



# **Corporate Presentation**

**February 2024**

[www.matinasbiopharma.com](http://www.matinasbiopharma.com)  
NYSE American: MTNB



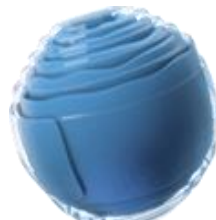
# Forward Looking Statements



This presentation contains "forward-looking statements" within the meaning of the Private Securities Litigation Reform Act of 1995, including those relating to the Company's product development, clinical and regulatory timelines, market opportunity, cash flow and other statements that are predictive in nature, that depend upon or refer to future events or conditions. All statements other than statements of historical fact are statements that could be forward-looking statements. Forward-looking statements include words such as "expects," "anticipates," "intends," "plans," "could," "believes," "estimates" and similar expressions. These statements involve known and unknown risks, uncertainties and other factors which may cause actual results to be materially different from any future results expressed or implied by the forward-looking statements. Forward-looking statements are subject to a number of risks and uncertainties, including, but not limited to, our ability to obtain additional capital to meet our liquidity needs on acceptable terms, or at all, including the additional capital which will be necessary to complete the clinical trials of our product candidates; our ability to successfully complete research and further development and commercialization of our product candidates; the uncertainties inherent in clinical testing; the timing, cost and uncertainty of obtaining regulatory approvals; our ability to protect the Company's intellectual property; the loss of any executive officers or key personnel or consultants; competition; changes in the regulatory landscape or the imposition of regulations that affect the Company's products; and the other factors listed under "Risk Factors" in our filings with the SEC, including Forms 10-K, 10-Q and 8-K. Investors are cautioned not to place undue reliance on such forward-looking statements. Except as may be required by law, the Company does not undertake any obligation to release publicly any revisions to such forward-looking statements to reflect events or circumstances after the date hereof or to reflect the occurrence of unanticipated events. Matinas BioPharma's product candidates are all in a development stage and are not available for sale or use.

# Matinas Investment Thesis: LNC Delivery Unlocks Therapeutic Value

## Lipid Nanocrystals (LNCs)



- Intracellular delivery
- Oral administration
- Less toxicity
- Targeting beyond the liver

### MAT2203

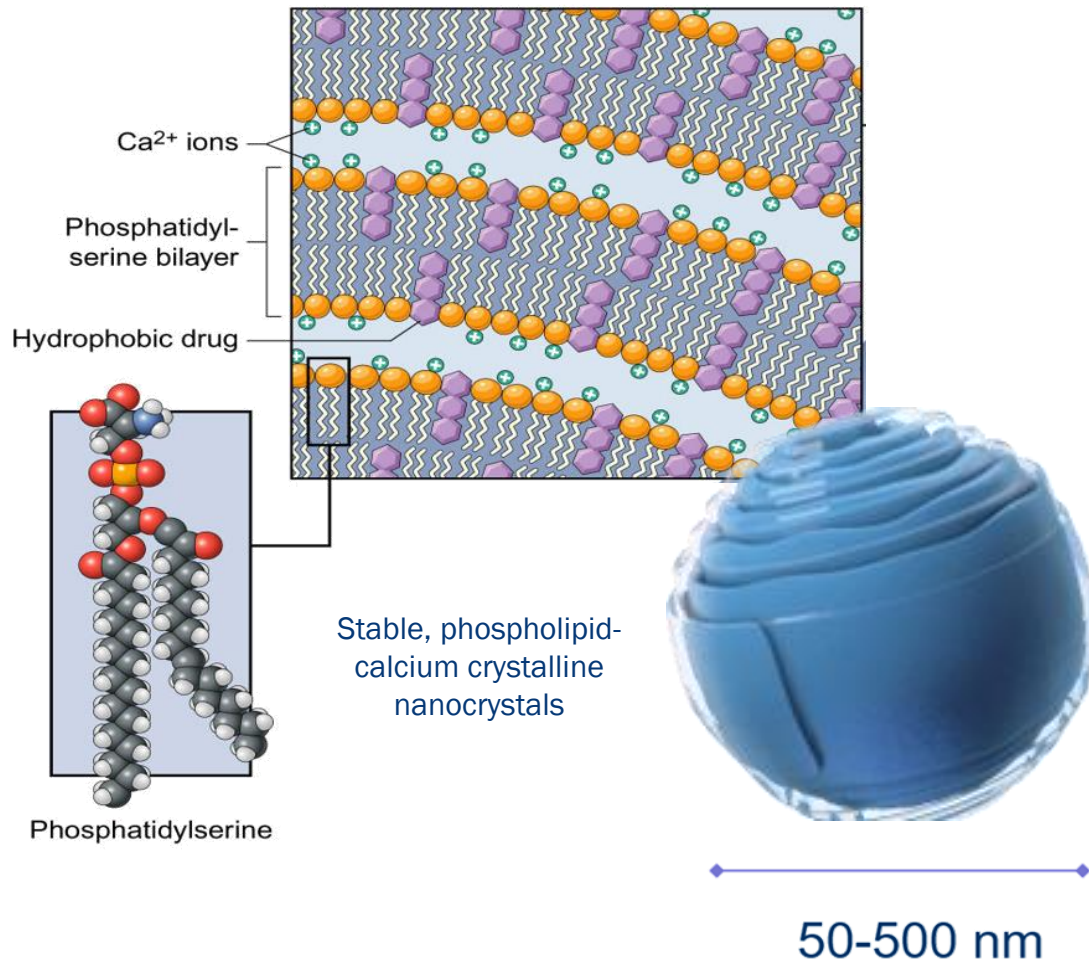
Clinical Validation  
of LNC Capabilities

### Pipeline Products and Opportunities

- Oral Amphotericin B - without nephrotoxicity
- Phase 3-ready (Invasive Aspergillosis in ~216 patients with limited or no treatment options)
- Provides effective longer-term fungicidal stepdown therapy for Invasive Fungal Infections
- **12** years of exclusivity\* \*QIDP and Orphan designations

- Current efforts expanding LNC application beyond small molecules to include delivery of smaller oligonucleotides (ASOs, siRNA, RNAi)
- Expanding primary therapeutic applications from infectious disease (antifungal, antibiotic, antiviral) to inflammation and cancer

# Lipid Nanocrystals (LNCs): A Clinically Validated Intracellular Delivery Platform



## Delivery of small molecules and small oligonucleotides

- Successful delivery of small molecules, proteins, small oligonucleotides (siRNA, ASOs), and vaccines

## Extra-hepatic targeting

- Selective uptake driven by phosphatidylserine enables delivery in infection, inflammation, oncology
- Validated Blood-Brain-Barrier penetration (MAT2203)

## Oral delivery

- Unique structure protects cargo in GI tract, avoids first-pass hepatic metabolism

## Safe & stable

- Deliver high-target tissue concentrations of drug with low plasma levels and no absorption by non-target tissues
- No evidence of immunogenicity or cytotoxicity

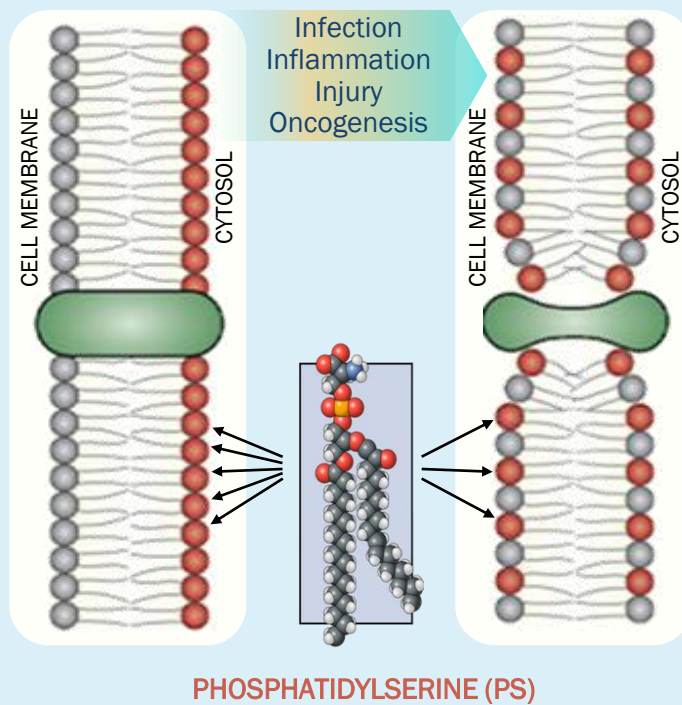
# Phosphatidylserine Enables Cellular Targeting and Intracellular Delivery

## Targeting

### Stressed Cells Externalize PS

Normally, **PS** is confined to the inner layer (facing cytosol)

With injury, **PS** moves from the inner layer to the outer layer of the cell membrane



With a wide variety of potential target cells

### PROFESSIONAL PHAGOCYTES

- Macrophages/monocytes
- Neutrophils
- Dendritic cells

### NON-PROFESSIONAL PHAGOCYTES

- Fibroblasts, epithelial cells, endothelial cells

### INJURED/STRESSED CELLS

- Infection
- Inflammation
- Other physiologic stressors

### TUMOR CELLS

### IMMUNE CELLS

- T-cells

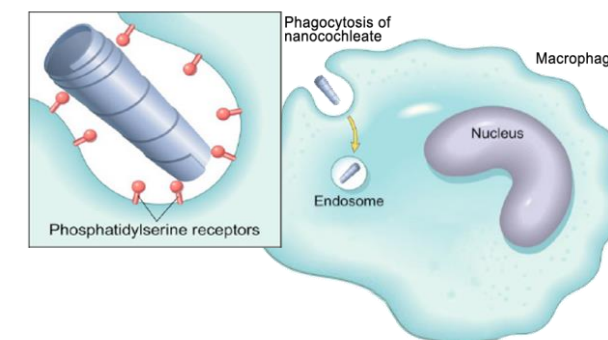
### OTHER ACTIVELY DIVIDING CELLS (including extracellular pathogens)

## Delivery

PS-containing LNCs deliver their cargo to the interior of cells by both phagocytosis and fusion

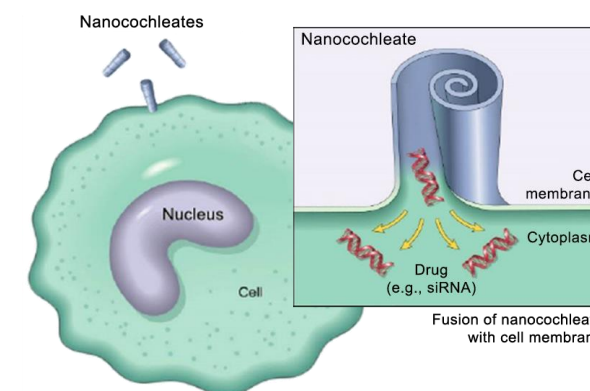
### PHAGOCYTOSIS

**PS** on the outer layer of injured cells is an “eat-me” signal enabling recognition and uptake by professional phagocytes. Cargo-carrying LNCs can be taken up in a similar fashion, with subsequent endosomal escape of cargo.




### FUSION

**PS** on the outer cell membrane is also a precursor for direct membrane-to-membrane fusion and more rapid direct cytosolic delivery by cargo-carrying LNCs to cells expressing PS on their outer membranes.







# **MAT2203**

# **Oral Amphotericin B**

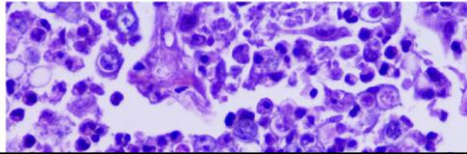
## **Clinical Validation of LNC Delivery**

# The Growing Threat of Invasive Fungal Infections (IFIs)

## WIRED

### The Battle Against the Fungal Apocalypse Is Just Beginning

Fungal infections are rising worldwide and climate change may be to blame. Medicine isn't ready.



**WHO fungal priority pathogens list to guide research, development and public health action**



## Wall Street Journal

### Deadly Fungal Infections Confound Doctors—'It's Going to Get Worse'

Once a freak occurrence, fungi resistant to standard drugs now threaten millions of vulnerable Americans

By Dominique Mosbergen [Follow](#)  
June 22, 2023 9:59 am ET

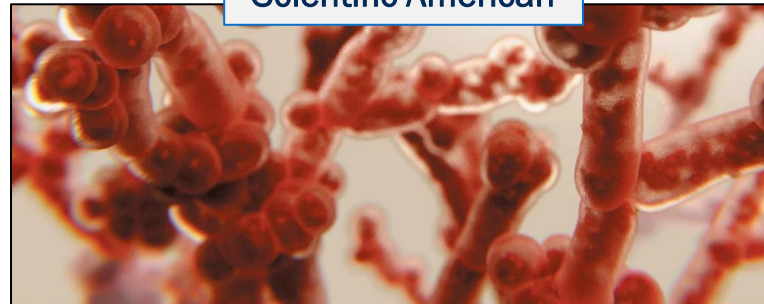
## Nature Reviews Microbiology 2022

### Tackling the emerging threat of antifungal resistance to human health

Matthew C. Fisher<sup>1,2,3</sup>, Ana Alastruay-Izquierdo<sup>2</sup>, Judith Berman<sup>2</sup>, Tihana Bicanic<sup>4</sup>, Elaine M. Bignell<sup>5</sup>, Paul Bowyer<sup>6</sup>, Michael Bromley<sup>6</sup>, Roger Brüggemann<sup>7</sup>, Gary Garber<sup>8</sup>, Oliver A. Cornely<sup>9</sup>, Sarah J. Gurr<sup>10</sup>, Thomas S. Harrison<sup>4,5</sup>, Ed Kuijper<sup>11</sup>, Johanna Rhodes<sup>1</sup>, Donald C. Sheppard<sup>12</sup>, Adilia Warris<sup>13</sup>, P. Lewis White<sup>14</sup>, Jianping Xu<sup>14</sup>, Bas Zwaan<sup>15</sup> and Paul E. Verweij<sup>11,16,17</sup>

**Abstract** | Invasive fungal infections pose an important threat to public health and are an under-recognized component of antimicrobial resistance, an emerging crisis worldwide. Across a period of profound global environmental change and expanding at-risk populations, human-infecting pathogenic fungi are evolving resistance to all licensed systemic antifungal drugs. In this Review, we highlight the main mechanisms of antifungal resistance and explore the similarities and differences between bacterial and fungal resistance to antimicrobial control. We discuss the research and innovation topics that are needed for risk reduction strategies aimed at minimizing the emergence of resistance in pathogenic fungi. These topics include links between the environment and One Health, surveillance, diagnostics, routes of transmission, novel therapeutics and methods to mitigate hotspots for fungal adaptation. We emphasize the global efforts required to steward our existing antifungal armamentarium, and to direct the research and development of future therapies and interventions.

## Scientific American



Fungus *Candida auris* lurks in hospitals, where it infects patients with impaired immune systems. Credit: Juan Gaertner Science Source

### Deadly Fungi Are the Newest Emerging Microbe Threat All Over the World

These pathogens already kill 1.6 million people every year, and we have few defenses against them

By Maryn McKenna

## New York Times



## Wall Street Journal

SCIENCE

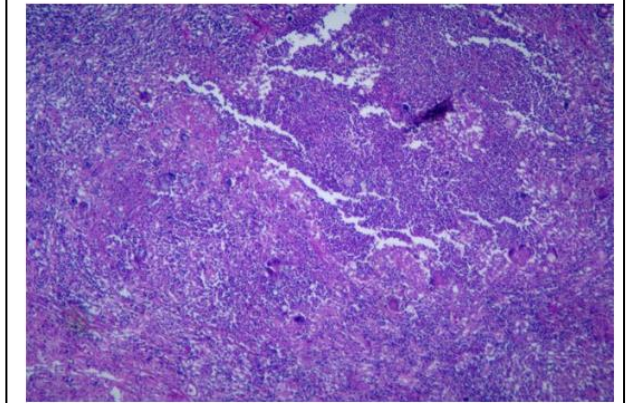
### Dangerous Fungi Are Spreading Across U.S. as Temperatures Rise

Some fungi such as the type that causes Valley Fever might be adapting to endure more heat stress

By Dominique Mosbergen [Follow](#)  
Feb. 1, 2023 10:08 am ET

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[Listen](#) (5 min)



## NBC Nightly News



# MAT2203: Unlocking the Full Potential of Amphotericin B

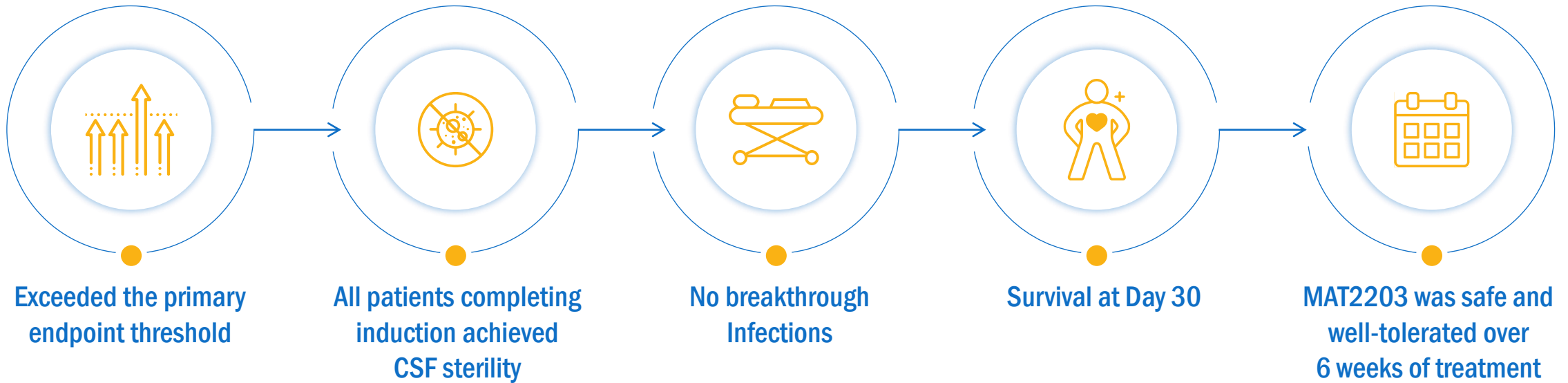
Innate Amphotericin B Characteristics	—	IV Amphotericin B Limitations	=	Unlocked Potential
✓ POTENT – broad-spectrum fungicidal		✗ Only available through IV administration		✓ Available systemically and orally (crosses BBB following oral administration)
✓ Minimal drug-drug interactions		✗ Significant toxicity and side effects		✓ Well-tolerated and safe
✓ Low propensity for resistance		✗ High systemic exposure distributed throughout the body		✓ Delivered directly into infected tissues
✓ Active against susceptible and emerging drug-resistant fungal infections		✗ Must be administered in hospital, increasing costs		✓ Cost-effective with potential for significant health economic benefits

**MAT2203 is a promising potential therapeutic option**  
for the treatment of MULTIPLE serious and life-threatening fungal infections



# EnACT: Phase 2 Clinical Validation of Safety and Efficacy

## EnACT Clinical Data in Cryptococcal Meningitis Validates the Use of the LNC Platform to Enable Oral Administration and Overcome Toxicity



- EFA for MAT2203 was 0.42  
(95% CI 0.29 to 0.55)  
(primary endpoint threshold was 0.20)

- 97% for patients receiving MAT2203
- 76% for patients receiving SOC

- Over 10 weeks, patients showed no breakthrough infections post-MAT2203 treatment

- 98% for patients receiving MAT2203
- 88% for patients receiving IV AMPB (SOC)

- Repeat dosing showed no renal toxicity or electrolyte abnormalities
- No discontinuations due to AEs nor MAT2203-related SAEs

# Results of EnACT Published in Highly-Regarded Peer-Reviewed Journal

*Clinical Infectious Diseases*

MAJOR ARTICLE

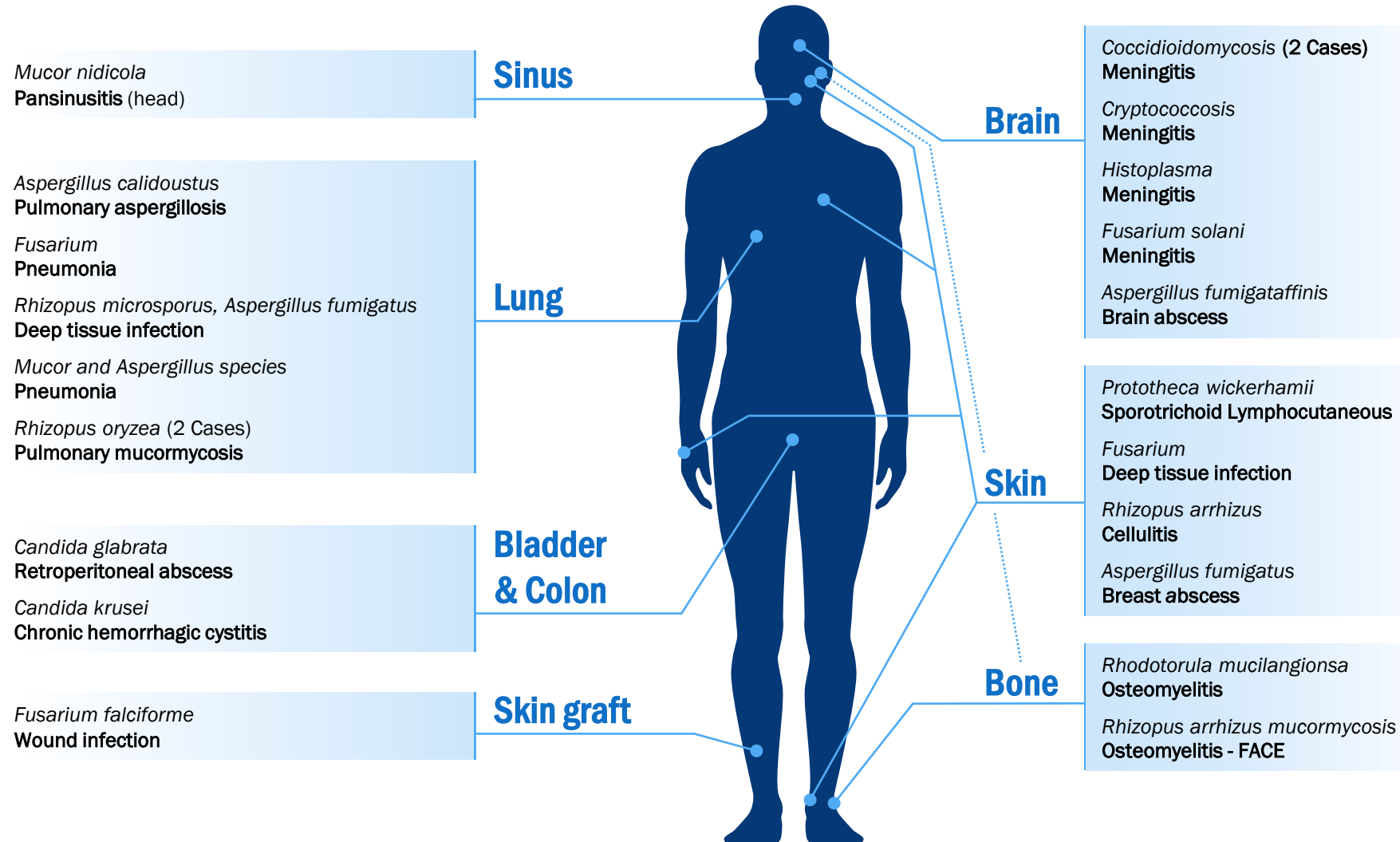


## Oral Lipid Nanocrystal Amphotericin B for Cryptococcal Meningitis: A Randomized Clinical Trial

David R. Boulware,<sup>1,a</sup> Mucunguzi Atukunda,<sup>2,a</sup> Enoch Kagimu,<sup>2</sup> Abdu K. Musubire,<sup>2</sup> Andrew Akampurira,<sup>2</sup> Lillian Tugume,<sup>2</sup> Kenneth Ssebambulidde,<sup>2,3</sup> John Kasibante,<sup>2</sup> Laura Nsangi,<sup>2</sup> Timothy Mugabi,<sup>2</sup> Jane Gakuru,<sup>2</sup> Sarah Kimuda,<sup>2</sup> Derrick Kasozi,<sup>2</sup> Suzan Namombwe,<sup>2</sup> Isaac Turyasingura,<sup>2</sup> Morris K. Rutakingirwa,<sup>2</sup> Edward Mpoza,<sup>2</sup> Enos Kigozi,<sup>4</sup> Conrad Muzoora,<sup>4</sup> Jayne Ellis,<sup>2</sup> Caleb P. Skipper,<sup>1</sup> Theresa Matkovits,<sup>5</sup> Peter R. Williamson,<sup>3</sup> Darlisha A. Williams,<sup>1</sup> Ann Fieberg,<sup>6</sup> Kathy H. Hullsiek,<sup>6</sup> Mahsa Abassi,<sup>1</sup> Biyue Dai,<sup>6</sup> and David B. Meya<sup>1,2</sup>

<sup>1</sup>Department of Medicine, University of Minnesota, Minneapolis, Minnesota, USA; <sup>2</sup>Infectious Diseases Institute, Makerere University, Kampala, Uganda; <sup>3</sup>Laboratory of Clinical Immunology and Microbiology, National Institute of Allergy and Infectious Diseases, National Institutes of Health, Bethesda, Maryland, USA; <sup>4</sup>Department of Medicine, Mbarara University of Science and Technology, Mbarara, Uganda; <sup>5</sup>Matinas Biopharma Nanotechnologies, Bedminster, New Jersey, USA; and <sup>6</sup>Division of Biostatistics, School of Public Health, University of Minnesota, Minneapolis, Minnesota, USA

# MAT2203 Expanded Access Program – Targeted Treatment of IFIs Throughout the Body



# MAT2203 Expanded Access/Compassionate Use Program

## Demonstrated Efficacy in Treatment of Patients with Limited Treatment Options

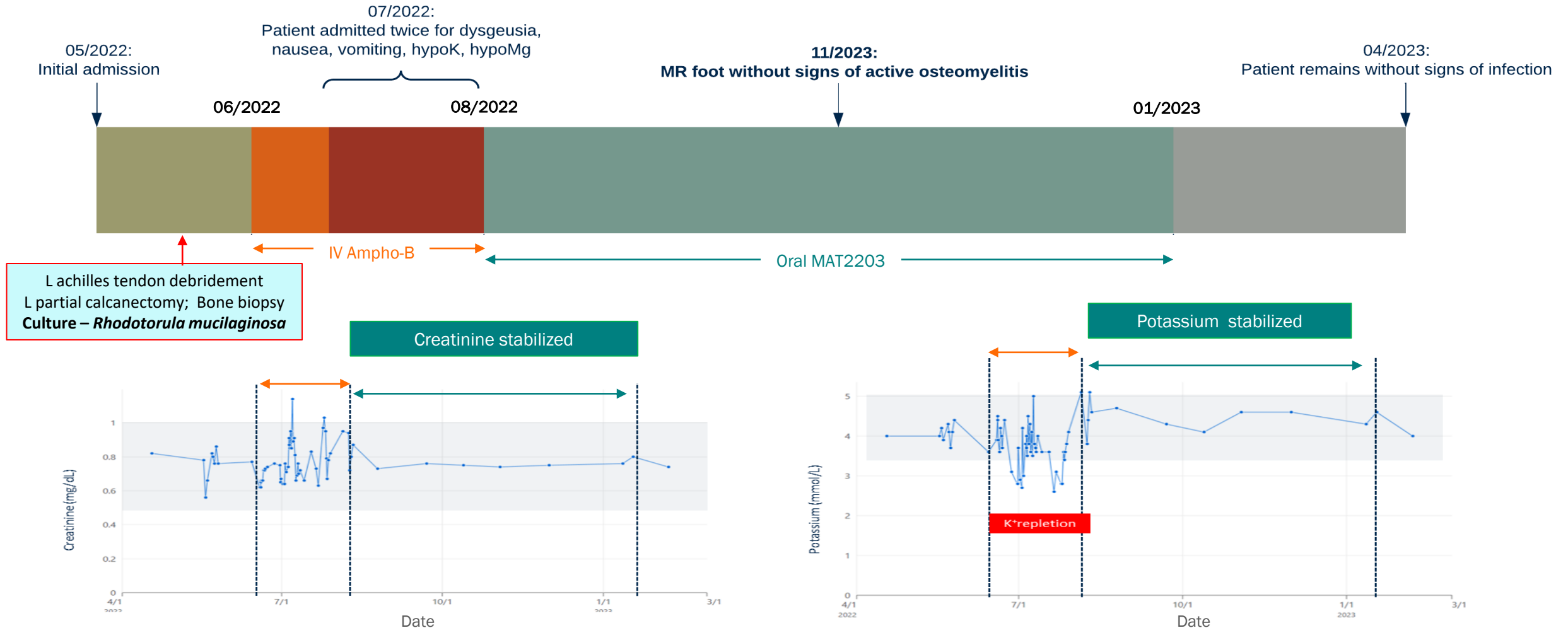
- 19 patients with no other treatment options have enrolled to receive or have completed treatment with MAT2203
  - **Notable Healthcare Institutions:** NIH, University of Michigan, Johns Hopkins, City of Hope, Nationwide Children's Hospital, Vanderbilt University Medical Center, Memorial Sloan Kettering, University of California at San Diego School of Medicine, Children's Hospital of Philadelphia
- Patients were not responding/resistant to, or unable to receive, azole therapy
- Patients were switched to treatment with IV Amphotericin B with clinical response but unable to tolerate treatment due renal toxicity
  - All patients hospitalized to monitor/manage renal safety and most received IV electrolyte supplementation
- Following oral MAT2203 initiation, patients were discharged to continue treatment at home
- Renal toxicity reversed and renal function returned to baseline after switching to MAT2203
- All patients who received at least two weeks or more of treatment had positive clinical outcomes with significant success stories of full recovery in majority of patients



# Compassionate Use - *Recovery from IV Amphotericin B Kidney Toxicity with MAT2203*



A 38 y/o female with systemic lupus erythematosus on chronic hydroxychloroquine and prednisone presented with a progressively enlarging wound on her left foot.



# High Unmet Medical Need in Treatment of Invasive Aspergillosis (IA)

- Invasive aspergillosis (IA) is a serious and life-threatening invasive fungal infection that occurs primarily in severely immunocompromised patients with hematological malignancies and transplant recipients
  - ~15,000 new cases per year in the U.S. alone
  - WHO, CDC, and FDA consider IA a critical priority and a global public health concern
- IDSA Guidelines recommend treatment with mold-active azoles as first-line treatment for 6-12 weeks
  - Azole use requires significant expertise to manage toxicities and significant drug-drug interactions that often limits duration of use
  - Resistance to azoles has been increasing globally
  - Recently, cases of breakthrough IA have been reported in patients receiving antifungal prophylaxis
    - Failures attributed to non-compliance, poor absorption, DDIs, or infection with a drug-resistant Aspergillus species
- Patients suffering from IA with little or no treatment options among the highest unmet medical need with approximately 3,000-5,000 cases per year (U.S. only)
  - Rare disease/orphan commercial opportunity

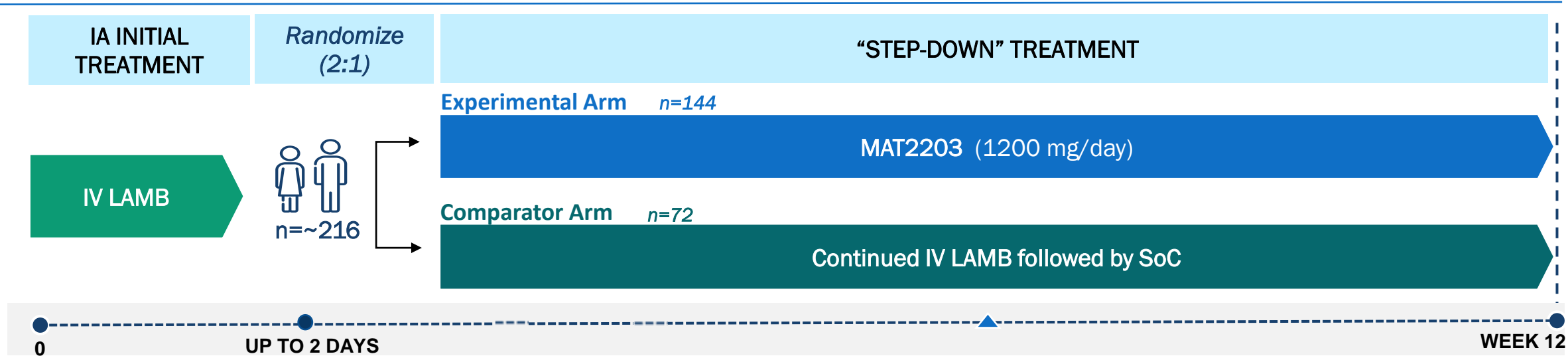
*\* Phase 3 trial dependent on securing partnership(s) or non-dilutive government funds*

# MAT2203 Regulatory and Development Strategy

- Near-term development strategy refined to narrow initial target indication: treatment of invasive aspergillosis in patients with limited treatment options (azole-intolerant, azole-resistant, or not effectively managed with an azole)
  - Potential registration through leveraging LPAD pathway
  - Other regulatory designations protected and maintained (QIDP, ODD, Fast-Track, potential for Breakthrough Therapy)
- In February 2024, reached agreement with FDA on a single Phase 3 Registration Trial in support of an NDA for the treatment of invasive aspergillosis in patients with limited treatment options (the “ORALTO” Trial)
- Preparations underway with global CRO in preparation for Phase 3 study implementation
- Development and commercial partnership discussions remain ongoing

# FDA-Agreed Phase 3 Study Design in Invasive Aspergillosis (IA) (the “ORALTO” Trial)

- To demonstrate that initial treatment with IV LAMB followed by step-down to oral MAT2203 is comparable to (noninferior) SoC treatment in adult patients with Invasive Aspergillosis (IA) who are unable to receive treatment with a mold-active azole and have limited alternative treatment options
- Patients will be randomized 2:1 to receive either oral MAT2203 (Experimental Arm) or continued IV LAMB followed by Standard of Care (Comparator Arm)



**Treatment of IA in patients with LIMITED TREATMENT OPTIONS**

- Fungal pathogen not susceptible to azoles
- Risk for toxicity or drug-drug interactions with azoles
- Other clinical contraindications for azole use

**PRIMARY ENDPOINT**

- All-Cause Mortality at Study Day 42

**SECONDARY ENDPOINTS**

- Global response to treatment
- All-cause mortality at 84 days
- Safety and tolerability
- Pharmacoeconomic impact

**KEY SECONDARY SUPERIORITY SAFETY ENDPOINT**

- Treatment discontinuation or dose adjustment due to treatment-related toxicity
  - Renal toxicity: serum creatinine >1.5X baseline
  - Hypokalemia: serum K<sup>+</sup> < 3.0 mmol/L or requiring K<sup>+</sup> supplementation
  - Infusion-related reaction
  - Hepatic toxicity
  - Clinically significant azole DDI requiring medical intervention
  - Severe GI intolerance requiring medical intervention
  - Other toxicity that resulted in discontinuation and/or change in dose



# MAT2203 Value Proposition

AMPHOTERICIN B + ORAL ADMINISTRATION + LACK OF NEPHROTOXICITY = Safe, long-term oral step-down Tx with a potent fungicidal agent

## LIFE CYCLE MANAGEMENT



VALUE

## Near-Term Development Strategy:

Focused strategy on Phase 3 registration trial in Invasive Aspergillosis in patients with limited or no treatment options under the Limited Population Pathway for Antifungal Drugs (LPAD)\*

\* NDA review issue



# LNCs Beyond MAT2203

## Efficient and Safe Delivery of Small Oligos

# LNC Therapeutic Cargo Experience to Date

Infection

Inflammation

Oncology

In Vivo Animal Studies		Human Clinical Trials	
Animal model	Cargo	Indication	Cargo
Cystic fibrosis mouse model	Amikacin	Mucocutaneous candidiasis Job's syndrome, NIH	Amphotericin - B
	ASO		
	Flu protein		
	siRNA		
BALB/c mouse flu model		Vulvovaginal Candidiasis (VVC) Phase 2	Amphotericin - B
[Multiple mouse fungal models]	Amphotericin- B		
Pneumocystis mouse model	Atovaquone	HIV / cryptococcal meningitis, Phase 2	Amphotericin - B
Mouse SARS-CoV-2 model	Remdesivir		
Rat footpad inflammation model	NSAID	* new	* new
GvHD mouse model	ST1959		
LPS mouse model	RNAi		
Psoriasis (IMQ) mouse model*	RNAi		
DSS colitis mouse model*	RNAi	* new	* new
Lymphocytic leukemia mouse model	ASO		
Syngeneic mouse melanoma model*	Docetaxel		

# Unlocking the Full Potential of the LNC Platform

Matinas is working internally and with third parties to broaden its pipeline of LNC-based therapeutics

## PROVEN

### Oral Formulations of Anti-Infectives

- MAT2203
- Remdesivir
- MAT2501

### Oral Formulations of Chemotherapeutics

- Docetaxel

## UNDER EVALUATION

### Intracellular Delivery of Nucleic Acids

- siRNA
- ASO

## FUTURE

### Potential Therapeutic Applications

#### INFECTION

- Anti-infectives
- Antivirals

#### \*INFLAMMATION

- Autoimmune diseases
- Neuro-inflammatory diseases
- Acute/chronic Inflammatory diseases

#### \*ONCOLOGY

- Hematologic & solid tumor malignancies

\* Key Areas of Focus for Internal LNC Oral Small Oligonucleotides

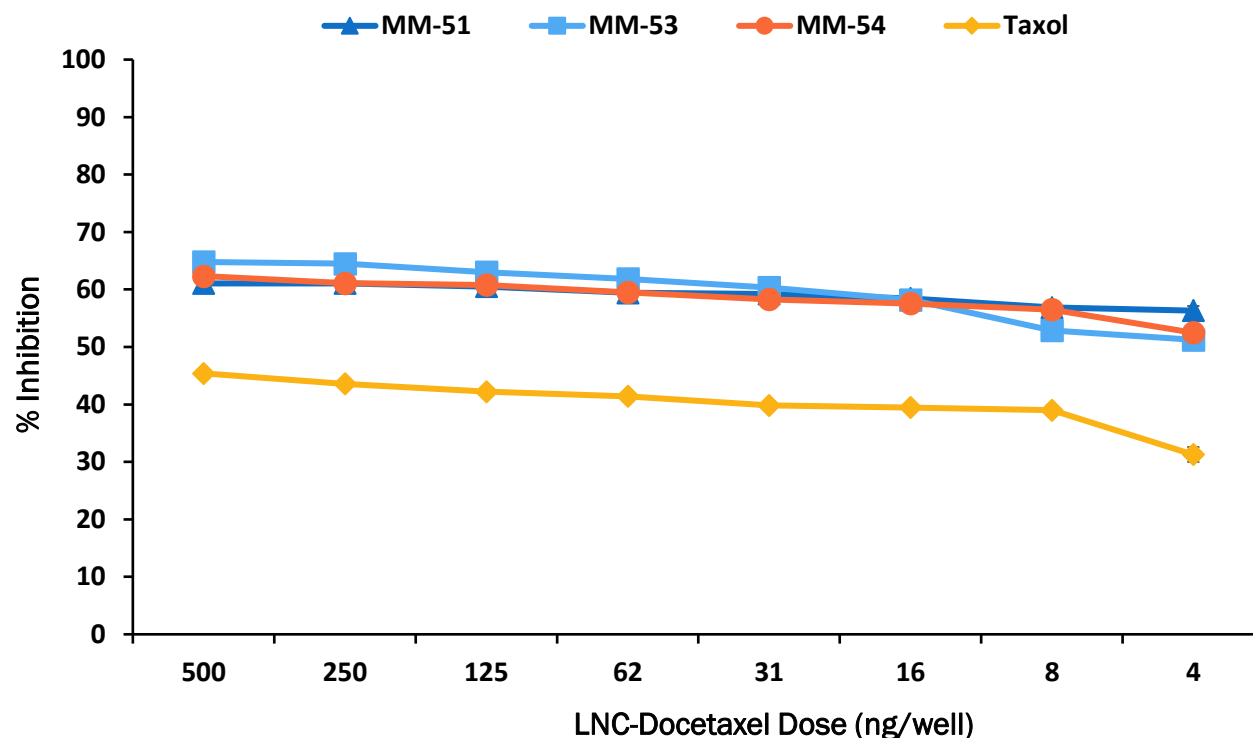


# *In vitro* Tumor Cell Inhibition with LNC Docetaxel Formulations

## RATIONALE AND NEXT STEPS

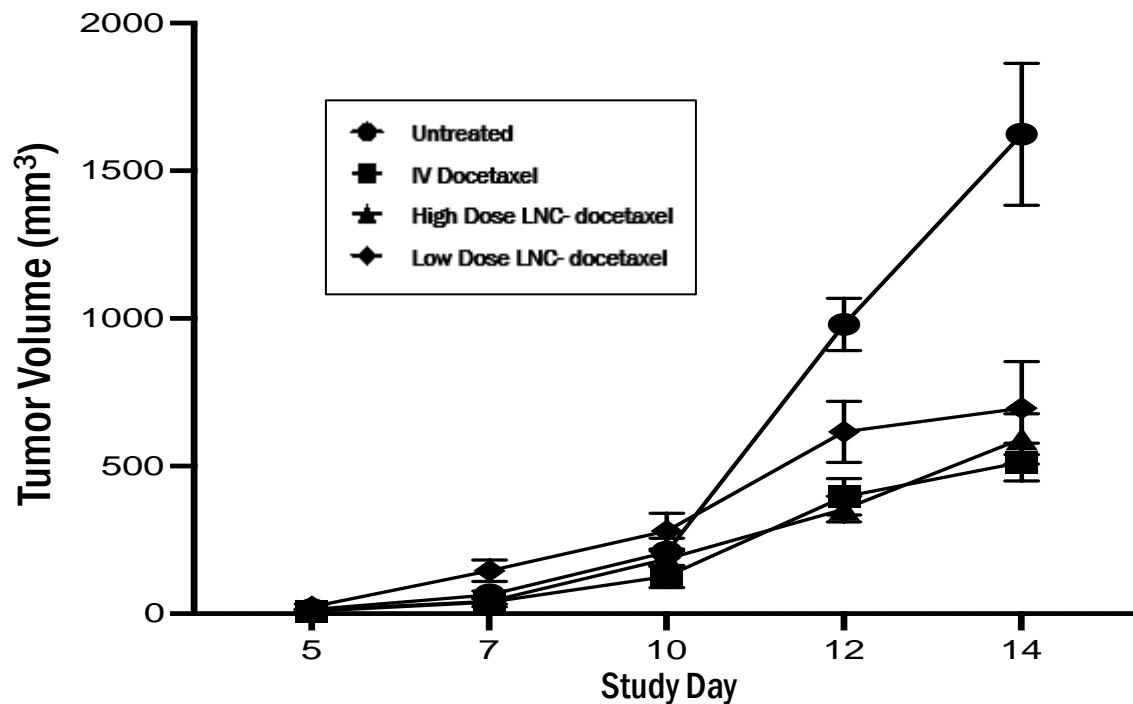
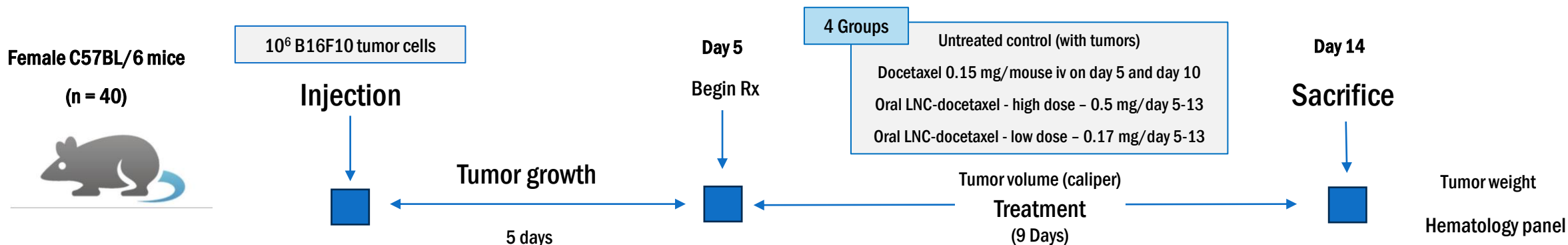
- LNC Docetaxel program also provides insight into tumor uptake of LNCs
  - Foundation for subsequent small oligo work
- *In vitro* results demonstrate strong LNC efficacy in inhibiting tumor cell growth
- *In vivo* studies planned in multiple tumor models

## 3 LNC Formulations Demonstrated Significantly Better Tumor Cell Inhibition Than Unformulated Drug



MCF7 breast cancer cells

# *In vivo* Therapeutic Efficacy of Oral Docetaxel LNCs in a Murine Syngeneic Melanoma Model



Reductions in tumor volume and tumor weight comparable to that seen with IV docetaxel

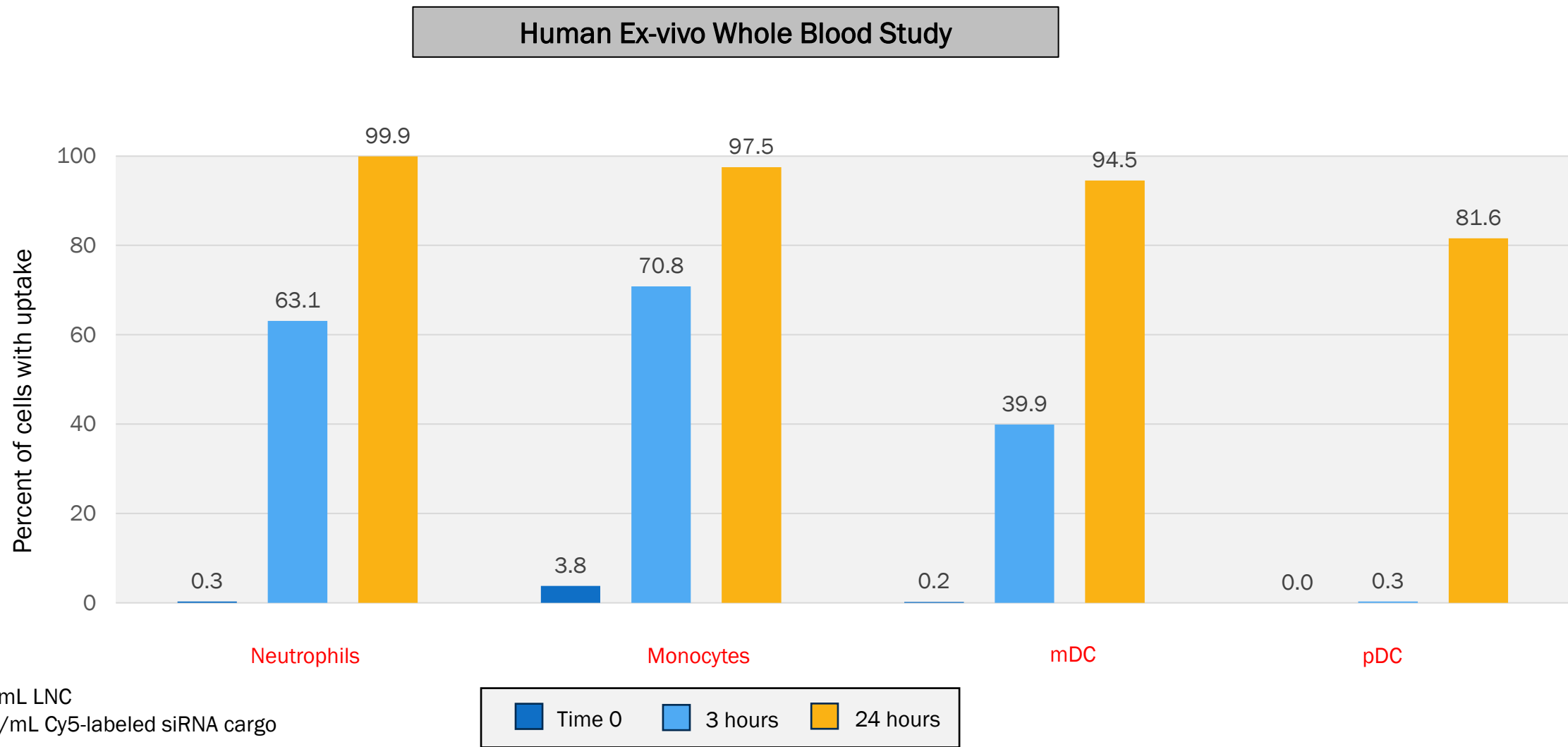
No significant effect of daily oral docetaxel treatment on body weight or hematologic parameters

## Next Steps

Further evaluation of safety vs IV Docetaxel  
Screening of multiple tumor cells lines for LNC uptake  
Evaluation in other tumor models  
Other potential chemotherapies

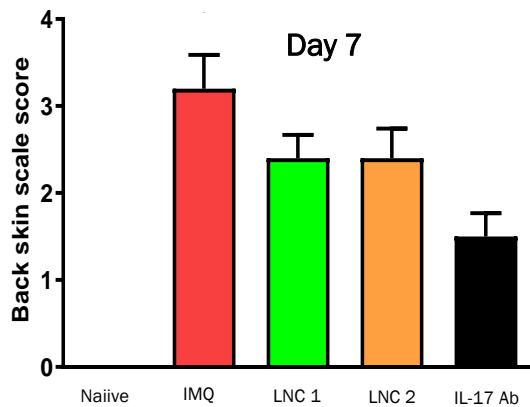
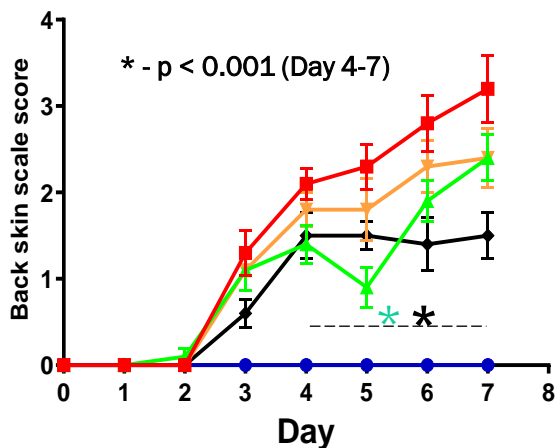
# LNC-formulated Small Oligos Show Strong Uptake in Innate Immune Cells

## Support Role in Treating Inflammation

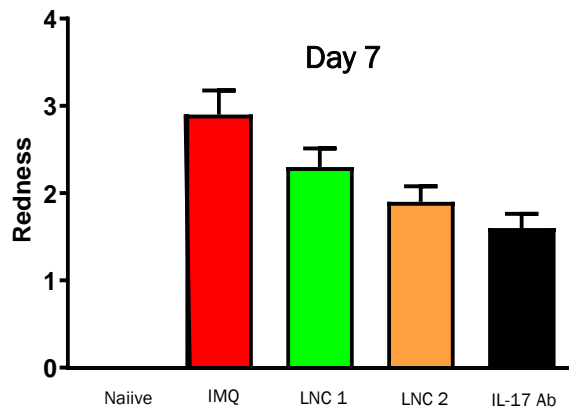
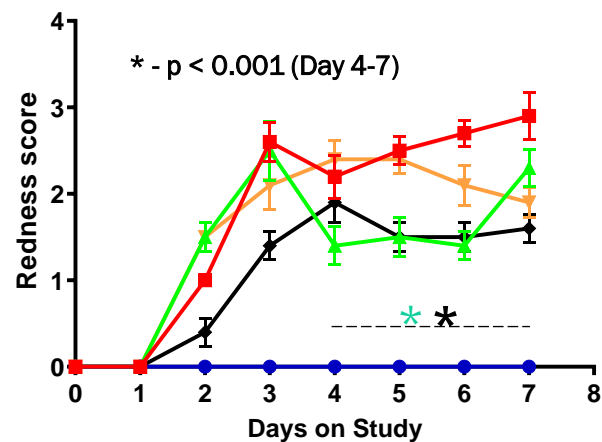


# Effect of Oral LNC IL-17A RNAi in a Murine Imiquimod (IMQ) Psoriasis Model

## Scaling



## Redness

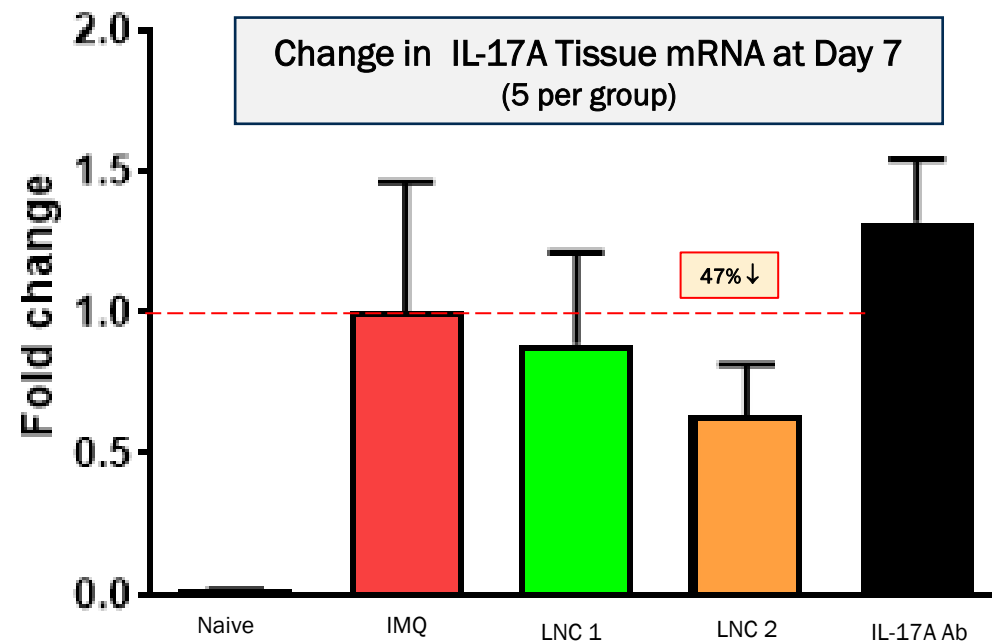


BalbC mice  
(n = 50)

5 Groups

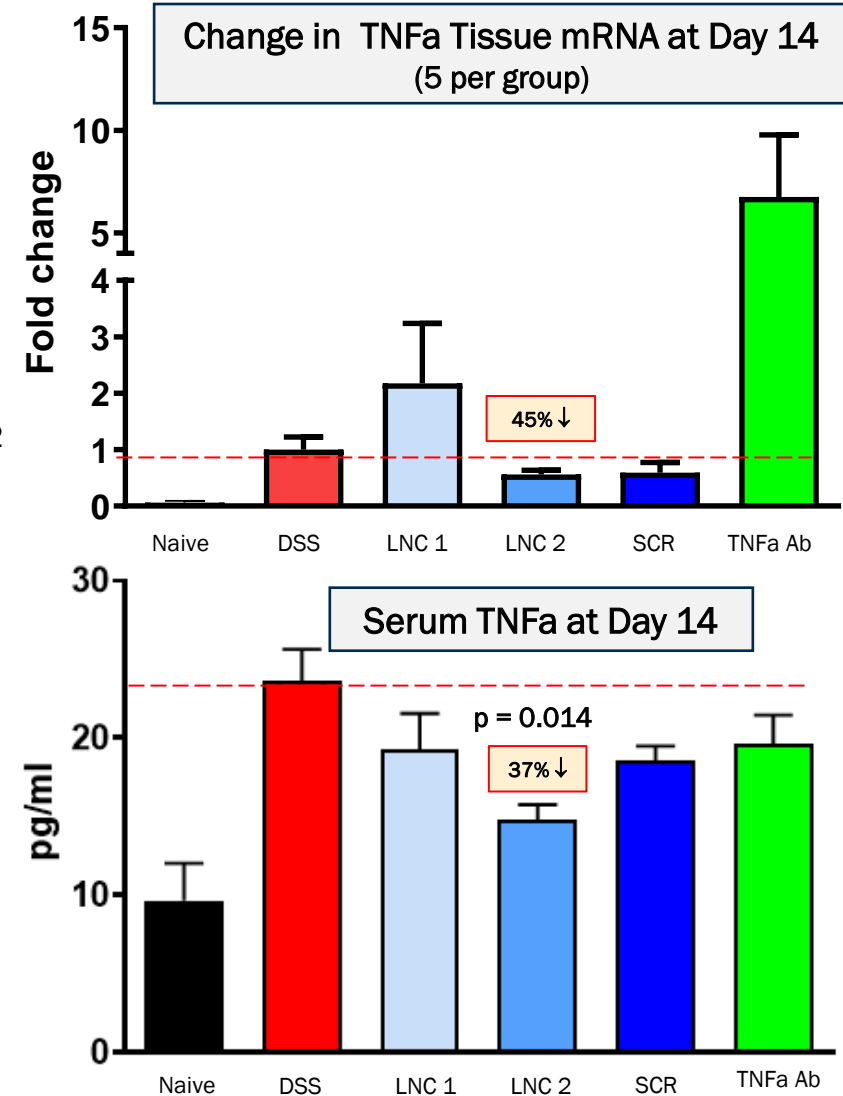
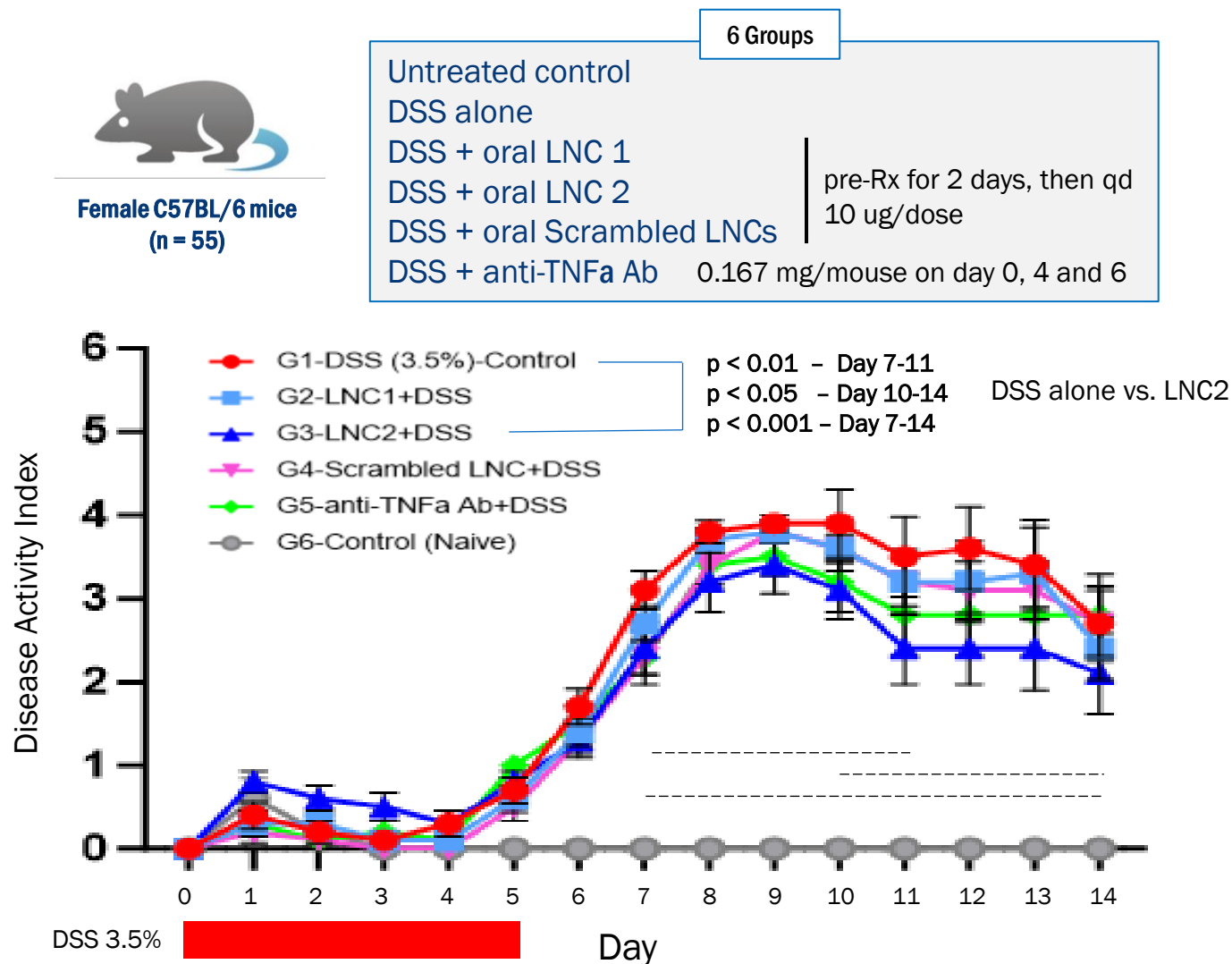
Naïve  
IMQ only  
LNC1  
LNC2  
IL-17A murine Ab

10 µg/mouse daily (oral gavage)  
1 mg/kg i.p. qod



[ Changes in serum cytokine levels not expected in this model ]

# Effect of Oral LNC-TNF $\alpha$ RNAi in a Murine DSS Acute Colitis Model



# Summary of Recent Key Advances

- Successful *in vivo* oral delivery of LNC-Docetaxel in a melanoma tumor model
  - Reductions in tumor weight and volume comparable to those seen with IV docetaxel
  - No adverse effects on body weight or hematologic parameters
- Successful *in vivo* oral delivery of LNC formulations of 2 different RNAi oligonucleotides targeting inflammatory cytokines
  - Documented biological activity and therapeutic impact in two different disease models.
    - Psoriasis (IL-17A)
      - Reduction in tissue IL-17A mRNA levels (skin)
      - Statistically significant improvement in clinical scoring of skin lesions
    - Colitis (TNFa)
      - Reductions in tissue TNFa mRNA levels (colon)
      - Statistically significant reductions in serum TNFα levels
      - Statistically significant improvement in disease activity scores

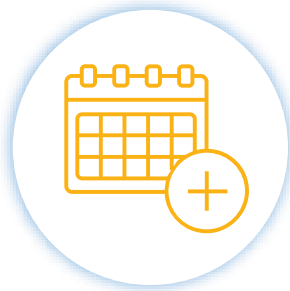
These results highlight and validate the unique capabilities of the LNC platform beyond MAT2203

- 1) Oral delivery
- 2) Delivery of active therapeutics (small molecule and small oligo) to diseased tissues outside the liver
- 3) Low blood levels of active drug, with potential for improved safety



# Expanding LNC Intellectual Property Portfolio

Continuingly increasing our patent suite to increase protection and exclusivity



MAT2203 potentially entitled to **12+ years of exclusivity** (QIDP & Orphan status)



Global Platform IP base **protection out to 2037** with 20 patents issued in last 5 years



Recent patent applications **based on formulation work with small oligonucleotides**

## Strong IP & Regulatory Designations

# Experienced Leadership Team

## EXECUTIVE TEAM



**Jerome D. Jabbour, J.D.**  
Chief Executive Officer



**Thomas Hoover, MBA**  
Chief Business Officer



**Theresa Matkovits, Ph.D.**  
Chief Development Officer



**James Ferguson, M.D.**  
Chief Medical Officer



**Keith Kucinski, CPA, MBA**  
Chief Financial Officer



**Hui Liu, Ph.D., MBA**  
Chief Technology Officer



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Director



**James Scibetta**  
Director



**Kathryn Corzo**  
Director



**Natasha Giordano**  
Director



**Matthew Wikler, M.D., MBA**  
Director



**Jerome D. Jabbour, J.D.**  
CEO



# Matinas Investment Thesis

## Financial Summary



Runway into Q3 2024



\$18.2M<sup>1</sup> in Cash,  
Cash Equivalents and  
Marketable Securities

<sup>1</sup> as of 09/30/23



Non-Dilutive  
Financing Options

## Near-Term Milestones – Setup for Strong Start to 2024



**Q1** – Successful Agreement with FDA for a Single Phase 3 Registration Trial to Support an NDA for MAT2203 for the Treatment of Invasive Aspergillosis (the “ORALTO” Trial)

**Q1** – LNC-docetaxel Maximum Tolerated Dose Longer Term Safety Study vs. IV-docetaxel

**Q1** – Additional *in vivo* Oral Delivery of Small Oligonucleotide Targeting Inflammation

**Q1** – Potential MAT2203 Domestic/Regional/Global Partnership

**Q1** – Screening of Tumor Cell Lines for LNC Uptake Including Breast, Brain, Liver & Lung

**Q1** – Evaluation of LNC-docetaxel in Additional Tumor Models

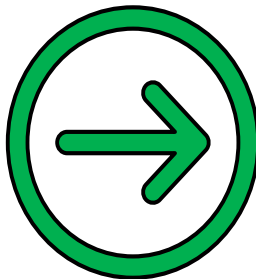
**Q2** – Potential New Platform Collaboration

**2024** – Potential BARDA Funding for MAT2203

## Solid Value “Foundation”

**MAT2203**

- Clinically Validated
- Phase 3-ready asset
- Highest unmet need



## Substantial UPSIDE

LNCs Facilitating ORAL Delivery  
of Small Oligonucleotides

And

Establishing Internal and  
External Pipelines