

# PURETECH

GIVING LIFE TO SCIENCE®

Corporate Presentation  
April 2024



# Important Information

The following presentation, including any printed or electronic copy of these slides, the talks given by the presenters, the information communicated during any delivery of the presentation and any question-and-answer session and any document or material distributed at or in connection with the presentation (together, the "Presentation"), has been prepared by PureTech Health plc (the "Company"). The information in the Presentation is not intended to form the basis of any contract. By attending (whether in person or by telephone) or reading the Presentation, you agree to the conditions set out below.

**THIS DOCUMENT AND THE PRESENTATION IS NOT A PROSPECTUS.** The Presentation does not constitute or form part of any offer or invitation to sell or issue, or any solicitation of any offer to purchase or subscribe for, any shares or other securities of the Company, nor shall there be any sale of these securities in any state or jurisdiction in which such offer, solicitation or sale would be unlawful prior to registration or qualification under the securities laws of any such state or jurisdiction. Any such offering of securities will only be made by means of a registration statement (including a prospectus) filed with the Securities and Exchange Commission (the "SEC"), after such registration statement becomes effective. No such registration statement has been filed as of the date of this presentation.

We report certain financial information using non-IFRS financial measures, as we believe these measures provide information that is useful to management and investors to assess financial performance. These non-IFRS financial measures do not have any standardized meaning and may not be comparable with similar measures used by other companies. For certain non-IFRS financial measures, there are no directly comparable amounts under IFRS. These non-IFRS financial measures should not be viewed as alternatives to measures of financial performance determined in accordance with IFRS. Please see slides 69-70 for a reconciliation of these measures to the most comparable IFRS measure.

This document and the Presentation contain statements that are or may be forward-looking statements. These statements are based on our management's current beliefs, expectations and assumptions about future events, conditions and results, and on information currently available to us. This document and the Presentation also contain estimates and other statistical data made by independent parties and by us relating to market size and growth and other data about our industry. This data involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such estimates. In addition, projections, assumptions and estimates of our future performance and the future performance of the markets in which we operate are necessarily subject to a high degree of uncertainty and risk.

All statements other than statements of historical facts included in this document may be forward-looking statements, including statements that relate to the Company's future prospects, developments and strategies. Words such as "expect," "anticipate," "intend," "plan," "believe," "seek," "estimate," "think," "may," "could," "will," "would," "should," "continue," "potential," "likely," "opportunity" and similar expressions or variations of such words are intended to identify forward-looking statements, but are not the exclusive means of identifying forward-looking statements. Additionally, statements concerning future matters such as our expectations of business and market conditions, development and commercialization of new products, enhancements of existing products or technologies, and other statements regarding matters that are not historical are forward-looking statements.

Such statements are based on currently available operating, financial and competitive information and are subject to various risks, uncertainties and assumptions that could cause actual results to differ materially from those anticipated or implied in our forward-looking statements due to a number of important factors including, but not limited to, those risks that are described in the Company's most recent Annual Report and Accounts which can be found on the Company's website at <https://investors.puretechhealth.com/financials-filings/reports> and in the Company's Annual Report on Form 20-F for the year ended December 31, 2022 filed with the Securities and Exchange Commission.

Given these risks, uncertainties and other factors, many of which are beyond the Company's control, you should not place undue reliance on these forward-looking statements.

Each forward-looking statement speaks only as at the date of this document. Except as required by law, we assume no obligation to update these forward-looking statements publicly, or to revise any forward-looking statements to reflect events or developments occurring after the date of this document, even if new information becomes available in the future.

The Presentation is confidential and should not be distributed, published or reproduced (in whole or in part) or disclosed by its recipients to any other person for any purpose, other than with the consent of the Company.

By attending any delivery of, or electronically accessing, the Presentation, you agree to be bound by the above limitations and conditions and, in particular, you represent, warrant and undertake to the Company that: (i) you will not retain in any manner the Presentation or forward the Presentation to any other person, or reproduce or publish this document, in whole or in part, for any purpose and (ii) you have read and agree to comply with the contents of this notice.

Our Founded Entities are comprised of our Controlled Founded Entities and our Non-Controlled Founded Entities, all of which are incorporated in the United States. References to our "Controlled Founded Entities" refer to Follica, Incorporated, and Entrega, Inc., for all periods prior to March 1, 2023, Vedanta Biosciences, Inc., for all periods prior to May 25, 2022, Sonde Health Inc., and for all periods prior to June 10, 2021, Alivio Therapeutics, Inc. References to our "Non-Controlled Founded Entities" refer to Akili Interactive Labs, Inc., Karuna Therapeutics, Inc., Vor Bio, Inc., Gelesis, Inc., for all periods following May 25, 2022, Sonde Health, Inc., for all periods following March 1, 2023, Vedanta Biosciences, Inc., and, for all periods prior to December 18, 2019, resTORbio, Inc. We formed each of our Founded Entities and have been involved in development efforts in varying degrees. In the case of our Controlled Founded Entities Follica, Incorporated and Entrega, Inc., we continue to maintain majority voting control. With respect to our Non-Controlled Founded Entities, we may benefit from appreciation in our minority equity investment as a shareholder of such companies.

# PureTech Team Has a Track Record of Outperforming

Oversaw R&D of therapeutics with 11 regulatory approvals; created several multibillion-dollar companies



**Bharatt Chowrira, PhD, JD**

Chief Executive Officer

30+ years of leadership roles as CEO, President, COO, and GC held in multiple biotech; including former COO at Auspex (acq. by Teva \$3.5B), COO at Nektar, GC at SIRNA (acq. by Merck \$1.1B), VP at Merck & Co.; Board Member



**Eric Elenko, PhD**

President

Co-inventor of KarXT & other key PureTech programs; oversaw 2 FDA & EU approvals; Co-founder & interim C-suite of multiple Founded Entities; Former McKinsey consultant for Fortune 500 & specialty pharmas



**Julie Krop, MD**

Chief Medical Officer

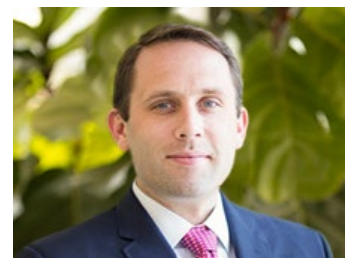
Former CMO at Freeline, AMAG (oversaw 3 FDA approvals; acquired by Covis group \$647M); Previously at Vertex, Millennium, Pfizer



**Robert Lyne**

Chief Portfolio Officer

Former CEO at Aris Bioscience (acq. by RTW Biotech \$250M); Previously at Touchstone Innovations, Bird & Bird; worked on >80 VC financings as well as multiple trade exits & IPOs



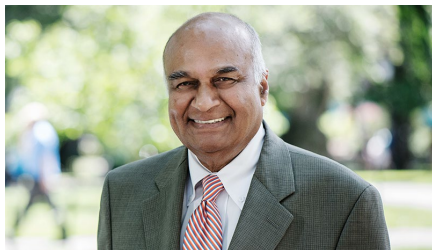
**Charles Sherwood, J.D.**

General Counsel

Former VP, Corporate Legal Counsel at Anika Therapeutics with extensive expertise in strategic transactions, IP, product & brand marketing, financing, securities compliance, etc.

# World Class Board of Directors Provides Strong Governance

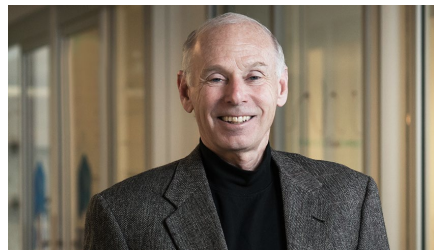
Our board contributed to regulatory approvals of approximately 20 drugs, led multi-billion-dollar strategic transactions & co-founded multiple companies



**Raju Kucherlapati, PhD**

Interim Board Chair

Harvard, Co-Founder of Millennium (acq. by Takeda \$8.8B) & Abgenix (acq. by Amgen \$2.2B)



**John LaMattina, PhD**

Board

Former President of Pfizer Global R&D, Forbes Contributor



**Robert Langer, ScD**

Board

MIT, Award winning materials science pioneer, Former member of the US FDA's SCIENCE Board, Co-founder of multiple biotech companies incl. Moderna & PureTech



**Daphne Zohar**

Senior Advisor & Board Observer

Founder & CEO of Seaport Therapeutics, BIO Board Member, Founder & former CEO of PureTech



**Robert Horvitz, PhD**

Board Observer & Chair of R&D Committee

Nobel Prize in Medicine, MIT, HHMI, neurobiologist at MGH, Former Novartis Scientific Advisory Board Member



**Kiran Mazumdar-Shaw**

Board

Founder & Chairperson of Biocon, Board of Trustees Member at MIT, Member of National Academy of Engineering



**Sharon Barber-Lui**

Board

CFO & Senior VP of Teva Pharma, Former CFO of Merck & Co. Inc. U.S. Oncology & Senior VP of EQRx





We are giving life to new  
classes of medicine to change  
the lives of patients with  
devastating diseases

# We Are Delivering on Our Mission to Change Patients' Lives

Outstanding track record of R&D productivity & clinical success

28



new therapeutics &  
therapeutic candidates  
generated to date

2



taken from inception to  
FDA & EU regulatory  
clearances

1

filed for FDA approval



>80%

of trials have been  
successful<sup>1</sup>

6x

better probability of clinical  
success compared to the  
industry average<sup>2</sup>

# Distinctive Approach

R&D engine is repeatable and scalable

## VALIDATED EFFICACY



Advancing new **medicines** with proven clinical **efficacy** previously held back by limitations

## CLEAR PATIENT BENEFIT



Applying **proprietary technologies** to address key limitations and **unlock drug potential** for **patients**

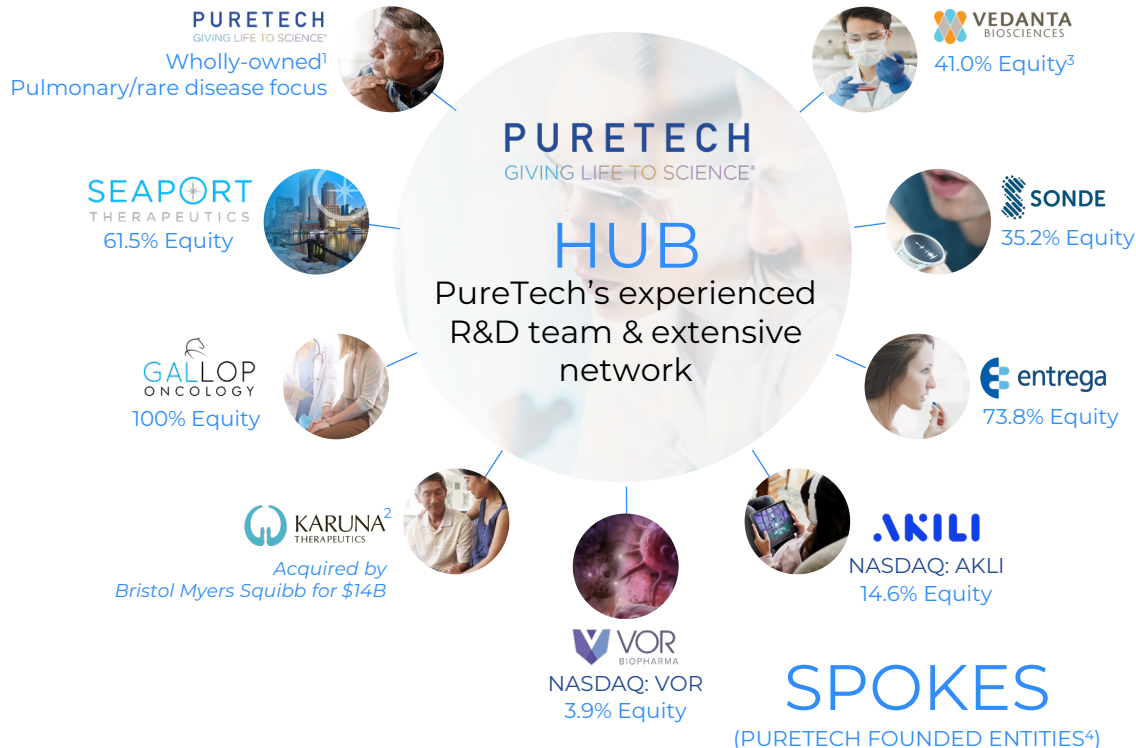
## EFFICIENT & DE-RISKED PATH



Building on **well-defined clinical & regulatory paths** backed by our **proven R&D track record**

# The PureTech-Pioneered Hub-and-Spoke Model

Robust pipeline of new medicines poised for tremendous growth



## CAPITAL EFFICIENT MODEL

- ✓ **\$320M** estimated Consolidated Cash, Cash Equivalents & Short-Term Investments<sup>5</sup>, excluding the ~\$293M proceeds from the BMS/Karuna transaction
- ✓ Operational runway into **2027**
- ✓ PureTech has not needed to raise capital in **~6 years**
- ✓ **\$17.8B** generated by Founded Entities since July 2018



# KarXT Case Study – Invented & Advanced by PureTech

1st new mechanism for treating schizophrenia in over 50 years

## PATIENT NEED

~**2.7M** living with schizophrenia in the US

~**3.2M** with Alzheimer's disease psychosis in the US

Current antipsychotics **have significant side effects and poor adherence**

**Xanomeline: clinical efficacy but was sitting on a shelf at Eli Lilly**



## PURETECH ROLE

Built top team of CNS experts & leaders

- ✓ **PureTech invented & filed patents** to cover the agonist/antagonist concept
- ✓ **Completed tolerability POC**
- ✓ Planned Phase 2 EMERGENT-1 study



**Xanomeline**  
CNS active agonist

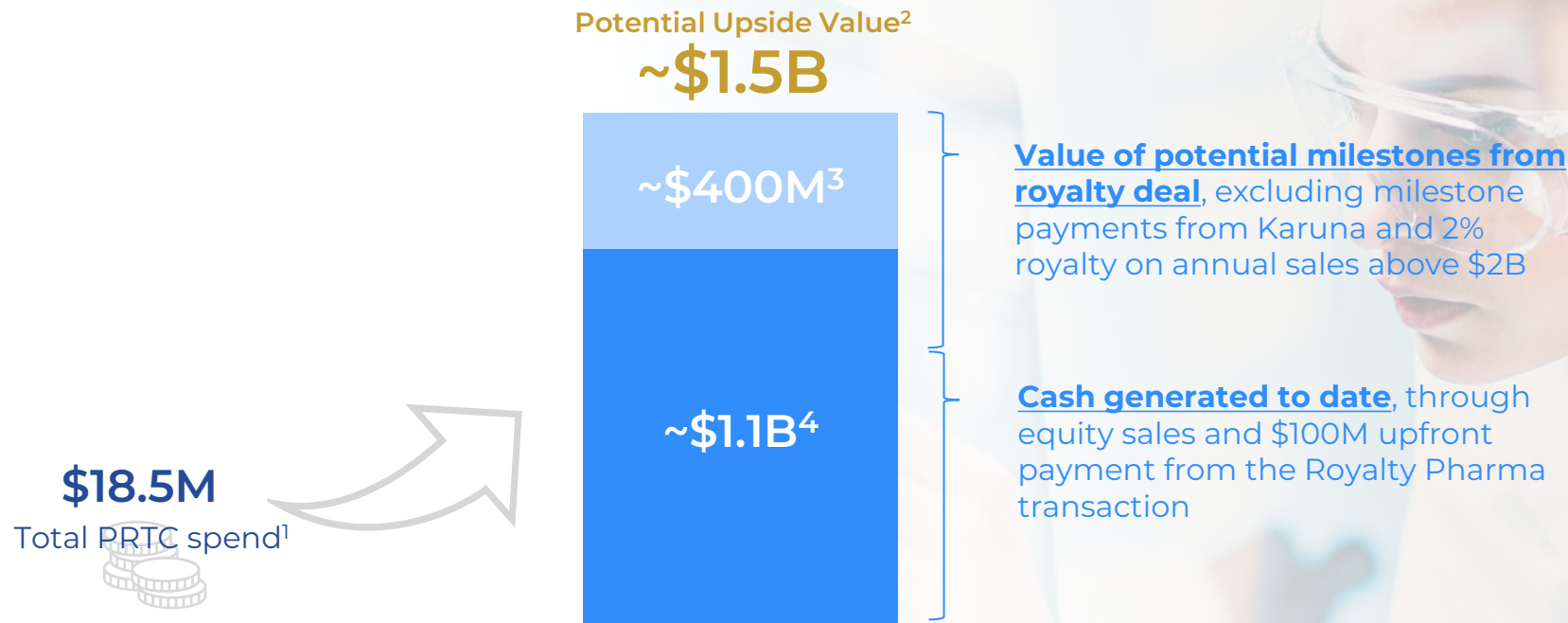
**Trosipium chloride**  
Peripheral antagonist  
blocks side effects of agonist

## VALUE REALIZATION

- ✓ Phase 2 EMERGENT-1, Phase 3 EMERGENT-2 & Phase 3 EMERGENT-3 trials **met primary endpoint** with clinically meaningful & significant **reduction in PANSS total score** vs. placebo
- ✓ NDA submission completed in September 2023; PDUFA date, September 26, 2024
- ✓ Ongoing Phase 3 programs in **psychosis in Alzheimer's disease**
- ✓ Karuna Therapeutics acquired by Bristol Myers Squibb for **\$14B**

# Generating Value for Patients and Shareholders

## KarXT Case Study Part 2



# The PureTech-Pioneered Hub-and-Spoke Model

Robust pipeline of new medicines poised for tremendous growth



# LYT-100 for Idiopathic Pulmonary Fibrosis (IPF)

ORPHAN DESIGNATION: ~120,000 patients in the US, ~110,000 in the EU<sup>1</sup>



## FATAL & PROGRESSIVE

Causes scar tissue in the lungs, leading to **shortness of breath and loss of lung function**<sup>2</sup>

Median survival 2 – 5 years<sup>3</sup>

## UNMET MEDICAL NEED

2 standard of care treatments proven to slow disease progression, but **have significant side effects, including nausea, vomiting and diarrhea**<sup>4,5</sup>

Phase 2b trial ongoing; Topline data expected in Q4 2024

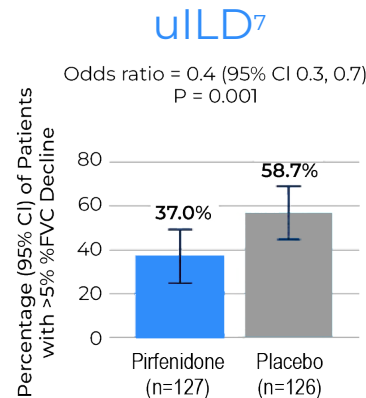
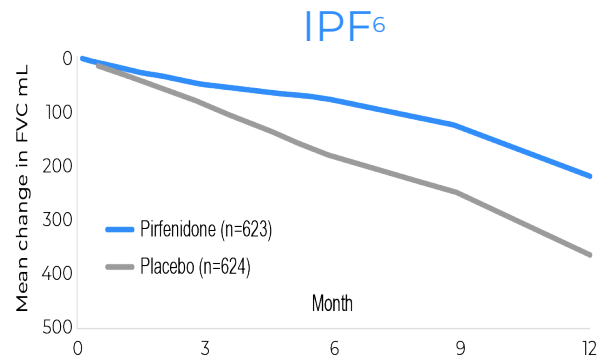
# Pirfenidone:

Clinically validated anti-fibrotic & anti-inflammatory

## VALIDATED EFFICACY

- ✓ Pirfenidone FDA-approved for IPF with breakthrough designation for Unclassifiable Interstitial Lung Disease (uILD); has been shown **to extend life in patients with IPF by an average of ~2.5 years**<sup>1</sup>
- ✓ Over a **dozen late-stage & real-world efficacy studies** demonstrate **efficacy in IPF**<sup>2</sup>
- ✗ BUT GI-related tolerability issues significantly limit its usage resulting in **~50% who discontinue, dose adjust, or switch**<sup>3</sup> & **3 out of every 4 patients are not on standard of care**<sup>4</sup>

Despite drawbacks, 2022 sales of both SOC treatments combined were ~\$4B<sup>5</sup>



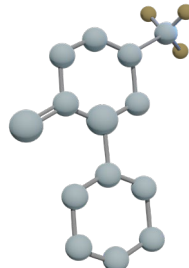
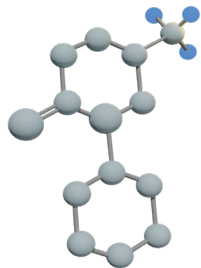


# LYT-100:

A potential game changer for IPF patients

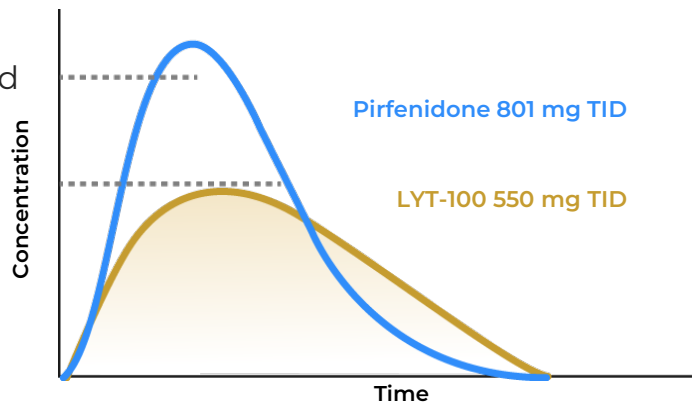
## PIRFENIDONE

- ✓ Clinically validated efficacy
- ✗ Associated with GI AEs
- ✗ Higher exposure limited by tolerability



## LYT-100

- ✓ Substantially improved adverse event profile<sup>1</sup>
- ✓ Pharmacology (activity) maintained
- ✓ Higher dose could improve efficacy; maximum tolerated dose not determined



# LYT-100: Efficient & De-risked Path

- ✓ **Composition of Matter Patent** exclusivity up to 2033 with PTE
- ✓ **Additional IP coverage** to ~2044
- ✓ Potential for **Orphan Drug Exclusivity for IPF** & other indications
- ✓ **Streamlined development program** using the same endpoints that have supported past approvals
- ✓ Potential to become the **frontline therapy for IPF**

# LYT-100:

Data to date (N>400)<sup>1</sup> demonstrate favorable safety & tolerability profile

## HEALTHY OLDER ADULT CROSSOVER TRIAL (N=49)

TEAE	LYT-100 550mg TID n (%)	Pirfenidone 801mg TID n (%)
<b>Gastrointestinal</b>	<b>8 (17.4%)</b>	<b>16 (34.0%)</b>
Nausea	7 (15.2%)	14 (29.8%)
Vomiting	2 (4.3%)	3 (6.4%)
Abdominal Pain/Distension	1 (2.2%)	3 (6.4%)
<b>Nervous System</b>	<b>8 (17.4%)</b>	<b>15 (31.9%)</b>
Headache	6 (13.0%)	9 (19.1%)
Dizziness	1 (2.2%)	7 (14.9%)
Somnolence	1 (2.2%)	2 (4.3%)

## LYT-100 DEMONSTRATED IMPROVED TOLERABILITY

Achieved **~50% reduction** in healthy older adults experiencing **GI-related AEs compared to pirfenidone** in crossover trial

## IN OTHER LYT-100 STUDIES

Multiple Ascending Dose trial (N=40): without dose titration<sup>2</sup>; well-tolerated at all doses studied<sup>3</sup>; all treatment-related **AEs** were **mild & transient**

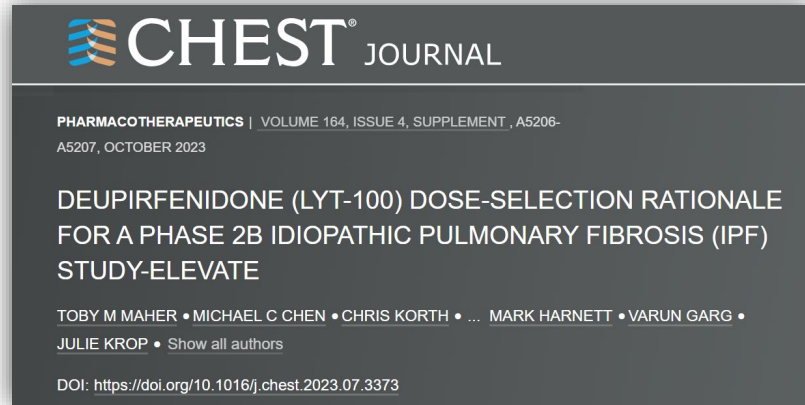
**Higher dose (824mg TID LYT-100)** in the 2<sup>nd</sup> Multiple Ascending Dose (N=24): well-tolerated with no additional incident<sup>4</sup>

Long COVID trial: strong safety & tolerability profile

# Potential for Higher Dose of LYT-100 Presented at CHEST '23

Higher dose could potentially translate to enhanced efficacy in IPF

- ✓ LYT-100 (824mg TID) achieved a 43% higher exposure level than approved dose of pirfenidone (801 mg TID)<sup>1</sup>
- ✓ LYT-100 (824mg TID) well-tolerated with no additional incidence of GI or CNS AEs when titrated up from LYT-100 550 mg TID



## WHAT DOES THIS MEAN?

**LYT-100 can be studied at a higher dose that could provide enhanced efficacy with favorable tolerability in IPF**, based on the pirfenidone Phase 3 CAPACITY trial where approved dose of pirfenidone demonstrated better efficacy than a lower dose

# Registration-enabling Program in IPF Guided by Leading Experts

PureTech's clinical advisory board for IPF & related lung disorders



**BILL BRADFORD, MD, PHD**

Former SVP InterMune;  
developed pirfenidone for the  
treatment of IPF



**VINCENT COTTIN, MD**

Professor at Université Claude  
Bernard Lyon; Coordinator of  
Center for Rare Pulmonary  
Diseases at Louis Pradel Hospital;  
Section Editor of the *European  
Respiratory Journal*



**KEVIN FLAHERTY, MD**

Professor at University of  
Michigan; PhIII trial of  
nintedanib in pILD (*NEJM*)



**TOBY MAHER, MD, PHD**

Professor & Director of ILD at  
Keck School of Medicine, USC;  
PhII trial of pirfenidone in  
uILDs (*Lancet RM*)



**PAUL NOBLE, MD**

Chair, Department of Medicine,  
Cedars-Sinai; results of two late-  
stage studies evaluating the  
effect of pirfenidone in patients  
w/ IPF (*Lancet*)



**MARLIES WIJSENBEEK,  
MD, PHD**

Chair of Erasmus Medical  
Center ILD program; PI on  
study to identify disease  
progression in patients with  
newly diagnosed pILDs



# LYT-100: Multiple Ways to Win

Current SOC has significant tolerability issues, with GI side effects being the most problematic

## BASE CASE: BETTER SAFETY

- ✓ LYT-100 **550 mg** demonstrates better safety



- ✓ Patients **can stay on the drug longer**



- ✓ Patients may achieve **more durable efficacy** compared to pirfenidone

## UPSIDE: BETTER EFFICACY

- ✓ LYT-100 **825 mg** demonstrates better efficacy



- ✓ Patients **retain more lung function**



- ✓ Patients may achieve **superior efficacy**

# LYT-100: Phase 2b Trial

1st of two potentially registration-enabling studies in patients with IPF

## PRIMARY AIM:

To evaluate activity of LYT-100 in patients with IPF

## PRIMARY ENDPOINT:

Slope of decline in FVC for LYT-100 compared to placebo over 6 months

## TRIAL DESIGN

**N= ~240 IPF patients**

- Placebo
- Pirfenidone 801 mg TID
- LYT-100 550 mg TID
- LYT-100 825 mg TID

**6-month treatment duration**



Phase 2b topline data expected in Q4 2024

# LYT-100: Potential to Address Multiple Underserved Diseases

## CURRENT INDICATION: IPF



Pirfenidone has been shown to improve survival by approximately 3 years compared to supportive care alone<sup>1</sup>

Pirfenidone reduces decline in lung function<sup>2</sup>

**Topline results from registration-enabling trial expected in 4Q 2024**

## NEXT INDICATION: PF-ILD



~650K non-IPF PF-ILD patients in the 16 major markets<sup>3</sup> are affected with few treatment options worldwide

Pirfenidone derisked due to similar pathophysiology of IPF and PF-ILD & clear development path

Pirfenidone showed initial efficacy signals in Ph2 RELIEF study

## MEDICAL COUNTERMEASURES



LYT-100 as medical countermeasure

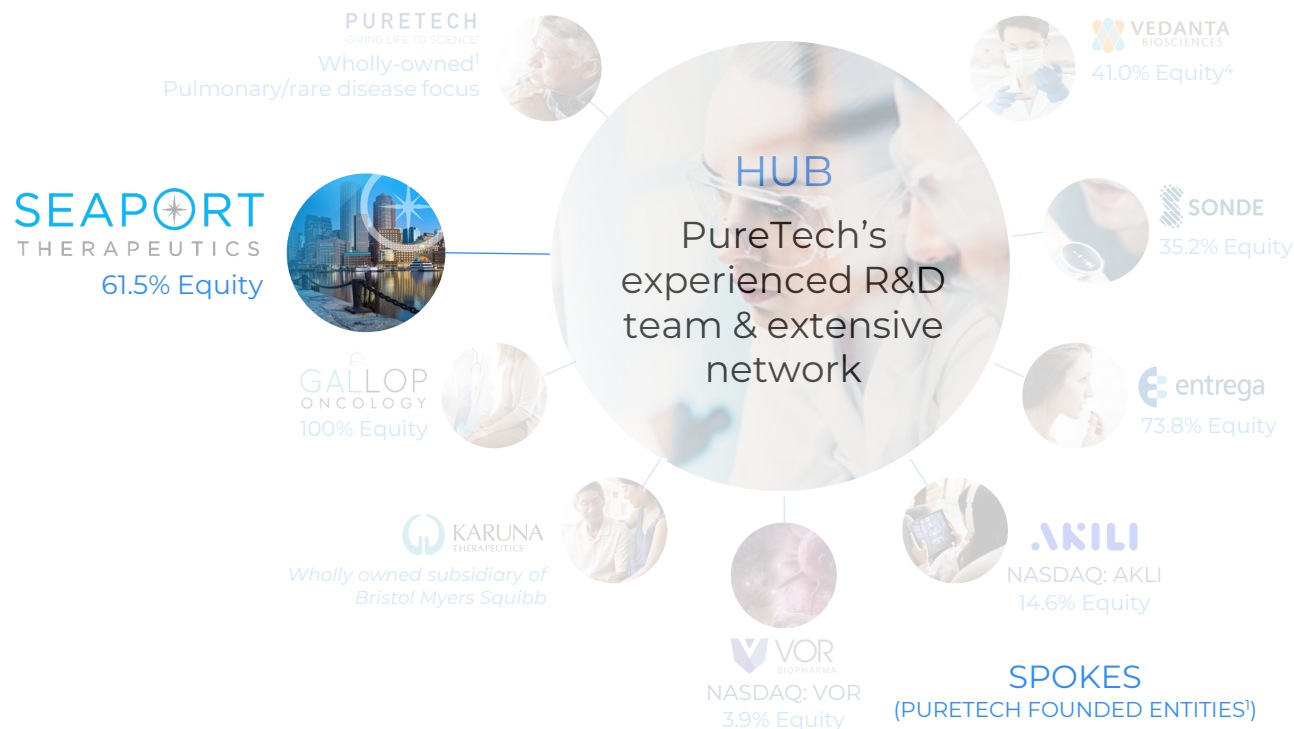
Pirfenidone mitigates radiation-induced lung fibrosis in preclinical study<sup>4</sup>

Subject to Animal Rule; allows for approval based on animal models when human efficacy studies are not feasible

Additional opportunities where pirfenidone has shown human efficacy signals (e.g. HFpEF, FSGS)

# The PureTech-Pioneered Hub-and-Spoke Model

Robust pipeline of new medicines poised for tremendous growth

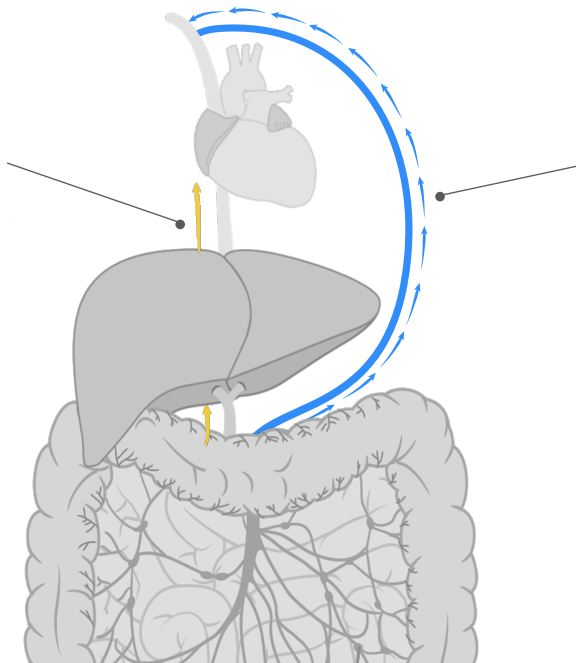


# Glyph™: Leveraging the Lymphatic System to Unlock New Medicines

Proprietary platform advances active drugs previously limited by low oral bioavailability/hepatotoxicity

## CONVENTIONAL

Oral drugs with high first-pass metabolism can have low bioavailability & hepatotoxicity



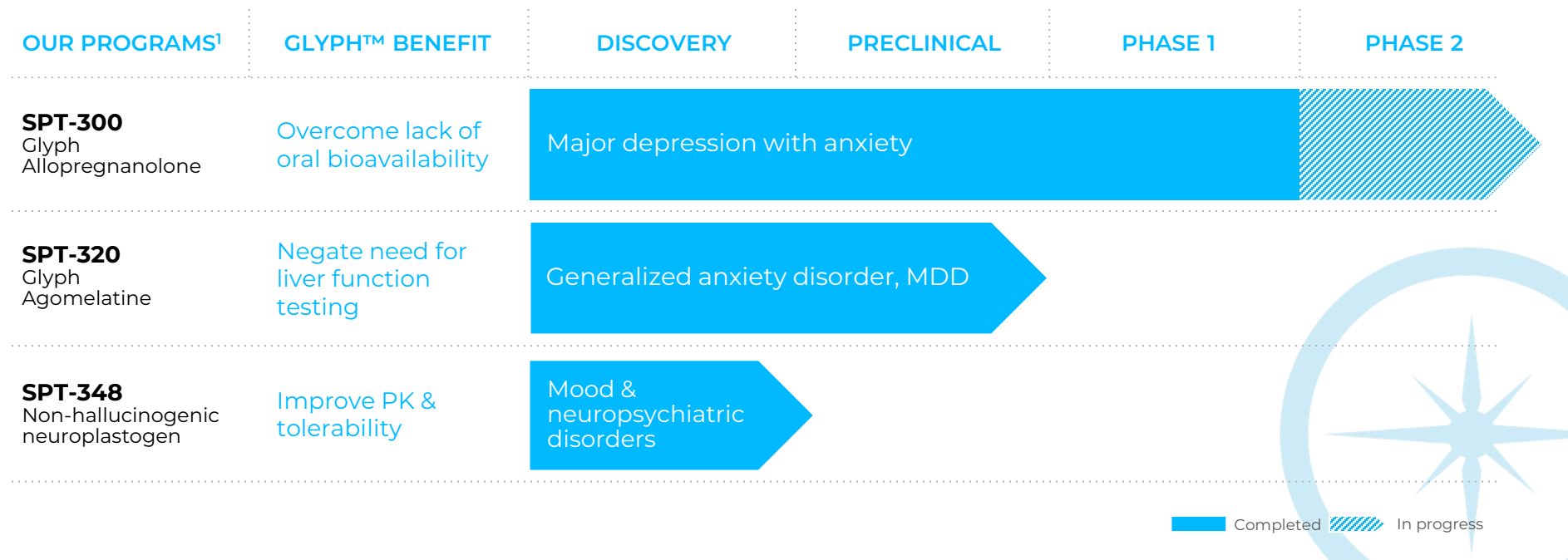
## GLYPH™

Employ the lymphatic system's natural lipid absorption and transport process to bypass the liver, as a result:

- ✓ Enhances oral bioavailability
- ✓ Reduces dose
- ✓ Reduces first-pass hepatotoxicity
- ✓ Provides novel composition IP



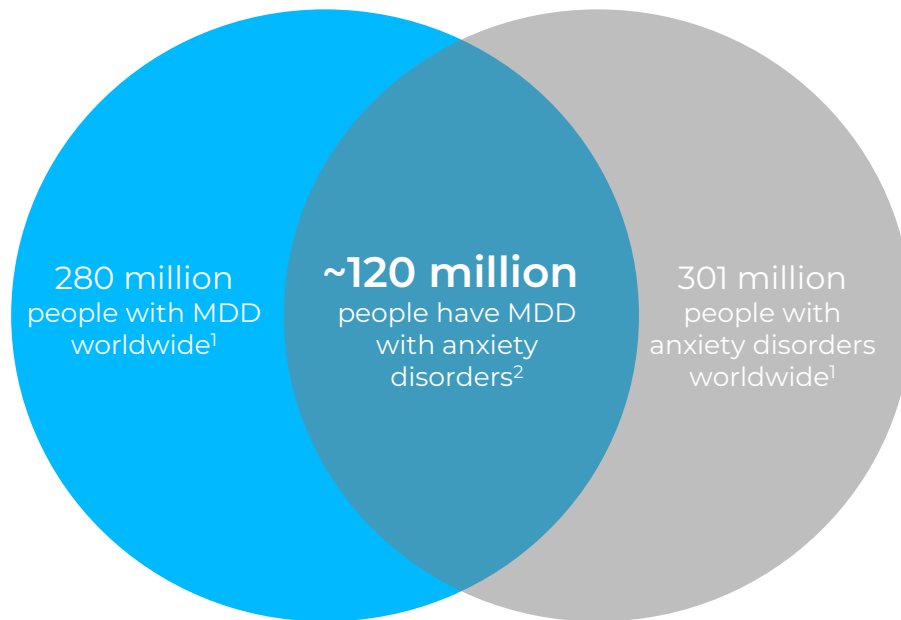
# Pipeline of First & Best-in-Class CNS Medicines



Multiple discovery/preclinical programs underway leveraging the Glyph™ platform

# SPT-300: First Tailored Treatment for MDD with Anxiety

Large unmet need for new therapies to address multiple mental health disorders



## MDD WITH ANXIETY

- MDD patients with anxiety, compared with MDD patients without anxiety<sup>3</sup>:
  - Less likely to achieve remission
  - Slower to respond to treatment
  - Poorer quality of life

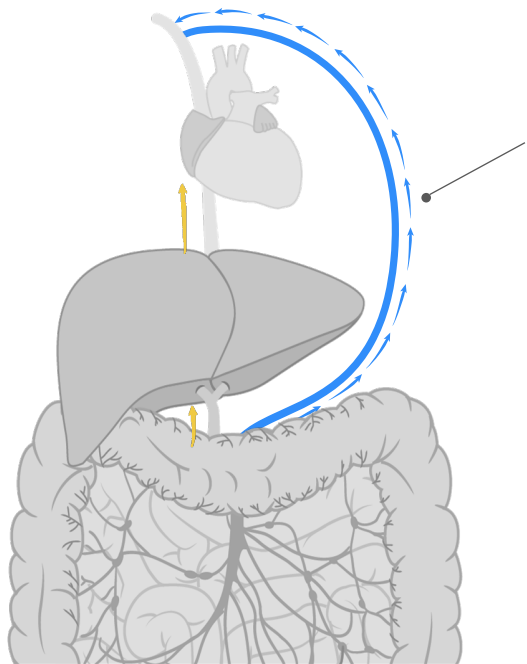
# SPT-300 (Glyph Allopregnanolone)

For major depressive disorder with anxiety



- ✓ Allopregnanolone is an endogenous (natural) neurosteroid with clinical validation in postpartum depression
- ✓ Lower levels of allopregnanolone have been documented in patients with mood disorders
- ✗ ...BUT method of administration (IV form) significantly limits patient uptake
- ✗ Oral chemical analogs have different composition than endogenous (natural) allopregnanolone and may not capture its full therapeutic potential
- ✓ **SPT-300 retains the activity & potency of endogenous allopregnanolone in an oral form**

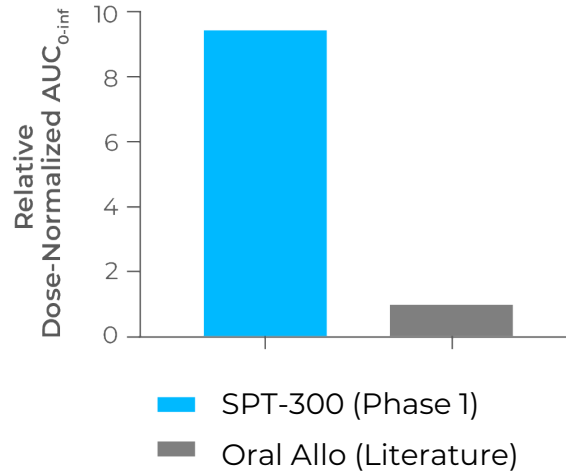
# SPT-300 (Glyph Allopregnanolone)



## GLYPH SPT-300 PHASE 1

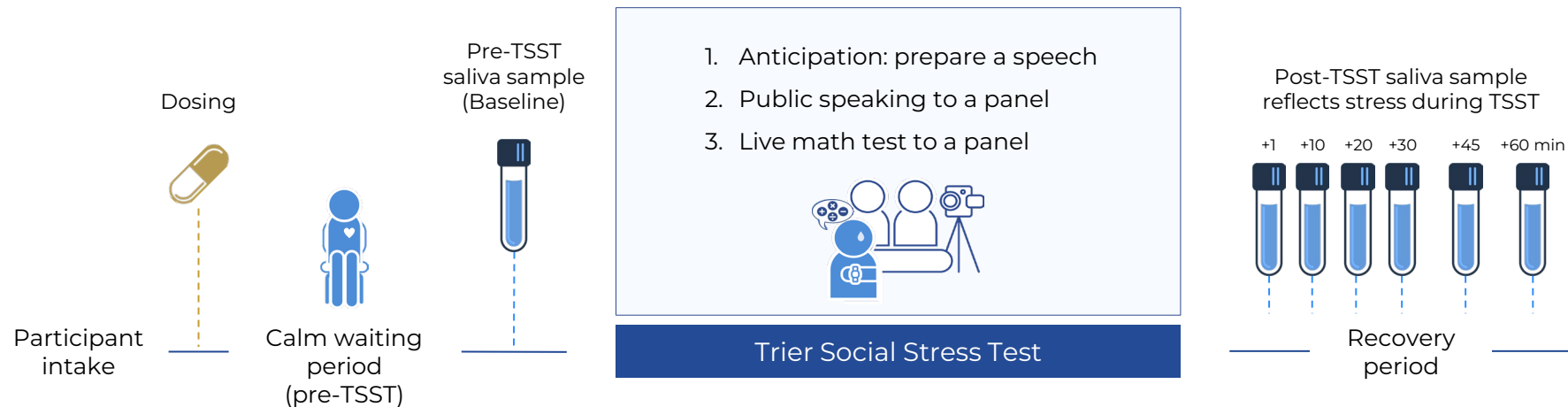
- ✓ Bioavailability >9X vs. oral allopregnanolone<sup>1</sup>
- ✓ Generally well-tolerated, AEs generally mild and transient
- ✓ Most common AE was somnolence (on-target effect of GABA<sub>A</sub>)
- ✓ No treatment-related severe or serious AEs
- ✓ No sudden loss of consciousness observed

## SPT-300 ORAL SYSTEMIC EXPOSURE (HUMAN) VS LITERATURE DATA<sup>1,2</sup>



# Phase 2a Trial Design in Acute Anxiety

Randomized, placebo-controlled trial in the Trier Social Stress Test (TSST)



## PRIMARY AIM:

To characterize pharmacology of SPT-300 for potential anxiety indications

## PRIMARY ENDPOINT:

Reduction in salivary cortisol, a stress hormone

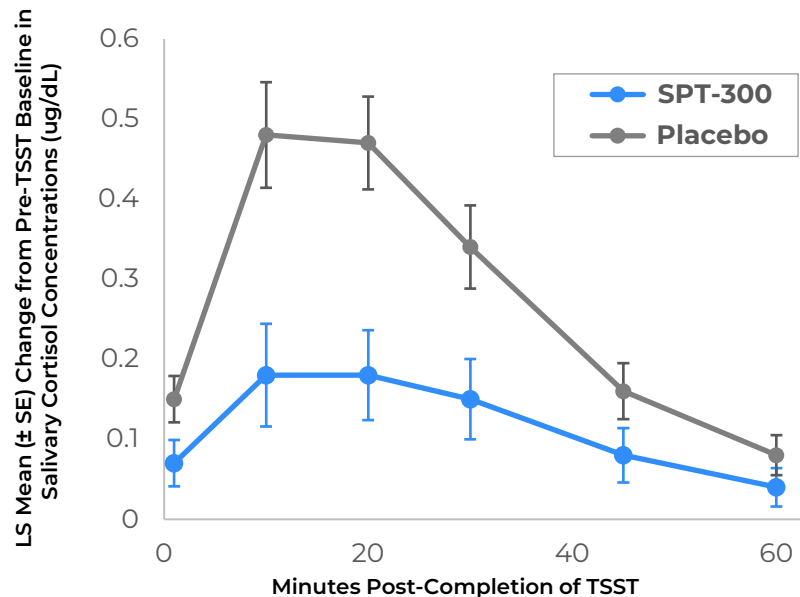
## TRIAL DESIGN:

N=80 randomized to SPT-300 or placebo

# Positive Phase 2a Study for SPT-300 in The Trier Social Stress Test

SPT-300 achieved primary endpoint ( $p=0.0001$ ) in stress hormone response<sup>1</sup>

## POSITIVE DATA



- ✓ SPT-300 had an effect size (Cohen's  $d = 0.72$ )<sup>2</sup>
- ✓ Generally well tolerated: All treatment-related adverse effects were transient, mild or moderate

## VALIDATION

- ✓ Further supports the potential of SPT-300 for anxiety disorders
- ✓ Further validates the Glyph platform



# SPT-320 (Glyph Agomelatine)

For anxiety & mood disorders

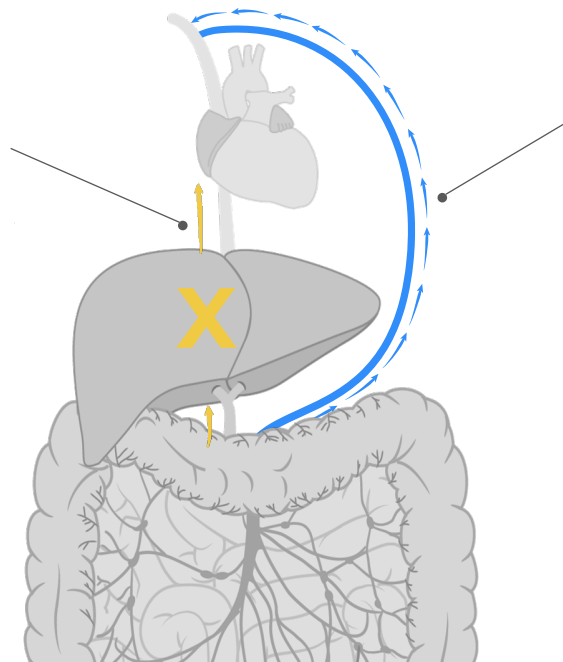


- ✓ Clinically validated and approved for MDD in the EU and MDD & GAD in Australia
- ✓ Differentiated mechanism of action
- ✓ Consistent and statistically significant against placebo in GAD (4/4 studies)
- ✓ Superior efficacy and tolerability vs. standard-of-care<sup>1,2</sup>
- ✗ ...BUT it has low oral bioavailability and is associated with hepatotoxicity necessitating liver function monitoring
- ✓ **SPT-320 has the potential to greatly reduce the risk of clinically significant liver enzyme elevations<sup>3</sup>**

# SPT-320: First-In-Class Potential for GAD & MDD

## AGOMELATINE

- ✗ Low oral bioavailability (~1%)
- ✗ Most of the drug does not reach the brain
- ✗ Liver enzymes increase likely linked to hepatic first-pass metabolism:  
~1.0% – 1.4% agomelatine vs. 0.7% with placebo<sup>1</sup>

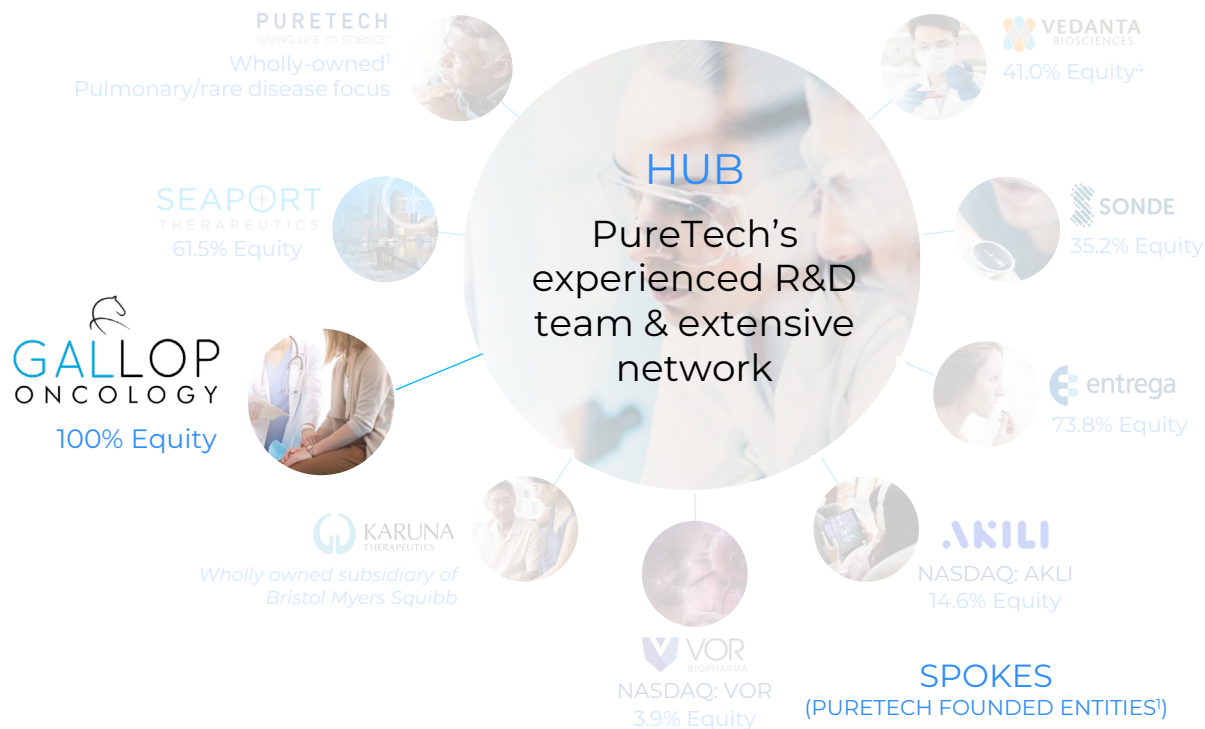


## SPT-320

- ✓ SPT-320 bioavailability is >10X of agomelatine based on non-human primate PK
- ✓ SPT-320 potential for therapeutic exposure with reduced risk of liver enzyme elevations

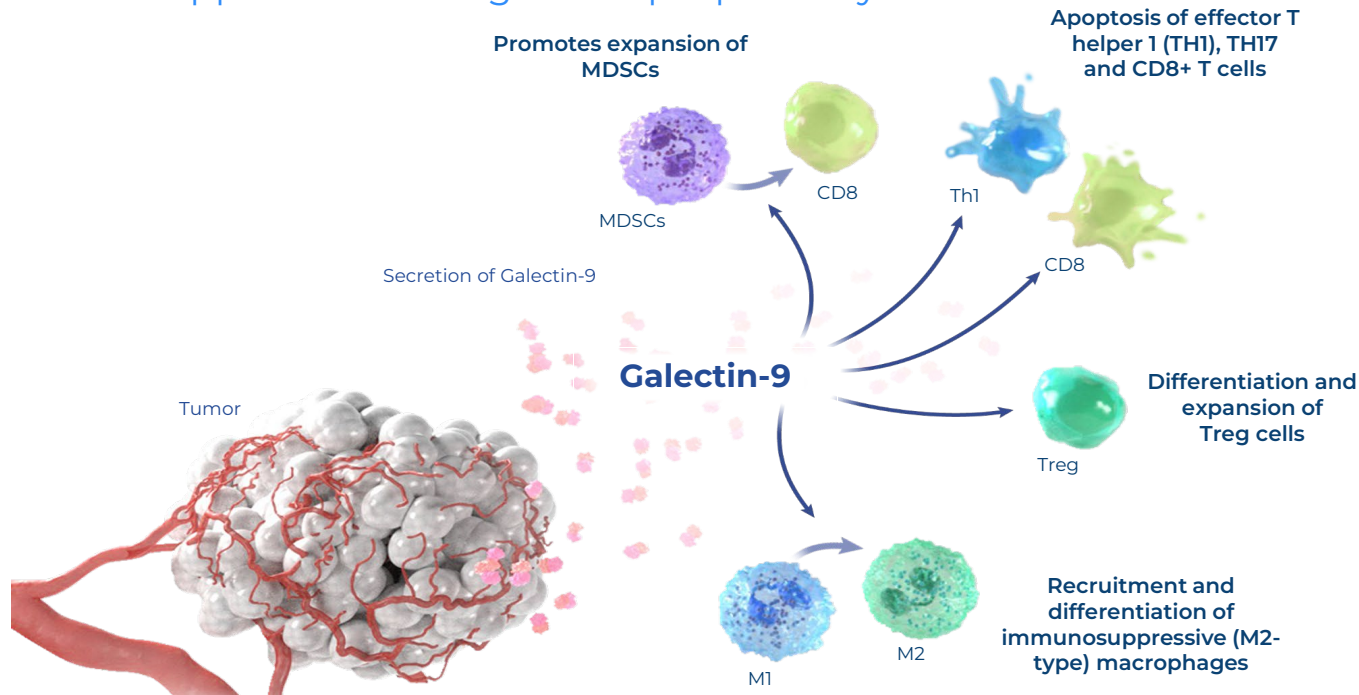
# The PureTech-Pioneered Hub-and-Spoke Model

Robust pipeline of new medicines poised for tremendous growth



# Gallop Oncology: Advancing Galectin-9 Targeting mAb, LYT-200

Driving immunosuppression through multiple pathways



Received **orphan drug designation** from the FDA  
for the treatment of AML

Received **Fast Track designation** from the FDA  
for the treatment of head and neck cancers

# Gallop Oncology:

LYT-200 (Anti-Galectin-9 mAb) for hematological malignancies and metastatic solid tumors



## LIFE ALTERING & PROGRESSIVE

### Hematological Malignancies

~**60,000 new cases of leukemia** diagnosed each year,<sup>1</sup> including  
~20,000 in acute myeloid leukemia (AML)<sup>2</sup>

### Solid Tumors

- ~ **82,000 new cases of bladder cancer** diagnosed each year<sup>3</sup>;  
~90% are urothelial carcinoma<sup>4</sup>
- ~**66,000 new cases of head and neck cancers** diagnosed each year<sup>5</sup>; ~10% metastatic disease at diagnosis & additional 20-30% will develop metastases<sup>6</sup>

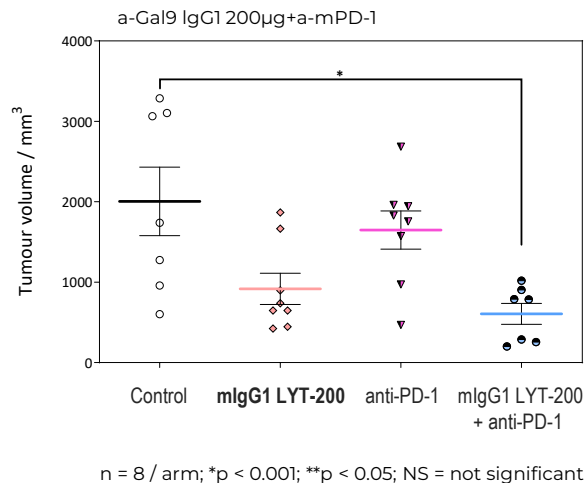
## UNMET MEDICAL NEED

**Over 50%** of AML patients either don't respond to initial treatment or experience relapse or death,<sup>7</sup> with ~12.6% five-year survival rate<sup>8</sup>

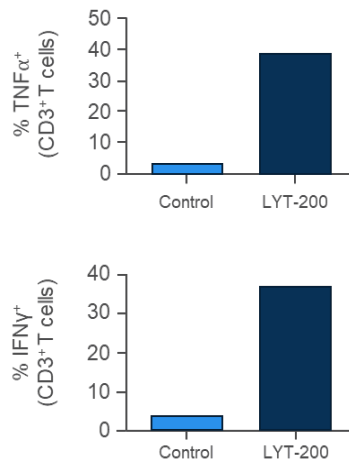
# LYT-200

## Multiple lines of preclinical data supporting therapeutic potential

### SINGLE AGENT ACTIVITY IN B16F10 MELANOMA MODEL



### T CELL ACTIVATION WITH LYT-200 IN PATIENT-DERIVED ORGANOID<sup>1</sup> MODEL



### LYT-200 DRUG PROPERTIES MAKE IT AN EXCELLENT CLINICAL CLONE:

#### High affinity & specificity for galectin-9

#### Robust activity in preclinical studies:

- Single agent causes tumor reduction in pancreatic models where anti-PD-1s don't work
- ~50% tumor reduction with LYT-200 vs. ~22% tumor reduction with anti-PD-1 in melanoma model
- Increase in intra-tumoral CD8 T cells in combination with anti-PD-1
- Activation of intra-tumoral immunity in patient-derived tumor models

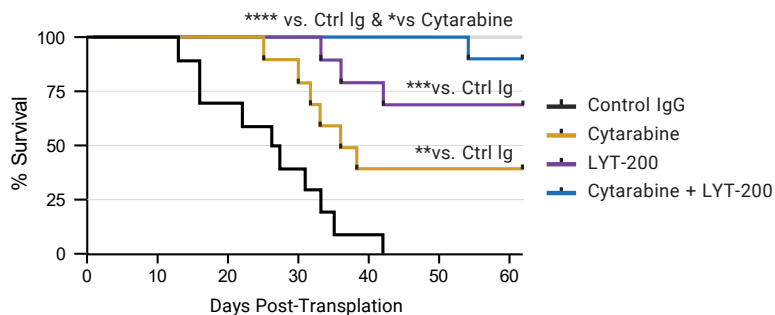
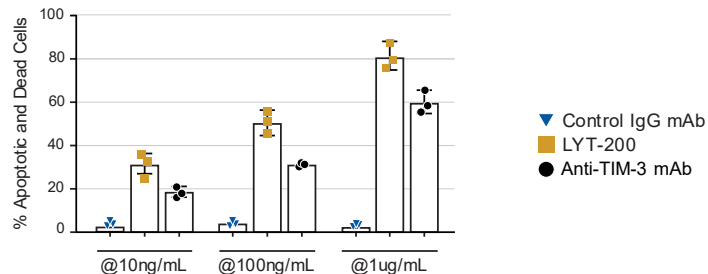


# LYT-200

Administration induces apoptosis of leukemia cells & extends survival of leukemia cell engrafted animals

## AML MODEL<sup>1</sup>

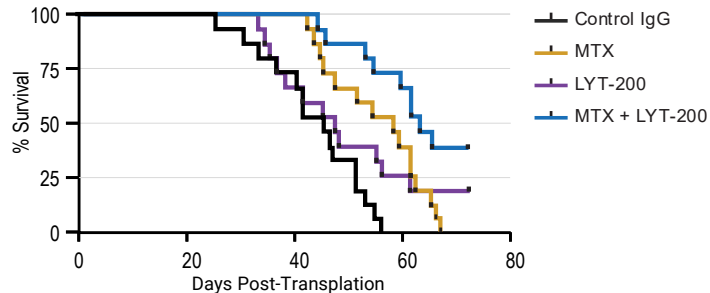
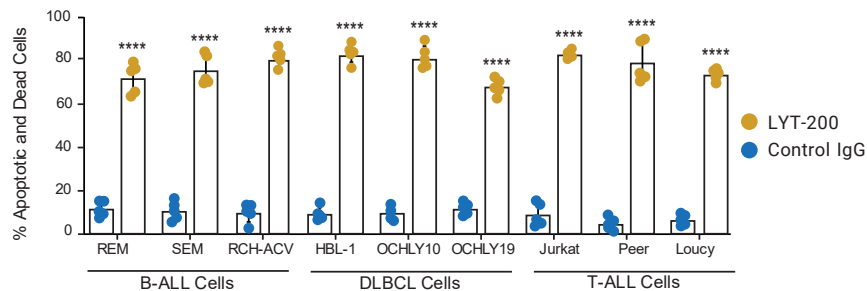
LYT-200 cause apoptosis of AML cells and is superior to anti-TIM-3 mAb



OCI-AML5  
Xenograft  
Experiment

## T-ALL, B-ALL & DLBCL MODEL

LYT-200 cause apoptosis of B-ALL, T-ALL and DLBCL cells



PDX  
Experiment  
for T-ALL<sup>2</sup>

# Phase 1b Clinical Trial in AML/MDS Ongoing

## DOSE ESCALATION TRIAL

Safety, tolerability, PK/PD, RP2D, Safety & efficacy + exploratory endpoints

### PATIENT POPULATION

AML R/R to at least one line of prior therapy with or without allogeneic system cell transplant

OR

Patients with a document-ed diagnosis of R/R, high-risk MDS after at least one line of treatment

AND

For whom no standard therapy that may provide clinical benefit is available

### DOSE FINDING (4+2 DESIGN)

Dose escalation:



Up to N=6 per cohort

If clinical benefit is observed & safety is maintained in any cohort, patients may be added to cohort(s) to further expand on safety/efficacy (Up to additional 6 patients)

In a heavily pre-treated patient population, early data demonstrated a favorable safety and tolerability profile of LYT-200 with no dose limiting toxicities;

Additional data from the study to be presented in a scientific forum in 2024

# Phase 1b Clinical Trial in Solid Tumors Ongoing

## SOLID TUMOR DOSE ESCALATION & DOSE EXPANSION TRIAL

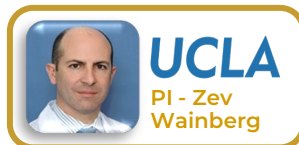
Dose Finding (CRM)  
(all comers), safety, tolerability, RP2D, PK/PD,  
exploratory

Up to 26 patients

- ✓ Completed bi-monthly, monotherapy dose escalation portion of Phase 1b/2a trial (no dose limiting toxicities)
- ✓ Completed evaluation of weekly dosing

Combination cohorts with tislelizumab (anti-PD-1 mAb) initiated in Q1 2023

## CLINICAL INVESTIGATORS



Beth Israel Deaconess Medical Center  
**Daniel Fein**



Health ONE  
**Gerald Falchook**



THE UNIVERSITY OF TEXAS  
**MD Anderson Cancer Center**  
Making Cancer History®  
**Siqing Fu**



COLUMBIA UNIVERSITY  
MEDICAL CENTER  
**Manji Gulam**



Memorial Sloan Kettering Cancer Center  
**Eric Sherman**

Other sites: Mayo, START, Sarah Cannon

Topline results expected in 2024

# LYT-200 Clinical Data to Date in AML & Solid Tumors

## AML/MDS DOSE ESCALATION COHORTS<sup>1</sup>

- Single agent LYT-200 in patients with relapse/refractory AML or high-risk MDS ongoing
- 16 patients dosed to date including 13 evaluable

- **Majority of patients have achieved stable disease<sup>2</sup>** per ELN guidelines
- At 7.5 mg/kg cohort (dose escalation continuing) a median duration of treatment was 77 days with **blast reduction observed**

## SOLID TUMOR COMBINATION COHORTS<sup>3</sup>

- Combination cohorts of LYT-200 + tislelizumab (PD-1) ongoing in head and neck cancer (HNSCC) and urothelial carcinoma (UC)
- HNSCC: 8 patients dosed to date including 4 evaluable
- UC: 3 patients dosed to date including 2 evaluable

- **Of the 4 evaluable HNSCC patients, 3 patients achieved disease control including 1 CR, 1 PR, and 1 SD and 1 PD**
- **Of the 2 evaluable patients with UC, both achieved SD** including one with near complete resolution of ascites and pleural effusion

Presented at ESMO IO '23

# PureTech Is a Respected Leader in The Boston Biotech Community: The World's #1 Biotech Hub

- >1000 biotech companies, with a critical mass of talent
- \$3.3B funding from National Institute of Health (NIH) in 2022; **top** NIH-funded state nationwide
- >100 academic institutions; MA is home to many leading universities and research institutes
- 25 hospitals with many **ranked among the best** in the U.S.

*"Massachusetts is known as the **most innovative square mile on the planet** and that's because we have such a rich density of ecosystem, for example, with over a thousand biotech companies, **18 of the top 20 biopharma**, world-class hospital system, and of course the best in class academic institutions."*

- Kendalle Burlin O'Connell, Esq., CEO & President of MassBio at the NYSE



Massachusetts  
Institute of  
Technology



HARVARD  
MEDICAL SCHOOL



MASSACHUSETTS  
GENERAL HOSPITAL



Boston  
Children's



BROAD  
INSTITUTE



Whitehead  
Institute



WYSS  
INSTITUTE

# Key Value Drivers

Multiple clinical milestones expected across Wholly Owned Programs<sup>1</sup> and Founded Entities<sup>2</sup>

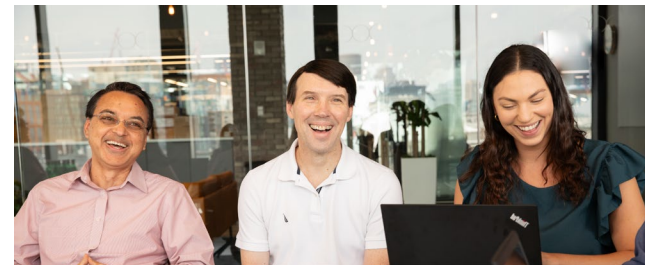
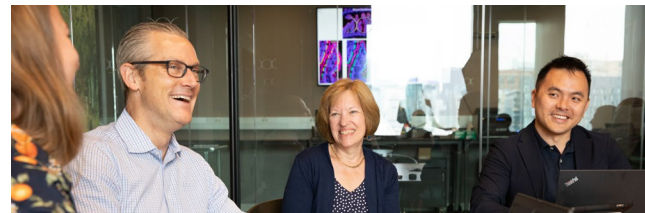
ENTITY	PURETECH OWNERSHIP	THERAPEUTIC CANDIDATE <sup>3</sup>		EXPECTED MILESTONES	
PureTech	100%	LYT-100	●	<b>Results from registration-enabling trial in IPF</b>	4Q 2024
Gallop	100%	LYT-200	●	Additional results from Phase 1b in combination w/ tislelizumab in solid tumors	2024
Karuna	Acquired by Bristol Myers Squibb for \$14B	KarXT <sup>5</sup>	●	<b>Results from Phase 3 EMERGENT-4 &amp; EMERGENT-5 trials for schizophrenia</b>	2H 2024
			●	<b>Results from Phase 3 ARISE trial for schizophrenia</b>	2H 2025
		KAR-2618	●	<b>Results from Phase 3 ADEPT-1 &amp; ADEPT-2 trials</b>	2026
			●	Initiation of Phase 1b trial for MDD	2024
Akili (NASDAQ: AKLI)	14.6%	EndeavorOTC <sup>6</sup>	●	Status update on the FDA marketing approval of EndeavorOTC as OTC treatment for adults with ADHD	2Q 2024
Vor (NASDAQ: VOR)	3.9%	Trem-Cel <sup>5</sup>	●	<b>Phase 1/2a (VBP101) clinical data update</b>	2H 2024
		VCAR33 <sup>ALLO5</sup>	●	<b>Phase 1/2 (VBP301) clinical data update</b>	2H 2024
Vedanta	41.0% <sup>3</sup>	VE303 <sup>5</sup>	●	Initiation of Phase 3 in <i>C. difficile</i>	1H 2024

✓ Indicates completed milestone

**B** Key anticipated milestones are **bolded**



# PureTech Team - Bringing Our Vision to Life





**Nasdaq Global Market & LSE  
Main Market / FTSE-indexed:**  
PRTC

Market capitalization \$815M  
(£646M) as of March 28, 2024; 1.26  
USD:GBP

**270,209,101** outstanding  
shares as of March 2024

**\$320M** estimated Consolidated  
Cash, Cash Equivalents & Short-  
Term Investments at year end 2023<sup>1</sup>

## ANALYST COVERAGE

### Leerink Partners LLC

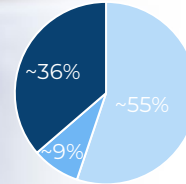
Thomas J. Smith

### Peel Hunt LLP

Miles Dixon

### Jefferies International Limited

Peter Welford



- Board & Management
- Disclosed Shareholders
- Other Shareholders

Headquartered in  
Seaport, Boston

Disclosed Shareholders as of December 31, 2023, include Invesco Asset Management, Lansdowne Partners LLP, Baillie Gifford & Co., M&G Investment Management, LTD., Vanguard Group, Patient Capital Management, Recordati S.p.A. Pharmaceutical Company.

<sup>1</sup>The preliminary selected financial results reported by the Company are unaudited, subject to adjustment, and provided as an approximation in advance of the Company's announcement of complete financial results in April 2024.

# Appendix Contents

- **Appendix A: Wholly Owned Pipeline**
  - ❑ LYT-100 Preclinical Data
  - ❑ LYT-100 Market Research
  - ❑ LYT-100 Payor Research
  - ❑ LYT-100 In the Face of Generics
  - ❑ Pirfenidone Safety Data
  - ❑ Lung Disease Prevalence
- **Appendix B: Founded Entities**
  - ❑ Vedanta
  - ❑ Akili
  - ❑ Vor
  - ❑ Sonde
  - ❑ Entrega
- **Appendix C: Supplemental Materials**
  - ❑ PureTech ESG Program
  - ❑ PureTech's Proven Expertise
  - ❑ PureTech is Executing & Delivering Results
  - ❑ Financial Highlights/Non-IFRS Measures

# Appendix A: Wholly Owned Pipeline

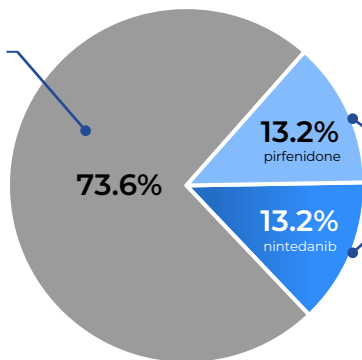
# IPF Treatment Rates are Low Due to Side Effects

OPTUM Study of 11,000 Patients with IPF<sup>1</sup>

October 2014 to July 2019

10,996 patients with IPF  
in a US health claims  
database (OPTUM)

Nearly 75% of  
patients in U.S. never  
receive antifibrotic  
therapy



Initiated pirfenidone  
or nintedanib since  
FDA approval in 2014

Over 40% of patients eventually  
discontinue antifibrotic therapy

Experienced nausea,  
diarrhea, or myalgias

21.2%

Switched to the  
other antifibrotic

10.5%

Discontinued therapy

42.8%

0% 10% 20% 30% 40% 50%

Patients, %

Mean duration  
of treatment



302

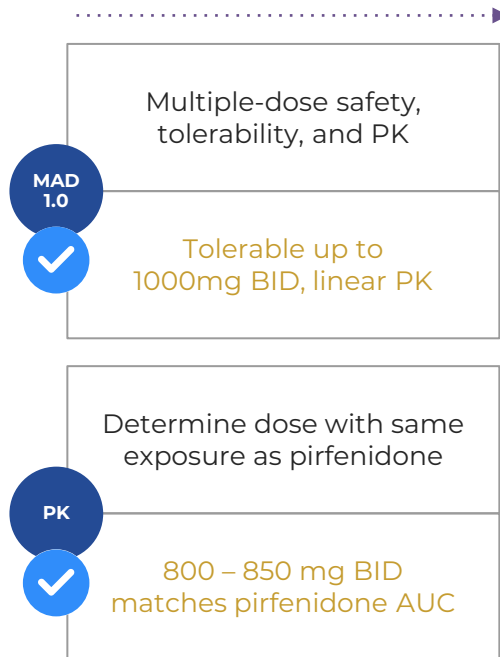
days



# LYT-100 Clinical Trials<sup>1</sup>

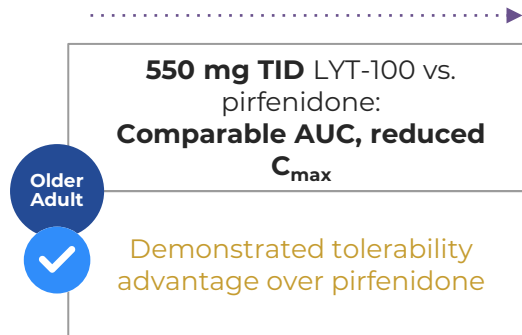
## 1. Initial PK studies

### FOUNDATIONAL PK DATA



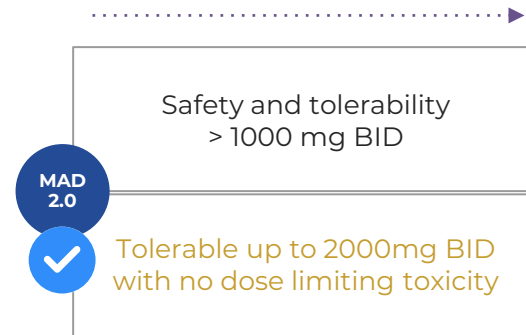
## 2. Head-to-head tolerability

### TOLERABILITY ADVANTAGE VS. PIRFENIDONE



## 3. High-dose studies

### EXPLORE FEASIBILITY OF HIGHER EXPOSURES

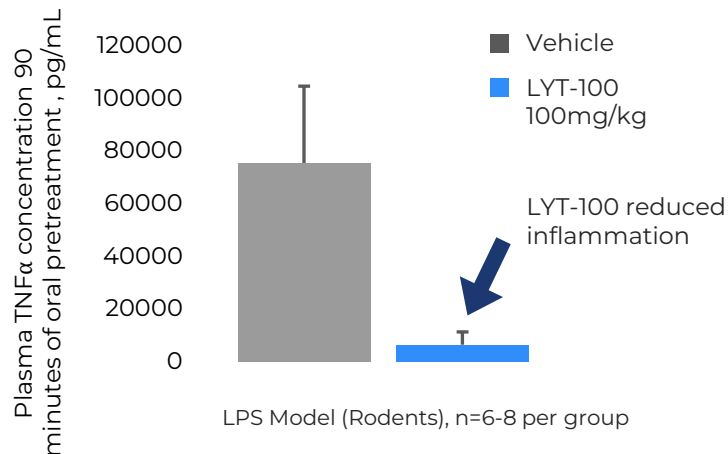




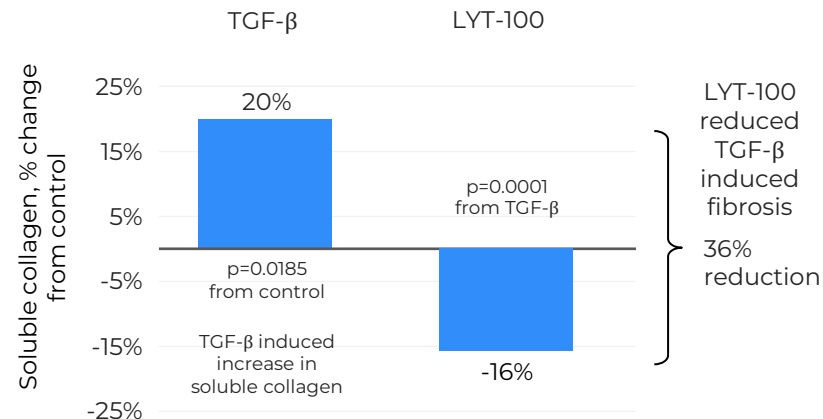
# LYT-100:

Preclinical POC demonstrates anti-inflammatory & anti-fibrotic pharmacology

## PRECLINICAL PLASMA CONCENTRATIONS OF TNF $\alpha$ WITH LYT-100 VERSUS CONTROL

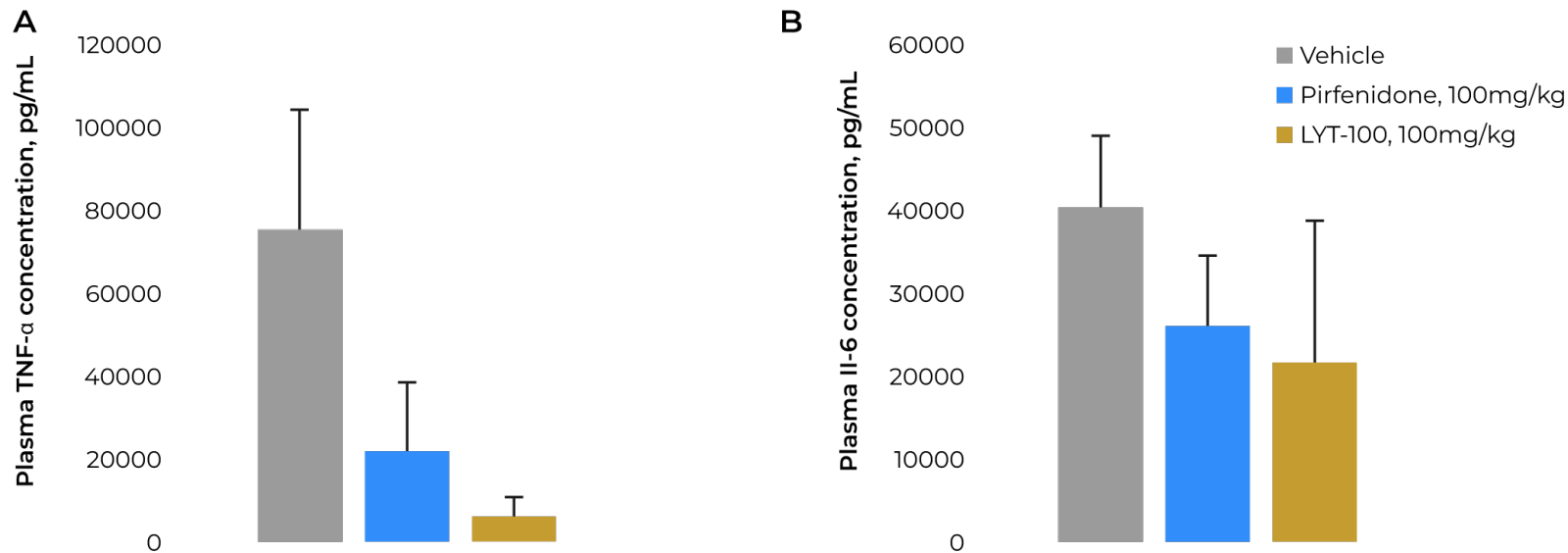


## IN VITRO REDUCTION OF TGF- $\beta$ INDUCED SOLUBLE COLLAGEN PRODUCTION (MOUSE FIBROBLASTS)



# LYT-100 Preserves Pharmacologic Effect of Pirfenidone:

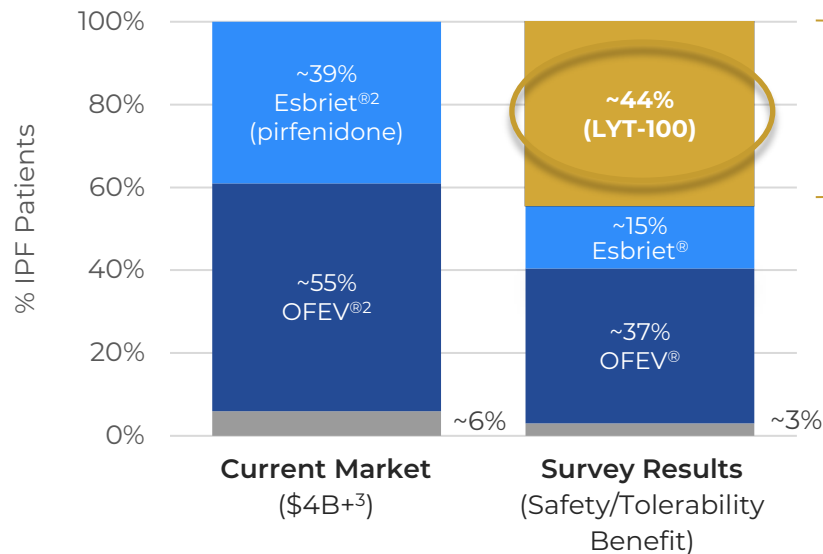
Preclinical data shows improved anti-inflammatory and anti-fibrotic activity vs pirfenidone



**Reduction in LPS-stimulated plasma concentrations of TNF-α and IL-6 by Pirfenidone or LYT-100.** Oral doses of vehicle, pirfenidone, or LYT-100 (100mg/kg) administered 60 minutes prior to LPS (30 µg/kg intravenous): TNF-α (A) and IL-6 (B) measured 90 min after LPS stimulation: N=6-8 animals per group. Data are presented as mean +/- standard deviation.

# LYT-100 Could Address Several Segments of IPF Patients

Independent research shows profile attractive to surveyed pulmonologists<sup>1</sup>



## 1 Newly diagnosed patients

Pulmonologists would prescribe LYT-100 to **~44% of newly diagnosed** patients with IPF

## 2 Patients currently on SOC treatment

Pulmonologists would **switch** some patients currently treated with SOC, particularly **ESBRIET (pirfenidone)**, to LYT-100

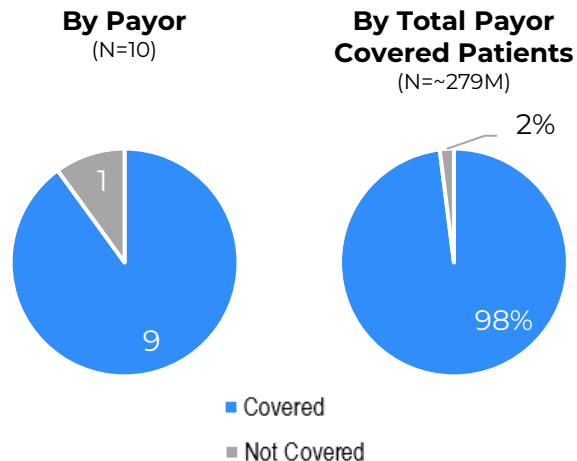
## 3 Currently untreated patients

Potentially address currently untreated patients who:  
(1) have never started treatment, AND  
(2) who started treatment but discontinued

# LYT-100 Payor Market Research

Independent research indicates payors in favor of profile<sup>1</sup>

## LYT-100 COVERAGE EXPECTATIONS<sup>2</sup>



- Payors are aware of unmet needs in IPF and recognize the clinical value of a product designed to provide improved safety/tolerability vs. pirfenidone, while maintaining efficacy
- ***Nine out of ten payors understood the LYT-100 clinical story and would consider putting it on formulary*** if successfully developed and approved, with a profile that shows superior safety/tolerability to pirfenidone<sup>3</sup>
- Payors indicated that coverage of LYT-100 could be similar to other branded IPF therapies (PA to label, Tier 3/Specialty Tier)

“...50% [tolerability improvement] would be really significant...30% would also be significant.”<sup>4</sup> – Regional Payor

Select quote from survey

# LYT-100 in The Face of Generics & Novel MOAs

## LYT-100 VS. GENERICS

- ✓ The **safety/tolerability advantage of LYT-100 remains attractive and meaningful to pulmonologists and payers** even in the face of generic competition<sup>1</sup>.
- ✓ **Only ~25% of patients in the U.S. are currently on SOC** primarily due to poor tolerability. Presence of generics is not likely to drive a dramatic increase in adoption.
- ✓ Even if all US payers require step edits through a generic antifibrotic, **~50% of IPF patients will still be eligible for LYT-100** due to the significant tolerability challenges with current standard of care.

## LYT-100 VS. NOVEL MODE OF ACTIONS (MOAS)

- ✓ There are several Phase 3 & a handful of notable Phase 2 programs evaluating novel MOAs in IPF. If successful, **nearly all of these programs are expected to be used on top of or after current SOC.**
- ✓ There is **potential for LYT-100 to be positioned as the preferred backbone antifibrotic** for future combination regimens.
- ✗ **Development of novel MOAs in IPF has proved difficult**, with many recent failures of late-phase programs. For all ongoing programs, it remains to be seen if **early Ph2 data can be replicated in Ph3 studies**

# LYT-100 in The Face of Generics & Novel MOAs (Cont'd)

Base Case: LYT-100 at equivalent dose to pirfenidone with improved safety/tolerability

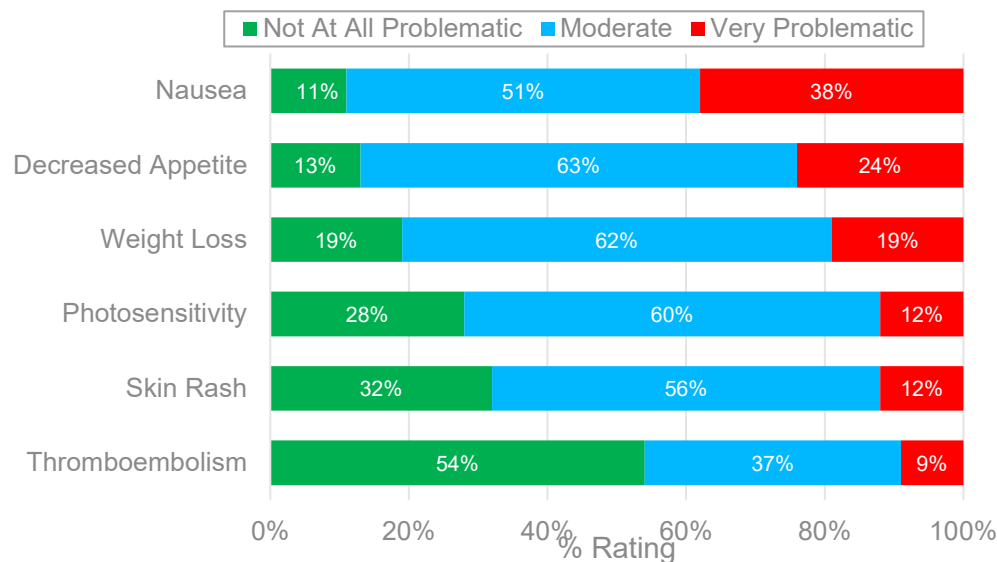
COMPETITOR	OVERVIEW	POSITIONING OF LYT-100
Generic pirfenidone and nintedanib	<ul style="list-style-type: none"><li>Both generic pirfenidone and generic nintedanib are expected to be on the market at time of LYT-100 launch<sup>1</sup></li><li>Assume all payers add generics to generic Tier<sup>2</sup>; some payers require step edits<sup>3</sup> of generics before allowing treatment with branded agents</li></ul>	<ul style="list-style-type: none"><li>Safety/tolerability advantage will enable <b>LYT-100 to complete for new patient starts in plans without step edits</b></li><li>In plans with step edits, LYT-100 will be used as second line of treatment for <b>patients who fail on generic antifibrotics</b></li><li>Even if all payers require step edits, <b>~50% of patients will be eligible for LYT-100</b></li></ul>
Reformulated pirfenidone and nintedanib	<ul style="list-style-type: none"><li>A few reformulated pirfenidone and nintedanib approaches, including inhaled and sustained release, are in early development</li></ul>	<ul style="list-style-type: none"><li>LYT-100 will offer oral systemic delivery of the medication, <b>without the AEs associated with inhaled (e.g., cough) and other reformulations of the currently approved drugs</b></li><li>None of the localized delivery candidates have demonstrated the same evidence of efficacy as systemic therapies</li></ul>
Novel Mechanisms	<ul style="list-style-type: none"><li>Nearly all new mechanisms are being studied on top of/or after the standard of care (currently pirfenidone &amp; nintedanib)</li></ul>	<ul style="list-style-type: none"><li><b>Potential for LYT-100 to be the backbone standard of care for future combination regimens</b></li><li>Pirfenidone and nintedanib remain key competitors for LYT-100</li></ul>

# ESBRIET (pirfenidone) and OFEV (nintedanib):

Both have significant tolerability issues, with GI side effects being the most problematic

- In a market research survey of 100 pulmonologists, respondents reported that, aside from efficacy, **safety/tolerability is the most important consideration** when treating patients with IPF
- However, **>70% of pulmonologists expressed they are “not at all satisfied” or only “moderately satisfied” with the safety/tolerability profile of SOC** today
- In particular, **GI side effects were noted as the most concerning/problematic** adverse events

## HOW PROBLEMATIC ARE AEs ASSOCIATED WITH IPF SOC? (N=100)

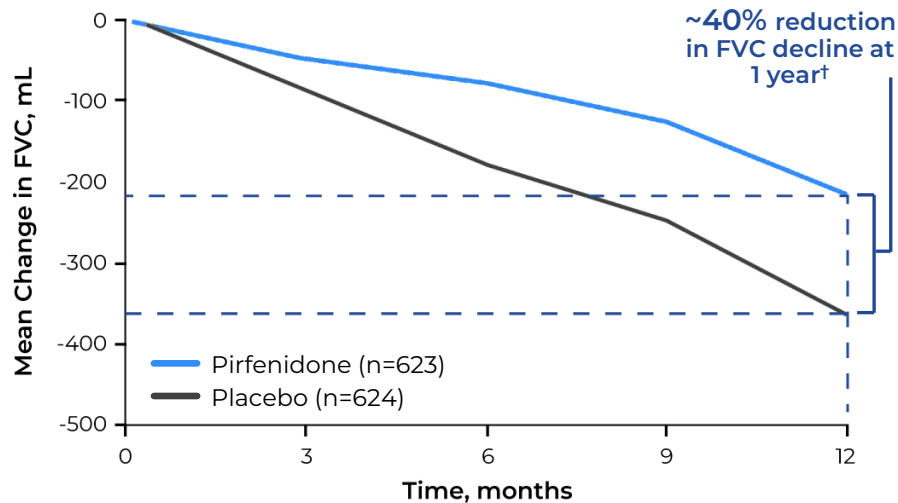




# Pirfenidone:

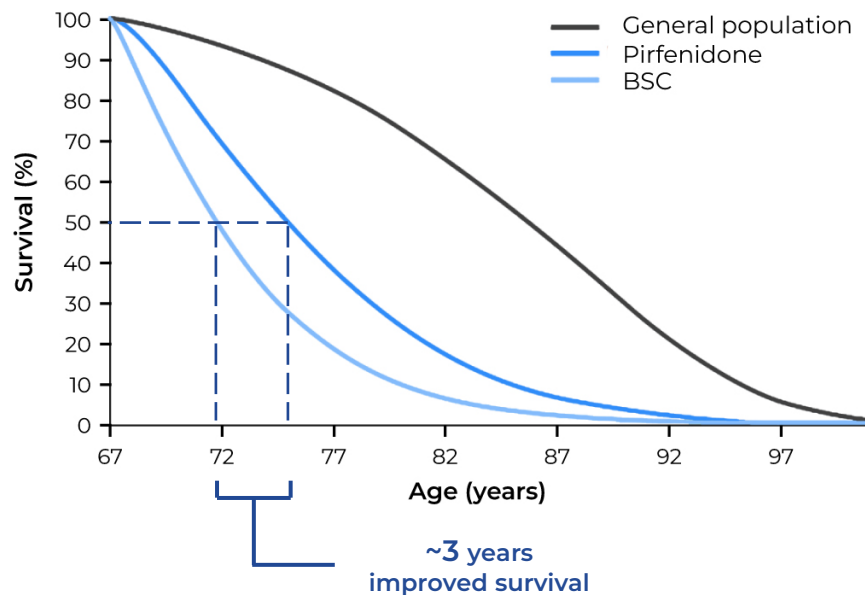
A clinically validated treatment for IPF with beneficial effects on FVC and survival

POOLED MEAN CHANGE FROM BASELINE IN FVC  
FROM THE **ASCEND** AND **CAPACITY** TRIALS<sup>1\*</sup>



\*FVC assessed at weeks 12, 24, 36, and 48 in CAPACITY and weeks 13, 26, 39, and 52 in ASCEND. †Mean change from baseline in FVC.

~3 YEAR IMPROVEMENT IN SURVIVAL WITH  
PIRFENIDONE VS BEST SUPPORTIVE CARE IN A  
MATCHED POPULATION FROM THE UK<sup>2</sup>



# Design & Tolerability Findings of Pirfenidone Studies

Pirfenidone discontinuations often related to gastrointestinal (GI) adverse events (AEs)<sup>1</sup>

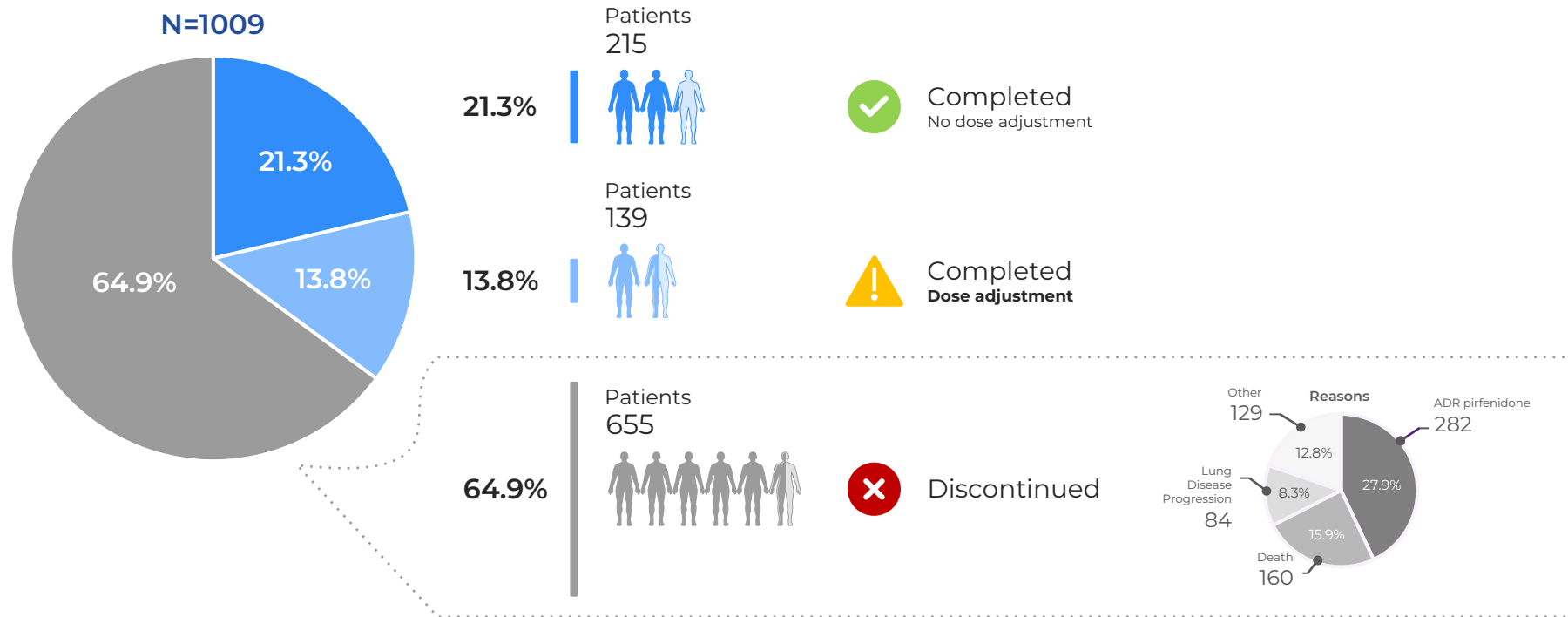
Pirfenidone GI AEs:

- Require titration in IPF and other studies
- More common in women<sup>2</sup>

	PIRFENIDONE FOOD EFFECT/ANTACID STUDY <sup>2</sup>		PIRFENIDONE FOOD EFFECT AND BIOEQUIVALENCE STUDY <sup>3</sup>		PIRFENIDONE PHASE 3 STUDIES <sup>1</sup>		
Design	801mg single-dose in healthy older adults, 44% women		801mg single-dose in healthy adults, 36% women		2403mg per day, IPF patients 26% women		
Most common AEs	Most common AEs	Pirfenidone N=16	Most common AEs	Pirfenidone N=44	Most common GI AEs <sup>Δ</sup>	Pirfenidone N=623	Placebo N=624
	Nausea	43.8%	Nausea	29.5%	Nausea	36%	16%
	Dizziness	37.5%	Dizziness	18.2%	Rash	30%	10%
	AEs more frequent in the fasted state AE rate higher in women		Headache	9.1%	Ab. pain	24%	15%
			Constipation	9.1%	Diarrhea	26%	20%
			Vomiting	4.5%	Headache	22%	19%
			Dyspepsia	4.5%	Dyspepsia	19%	7%
			AEs more frequent in the fasted state		Dizziness	18%	11%
					Vomiting	13%	6%
					Anorexia	13%	5%

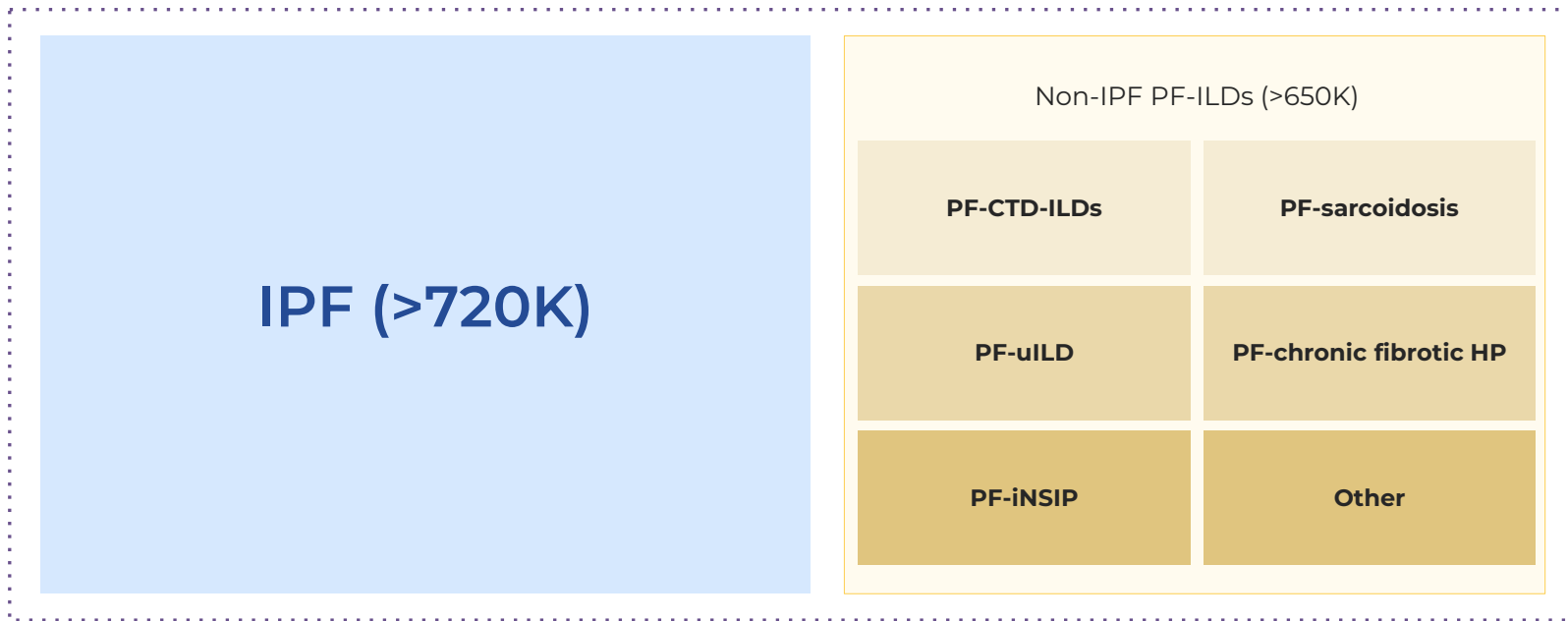
<sup>Δ</sup>Other most common AEs observed in the Phase 3 studies (pirfenidone vs. placebo) include upper resp. infect (27% vs. 25%), fatigue (26% vs. 19%), GERD (11% vs. 7%), sinusitis (11% vs. 10%), insomnia (10% vs. 7%), weight decrease (10% vs. 5%), arthralgia (10% vs. 7%)

# Prospective Registry Found Only 21% of Patients Who Started Pirfenidone Remained on Full Dose After 2 Years



# Enduring High Unmet Need in Interstitial Lung Diseases Including IPF

Progressive Fibrosing ILDs (PF-ILDs) are estimated to affect >1.3M patients in the 16 Major markets<sup>1,2</sup>



Major potential to improve care in IPF & address other interstitial lung diseases

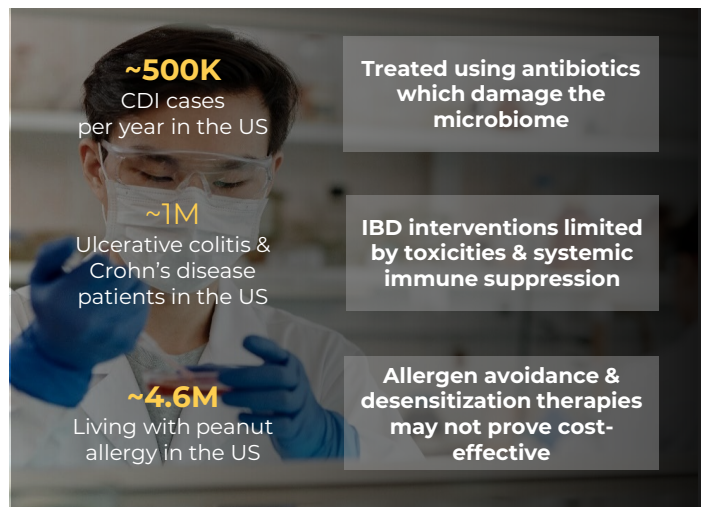
# Appendix B: Founded Entities

# Vedanta

Developing a new class of drugs to modulate the human microbiome

## INNOVATION

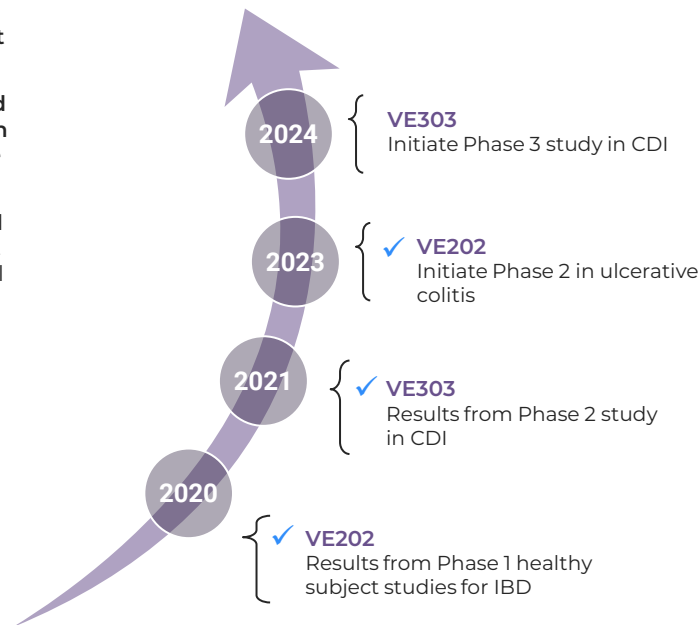
Rationally-defined consortia of **gut bacteria**; manufactured from **pure cell banks** to produce drug product of **known bacterial isolates**; **orally administered** to modulate microbial communities and immune responses



## VALIDATION

- **Four clinical-stage programs in development**
- VE303 (*C. difficile*) demonstrated **accelerated gut microbiota restoration** after antibiotics in a **Phase 1a/1b study**
- VE202 (IBD) demonstrated durable & dose dependent colonization in Phase 1 trial in healthy volunteers
- VE416 (food allergy) being evaluated in Phase 1/2 study
- Strong IP portfolio
- \$71.1M in total Series C

## UPCOMING MILESTONES<sup>2</sup> & VALUE REALIZATION



VE303 & VE202 received Fast Track designation from the U.S. FDA

# Akili

First game-based digital therapeutic cleared by the FDA for ADHD

(PRTC Ownership: 14.6%<sup>1</sup>)

## INNOVATION

~6.4M **pediatric ADHD patients in the US**

Treatment of many neuropsychiatric disorders is only partially served, or not served at all, by current medications or in-person behavioral therapy

**Engaged with leading experts who had been studying the effects of video games on cognition**



- ✓ In-licensed from University of California, San Francisco the intellectual property invented by Adam Gazzaley, MD, PhD
- ✓ Oversaw initial product development & design

## VALIDATION

Helped build top development & commercial team & raise funds

- ✓ **Planned & completed initial pilot & POC studies**



## VALUE REALIZATION

**FDA Clearance & European CE Mark**

- ✓ **FDA cleared & granted European marketing authorization** for pediatric patients age 8-12 years old with primarily inattentive or combined-type ADHD
- ✓ **EndeavorRx®<sup>2</sup> (AKL-T01)** showed statistically significant improvement compared to active control ( $p=0.006$ ) on T.O.V.A.® in pivotal study; recently showed statistically significant improvement in ADHD when used with & without stimulants
- ✓ **Achieved primary endpoint**, improving cognitive impairments in MDD
- ✓ Development & commercialization partnership with **Shionogi** in Japan & Taiwan (\$20M up front; milestones up to \$105M plus royalties)

## UPCOMING MILESTONES

- Plans to pursue regulatory approval for OTC labeling of its treatment products
- Data submission to the FDA to convert EndeavorRx to OTC in 2024
- Shionogi pivotal trial data in 6-17 year old children with ADHD in Japan expected in 2024

EndeavorOTC for adults 18 y.o. and older with ADHD now available without a prescription nationwide



# Vor

Selectively protecting healthy cells from targeted cancer therapies

## INNOVATION

~42.5K **new diagnoses of AML patients each year in the US, Europe & Japan**

Prognosis for relapsed & refractory blood-borne malignancies is very poor

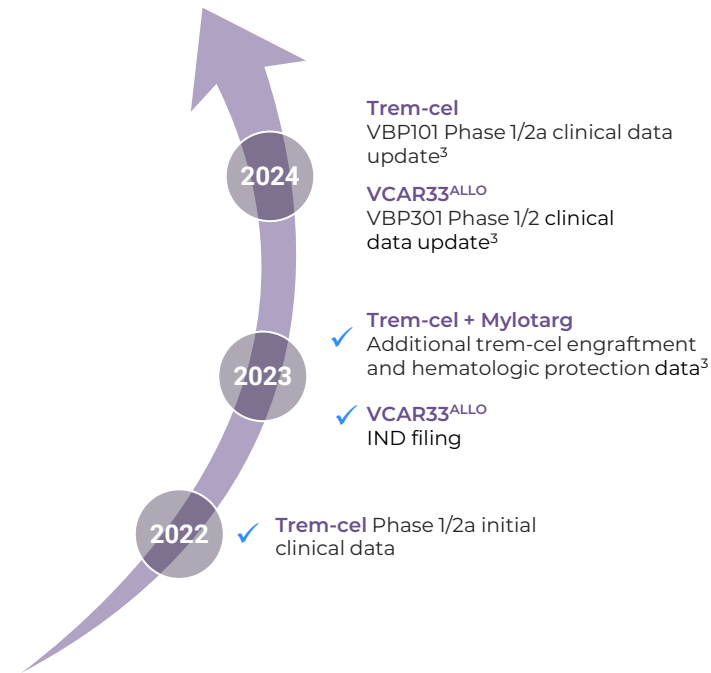
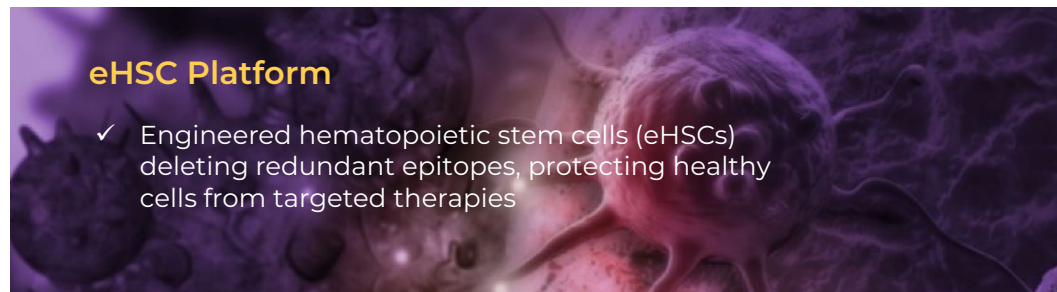
Median 5 year survival rate for patients with AML is <30%, but there are significant differences in prognosis depending on several factors, including the age of the patient at diagnosis

## VALIDATION

- **Ex vivo & mouse proof-of-concept studies** led by Siddhartha Mukherjee, MD, PhD; Also published in *PNAS*
- Optimize targeted therapies including **ADCs, T cell engager / bispecific antibodies, conventional mAbs & CAR-T cells**
- May lead to **limited on-target toxicity & durable antitumor activity**
- Conducting ongoing discovery efforts for non-myeloid malignancies
- **Announced \$110M Series B financing in July 2020**
- **Completed \$176.9M IPO in February 2021**
- **Completed \$115.8M follow-on offering in December 2022**

(PRTC Ownership: 3.9%<sup>1</sup>)

## UPCOMING MILESTONES<sup>2</sup> & VALUE REALIZATION



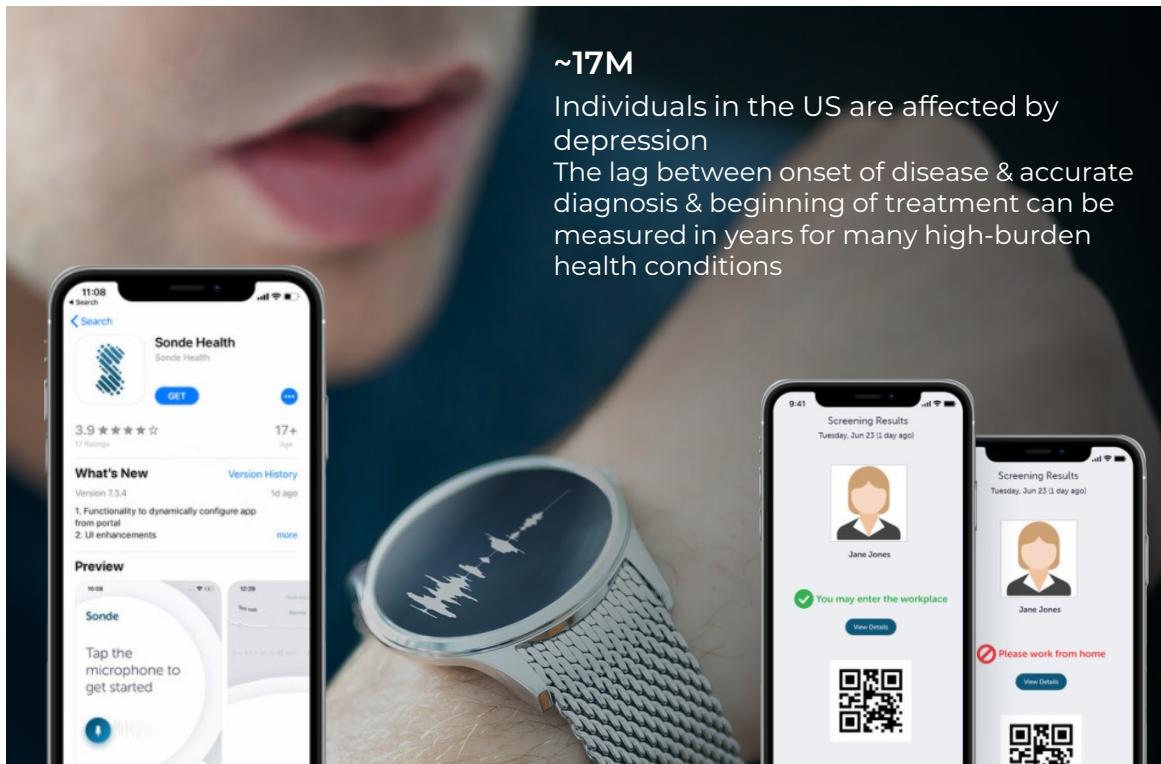
# Sonde

Voice-based artificial intelligence (AI) platform with the potential to transform how we monitor health

(PRTC Ownership: 35.2%<sup>1</sup>)

~17M

Individuals in the US are affected by depression  
The lag between onset of disease & accurate diagnosis & beginning of treatment can be measured in years for many high-burden health conditions



## SONDE

Developing proprietary technology to sense & analyze subtle changes in the voice to create a range of persistent brain, muscle & respiratory health measurements that provide a more complete picture of health in just seconds

## KEY HIGHLIGHTS

- Technology has demonstrated the **potential to screen & monitor for disease** in individuals from brief samples of speech
- **Ongoing collaborations** with multiple US & ex-US hospitals, clinics & academic medical centers
- Partnership with Qualcomm Technologies for vocal biomarker technology
- Collected **voice data** from over 80,000 subjects as part of ongoing validation of platform
- Expanded development of its proprietary technology into respiratory & other **health & wellness conditions, including mental health**

# Entrega

Engineering hydrogels to enable the oral administration of peptide therapeutics (e.g., GLP-1 agonists)

(PRTC Ownership: 73.8%<sup>1</sup>)



Entrega is focused on the oral administration of biologics, vaccines and other drugs that are otherwise not efficiently absorbed when taken orally.

The vast majority of biologic drugs, including peptides, proteins and other macromolecules are currently administered by injection, which can present challenges for healthcare administration and compliance with treatment regimes.

## MILESTONES ACHIEVED

- To validate its technology, Entrega generated preclinical proof-of-concept data demonstrating administration of therapeutic peptides into the bloodstream of large animals.

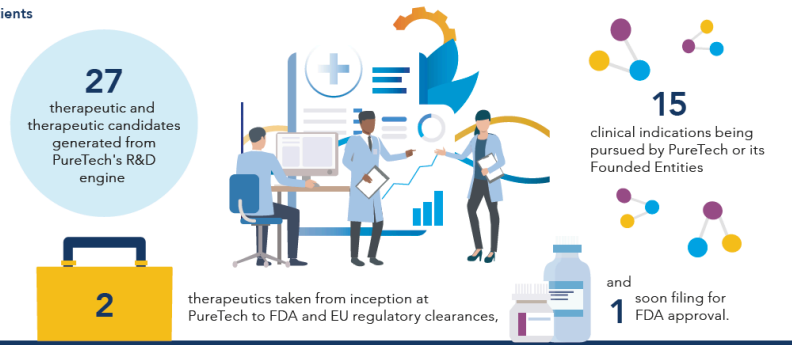
# Appendix C: Supplemental Materials

# PureTech ESG Program

3 areas of focus: patients, people & planet

Named as one of the 2022 top-rated ESG companies by Sustainalytics<sup>3</sup> & ranked in the top 3 percent of pharmaceutical companies

## Patients



## People






## Planet



# PureTech's Proven Expertise

We give life to classes of medicine with proven efficacy by addressing key limitations

	PROGRAM	VALIDATED EFFICACY	PROBLEM	PURETECH INSIGHT/IP
<b>KarXT</b> , invented by PureTech Team	<b>Karuna's KarXT</b> for schizophrenia & psychosis in Alzheimer's disease	Xanomeline is highly effective in reducing psychosis	Xanomeline has GI tolerability issues	Pairing xanomeline with peripherally-restricted muscarinic antagonist improved AE profile and unlocked 1st new class in 50+ years 
Wholly Owned Pipeline includes multiple clinical programs building on PureTech's expertise & platforms. Certain CNS programs & relevant Glyph intellectual property to be advanced via Seaport Therapeutics	<b>LYT-100</b> for inflammation and fibrosis, including IPF	Pirfenidone extends life in patients with IPF by an average of ~2.5 years <sup>1</sup>	GI tolerability issues negatively impact patient compliance & efficacy	Retain clinically-validated activity of pirfenidone w/ improved tolerability & potential for improved efficacy 
	<b>SPT-300</b> for neuropsychiatric & rare CNS conditions	Allopregnanolone has demonstrated efficacy in mental health conditions	Marketed allopregnanolone requires 60-hr IV infusion & chemical analogs may have different pharmacological effects than endogenous allopregnanolone	Using proprietary Glyph technology, achieved blood levels of allopregnanolone at/above those associated w/ therapeutic effect & demonstrated exposure-dependent target engagement w/ GABA <sub>A</sub> receptors <sup>2</sup> . Approach may have advantages vs. oral chemical analogs 

# PureTech is Executing & Delivering Results

## REGULATORY

FDA Clearance & **EndeavorRx<sup>®1</sup>** (AKL-T01)  
European CE Mark **Plenity<sup>®2</sup>** (Gelesis100)

## R&D & DATA PRESENTATIONS

- ✓ **Phase 2 & Phase 3 results** for Karuna's KarXT
- ✓ **Phase 1 results** for Vedanta's VE303 & VE202
- ✓ **Phase 2 results** for Vedanta's VE303
- ✓ Pivotal data for AKL-T01 ADHD study published in **Lancet Digital Health**
- ✓ Results for Akili's AKL-T01 in children with ADHD alone or as an adjunct to stimulants
- ✓ Akili's AKL-T03 data on MDD presented at ACNP
- ✓ Vedanta's IO candidate selected & being **advanced with BMS**
- ✓ PureTech programs published in **Nature & Nature Neuroscience**
- ✓ POC study for Vor published in **PNAS**
- ✓ Presentations on PureTech's LYT-200 at **ESMO & ASH & SITC & AACR**
- ✓ Presentations on PureTech's LYT-100 at **CHEST & ATS & ERS**
- ✓ PureTech's LYT-100 MAD study published in **Clinical Pharmacology in Drug Development**

## PARTNERSHIPS

- ✓ **Akili's partnership with Shionogi**  
*Up to \$20M in upfront payments with the potential to receive milestone payments for Japan & Taiwan commercialization of up to an additional \$105M in addition to royalties on product sales*
- ✓ **PureTech's partnership with Imbrium Therapeutics to advance LYT-503/IMB-150**  
*\$6.5 million in upfront payment and eligible to receive up to \$53 million in additional development milestone payments for this program as well as royalties on product sales*
- ✓ **PureTech's royalty agreement with Royalty Pharma for up to \$500M**  
*\$100 million up front and up to \$400 million in additional payments for PureTech's 3% royalty in Karuna's KarXT. After \$2 billion sales threshold, PureTech to retain 67% of royalty payments*

## FINANCINGS

- ✓ **Seaport's \$100M Series A financing**  
*Key investors include ARCH Venture Partners, Sofinnova Investments, Third Rock Ventures*
- ✓ **Karuna's \$124M Series A+B financings; \$103M IPO; \$14B acquisition by BMS**  
*Key investors include ARCH Venture Partners, Fidelity, Eventide, Pivotal bioVenture Partners, Partner Fund*
- ✓ **Vor's \$153M Series A+B financings; \$203.4M IPO**  
*Key investors include RA Capital Management, Fidelity Management & Research Company, Pagliuca Family Office, Alexandria Venture Investments, SAM Ventures, Johnson & Johnson Innovation—JJDC, Inc. (JJDC), Osage University Partners, Novartis Institutes for BioMedical Research*
- ✓ **Vedanta's \$71M Series C financing; \$68M Series D financing**  
*Key investors include Bill & Melinda Gates Foundation, Bristol-Myers Squibb, Rock Springs Capital, affiliates of Magnetar Capital*
- ✓ **Sonde's \$16M Series A financing**  
*Key investors include M Ventures, MP Healthcare Venture Management, Neoteny 4*
- ✓ **Akili's Nasdaq debut ('AKLI') via SPAC merger**  
*Transaction generated more than \$163M in gross proceeds, which will be used to fund the commercial launch of EndeavorRx<sup>®</sup>, its potential expansion into additional ADHD patient populations, and will also support the advancement of the company's late-stage pipeline*
- ✓ **Vedanta's \$106.5M financing**  
*Syndicate led by new investors AXA IM Alts and The AMR Action Fund along with existing investors Bill & Melinda Gates Foundation, Skyviews Life Science, and others*



# Financial Highlights

Estimated  
December 31, 2023  
\$ millions

## Cash Flow and Liquidity<sup>1</sup>

<b>Consolidated Cash, Cash Equivalents, and Short-Term Investments</b>	<b>320.0</b>
Less: Cash and Cash Equivalents held at non-wholly-owned subsidiaries	(1.2)
<b>PureTech Level Cash, Cash Equivalents, and Short-Term Investments<sup>2</sup></b>	<b>318.8</b>

# Non-IFRS Measures

## Reported Performance

Reported performance considers all factors that have affected the results of our business, as reflected in our consolidated financial statements.

## Core Performance

Core performance measures are alternative performance measures (APM) which are adjusted and non-IFRS measures. These measures cannot be derived directly from our Consolidated Financial Statements. We believe that these non-IFRS performance measures, when provided in combination with reported performance, will provide investors, analysts and other stakeholders with helpful complementary information to better understand our financial performance and our financial position from period to period. The measures are also used by management for planning and reporting purposes. The measures are not substitutable for IFRS financial information and should not be considered superior to financial information presented in accordance with IFRS.

## Cash flow and liquidity

### PureTech Level Cash and cash equivalents

**Measure type:** Core performance.

**Definition:** Cash and cash equivalents held at PureTech Health plc and only wholly-owned subsidiaries (PureTech LYT, PureTech LYT-100, Alivio Therapeutics, Inc., PureTech Management, Inc., PureTech Health LLC, PureTech Securities Corp, PureTech Securities II Corp)

**Why we use it:** PureTech Level Cash and cash equivalents is a measure that provides valuable additional information with respect to cash and cash equivalents available to fund the Wholly Owned Programs and make certain investments in Founded Entities