

Clinical stage biopharmaceutical company focused on developing first-in-class, oral therapeutics for autoimmune disease May 2023



## **Forward Looking Statements**

This presentation contains forward-looking statements that involve substantial risks and uncertainties. All statements, other than statements of historical facts, contained in this presentation, including statements regarding our strategy, future operations, prospects, plans and objectives of management, are forward-looking statements. The words "anticipate," "believe," "estimate," "expect," "intend," "may," "might," "plan," "predict," "project," "target," "potential," "will," "would," "could," "could," "continue," and similar expressions are intended to identify forward looking statements, although not all forward-looking statements contain these identifying words. These forward-looking statements are subject to a number of risks, uncertainties and assumptions. Risks regarding our business are described in detail in our Securities and Exchange Commission filings, including in our Annual Report on Form 10-K for the year ended December 31, 2022. We may not actually achieve the plans, intentions or expectations disclosed in our forward-looking statements, and you should not place undue reliance on our forward-looking statements. Actual results or events could differ materially from the plans, intentions and expectations disclosed in the forward-looking statements we make. The forward-looking statements contained in this presentation reflect our current views with respect to future events, and we assume no obligation to update any forward-looking statements except as required by applicable law.

This presentation includes statistical and other industry and market data that we obtained from industry publications and research, surveys and studies conducted by third parties as well as our own estimates of potential market opportunities. All of the market data used in this prospectus involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such data. Industry publications and third party research, surveys and studies generally indicate that their information has been obtained from sources believed to be reliable, although they do not guarantee the accuracy or completeness of such information. Our estimates of the potential market opportunities for our product candidates include several key assumptions based on our industry knowledge, industry publications, third-party research and other surveys, which may be based on a small sample size and may fail to accurately reflect market opportunities. While we believe that our internal assumptions are reasonable, no independent source has verified such assumptions.



## Landos Biopharma is Singularly Focused on Advancing NX-13 Clinical Development in UC

#### NX-13

#### Potentially transformative oral, once-daily therapy for moderate to severe ulcerative colitis (UC)

- Addresses multiple causes of UC through novel, bimodal MOA targeting NLRX1
- Promising safety profile and early signals of clinical improvement in Phase 1b study
- NEXUS Phase 2 proof of concept trial initiated Q2 2023; Top-line results expected Q4 2024



Experienced management team with significant gastroenterology, immunology and drug development expertise



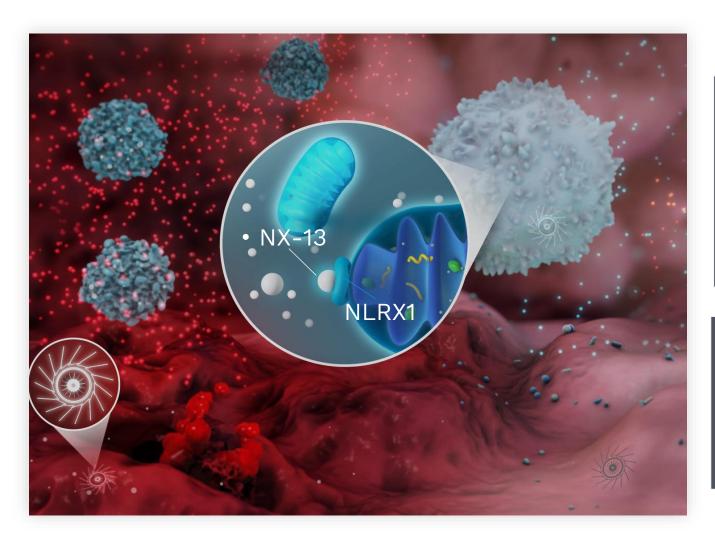
Strong IP position Significant optionality portfolio-wide for partnerships, development & investment



Capital efficient with sufficient cash to fund planned operations into first half of 2025



# NX-13 Unique Bimodal MOA Targets NLRX1 Pathways for Treatment of Ulcerative Colitis (UC)



NX-13 is an oral, once-daily therapy being developed for moderate to severe UC

#### Novel NLRX1 agonist

Bimodal MOA aims to reduce reactive oxygen species **intracellularly** and inflammatory pathways **extracellularly** to reduce UC symptoms and flares

#### NLRX1: the NEXUS of Immunometabolism

mitochondrial-associated anti-inflammatory NOD-like receptor (NLR)

- Direct metabolic role in mitochondria
- Direct anti-inflammatory role as NLR



## **Therapeutic Challenges Present Large Unmet Need for UC Patients**

#### **Ulcerative Colitis**

Chronic colonic inflammation with rectal bleeding and diarrhea

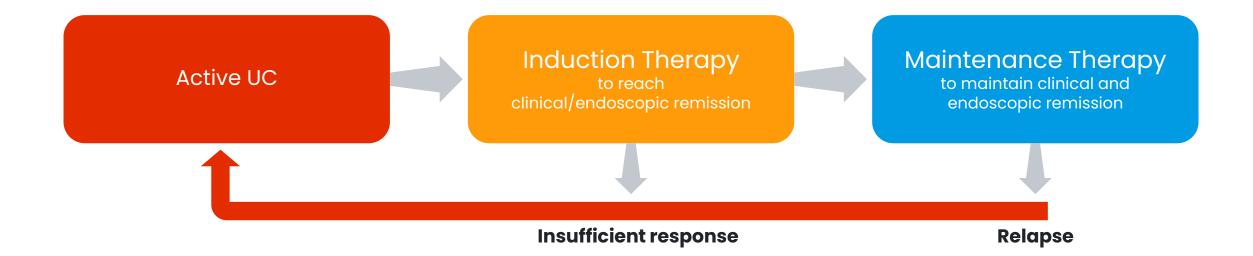
Patients experience relapsing (flares) and remitting episodes of disease severity

#### **Therapeutic Goals**

- induce and maintain steroid-free symptom relief
- healing of colon lining
- improved quality of life

#### **Therapeutic Challenges**

Limited Efficacy: many patients do not respond or lose response to treatment Safety Risks: infections, cancer, blood clots or cardiac events



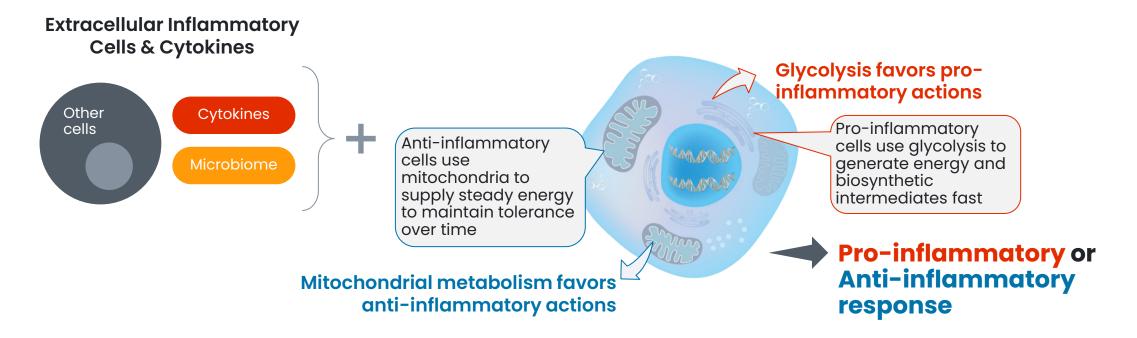


## Current Therapies Focus Exclusively on Extracellular Actions or Signals Falling Short of Effectively Treating a Multifactorial Disease Like UC

Drug Classes	ΜΟΑ	Extracellular (External)		Intracellular (Internal)
		Cytokines	Specific Cells	Environment
NX-13 Bimodal targeting (Immunometabolism)	Reduce intracellular reactive oxygen species (ROS) & extracellular immune response	$\checkmark$	$\checkmark$	$\checkmark$
Anti-Inflammatory / Immunosuppressants	Reduce entire immune response	X	X	
Anti-TNFs, Anti-ILs	Block cytokine binding	X		
Anti-integrins	Inhibit entrance of immune cells to the gut tissue from the circulation		X	
SIP modulators	Inhibit exit of immune cells from immune organs to circulation & gut		X	
JAK Inhibitors	Block cytokine cascades (TNF, IL-17, IFN, etc)	X	X	



## Immune Function is Intimately Tied to the Intracellular Environment of Processing & Using Energy



- The intracellular immunometabolic state (the processing & using of energy through glycolysis or mitochondrial metabolism) provides a baseline, and can affect cellular response as pro- or anti-inflammatory
- Many proteins, molecules & substrates have dual action on cellular metabolism AND immune function
- The underlying intracellular (internal) immunometabolic environment can affect the response of multiple cells involved in UC and gut homeostasis (including T cells, antigen presenting cells, and epithelial cells)



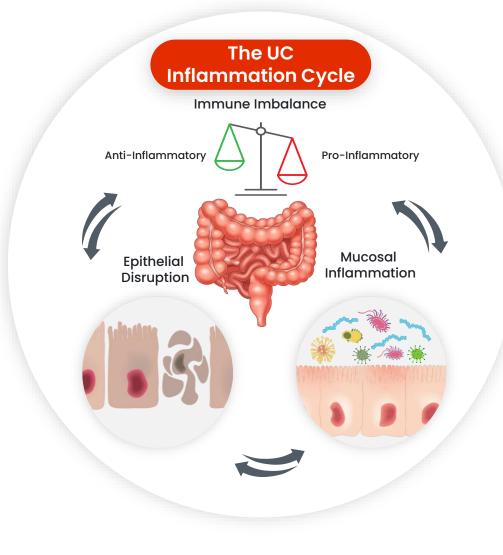
## The Role of Immunometabolism in Immunology & UC

## Immunometabolic response in inflammatory diseases in the immunology universe & UC:

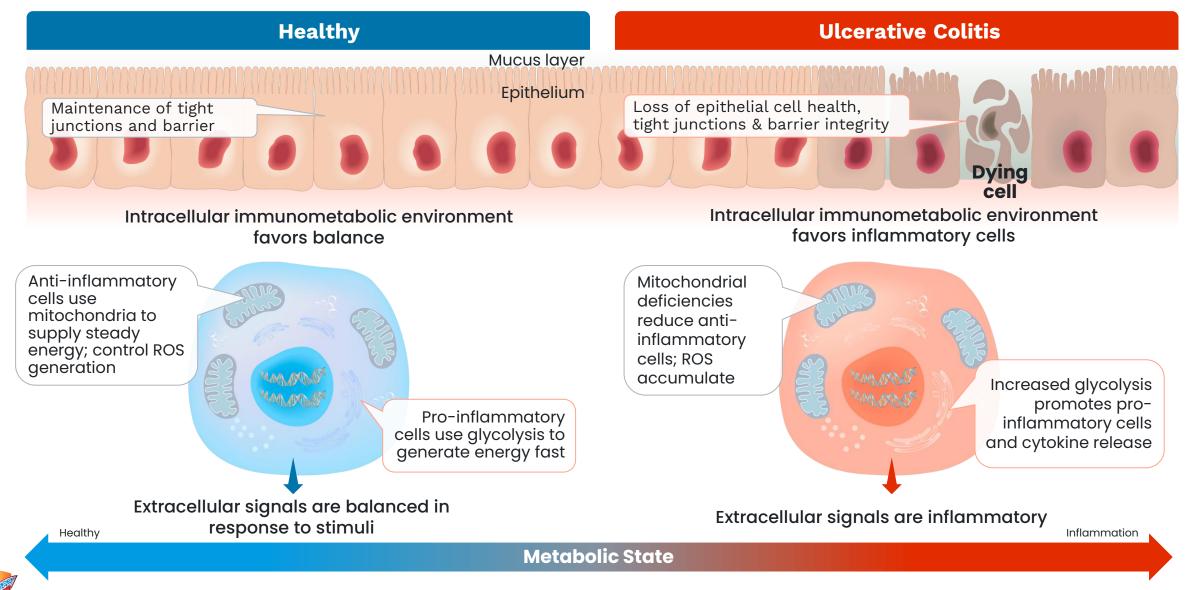
- Abnormal or imbalanced immune activation of the response resulting in over abundance of proinflammatory cells & cytokines with lack of antiinflammatory control.
- In UC, Pathogens cross the damaged epithelial barrier, activating immune response
- Immune activation is energetically costly, requiring the cell to use fast & inefficient glycolytic metabolism.

#### Multiple Factors contribute to the UC Inflammation Cycle:

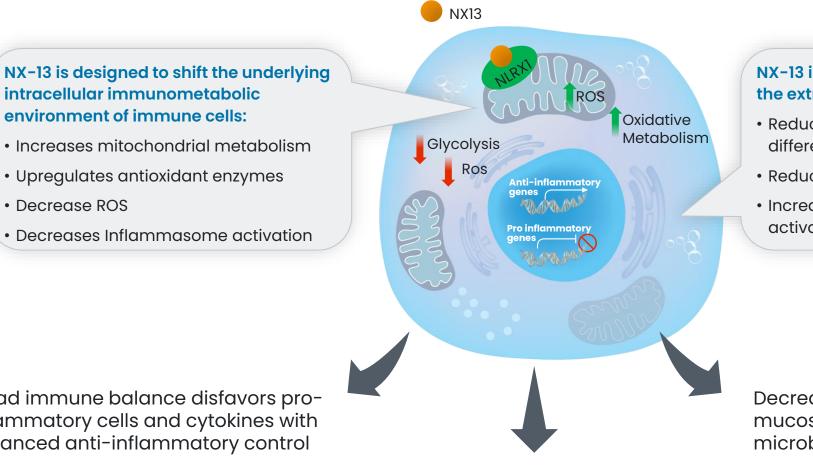
- Low grade Mucosal Inflammation and microbiome dysbiosis
- Epithelial Cell Damage and barrier disruption
- Broad Immune Activation favoring pro-inflammatory cells and cytokines



### Bimodal Targeting of the Intracellular Environment & Extracellular Inflammatory Response Aims to Control Multiple Factors in the UC Inflammation Cycle



## NX-13 Bimodal MOA Addresses Both Extracellular Signals and Intracellular **Environment to Reduce UC Inflammation Cycle**



#### NX-13 is designed to modulate the extracellular response:

- Reduces inflammatory cell differentiation
- Reduces TNFα, IFNγ, IL-17, IL-1.
- Increases anti-inflammatory activation

Broad immune balance disfavors proinflammatory cells and cytokines with enhanced anti-inflammatory control

• Decrease ROS

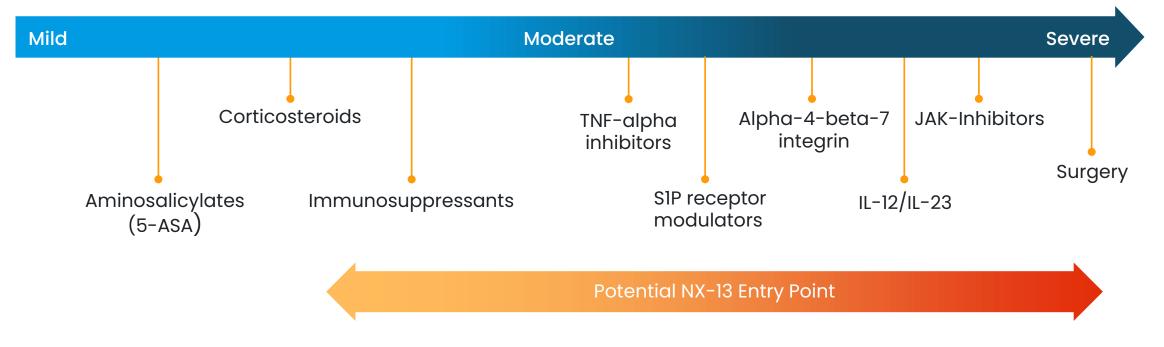
Improved epithelial barrier integrity to reduce exposure to inflammatory microbes

Decreased low grade mucosal inflammation and microbiome dysbiosis

## NX-13 Poised for Broad Utilization in Both Early & Late-Stage Disease

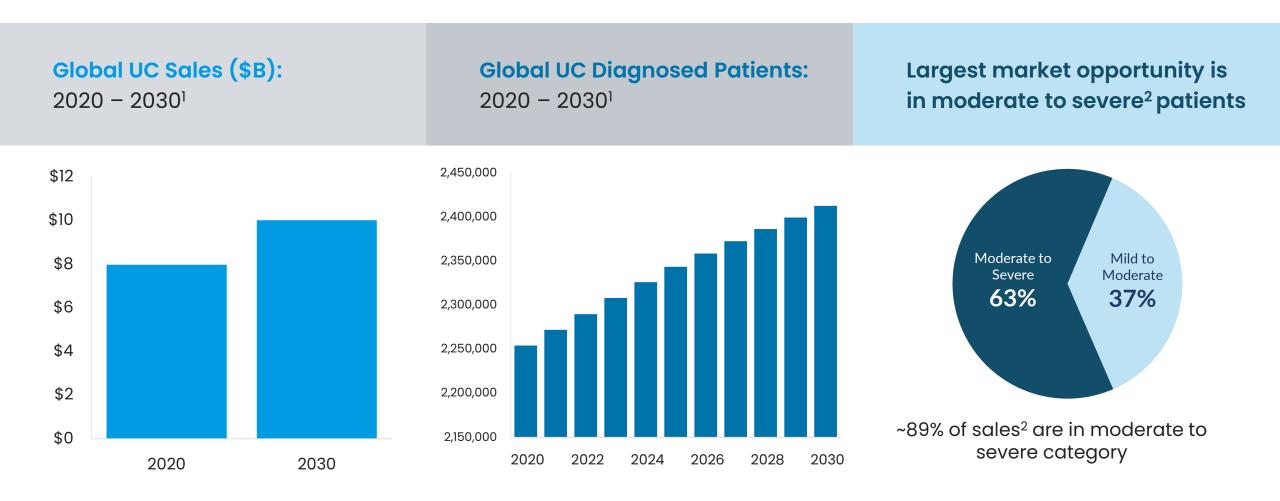
#### Potential benefits may help transform the current treatment paradigm:

- Gut selective allowing target engagement with the GI tract
- Novel MOA, with oral, once-daily dosing
- MOA may allow for improved efficacy, greater mucosal healing, and safety for long-term use
- No on-target toxicities associated with NLRX1, with Adverse Event incidence in Phase 1a & 1b similar to placebo



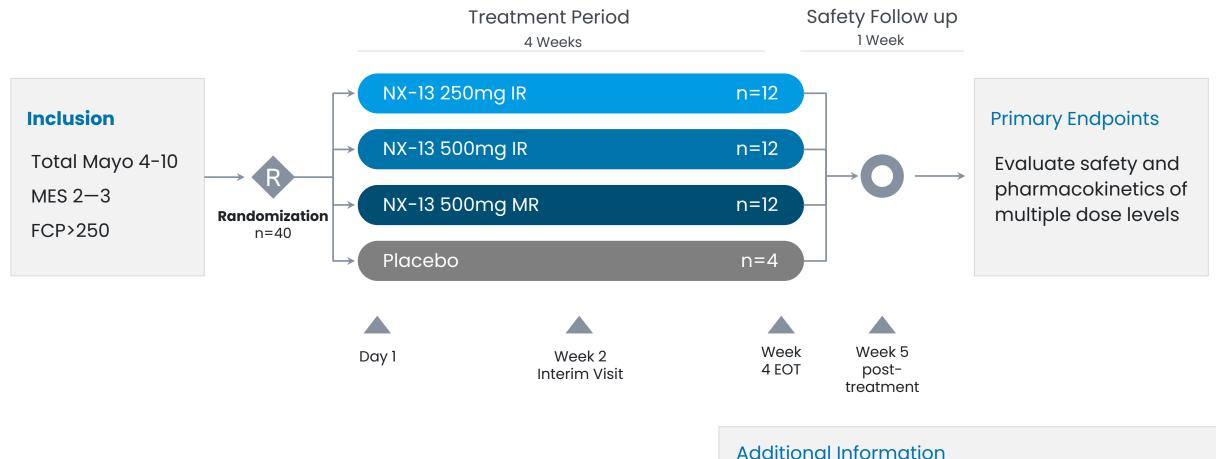


## Attractive & Growing Market Opportunity in UC



2022 Decision Resource Group (Clarivate) UC Disease Landscape and Forecast 2020-2030
April 2023 Global Data Ulcerative Colitis: Eight-Market Drug Forecast & Market Analysis 2021-2031; Severe category includes fulminant

## Phase 1b Study Design of NX-13 in Active UC



Iandosbiopharma.com/events-presentations (NX-13 Phase 1b Topline Data Presentation)



Note: Study was not designed or powered for exploratory clinical endpoints therefore results are hypothesis-generating only

## Phase 1b Results: NX-13 Demonstrated Favorable Endoscopic and Histologic Responses with Reductions in Multiple Clinical Measures After 4 Weeks

Patients receiving NX-13 IR doses responded best:

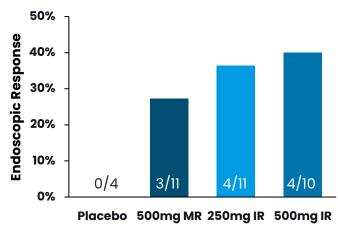
- Drug activity with IR formulation; study not designed for dose selection
- 72% of 250mg group achieved clinical response; 40% of 500mg IR group achieved clinical response
- 36-40% endoscopic response after just 4 weeks treatment across IR dosage groups
- 36-40% of patients receiving IR achieved histologic remission after 4 weeks of treatment

#### **Clinical Response**

Defined as CFB of at least -3, or -30% in Mayo Score **80% 60% 40% 20% 0**/4 3/11 8/11 4/10 Placebo 500mg MR 250mg IR 500mg IR

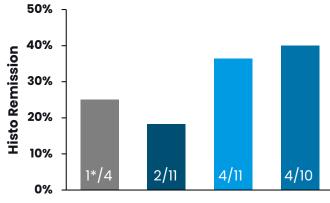
#### **Endoscopic Response**

MES CFB of at least -1



#### **Histologic Remission**

Geboes <3.1, no increased neutrophils in the LP



Placebo 500mg MR 250mg IR 500mg IR

\*Placebo patient started trial with Geboes < 3.1



Primary endpoints were safety and tolerability; Exploratory endpoints were efficacy and biomarkers;

IR= Immediate Release; MR= modified release designed to dissolve at the terminal ileum

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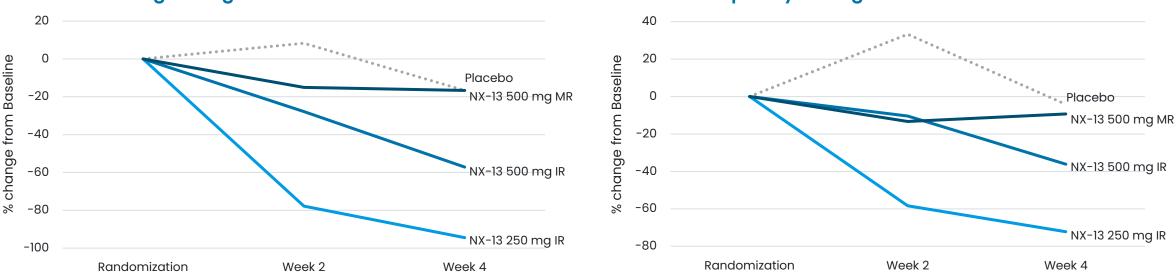
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## Phase 1b Results: Fast Onset of Action for NX-13 Supported Symptomatic Remission in Rectal Bleeding & Stool Frequency

250mg group had greatest reduction of Rectal Bleeding and Stool Frequency at 2 weeks, with further reduction at 4 weeks

Majority of patients treated once daily with 250mg NX-13, saw complete resolution of BOTH rectal bleeding and stool frequency after 4 weeks of treatment

**Stool Frequency Change from Baseline** 



#### Rectal Bleeding Change from Baseline

## Phase 1b Results: NX-13 Was Well-Tolerated & Shows Promising Signs of Clinical Improvement in Active UC



#### Generally well tolerated, consistent with non-clinical, Phase la data

- No Serious Adverse Events
- 3 unrelated Adverse Events (AEs) of note

### Pharmacokinetics

#### NX-13 was gut-selective with low systemic exposure

- IR dosing peaks ~1 hour post-dose
- No signs of NX-13 accumulation

## Efficacy



NX-13 induced early signs of clinical improvement in patient's symptoms by 2 weeks and endoscopy at 4 weeks:

 Positive signals of target engagement and downstream immunometabolic effects



## NEXUS Phase 2 Proof of Concept Trial

Goal	Evaluate safety, efficacy and pharmacokinetics of NX-13 in moderate to severe UC patients in 12-week induction trial		
Timing	Initiated in Q2 2023; Expecting to report topline results by Q4 2024		
Additional Phase 2 Learnings	Dose-Exposure-Response and PK/PD relationships (including site and MOA)		
Dosing	Oral, once daily treatment with either: 250 mg IR dose of NX-13   750 mg IR dose of NX-13   Placebo		
Blinded Key Design Principles	Powered Placebo Controlled Dose-Ranging   Image: Control in the second se		

## NEXUS Phase 2 Proof of Concept Study Design: NX-13 in Moderate to Severe UC

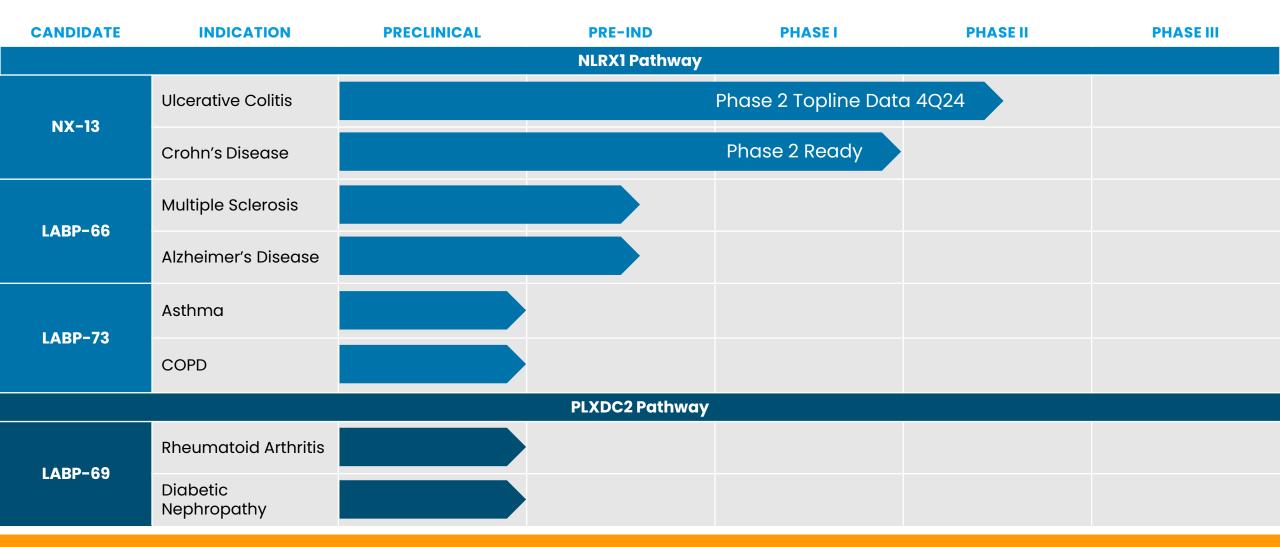


#### Additional Information

clinicalTrials.gov: NCT05785715



## Landos Pipeline Focused on Novel, Immunometabolic Targets



#### Significant optionality portfolio-wide for partnerships, development & future investment



Note: The Company is focused on advancing NX-13 clinical development in UC; Development and potential commercialization rights of NX-13 in China and select Asian markets licensed to LianBio; Research collaboration with Johns Hopkins University School of Medicine focused on advancing LABP-66 as a potential oral, once-daily therapy for MS and other disorders.

## **Experienced Management Team in Immunology & Drug Development**



#### **GREGORY OAKES**

President & Chief Executive Officer





#### FABIO CATALDI, MD

**Executive Vice President &** Chief Medical Officer





**CLAUDIA LOPEZ, DVM** Vice President, **Clinical Development** 

PHARMACEUTICALS





**DAWN LOURO** Vice President, Clinical Operations

> STATERA Supernus<sup>®</sup>



**REBECCA MOSIG, PHD** Executive Director, Corporate Development

> Mount Sinai



**DAVID PEREIRA, PHD** Vice President, CMC

cempra





**AMY PLACE, PHD** Vice President, Project Leadership & Site Engagement Chire *Cpfizer* 





Vice President, Controller & Principal Accounting Officer

🕑 PhaseBio gemini 🧐

**Deloitte** 



## **Top-Tier Advisory Teams**

### **Board of Directors**

### Scientific & Steering Committee

GREGORY OAKES President & Chief Executive Officer

CHRIS GARABEDIAN Chairman

#### ROGER ADSETT Chief Operating Office of Insmed, Inc.

FRED CALLORI Xontogeny, Perceptive Advisors

TIAGO GIRÃO CFO of Proteovant Therapeutics

TIM M. MAYLEBEN Director JEAN-FREDERIC COLOMBEL, MD Icahn School of Medicine at Mount Sinai

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BRAM VERSTOCKT, MD, PHD University Hospitals Leuven, KU Leuven

ANDRES YARUR, MD Cedars Sinai Medical Center



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## Thank you

**Contact:** IR@landosbiopharma.com