SEPTEMBER 21, 2022

Evolutionary Cytokines Revolutionary Medicines





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Investment Highlights

Clinical Data Updates from MDNA11 Program Expected Throughout 2022

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-2Rβ binding and lack of IL-2Rα affinity position MDNA11 to be best-in-class Clinical data updates expected throughout 2022	
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 Pursing a partnership to advance development	
BiSKIT Platform: <u>Bi</u> functional <u>S</u> uper <u>K</u> ines for <u>I</u> mmuno <u>T</u> herapy	Fusion of two Superkines or a Superkine and an antibody (e.g. a checkpoint inhibitor) Incorporate two synergistic MOAs into a single molecule	

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

MDNA11

"Beta-only" & Longacting IL-2 Super-Agonist for Solid Tumors



Targeting IL-2 Receptor Subunits in Cancer Therapy



Mullard, Nat Rev Drug Discovery 20(3): 163-165, 2021.

- > The IL-2 receptor (IL-2R) consists of three subunits
 - CD25 (IL-2Rα)
 - CD122 (IL-2Rβ)
 - CD132 (IL-2Rγ)

Stimulation of CD122

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

Stimulation of CD25

- Leads to activation of immunosuppressive Tregs, which abrogate the anti-tumor response
- Causes extreme toxicity (i.e., pulmonary edema, vascular leak syndrome)

Proleukin® (recombinant human rhIL-2), which selectively stimulates CD25, is approved for the treatment of metastatic melanoma and renal cell carcinoma

IL-2 Transaction Landscape

Bristol Myers and Nektar

NEKTAR[®] Bristol-Myers Squibb

\$3.6 Billion licensing transaction

Sanofi and Synthorx

synth@rx

sanofi

\$2.5 Billion acquisition – Phase 1

Merck and Pandion



\$1.85 Billion acquisition – Phase 1

Roche and Good Therapeutics

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\$250M acquisition – Preclinical

Q3 2022 Medicenna Corporate Overview

MDNA11 Compares Well With Other IL-2 Programs

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

	MDNA11	CLINIGEN Proleukin ¹	SOR'245 ²	Alkermes ALKS 4230 ³	NL-2014	Werewolf THERAPEUTICS WTX-124 ⁵	XTX2026	Synthekine STK-0127
No binding to IL-2Rα	V	X	V	V	V	×	V	×
Enhanced IL-2Rβγ Binding	V	X	X	X	V	×	×	×
Q2W or Q3W Dosing	V	X	V	×	V	Unknown	V	V
Albuminated	V	×	×	×	×	×	×	×
No Pegylation Liabilities	V	V	X	V	×	V	V	X

[1] Nature Rev. Drug Discovery (2021). [2] Ptacin et al., Nat Comm (2021). [3] Lopes et al., JITC (2020). [4] Da Silva et al., Nature (2019).
[5] Nirschl et al, Cancer Immunol Res (2022). [6] O'Neil et al., ASCO (2021). [7] Oft et al, AACR (2021). Additional information from https://clinicaltrials.gov/



Phase 1/2 ABILITY Study Schema: Enrolling Dose Level 5



before target dose

Protocol Version 5

MDNA11 Induced Lymphocyte Expansion



DL4 patients received 2 priming doses (30 $\mu g/kg$ Q2W) prior to target dose (60 $\mu g/kg$ Q2W) Graph shows mean ± SEM.

AUC measured as area between minimum lymphocyte count values

Q3 2022 Medicenna Corporate Overview

Graph shows mean ± SEM

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

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

No Evidence of Eosinophilia Associated with VLS



DL4 patients received 2 priming doses (30 μ g/kg Q2W) prior to start of target dose of 60 μ g/kg (Q2W) Graphs show mean ± SEM.

Q3 2022 Medicenna Corporate Overview

Duration of Treatment and Summary of Response

 Tumor control in 4 of 10 evaluable patients (including 1 unconfirmed PR) despite low dose levels and heavily pre-treated patients



¹ 1 – Patient demonstrated stable disease at 12 week scan and an unconfirmed partial response (per RECIST 1.1) at a 16 week scan on August 1, 2022 which is preliminary and must be confirmed by a second scan at equal to or greater than 28 days

MDNA55

Empowered IL-4 Superkine Targeting Glioblastoma



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



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





Planned MDNA55 Phase 3 Trial – Hybrid Design With ECA

Hybrid Design Trial with an External Control Arm



Study Assumptions

- 90% power
- HR of MDNA55 vs. pooled control = 0.65
- 2-sided alpha = 0.05
- Effect size = 4.6 months in mOS time
- Drop-out rate = $\sim 5\%$

Upcoming Anticipated Milestones & Financial Summary

2022/2023 Anticipated Milestones

- Fourth Dose Cohort Initial Anti-Tumor Activity Data (September 2022)
- Fifth Dose Cohort Initial Anti-Tumor Activity Data (Q4 2022)
- Single Agent Expansion Anti-Tumor Activity Data (Mid 2023)
- Combination Study Top-Line Anti-Tumor Activity Data (2H 2023)

Financial Highlights

Nasdaq/TSX
Headquarters
Cash
Debt
Preferred Shares
Issued and Outstanding
Fully Diluted

MDNA Toronto, CA CDN \$19.3M** +US\$20M 08/22 \$0 None ~70 Million*

~92 Million*

*As of August 12, 2022 **As of June 30, 2022



Thank you

Fahar Merchant, PhD President and CEO

Elizabeth Williams Chief Financial Officer

