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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 <a href="https://investors.puretechhealth.com/financials-filings/reports">https://investors.puretechhealth.com/financials-filings/reports</a> 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.

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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



Daphne Zohar
Founder & Chief Executive
Officer

Built team, scientific network & pipeline; Recognized as a top leader in biotech by EY, Fierce Pharma, Scientific American, BioWorld; BIO Board Member & Strategy & Policy Committee cochair



Bharatt Chowrira, PhD, JD
President

Former COO at Auspex
(acq. by Teva \$3.5B), COO
at Nektar, GC at SIRNA
(acq. by Merck \$1.1B), VP at
Merck & Co.; 20+ years of
leadership roles in
multiple biotechs;
Board Member



Eric Elenko, PhD

Co-founder & Chief
Innovation
& Strategy Officer

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 Arix Bioscience (acq. by RTW Biotech \$250M); Previously at Touchstone Innovations, Bird & Bird

### World Class Board of Directors Provides Strong Governance

Our board contributed to regulatory approvals of approximately 20 drugs, led multi-billiondollar 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)



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



John LaMattina. PhD Board

Former President of Pfizer Global R&D. Forbes Contributor



Kiran Mazumdar-Shaw Board

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



Robert Langer, ScD Board

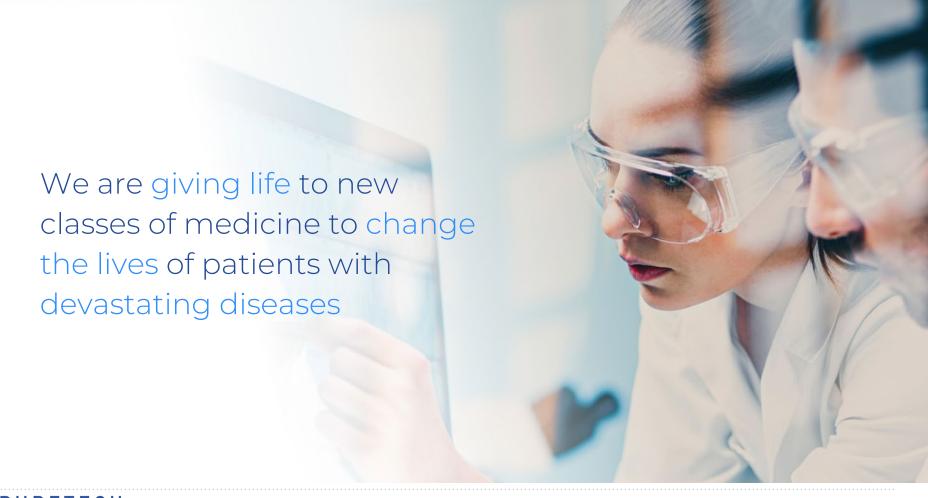
MIT, Award winning materials science pioneer, Former member of the US FDA's SCIENCE Board. Co-founder of multiple biotech companies incl. Moderna & PureTech



Sharon Barber-Lui Board

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





### 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

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







### The PureTech-Pioneered Hub-and-Spoke Model

Robust pipeline of new medicines poised for tremendous growth



SEAP(\*)R

100% Equity











PureTech's experienced R&D team & extensive network





















### CAPITAL EFFICIENT MODEL

- \$320M estimated Consolidated Cash, Cash Equivalents & Short-Term Investments<sup>5</sup>; cash figures & capital return strategy to be updated following the closing of the Karuna/BMS transaction. Estimated gross proceeds from PureTech's KRXT equity holdings is ~\$294M.
- Operational runway into 2027
- PureTech has not needed to raise capital in ~6 years
- \$3.8B raised by Founded Entities since July 2018, of which 96% was from 3<sup>rd</sup> parties



### 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

**~3.2**M 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

**Trospium 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 is expected to be acquired by Bristol Myers Squibb for \$14B



### Generating Value for Patients and Shareholders

KarXT Case Study Part 2

\$18.5M

Total PRTC spend<sup>1</sup>



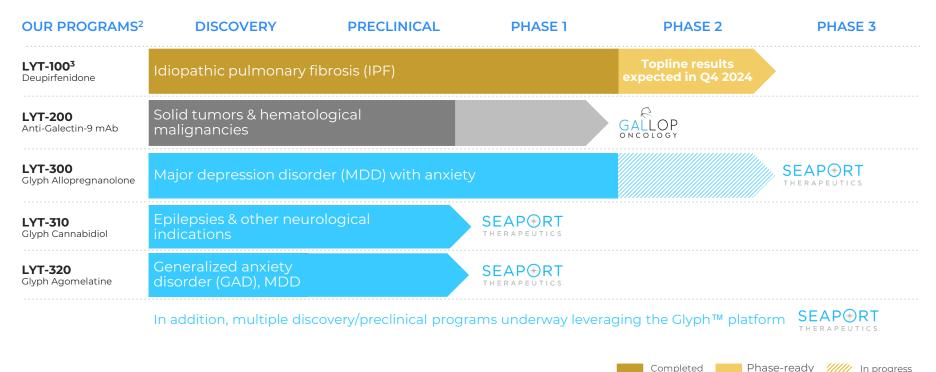
Value of remaining Karuna
holdings plus potential milestones
from royalty deal, excluding
milestone payments from Karuna
and 2% royalty on annual sales
above \$2B

Cash generated to date, through equity sales and \$100M upfront payment from the Royalty Pharma transaction



### Wholly Owned Pipeline<sup>1</sup>

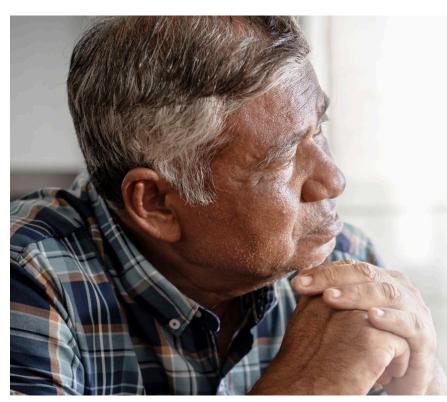
Certain assets to be advanced by new Founded Entities





### LYT-100 for Idiopathic Pulmonary Fibrosis (IPF)

ORPHAN DESIGNATION: ~120,000 patients in the US, ~110,000 in the EU51



#### **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>



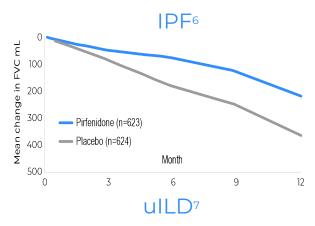
### Pirfenidone:

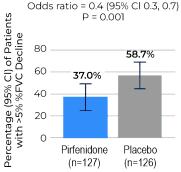
Clinically validated anti-fibrotic & anti-inflammatory

#### 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 years¹
- Over a dozen late-stage & real-world efficacy studies demonstrate efficacy in IPF<sup>2</sup>
- <u>BUT</u> 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>





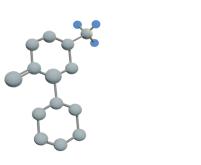


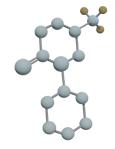
### LYT-100:

#### A potential game changer for IPF patients

#### PIRFENIDONE

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







Time

#### 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 (%)
Gastrointestinal	8 (17.4%)	16 (34.0%)
Nausea	7 (15.2%)	14 (29.8%)
Vomiting	2 (4.3%)	3 (6.4%)
Abdominal Pain/Distension	1 (2.2%)	3 (6.4%)
Nervous System	8 (17.4%)	15 (31.9%)
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** 

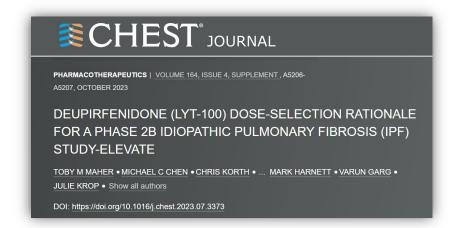
**Higher dose (824mg TID LYT-100)** in the 2<sup>nd</sup> Multiple Ascending Dose (N=24): well-tolerated with no additional incident<sup>4</sup>

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)¹
- 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



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 pfILD (NEJM)



TOBY MAHER, MD, PHD Professor & Director of ILD at Keck School of Medicine. USC: PhII trial of pirfenidone in uILDs (Lancet RM)



Chair. Department of Medicine. Cedars-Sinai: results of two latestage studies evaluating the effect of pirfenidone in patients w/ IPF (Lancet)

PAUL NOBLE, MD



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

### 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 registrationenabling 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

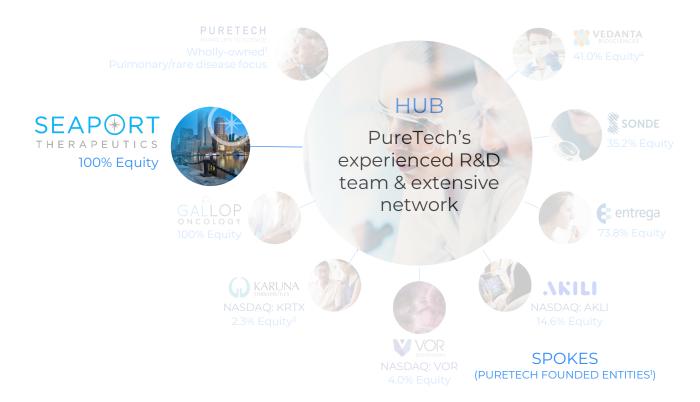
Pirfenidone mitigates radiationinduced lung fibrosis in preclinical study<sup>4</sup>

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

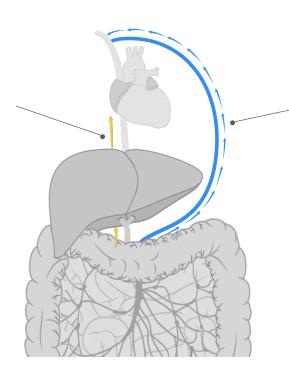


### Glyph™: Leveraging the Lymphatic System to Unlock New Medicines

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

#### CONVENTIONAL

Oral drugs with high firstpass metabolism can have low bioavailability & hepatotoxicity

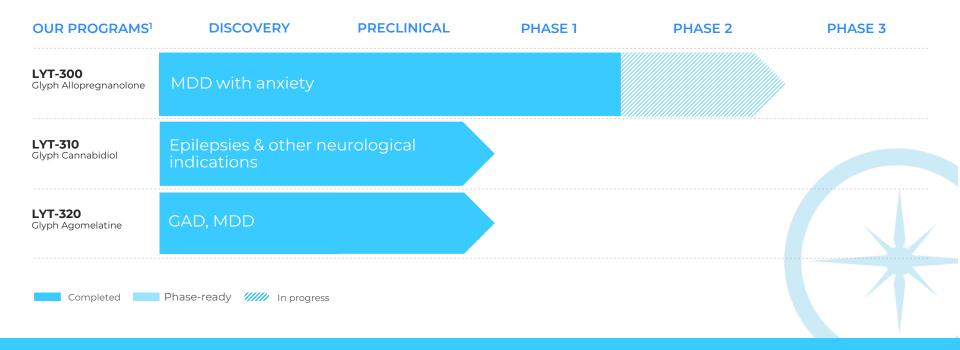


#### GLYPH™

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

### Seaport Therapeutics: First & Best-in-Class CNS Medicines



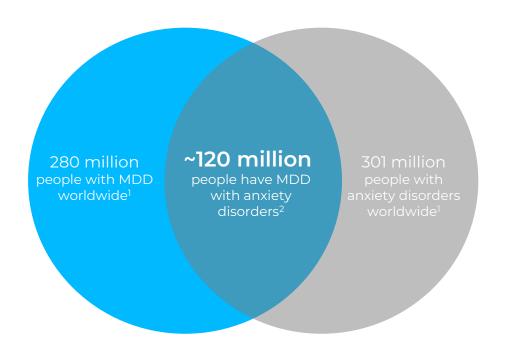
In addition, multiple discovery/preclinical programs underway leveraging the Glyph™ platform





### LYT-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

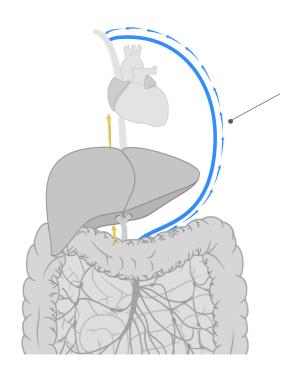
### LYT-300 (Glyph Allopregnanolone)

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
- LYT-300 retains the activity & potency of endogenous allopregnanolone in an oral form

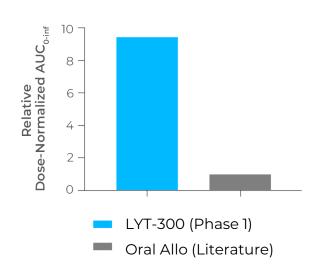
### LYT-300 (Glyph Allopregnanolone)



#### GLYPH LYT-300 PHASE 1

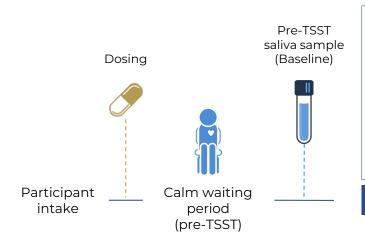
- ✓ Bioavailability >9X vs. oral allopregnanolone<sup>1</sup>
- Generally well-tolerated, AEs generally mild and transient
  - Most common AE was somnolence (on-target effect of GABA<sub>A</sub>)
- No treatment-related severe or serious AEs
- No sudden loss of consciousness observed

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



### Phase 2a Trial Design in Acute Anxiety

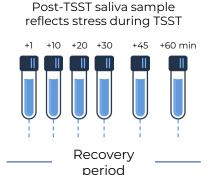
Randomized, placebo-controlled trial in the Trier Social Stress Test (TSST)



- 1. Anticipation: prepare a speech
- 2. Public speaking to a panel
- 3. Live math test to a panel



**Trier Social Stress Test** 



#### PRIMARY AIM:

To characterize pharmacology of LYT-300 for potential anxiety indications

#### PRIMARY ENDPOINT:

Reduction in salivary cortisol, a stress hormone

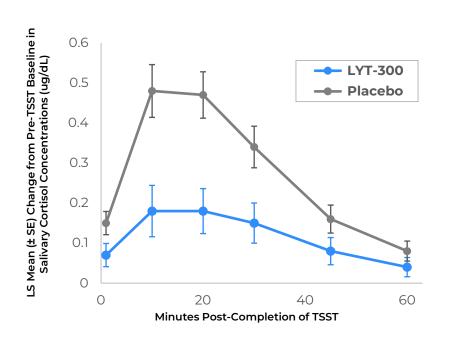
#### TRIAL DESIGN:

N=80 randomized to LYT-300 or placebo



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

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



#### **POSITIVE DATA**

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

#### **VALIDATION**

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

### LYT-320 (Glyph Agomelatine)

#### For anxiety & mood disorders



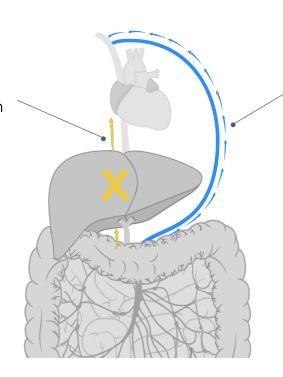
- Clinically validated and approved for MDD in the EU and MDD & GAD in Australia
- Oifferentiated mechanism of action
- Consistent and statistically significant against placebo in GAD (4/4 studies)
- Superior efficacy and tolerability vs. standard-of-care<sup>1,2</sup>
- ...BUT it has low oral bioavailability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

#### LYT-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>

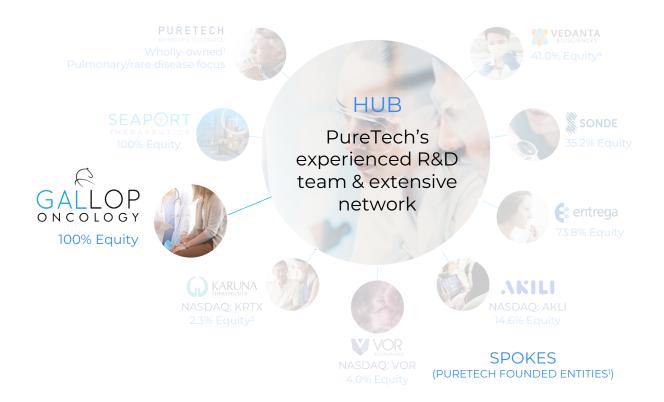


#### LYT-320

- LYT-320 bioavailability is >10X of agomelatine based on nonhuman primate PK
- LYT-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

#### Apoptosis of **Promotes expansion** effector T helper 1 of MDSCs (TH1), TH17 and FOUNDATIONAL BIOLOGY CD8+ T cells Galectin-9 modulates multiple pathways CD8 of cancer immunosuppression, including **MDSCs** those modulated by PD-1 & TIM-3 Secretion of Galectin-9 Galectin-9 Differentiation and expansion of Trea cells Treg Recruitment and differentiation of immunosuppressive M2 (M2-type) macrophages





### Gallop Oncology:

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



#### LIFE ALTERING & PROGRESSIVE

#### Solid Tumors

- ~ 82,000 new cases of bladder cancer diagnosed each year<sup>1</sup>;
   ~90% are urothelial carcinoma<sup>2</sup>
- ~66,000 new cases of head and neck cancers diagnosed each year<sup>3</sup>; ~10% metastatic disease at diagnosis & additional 20-30% will develop metastases<sup>4</sup>

#### Hematological Malignancies

**~60,000 new cases of leukemia** diagnosed each year,<sup>5</sup> including ~20,000 in acute myeloid leukemia (AML)<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>





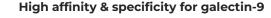
### LYT-200

#### Multiple lines of preclinical data supporting therapeutic potential

### SINGLE AGENT ACTIVITY IN B16F10 MELANOMA MODEL

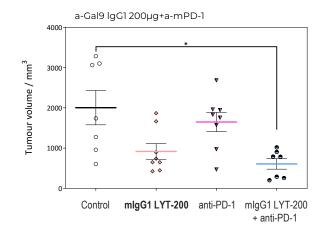
# T CELL ACTIVATION WITH LYT-200 IN PATIENT-DERIVED ORGANOID<sup>1</sup> MODEL



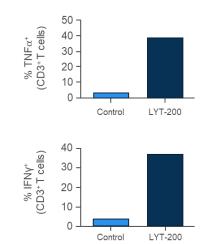


#### Robust activity in preclinical studies:

- 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



n = 8 / arm; \*p < 0.001; \*\*p < 0.05; NS = not significant

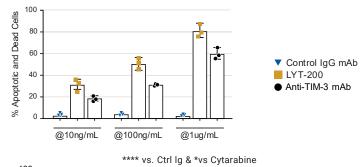


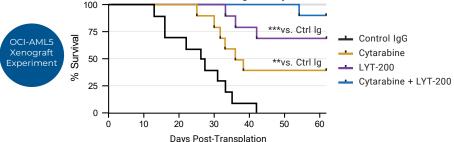
### LYT-200

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

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

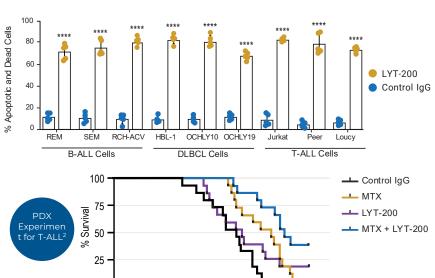
LYT-200 cause apoptosis of AML cells and is superior to anti-TIM-3 mAb





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

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



40

Davs Post-Transplation





### 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**





















Other sites: Mayo, START, Sarah Cannon

Topline results expected in 2024

### 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



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

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

#### SOLID TUMOR COMBINATION COHORTS<sup>1</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

#### AML/MDS DOSE ESCALATION COHORTS<sup>2</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>3</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



# 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

















Source: 2022 Industry Snapshot Report by MassBio.

### Why Invest in Biotech?

### Source: Stifel 2023 Healthcare Report

		Rus
1. HISTORY:	stocks up 20x over past 30 years. Following past	1400% 1200% 1000%
2. VALUATIONS:	Valuations are highly attractive now with 70% discount on the entire biotech sector since 2021 highs	600% 400% 200% 0%
3. MACRO:	Macro picture shifting to favor biotech. In past, NBI has weathered the Fed Funds rate tightening well, up 75% of years with rate hikes since 1994	Jan-07-2000 Jul-07-1999
4. PHARMA:	Pharma needs to acquire biotechs due to ~\$182B revenue at risk with pharma patent expiration; there is >\$500B of M&A firepower at top 18 pharmas	Company  Obbvie  Johnson-John  Jazz Paracoo
5. DEMAND:	Medical spend will accelerate in the future; e.g., >90M Americans in Medicare by 2060	Sunovid

#### NBI, S&P 500, Tech Select (XLK) and Russell 2000 Returns, Jan 1999 to Nov 3, 2023



#### Key Drugs Losing Patent Exclusivity by 2028



### Key Value Drivers

Multiple clinical milestones expected across Wholly Owned Programs, Seaport & Gallop

#### THERAPEUTIC CANDIDATE<sup>1</sup>

#### **MILESTONES**

LYT-100	Deupirfenidone		Results from registration-enabling trial in IPF	4Q 2024
LYT-200	Anti-Galectin-9 Mab	•	Additional results from Phase 1b in combination w/ tislelizumab in solid tumors	2024
LYT-300	Glyph Allopregnanolone		Initiation of Phase 2 clinical trial in FXTAS	2024
LYT-310	Glyph Cannabidiol	•	Initiation of clinical trial	1H 2024
LV/T 720			Initiation of first-in-human enabling studies	2024
LYT-320	Glyph Agomelatine		Initiation of clinical studies	1H 2025

Key anticipated milestones are **bolded** 

### Multiple additional catalysts across Founded Entities



## PureTech Team - Bringing Our Vision to Life











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

**269,949,741** outstanding shares as of February 2024

Market capitalization \$673M (£532M) as of February 29, 2024; 1.27 USD:GBP

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

### **ANALYST COVERAGE**

**Peel Hunt LLP** 

Miles Dixon

Liberum

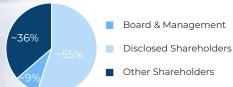
**Edward Thomason** 

**Leerink Partners LLC** 

Thomas J. Smith

**Jefferies International Limited** 

Peter Welford



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.

¹ 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 ☐ I YT-100 Market Research ☐ LYT-100 Payor Research ☐ LYT-100 In the Face of Generics ☐ Pirfenidone Safety Data ☐ Lung Disease Prevalence ☐ Glyph Technology Platform ☐ LYT-200 Supplemental Data Appendix B: Founded Entities ■ Seaport □ Vedanta □ Akili □ Vor ■ Sonde Entrega ☐ Founded Entities Upcoming Catalysts Appendix C: Supplemental Materials ■ PureTech ESG Program ■ PureTech's Proven Expertise ☐ PureTech is Executing & Delivering Results ☐ Financial Highlights/Non-IFRS Measures

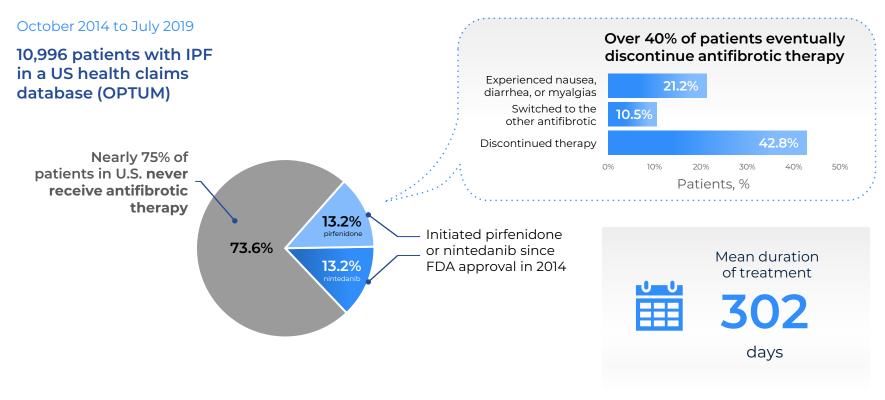


# Appendix A: Wholly Owned Pipeline



### IPF Treatment Rates are Low Due to Side Effects

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





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

1. Initial PK studies

FOUNDATIONAL PK DATA

Multiple-dose safety. tolerability, and PK

MAD 1.0

> Tolerable up to 1000mg BID, linear PK

Determine dose with same exposure as pirfenidone

matches pirfenidone AUC

2. Head-to-head tolerability TOI FRABILITY ADVANTAGE VS. **PIRFENIDONE** 

> 550 mg TID LYT-100 vs. pirfenidone:

Comparable AUC, reduced C<sub>max</sub>

Older

Adult

Demonstrated tolerability advantage over pirfenidone

3. High-dose studies **EXPLORE FEASIBILITY OF** HIGHER EXPOSURES

> Safety and tolerability > 1000 mg BID

MAD 2.0

> Tolerable up to 2000mg BID with no dose limiting toxicity



800 - 850 mg BID

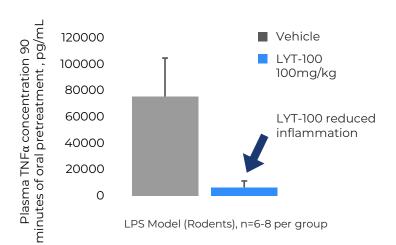


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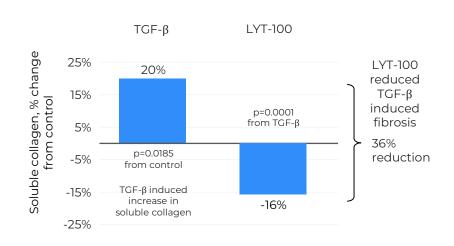
### LYT-100:

Preclinical POC demonstrates anti-inflammatory & anti-fibrotic pharmacology

# PRECLINICAL PLASMA CONCENTRATIONS OF TNFA WITH LYT-100 VERSUS CONTROL

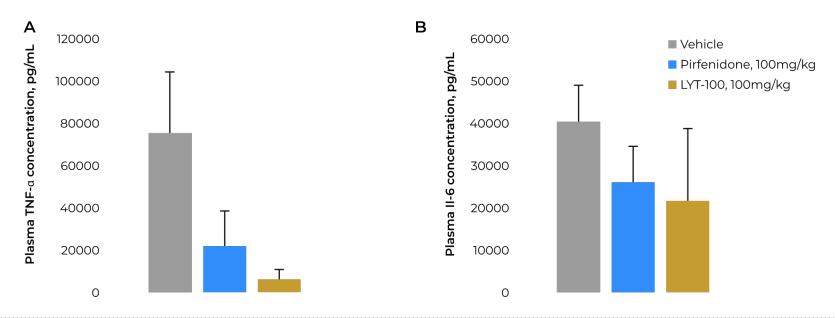


# IN VITRO REDUCTION OF TGF-B INDUCED SOLUBLE COLLAGEN PRODUCTION (MOUSE FIBROBLASTS)



### LYT-100 Preserves Pharmacologic Effect of Pirfenidone:

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

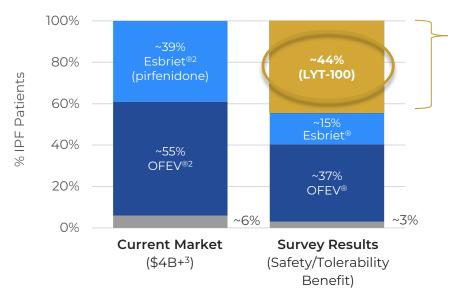


Reduction in LPS-stimulated plasma concentrations of TNF- $\alpha$  and IL-6 by Pirfenidone or LYT-100. Oral doses of vehicle, pirfenidone, or LYT-100 (100mg/kg) administered 60 minutes prior to LPS (30  $\mu$ g/kg intravenous): TNF- $\alpha$  (A) and IL-6 (B) measured 90 min after LPS stimulation: N=6-8 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>



- Newly diagnosed patients

  Pulmonologists would prescribe LYT-100 to

  ~44% of newly diagnosed patients with IPF
- 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:

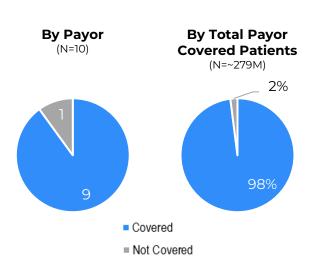
- (1) have never started treatment, AND
- (2) who started treatment but discontinued



### 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.."4 – 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.
- 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> <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> <li>Safety/tolerability advantage will enable LYT-100 to complete for new patient starts in plans without step edits</li> <li>In plans with step edits, LYT-100 will be used as second line of treatment for patients who fail on generic antifibrotics</li> <li>Even if all payers require step edits, ~50% of patients will be eligible for LYT-100</li> </ul>
Reformulated pirfenidone and nintedanib	<ul> <li>A few reformulated pirfenidone and nintedanib approaches, including inhaled and sustained release, are in early development</li> </ul>	<ul> <li>LYT-100 will offer oral systemic delivery of the medication, without the AEs associated with inhaled (e.g., cough) and other reformulations of the currently approved drugs</li> <li>None of the localized delivery candidates have demonstrated the same evidence of efficacy as systemic therapies</li> </ul>
Novel Mechanisms	<ul> <li>Nearly all new mechanisms are being studied on top of/or after the standard of care (currently pirfenidone &amp; ninetedanib)</li> </ul>	<ul> <li>Potential for LYT-100 to be the backbone standard of care for future combination regimens</li> <li>Pirfenidone and nintedanib remain key competitors for LYT-100</li> </ul>

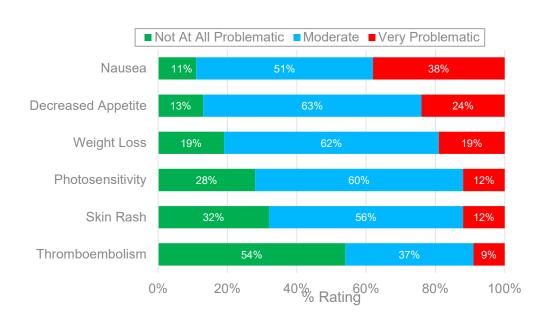


### ESBRIET (pirfenidone) and OFEV (nintedanib):

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

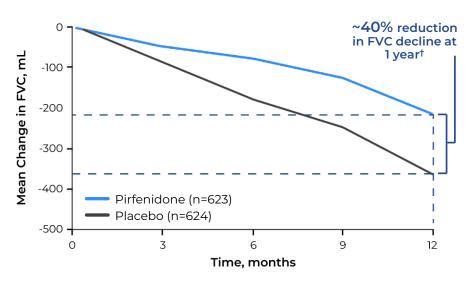
# HOW PROBLEMATIC ARE AES ASSOCIATED WITH IPF SOC? (N=100)



### Pirfenidone:

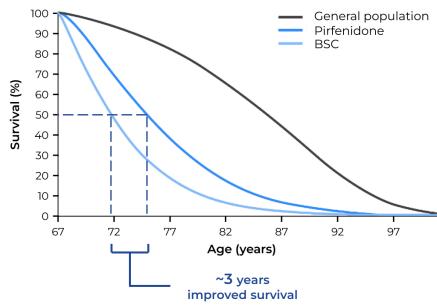
A clinically validated treatment for IPF with beneficial effects on FVC and survival

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)1

#### Pirfenidone GLAEs:

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

#### PIRFENIDONE FOOD **EFFECT/ANTACID STUDY<sup>2</sup>**

801mg single-dose in healthy older

PIRFENIDONE FOOD EFFECT AND **BIOEOUIVALENCE STUDY**<sup>3</sup>

801mg single-dose in healthy adults, 36% women

PIRFENIDONE PHASE 3 STUDIES<sup>1</sup>

2403mg per day, IPF patients 26% women

Design

adults, 44% women

Most common AEs	Pirfenidone N=16
Nausea	43.8%
Dizziness	37.5%

Most common **AEs** 

AEs more frequent in the fasted state AE rate higher in women

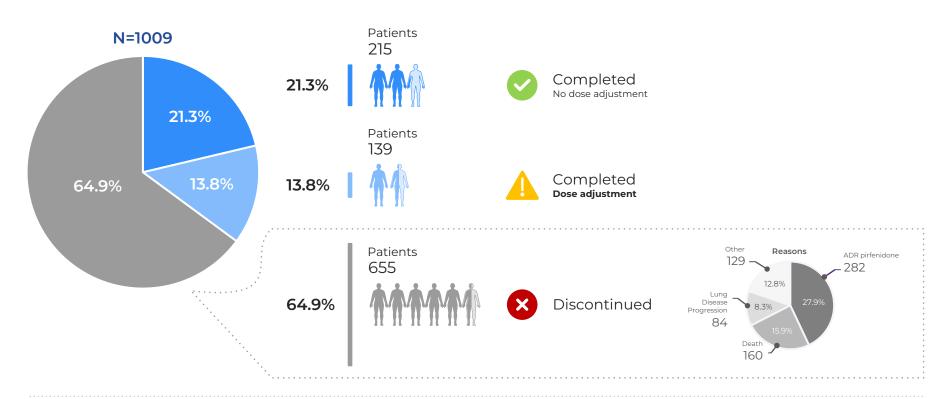
Most common AEs	Pirfenidone N=44			
Nausea	29.5%			
Dizziness	18.2%			
Headache	9.1%			
Constipation	9.1%			
Vomiting	4.5%			
Dyspepsia	4.5%			
AEs more frequent in the fasted state				

Most common GI AEs^	Pirfenidone N=623	Placebo N=624
Nausea	36%	16%
Rash	30%	10%
Ab. pain	24%	15%
Diarrhea	26%	20%
Headache	22%	19%
Dyspepsia	19%	7%
Dizziness	18%	11%
Vomiting	13%	6%
Anorexia	13%	5%

^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%), arthalgia (10% vs. 7%)



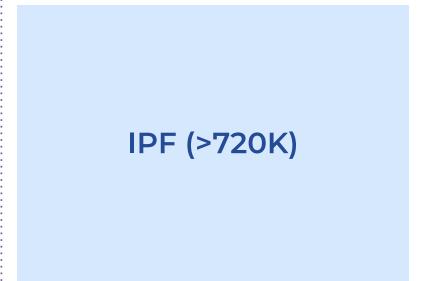
### Prospective Registry Found Only 21% of Patients Who Started Pirfenidone Remained on Full Dose After 2 Years

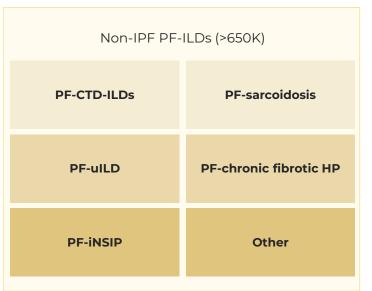




### Enduring High Unmet Need in Interstitial Lung Diseases Including IPF

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

### Proprietary Technology Platforms

Designed to harness the lymphatic system & administer therapeutics to immune, lymphatic and inflamed tissue



### APPLICATION/FOCUS

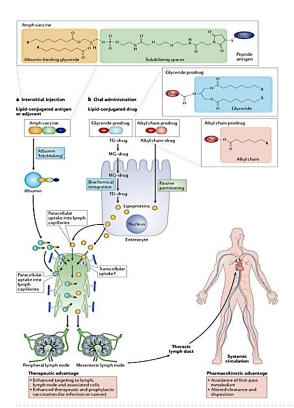
Employs the body's natural lipid absorption & transport process to **orally administer drugs** via the lymphatic system by **bypassing first-pass metabolism** 

Enable **oral administration** of therapeutic payloads, such as **mRNA**, **biologics**, **vaccines and other drugs**, that are otherwise not efficiently absorbed when taken orally or are otherwise administered exclusively by injection

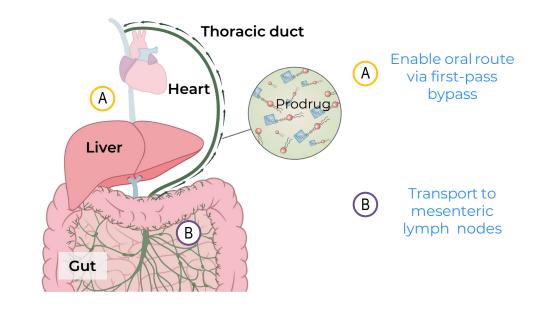


### Glyph™ Technology Platform:

Designed to utilize natural lipid transport system to enable lymphatic targeting



# LIPID PRODRUGS PROVIDE MULTIPLE OPPORTUNITIES TO ENHANCE SMALL MOLECULE DRUGS



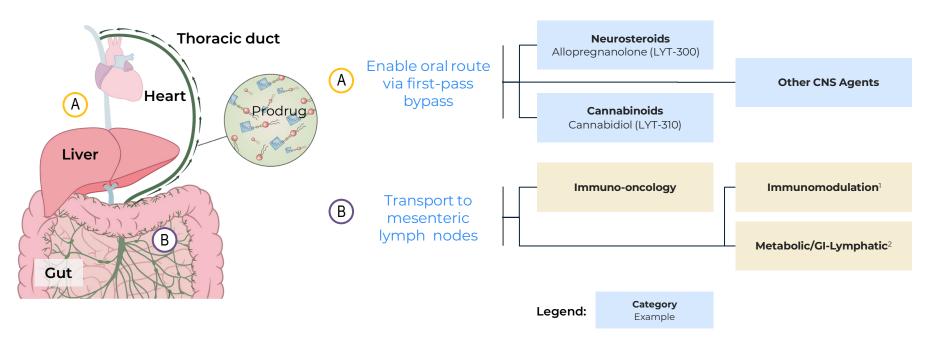


Trevaskis, N. L., et al. Nat. Rev. Drug Discov. 14, 781-803 (2015).

### Glyph™ Technology Platform:

Exploring therapeutic approaches enabled by transporting via the lymphatic system

# LIPID PRODRUGS PROVIDE MULTIPLE OPPORTUNITIES TO ENHANCE SMALL MOLECULE DRUG DISTRIBUTION

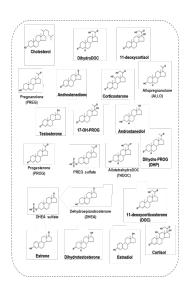




### Glyph™ Technology

Technology enables oral administration of natural neurosteroids

#### NATURALLY-OCCURRING NEUROSTEROIDS

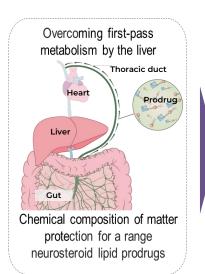


#### UNTAPPED OPPORTUNITIES DUE TO CLINICAL TRANSLATION HURDLES

- a) Not orally bioavailableb) Properties not druglike
- c) No composition-ofmatter IP

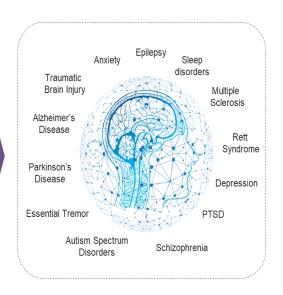
Chemical modifications may alter target engagement

### UNLOCKING THERAPEUTIC POTENTIAL / VALUE



Harnesses natural mechanisms (validated efficacy)

### POTENTIALLY APPLICABLE TO A RANGE OF CNS CONDITIONS



### LYT-300: Potential First-in-Disease Therapy for FXTAS

PureTech awarded ~\$11.4M grant in competitive process



Phase 2 trial initiation in 2024

### LATE ONSET & DEVASTATING RARE DISEASE

### Fragile X-associated Tremor/ Ataxia Syndrome

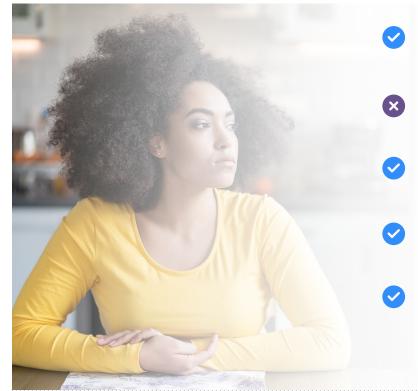
- Closely related to, but distinct from, fragile X syndrome (FXS);
   both conditions are the result of repeated elements in the FMRI gene
- Clinical signs, including tremor, balance problems and cognitive decline

### UNMET MEDICAL NEED

- Currently, there are no primary treatments for FXTAS
- Only **one treatment has shown clinical benefit**: intravenous allopregnanolone significantly (p=0.009) improves executive cognitive / motor function (BDS-2), N=6, open label

### LYT-310 (Glyph Cannabidiol)

### For epilepsies & other neurological indications



A CBD-based product is approved in the U.S. and the EU to treat seizures resulting from certain rare conditions

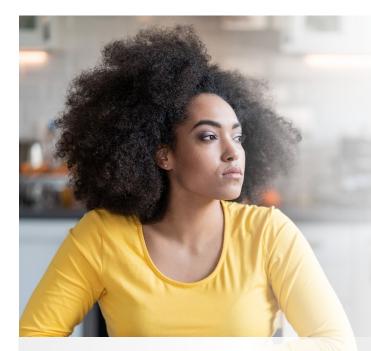
...BUT its dosing (large volume of oily solution via syringe) limits its use in broader indications and age groups

LYT-310 has the potential to enable oral administration, improve safety, and reduce GI side effects

LYT-310 may allow for a readily scalable, consistent product in a cost-effective manner

LYT-310 could expand the therapeutic application of CBD to a wider range of age groups and indications

### LYT-310: Potential to Be Highly Differentiated in Epilepsies & Other Neurological Indications



Initiation of Phase 1 trial expected in 1H 2024

### **Epilepsies**

- ~3 million adults and 470,000 children are affected by epilepsy in the U.S.<sup>1</sup>
- 20-33% of patients with epilepsy have drug-resistant epilepsy<sup>2</sup>

#### UNMET MEDICAL NEED

Despite the many approved antiseizure medications, patients are often refractory to treatments or discontinue effective treatments.

There is a need for treatment options with better safety, tolerability, efficacy, and convenience.

### LYT-310: Oral CBD A Highly Differentiated Profile for Broad Application

#### APPROVED CBD ORAL SOLUTION



Target Population

Target Indications

#### **PURETECH'S LYT-310**



1 yr+, Primarily Teens/Adults

Pediatric form planned with target down to 1 year old

#### Approved in US for 3 Rare Epilepsies

Approved in US and EU for Lennox-Gastaut syndrome, Dravet syndrome, or tuberous sclerosis complex

#### Large Volume of Oily Solution

Administered via syringe twice per day



Dosina

#### Side Effects



#### **Epilepsies & Other CNS Indications**

Exploring range of rare and more common forms of epilepsy as well as other CNS disorders

#### Capsules & Other Flexible Routes

Capsule dosage form plus water-based emulsion planned for pediatric patients

#### Potential for Reduced Side Effects

...due to lower dose to GI & liver, a result of bypassing first-pass metabolism in liver

#### Standard Manufacturing

Standard pharmaceutical manufacturing techniques

Processed from extraction of cannabis plant

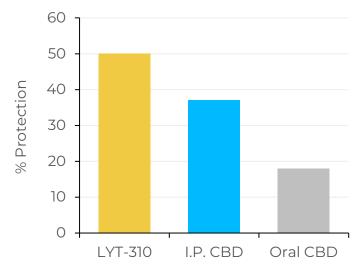




### LYT-310: Positive Data in Gold Standard MES Seizure Model

- LYT-310 was more effective at preventing seizures at a dose 3x lower than an oral CBD formulation<sup>1</sup>
- ED<sub>50</sub> for LYT-310 corresponds to an ED<sub>18</sub> for synthetic oral CBD
- Over 85% (18 of 21) of the approved anti-seizure medications for focal seizures were active in MFS<sup>2</sup>

LYT-310 ED50 DOSE DEMONSTRATED GREATER SEIZURE PROTECTION AT EQUIVALENT CBD DOSE (N=12/GROUP)



Initiation of Phase 1 trial expected in 1H 2024



# 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

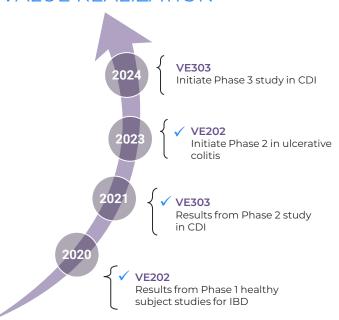


#### **VALIDATION**

- Four clinical-stage programs in development
- VE303 (C. difficile) demonstrated accelerated gut microbiota restoration after antibiotics in a Phase la/lb 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

### (PRTC Ownership: 41.0%<sup>1</sup>)

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



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



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

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

#### 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 inperson 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®² (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

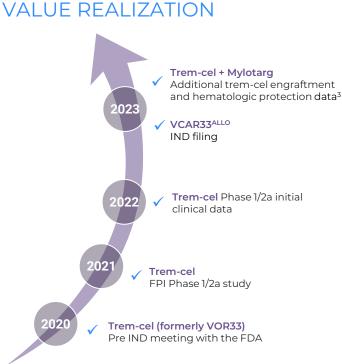
### **VALIDATION**

- Ex vivo & mouse proofof-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 ontarget 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

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

(PRTC Ownership: 4.0%<sup>1</sup>)



#### eHSC Platform

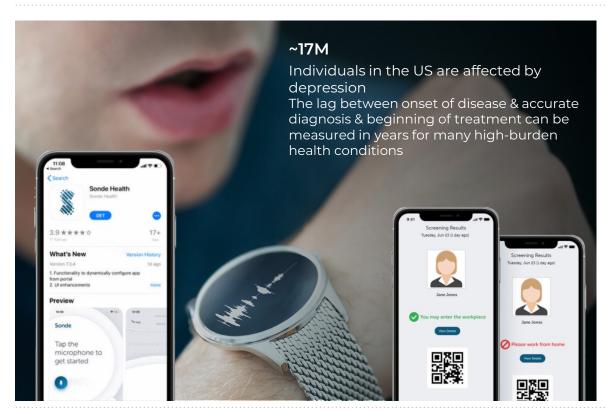
✓ Engineered hematopoietic stem cells (eHSCs) deleting redundant epitopes, protecting healthy cells from targeted therapies



### Sonde

### (PRTC Ownership: 35.2%1)

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



### 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**

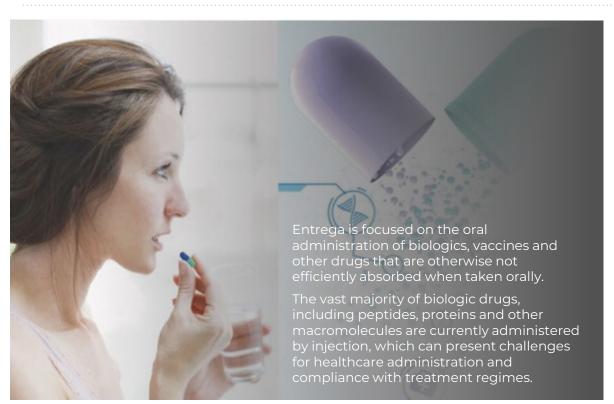
- Technology has demonstrated the potential to screen & monitor for disease in individuals from brief samples of speech
- Ongoing collaborations with multiple US & ex-US hospitals, clinics & academic medical centers
- Partnership with Qualcomm Technologies for vocal biomarker technology
- Collected voice 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

### (PRTC Ownership: 73.8%1)

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



#### MILESTONES ACHIEVED

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



### Upcoming Value Drivers: Founded Entities

Potential launch momentum & source of capital to further fund advancement of Wholly Owned Programs

ENTITY	PURETECH OWNERSHIP <sup>1</sup>	THERAPEUTIC CANDIDATE	EXPECTED MILESTONES								
				NDA submission for KarXT in schizophrenia	3Q 2023						
		KarXT <sup>3</sup>	•	Results from Phase 3 EMERGENT-4 & EMERGENT-5 trials for schizopherenia	2H 2024						
<b>Karuna</b> (NASDAQ: KRTX)	2.3%			Results from Phase 3 ARISE trial for schizophrenia	2H 2025						
				Results from Phase 3 ADEPT-1 & ADEPT-2 trials	2026						
		KAR-2618		Initiation of Phase 1b trial for MDD	2024						
Akili	14.604	AKL-T01 <sup>4</sup>	€	Secure label expansion for 13-17 y/o children with ADHD	2023						
(NASDAQ: AKLI)	14.6%	14.0%	EndeavorOTC <sup>4</sup>	•	Data submission to FDA to market EndeavorOTC as OTC treatment for adults with ADHD	2024					
Vor	4.0%	Trem-Cel <sup>3</sup>	<b></b>	Additional trem-cel engraftment and hematological protection data updates	2023						
(NASDAQ: VOR)								VCAR33 <sup>ALLO3</sup>		IND application to support Phase 1/2 clinical trial	1H 2023
Vedanta	41.0% <sup>2</sup>	VE303 <sup>3</sup>	•	Initiation of Phase 3 in <i>C. difficile</i>	1H 2024						
Sonde	35.2%	Sonde App	•	Launch of key pilot programs	2023						



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



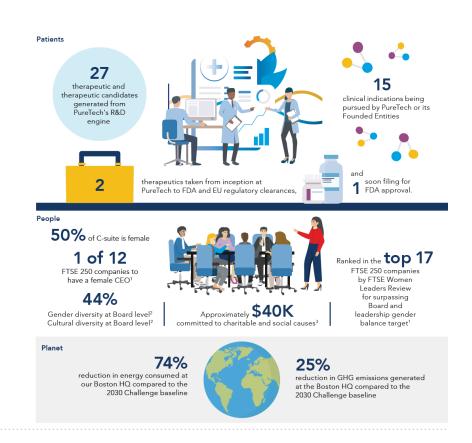
# Appendix C: Supplemental Materials



### 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



### 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
KarXT, invented by PureTech Team	Karuna's KarXT 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  Muscarinic agonist Muscarinic antagonist Trospium chloride
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
	LYT-300 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
	LYT-310 for epilepsies & other neurological indications	Cannabidiol (CBD) effective in several epilepsies	GI tolerability & liver safety issues as well as undesirable dosing of large amounts of oily liquid via oral syringe	Using proprietary Glyph technology, developed CBD prodrug that enables capsule formulation w/ higher bioavailability, reducing Gl/liver exposure & potentially related AEs



### PureTech is Executing & Delivering Results

#### **REGULATORY**

FDA Clearance & EndeavorRx®1 (AKL-TO1)

European CE Mark

Plenity\*2 (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
- √ Karuna's strategic collaboration with Zai Lab
  - \$35 million in upfront payment for development, manufacturing, & commercialization of KarXT in Greater China, & up to an additional \$80 million in development and regulatory milestones; Karuna also eligible to receive up to \$72 million in sales milestones & low-double-digit to high-teens tiered royalties based on annual net sales of KarXT in Greater China
- 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**

- ✓ Karuna's \$124M Series A+B financings; \$103M IPO
  - 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,
  - 5AM Ventures, Johnson & Johnson Innovation—JJDC, Inc. (JJDC), Osage University Partners, Novartis Institutes for BioMedicalResearch
- 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®, 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

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PURETECH
GIVING LIFE TO SCIENCE\*

### Financial Highlights

Estimated December 31, 2023 \$ millions

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

Consolidated Cash, Cash Equivalents, and Short-Term Investments

320.0

Less: Cash and Cash Equivalents held at non-wholly-owned subsidiaries

(1.2)

PureTech Level Cash, Cash Equivalents, and Short-Term Investments<sup>2</sup>

318.8



### 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



### Non-IFRS Measures Reconciliation – Karuna ROI

Investments Held at Fair Value @ 12/31/2022 in audited consolidated balance sheet	251.9
(-) Other Investments Held at Fair Value @ 12/31/2022	(44.7)
Karuna Investment Held at Fair Value @ 12/31/2022	207.2
(+) Sale of 167,579 shares of Karuna in October through December 2023	(33.3)
(+/-) Karuna Fair Value Gain/ Loss for the period 12/31/2022 to 12/31/2023	118.8
(a) Karuna Investment Held at Fair Value @ 12/31/2023	292.7
Proceeds From Sale of Investments Held at Fair Value @ 12/31/2020	350.6
(-) Sale of 2,119,696 shares of resTORbio	(3.0)
Proceeds From Sale of Karuna @ 12/31/2020	347.5
(+) Sale of 1,000,000 shares of Karuna @ 2/9/2021	118.0
(+) Sale of 750,000 shares of Karuna @ 11/10/2021	100.1
(+) Sale of 602,100 shares of Karuna during August and September 2022	115.5
(+) Sale of 167,579 shares of Karuna in October through December 2023	33.3
Proceeds From Sale of Karuna	714.4
b) Total Karuna Investment Held at Fair Value and Proceeds @ 12/31/2023	1,007.1
c) Total PureTech Principal Investment in Karuna	18.5
c 1 Return on Investment (ROI)	53.5

