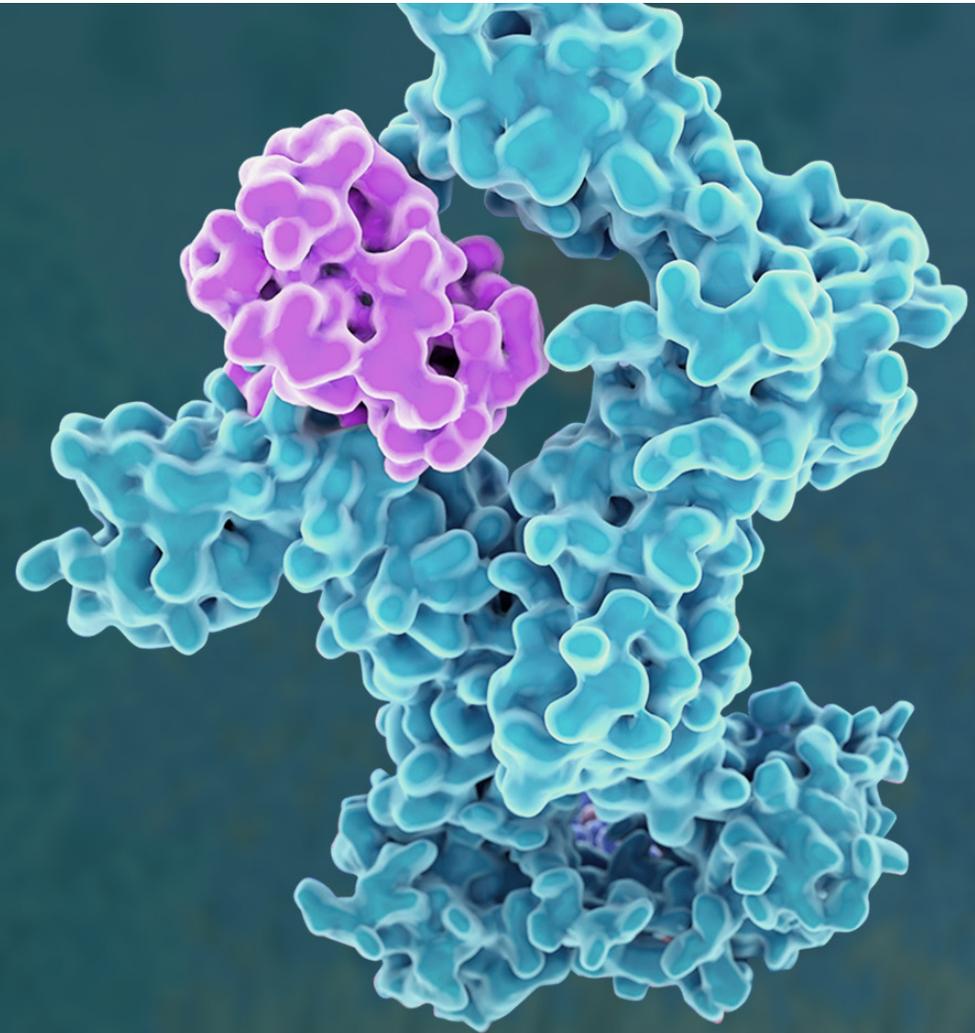


Q3, 2023

Evolutionary Cytokines Revolutionary Medicines



MEDICENNA

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Investment Highlights

Multiple Significant Catalysts Are Expected in The Next Six Months



Superkine Platform: Drug Discovery Engine

Directed evolution **enhances the desired properties** of IL-2, IL-4, & IL-13 to generate Superkines
Protein fusion can **improve PK, add an MOA, or confer new capabilities** to Superkines
IL-2, IL-4, & IL-13 are known to modulate immune activity against **2,000 different diseases**



MDNA11: "Beta-only" & Long-acting IL-2 Super- agonist in Phase 1/2

Super-agonist against IL-2R, a **clinically validated anti-cancer target**
Enhanced IL-2R β binding and lack of IL-2R α affinity position MDNA11 to be **best-in-class**
MDNA11 has **successfully completed** the dose escalation phase 1 with grade 1-2 AEs only
MDNA11 **commences dose expansion** in Q3 2023 as monotherapy and Q4 with pembrolizumab



MDNA55: Phase 3 Ready Empowered IL-4 Superkine

Targeting recurrent glioblastoma, the most aggressive form of brain cancer
Phase 2b data show **~100% improvement in median OS** vs. a matched external control arm
Pursuing a **partnership** to advance development



BiSKIT Platform: Bifunctional SuperKines for ImmunoTherapy

Fusion of two Superkines or a Superkine and an antibody (e.g. a checkpoint inhibitor)
Incorporate **two synergistic MOAs** into a single molecule





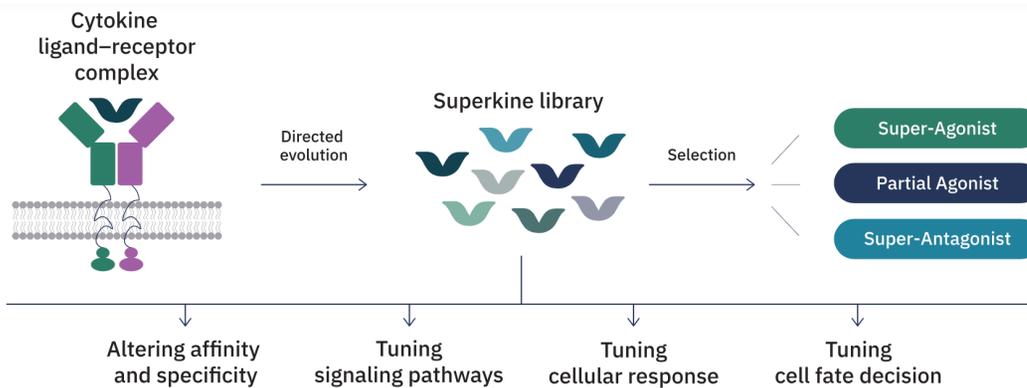
Our Superkine Platform

Our scientific platform enables us to transform natural interleukins into Superkines, which are enhanced to combat a specific disease. For example, Superkines could alter the immunosuppressive tumor microenvironment, deliver cell-killing agents without harming healthy cells, or turn off destructive autoimmune processes.



Superkine Platform Powers Drug Discovery Engine

Transforming IL-2, IL-4 and IL-13 into Druggable Superkines Using Directed Evolution



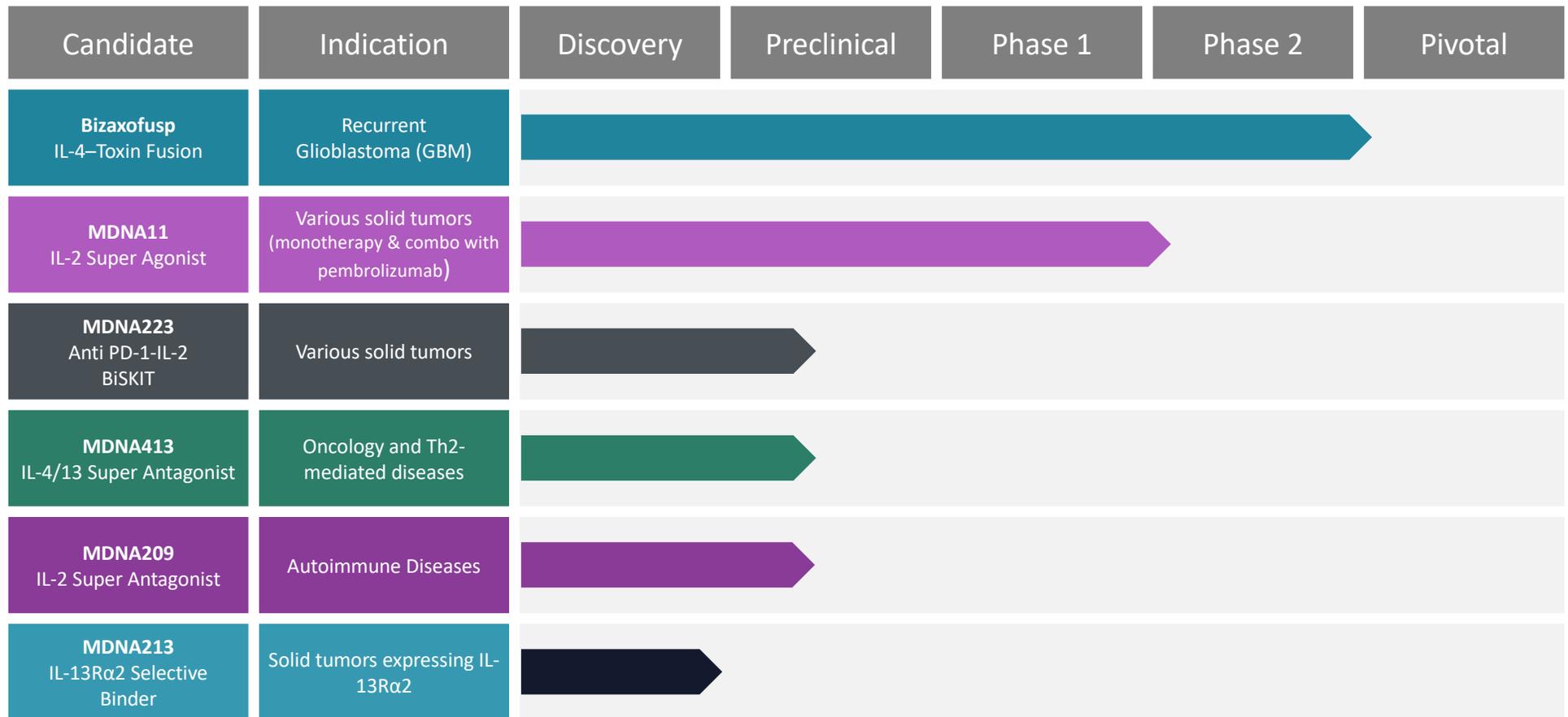
Our IL-2, IL-4 and IL-13 Superkines are known to modulate immune activity in many diseases, each providing “A Pipeline in a Product” opportunity

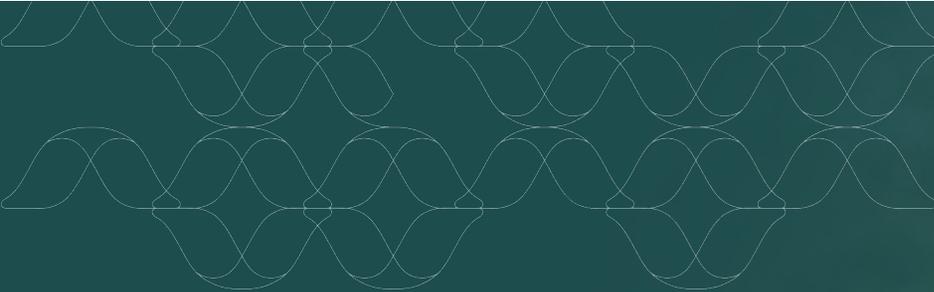
Superkine Design and Development

-  **Generate Tunable Superkine Library**
Transform interleukins using directed evolution to enhance desired properties
-  **Enhance via Protein Fusion**
To improve PK, add a second MOA, or confer new capabilities
-  **Lead Selection & Development**
Advance the most promising candidates towards clinical studies



Our Diversified Pipeline Of Next-Generation Superkines





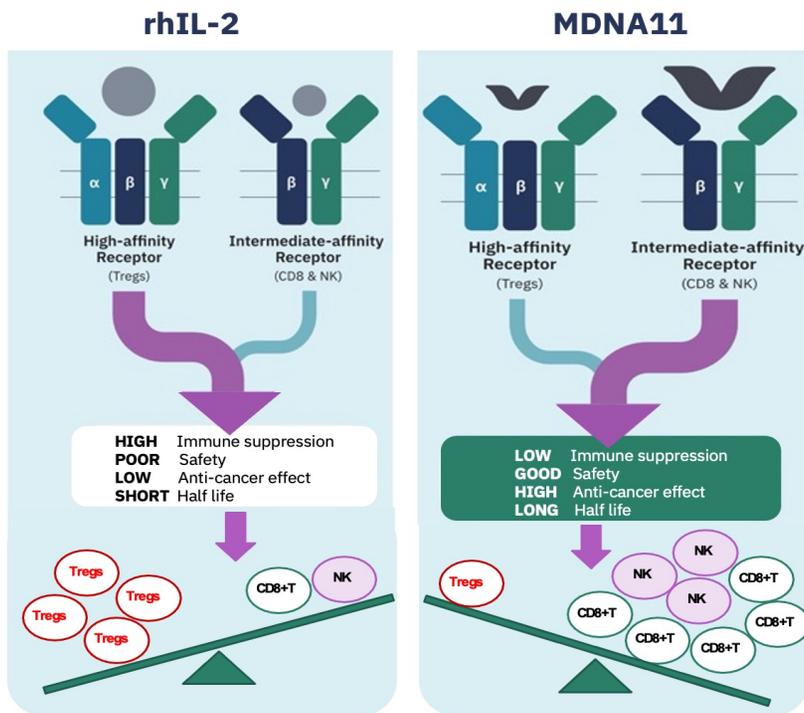
MDNA11

“Beta-only” & Long-acting IL-2
Super-Agonist for Solid Tumors



Targeting IL-2 Receptor Subunits in Cancer Therapy

IL-2 Receptor



The IL-2 receptor (IL-2R) consists of three subunits

- IL-2R α (CD25)
- IL-2R β (CD122)
- IL-2R γ (CD132)

Stimulation of IL-2R β

- Key for the activation of cancer killing immune cells such as CD8+ T cells, naïve T cells, and NK cells.

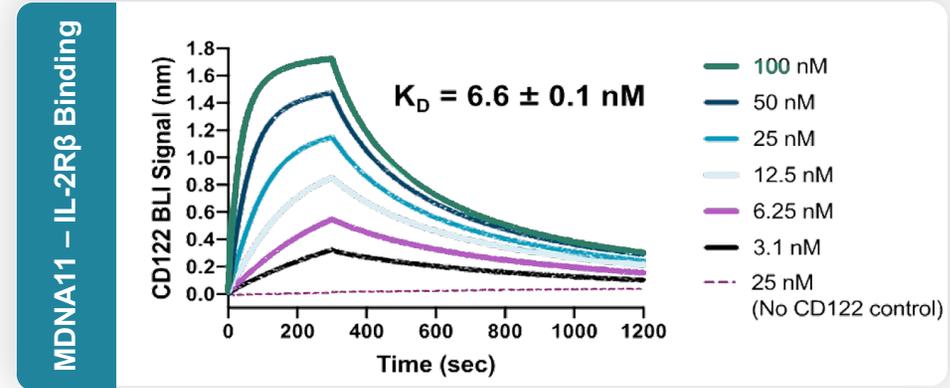
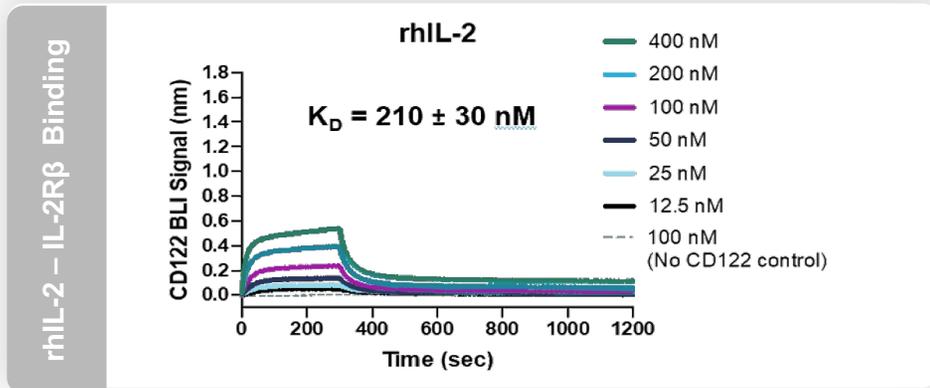
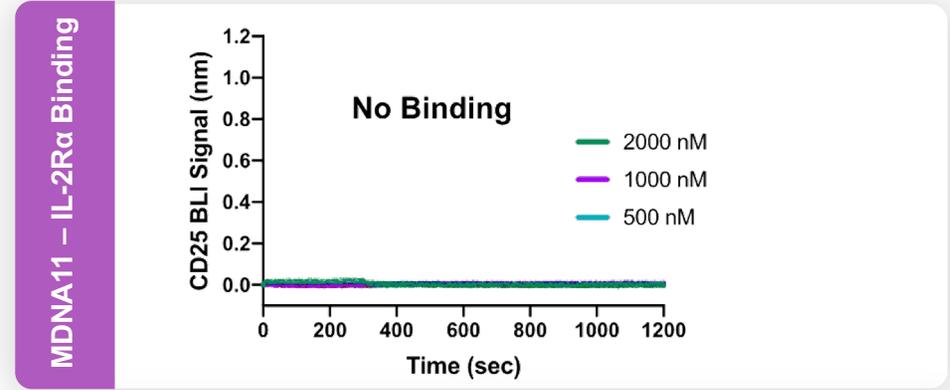
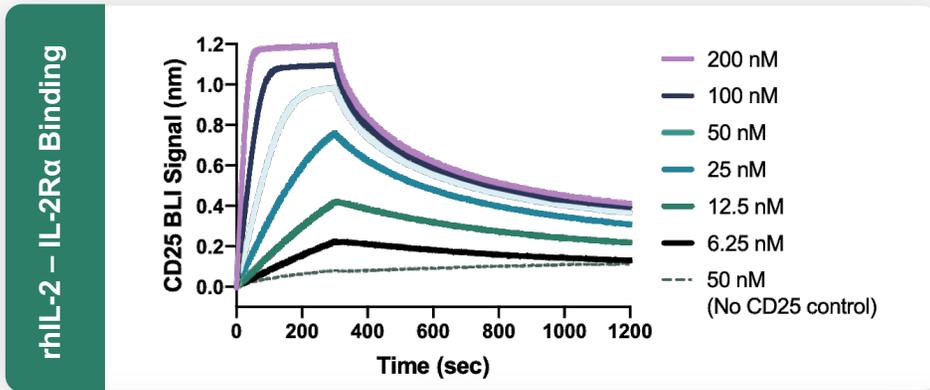
Stimulation of IL-2R α

- Leads to activation of immunosuppressive Tregs, which abrogate the anti-tumor response
- Causes extreme toxicity

Proleukin (recombinant human [rh] IL-2), which selectively stimulates IL-2R α , is approved by the FDA for the treatment of metastatic melanoma and renal cell carcinoma.

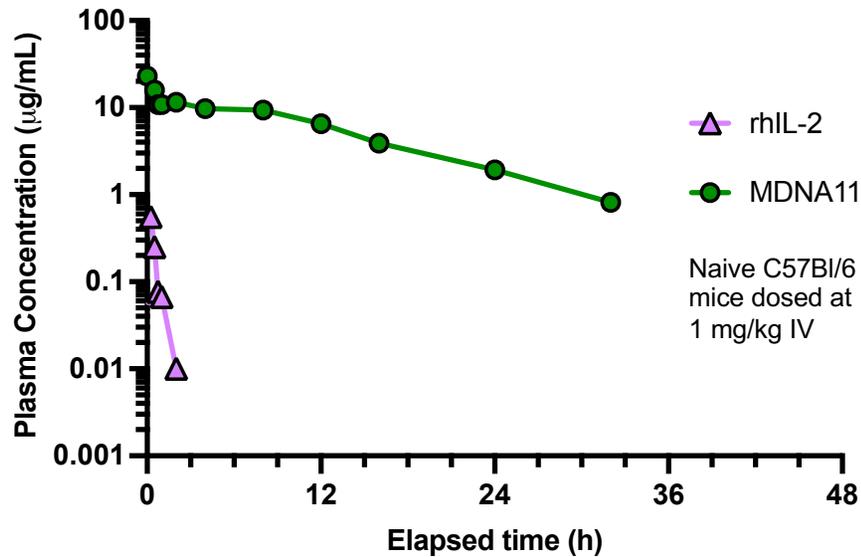
MDNA11's IL-2 Binding is Highly Differentiated vs. Proleukin (rhIL-2)

No IL-2R α (CD25) Binding and Enhanced Affinity and Selectivity for IL-2R β (CD122) Compared to rhIL-2



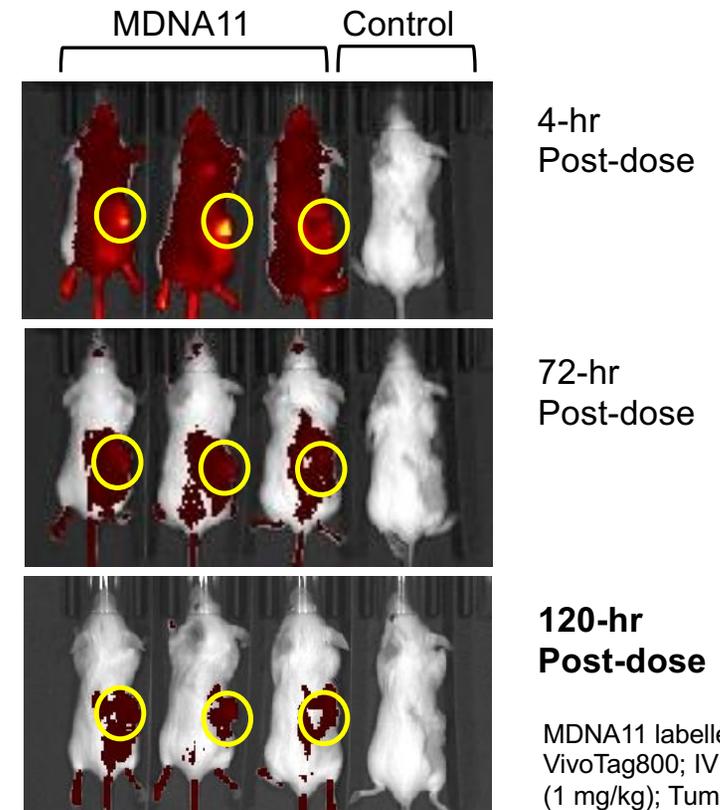
MDNA11 Durably Accumulates In Solid Tumors In Vivo

PK Profile in Mice



	C_{max} (µg/mL)	AUC (µg.hr/mL)	T_{half} (hr)
rhIL-2	5.77	1.07	0.28
MDNA11	23.02	182.3	6.83

MDNA11 Imaging in CT26 Tumor Model



MDNA11 labelled with VivoTag800; IV dosing (1 mg/kg); Tumor size: 150-200 mm³

Tumor exposure of MDNA11 is > 15x longer than its serum half-

MDNA11 – Best in Class Potential

MDNA11's strong anti-tumor activity, well-tolerated safety profile and convenient outpatient dosing regimen paves the way for a potential best-in-class therapy with significant commercial potential

	 MDNA11	 Proleukin ¹	 NKTR-214	 SAR'245 ²	 ALKS 4230 ³	 WTX-124 ⁵	 XTX202 ⁶	 STK-012 ⁷	 TransCon IL-2 β/γ ⁸
No binding to IL-2R α	✓	X	X	✓	✓	X	✓	X	Minimal binding
Enhanced IL-2R β/γ Binding	✓	X	X	X	X	X	X	X	✓
QW, Q2W or Q3W Dosing	✓	X	✓	✓	X	Unknown	✓	✓	✓
Tumor Accumulation	✓	X	X	X	X	X	X	X	X
No Pegylation Liabilities	✓	✓	X	X	✓	✓	✓	X	✓
Pipeline Potential	✓	✓	X	X	X	X	X	✓	✓



ABILITY: Phase 1/2 Dose Escalation & Expansion Study

Monotherapy Dose Escalation

N = 20 patients with advanced, treatment-refractory solid tumors

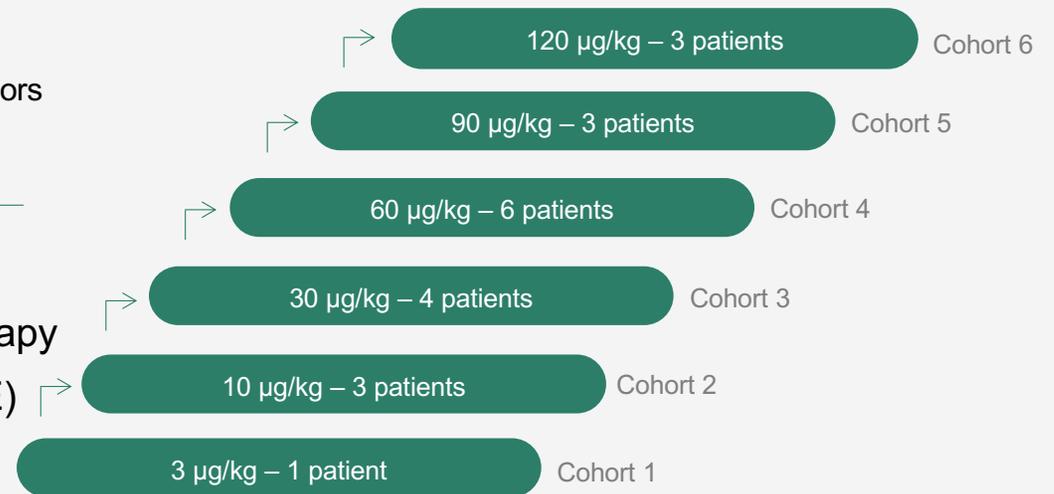
MDNA11 Q2W IV; cut-off date: June 20, 2023

Modified 3+3 design. Open-label

Assess safety & tolerability of MDNA11 monotherapy

Identify Recommended Dose for Expansion (RDE)

[NCT05086692](https://www.clinicaltrials.gov/ct2/show/study/NCT05086692)



MDNA11 Monotherapy Dose Expansion

N≈40: Melanoma and other selected solid tumors

MDNA11 Q2W IV at RDE

Further evaluate safety and tolerability

Evaluate single-agent anti-tumor activity

MDNA11 + Anti-PD-1 (Pembrolizumab) Dose Expansion

N≈40: Melanoma and other selected solid tumors

MDNA11 + anti-PD-1 (pembrolizumab)

Evaluate safety and tolerability of MDNA11 / anti-PD-1 combination

Evaluate combination anti-tumor activity



ABILITY: Patient Baseline Characteristics in Dose Escalation

Demographics/Performance

Median age, years (range)	61 (27-78)
Male (%)	16/20 (80%)
ECOG 0	14/20 (70%)
ECOG 1	6/20 (30%)

Prior Systemic Therapies

Prior Lines of Therapy: 1	5/20 (25%)
Prior Lines of Therapy: 2-4	15/20 (75%)
Prior Immunotherapy	15/20 (75%)
Prior Targeted Therapy	5/20 (25%)
Prior Chemotherapy	9/20 (45%)

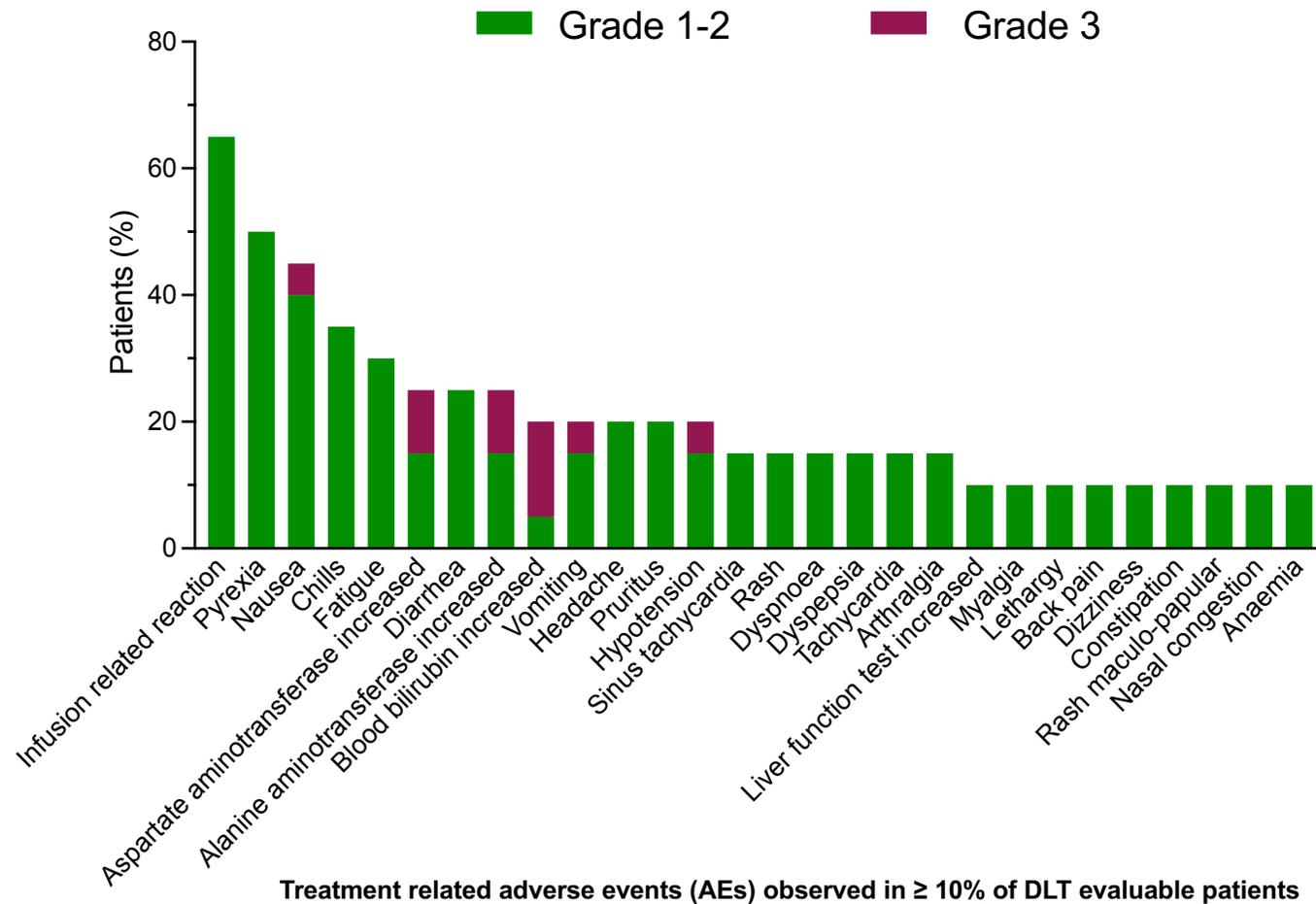
Primary Cancer Diagnosis

Melanoma	11/20 (55%)
Renal Cell Carcinoma (non-clear cell)	2/20 (10%)
Pancreatic Ductal Adenocarcinoma (PDAC)	2/20 (10%)
Sarcoma	2/20 (10%)
Squamous Cell Carcinoma	1/20 (5%)
Gastro-esophageal Adenocarcinoma	1/20 (5%)
Lung Adenocarcinoma	1/20 (5%)

Most patients received multiple prior lines of anti-cancer therapy, including immunotherapy



Single Agent Safety Profile Across all Dose Escalation Cohorts



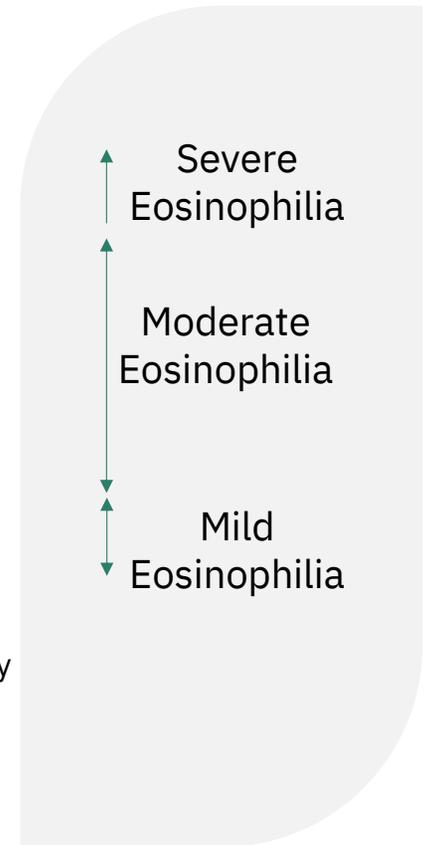
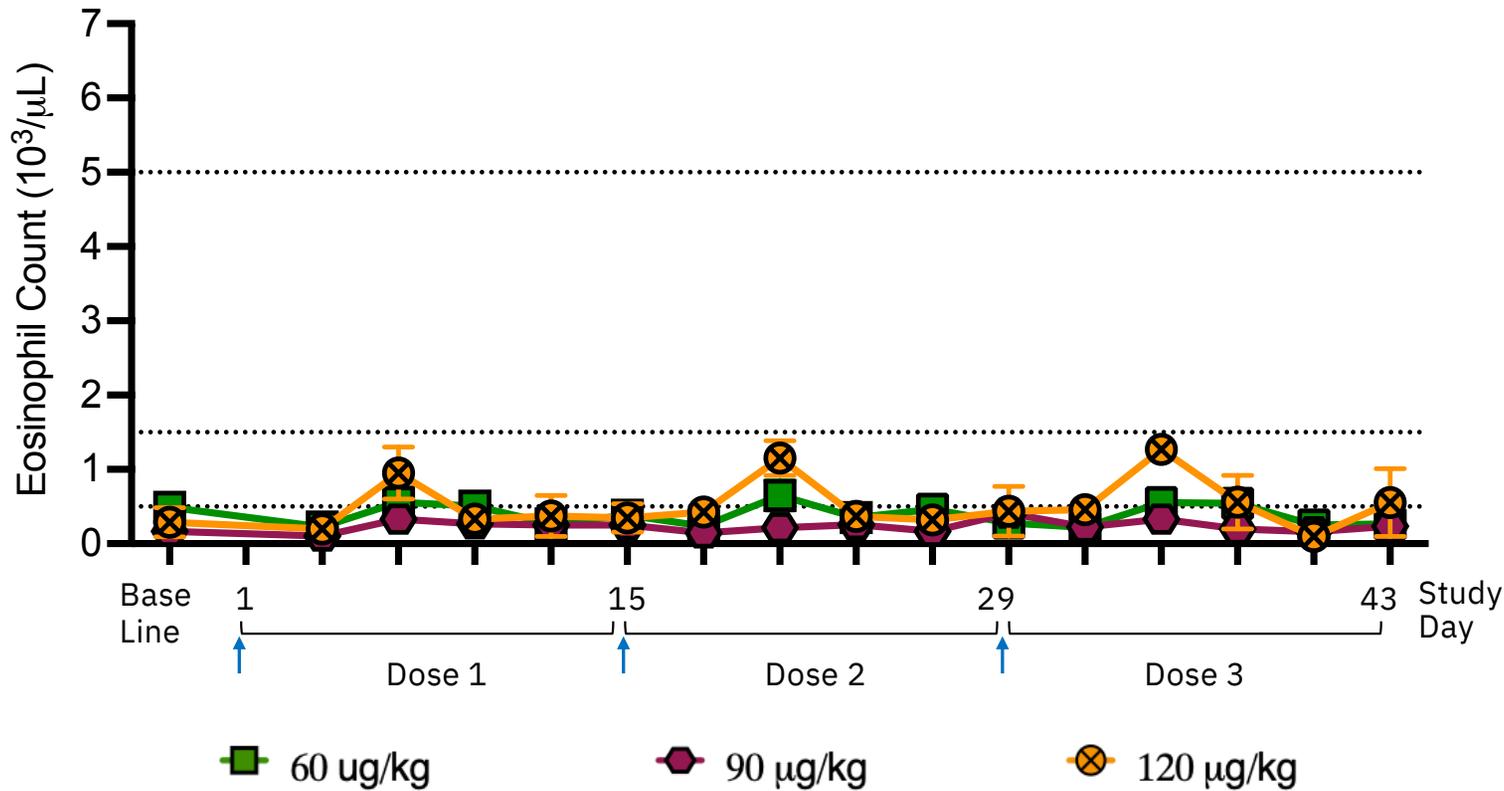
No DLTs or MTD reached
 No Grade 4/5 events
 Majority of AEs Grade 1/2 and resolved within 1-2 days
 Data cut-off: June 20, 2023

MNDA11 generally well-tolerated across cohorts

Treatment related adverse events (AEs) observed in ≥ 10% of DLT evaluable patients

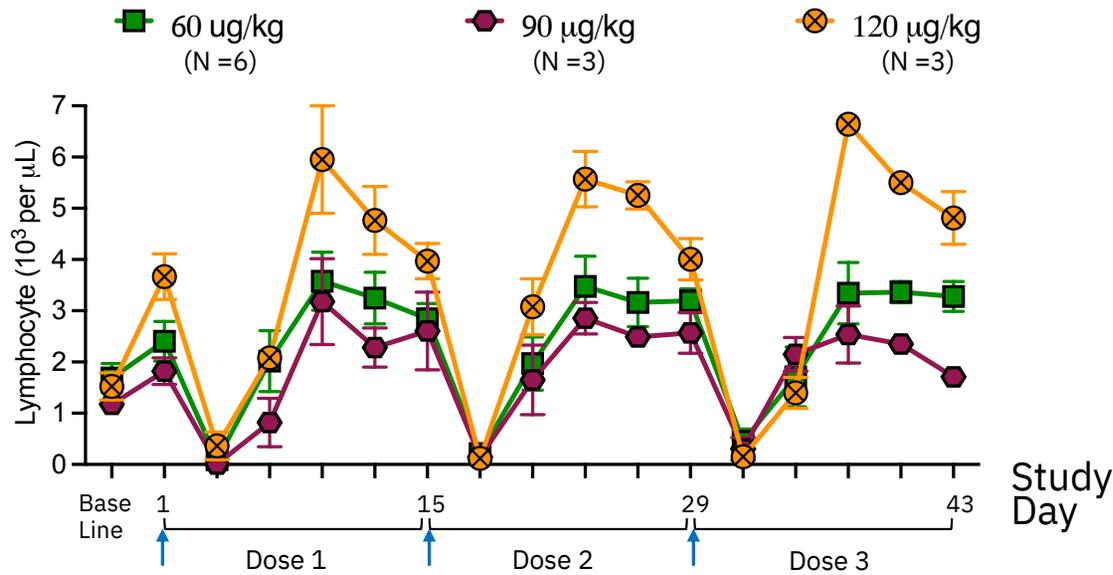
No Significant Eosinophilia (Associated with VLS)

Vascular Leak Syndrome (VLS) is a hallmark dose-limiting toxicity of IL-2



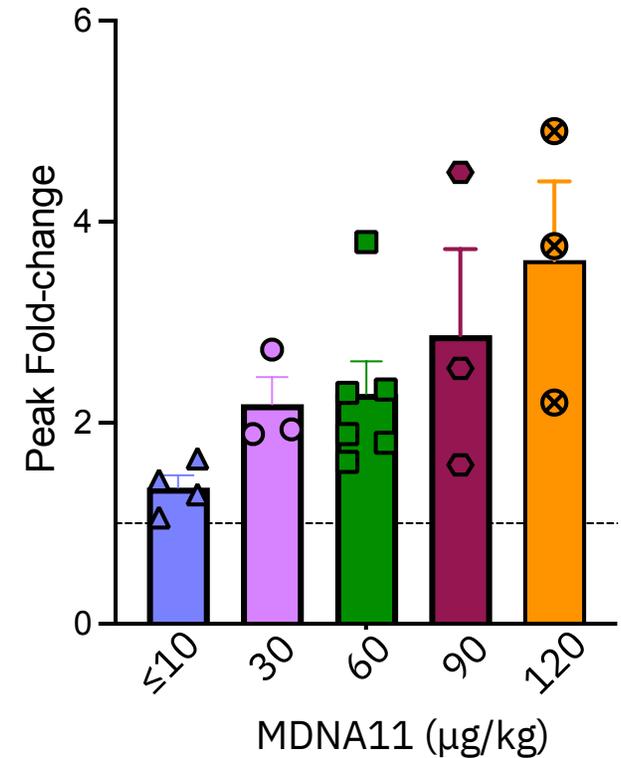
MDNA11-Induced Sustained Dose-Dependent Lymphocyte Expansion

Expansion of cancer killing immune cells



	60 µg/kg	90 µg/kg	120 µg/kg
Average Baseline Lymphocyte (10 ³ per µL)	1.48	1.17	1.52
Median Age, years (Range)	58 (27-78)	74 (73-75)	49 (41-67)

Lymphocyte Peak Fold-Change



Peak fold-change relative to baseline.

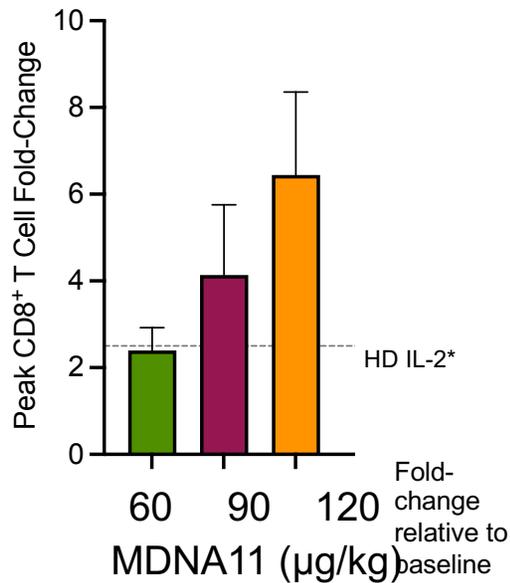
Graph shows mean ± SEM

For < 10 µg/kg and 30 µg/kg, peak data for Dose 3

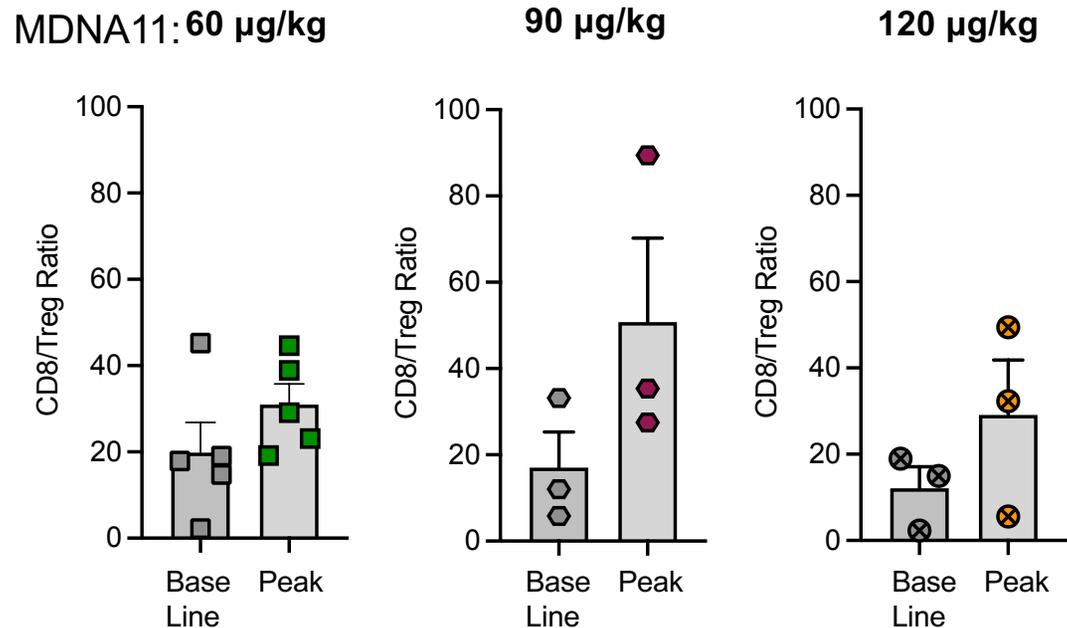
For 60 µg/kg, 90 µg/kg and 120 µg/kg, peak data for Target Dose 1

MDNA11 Preferentially Induced CD8⁺ T Cell Expansion Over Tregs

Peak Increase: CD8⁺ T Cell



Peak Increase: CD8⁺/Treg Ratio

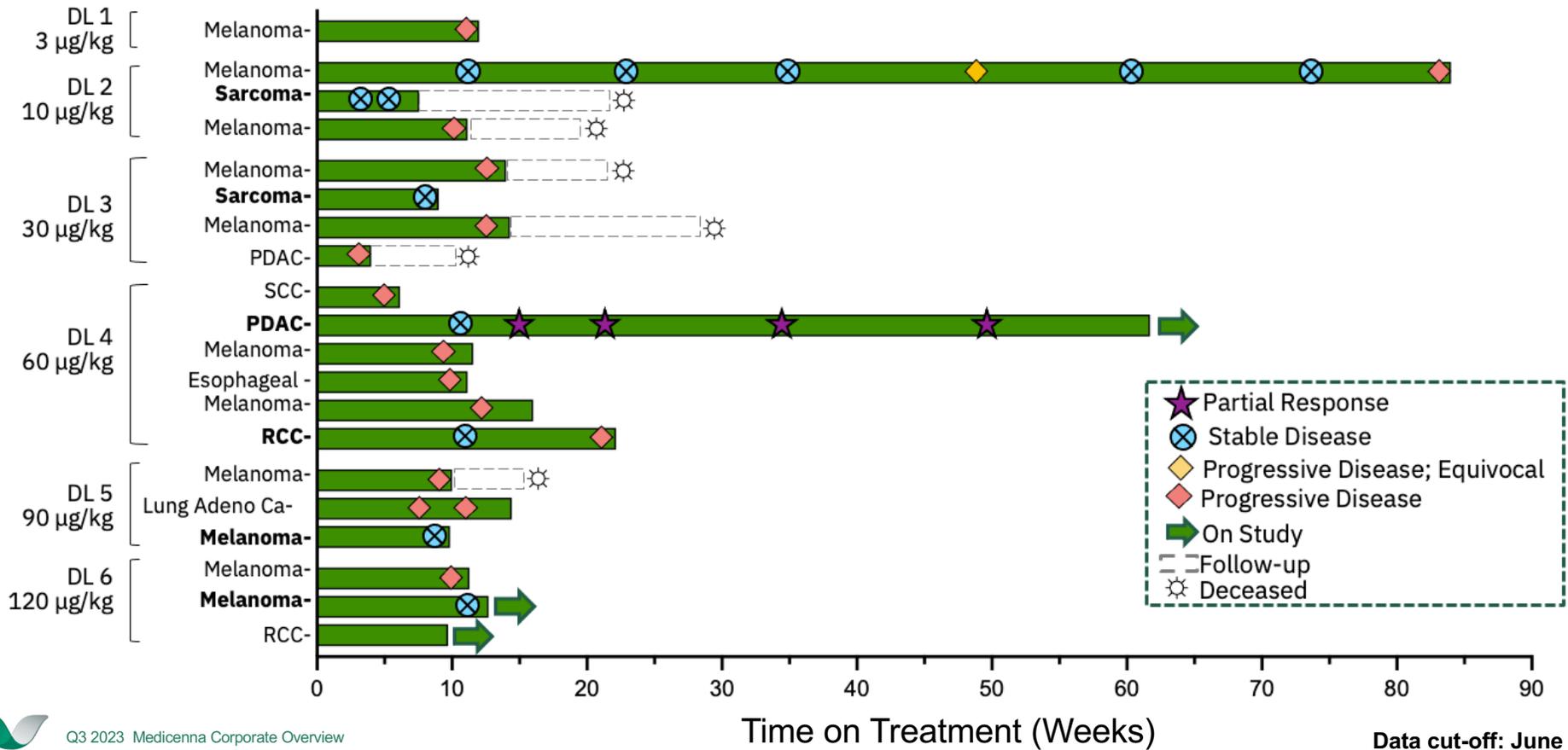


CD8⁺ T cells are powerful effectors of the anti-cancer immune response

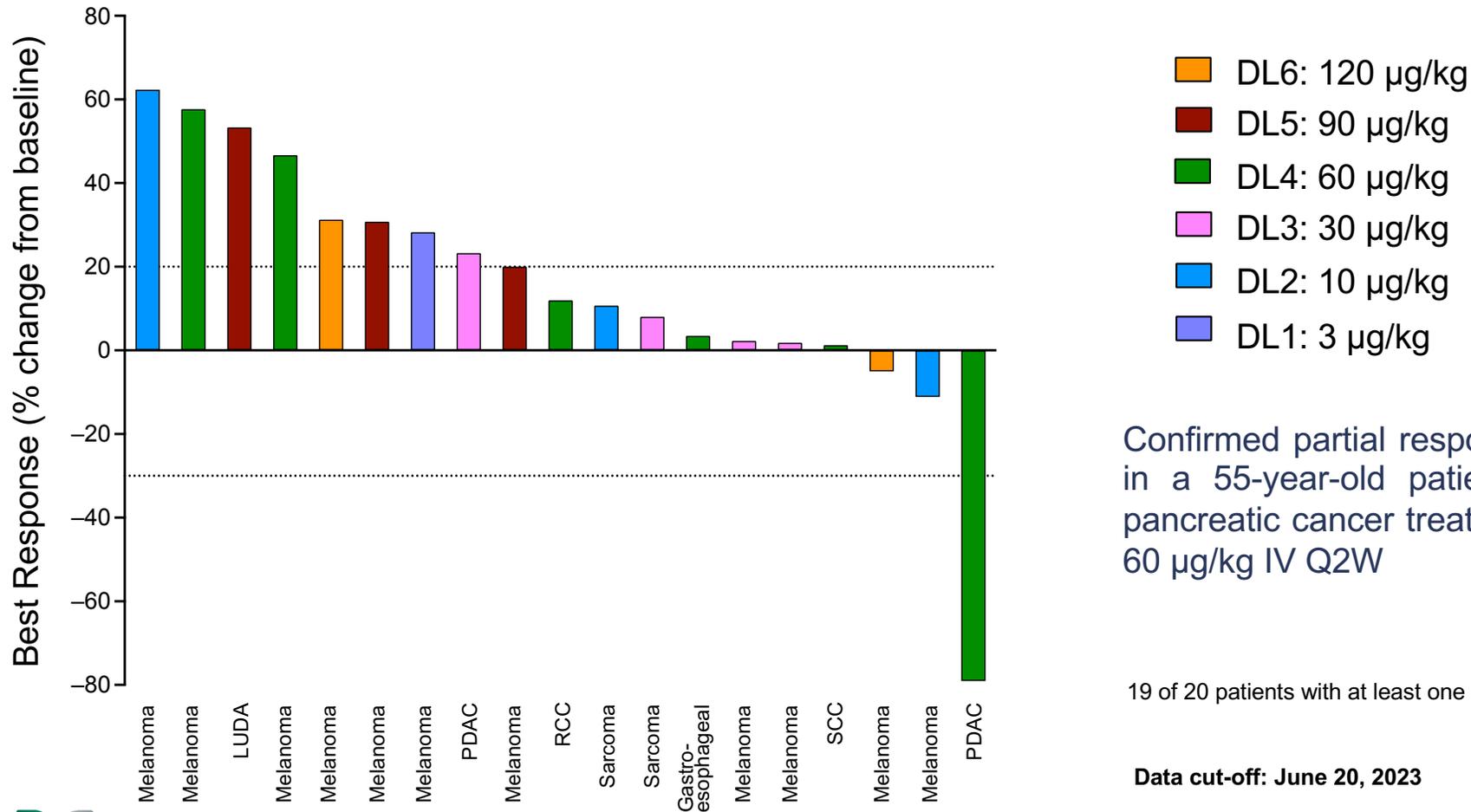


Durable Responses Observed During Dose Escalation

SD lasting ~18 months in 71-year-old patient with metastatic melanoma treated with 2 prior lines of IO



MDNA11 Shows Single Agent Clinical Activity



Confirmed partial response (PR) achieved in a 55-year-old patient with metastatic pancreatic cancer treated with MDNA11 at 60 µg/kg IV Q2W

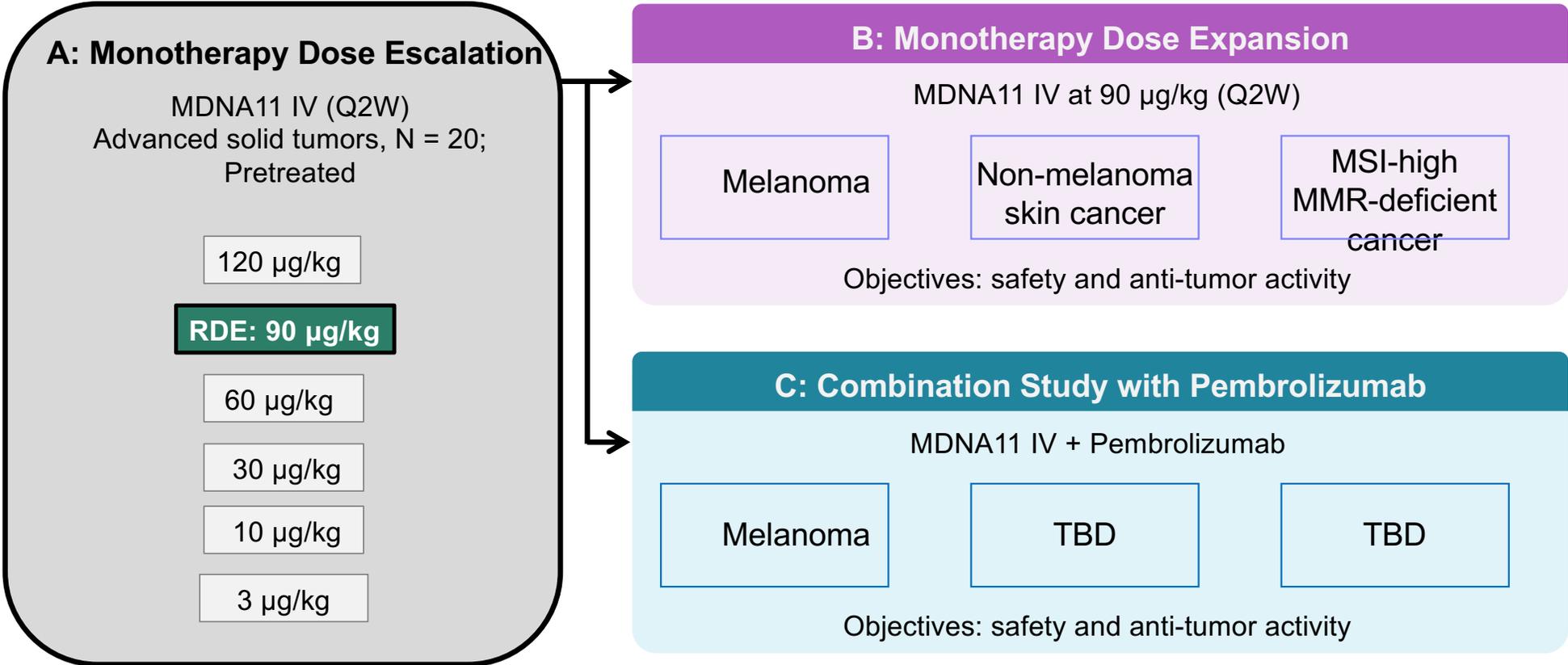
19 of 20 patients with at least one follow-up assessment

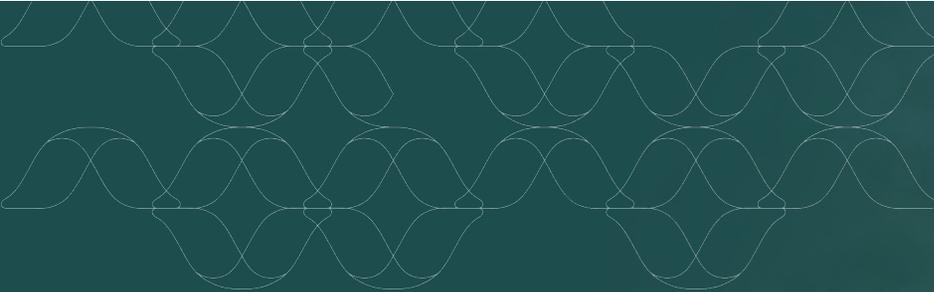
Data cut-off: June 20, 2023

ABILITY Study Plan – Dose Expansion & Combination Phase

Global, multi-center, open-label Phase 1/2 study

Monotherapy dose escalation and expansion; combination with Pembrolizumab



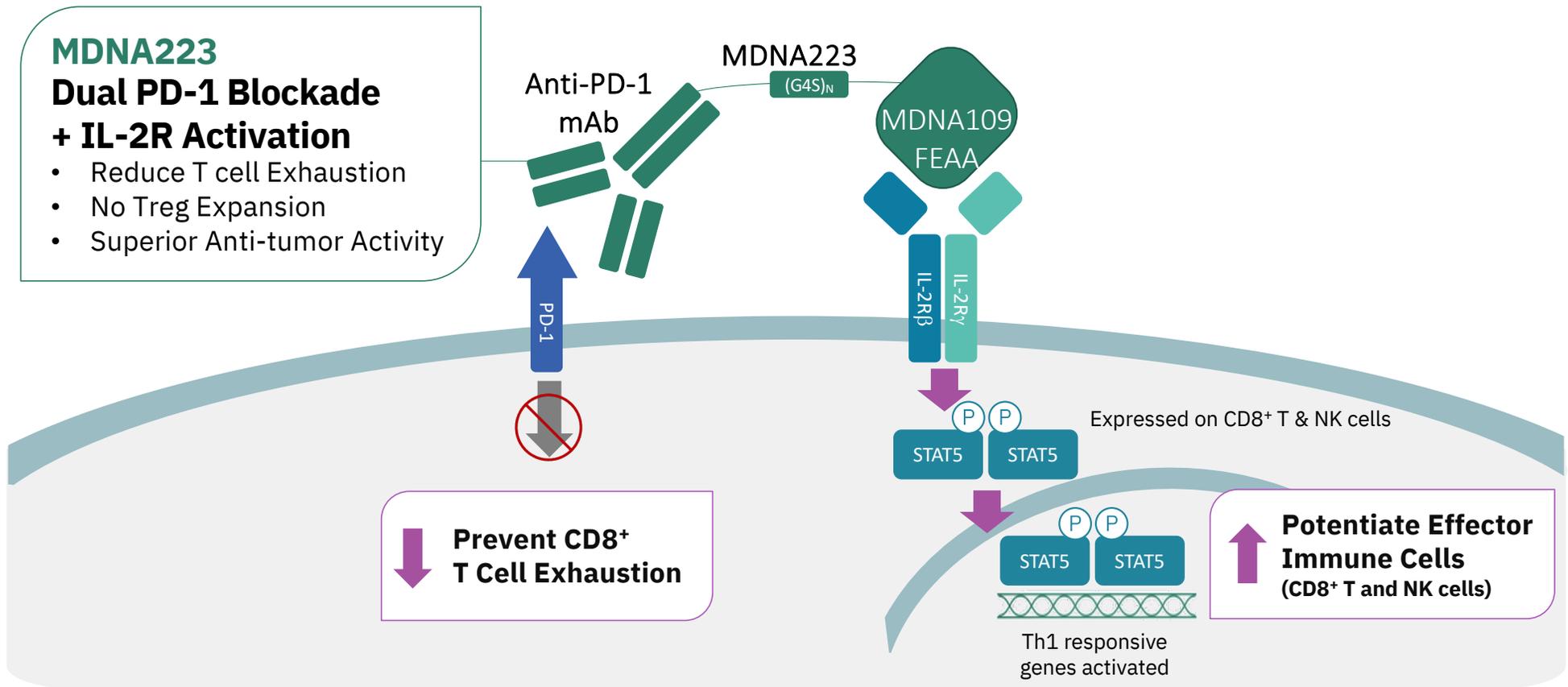


Bifunctional SuperKines for ImmunoTherapy (BiSKIT)



MDNA223: Anti-PD1-IL-2 Superkine BiSKIT

Synchronized cis-binding for PD-1 blockade and IL-2R activation on same CD8⁺ T or NK cell



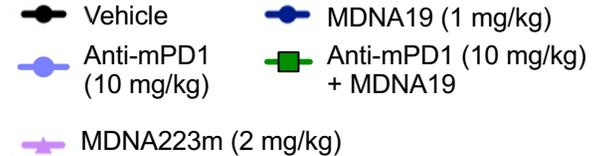
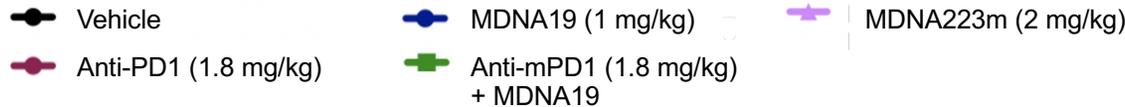
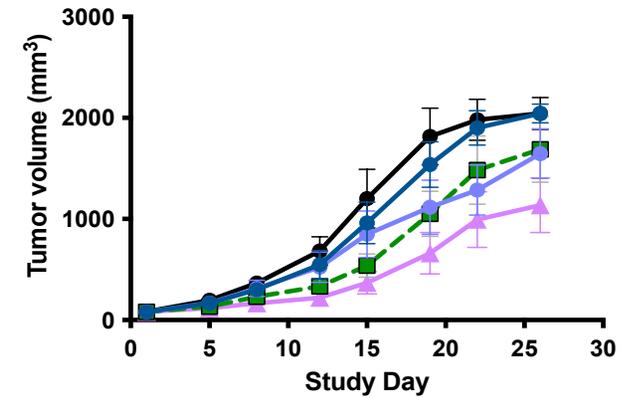
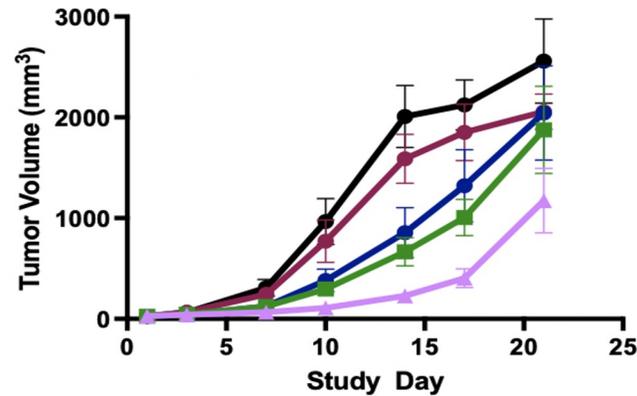
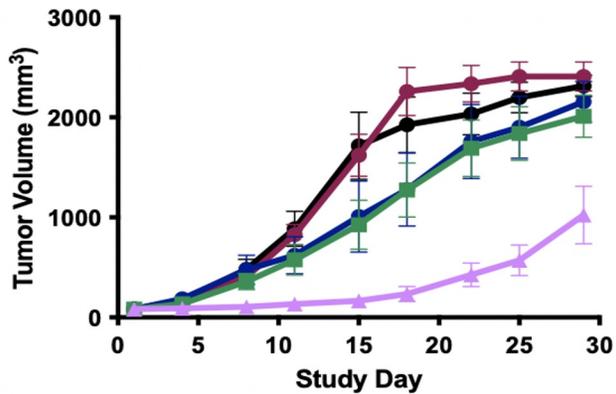
MDNA223m Demonstrated Superior Anti-Tumor Activity

MDNA223m showed higher levels of anti-tumor activity than co-administration in pre-clinical studies

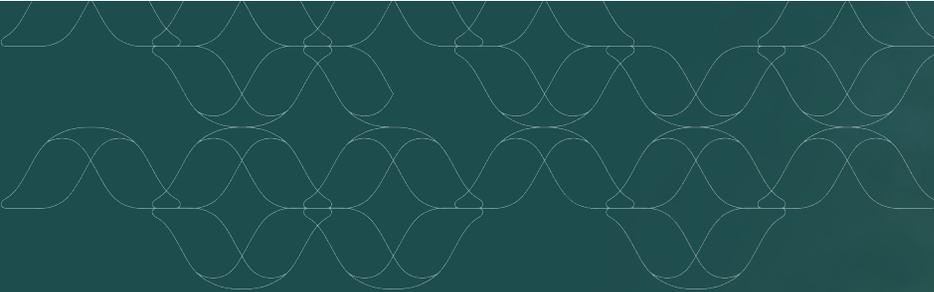
CT26 Colon Tumor Model

B16F10 Melanoma Model

E0771 Breast Tumor Model



Treatment with molar equivalent doses of anti-PD1 (150 Kda), MDNA19 (83 Kda) or MDNA223m (165 Kda).
 IP treatment QWx2 (CT26 and E0771) and QWx3 (B16F10)
 Avg tumor size at initiation of dosing: 127 mm³ (CT26), 80 mm³ (E0771) or 30 mm³ (B16F10)



MDNA55

Empowered IL-4
Superkine Targeting
Glioblastoma



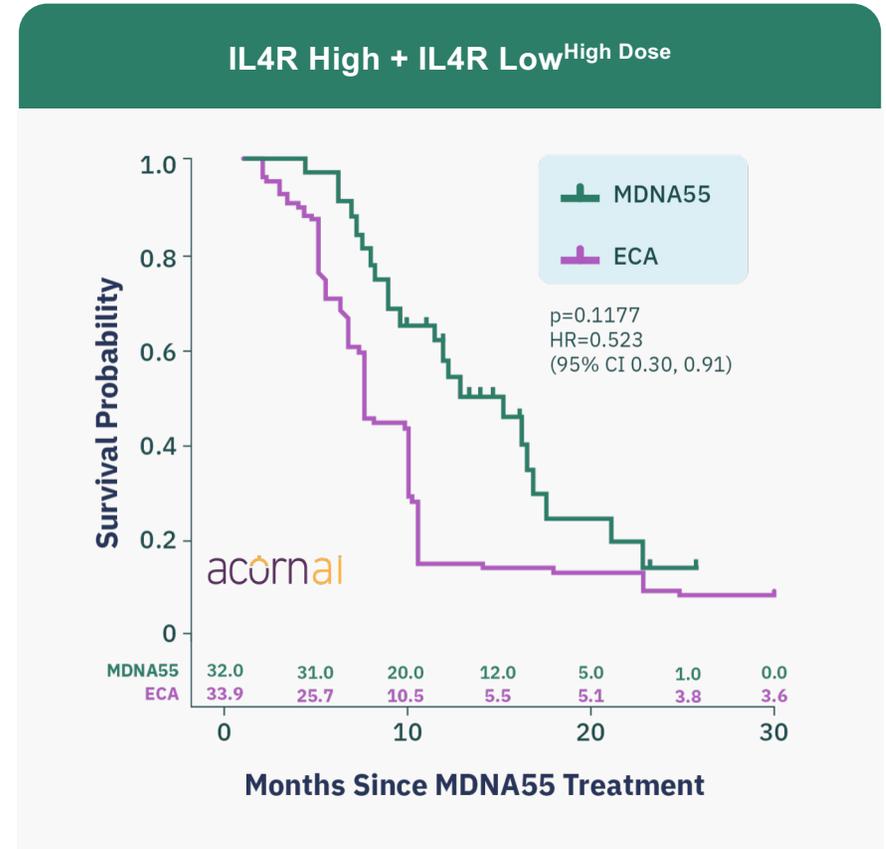
Improvement of ~ 100% in mOS vs External Control Arm (ECA)

Results*

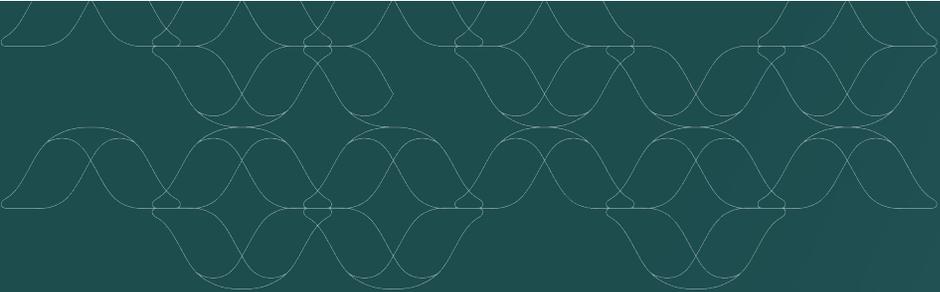
Weighted IL4R High + IL4R Low^{High Dose} (n=32)
mOS is 15.7 months vs 7.2 months in ECA

→ Survival time more than doubled in the IL4R High + IL4R Low^{High Dose} group compared to ECA

Bisaxofusp (MDNA55) is Phase 3 ready in recurrent glioblastoma



*Survival was calculated from time of relapse



Next Milestones



Upcoming Anticipated Milestones & Financial Summary

ABILITY Study Fully Funded – Cash Runway Through Q3 2024

Anticipated Milestones

Start of ABILITY monotherapy expansion	Q3 2023
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Update from ABILITY monotherapy expansion	Q4 2023
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Commence combination phase of ABILITY with MDNA11 & pembrolizumab	Q4 2023
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Update from ABILITY mono- and combination phases	Q1 2024
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Financial Highlights

Nasdaq/TSX	MDNA
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Headquarters	Toronto, CA
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Cash	CDN \$29.6M*
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Debt	\$0
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Preferred Shares	None
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Issued and Outstanding	~70 Million*
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Fully Diluted	~92 Million*
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* As of June 30, 2023.



Thank you

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