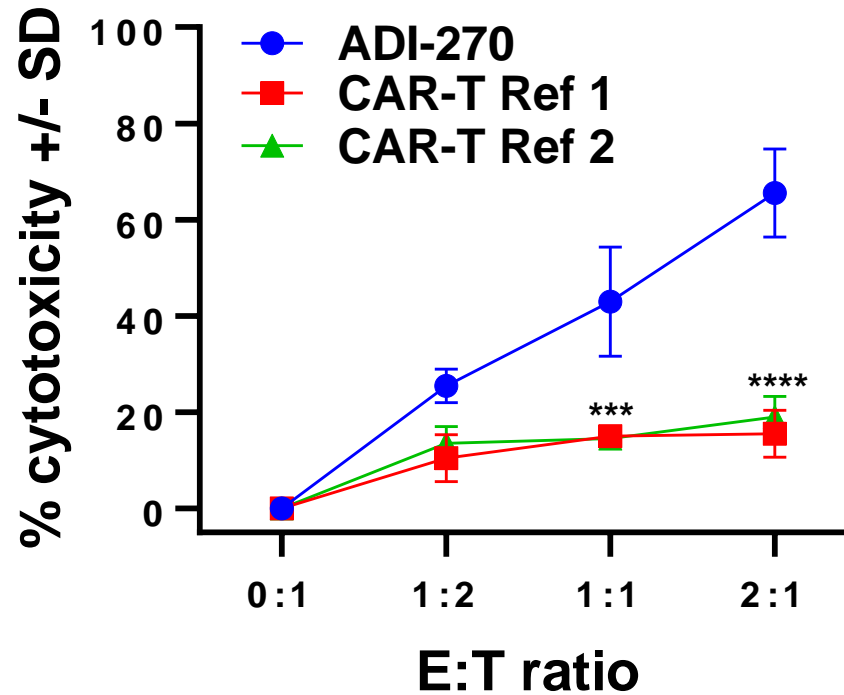
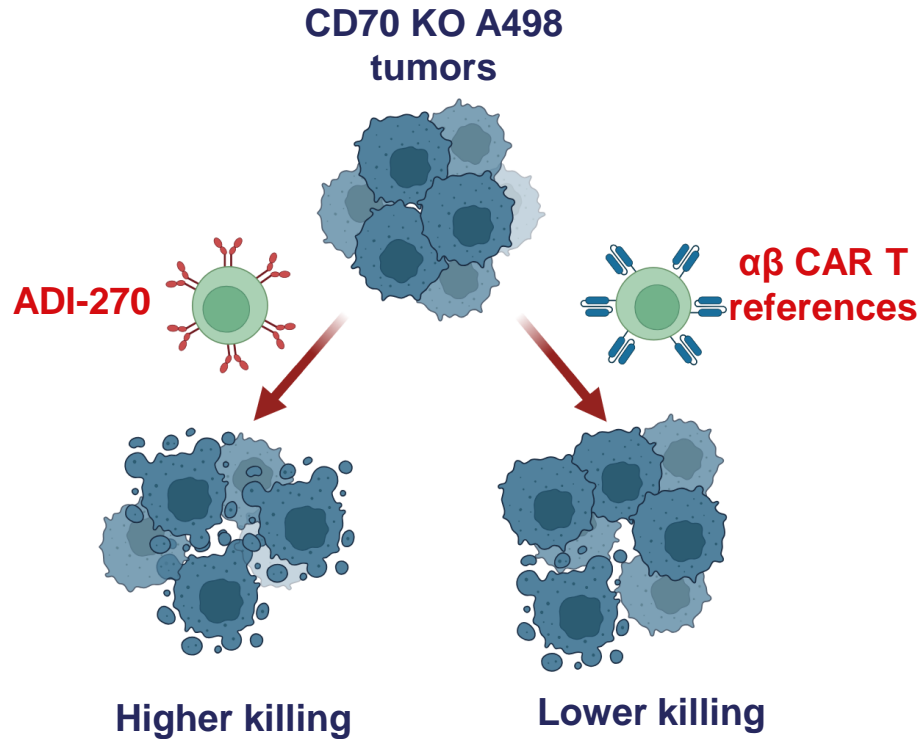
# ADI-270 retained potent activity in the context of CD70-low tumors compared to clinically relevant CD70-targeting $\alpha\beta$ CAR T cell benchmarks



● ADI-270  
 ■  $\alpha\beta$  CAR-T Ref 1  
 ▲  $\alpha\beta$  CAR-T Ref 2  
 E:T ratio = 1:1  
 N=3 donors  
 \*\*\*\* $p < 0.0001$   
 test materials derived from same set of donor PBMCs



# ADI-270 demonstrated higher innate cytolytic activity against CD70 negative tumor cells compared to CAR-T cell references

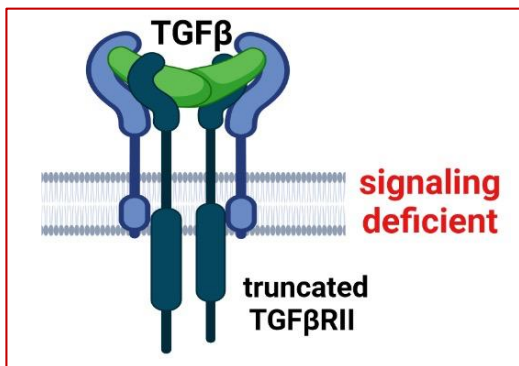
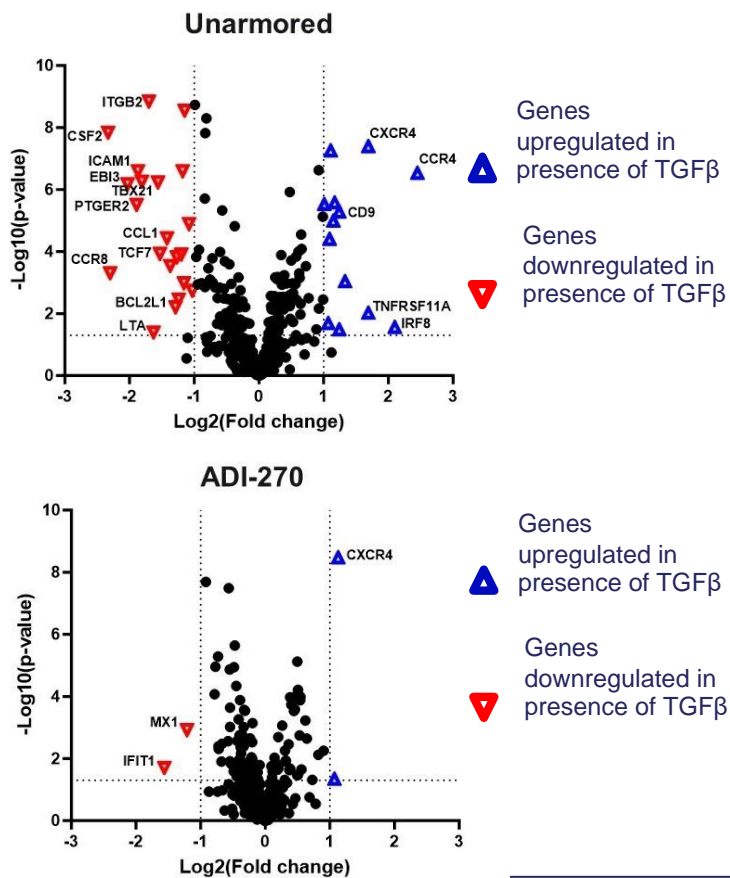


\*\*\* $p < 0.001$ , \*\*\*\* $p < 0.0001$

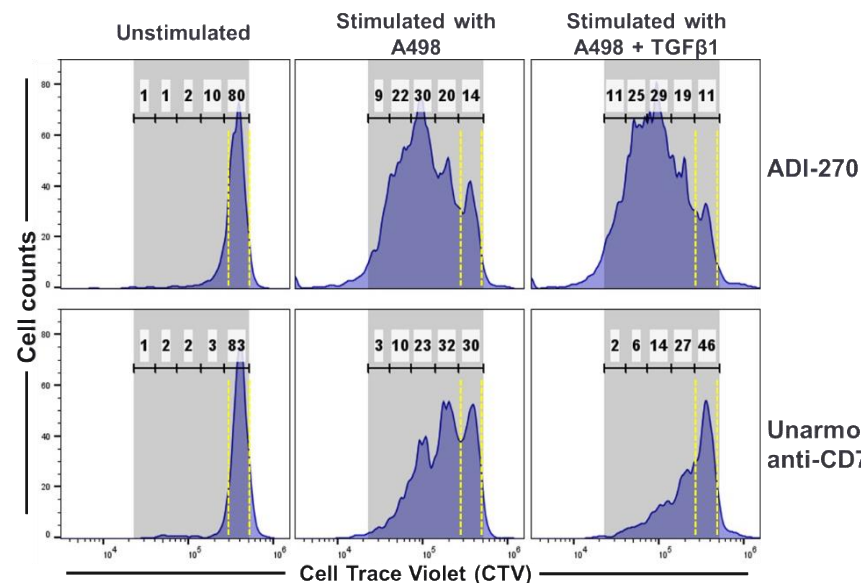
test materials derived from same donor PBMCs

# ADI-270 is resilient to the inhibitory effects of TGFβ

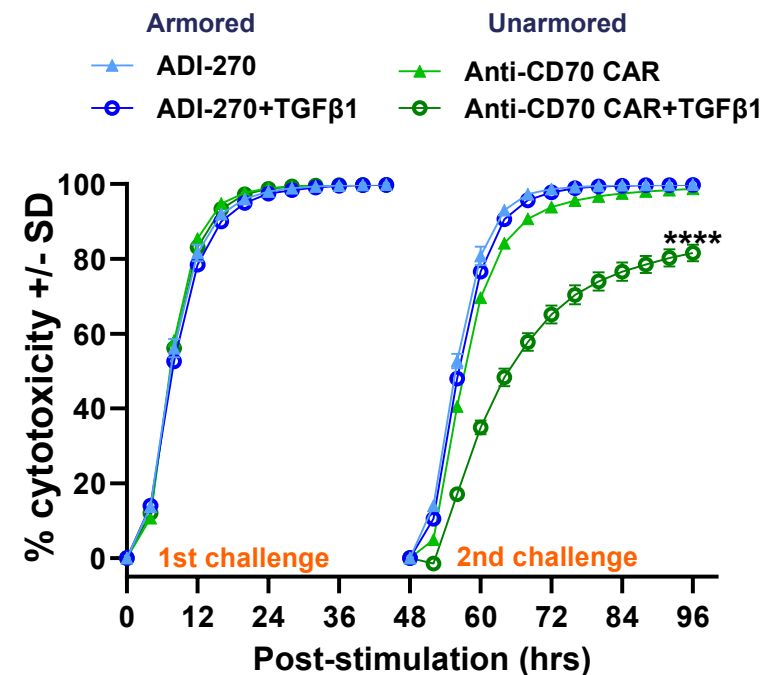
ADI-270 showed resilience to transcriptional changes driven by TGFβ signaling



ADI-270 maintained proliferation in the presence of TGFβ

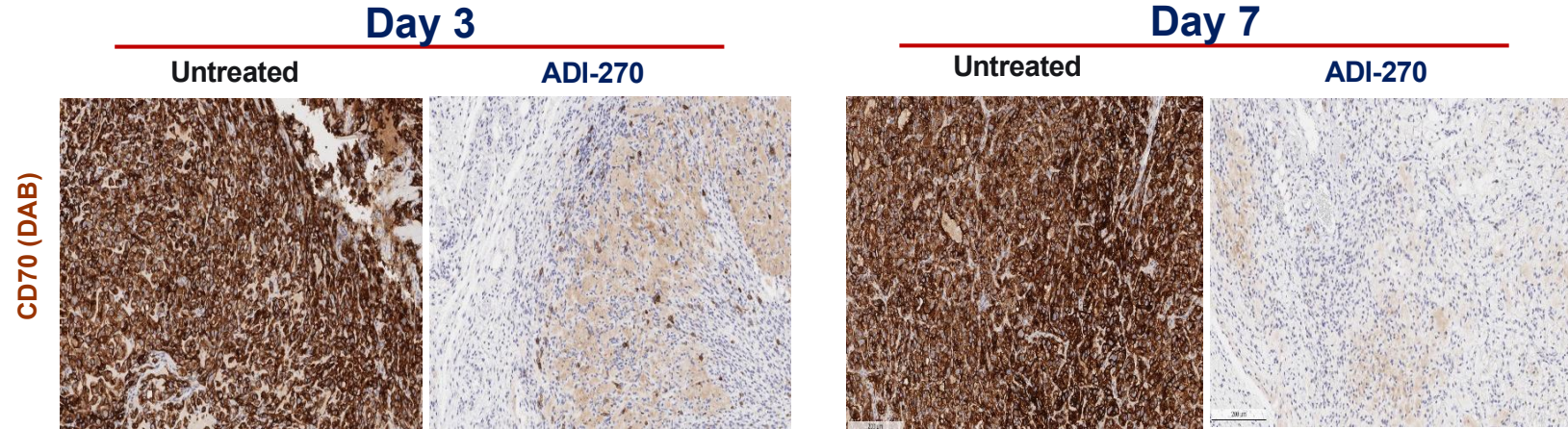
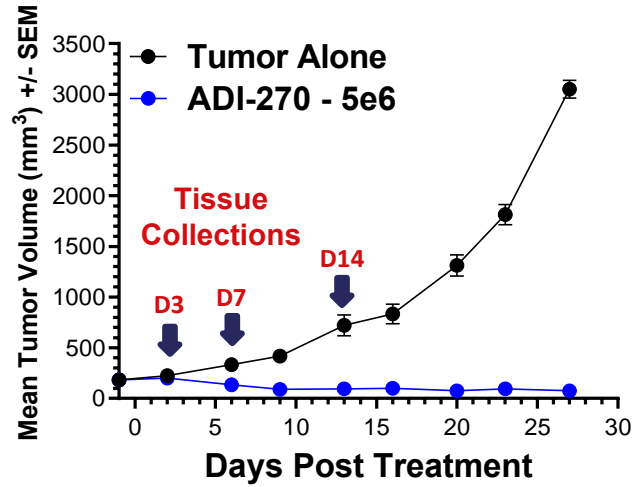


ADI-270 maintained cytotoxicity in the presence of TGFβ

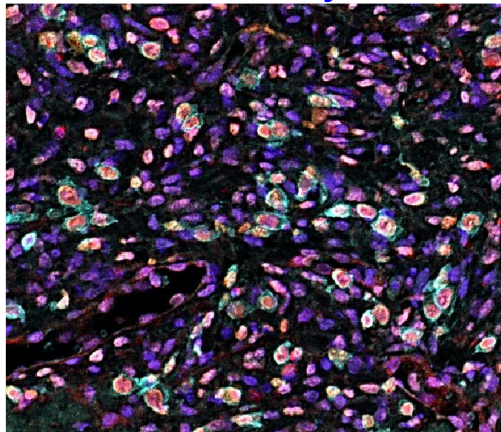


# ADI-270 demonstrated rapid homing, activation and killing kinetics in ccRCC xenografts resulting in tumor and target eradication

A single dose of ADI-270 showed potent efficacy in A498 tumors, rapidly eradicating CD70+ cells

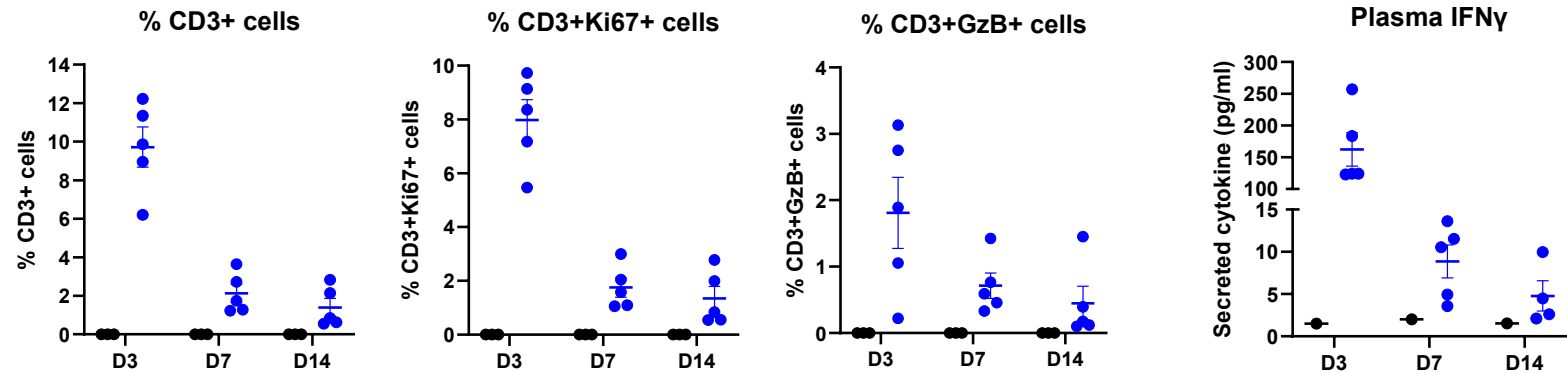


ADI-270 Day 3



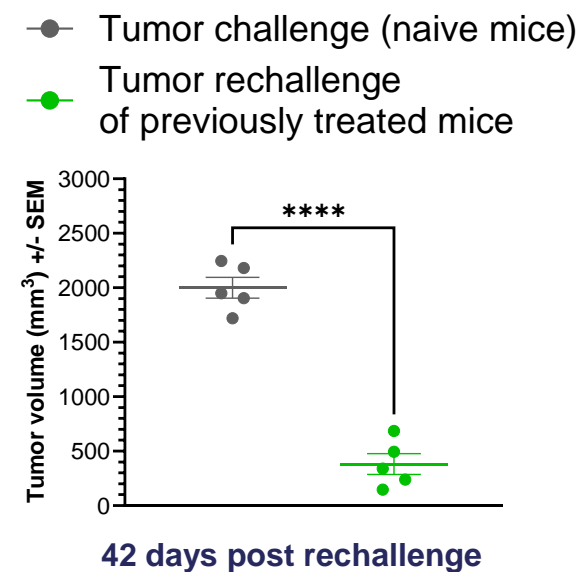
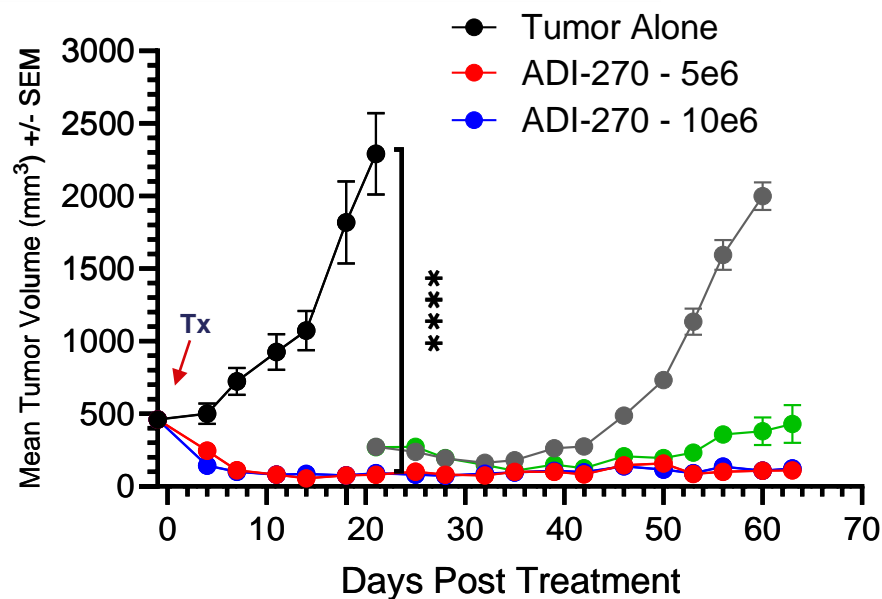
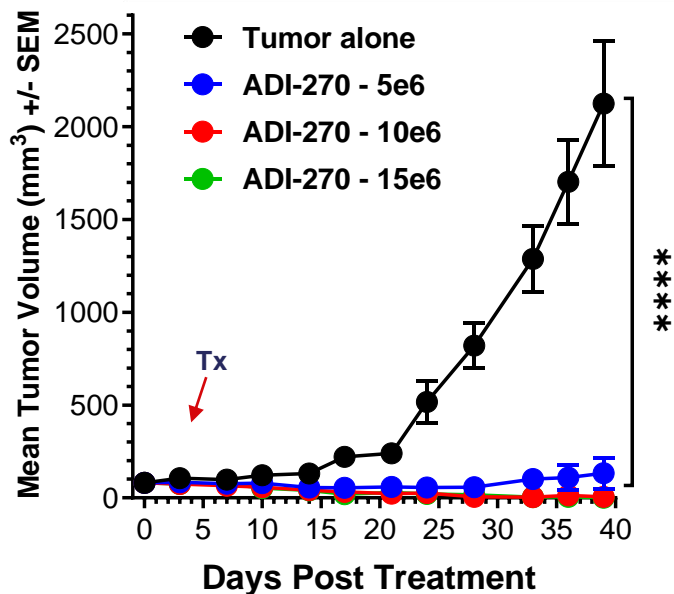
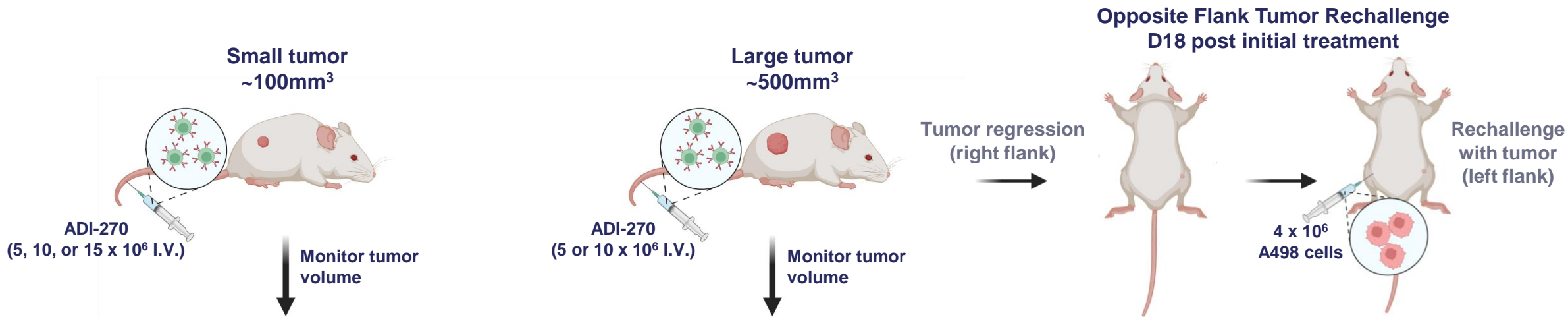
DAPI + CD3 (cyan) + Ki67 (orange)  
+ Granzyme B (red)

ADI-270 infiltrated and proliferated with effector function as early as Day 3





# A single dose of ADI-270 showed potent regression and sustained systemic anti-tumor activity in ccRCC xenograft models



\*\*\*\*p<0.0001

# Next steps: ADI-270

- ADI-270 represents potential evolution of  $\gamma\delta$  CAR T-cell based therapeutics
- CD27-based 3<sup>rd</sup> gen CAR demonstrated significant potency advantages<sup>1,2,3,4</sup>
- Armoring against TGF $\beta$  and alloreactive T cells confirmed and characterized preclinically
- Robust efficacy maintained across multiple relevant tumor models of varying stringency
- Desirable preclinical safety profile with lower potential for CRS and macrophage activation syndrome
- IND submission in ccRCC expected Q2 2024

<sup>1</sup>Shaffer et al., Blood 2011  
<sup>3,4</sup>Acharya et al., Blood 2023

<sup>3</sup>Leick et al., Cancer Cell 2022  
<sup>4</sup>Kasap et al., BioRxiv 2024

# Potential Near-Term Milestones

	2024		2025	
	1H	2H	1H	2H
<b>ADI-001 Autoimmune Diseases</b>	<ul style="list-style-type: none"> <li>Initiate LN Phase 1</li> <li>Expand to 1-2 Additional Autoimmune Indications</li> </ul>	<ul style="list-style-type: none"> <li>Preliminary Clinical Data in LN</li> <li>Preliminary Clinical Data in 1-2 Additional Autoimmune Indications</li> </ul>		<ul style="list-style-type: none"> <li>Clinical Data in 3 Autoimmune Indications</li> </ul>
<b>ADI-001 MCL</b>	<ul style="list-style-type: none"> <li>Study Enrollment Ongoing</li> </ul>	<ul style="list-style-type: none"> <li>Clinical Update</li> </ul>	<ul style="list-style-type: none"> <li>Define Regulatory Path for Potentially Pivotal Phase 2 Study</li> </ul>	<ul style="list-style-type: none"> <li>Clinical Update</li> </ul>
<b>ADI-270 RCC</b>	<ul style="list-style-type: none"> <li>ADI-270 IND Submission</li> </ul>		<ul style="list-style-type: none"> <li>Clinical Data RCC and Potential Expansion to additional CD70+ Tumors</li> </ul>	<ul style="list-style-type: none"> <li>Clinical Update</li> </ul>

Proforma adjusted cash and cash equivalents : ~\$247.6M (3/31/24)  
 Projected cash runway into 2H 2026



**Leaders in Developing Allogeneic  
CAR  $\gamma\delta$ 1 T Cell Therapies to Fight  
Autoimmune Diseases and Cancer**

