



# Provascor

PROTECTING THE HEART

MRS2339 AS A FIRST-IN-CLASS NOVEL PROTECTIVE DRUG

[Provascor.com](http://Provascor.com)

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# MRS2339 is a first-in-class drug to treat patients with systolic heart failure without causing any drop in blood pressure or change in heart rate.



- The rationale: The heart's pumping ability is severely reduced, associated with low blood pressure, particularly in those with Class IIIb and Class IV heart failure.
- The problem: All current drugs lower blood pressure, causing concern by providers and lack of compliance from patients, particularly those in late/end stage HF;
- Unmet need: There is a need to develop new medical therapy for patients with late/end stage HF;
- The solution: MRS2339 can benefit these patients and address an unmet medical need with oral, SQ & IV formulations to treat the full spectrum of heart failure;
- Very high disease burden: 3-month readmission rate is up to 40% and 4-year mortality rate after advanced heart failure diagnosis is nearly 80%;
- Very high economic burden: cost projected to reach \$70B by 2030 in the U.S. alone;
- We aim to raise \$30m: Phase 1b (\$5m) with dose escalation on Class IIIb/Class IV patients and Phase 2 trial (\$25m) for POC as a therapy for hospitalized patients with worsening heart failure and as a SQ therapy for bridging to outpatient setting.

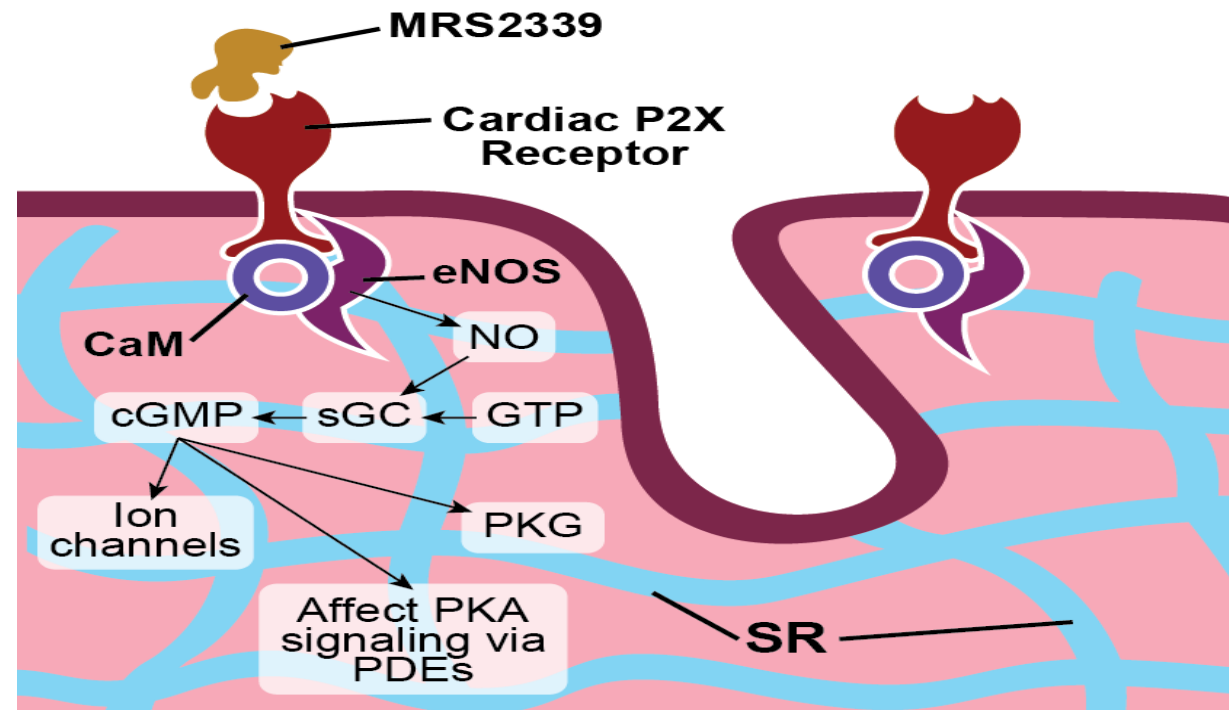


MRS2339 has a unique mechanism of action with cardiac-specific increase in cyclic GMP which is an established cardioprotective molecule

- MRS2339 is a cardiac P2X receptor agonist that is a hydrolysis-resistant adenosine monophosphate derivative;
- Physical association of P2X receptors with endothelial Nitric Oxide Synthase (eNOS) in the heart;
- MRS2339 is without vasodilator effect or effect on immune cells;
- Increase stroke volume and cardiac output & reverse the maladaptive remodeling;
- Cardioprotective with benefits lasting for at least one month post-dose;
- Efficacious in rescuing HF in multiple animal models of heart failure;
- Can be additive to other heart failure drugs and not replacing them.

MRS2339 differs from nonselective cyclic GMP stimulating drug which tends to cause hypotension in sick patients

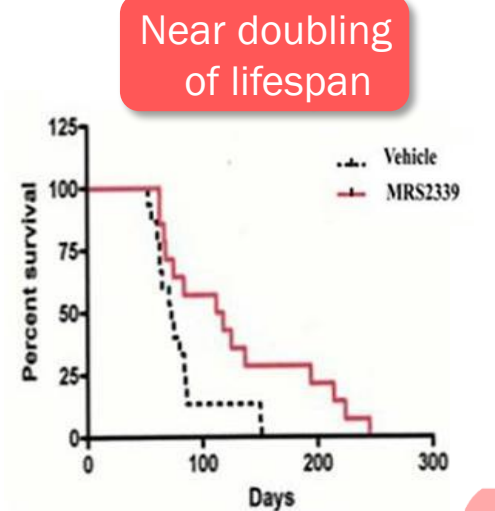
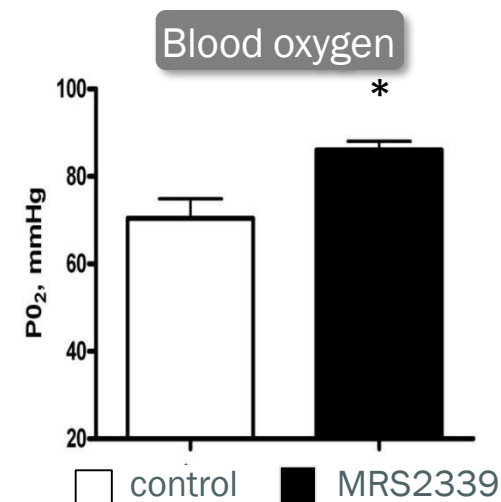
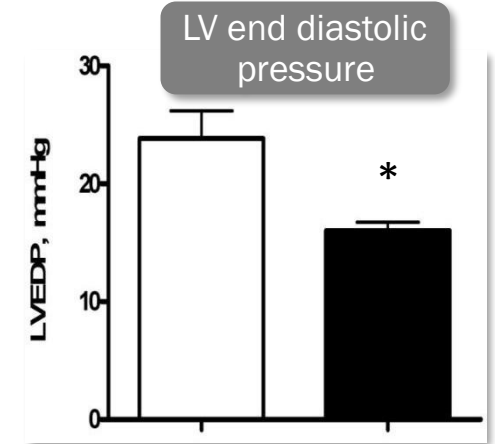
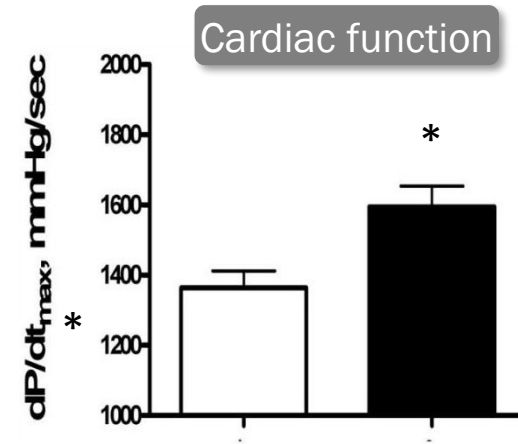
### Proposed Mechanism of Action



MRS2339 increases nitric oxide and subsequent increase in cyclic GMP with both being localized to heart

# MRS2339 improves heart function and prolongs lifespan in animal models

- MRS2339 improved contractile function, oxygenation and reduced lung congestion due to decreased left ventricular end diastolic pressure (LVEDP) which is equal to the pressure in the lungs (higher pressure impairs oxygenation)-saw improvement in just 4-5 dogs.
- MRS2339 preserved LV wall thickness, increased stroke volume and decreased LV remodeling
- Extended survival in a mouse model of severe dilated cardiomyopathy
- Wide therapeutic range: up to 22.5-fold



\* P<0.05 vs control dogs

Mouse



# No Competition in Class IIIb and Class IV Heart Failure Patients

| Company              | Product                                              | Likely Effect on BP in Class III & IV | % Class IV in Phase 3 Trial | Clinical Benefit in Class IV Patients |
|----------------------|------------------------------------------------------|---------------------------------------|-----------------------------|---------------------------------------|
| AstraZeneca          | farxiga®<br>(dapagliflozin) 5mg & 10mg tablets       | ↓                                     | < 1% in DAPA-HF             | No                                    |
| Boehringer Ingelheim | Jardiance®<br>(empagliflozin) tablets 10 mg/25 mg    | ↓                                     | 0.5% in EMPEROR-REDUCED     | No                                    |
| MERCK                | Verquvo®<br>(vericiguat) tablets 2.5 mg, 5 mg, 10 mg | ↓                                     | 1.4% in VICTORIA            | Not Shown for Class IV                |
| novo nordisk         | Victoza®<br>liraglutide injection 1.2 mg   1.8 mg    |                                       | FIGHT                       | Worse                                 |
| NOVARTIS             | Entresto®<br>Sacubitrilo-Valsartan                   | ↓                                     | 100% in LIFE                | No                                    |

MRS2339 does not cause hypotension while producing benefits in many animal models of systolic HF. MRS2339 is different from Verquvo which non-selectively increases cyclic GMP throughout and risks causing decreased blood pressure. We expect that MRS2339 will have little competition in late/end stage HF.



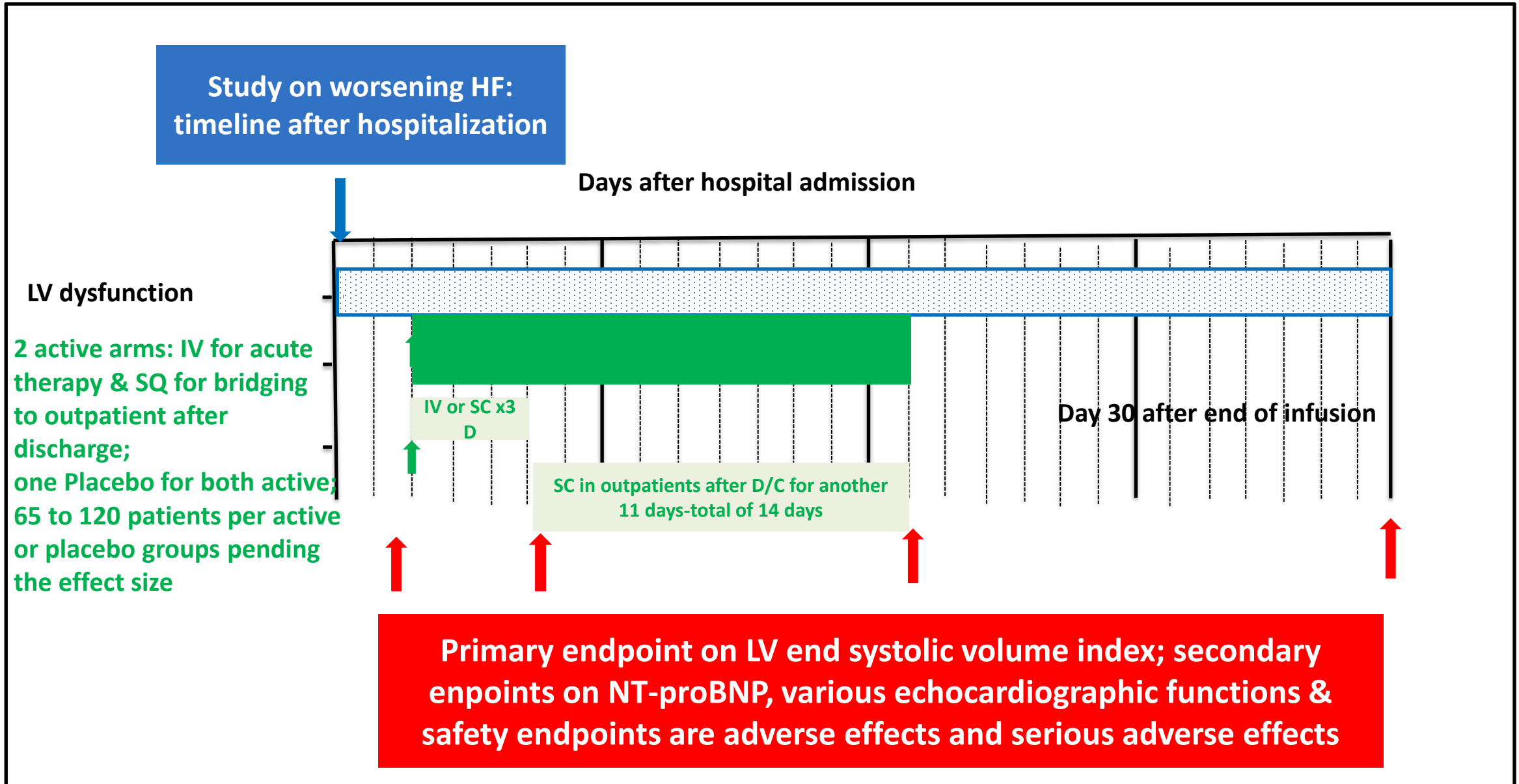
# MRS2339 testing is completed in Phase 1a study on healthy human volunteers without safety concerns or serious adverse effect

## Phase 1a – First-in-Human Study

- Randomized, double blind, single ascending dose (SAD) in healthy volunteers (HV)
- Assess the safety, tolerability, and pharmacokinetics (PK)
- 24hr IV infusion and 24hr monitoring (in-house) for 3 days: day-1, Day 1, Day 2
- 3 cohorts with 8 healthy subjects (4 males and 4 females) per cohort and total of 5 dosings.

*The highest of the 5 dosing levels is without any safety concerns or change in blood pressure or heart rate. This highest dosing level is 22.5 -fold or even higher than the anticipated efficacious dosage in humans. We expect a wide therapeutic range in patients. The lack of blood pressure or heart rate changes is a unique property, making the drug candidate suitable for all patients including severe heart failure. FDA agrees with the results of the 1a study and deems the drug safe in HV. **PD effect with target engagement is already evident.***

# Phase 2 POC for those with Class IIb & IV heart failure using the \$25m FDA advice requested, obtained and incorporated



# Timeline on the Phase 1b/Phase 2 trial for the indication of treating worsening HF- using the \$25-\$30m:

SC formulation developed with approval by FDA & SC delivery device testing done

*Mid 2024 – Mid/Late 2025-completed*

## **Milestone 4**

Phase 1a Clinical trial in  
Healthy Subjects IV infusion

*Mid 2026 – Late 2027*

## **Milestone 5 FDA approved protocol**

Phase 1b Clinical Trial in Class III, IV  
with IV infusion for safety, PK and PD

*2027 – 2029*

## **Milestone 6 FDA approved protocol**

Phase 2 POC using IV and/or SC  
formulations for worsening HF

*2029-2030 – exit options*

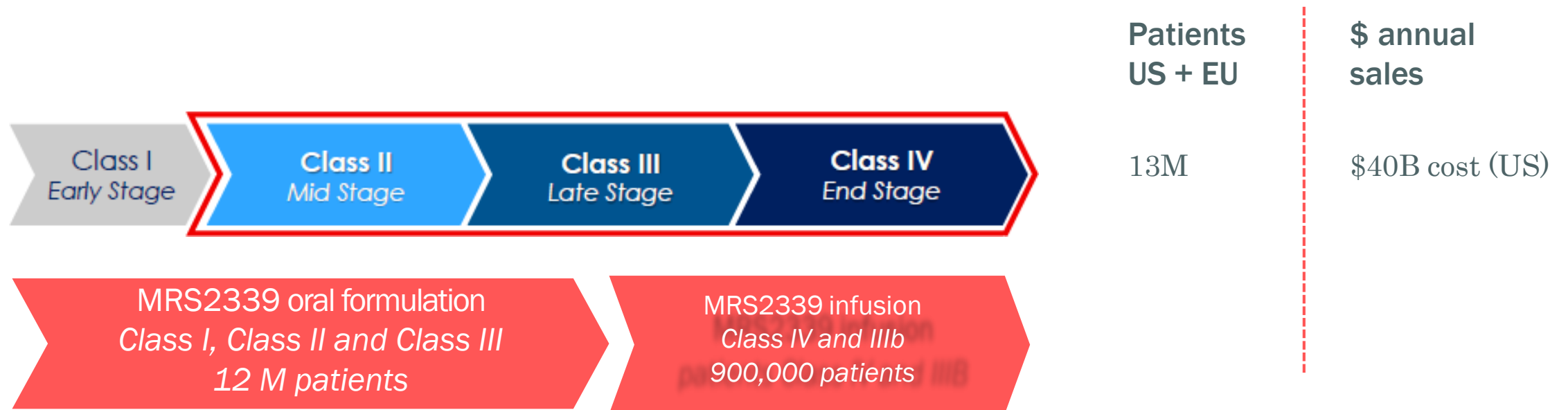
## **Milestone 7**

Partnership or acquisition by pharma, IPO

**IV or SQ infusion is for Class IIIb & Class IV patients;**  
**Oral ingestion is for chronic use in milder cases**

The annual gross revenue is based on \$5,000-\$7,000 for one infusion cycle of and 20% market penetration for \$0.9B-\$1.26B. Repeat cycle can be considered for each episode of worsening heart failure with doubling or tripling of gross sale. Repeat cycles are considered after the benefits waned after each cycle.

Oral therapy anticipates \$16.8B of annual gross sale.



# Provascor Leadership and Operation



Bruce Liang, MD  
Cardiologist/ Scientist,  
Dean of UConn School of  
Medicine,  
Co-Founder & Lead  
Advisor



Mark Van Allen  
Chief Operating  
Officer, Former head  
of UConn R&D  
Aided by 3 staff



Martin Bexon, MD,  
Bexon Clinical, Inc.  
Medical Monitor,  
Regulatory Support,  
Previously at CSL  
Behring and Roche



William White, MD  
Clinical  
Pharmacologist,  
Point on Clinical and  
Cardiovascular  
Safety Endpoint



Mary E ("Beth")  
P Goad, DVM,  
PhD, Board-  
Certified  
Veterinary  
Pathologist and  
Toxicologist  
With >30 Years  
of Experience.



Donna Mancini,  
MD, Clinical  
Development  
and Strategy  
Advisor on  
Heart Failure,  
Previously Led  
Advanced Heart  
Failure and  
Cardiac  
Transplant  
Programs at  
Columbia and  
Mount Sinai

James  
Heym, PhD

Science Advisor,  
Led Drug  
Discovery and  
Development As  
Vice President  
at Pfizer



# Preclinical and clinical plans on the oral formulation-need \$5m for Phase 1 & \$25m-\$27m for Phase 2 POC in Class II and Class III heart failure

*Mid 2026 - Mid 2027*

## **Milestone 1**

Successful development and testing in preclinical animal model

*Late 2027 – Mid 2028*

## **Milestone 2**

IND submission for an open IND in Phase 1 Clinical Trial on Class I, II or III heart failure patients

*Mid 2029 – Late 2029*

## **Milestone 3**

Analyze data on clinical POC from Phase 2 IV&SC trial & Ready for Phase 1 trial using oral drug

*Late 2029 – Early 2030*

## **Milestone 4**

Fund raising and conducting Phase 1 trial on oral drug in patients with mild HF

Lead and co-lead oral formulations as stable emulsions have been developed and are feasible to manufacture.