

Phase 2 TORREY Study Topline Results

December 6, 2022

Forward Looking Statements

This presentation contains forward-looking statements. All statements other than statements of historical facts contained in this presentation, including statements regarding the potential of seralutinib to serve patients with pulmonary arterial hypertension (PAH), the potential for seralutinib to be differentiated from other PAH therapies, plans to complete regulatory interactions regarding the Phase 2 TORREY study and the timing thereof, plans to commence a global registrational Phase 3 Program in PAH and the timing thereof, and plans to commence a development program in World Health Organization Group 3 pulmonary hypertension (PH) and the timing thereof, are forward-looking statements.

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Presenters for Today's Call

Gossamer Bio		
Faheem Hasnain	Co-Founder, Chairman, & Chief Executive Officer	
Richard Aranda, MD	Chief Medical Officer	
Robert Roscigno, PhD	VP, Clinical Development	
Larry Zisman, MD	Sr Dir, Clinical Development	
Ed Parsley, DO	Consultant Pulmonologist	
Matt Cravets	SVP, Biometrics	
Laura Carter, PhD	Chief Scientific Officer	
Caryn Peterson	EVP, Regulatory Affairs	
Bryan Giraudo	COO & CFO	

Guest Speakers



Ardeschir Ghofrani, MD* Professor of Pulmonary Vascular Research, Justus Liebig University; Head of the Pulmonary Hypertension Division, University Hospital Giessen



Raymond Benza, MD*

Professor of Medicine, Division of
Cardiovascular Medicine,
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TORREY Study Topline Results Highlights

- Met primary endpoint of statistically significant reduction in PVR in a heavily treated, prevalent study population
- Functional outcome, six-minute walk test, favored seralutinib, though study was neither powered nor designed for this endpoint
- Statistically significant reduction in NT-proBNP, coupled with significant changes observed in right heart parameters[†]
- Well tolerated, avoiding side effect profile associated with systemic imatinib in PAH



TORREY Study Topline Results Highlights, Cont.

 Consistent benefit across pre-specified sub-groups in favor of seralutinib with enhanced effects in patients with more severe disease at baseline§

Overall Stud	y Population	Function	al Class III	REVEAL 2.0 R	isk Score ≥ 6
PVR	6MWD	PVR	6MWD	PVR	6MWD
-14%*	+6.5m	-21%*	+37.3m*	-23%*	+21.9m
p = 0.0310	p = 0.5972	p = 0.0427	p = 0.0476	p = 0.0134	p = 0.2482

Consistently favorable results for hemodynamic and ECHO endpoints

^{* =} p-value \leq 0.05. All p-values in this presentation are nominal, aside from primary endpoint (Overall study population delta in PVR). § At baseline, as determined by Functional Class and REVEAL 2.0 Risk Score (pre-specified subgroups).

Disease Overview and Available Treatments



PAH is a Rare and Progressive Disease

- PAH is a rare, progressive disease^{1,2} with an estimated worldwide prevalence of 5-25 cases per million per year³
- PAH has no known cure, is associated with poor survival, and has a debilitating impact on the health-related quality of life of patients and caregivers^{1,2,4,5}
- PAH is characterized by vascular remodeling^{6,7}
 - cellular overgrowth, narrowing and thickening of pulmonary arterioles, and formation of pathologic lesions
 - underlying pathologic mechanisms include inflammation, proliferation, and fibrosis
 - leads to obstructed pulmonary blood flow, increased PVR, ultimately right heart failure and death
- Current treatment approaches are primarily vasodilatory⁸⁻¹⁰
- A significant unmet need exists for new therapies that address the underlying pathological mechanisms of PAH

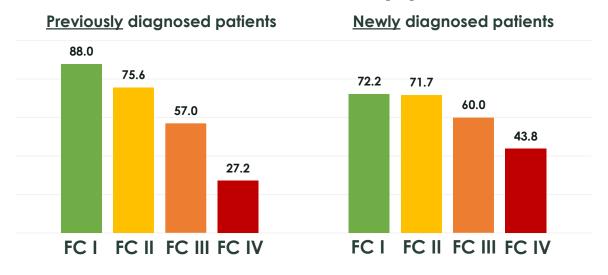


Functional Class Correlates With Risk Status & Predicts Survival

- Analysis of REVEAL Registry shows that 5-year survival remains poor despite progress in PAH-specific therapy options and improved patient support strategies
- Functional Class is predictive of survival

WHO Functional Assessment for Pulmonary Hypertension			
Class I	Class II	Class III	Class IV
No limitation of physical activity	Slight limitation of physical activity	Marked limitation of physical activity	Inability to carry out physical activity

5 Year Survival* (%)



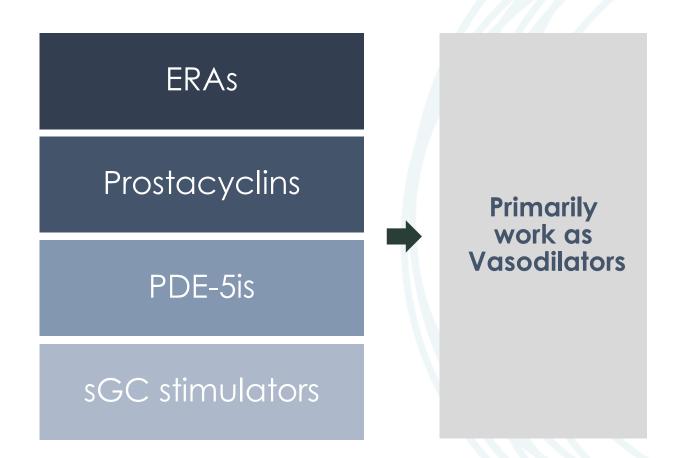


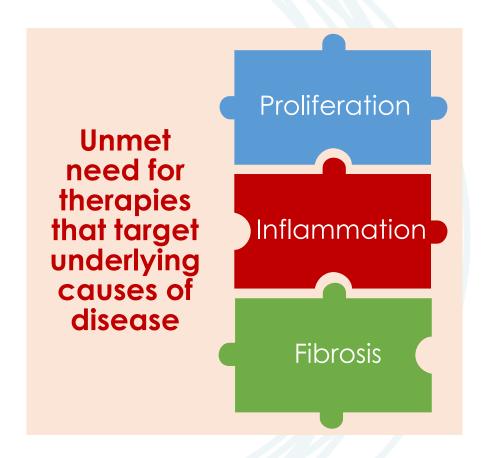
The Clinical Goal of PAH Therapy is to Achieve Low Risk Status¹

- Low risk status usually associated with¹
 - Good exercise capacity
 - Good quality of life
 - Good right ventricular function
 - Low mortality risk
- Patients categorized as low risk¹
 - Have est. 1-year mortality <5%
 - Present with non-progressive disease in WHO-FC I/II with 6MWD >440m and no signs of clinically relevant RV dysfunction



What Do Currently Available Therapies Do?



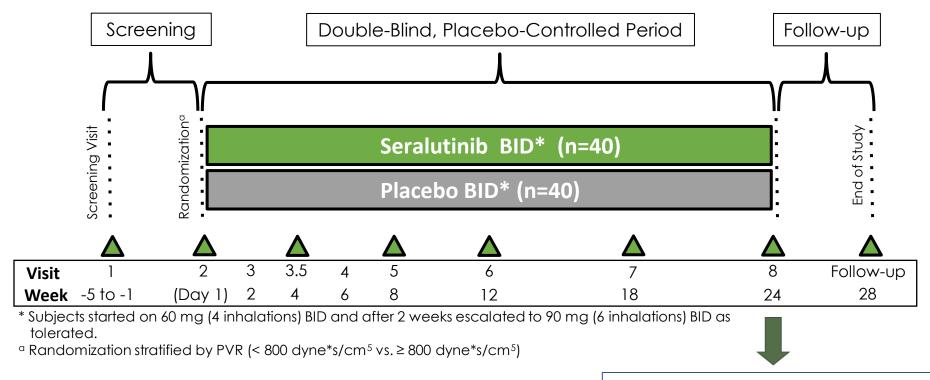




Study Design and Baseline Characteristics

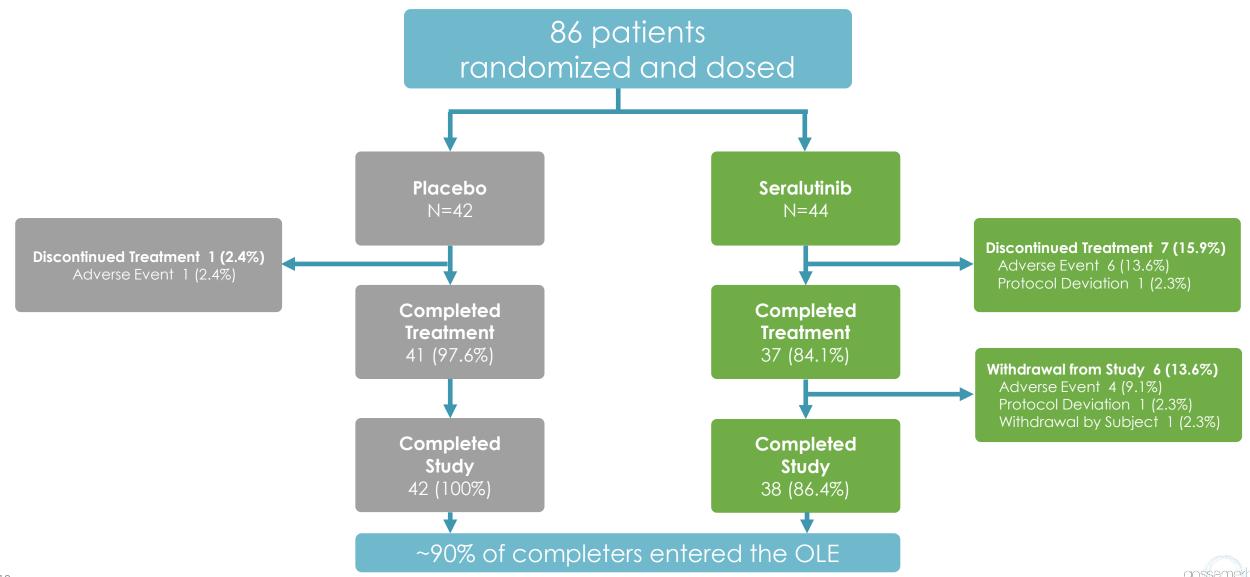


Study Design



After completing the Week 24 visit, subjects have the option to roll into an open-label extension study

Patient Disposition



Baseline Demographics (ITT Population)

Characteristic	Placebo (N=42)	Seralutinib (N=44)	Total (N=86)
Age (years) – mean (SD)	49.5 (11.81)	48.3 (12.70)	48.8 (12.22)
Sex - n (%)			
Female	38 (90.5)	40 (90.9)	78 (90.7)
Male	4 (9.5)	4 (9.1)	8 (9.3)
Race - n (%)			
White	37 (88.1)	37 (84.1)	74 (86.0)
Black or African American	1 (2.4)	0	1 (1.2)
Asian	2 (4.8)	4 (9.1)	6 (7.0)
Other	2 (4.8)	3 (6.8)	5 (5.8)
Ethnicity – n (%)			
Hispanic or Latino	6 (14.3)	8 (18.2)	14 (16.3)
Not Hispanic or Latino	34 (81.0)	36 (81.8)	70 (81.4)
Not reported	2 (4.8)	0	2 (2.3)
Region - n (%)			
North America	30 (71.4)	29 (65.9)	59 (68.6)
Western Europe	10 (23.8)	11 (25.0)	21 (24.4)
Asia Pacific	1 (2.4)	4 (9.1)	5 (5.8)
Eastern Europe	1 (2.4)	0	1 (1.2)

Baseline Background PAH Medication Use (ITT Population)

Characteristic	Placebo (N=42)	Seralutinib (N=44)	Total (N=86)
Number of background therapies – n (%)			
1	2 (4.8)	1 (2.3)	3 (3.5)
2	16 (38.1)	18 (40.9)	34 (39.5)
3	24 (57.1)	25 (56.8)	49 (57.0)
Prostacyclin/Prostacycin Receptor Agonist use – n (%)			
None	13 (31.0)	15 (34.1)	28 (32.6)
Monotherapy	1 (2.4)	1 (2.3)	2 (2.3)
Double therapy	4 (9.5)	3 (6.8)	7 (8.1)
Triple therapy	24 (57.1)	25 (56.8)	49 (57.0)
Parenteral Prostacyclin	19 (45.2)	19 (43.1)	38 (44.2)
Oral	10 (23.8)	10 (22.7)	20 (23.3)

Baseline Disease Characteristics (ITT Population)

Characteristic	Placebo (N=42)	Seralutinib (N=44)	Total (N=86)
Age at PAH diagnosis (years) – mean (SD)	41.2 (11.65)	40.7 (15.84)	40.9 (13.87)
Years since PAH diagnosis – mean (SD)	8.78 (7.218)	8.07 (7.074)	8.41 (7.111)
PAH classification – n (%)			
Idiopathic	22 (52.4)	20 (45.5)	42 (48.8)
Heritable	5 (11.9)	10 (22.7)	15 (17.4)
Associated with:			
CTD	11 (26.2)	6 (13.6)	17 (19.8)
Anorexigen use	0	1 (2.3)	1 (1.2)
Methamphetamine use	4 (9.5)	4 (9.1)	8 (9.3)
Corrected congenital shunts	0	3 (6.8)	3 (3.5)
WHO FC - n (%)			
Class II	20 (47.6)	30 (<mark>68.2</mark>)	50 (58.1)
Class III	22 (<mark>52.4</mark>)	14 (31.8)	36 (41.9)
REVEAL 2.0 Risk Score ≥ 6 – n (%)	17 (40.5)	20 (45.5)	37 (43.0)
PVR (dyne*s/cm ⁵) – mean (SD)	661.3 (164.91)	675.8 (240.35)	<mark>668.7</mark> (205.90)
6MWD (m) – mean (SD)	407.1 (107.02)	408.6 (75.11)	<mark>407.9</mark> (91.54)
NT-proBNP (ng/L) – mean (SD)	645.6 (1158.75)	611.0 (714.58)	628.3 (956.83)

Abbreviations: 6MWD, six-minute walk distance; CTD, connective tissue disease; FC, functional class; NT-proBNP, N-terminal pro B-type natriuretic peptide; PAH, pulmonary arterial hypertension; PVR, pulmonary vascular resistance; WHO, World Health Organization; ITT, Intention-to-treat.



Baseline Demographics and Disease Characteristics by Baseline WHO FC (ITT Population)

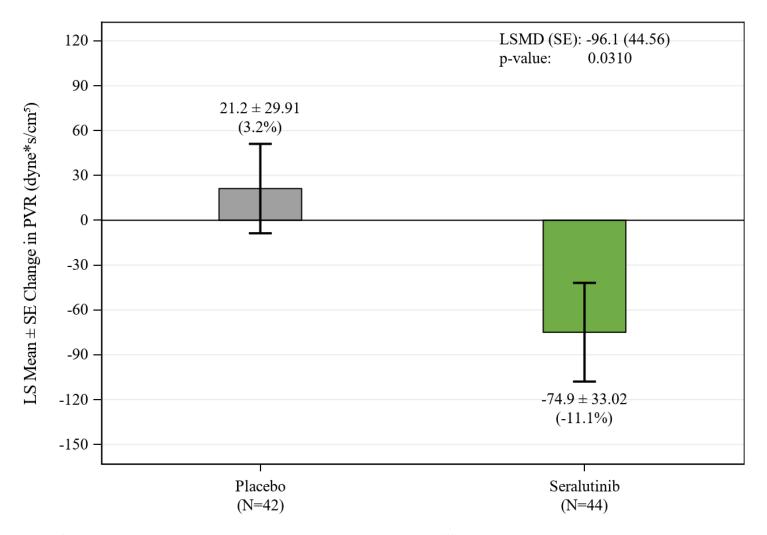
	Baseline WHO FC Class II		
Characteristic	Placebo (N=20)	Seralutinib (N=30)	Total (N=50)
Age (years) – mean (SD)	47.6 (11.69)	47.7 (13.42)	47.7 (12.63)
Female – n (%)	19 (95.0)	27 (90.0)	46 (92.0)
Race, White – n (%)	19 (95.0)	24 (80.0)	43 (86.0)
Region, North America – n (%)	13 (65.0)	20 (66.7)	33 (66.0)
Years since PAH diagnosis – mean (SD)	9.60 (7.262)	8.40 (6.961)	8.88 (7.034)
PAH classification – n (%)			
Idiopathic	11 (55.0)	16 (53.3)	27 (54.0)
Heritable	4 (20.0)	6 (20.0)	10 (20.0)
Associated with CTD	5 (25.0)	5 (16.7)	10 (20.0)
REVEAL 2.0 Risk Score ≥ 6 - n (%)	4 (20.0)	11 (36.7)	15 (<mark>30.0</mark>)
PVR (dyne*s/cm ⁵) – mean (SD)	638.3 (161.85)	689.9 (265.72)	669.3 (229.34)
6MWD (m) – mean (SD)	455.5 (63.96)	425.5 (62.98)	<mark>437.5</mark> (64.45)
NT-proBNP (ng/L) – mean (SD)	406.8 (798.39)	609.9 (715.31)	<mark>525.3</mark> (749.58)
On 3 background therapies – n (%)	11 (55.0)	18 (60.0)	29 (58.0)
ERA + PDE-5i + Prostacyclins/PRA	8 (40.0)	16 (53.3)	24 (48.0)
ERA + sGC + Prostacyclins/PRA	3 (15.0)	2 (6.7)	5 (10.0)

Baseline WHO FC Class III			
Placebo (N=22)	Seralutinib (N=14)	Total (N=36)	
51.1 (11.94)	49.4 (11.40)	50.4 (11.60)	
19 (86.4)	13 (92.9)	32 (88.9)	
18 (81.8)	13 (92.9)	31 (86.1)	
17 (77.3)	9 (64.3)	26 (72.2)	
8.02 (7.263)	7.36 (7.527)	7.76 (7.266)	
11 (50.0)	4 (28.6)	15 (41.7)	
1 (4.5)	4 (28.6)	5 (13.9)	
6 (27.3)	1 (7.1)	7 (19.4)	
13 (59.1)	9 (64.3)	22 (<mark>61.1</mark>)	
682.2 (168.62)	645.7 (179.29)	668.0 (171.25)	
363.2 (120.05)	372.4 (87.97)	<mark>366.8</mark> (107.43)	
873.0 (1403.06)	613.3 (742.17)	<mark>773.7</mark> (1187.34)	
13 (59.1)	7 (50.0)	20 (55.6)	
10 (45.5)	6 (42.9)	16 (44.4)	
3 (13.6)	1 (7.1)	4 (11.1)	

Primary
Endpoint:
Change From
Baseline in PVR

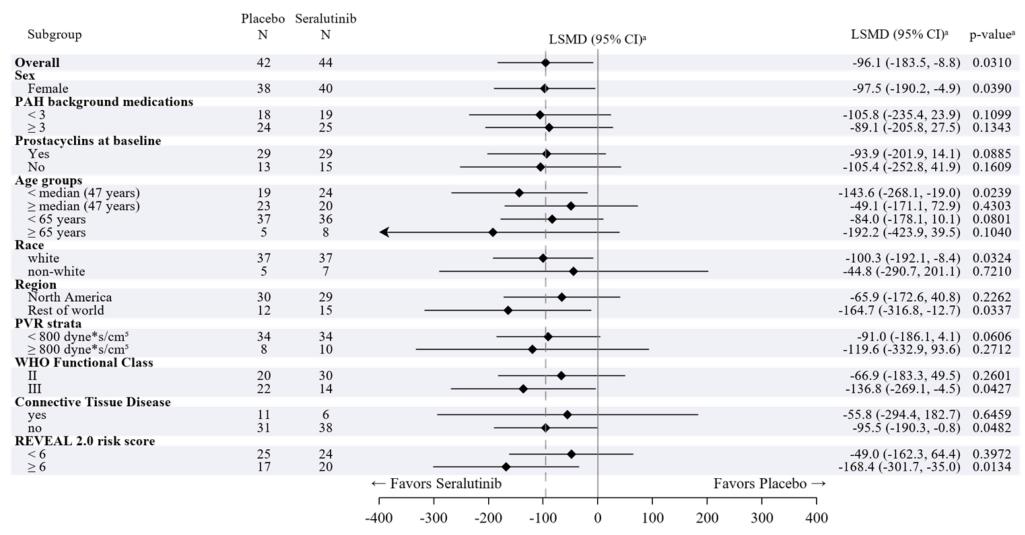


Primary Endpoint: Seralutinib Significantly Reduced PVR at Week 24 (ITT Population)





Seralutinib Consistently Reduced PVR Across All Pre-Specified Sub-Groups (ITT Population)



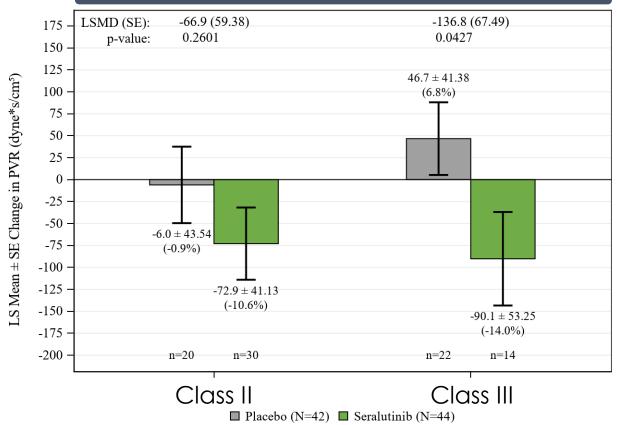
Abbreviations: ANCOVA, analysis of covariance; FC, functional class; PVR, pulmonary vascular resistance; WHO, World Health Organization; LSMD, least squares mean difference.

^a Based on an ANCOVA model with multiple imputation.

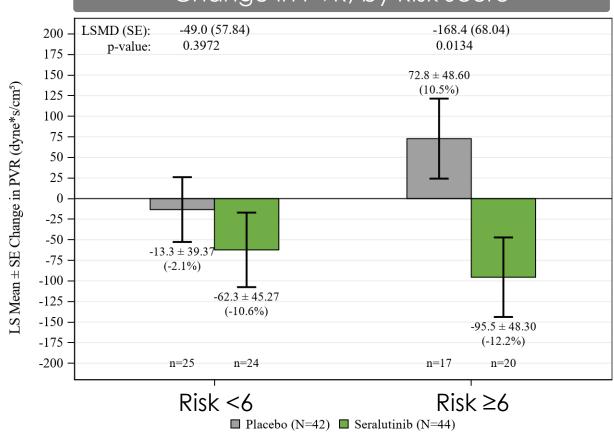
Source: Data on file.

Seralutinib's Effect on PVR was More Pronounced in Patients with More Severe Disease at Baseline (ITT Population)

WHO Functional Class Change in PVR, by Functional Class



REVEAL 2.0 Risk ScoreChange in PVR, by Risk Score

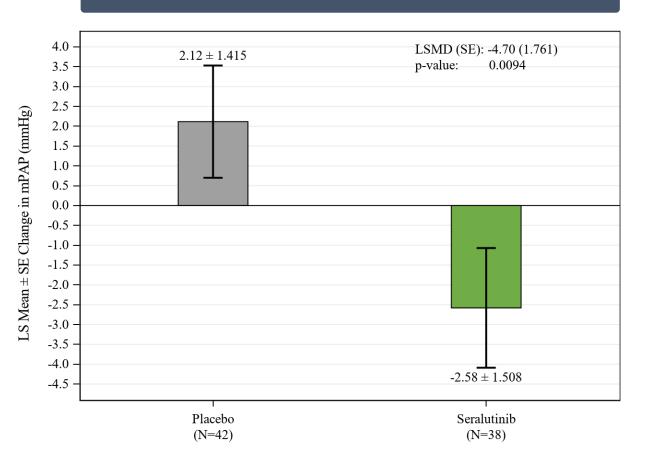


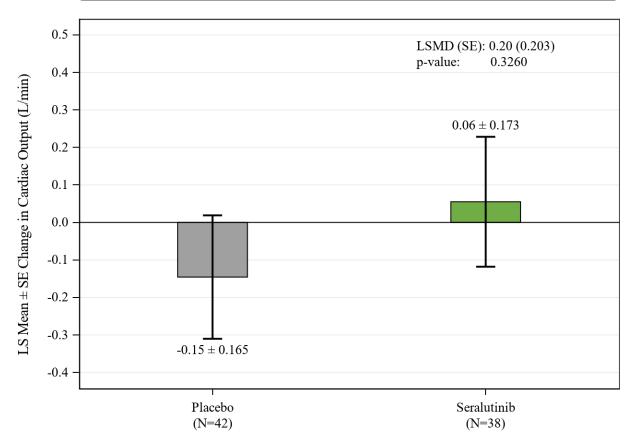
Abbreviations: ANCOVA, analysis of covariance; LS, least squares; LSMD, least squares mean difference; PVR, pulmonary vascular resistance; WHO, World Health Organization. Note: Based on ANCOVA modelling.

Observed Reduction in PVR Mainly Driven by Reduction in mPAP

Change in mPAP from Baseline to Week 24

Change in CO from Baseline to Week 24





Abbreviations: ANCOVA, analysis of covariance; CO, cardiac output; LS, least squares; LSMD, least squares mean difference; mPAP, mean pulmonary arterial pressure; RHC, right heart catheterization; PVR, pulmonary vascular resistance.

Note: Based on ANCOVA modelling using observed cases.

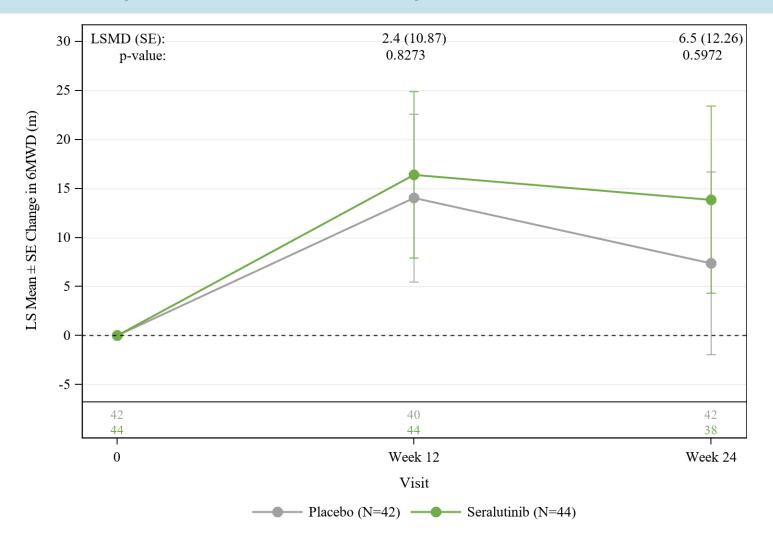
Source: Data on file.



Secondary Endpoint: Change From Baseline in 6MWD

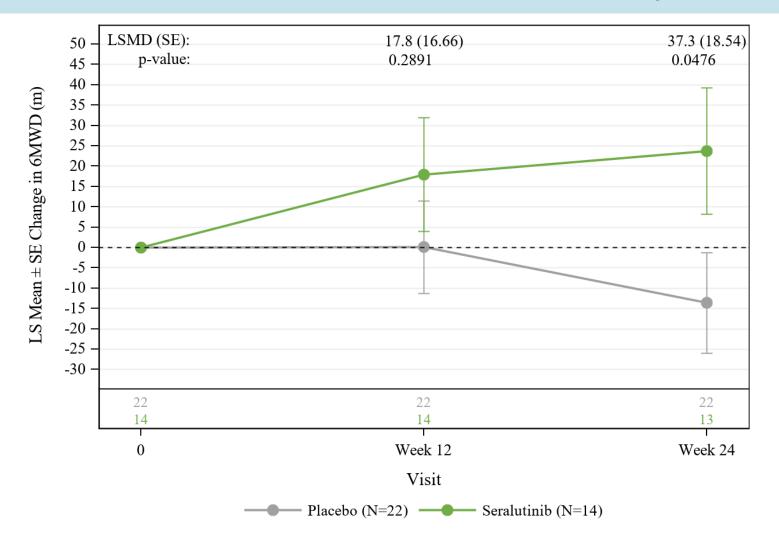


Secondary Endpoint: Change in 6MWD from Baseline to Each Visit (ITT Population)





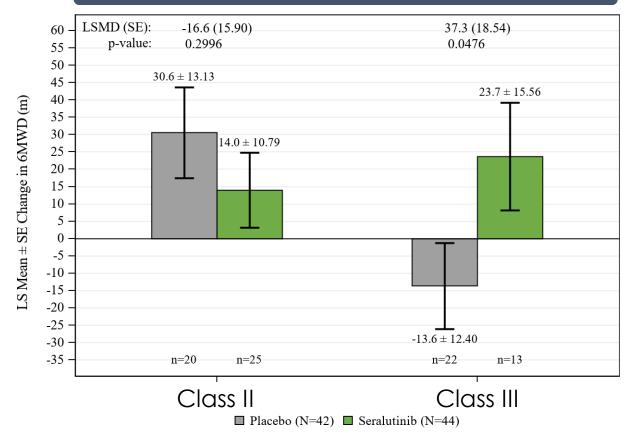
Secondary Endpoint: Change in 6MWD from Baseline to Each Visit for Baseline FC III Patients (ITT Population)



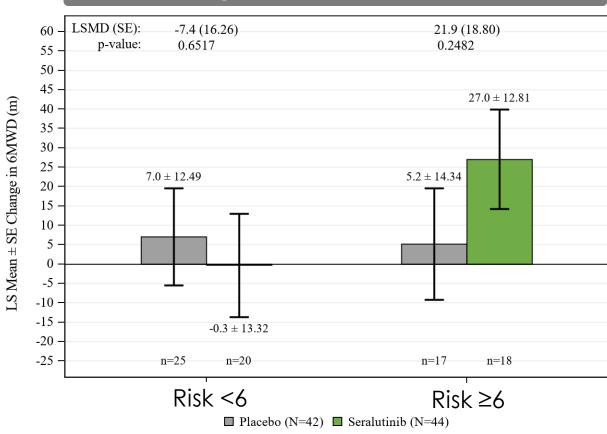


Change in 6MWD by Functional Class and REVEAL 2.0 Risk Score (ITT Population)

Functional Class Change in 6MWD, by Functional Class



REVEAL 2.0 Risk ScoreChange in 6MWD, by Risk Score



Abbreviations: 6MWD, six-minute walk distance; FC, functional class; LS, least squares; LSMD, least squares mean difference; MMRM, mixed-effects model with repeated measures; WHO, World Health Organization.

Note: Based on MMRM modelling.

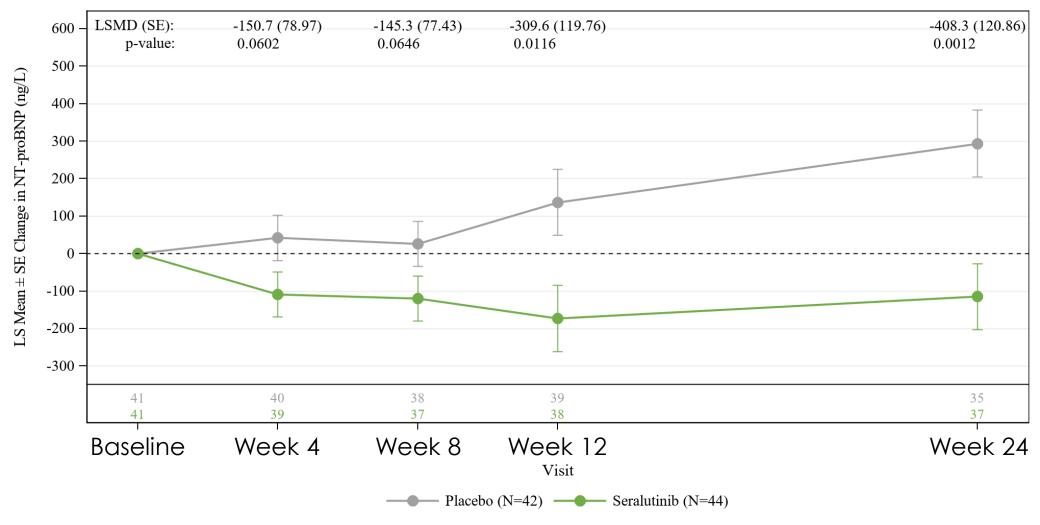
Source: Data on file.



Exploratory
Endpoints &
REVEAL 2.0 Risk
Score Change



Seralutinib Treatment Led to Statistically Significant Reduction in NT-proBNP (ITT Population)



Abbreviations: FC, functional class; LS, least squares; LSMD, least squares mean difference; MMRM, mixed-effects model with repeated measures; NT-proBNP, N-terminal pro B-type natriuretic peptide.

Note: Based on a MMRM model.

Source: Data on file.



Centrally-Read RHC and ECHO Results at Week 24 Consistently Favored Seralutinib (ANCOVA – Observed Cases)

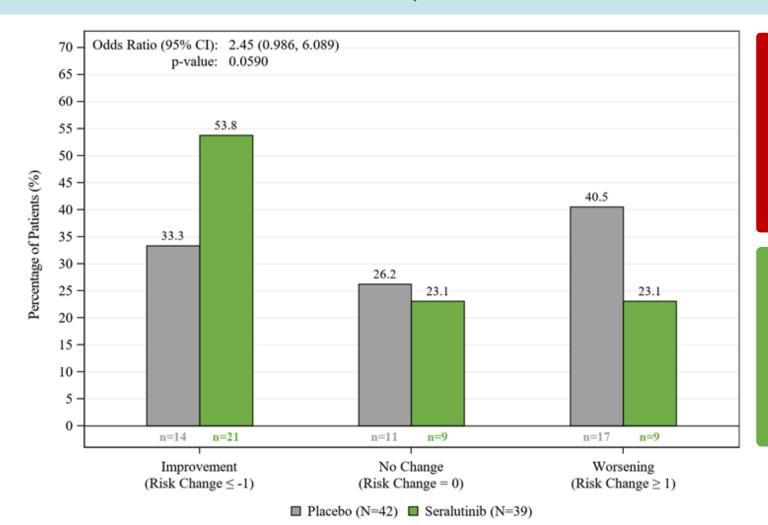
Endpoint	LS Mean Difference (95% CI)	Statistically Significant Result Favoring Seralutinib (p ≤ 0.05)	Point Estimate Favoring Seralutinib	p-value
Right Atrium Area (cm²)	-1.99 (-3.783, -0.206)			0.0293*
RV Free Wall Strain (%)	-2.64 (-5.172, -0.098)			0.0420*
PA Compliance (mL/mmHg)	0.22 (0.009, 0.423)			0.0410*
RV Systolic Pressure (mmHg)	-8.10 (-13.877, -2.317)			0.0067*
PA Systolic Pressure (mmHg)	-6.98 (-12.774, -1.187)			0.0189*
PA Diastolic Pressure (mmHg)	-3.43 (-6.211, -0.643)			0.0165*
RV Fractional Area Change	2.62 (-1.405, 6.652)			0.1983
PVR index (dyne*s/cm ⁵ /m ²)	-160.42 (-333.970, 13.138)			0.0695
mRAP (mmHg)	-0.99 (-2.350, 0.367)			0.1503
Stroke Volume Index (mL/m²)	2.19 (-0.917, 5.299)			0.1644
Cardiac Index (L/min/m²)	0.13 (-0.100, 0.359)			0.2658

^{*} $p \le 0.05$.

Abbreviations: mRAP, mean right atrial pressure; PA, pulmonary artery; PVR, pulmonary vascular resistance; RV, right ventricle; LS, least squares; RHC, right heart catheterization; ECHO, echocardiography.

Source: Data on file.

The Majority of Patients Receiving Seralutinib Demonstrated an Improvement in REVEAL 2.0 Risk Score at Week 24



1-point improvement in REVEAL 2.0 Risk Score at baseline associated with⁽¹⁾:

- 23% reduction in relative risk of death
- 20% reduction in relative risk of clinical worsening

Seralutinib patients have 2.45 times the odds of achieving a REVEAL 2.0 Risk Score improvement compared to placebo patients

30 of 39 seralutinib patients improved or maintained baseline REVEAL 2.0 Risk Score

Post hoc analysis. Odds ratio, 95% CI, and p-value from a stratified Cochran-Mantel-Haenszel chi-square test of improvement (yes vs. no).

1) A 1-point improvement in REVEAL 2.0 Risk Score (RRS) at PATENT-1 baseline was associated with a 23% reduction in the relative risk of death and a 20% reduction in the relative risk of clinical worsening in PATENT-2. Similarly, a 1-point improvement in RRS 2.0 at PATENT-1 Week 12 was associated with a 26% reduction in the relative risk of death and a 23% reduction in the relative risk of clinical worsening in PATENT-2. Source: https://doi.org/10.1016/j.ijcard.2021.03.034

Source: Data on file.



Safety and Tolerability



Top AEs From Imatinib IMPRES Study Not Observed at High Incidence in TORREY

Preferred Term ^a
Nausea
Peripheral edema ^b
Diarrhea
Vomiting
Periorbital edema ^c
Dyspnea
Hypokalemia
Anemia
Face edema d
Muscle spasms

IMPRES Study (Phase 3) Imatinib		
Placebo (N=98)	Imatinib (N=103)	
23 (24)	57 (55)	
20 (20)	45 (44)	
19 (19)	36 (35)	
10 (10)	31 (30)	
7 (7)	30 (29)	
13 (13)	19 (18)	
3 (3)	16 (16)	
3 (3)	14 (14)	
1 (1)	10 (10)	
2 (2)	10 (10)	

TORREY Study (Phase 2) Seralutinib		
Placebo (N=42)	Seralutinib (N=44)	
6 (14)	5 (11)	
1 (2)	2 (5)	
3 (7)	6 (14)	
3 (7)	2 (5)	
0 (0)	1 (2)	
5 (12)	4 (9)	
1 (2)	2 (5)	
0 (0)	1 (2)	
0 (0)	1 (2)	
0 (0)	1 (2)	

Abbreviations: AE: adverse event; MedDRA, Medical Dictionary for Regulatory Activities; PT, preferred term.

Note: AEs in IMPRES with an incidence ≥ 10% in Imatinib and ≥5% higher in Imatinib than Placebo are summarized for both IMPRES and TORREY.

Note: The above tables are for illustrative purposes only and are not a head-to-head comparison. Differences exist between study designs and methodologies, and caution should be exercised when comparing data across studies.

^a Coded using MedDRA (v 24.0 in TORREY).

^b Includes AE PTs of oedema, oedema peripheral, and peripheral swelling in TORREY.

c Includes AE PT of periorbital edema in IMPRES and AE PT of periorbital swelling in TORREY.

^d Includes AE PT of face edemain IMPRES and AE PT of swelling face in TORREY. Source: Data on file.

Incidence of TEAEs by Preferred Term: ≥ 5% in Seralutinib (Safety Population)

Preferred Term ^a	Placebo (N=42)	Seralutinib (N=44)
Number of subjects with a TEAE	36 (85.7)	41 (93.2)
Cough COVID-19	16 (38.1) 7 (16.7)	19 (43.2) 6 (13.6)
Diarrhea	3 (7.1)	6 (13.6)
Headache Dizziness	8 (19.0) 2 (4.8)	6 (13.6) 5 (11.4)
Fatigue Nausea	3 (7.1) 6 (14.3)	5 (11.4) 5 (11.4)
Dyspnea	5 (11.9)	4 (9.1)
Nightmare Abdominal pain lower	1 (2.4) O	4 (9.1) 3 (6.8)
Arthralgia Back pain	1 (2.4) 2 (4.8)	3 (6.8) 3 (6.8)
Chest discomfort Nasal congestion	1 (2.4) 1 (2.4)	3 (6.8) 3 (6.8)
Nasopharyngitis	0	3 (6.8)
Rash Throat irritation	1 (2.4) 0	3 (6.8) 3 (6.8)

All TEAEs in the table above were mild or moderate in severity.

Abbreviations: MedDRA, Medical Dictionary for Regulatory Activities; TEAE, treatment-emergent adverse event.

Source: Data on file.



 $^{^{\}mbox{\tiny a}}$ Coded using MedDRA v 24.0

Incidence of Adverse Events Leading to Treatment Discontinuation (Safety Population)

Preferred Term ^a	Placebo (N=42)	Seralutinib (N=44)
Number of subjects with a TEAE leading to treatment discontinuation	1 (2.4)	6 (13.6)
Abdominal pain lower	0	1 (2.3)
Cough	0	1 (2.3)
Dry mouth	0	1 (2.3)
Haemoptysis	0	1 (2.3)
Alanine aminotransferase increased	0	1 (2.3)^
Aspartate aminotransferase increased	0	1 (2.3)^
Transaminases increased	0	1 (2.3)
Liver function test abnormal	1 (2.4)	0

Abbreviations: MedDRA, Medical Dictionary for Regulatory Activities; TEAE, treatment-emergent adverse event.

Source: Data on file.

^a Coded using MedDRA v 24.0

[^] Events occurred in same patient.

Summary of TORREY Topline Results



Summary of Topline Results

- Primary endpoint met with concordant statistically significant and directional improvements from baseline across multiple endpoints, including hemodynamics, NT-proBNP, and right heart structure and function
- Drug characteristics, limited systemic PK, and route of administration led to the avoidance of safety/tolerability issues seen with systemic imatinib administration
- ✓ 6MWD improvement in more severe patient groups provides clear path forward for Phase 3 development program in PAH
- Statistically significant study in PAH and strong mechanistic rationale support development in Group 3 PH



Acknowledgements

We thank all patients, their families, and all the TORREY study investigators who participated in the trial





Next Steps

□ 1H:23 – Complete End of Phase 2 Regulatory Interactions

□ 2H:23 – Commence Global Registrational Phase 3 Program in PAH

☐ 2H:23-1H:24 – Commence Development Program in WHO Group 3 PH



Participants for Q&A Session

Gossamer Bio		
Faheem Hasnain	Co-Founder, Chairman, & Chief Executive Officer	
Richard Aranda, MD	Chief Medical Officer	
Robert Roscigno, PhD	VP, Clinical Development	
Larry Zisman, MD FACC	Sr Dir, Clinical Development	
Ed Parsley, DO	Consultant Pulmonologist	
Matt Cravets	SVP, Biometrics	
Laura Carter, PhD	Chief Scientific Officer	
Caryn Peterson	EVP, Regulatory Affairs	
Bryan Giraudo	COO & CFO	

Guest Speakers



Ardeschir Ghofrani, MD* Professor of Pulmonary Vascular Research, Justus Liebig University; Head of the Pulmonary Hypertension Division, University Hospital Giessen



Raymond Benza, MD*

Professor of Medicine, Division of
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