

Corporate Presentation

January 2024



Cautionary Note Regarding Forward-Looking Statements

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Sana Biotechnology

Changing the Possible for Patients

Sana's hypoimmune technology goal is to overcome allogeneic rejection

- HIP technology provides foundation for potential multiple drugs across many therapeutic areas

Begin 2024 with four clinical programs treating seven diseases

- SC291 oncology – NHL and CLL
- SC291 B-cell mediated autoimmune – lupus nephritis, extrarenal lupus, and ANCA-associated vasculitis
- SC262 oncology – r/r NHL, initially in CD19 CAR T failures
- HIP primary islet cells in patients with type 1 diabetes

Pipeline positioned to deliver additional clinical data over time

- Regenerative medicine: SC379 (CNS disorders) and SC451 (type 1 diabetes)
- Hypoimmune allogeneic CAR T cells: SC255 (BCMA) and beyond

Balance sheet allows potential for multiple data readouts

Sana pipeline positioned to deliver meaningful clinical data

PRODUCT CANDIDATE	MECHANISM	INDICATIONS	PRECLINICAL IND-ENABLING	PHASE 1	PHASE 2/3	SANA'S RIGHTS
Oncology						
SC291	CD19-directed allo CAR T	NHL	ARDENT			WW
SC291	CD19-directed allo CAR T	CLL	ARDENT			WW
SC262	CD22-directed allo CAR T	NHL (CD19 failures)	VIVID			WW
SC255	BCMA-directed allo CAR T	MM				WW
B-cell Mediated Autoimmune Diseases						
SC291	CD19-directed allo CAR T	LN	GLEAM			WW
SC291	CD19-directed allo CAR T	ERL	GLEAM			WW
SC291	CD19-directed allo CAR T	AAV	GLEAM			WW
SC291	CD19-directed allo CAR T	Other indications				WW
Regenerative Medicine						
UP421	HIP primary islet cells ¹	T1D				WW
SC451	Stem-cell derived pancreatic islet cells	T1D				WW
SC379	Glial progenitor cells	HD, PMD, SPMS				WW

¹Investigator sponsored trial.

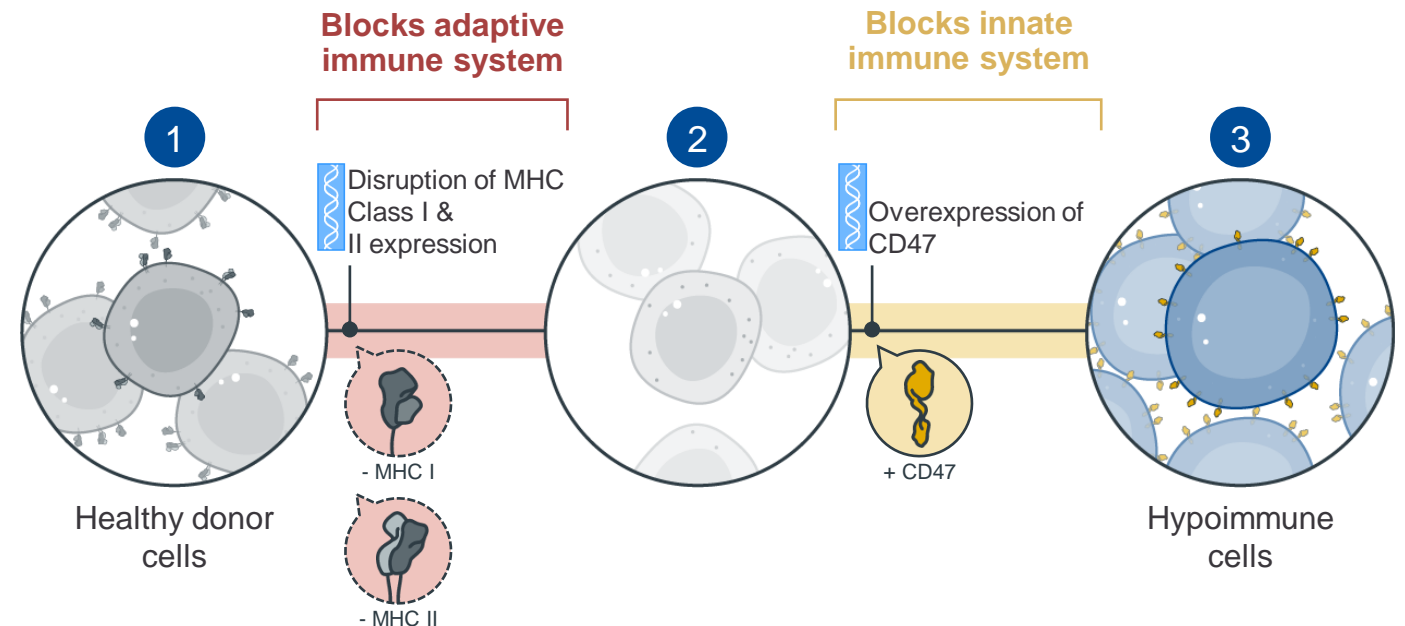
Abbreviations: AAV, ANCA-associated vasculitis; CLL, chronic lymphocytic leukemia; ERL, extrarenal systemic lupus erythematosus; HD, Huntington's disease; LN, lupus nephritis; MM, multiple myeloma; NHL, non-Hodgkin lymphoma; PMD, Pelizaeus-Merzbacher Disease; SPMS, secondary progressive multiple sclerosis; T1D, type 1 diabetes; WW, worldwide.

Overcoming allogeneic immune rejection has been key limitation in transplant and cellular medicine

Allogeneic cell rejection

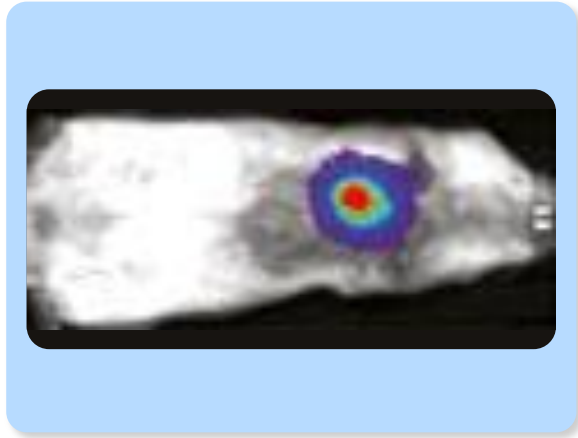
- ~75 years of transplants – immune rejection remains the largest problem
- Lifelong immunosuppression is current standard
- Genome modification efforts to date have generally been incomplete
- Autologous therapies have limited scalability and are only available for a small number of cell types

Sana's hypoimmune approach



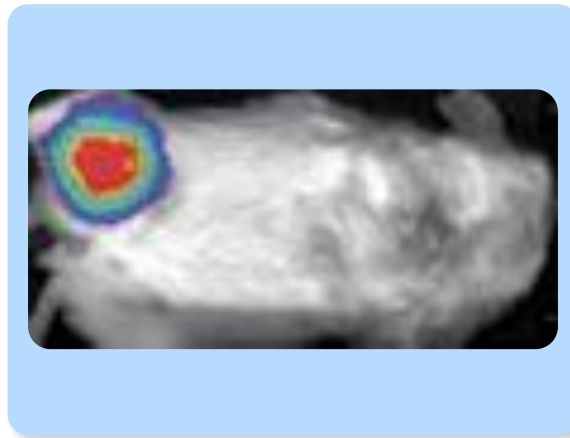
Current clinical platform with multiple ongoing approaches in research phase.

HIP-modified cells successfully transplanted in allogeneic models across various species and cells types



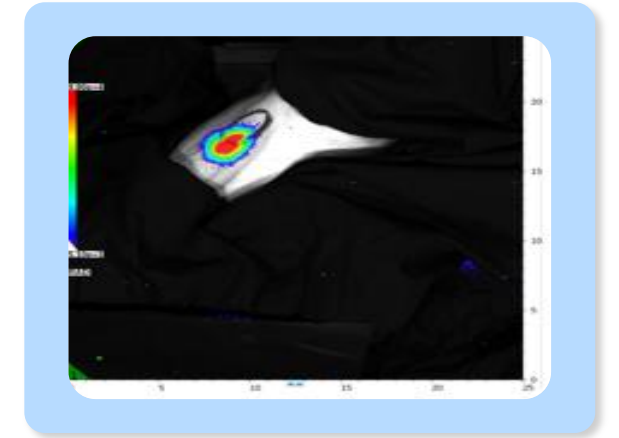
Mice

- Mouse iPSCs
- Mouse iPSC-derived endothelial cells
- Mouse iPSC-derived smooth muscle cells
- Mouse iPSC-derived cardiomyocytes



Humanized mice

- Human iPSCs
- Human iPSC-derived endothelial cells
- Human iPSC-derived smooth muscle cells
- Human iPSC-derived cardiomyocytes
- Human iPSC-derived pancreatic islet cells
- Human donor-derived islet cells
- Human donor-derived CAR T cells

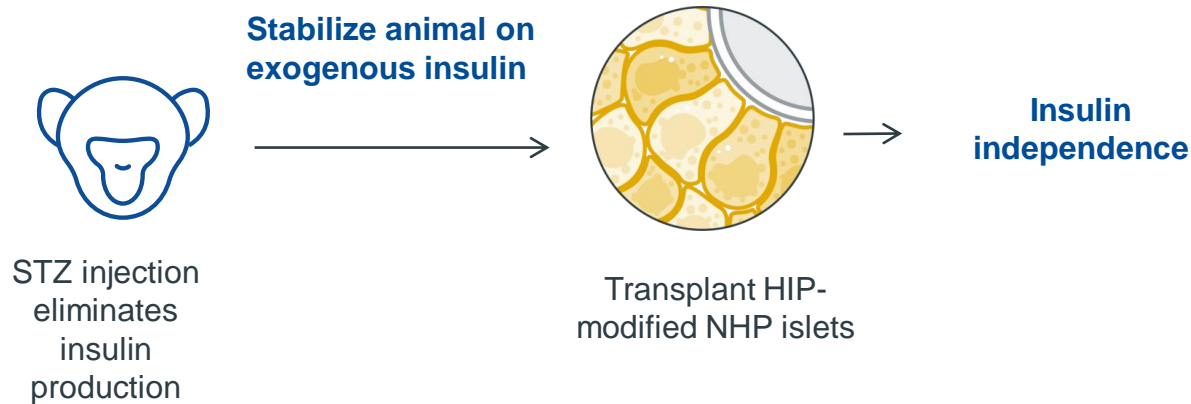


NHPs

- NHP iPSCs (16 weeks follow-up)
- NHP donor-derived islets (40 weeks follow-up)
- NHP iPSC-derived cardiomyocytes
- NHP iPSC-derived retinal pigment epithelium (RPE) cells

HIP-modified allogeneic islet cells to control glucose in a type 1 diabetic NHP model

Type 1 diabetes is a disease of missing pancreatic beta cells



Study Design (N=1)

- NHP treated with STZ
- Glucose stabilized with exogenous insulin
- Allogeneic NHP primary islet cells isolated and HIP-modified
- Cells injected intramuscularly without immunosuppression

Key goals of study

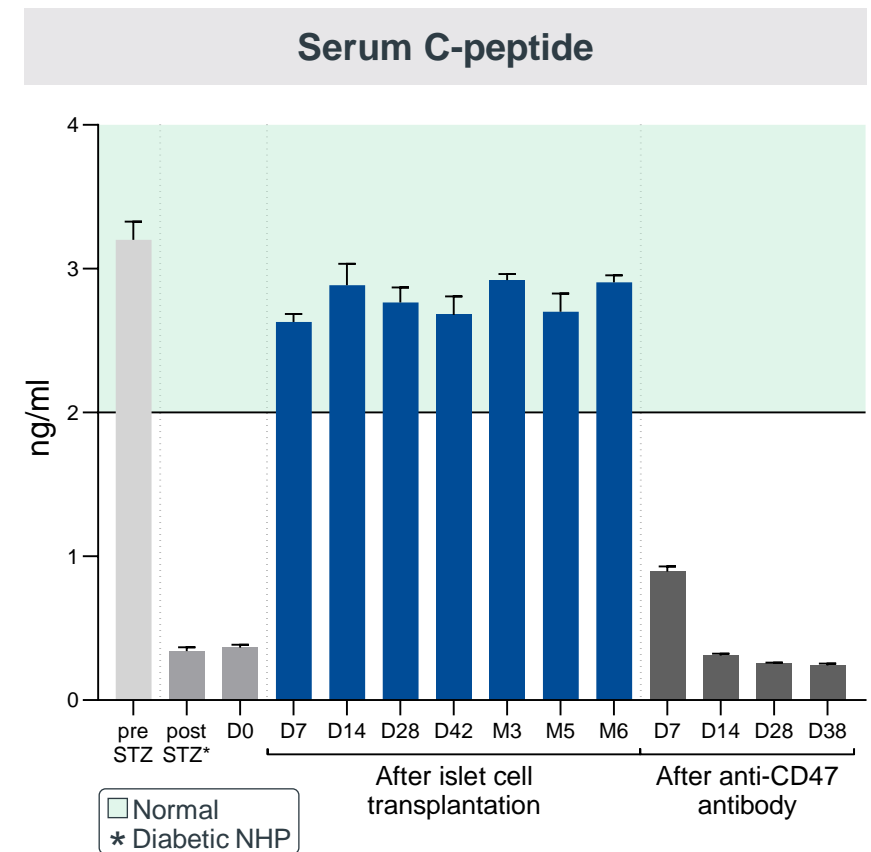
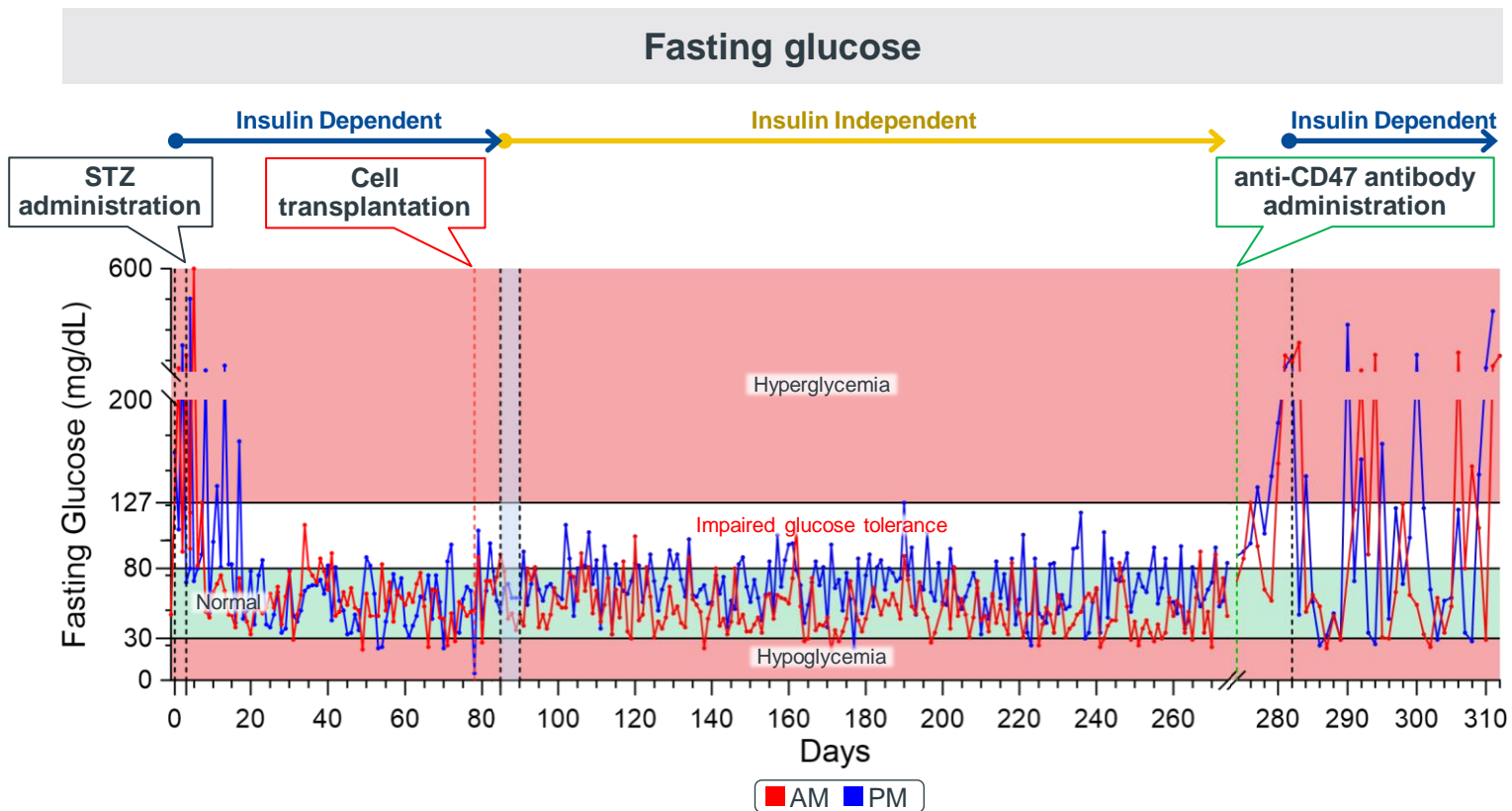
- Demonstrate survival and function of HIP-modified allogeneic islet cells in diabetic NHP without immunosuppression
- Demonstrate long-term glucose normalization in diabetic NHP without exogenous insulin or immunosuppression
- Demonstrate the principle of graft ablation/safety switch with anti-CD47 antibody

Abbreviations: NHP, non-human primate; STZ, Streptozotocin.

Survival and function of allogeneic hypoimmune pancreatic islet cells in diabetic NHP 6 months without immunosuppression

Study Design (N=1)

- NHP primary islet cells isolated and HIP-modified
- Cells injected intramuscularly into a diabetic, allogeneic NHP without immunosuppression



Near-term opportunities to apply HIP modifications to validated mechanisms with unmet need

Blood cancers:

>100,000 patients/year^{1,2}



B-cell mediated autoimmune diseases:

>5 million patients³



Type 1 diabetes:

>8 million patients⁴



¹Avezbakiyev et al. *Blood*. 2022

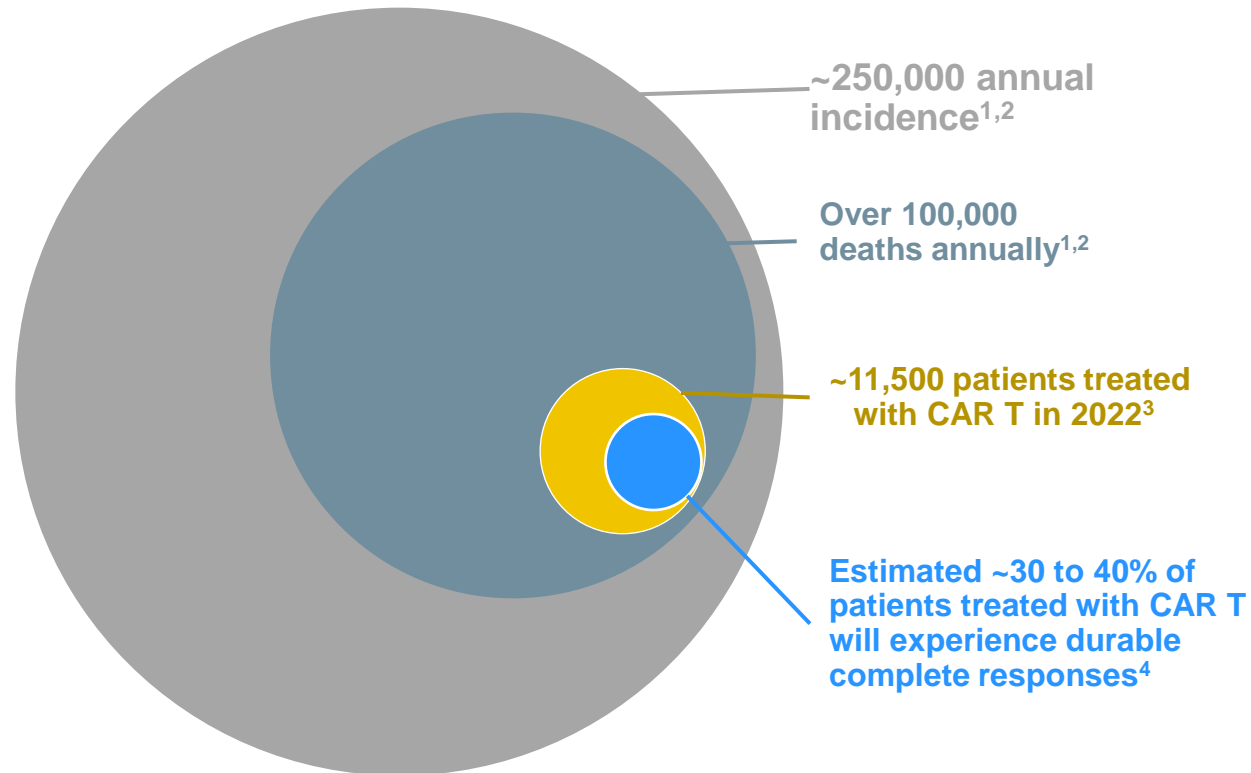
²Durie et al. *The Oncologist*. 2020

³NIH Autoimmune Diseases Coordinating Committee: Autoimmune Diseases Research Plan, October 2017, U.S.

⁴t1dindex.org

Hematologic cancers continue to have a high unmet need

High mortality in lymphoma and myeloma in the US and EU5



¹Avezbakiyev et al. *Blood*. 2022

²Durie et al. *The Oncologist*. 2020

³Clarivate DRG NHL and MM Market Forecast Nov 2022; internal analysis of secondary EPI data.

⁴Scivida 2022 NHL Factbook

Abbreviations: EU5, France, Germany, Italy, Spain, UK

Challenges

- Autologous CAR T cell scalability
- Many patients fail CAR T treatment
- Allogeneic CAR T cell immune rejection limits persistence and efficacy

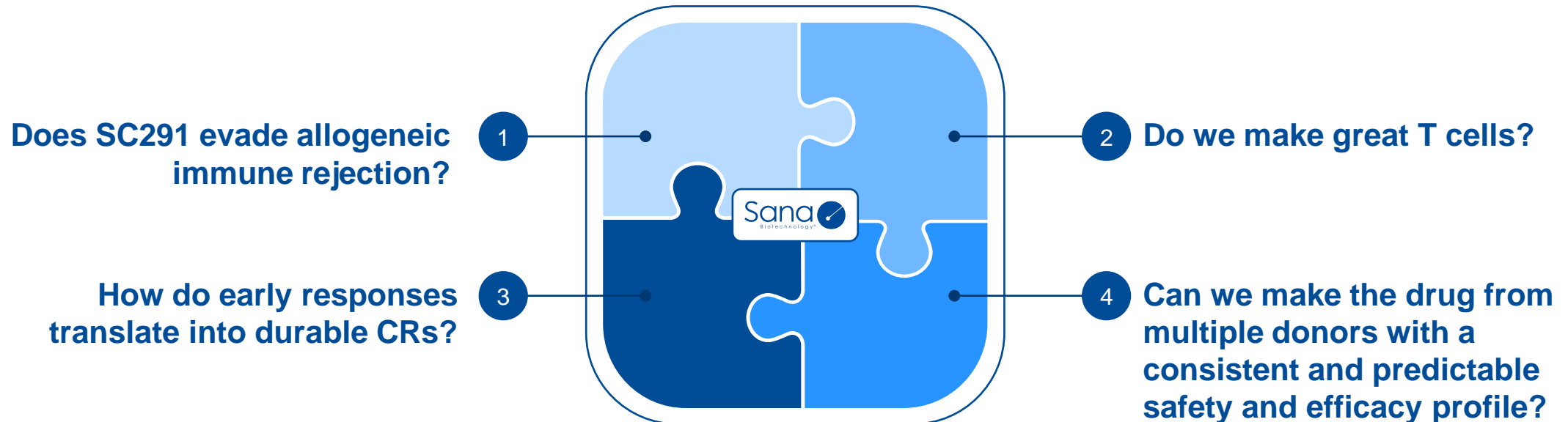
Opportunity

- Known targets
- Known efficacy and safety bar

Sana's HIP CAR T platform can address challenges and exploit opportunities

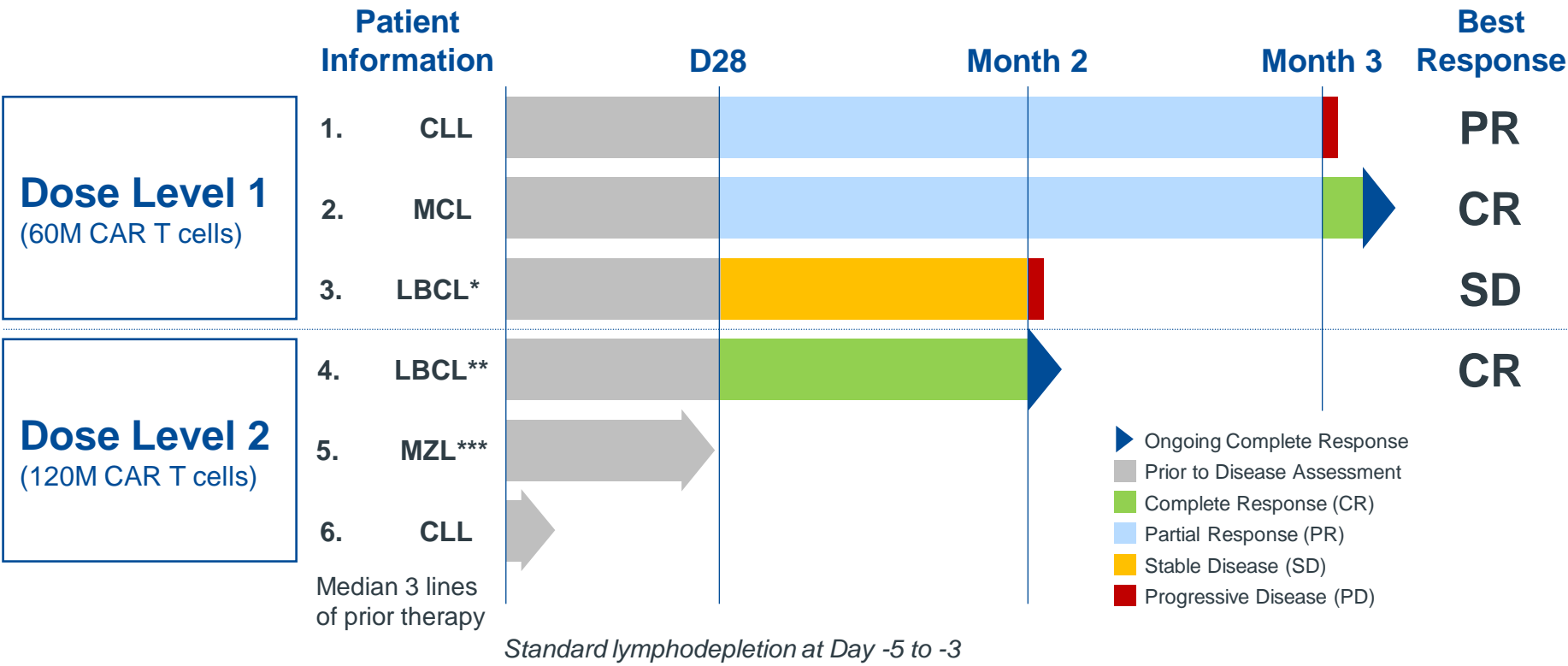
Defining success for SC291 in oncology

Understanding levels of evidence as data mature



ARDENT: 3 of 4 evaluable patients had at least a partial response, with 2 ongoing complete responses

6 patients treated to date; dose escalation ongoing



Safety

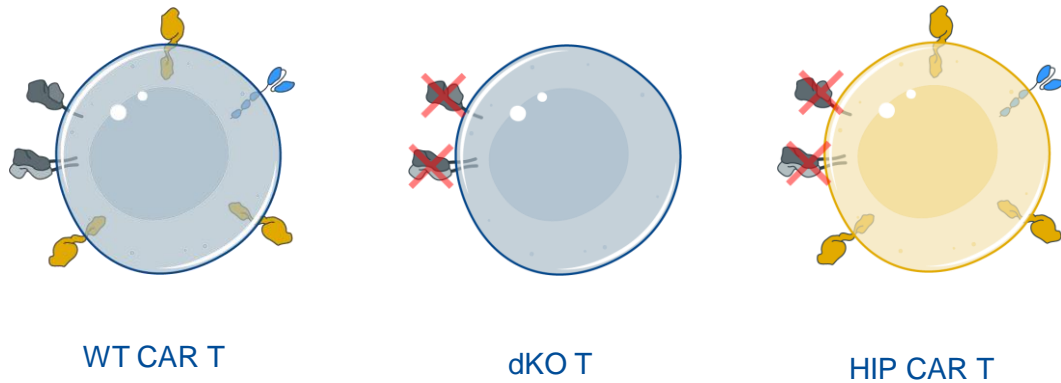
- No dose limiting toxicities
- No SC291-related SAEs
- No CRS or ICANS
- No Grade 3 or higher infections

Clinical data as of: January 5, 2024
 "evaluable" defined as patients treated with SC291 and had at least one disease assessment
 *Transformed DLBCL from FL. **Transformed DLBCL from MZL. ***Assessment ongoing as of January 5, 2024.

Immune response data provide important early insights

Translating preclinical data to people

1 SC291 is a mixture of HIP and non-HIP CAR T cells



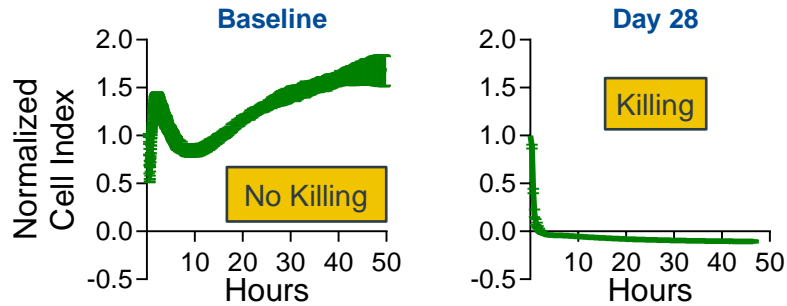
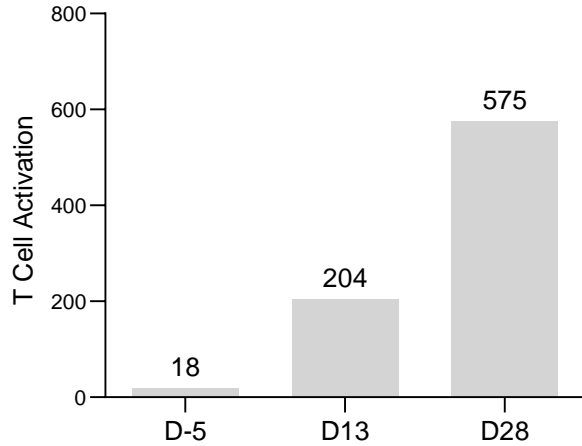
T Cell Population	Genetic Modifications
WT CAR T	CD47-CD19 CAR
dKO T	HLA I/II deficient
HIP CAR T	CD47-CD19 CAR; HLA I/II deficient

2 Test the patient's immune system against SC291

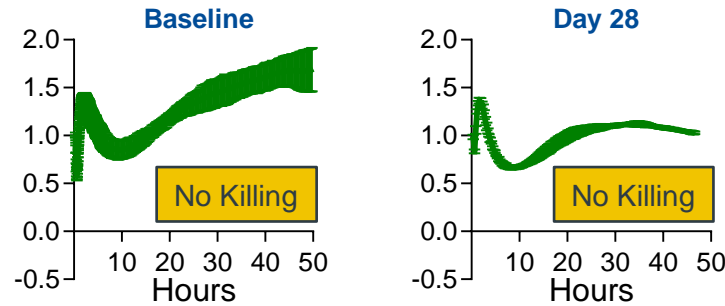
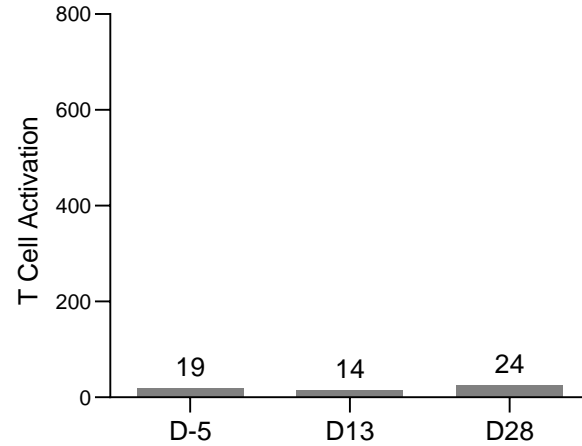
Cell	Day 28			
	T cell	Ab	NK cell	Blood
WT CAR T	Red	Red	Green	Red
dKO T	Green	Green	Red	Red
HIP CAR T	Green	Green	Green	Green

Patient T cells kill WT CAR T cells but do not kill dKO T cells or HIP CAR T cells

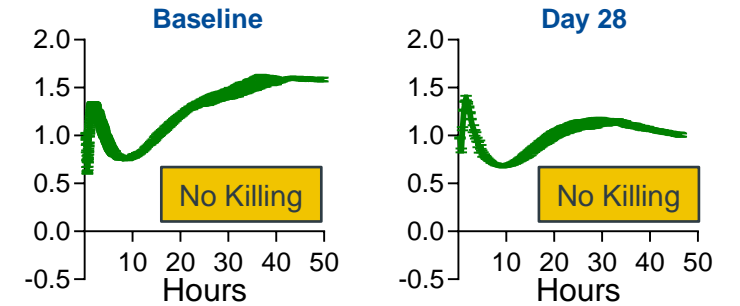
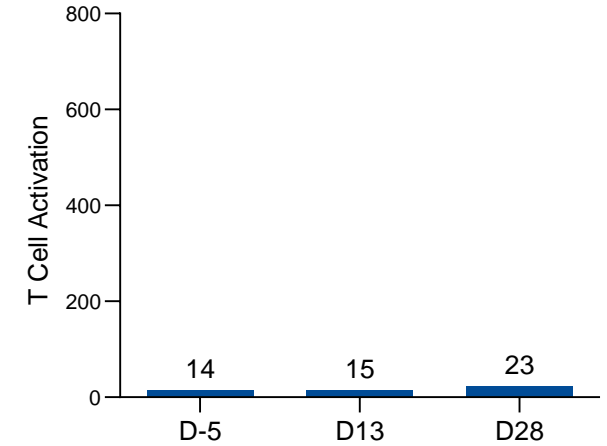
Patient T cells kill WT CAR T cells



Patient T cells do not kill dKO T cells



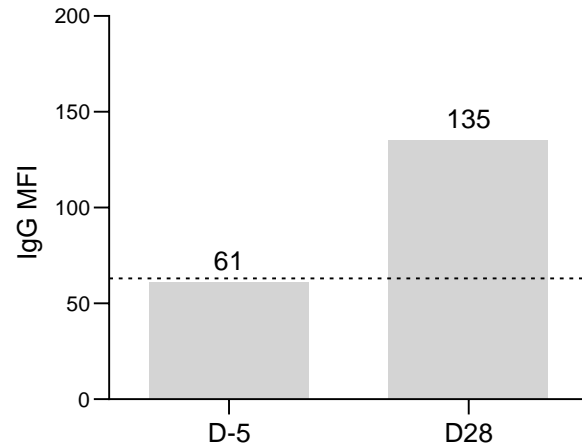
Patient T cells do not kill HIP CAR T cells



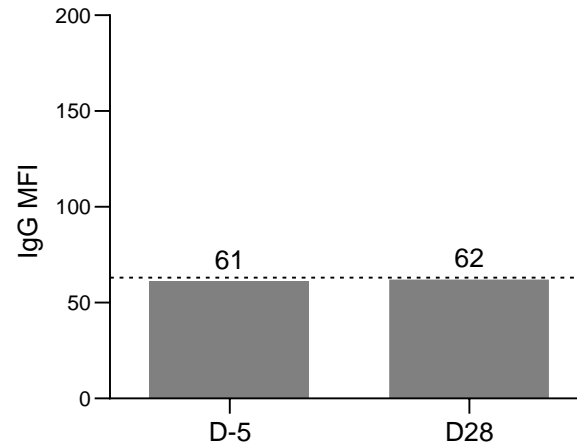
From Patient #1 in the ongoing ARDENT trial.

Patient generates antibodies against WT CAR T cells but not dKO T cells or HIP CAR T cells

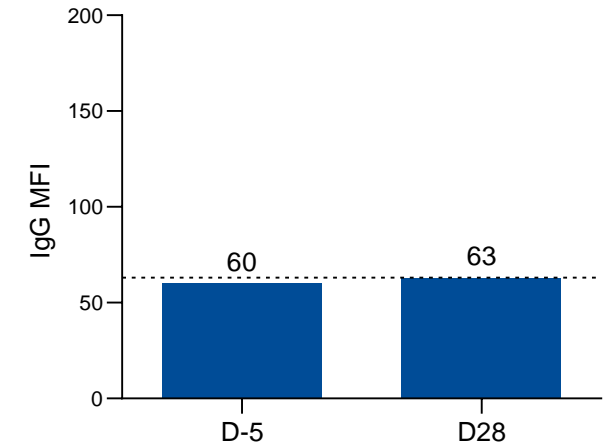
WT CAR T cells induce an antibody response



dKO T cells do not induce an antibody response



HIP CAR T cells do not induce an antibody response



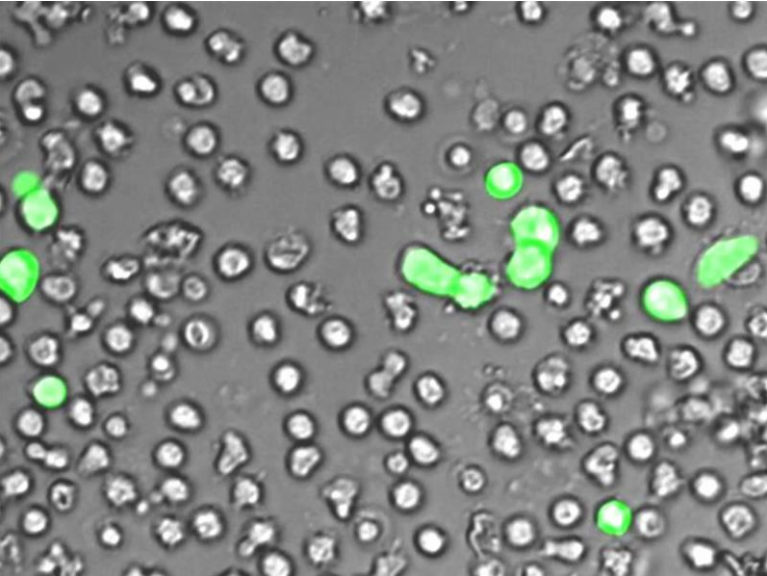
From Patient #1 in the ongoing ARDENT trial.

Only HIP CAR T cells avoid NK cell killing

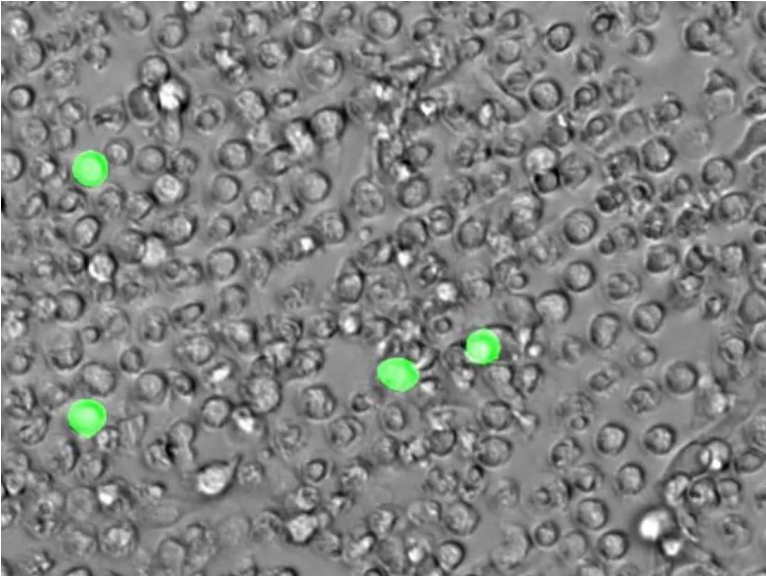
NK cells taken from patient's blood

Patient's NK cells from day 13 after SC291 dosing

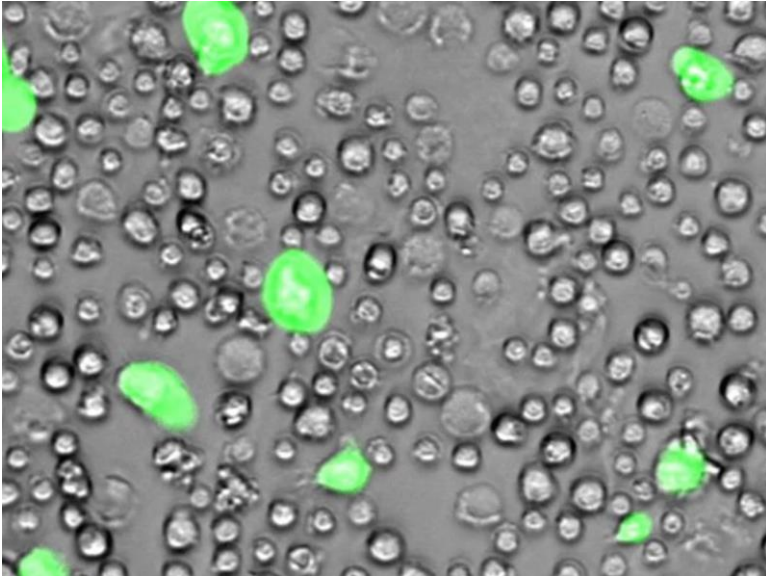
NK cells kill dKO T cells





NK cells kill dKO T cells with HLA-E overexpression



NK cells do NOT kill HIP CAR T cells



Actual assay time = 4 hours.

 T cell with editing profile in column title  NK cells

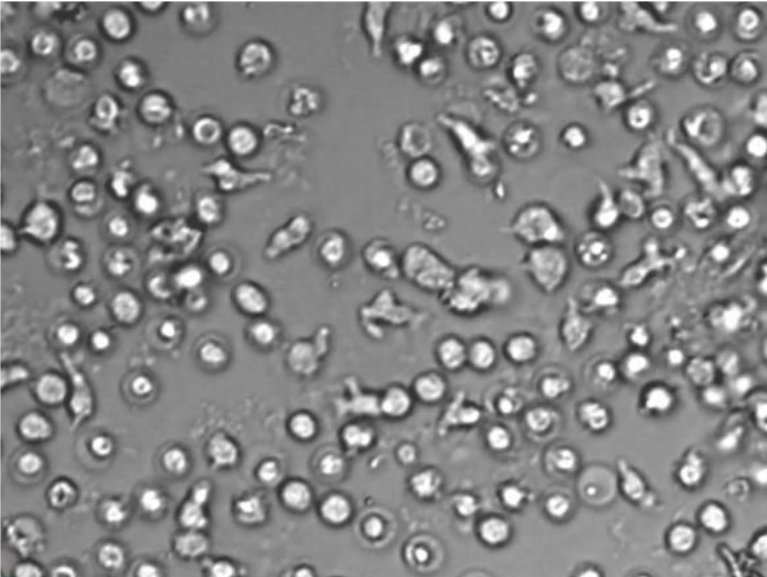
From Patient #1 in the ongoing ARDENT trial.

Only HIP CAR T cells avoid NK cell killing

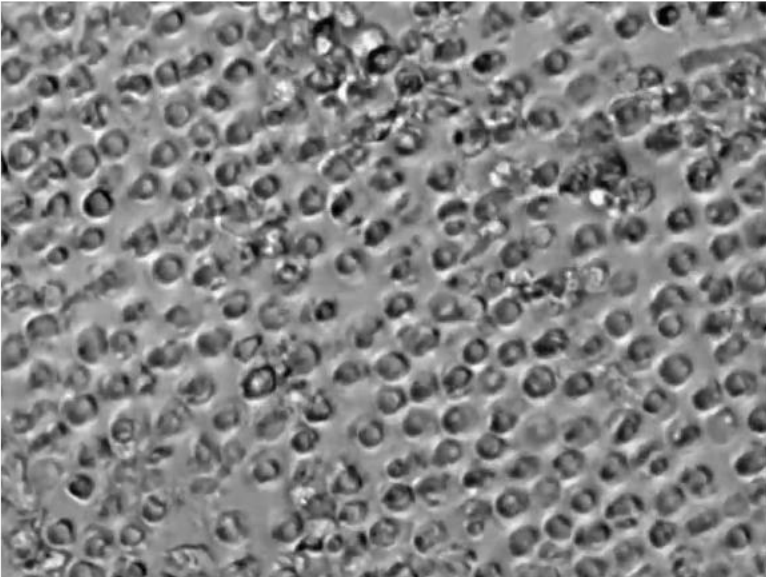
NK cells taken from patient's blood

Patient's NK cells from day 13 after SC291 dosing

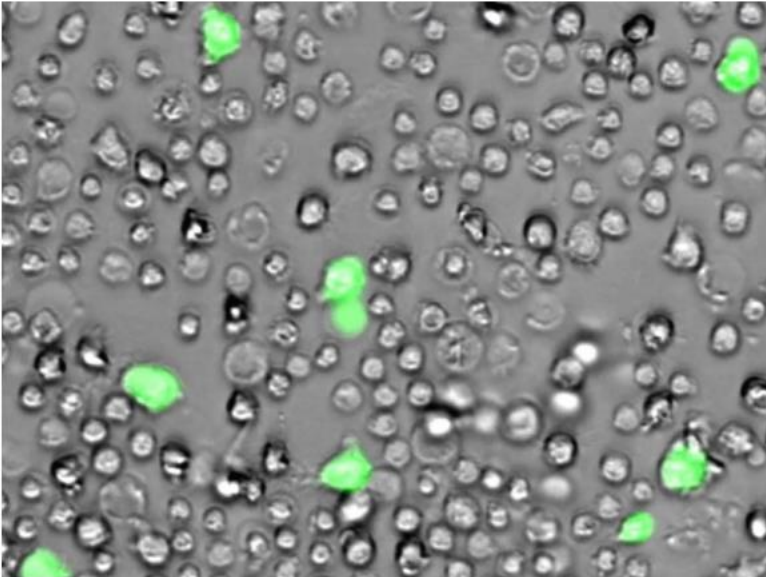
NK cells kill dKO T cells



NK cells kill dKO T cells with HLA-E overexpression



NK cells do NOT kill HIP CAR T cells



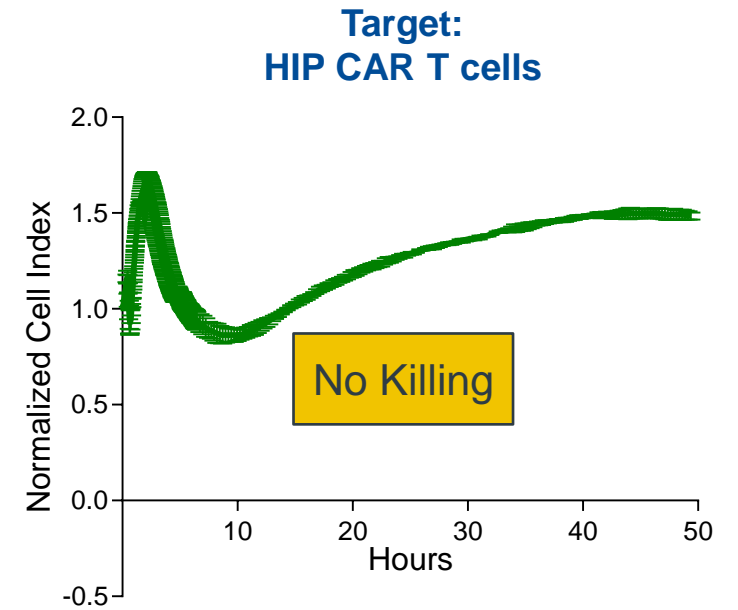
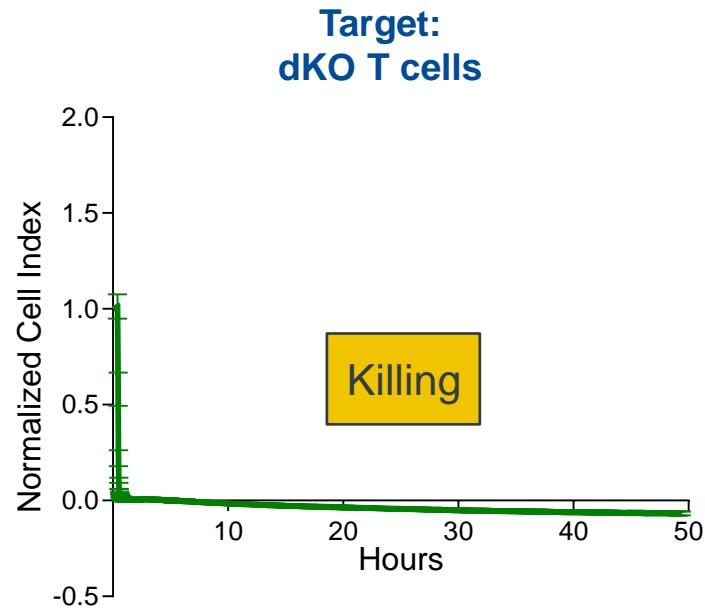
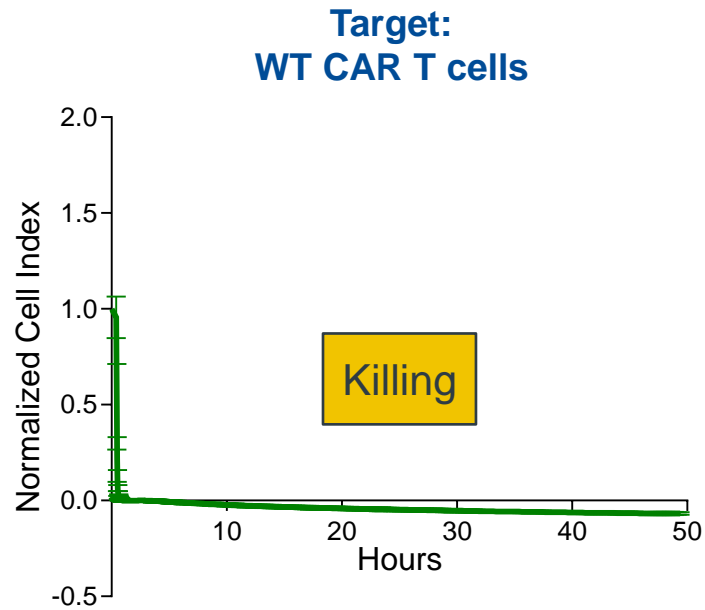
Actual assay time = 4 hours.

 T cell with editing profile in column title  NK cells

From Patient #1 in the ongoing ARDENT trial.

No detectable immune response in the patient toward HIP CAR T cells

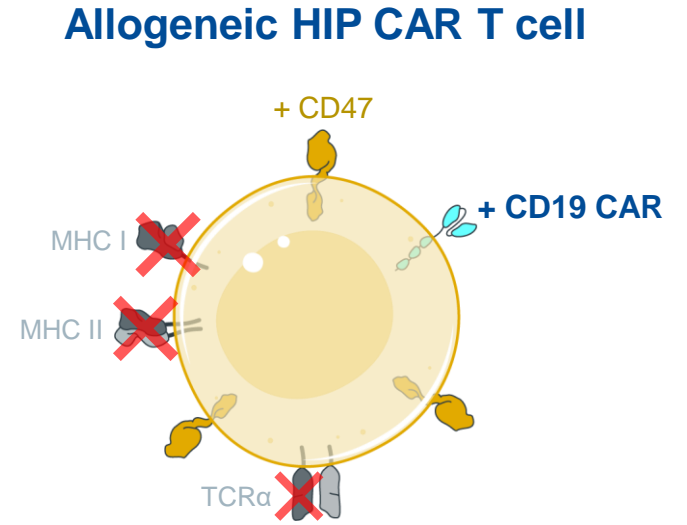
D28 blood sample



From patient #1 in the ongoing ARDENT trial.

SC291: ARDENT trial continues enrollment with more data expected in 2024

- Early data suggest ability to dose safely, the desired immune evasion profile, and clinical efficacy
- More data to come
 - Immune evasion
 - Safety profile
 - Response rate
 - Cell persistence
 - Durability of responses



An effective allogeneic CAR T cell therapy offers potential to transform outcomes for patients

Autoimmune diseases have emerged as promising opportunity

1 B-cell targeting therapies have been efficacious across many autoimmune diseases¹

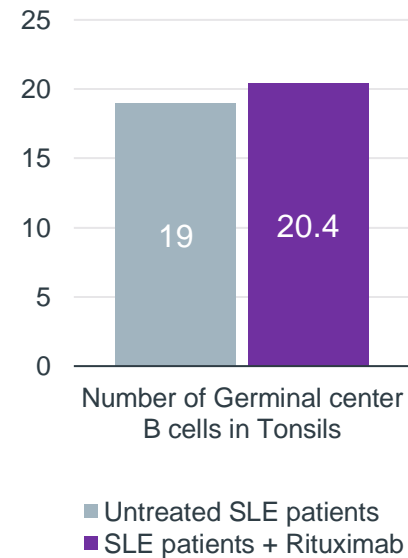
- SLE
- Vasculitis (granulomatosis with polyangiitis & microscopic polyangiitis)
- Neuromyelitis optical spectrum
- Pemphigus
- Relapsing and progressive MS
- Rheumatoid arthritis
- Lupus nephritis
- Sjogren syndrome
- NMDAR encephalitis
- Thrombocytopenic purpura
- Amyloidosis
- Scleroderma
- Autoimmune hemolytic anemia
- Chronic immune demyelinating polyradiculoneuropathy
- Immune-mediated necrotizing myopathy
- Membranous nephropathy

¹Adapted from Zhang et al. *Frontiers in Immunology*. 2023; Oh et al. *Immune Network*. 2023; Lee et al. *Nature Reviews Drug Discovery*. 2021.

²Anolik et al. *Arthritis and Rheumatism* 2007

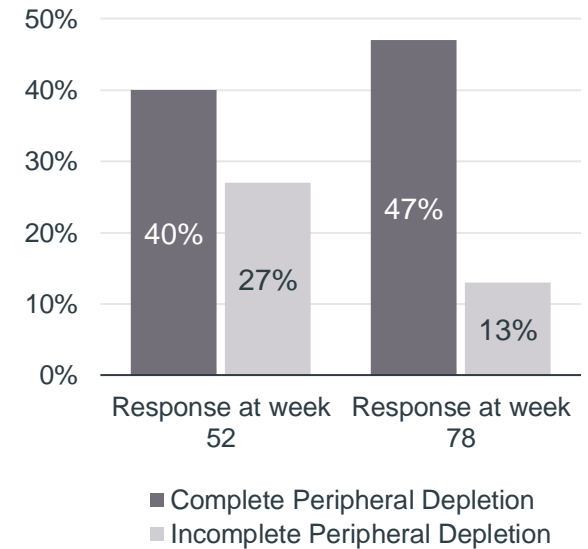
³Mendez et al. *Clinical Journal of the American Society of Nephrology* 2018

2 Germinal center B cells are unaffected by rituximab treatment²



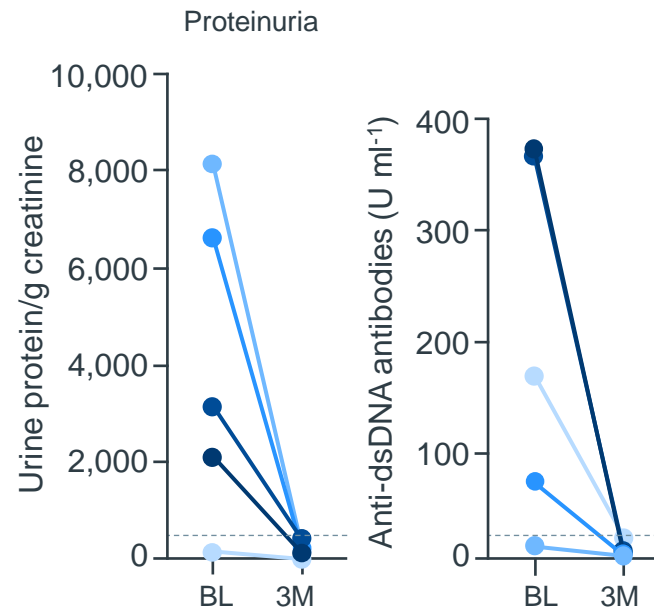
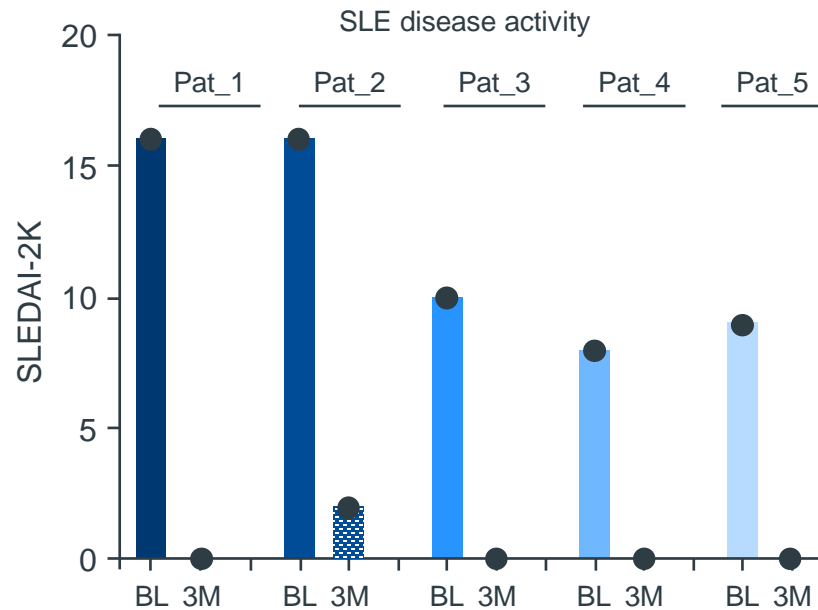
3 Depth of B cell depletion with treatment predicts efficacy in early trials³

Complete B-cell depletion resulted in greater complete responses in Lupus Nephritis patients²



Autologous CD19 CAR T therapy results in durable drug-free remission in refractory SLE patients

Improvement in signs and symptoms of SLE after CD19 CAR T treatment

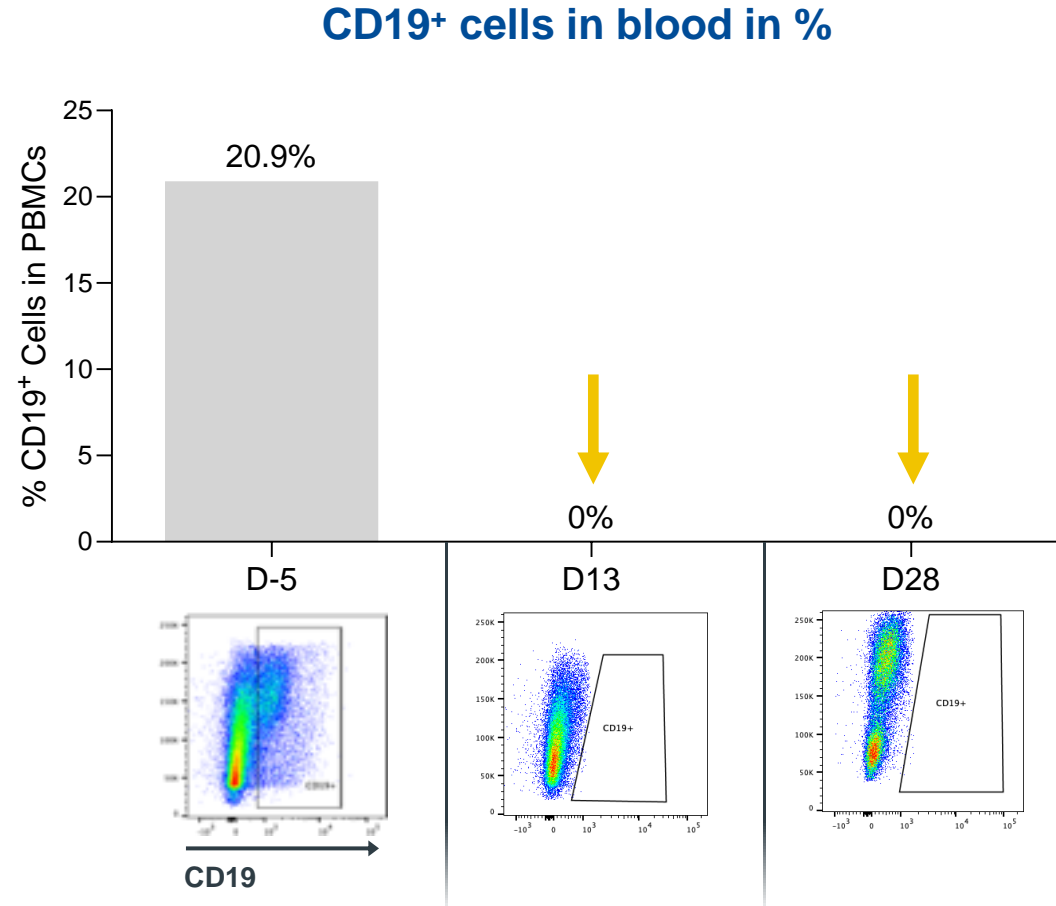


- Well tolerated – mild CRS (Grade 1) and no ICANS seen
- 5/5 patients are in a durable complete remission and off all drugs
- All known disease biomarkers rapidly normalized and have remained so thus far
- 24+ months of drug-free remission seen in patients constituting a potential functional cure
- B-cell recovery and immune system reset in ~3 months with sustained SLE remission

Mackensen et al. *Nature Medicine*. 2022

Abbreviations: BL, baseline; CRS, cytokine release syndrome; ICANS, immune effector cell-associated neurotoxicity syndrome; M, months; SLE, systemic lupus erythematosus.

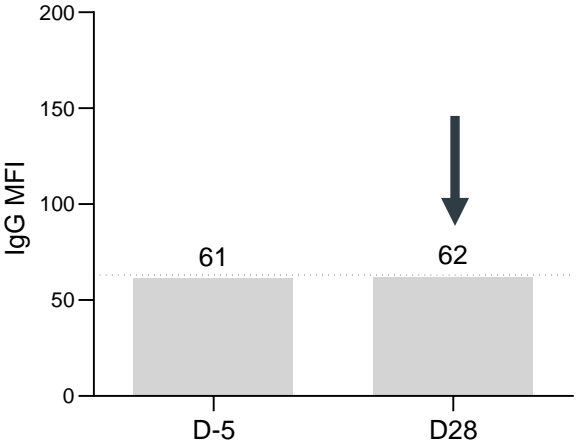
ARDENT trial: SC291 treatment leads to deep B cell depletion in oncology patient



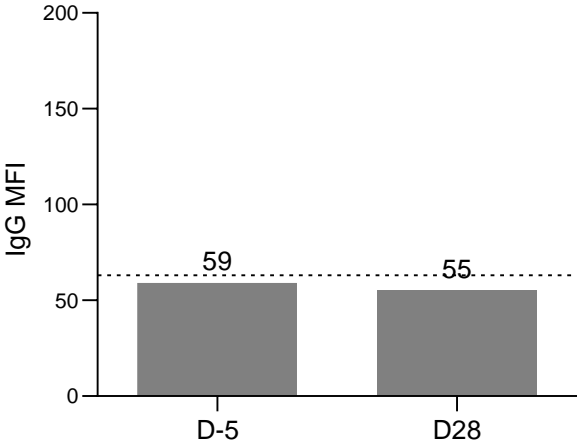
From Patient #4 in the ongoing ARDENT trial.

Complete B cell depletion may be even more important in autoimmune than oncology patients

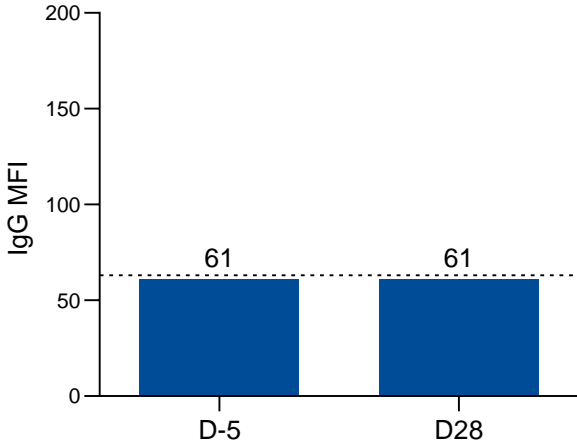
No anti-HLA antibody production against WT CAR T cells suggests complete B cell depletion



dKO T cells do not induce an antibody response



HIP CAR T cells do not induce an antibody response



From Patient #4 in the ongoing ARDENT trial.

SC291 offers potential for transformative treatment for B-cell mediated autoimmune diseases

Targeting multiple indications

Phase 1 trial – multiple autoimmune disorders

- 1 Lupus nephritis >230K^{1,2} patients³
- 2 Extrarenal SLE >200K¹ patients³
- 3 ANCA-associated vasculitis >60K⁴ in US

SC291 benefits versus autologous therapies

- 1 No patient apheresis
- 2 Product availability
- 3 Scaled manufacturing
- 4 Consistent T cell quality

¹Lu et al. *Annals of Rheumatic Diseases*. 2023

²Guzman et al. *Arthritis Rheum*. 2013

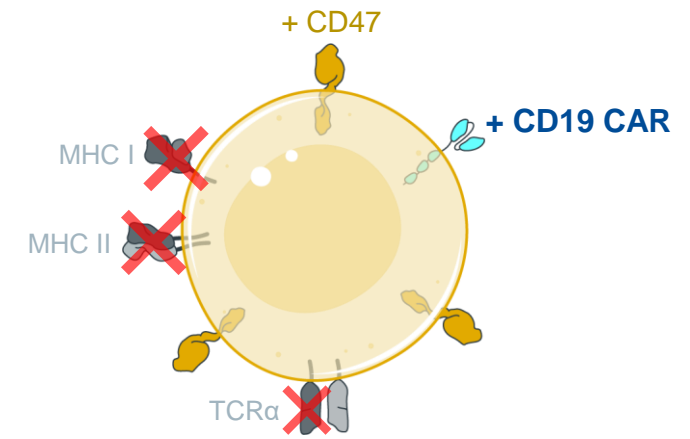
³US, EU5, and Japan

⁴Jayne et al. ANCA-Associated Vasculitis: An Update

SC291: GLEAM Phase 1 trial goals are to understand safety, dose, and early efficacy

- Key features of Phase 1 trial (GLEAM)
 - Patients with refractory lupus nephritis, extrarenal SLE, and AAV
 - Starting dose of 90 million CAR T cells
 - Potential to expand beyond these indications over time
- Data expected in 2024 from multiple indications
 - Safety and tolerability
 - Early response rates

Allogeneic HIP CAR T cell



An effective allogeneic CAR T offers potential to transform outcomes for patients

SC262: Targeting growing population of patients with inadequate response to CD19 therapy

CD19 CAR T relapsed patients represent large and growing unmet need¹

Estimated ~12,000 B cell malignancy patients treated with CD19 CAR T in 2027²

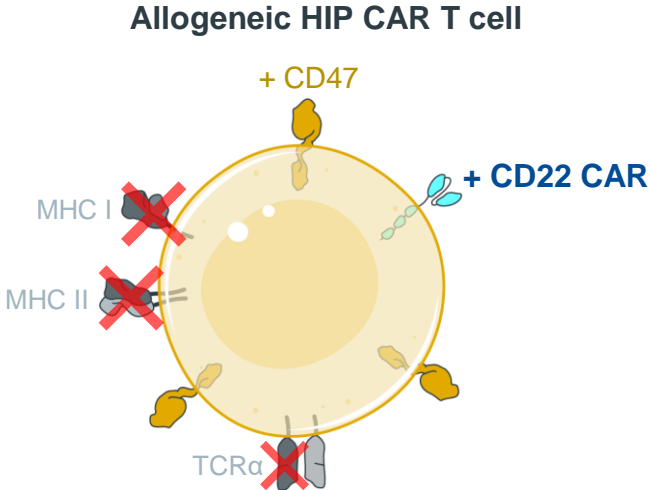


- Potential of ~7,500 CAR T failures annually in 2027²
- Median survival of ~5 months post-CD19 CAR T therapy failure³

Estimated ~35-40% of CAR T patients with durable complete responses⁴

= 1,000 people

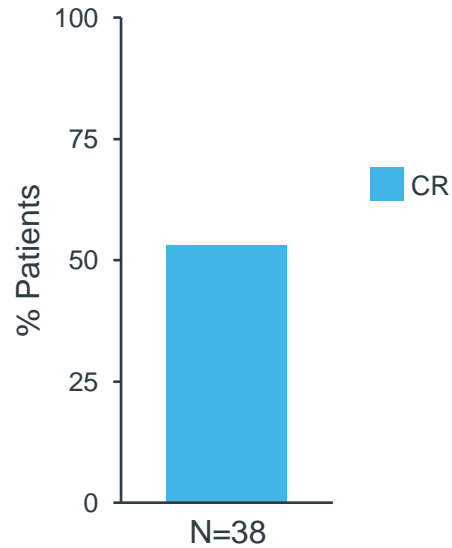
Expand our allo T platform to CD22 with Sana's SC262 candidate



¹US, EU5, and Japan. ²Clarivate DRG NHL Market Forecast Nov 2021; 2027 Forecast is 2L+ LBCL patients; internal analysis of secondary EPI data. ³Di Blasi et al. *Blood*.2022; DESCAR-T registry.

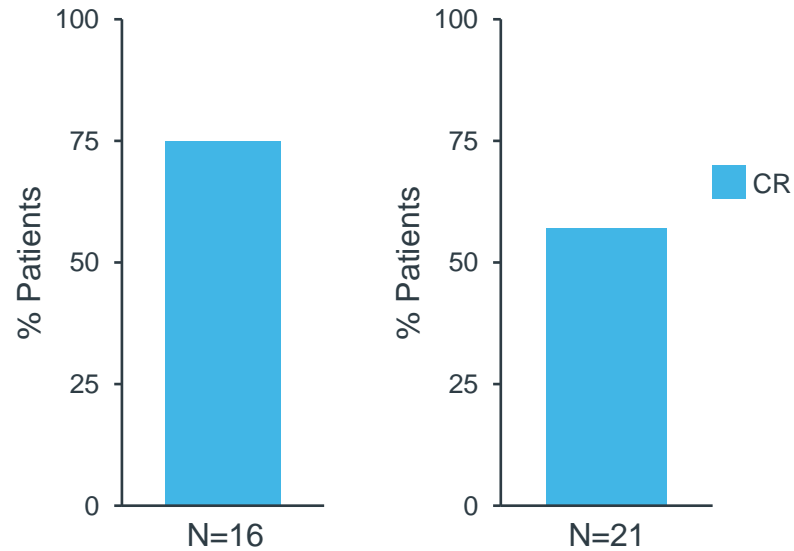
SC262: Licensed CD22 CAR produced strong clinical data in CD19 failures when part of autologous CAR T

>50% 6-month CR rate in CD19 CAR failure DLBCL patients



2023 ASH Yi-Jiun Su

High rate of CRs in CD19 failure ALL patients ~80% patients with prior CD19 therapy



2022 ASH Miklos/Stanford

2018 Nature Med Fry, et al.

VIVID Phase 1 Trial

- CD19 CAR T exposed relapsed and/or refractory NHL
- Adult subjects
- Dose escalation study
- Cell dose: 90M, 150M, and 250M
- Standard lymphodepletion
- Primary Endpoints: Safety and tolerability
- Secondary Endpoints: Patient response

Type 1 diabetes represents a large unmet need with a loss of ~15 years of life¹

- Disease caused by autoimmune destruction of insulin-producing pancreatic beta cells, resulting in no insulin production
- Type 1 diabetes is a large unmet need with >8M WW²
- Short-term complications result from hypo- and hyperglycemia
- Long-term complications result from micro- and macrovascular disease and end-organ damage: including heart attack, stroke, blindness, and kidney failure
- SC451 goal is euglycemia without any immunosuppression or exogenous insulin

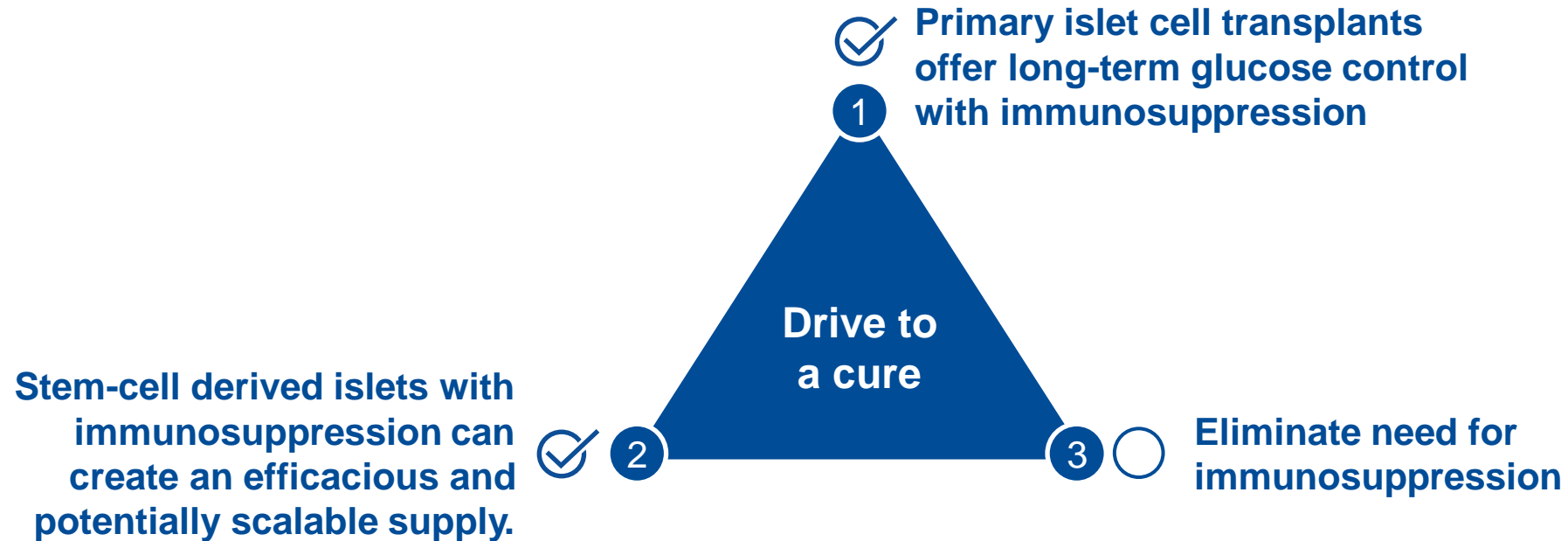


¹Rawshani et al. *Lancet*. 2018

²t1dindex.org

Emerging data suggest a cure is possible

Sana – combining stem cell, gene editing, and immunology expertise

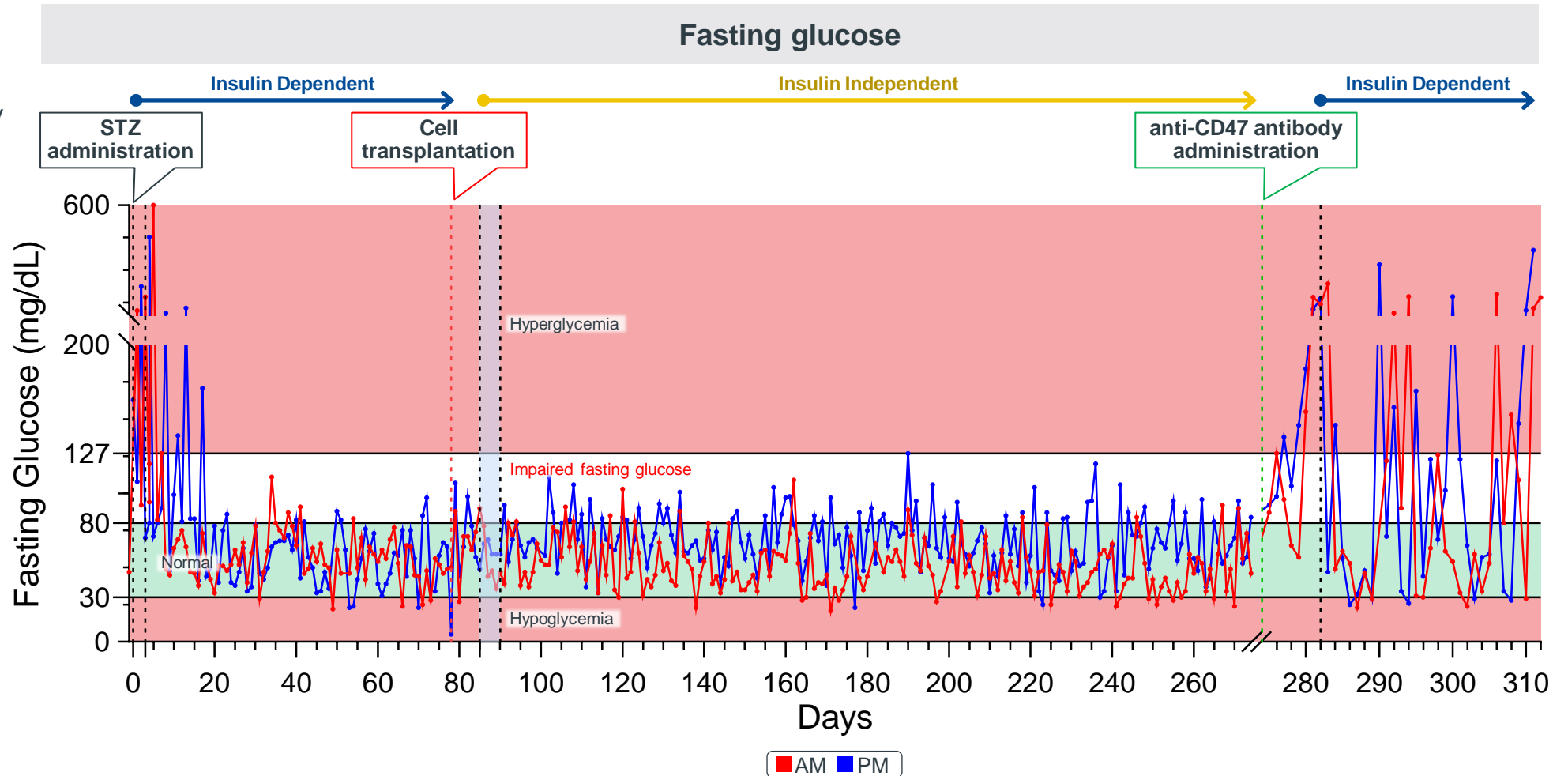


Goal – single treatment with long-term normal blood glucose without immunosuppression or insulin

Survival and function of allogeneic hypoimmune pancreatic islet cells in diabetic NHP 6 months without immunosuppression

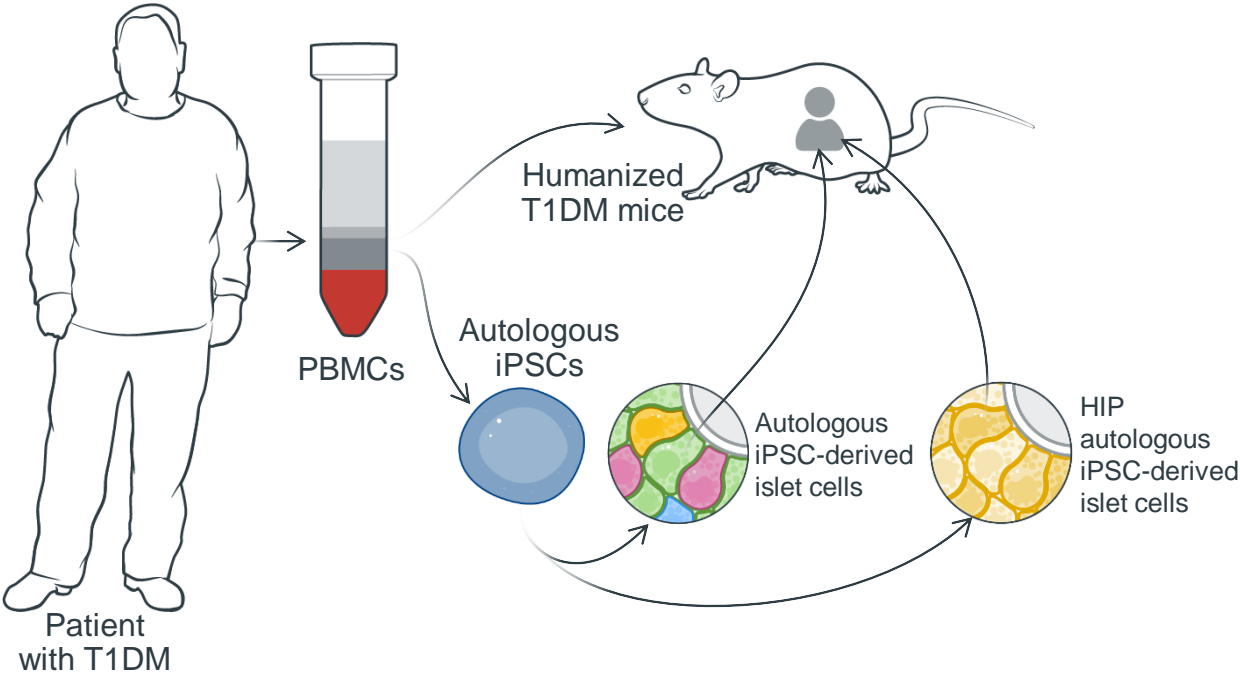
Study Design (N=1)

- NHP primary islet cells isolated and HIP-modified
- Cells injected intramuscularly into a diabetic, allogeneic NHP without immunosuppression



Type 1 diabetes model highlights potential to overcome autoimmune rejection of pancreatic beta cells

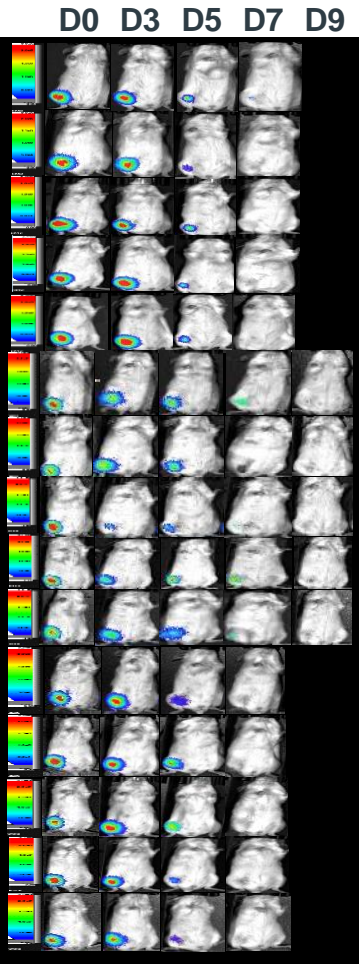
PBMCs from patient with T1DM used to generate stem cell-derived islet cells and to humanize immune system in mice



Unmodified stem cell-derived islet cells from patient with T1DM do not survive

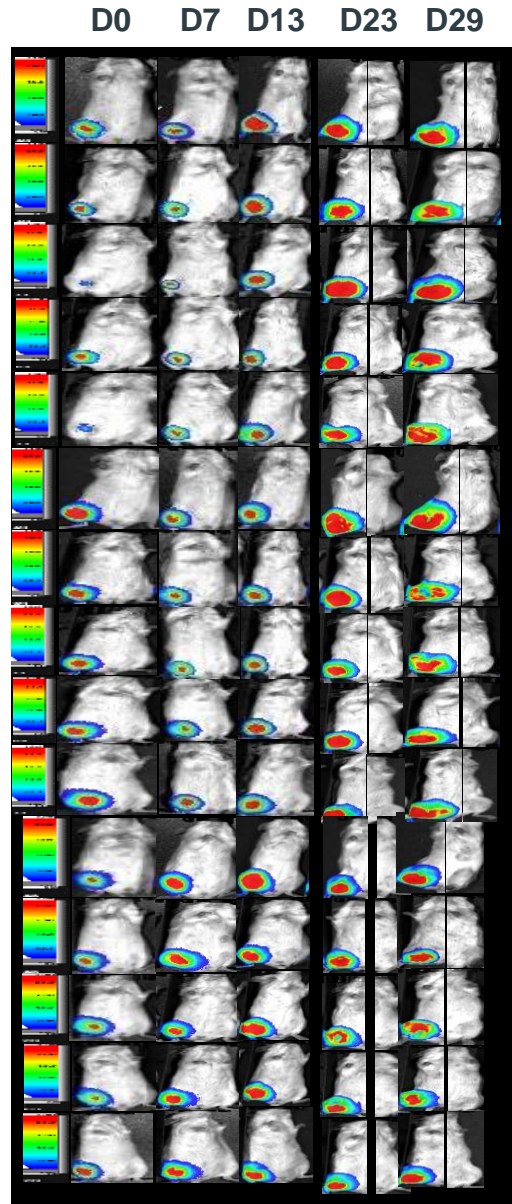
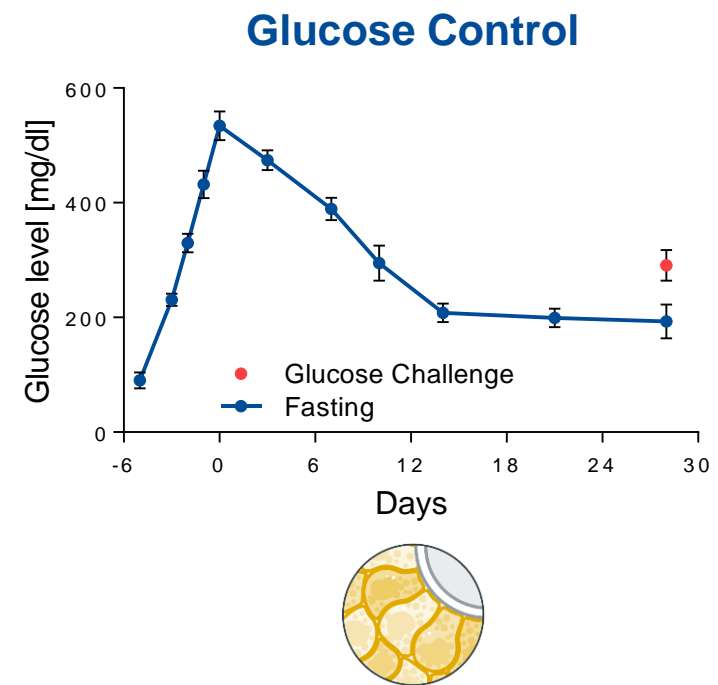
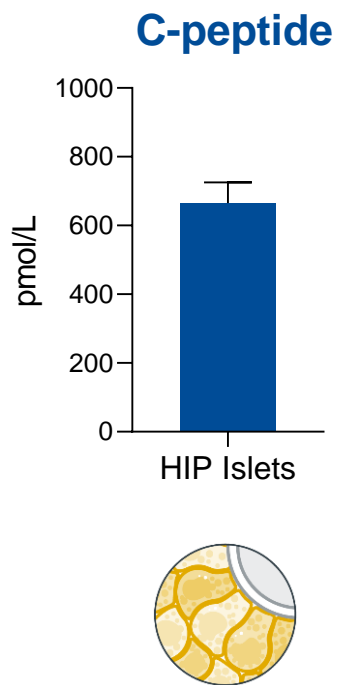
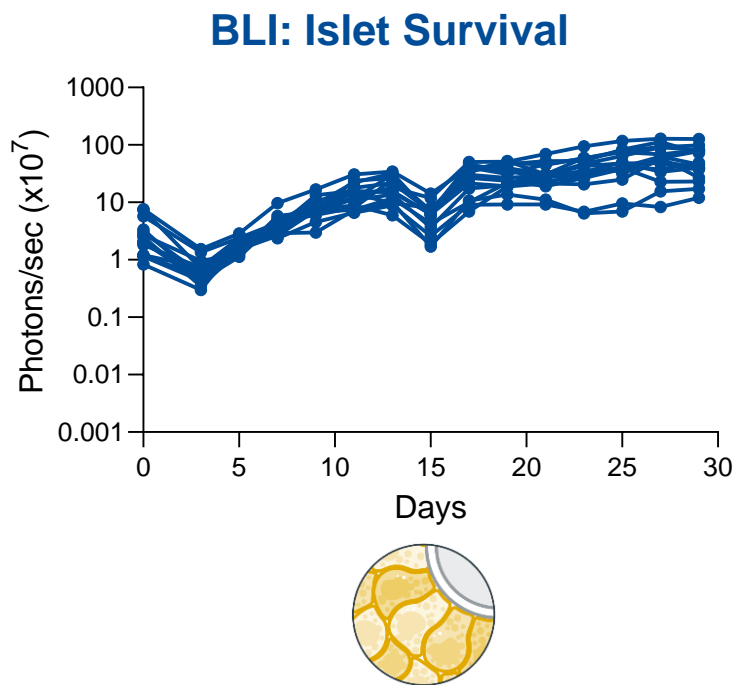


Patient T cells eliminate islet cells due to autoimmunity



Abbreviations: T1DM, type 1 diabetes mellitus
Hu et al. *Sci Transl Med.* 2023

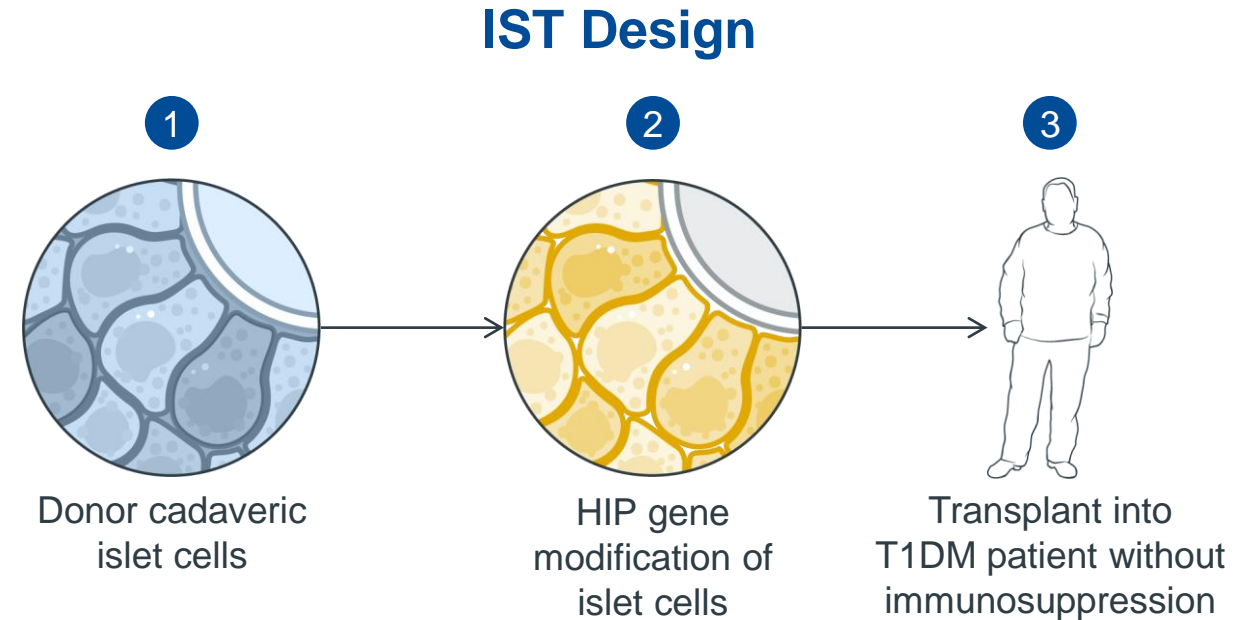
HIP iPSC-derived pancreatic islet cells from patient with T1DM evade autoimmune killing and control glucose



Abbreviations: BLI, bioluminescence imaging
Hu et al. *Sci Transl Med.* 2023.

Potential clinical validation of hypoimmune islet cells in T1DM patients

- Trial authorized at Uppsala University Hospital
- Primary human HIP islet cells transplantation in type 1 diabetes patients
- Intramuscular administration in forearm
- No immunosuppression
- Insights for SC451

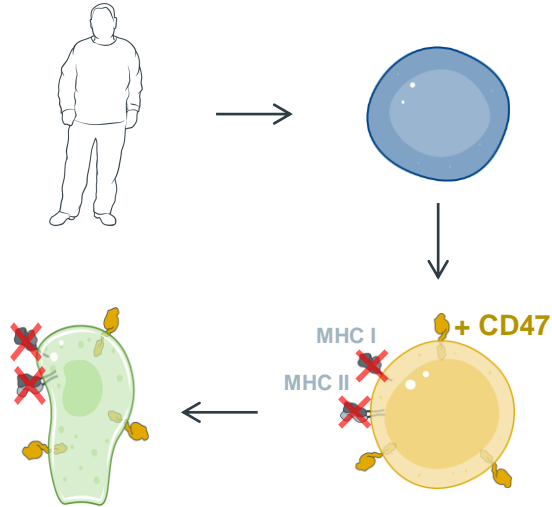


Key Measured Outcomes

Cell survival & immune evasion
C-peptide
Glycemic control

Sana's approach to treat type 1 diabetes

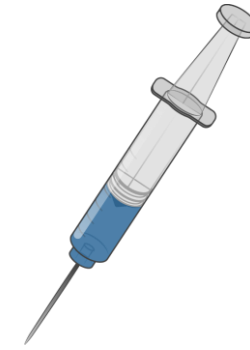
1 Make hypoimmune islet cells from stem cells



2 Manufacture at scale

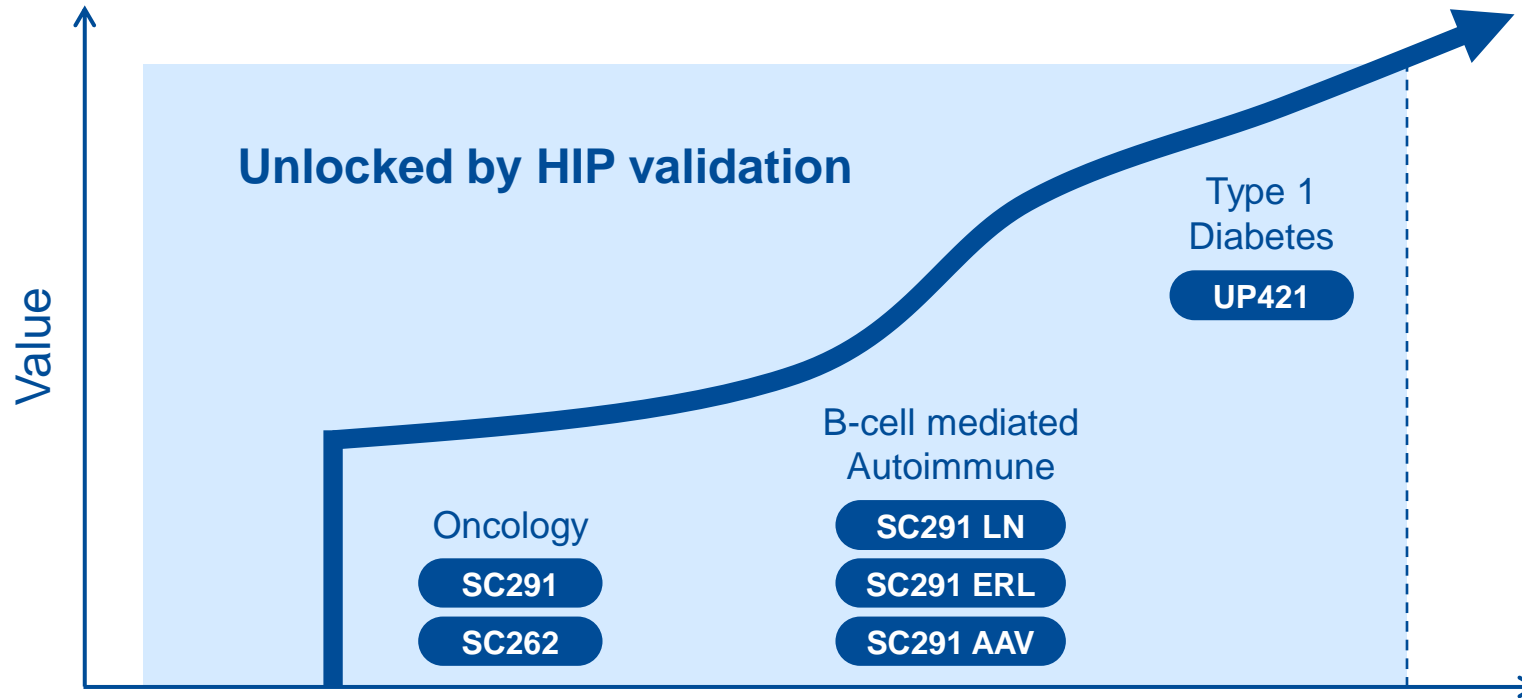


3 Deliver as a single therapy



SC451 program – HIP stem cell-derived islet cell therapy – delivered with no immunosuppression

Meaningful clinical data in multiple diseases in 2024



Unlocking the potential of our hypoimmune platform across multiple patient populations

Thank You

Sana Biotechnology
www.sana.com

