

Introduction

February 2025

Pharmazz, Inc. At A Glance



Pharmazz is a late-stage biopharmaceutical company that addresses the underserved critical care market. It has two first-in-class drug candidates in Phase 3 in the US, two drugs approved in India for marketing, and partnerships with major pharmaceutical companies.

Sovateltide, an endothelin-B agonist, is a first-in-class drug product for acute cerebral ischemic stroke.

- Approved Phase 3 IND and a SPA agreement with the FDA
- India Phase 3 data showing statistically significant improvement in neurological outcomes.
- Approved and marketed in India as Tyvalzi[™] by our strategic partner, Sun Pharmaceuticals, with 60,000+ patients treated since the launch in September 2023

Centhaquine is a resuscitative agent without arterial constriction for hypovolemic shock

- US IND for Phase 3 approved for hypovolemic shock
- US IND for Phase 2 approved for Acute Respiratory Distress Syndrome (ARDS)
- Approved in India and marketed in India as Lyfaquin® by our strategic partner, Dr. Reddy's Laboratories
- Clinical trial results from India met all four primary endpoints in Phase 3 and showed a 75% reduction in 28-day mortality



Business Highlights

Large and Growing Addressable Market Opportunity (5-year estimated CAGR from 2028E to 2033E).

- In the US, the treatment of ACIS is projected to generate \$3.6bn of net revenues with a 5-year CAGR of 132% by 2033
- In the US, the treatment of Hypovolemic Shock is projected to generate \$1.0bn of net revenues with a 5-year CAGR of 161% by 2033

Experienced Management Team with a Proven Track Record:

- Chairman & CEO has 40+ years of drug discovery, development, clinical, and management experience
- President has 30+ years of clinical, industry, and business experience
- COO has 35+ years of industry experience with a track record of conducting and leading numerous successful in-house clinical trials

Clear Visibility on Critical Milestones (projected)

- Sovateltide Milestones: Full data (4Q'26), File NDA (1Q'27), NDA Approved (3Q'27), Launch (4Q'27)
- Centhaquine Milestones: Full data (2Q'27), File NDA (3Q'27), NDA Approved (1Q'28), Launch (2Q'28)
- Established market receptivity to drug candidates in the Indian Market supported by high-profile strategic partners

Product Pipeline



	Indication	Pre-clinical	Phase 1	Phase 2	Phase 3	Market
Sovateltide	Acute Cerebral Ischemic Stroke					
	Hypoxic-Ischemic Encephalopathy					
	Alzheimer's Disease					
Centhaquine	Hypovolemic Shock					
	ARDS			13		
	Septic Shock					
PMZ-2123	Undisclosed Indication		nigh			
		US	S India			

Centhaquine

A resuscitative agent that is free of arterial constriction







Centhaquine: Hypovolemic / Hemorrhagic Shock



Hypovolemic / Hemorrhagic Shock is a life-threatening condition with high mortality rates. The annual incidence is 0.3 to 0.7 per 1,000 in the US with a 15% to 20% mortality rate

Decreased Cardiac Output	Caused by severe blood or fluid loss	Blood Pressure
Hypoperfusion of Organs	Due to poor cardiac output and perfusion of vital organs	Oxygen Levels
Multiple Organ Failure	This is the critical driver of mortality	Organ Failure
	DEATH	

Reference: Uptodate; StatPearls, Annane et al. 2013 JAMA; Standl et al. 2018 Dtsch Arztebl Int.; Canon et al. 2018 NEJM; Standl et al. 2018 Dtsch Arztebl Int.;

Progression

Centhaquine: Current Treatment Protocol



The current treatment protocol for hypovolemic shock includes a mix of fluid replacement and vasopressors

Current Treatment: Hypovolemic / Hemorrhagic Shock

Fluid Replenishment: Colloid / Crystalloid Solutions +/- Blood Products



Challenges with Current Treatment Protocol

- Arterial constriction, reduced tissue blood perfusion
- Cardiac Arrhythmias



- Fluid Extravasation
- Vasopressor Infusion requires careful titration

The administration of Centhaquine does not require the insertion of a Central Venous Line (peripheral IV administration instead)

Note: The average time to therapeutic intervention in the US is < 30 minutes (within the golden hour) versus > 60 minutes in India. This difference in therapeutic intervention is due to external factors

Centhaquine: Mechanism of Action

Centhaquine's MOA is distinct among resuscitative agents as it increases cardiac output while decreasing vascular resistance



Centhaquine increases cardiac output due to an increase in venous return in hypovolemic shock patients





Phase 3 Primary and Secondary Endpoints

Centhaquine's Phase 3 trial in India met all four primary efficacy endpoints. The trial's secondary endpoint, 28-day mortality, also trended toward the benefit

Key Parameters	Overview	Endpoints	Results (% of patients)		P Value
	71 patients: experimental arm:		Control	Centhaquine	
Treatment Arms	 Centhaquine + standard of care 34 patients: comparator arm: standard of care 	SBP ≥ 110 mmHg at 24 hrs.	60.6	79.7*	P=0.0444
		DBP ≥ 70 mmHg at 24 hrs.	51.5	76.6*	P=0.0122
Dosage	Centhaquine administered at 0.01mg/kg, i.v. in 100 mL of normal saline	Blood Lactate of ≤ 1.5	46.9	69.4*	P=0.0336
		Base-Deficit <- 2.0 (mmol/L)	43.8	69.8*	P=0.0137
Efficacy Assessment	SBP, DBP, Blood Lactate, base-deficit Secondary endpoint: 28-day Mortality	28-day Mortality	11.8	2.94	P=0.0742

Study Design Summary

Clinical Trials Identifier: CTRI/2019/01/017196 and NCT04045327

Reference: Gulati et al., (2021) Drugs. 2021 Jun;81(9):1079-1100; doi: 10.1007/s40265-021-01547-5; Gulati et al., (2021) Advances in Therapy 38 (6), 3223-3265. doi: 10.1007/s12325-021-01760-4.; Gulati et al., (2020) Drugs Fut 2020, 45(3): 153; doi: 10.1358/dof.2020.45.3.3098155.; Lyfaquin® clinical data

Centhaquine: Trial Protocol (US Phase 3 IND Approved)



Study Design		Primary	All cause mortality at day 28		
Design Parameters	Multi-Center, Randomized, Double- Blinded, Placebo-controlled	Endpoint	Mortality 60 days		
Dosage	0.01 mg/kg of Centhaquine + Standard of Care	Secondary Endpoints	 Days in hospital Days in ICU Days on organ support 		
No. of Participants	430 patients, randomly assigned equally to both arms		 Systolic and diastolic blood pressure Blood lactate 		
Time Frame	Enrollment period 12 months and total duration 24 months	Exploratory Endpoints	 Amount of fluid or blood infused Change in Multiple Organ Dysfunction Syndrome score 		

The India Phase 3 study showed a ~75% reduction in mortality. A Phase 2 and 3 data meta-analysis reached statistical (p=0.03) significance. A prospective, multi-centric, open-labeled study of 400 patients to assess the safety and efficacy of centhaquine is ongoing; more than 270 patients enrolled.

Sovateltide

A first-in-class drug candidate to demonstrate statistically significant results in acute cerebral ischemic stroke since tPA







Sovateltide: Mechanism of Action



A highly selective endothelin-B receptor agonist



- Increases cerebral blood flow
- Has anti-apoptotic activity with protection of neural mitochondria and enhances their biogenesis
- Produces neurovascular remodeling through the formation of new neurons and blood vessels
- Reduces infarct volume and improves neurological outcomes in an animal model of ACIS*

Sovateltide enhances the expression of markers for neural progenitor cells and neuronal cells, but not the stem cell markers

Reference: Ranjan et al., Sci Rep. 2020 Jul 29;10(1):12737. PMID: 32728189; Ranjan et al., Can J Physiol Pharmacol. 2020 Sep;98(9):659-666. PMID: 32574518; Briyal et al., Sci Rep. 2019 Jul 18;9(1):10439. PMID: 31320660; *Leonard et al., Brain Res. 2011;1420:48–58; Brain Res. 2012;1464:14–23; Brain Res. 2013;1528:28–41; Gulati Curr. Neuropharmacol. 2016;14(6):619–26; Gulati et al., (2021) CNS Drugs 35; 85–104. PMID: 33428177;; <u>https://rdcu.be/cdps6</u>

Sovateltide: SPA agreement with US FDA for Phase 3 Trial



Sovateltide Phase 3 IND clinical trial application approved by the US FDA (02/08/2023)



(NCT05691244)



Differences and similarities between India and US studies

Parameter	US Study (Special Protocol Assessment)	India Study
Primary endpoint	The proportion of patients with mRS of 0-2 at 90 days	The proportion of patients with improved neurological outcomes (mRS, NIHSS, BI) at 90 days.
Inclusion criteria	Age 18-80, Either sex; Ischemic stroke; Within 24 hours of stroke onset; NIHSS ≥8 to <20;	Age 18-78, Either sex; Ischemic stroke; Within 24 hours of stroke onset; NIHSS >5;
Exclusion criterion	Endovascular therapy, surgical intervention, intracranial hemorrhage, comatose, pregnancy	Endovascular therapy, surgical intervention, intracranial hemorrhage, comatose, pregnancy
Sample size; Randomization; Time from onset of stroke	514; 1:1 randomization; 50% within 12 hours (minimum 200 (40%) patients)	158; 1:1 randomization; within 12 hours 24% (38, 17 control and 21 sovateltide) patients
Interim analysis	No interim analysis	Trial complete, approved for marketing
Data analysis (Statistical Analysis Plan (SAP))	Multiple imputation for missing data, intention-to-treat (ITT) patients. SAP approved by FDA	No SAP. The next Slide Table is the data analyzed as per SAP with FDA, multiple imputation + ITT patients
Standard of care	SOC (thrombolytics, anti-coagulants, anti- hypertensive, anti-diabetic, mannitol, and other medication as needed)	SOC (thrombolytics, anti-coagulants, anti- hypertensive, anti-diabetic, mannitol, and other medication as needed)

Sovateltide: Phase III analysis of results



Data from 158 patients analyzed as per the agreed Special Protocol Assessment with the FDA

Number of patients with mRS of 0-2					
	Control (N=78)	Sovateltide (N=80)	P value		
Day 90 (Primary end point)	53.58% (N=42)	76.25% (N=61)	0.0031		
Day 30	41.03% (N=32)	63.75% (N=51)	0.0042		
Day 6	20.51% (N=16)	32.50% (N=26)	0.0882		
Number of patients with NIHSS of 0-5					
	Control (N=78)	Sovateltide (N=80)	P value		
Day 90 (Secondary end point)	67.95% (N=53)	85.00% (N=68)	0.0114		
Day 30	58.97% (N=46)	78.75% (N=63)	0.0072		
Day 6	37.18% (N=29)	56.25% (N=45)	0.0163		
Number of patients with BI of 90-100					
	Control (N=78)	Sovateltide (N=80)	P value		
Day 90 (Secondary end point)	43.59% (N=34)	57.50% (N=46)	0.0804		
Day 30	30.77% (N=24)	50.00% (N=40)	0.0138		
Day 6	8.97% (N=7)	20.00% (N=16)	0.0495		

Missing values imputed using Multivariate Imputation by Chained Equations (MICE)

Sovateltide: Phase III analysis of results





Distribution of scores on the Modified Rankin Scale at 90 days in the Intention-to-Treat population The modified Rankin Scale (mRS) score is the most widely used primary outcome measure in trials for acute stroke interventions. A modified Rankin scale score of 0 indicates no disability, 1 no clinically significant disability, 2 slight disability, 3 moderate disability but able to walk unassisted, 4 moderately severe disability, 5 severe disability, and 6 death.

Distribution of scores on the NIHSS Scale at 90 days in the Intention-to-Treat population. The National Institutes of Health Stroke Scale (NIHSS) is used to assess the severity of a stroke and the neurological deficit in stroke patients. The NIHSS of 1-4 = minor stroke. 5-15 = moderate stroke. 15-20 = moderate/severe stroke. 21-42 = severe stroke.

Data analyzed as per the Statistical Analysis Plan in the SPA agreed with the FDA

Status of US Phase 3 trial



A multicenter, randomized, double-blind, parallel, placebo-controlled phase III study will be conducted to assess the safety and efficacy of Sovateltide in patients with acute cerebral ischemic stroke. Control Group: Placebo + standard of care and Active Group: Sovateltide + standard of care (NCT05691244)



Upcoming Milestones



\$35M projected to fund through Sovateltide commercialization



Patents and Licenses

- Exclusive worldwide rights of intellectual property from Midwestern University with, single-digit royalties due once commercialized
- Several patent applications related to Sovateltide and Centhaquine composition and methods Inc. under examination.
- New patent application in filing process currently

Over 50 Issued Patents Covering Relevant Geographies With Expiry Between 2028 and 2044

The Team

Experienced team with extensive drug development and clinical expertise

Anil Gulati, MD, PhD Chairman and Chief Executive Officer » Inventor with 40 years of drug discovery,

development, clinical and management experience.

» 300 peer-reviewed publications and >50 issued patents

David Costello Vice President and Controller

- » 25 years of financial and accounting experience
- » Assisted closing of >\$500 million in structured finance and equity transactions

Neil Marwah, MD President

 » 30 years of experience in large healthcare provider organizations, government relations, managed care, private equity, and senior management at Global 500 enterprises

Sunil Gulati, PhD Chief Operating Officer

- » 35 years of running medium sized companies with governance and compliance expertise
- » In house development of clinical trials team and successful completion of numerous trials

Manish Lavhale, PhD Managing Director, India

- » 20 years of pharmaceutical industry experience
- » Expertise in regulatory strategy, with lead role in development of Centhaquine and Sovateltide

Dharmesh Shah, MD, DM

Assistant Medical Director

- » 15 years of clinical and pharmaceutical industry experience
- » Expertise in medical affairs with a role in the development of Centhaquine and Sovateltide

Late-stage biopharmaceutical company with two US FDA approved Phase 3 INDs for clinical programs addressing the underserved critical care market

Lead asset (Sovateltide) designed to transform the treatment of acute cerebral ischemic stroke, supported by the first statistically significant clinical data in 25+ years

Secondary asset (Centhaquine) designed to reduce mortality as a resuscitative agent and improving cardiac output and blood pressure without arterial constriction in

hypovolemic shock patients

Lead pipeline programs designed to address multibillion dollar end markets and line of sight on market debut by early 2027

Worldwide rights in hand with potential to partner both Sovateltide and Centhaquine in selected geographies

Validating and functional partnerships for sales and distribution in India

Dr.Reddy's Centhaquine

Thank You