



# Harnessing fibrosis, unleashing life

Corporate Presentation | January 2026

# Forward Looking Statements

This presentation and various remarks we make during this presentation contain forward-looking statements of Rein Therapeutics, Inc. (“Rein”, the “Company”, “we”, “our” or “us”) within the meaning of the Private Securities Litigation Reform Act of 1995, including statements with respect to: Company’s RENEW Phase 2 clinical trial of LTI-03, including with respect to the timing of the trial and the assumption that Company will raise the funds necessary to conduct the trial; future expectations, plans and prospects for the Company; the sufficiency of the Company’s cash resources; and the potential commercial opportunity of LTI-03 and LTI-01. We use words such as “anticipate,” “believe,” “estimate,” “expect,” “hope,” “intend,” “may,” “plan,” “predict,” “project,” “target,” “potential,” “would,” “can,” “could,” “should,” “continue,” and other words and terms of similar meaning to help identify forward-looking statements, although not all forward-looking statements contain these identifying words. Actual results may differ materially from those indicated by such forward-looking statements as a result of various important factors, including risks and uncertainties related to: the ability of the Company to obtain the cash resources to fund the RENEW Phase 2 clinical trial and the Company’s operations for the necessary periods of time, and the Company’s ability to manage unplanned cash requirements; changes in applicable laws or regulations; the possibility that the Company may be adversely affected by other economic, business, and/or competitive factors, including risks inherent in pharmaceutical research and development, such as: adverse results in the Company’s drug discovery, preclinical and clinical development activities, the risk that the results of preclinical studies and early clinical trials may not be replicated in later clinical trials, including the RENEW Phase 2 clinical trial of LTI-03, or that partial results of a trial will be indicative of the full results of the trial, the Company’s ability to enroll patients in its clinical trials, and the risk that any of its clinical trials may not commence, continue or be completed on time, or at all; and decisions made by the U.S. Food and Drug Administration and other regulatory authorities, investigational review boards at clinical trial sites and publication review bodies with respect to our development candidates; as well as the risks and uncertainties discussed in the “Risk Factors” section of the Company’s Annual Report on Form 10-K for the year ended December 31, 2024, which is on file with the United States Securities and Exchange Commission (the “SEC”), and in the subsequent filings that the Company files with the SEC. These forward-looking statements should not be relied upon as representing the Company’s view as of any date subsequent to the date of this presentation, and we expressly disclaim any obligation to update any forward-looking statements, whether as a result of new information, future events or otherwise, except as required by law. This Presentation is for informational purposes only and shall not constitute an offer to sell or the solicitation of an offer to buy any securities, or a solicitation of any vote or approval, nor shall there be any sale of securities in any states or jurisdictions in which such offer, solicitation or sale would be unlawful prior to registration or qualification under the securities laws of any such jurisdiction or an exemption therefrom.

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# Pioneering Best-in-Class Therapies for Pulmonary & Fibrosis Indications

- Clinical-stage biotech company **pioneering potential first-in-class multi-pathway therapies in orphan pulmonary and fibrosis indications**
- **LTI-03** is a **potential blockbuster treatment** which has demonstrated antifibrotic and regenerative properties
  - **Unique dual mechanism** promoting alveolar epithelial cell survival & inhibiting profibrotic signaling
  - Favorable safety profile to date
  - KOL support for new, safer and effective therapies
  - Initial interim data anticipated second half 2026

# Therapies for Underserved Fibrosis and Pulmonary Conditions

## LTI-03

*Idiopathic Pulmonary  
Fibrosis*

*Phase 2*

- Phase 1 clinical met primary endpoint; High dose LTI-03 (5mg BID) was **well-tolerated** with **no identified safety issues**
- Evaluated a robust set of **exploratory biomarkers** with **predictive value of lung health**
- Results **validated preclinical & clinical findings** for key biomarkers with **dose-dependent effects**

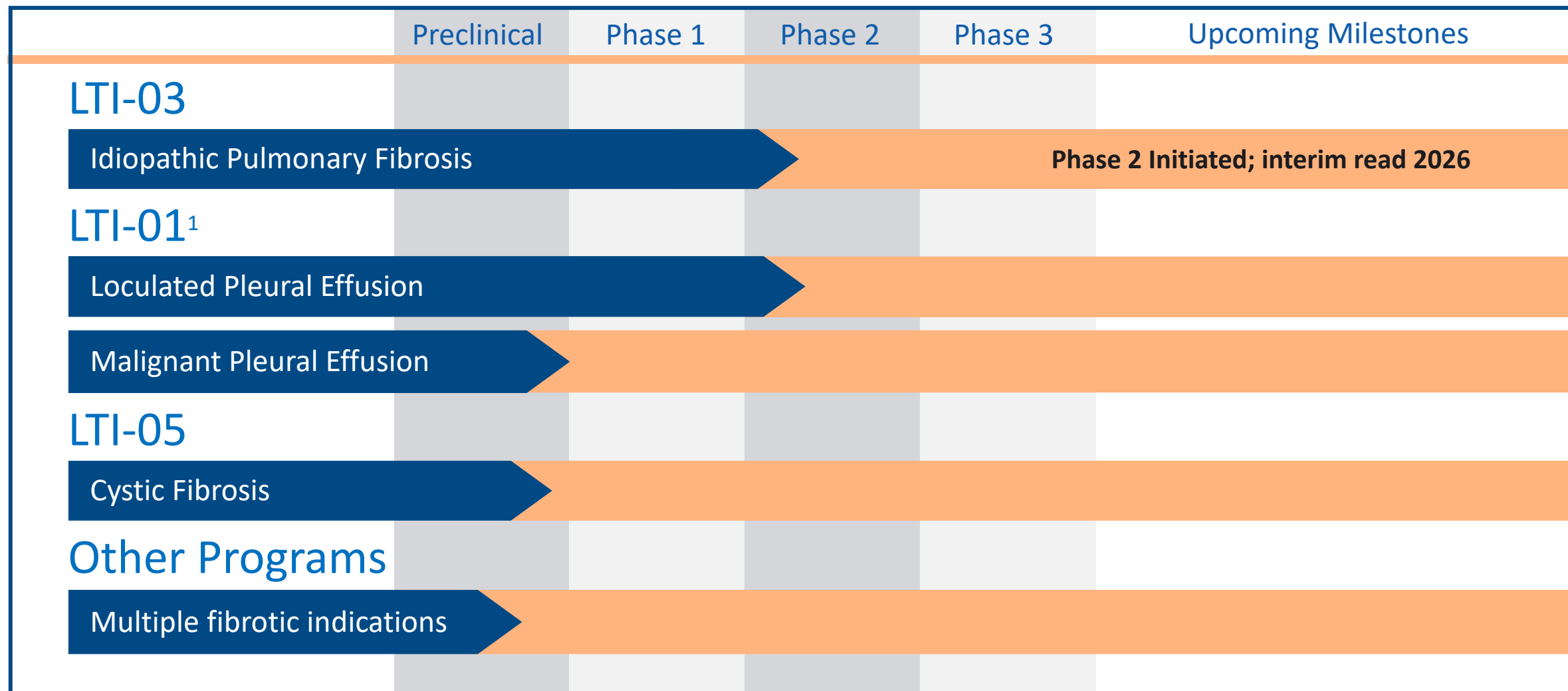
## LTI-01

*Loculated Pleural  
Effusions*

*Phase 2b  
ready*

- **Potentially fatal disease with no approved drugs**
- Completed Phase 1b and Phase 2 trials; similar mechanism as existing, off label therapeutic use

# Multiple Orphan Disease Programs Ready for Phase 2 Clinical Trials



<sup>1</sup>In June 2024, the Company decided to temporarily delay clinical development of LTI-01 in an effort to focus its resources on clinical development of LTI-03 and until additional funds are raised.

# Led by Experienced Biotech and Pulmonary Team

## Key Management and Board of Directors



Brian Windsor, Ph.D.  
*Chief Executive Officer  
and Director*



Joe von Rickenbach  
*Chairman*



Cory H. Hogaboam, Ph.D.  
*Chief Scientific Officer*



Bill Fairey  
*Director*



Tim Cunningham  
*Chief Financial Officer*



Alan Musso  
*Director*



Kristie Lauterbach  
*VP Quality*



Reinhard Ambros, Ph.D.  
*Director*



# **LTI-03: A Novel Treatment Designed to Reverse the Course of IPF**

# Idiopathic Pulmonary Fibrosis – a Deadly Diagnosis

- Idiopathic Pulmonary Fibrosis, or IPF, is a fatal age-related disease characterized by progressive scarring in the lungs<sup>1</sup>
- IPF is part of a larger group of diseases known as Interstitial Lung Diseases, or ILDs, which are diseases characterized by lung inflammation and/or scarring. There are more than 200 types of Pulmonary Fibrosis conditions within ILDs<sup>2</sup>
- Approximately 100,000 patients in the US alone are living with IPF each year<sup>3</sup> and more than 250,000 Americans live with some sort of pulmonary fibrosis<sup>2</sup>
- Median survival from the time of diagnosis is 3-5 years



<sup>1</sup>Mora, A., Rojas, M., Pardo, A. et al. Emerging therapies for idiopathic pulmonary fibrosis, a progressive age-related disease. Nat Rev Drug Discov 16, 755–772 (2017).

<sup>2</sup> <https://www.pulmonaryfibrosis.org/understanding-pff/about-pulmonary-fibrosis/what-is-pulmonary-fibrosis>

<sup>3</sup> <https://www.healthline.com/health/managing-idiopathic-pulmonary-fibrosis/ipf-facts#prevalence>

<sup>4</sup> Kirby, Living with idiopathic pulmonary fibrosis, The Lancet VOLUME 9, ISSUE 2, P136-138, FEBRUARY 2021

# Sizable Global Opportunity with Potential Upside









- Only 3 drugs are approved for IPF as of 2025
- Ofev<sup>®</sup>, the global leader, did \$4.0B in sales for 2024<sup>1</sup>
- The estimated global market for IPF alone is projected to be \$11.7B<sup>2</sup>
- Other PF and ILDs represent upside for a successful IPF drug
- The mechanism of LTI-03 could potentially address other fibrosis conditions, such as those associated with the heart, kidneys, liver, skin, or eyes



<sup>1</sup> <https://www.boehringer-ingenelheim.com/us/about-us/who-we-are/2024-results-research-and-development-investment-rise>

<sup>2</sup> iHealthcareAnalyst Global Idiopathic Pulmonary Fibrosis Market \$11.7 Billion by 2031 January 5, 2024 by iHealthcareAnalyst, Inc. <https://www.ihealthcareanalyst.com/global-idiopathic-pulmonary-fibrosistreatment-market/>

# Select Competitive Landscape<sup>1</sup>

Company	Compound	Single or Multi-pathway	Mechanism	Target Cell Types	Clinical Stage
	LTI-03	Multi-pathway	Caveolin-1 CSD mimic	Fibroblasts, type 2 Epithelial cells, Aberrant basaloid cells, Macrophages	P2
	Nintedanib	Multi-pathway	Broad tyrosine kinase inhibitor	Fibroblasts	Approved
	Pirfenidone	Multi-pathway	unknown	Fibroblasts	Approved
	Nerandomilast	Multi-pathway	PDE-4B inhibitor	Aberrant basaloid cells, Fibroblasts	Approved
	Tyvaso	Multi-pathway	Prostacyclin mimic	Smooth Muscle, endothelium, immune cells	P3
	Buloxibutid	Single	Angiotensin II type 2 receptor agonist	Type 2 Epithelium	P2
	APO-1	Multi-pathway	unknown	Fibroblasts	P2
	TTI-101	Multi-pathway	STAT3 inhibitor	Aberrant basaloid cells, Fibroblasts	P2 halted

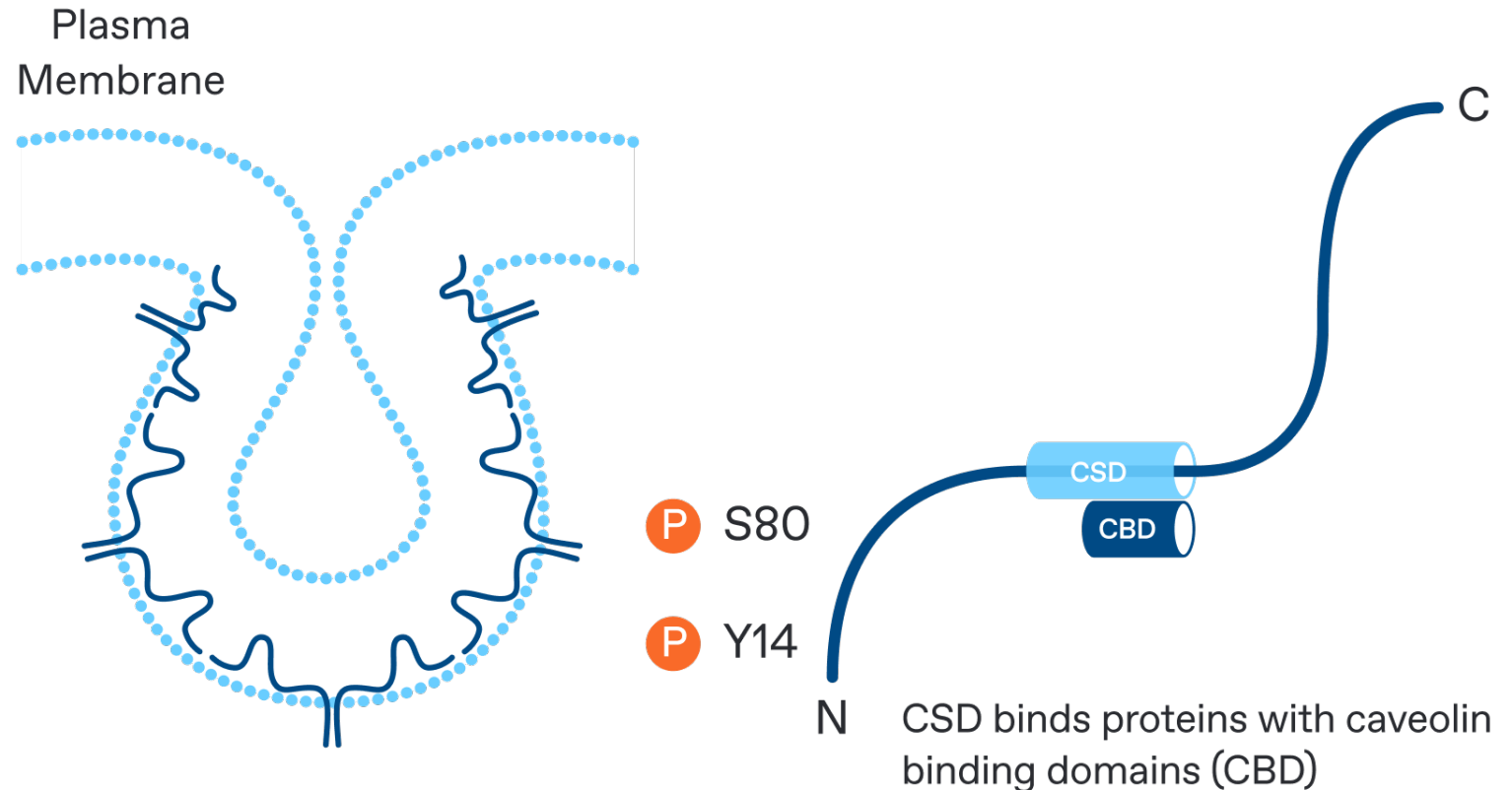
<sup>1</sup>Libra, A., Sciacca, E., Muscato, G., Sambataro, G., Spicuzza, L., & Vancheri, C. (2024). Highlights on Future Treatments of IPF: Clues and Pitfalls. International Journal of Molecular Sciences, 25(15). <https://doi.org/10.3390/IJMS25158392>




# LTI-03: the Critical Portion of the CSD Region of Cav1

## Mimics the Regulatory Activity of Cav1, Affecting a Wide Range of Proteins Involved in Fibrosis

- LTI-03 is a seven amino acid peptide comprising a portion of the Cav1 CSD. Substitution/deletion analysis revealed it is the smallest CSD fragment that retains functionality
- This hydrophobic peptide can enter cells, and it may be acting both at the cell membrane and intracellularly
- Studies have shown that the LTI-03 peptide can affect phosphorylation of dozens of profibrotic proteins
- LTI-03 is dosed direct to the lungs by dry powder inhaler, and studies have shown that intact peptide can be detected in the lungs 24 hours after administration



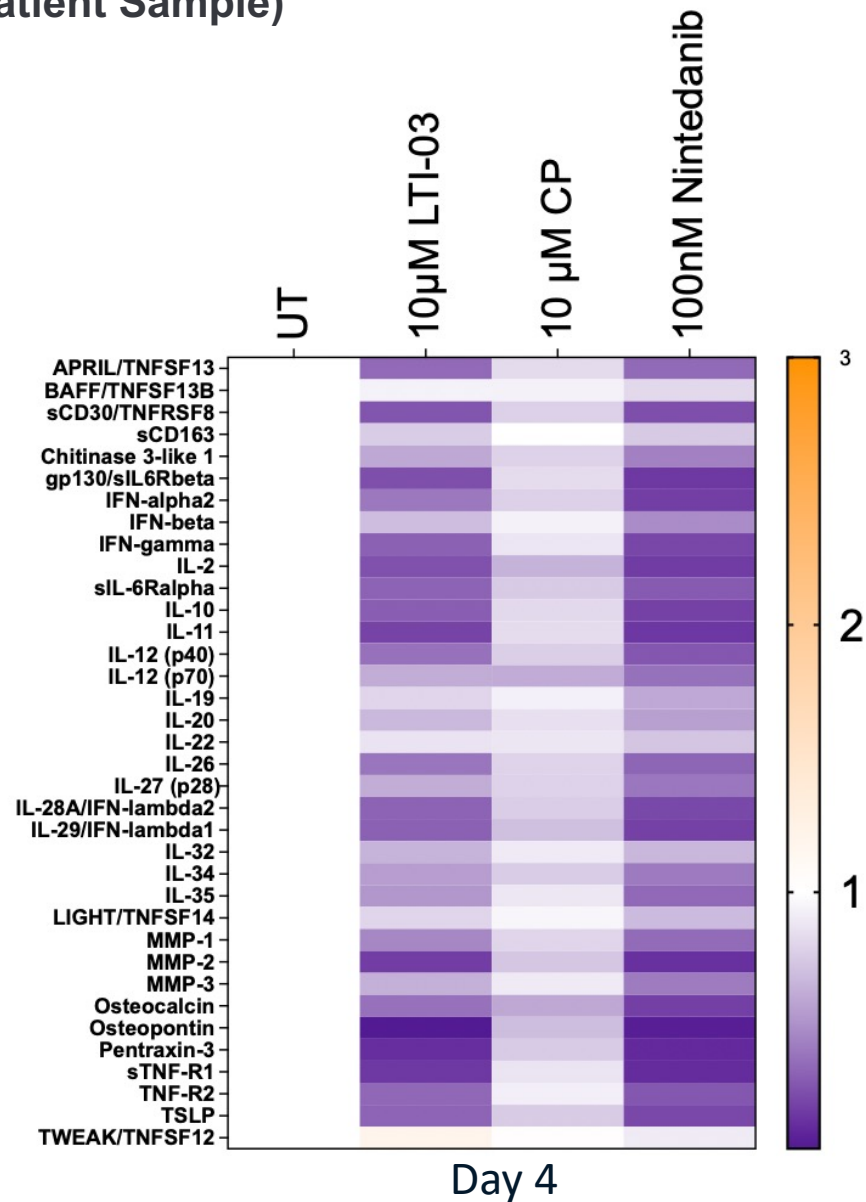
 full CSD (20-mer): N-DGIWKASFTTFTVTKYWFYR-C

 **LTI-03 (7-mer): FTTFTVT**  
predicted molecular weight: 815.92 Da

For a review on CSD/CBD binding domain list, see: Byrne et. al. PLOS One 2012

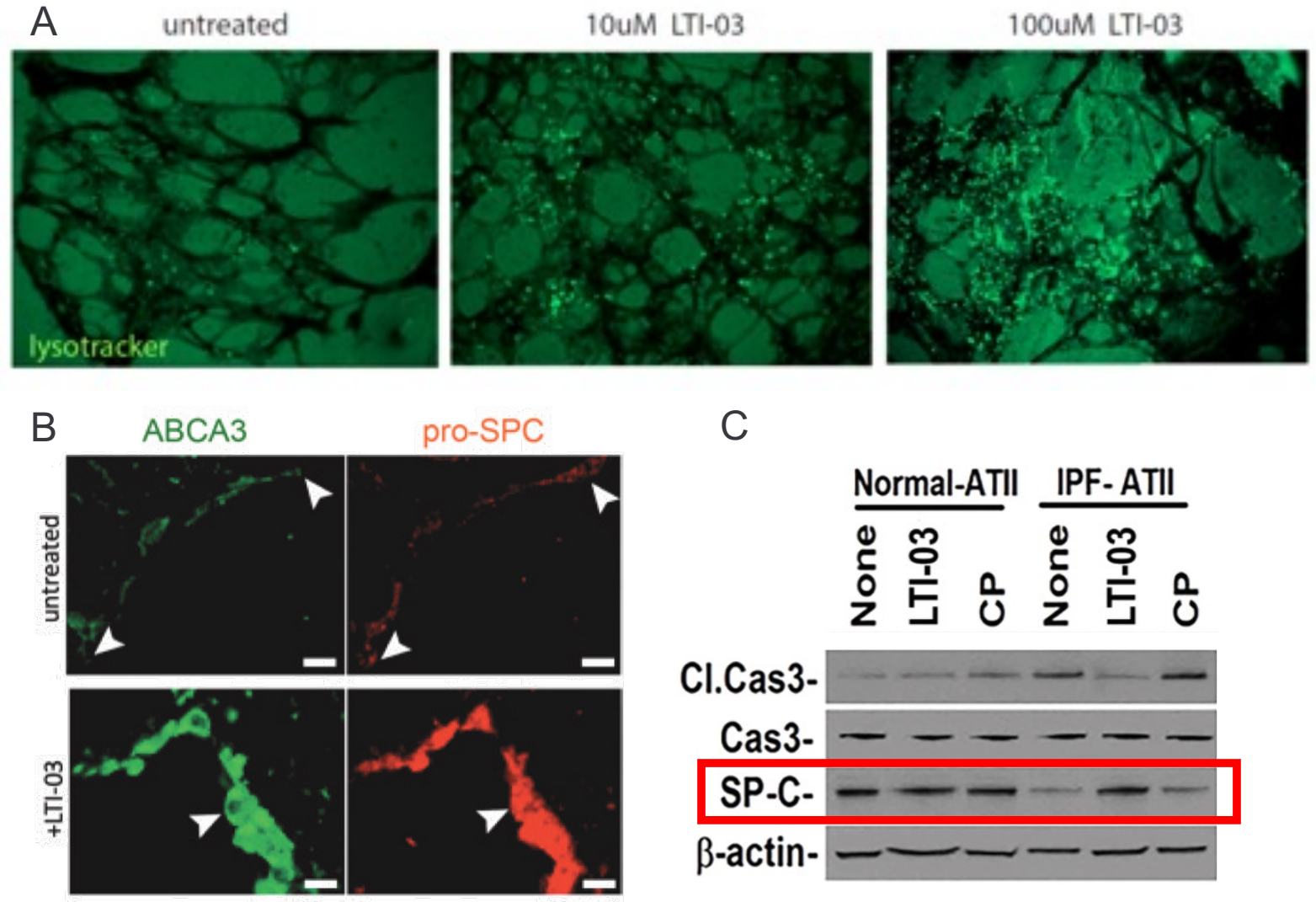
## Antifibrotic Activity: Single dose LTI-03 inhibits multiple profibrotic proteins similar to Ofev® (Every 12hrs in Precision Cut Lung Slices (PCLS)—Single Patient Sample)

- As an **antifibrotic**, LTI-03 inhibits large panels of profibrotic proteins in a manner similar to the standard of care drug Ofev® (nintedanib)
  - Darker purple = more inhibition of the protein
- The PCLS tissue culture system uses actual biopsied tissue from an IPF lung (removed due to lung transplant), preserving all cell types in the IPF lung
- 10 µM LTI-03 is equivalent to an approximate dose of 1 mg in a dry powder inhaler. Phase 1b trial tested 5mg and 10mg, both of which were well tolerated with no safety issues identified
- The equivalent human dose of 100nM nintedanib is very poorly tolerated, with significant GI side effects



# Regenerative Activity: LTI-03 Preserves Critical Progenitor Cells in the Lung (PCLS Studies. Effects 48 Hours After Administration)

- LysoTracker dye (Panel A, bright green dots) localizes to AEC2 cells, the progenitor cells of the lung, which are responsible for making new lung tissue. LTI-03 resulted in an increase in staining, meaning an increase in these critical progenitor cells
- Increases in lysoTracker staining (Panel B) also correlated with increases in surfactant protein C (pro-SPC) and ABCA3 (the pro-SPC transporter)
- Western blots (Panel C) confirm that in the IPF lung SPC levels are diminished, but that LTI-03 causes levels to increase

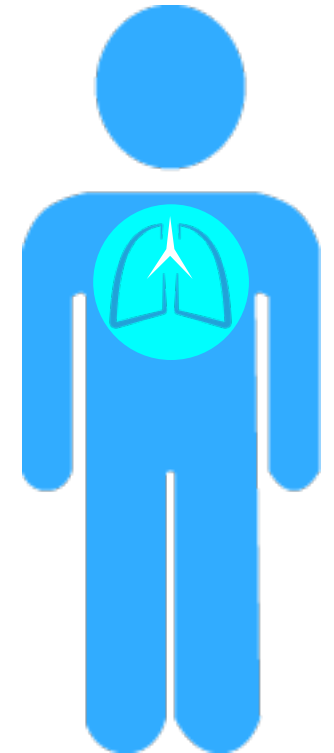


# **LTI-03: Clinical Evidence Shows Favorable Safety Profile and Potential for Efficacy**

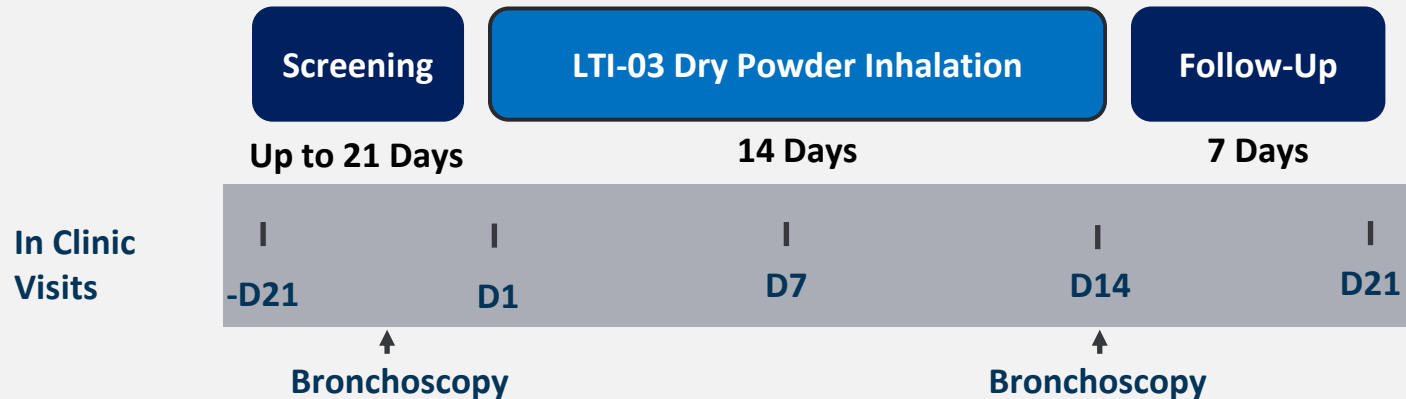
# Phase 1a Clinical Trial—at 20 mg and Below, LTI-03 Demonstrated Favorable Safety Profile and was Well Tolerated (Status: Complete)

## Healthy Human Volunteer Clinical Trial

- Objectives
  - Primary – Safety and Tolerability
  - Secondary – Pharmacokinetics
- Design
  - Single Ascending Dose (32 subjects / 3 doses)
    - Doses: 20mg, 40mg, 80mg
  - Multiple Ascending Dose (40 subjects / 5 doses)
    - Doses: 2.5mg, 5mg, 10mg, 20mg, 40mg



## Phase 1b Clinical Trial Design—Focus on Safety, Tolerability, and Biomarkers (Status: Complete)



### Study Design

- IPF diagnosis  $\leq$  3 years; no previous antifibrotic therapy w/in 2 months of baseline
- 24 patients total (18 active, 6 placebo)
  - Low (2.5mg BID) and high (5mg BID) dose cohorts, sequential daily dosing for 14 days
- Bronchoscopy at screening and Day 14
- Primary endpoint: Safety/tolerability
- Key exploratory endpoints: Biomarkers (blood, BAL, brushings)

# Robust Biomarker Evaluation for De-Risking of LTI-03

## Several Markers Linked to Lung Function

- All of the biomarkers selected for evaluation
  - ✓ Have literature suggesting their involvement
  - ✓ Are primarily found in important cell types in the IPF lung
  - ✓ Were shown in preclinical studies to be attenuated by LTI-03
- Attenuation of markers in the Phase 1b trial would demonstrate
  - ✓ That LTI-03 is reaching important cells in the deeply fibrosed lung
  - ✓ Surrogate target engagement
  - ✓ That LTI-03 is positively affecting pathogenic factors in the IPF lung

### Statistically Significant Biomarkers from Phase 1b Trial

#### Associated with Fibroblasts/myofibroblasts cell-type

Interleukin 11 (IL-11)

A predictor of prognosis and acute exacerbation in IPF patients

CXCL7

Proinflammatory and pro-fibrotic chemokine

#### Associated with Basal-like cell-type

Thymic Stromal Lymphopoietin Protein (TSLP)

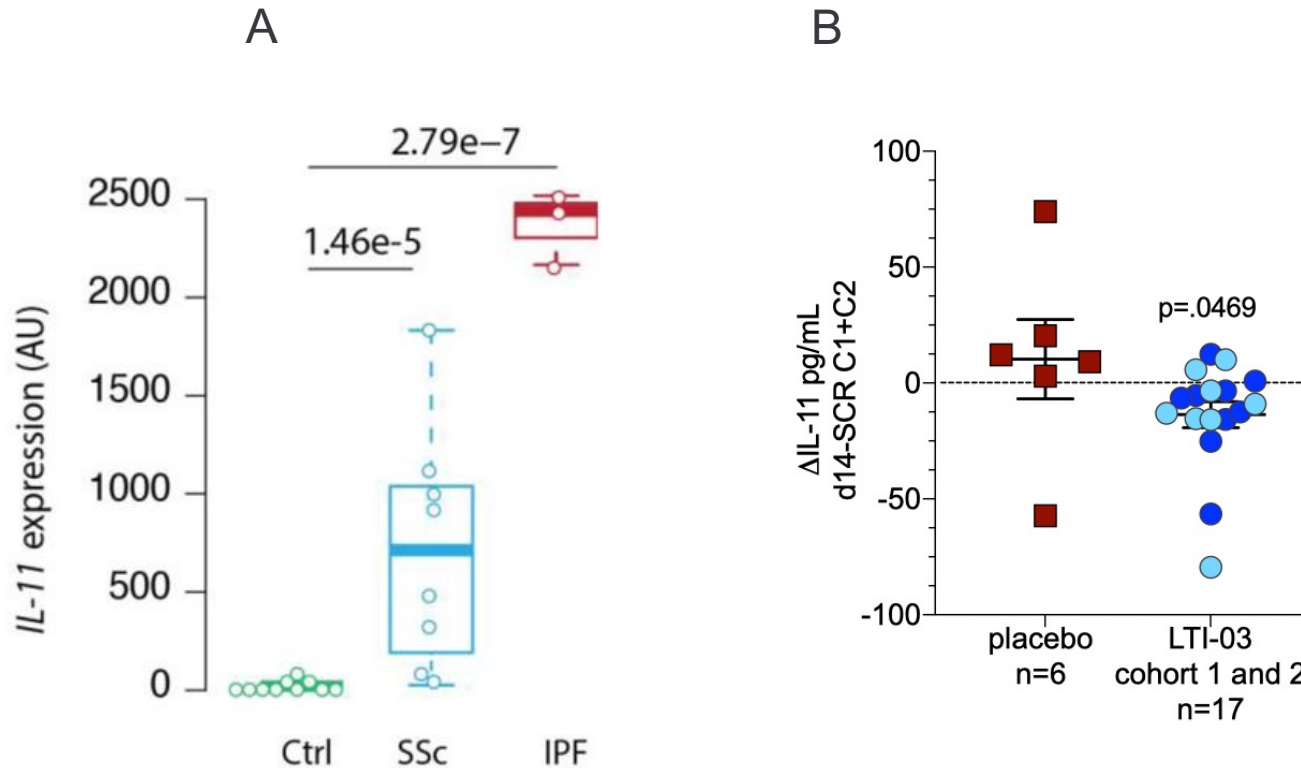
Expressed in fibroblasts and basal like epithelium of IPF UIP lesions

Galectin 7 (Gal7)

Highly expressed in Caveolin-1 deficient bronchiolized areas in the IPF lung

# IL-11: Example of Strong Evidence of LTI-03 Activity and Potential

- A. IL-11 is highly expressed in IPF\*
- B. LTI-03 significantly reduced IL-11 in IPF patients (vs placebo)
- C. IL-11 is a predictor of prognosis and acute exacerbation in IPF patients\*

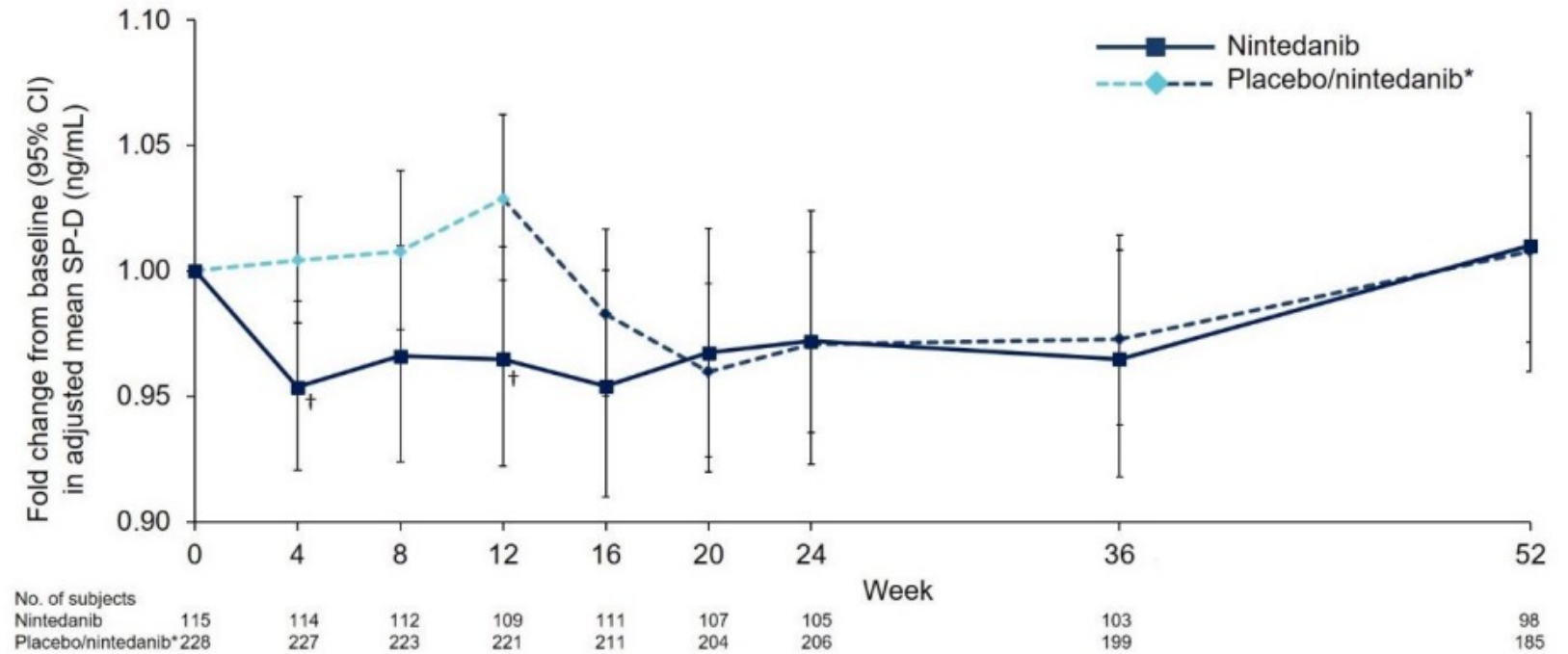


C

IPF and investigated its predictive significance for survival and AE occurrence. The serum IL-11/%FVC value was an independent predictor of prognosis and AE occurrence in patients with IPF, and the IL-11 level appeared to show pathophysiologic value in IPF.

# Surfactant Protein D (SPD) is an important biomarker for the approved IPF drug Ofev<sup>®</sup>\* and now for LTI-03

- SPD is an indicator of epithelial cell health, an important cell type for proper lung function
- SPD has been significantly linked to decline in lung function
- SPD was reduced by 4% by Ofev over 12 weeks in the INMARK clinical trial
- LTI-03 (5 mg BID) decreased SPD by 5% over two weeks in the Phase 1b trial






No. of subjects	0	4	8	12	16	20	24	36	52
Nintedanib	115	114	112	109	111	107	105	103	98
Placebo/nintedanib*	228	227	223	221	211	204	206	199	185

\*Subjects received placebo (blinded) for 12 weeks followed by nintedanib (open-label) for 40 weeks.  
 †p<0.05 for adjusted difference in change from baseline between groups.

Nintedanib versus placebo. Fold changes from baseline in SP-D at week 12 corresponded to a 4% decrease and 3% increase in the nintedanib and placebo groups, respectively (ratio 0.94 [95% CI: 0.89, 0.99]; p=0.024).



# Surfactant Protein D (SPD) Reduction<sup>1, 2, 3</sup>

Company	Trial	Compound	Percent Reduction	Duration of Treatment
	Phase 1b	LTI-03	5%	2 weeks
	INMARK (P2) <sup>1</sup>	Nintedanib	4%	12 weeks
	CAPACITY (P3) <sup>2</sup>	Pirfenidone	5%	12 weeks

<sup>1</sup>Maher, T. M., Gisli Jenkins, R., Cottin, V., Nishioka, Y., Noth, I., Selman, M., Song, J. W., Ittrich, C., Diefenbach, C., Stowasser, S., & White, E. S. (2024). Circulating biomarkers and progression of idiopathic pulmonary fibrosis: data from the INMARK trial. *ERJ Open Research*, 10(4). <https://doi.org/10.1183/23120541.00335-2023>

<sup>2</sup>Ikeda, K., Chiba, H., Nishikiori, H., Azuma, A., Kondoh, Y., Ogura, T., Taguchi, Y., Ebina, M., Sakaguchi, H., Miyazawa, S., Suga, M., Sugiyama, Y., Nukiwa, T., Kudoh, S., & Takahashi, H. (2020). Serum surfactant protein D as a predictive biomarker for the efficacy of pirfenidone in patients with idiopathic pulmonary fibrosis: a post-hoc analysis of the phase 3 trial in Japan. *Respiratory Research*, 21(1), 1

<sup>3</sup>The INMARK and CAPACITY trials were not conducted by the Company, no trials have been conducted comparing these compounds and the referenced trials had different trial designs, patient enrollment criteria and treatment regimens. In addition, the applicable measurements for the referenced trials were observed over different time periods and using different assays. As a result, the data from these trials may not be directly comparable.

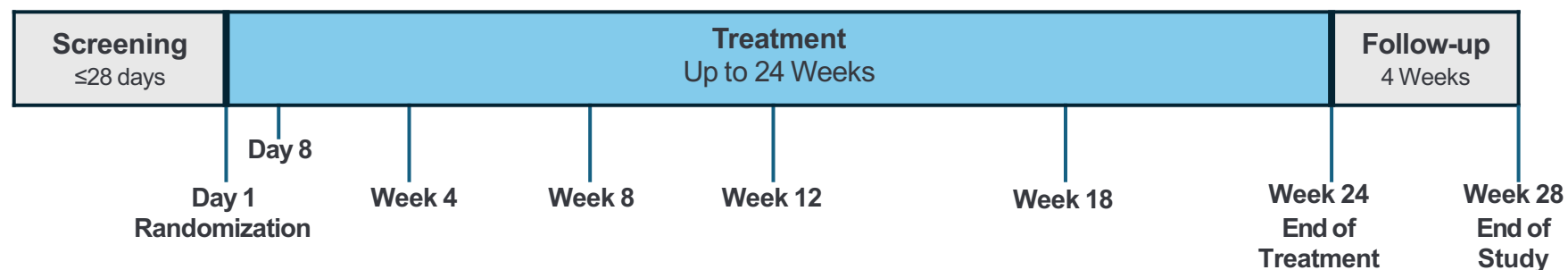
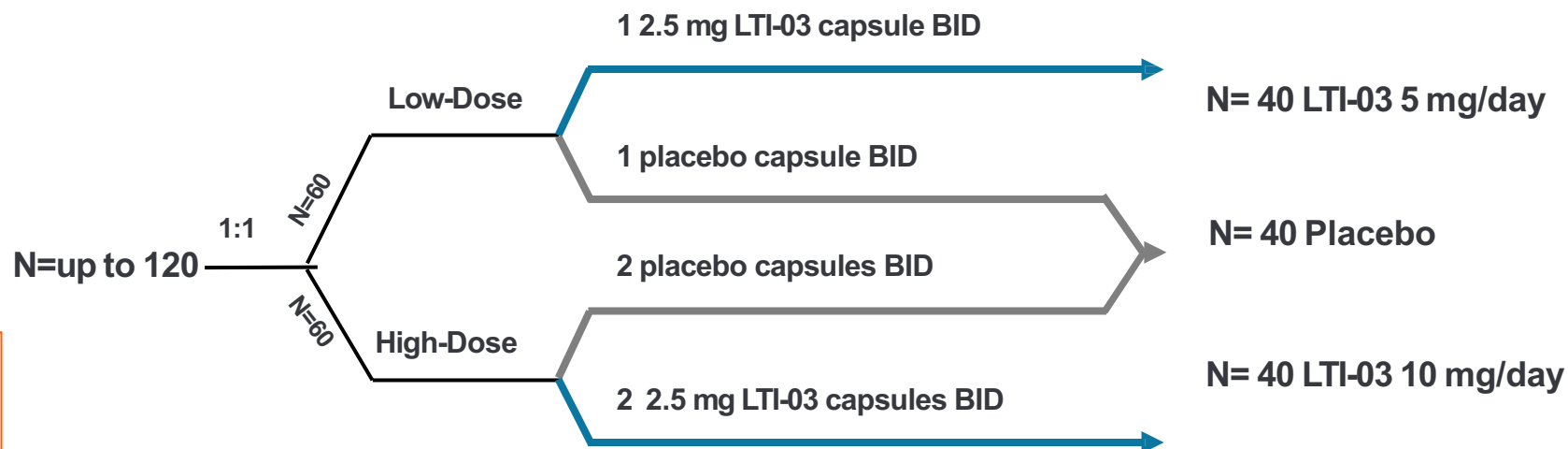
# LTI-03 – Phase 2 Trial Measuring Lung Function

## Primary endpoints

Safety and tolerability measured by incidence of treatment emergent adverse events

Efficacy of inhaled LTI-03 measured by:

- Change from baseline in FVC in mL
- Change from baseline in percent predicted FVC
- Change from baseline in lung fibrosis measure by HRCT

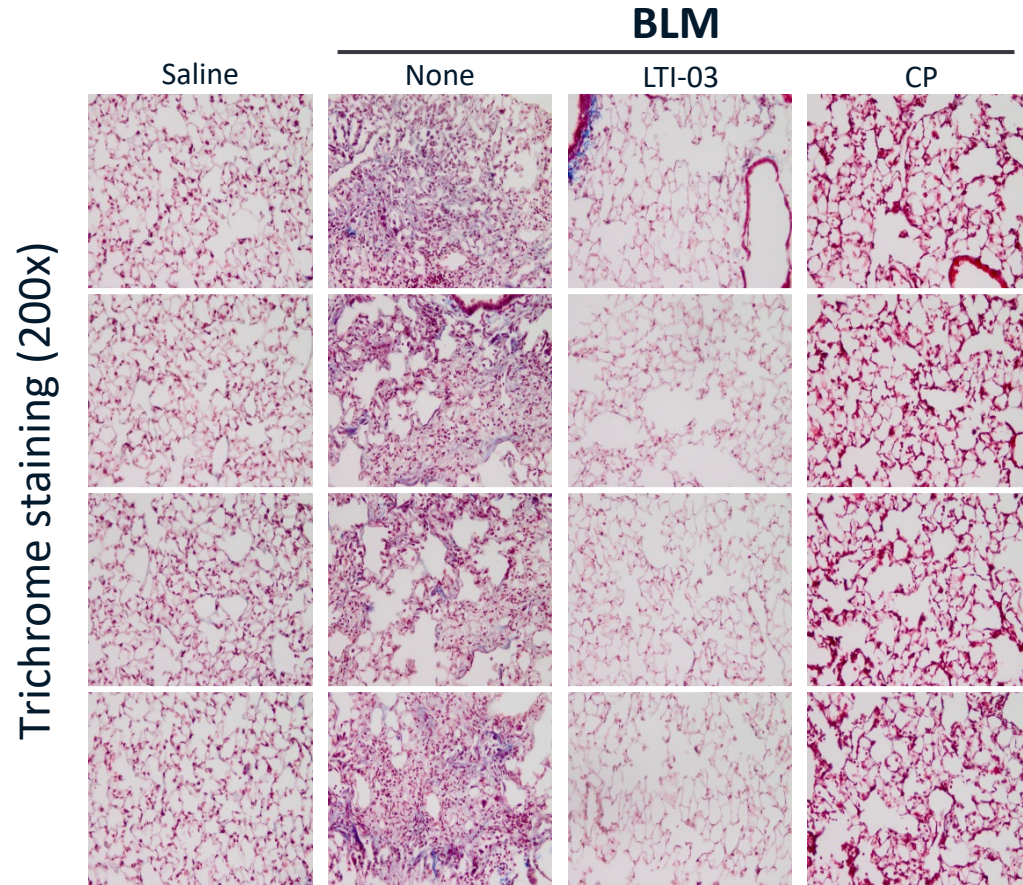




Nasdaq: RNTX

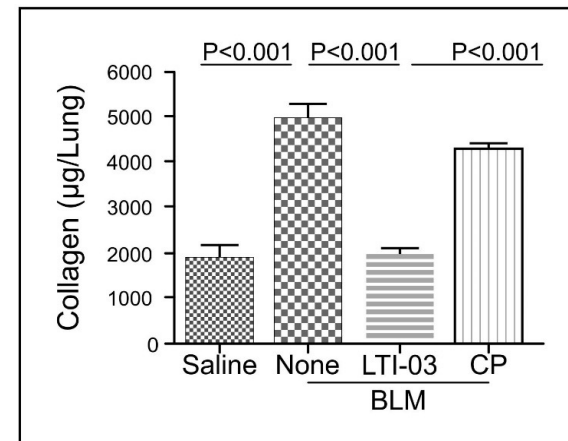
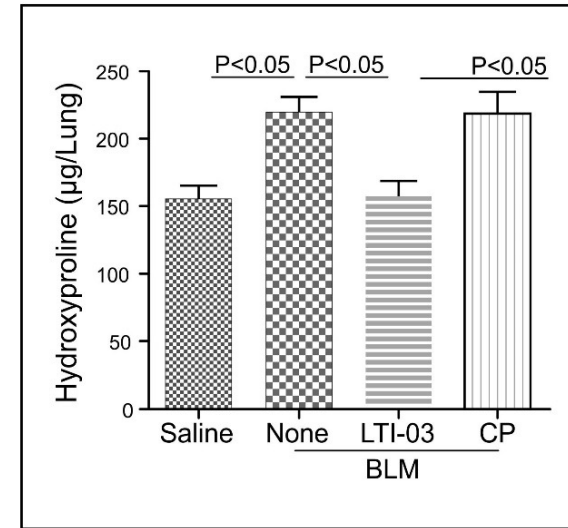
# Appendix

# Demonstrated Anti-Fibrotic Properties in the 21-day Bleomycin Mouse Model of IPF

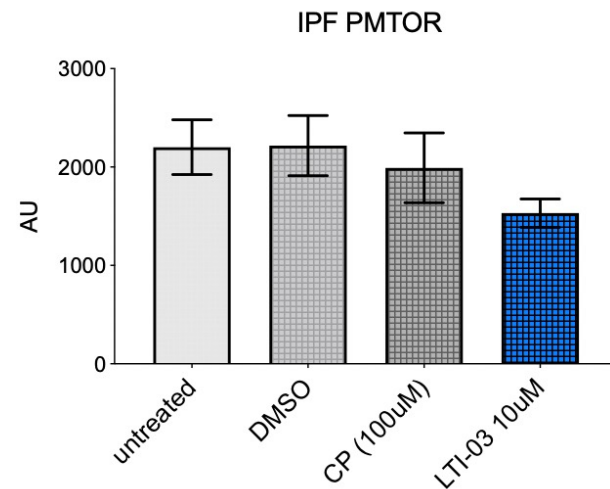
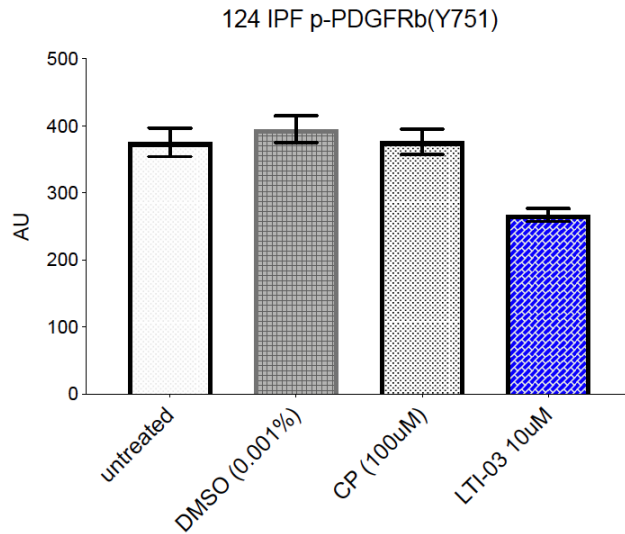


***The bleomycin mouse model is an established murine model for characterizing and assessing the impact of novel IPF therapies***

Fibrotic biomarkers



# LTI-03 Attenuates the overexpression of many profibrotic proteins



## RTK and associated signaling ↓

ALK(D5F3)	*
p-ALK(3B4)(Y1586)	***
c-Jun	*
p-c-Myc(T58)	**
Herb2/ErbB3	***
p-EGFR(Y1173)(53A5)	***
p-MEK (1/2)	****
p44/42 MAPK (ERK1/2)	*
p-PDK1(S241)	****
<b>p-PDGFRb(Y761)</b>	****
p-RafB(S445)	****
p-Ret(Y905)	**
Stat5a	*
p-Stat5(Y694)	***
PI3Kp110a	**
PTEN	*
p-SRC	****
SRC-1	***
YAP	**

## Metabolic signaling ↓

AMPKa	****
p-AMPKb1(S108)	****
Deptor	**
LDHA	**
<b>p-mTOR</b>	****
p-Raptor	****
Raptor	**
p-Tuberin	****

## Invasion associated markers ↓

TWIST2	*
Wnt5ab	**

## HDACs ↓

HDAC4	*
HDAC6	***

RPPA assay, Shixia Huang; Baylor College of Medicine; Note(s) Data expressed as mean values and SDs. \* $p < .05$ ; \*\*  $p < .01$ ; \*\*\* $p < .001$ ; \*\*\*\* $p < .0001$

# Cohort Two Biomarker Results

Biomarker	Positive Trend C2	Statistically Significant (p<0.05) C2	Positive Trend C1+C2	Statistically Significant (p<0.05) C1+C2	dose dependency
<b>Fibroblasts/ myofibroblasts</b>					
COL1A1	✓		✓		✓
IL-11	✓		✓	✓	
CXCL7	✓	✓	✓	✓	✓
pSMAD/ tSMAD					
<b>Basal-like cells</b>					
TSLP	✓	✓	✓	✓	✓
GAL7	✓	✓	✓	✓	✓
<b>Alveolar epithelial health</b>					
SPD	✓		✓		✓
<b>Inflammation/ safety</b>					
%pAKT	✓	N/A	✓	N/A	N/A

# LTI-01: the First Drug Being Developed for Loculated Pleural Effusion

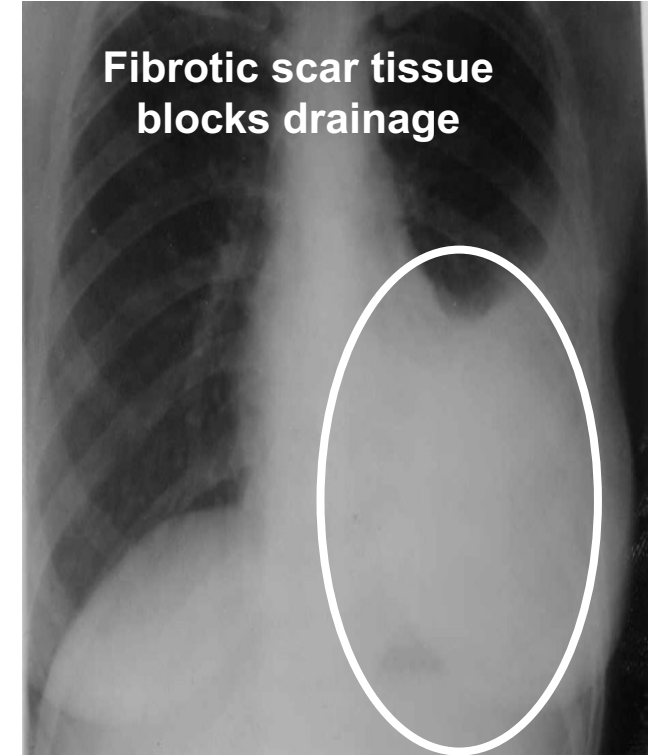
# There Are No Approved Drug Treatments for Loculated Pleural Effusion

- Loculated pleural effusion, or LPE occurs when fibrotic scar tissue forms in the pleural cavity, preventing effective drainage of fluid
- LPE is a frequent pneumonia complication in the elderly with a ~20% mortality rate
- LPE is managed with tPA/DNase (off-label) and/or surgery (costly and invasive)
- Surgery can be effective but can also result in lengthy hospital stays. This is why off-label fibrinolytics is widely regarded as first line therapy
- Off-label therapy
  - Not FDA approved for LPE
  - Risk of intrapleural hemorrhage
  - Problematic dosing (at least twice daily, 12 hours apart)

Healthy Lungs

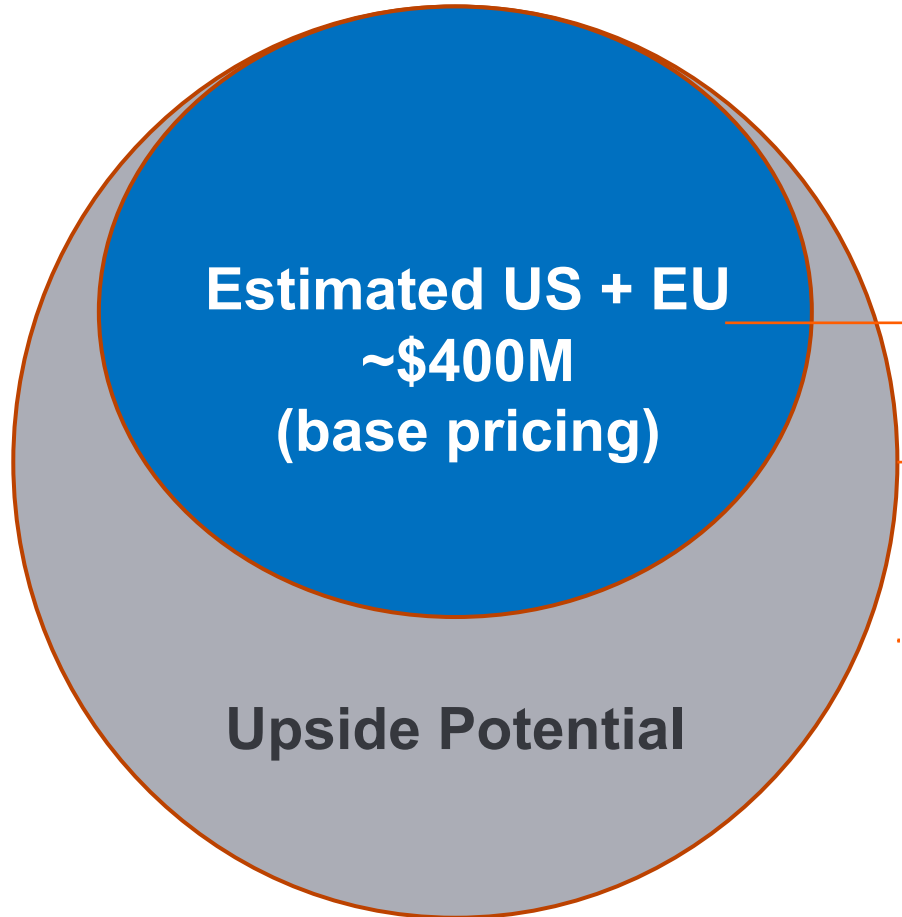


Loculated Pleural Effusion



# Sizeable US and EU Commercial Opportunity with Potential Upside

Addressable market



## Current US and EU Opportunity

- 30,000 US fibrinolytic patients
- Up to 30,000 additional US LPE patients
- tPA/DNase priced at \$6,700 per patient in US
- Estimate similar EU market opportunity to US market

**Key Catalyst: Substitution of tPA/DNase with on-label therapeutic**

## Upside Market Potential in the US and EU

- Premium Pricing
- Ability to drive beneficial clinical and economic outcomes

**Key Catalyst: On-label therapy with clear efficacy, safety and dosing benefits**

+ Japan partnership with  TAIHO PHARMA

Source: Management estimates, industry publications and MME market access research study for Rein Tx