

UNITED STATES  
SECURITIES AND EXCHANGE COMMISSION  
WASHINGTON, D.C. 20549

FORM 8-K

CURRENT REPORT

Pursuant to Section 13 or 15(d) of the Securities Exchange Act of 1934

Date of Report (Date of earliest event reported): August 10, 2016

**Mast Therapeutics, Inc.**

(Exact name of Registrant as Specified in Its Charter)

Delaware  
(State or Other Jurisdiction  
of Incorporation)

001-32157

(Commission File Number)

84-1318182  
(IRS Employer  
Identification No.)

3611 Valley Centre Drive, Suite 500,  
San Diego, CA  
(Address of Principal Executive Offices)

92130  
(Zip Code)

Registrant's Telephone Number, Including Area Code: (858) 552-0866

Not Applicable

(Former Name or Former Address, if Changed Since Last Report)

Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions (see General Instructions A.2. below):

- Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)
- Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)
- Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))
- Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))

#### Item 8.01 Other Events.

The information attached as Exhibit 99.1 to this report, which relates to Mast Therapeutics, Inc. (the “Company”) and its development programs, may be presented from time to time by the Company at various investor and analyst meetings, including at the 36<sup>th</sup> Annual Canaccord Genuity Growth Conference on August 10, 2016.

#### Item 9.01 Financial Statements and Exhibits.

(d) Exhibits.

The list of exhibits called for by this Item is incorporated by reference to the Exhibit Index immediately following the signature page of this report.

By filing this report, including the information contained in Exhibit 99.1 attached hereto, the Company makes no admission as to the materiality of any information in this report. The information contained in Exhibit 99.1 hereto is summary information that is intended to be considered in the context of the Company’s filings with the U.S. Securities and Exchange Commission (the “SEC”), including its Annual Report on Form 10-K filed on March 14, 2016, Quarterly Report on Form 10-Q filed on August 9, 2016, and other public announcements that the Company makes, by press release or otherwise, from time to time. The Company undertakes no duty or obligation to publicly update or revise the information contained in this report, although it may do so from time to time as it believes is appropriate. Any such updating may be made through the filing of other reports or documents with the SEC, through press releases, or through other public disclosure.

#### Forward-Looking Statements

Mast Therapeutics cautions you that statements in this report, including in Exhibit 99.1 attached hereto, that are not a description of historical facts are forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995. Forward-looking statements may be identified by the use of words referencing future events or circumstances such as “expect,” “intend,” “plan,” “anticipate,” “believe,” and “will,” among others. Examples of forward-looking statements include, but are not limited to, statements regarding the Company’s development, regulatory and commercialization strategies and plans for its investigational new drugs, vepoloxamer (also known as MST-188) and AIR001, as well as the timing of activities and events related to those plans, including commencement and completion of clinical studies, announcements of study results, submission of applications to regulatory authorities for marketing approval, and product launch, and prospects for clinical, regulatory and commercial success. Forward-looking statements should not be read as guarantees of future performance or results because they involve the Company’s beliefs and assumptions based on currently available information and are subject to significant known and unknown risks and uncertainties that may cause actual performance and results to differ materially from expectations indicated by the forward-looking statements. Some of the factors that could cause actual performance or results to differ include, without limitation: the potential for additional delays in EPIC study closeout procedures, including blinded data validation and quality assurance/quality control procedures; the inherent uncertainty of outcomes in clinical studies and the risk that the Company’s product candidates may not demonstrate adequate safety, efficacy or tolerability in one or more such studies, including vepoloxamer in the ongoing EPIC study; the risk that, even if EPIC results are positive, the FDA or other regulatory agencies may determine they are not sufficient to support a new drug application; delays in the commencement or completion of clinical studies, including the EPIC study, the Phase 2 study of vepoloxamer in heart failure, and the Phase 2 studies of AIR001 in HFpEF, including as a result of difficulties in obtaining regulatory agency agreement on clinical development plans or clinical study design, opening trial sites, enrolling study subjects, manufacturing sufficient quantities of clinical trial material, completing manufacturing process development activities, being subject to a “clinical hold,” and/or suspension or termination of a clinical study, including due to patient safety concerns or lack of funding; risks associated with the Company’s ability to manage operating expenses and obtain additional capital as needed; the Company’s potential inability to continue as a going concern if it does not raise sufficient additional capital as needed; the risk that the Company may be required to repay its outstanding debt obligations on an accelerated basis and/or at a time that could be detrimental to its financial condition, operations and/or business strategy, including prepayment of \$10 million of the principal balance of its debt facility if results from EPIC are not positive and/or are not available on or before October 14, 2016; the Company’s reliance on contract research organizations (CROs), contract manufacturing organizations (CMOs), and other third parties to assist in the conduct of important aspects of development of its product candidates, including clinical studies, manufacturing, and regulatory activities for its product candidates and that such third parties may fail to perform as expected, leading to delays in product candidate development, regulatory approval, and/or commercial launch; the potential for the Company to significantly delay, reduce or discontinue current and/or planned development, regulatory and commercial-readiness activities or sell or license its assets at inopportune times if it is unable to raise sufficient additional capital as needed; the risk that the FDA and regulatory agencies outside of the U.S. do not grant marketing approval of a product candidate, on a timely basis, or at all; the risk that, even if the Company successfully develops a product candidate in one or more indications, it may not realize commercial success and may never achieve profitability; the risk that the Company is not able to adequately protect its intellectual property rights, through patents or otherwise, and prevent competitors from

duplicating or developing equivalent versions of its product candidates or that the use or manufacture of its products or product candidates infringe the proprietary rights of others; and other risks and uncertainties more fully described in the Company's press releases and periodic filings with the Securities and Exchange Commission.

You are cautioned not to place undue reliance on forward-looking statements, which speak only as of the date they are made. Mast Therapeutics does not intend to revise or update any forward-looking statement set forth in this report to reflect events or circumstances arising after the date hereof, except as may be required by law. This caution is made under the safe harbor provisions of Section 21E of the Securities Exchange Act of 1934, as amended, and Section 27A of the Securities Act of 1933, as amended.

**SIGNATURES**

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

Mast Therapeutics, Inc.

Date: August 10, 2016

By: /s/ Brandi L. Roberts  
Brandi L. Roberts  
Chief Financial Officer and Senior Vice President

Exhibit Index

Exhibit  
Number

Description

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99.1 Mast Therapeutics, Inc. corporate presentation, August 10, 2016



# Corporate Overview

## August 10, 2016

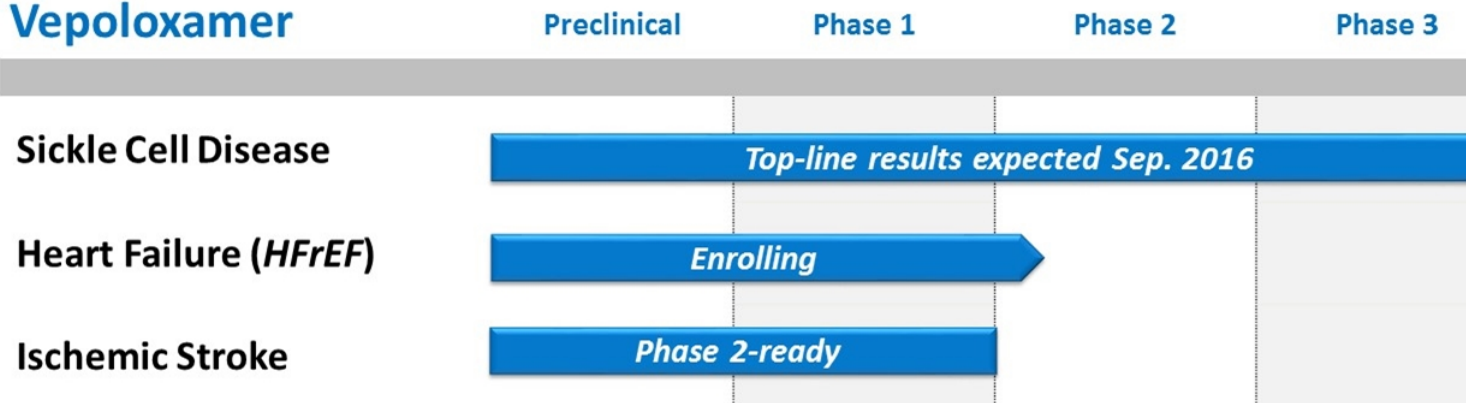
# Safe Harbor Statement

This presentation includes forward-looking statements about our business prospects, financial position, and development of vepoloxamer and AIR001 for therapeutic use in humans. Any statement that is not a statement of historical fact should be considered a forward-looking statement. Because forward-looking statements relate to the future, they are subject to inherent risks, uncertainties and changes in circumstances that are difficult to predict. Actual events or performance may differ materially from our expectations indicated by these forward-looking statements due to a number of factors, including, but not limited to, results of our pending and future clinical studies, the timeline for clinical and manufacturing activities and regulatory approval; our dependency on third parties to conduct our clinical studies and manufacture our clinical trial material; our ability to raise additional capital, as needed; our ability to repay outstanding debt as payments come due; our ability to establish and protect proprietary rights related to our product candidates; and other risks and uncertainties more fully described in our press releases and our filings with the SEC, including our quarterly report on Form 10-Q filed with the SEC on August 9, 2016.

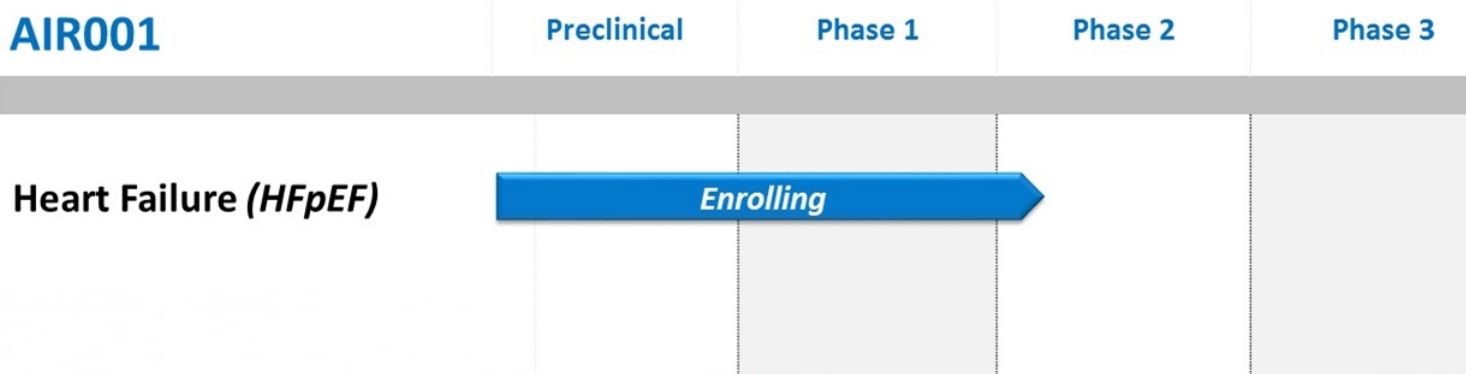
We caution you not to place undue reliance on any of these forward-looking statements, which speak only as of the date of this presentation. We do not intend to update any forward-looking statement included in this presentation to reflect events or circumstances arising after the date of the presentation, except as may be required by law.

# Product Candidate Pipeline

## Vepoloxamer



## AIR001

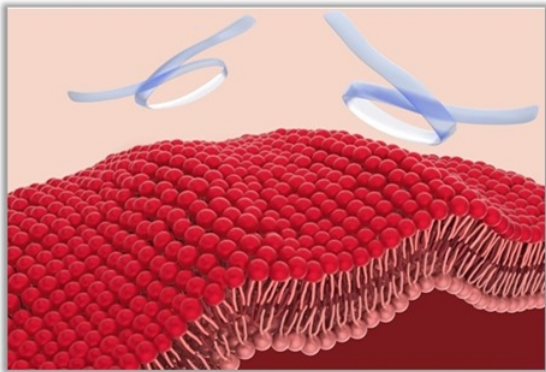




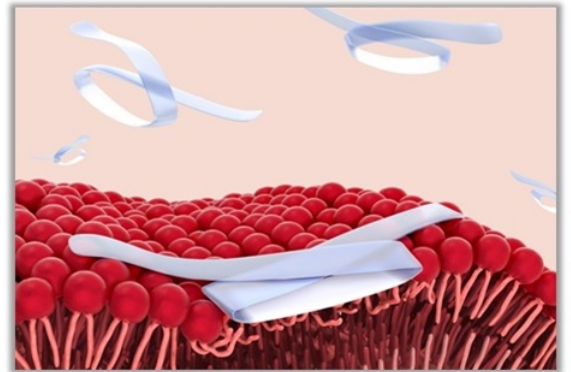
# Vepoloxamer: A Novel Biophysical Agent

- **Corrects defects in membrane surface tension, an underlying feature of multiple diseases**
- **“Bio-physical” mechanism of action offers lower development risk;**
  - Targets damaged tissue; little or no activity in healthy tissue
  - Not metabolized; no active metabolites to track, no difference in fast vs slow metabolizers, less susceptible to drug-drug interactions
  - Less susceptible to genetic variation; independent of receptors, etc.

*Healthy Cell Membrane? Vepoloxamer Inactive*



*Damaged Cell Membrane? Vepoloxamer Adheres*



# Pathologies and Conditions Related to Defects in Membrane Surface Tension

## Surface Tension Pathologies

Elevated blood viscosity

Cellular aggregation

Cellular adhesion

Loss of membrane barrier function / repair capacity

Dysfunctional coagulation

## Examples

Sickle Cell Disease

Chronic & Acute  
Heart Failure

Ischemic Stroke

# Vepoloxamer in Sickle Cell Disease

**Objective: Improve blood flow to prevent ischemic injury and shorten duration of crisis**

**Phase 3 Study – Enrollment Complete**

# Overview of Sickle Cell Disease

- **A chronic genetic disorder and rare (orphan) disease**
  - Affects ~100,000 people in the U.S. and is characterized by severe deformation of red blood cells
- **Hallmark of disease is a “vaso-occlusive crisis” (VOC)**
  - Reduces blood flow and causes ischemic injury to tissues and organs
  - Exceedingly painful condition and a leading cause of hospitalization
  - 80-100k hospitalizations annually in the U.S. with average hospital stay of 4-5 days
- **Current treatment is palliative**
  - Consisting of hydration and IV opioids
  - Does not treat underlying pathophysiology of the disease
  - No VOC interventional therapy available
- **Significant unmet need**
  - No approved agents to shorten duration or severity of crisis
  - Shorter life expectancy due to ischemic injury (~45y)

Sources: Life-expectancy: Hassell, K. Am J Prev Med 2010; 38(4S): S512-521; Utilization data calculated from data in Brousseau, D., et al., Acute Care Utilization and Rehospitalizations for Sickle Cell Disease. Apr 7 2010 and HCUP 2010-2012; SCD prevalent pop: Brousseau DC, Panepinto JA, Nimmer M, Hoffmann RG. The number of people with sickle-cell disease in the United States: national and state estimates Hematol 2010;85:77-8; Steinberg, M.H., Management of Sickle Cell Disease, New Eng J Med; 1999; Vol 340, No 13; Platt, et al., Mortality in Sickle Cell Disease (N Engl J Med 1994;330:1639-44)

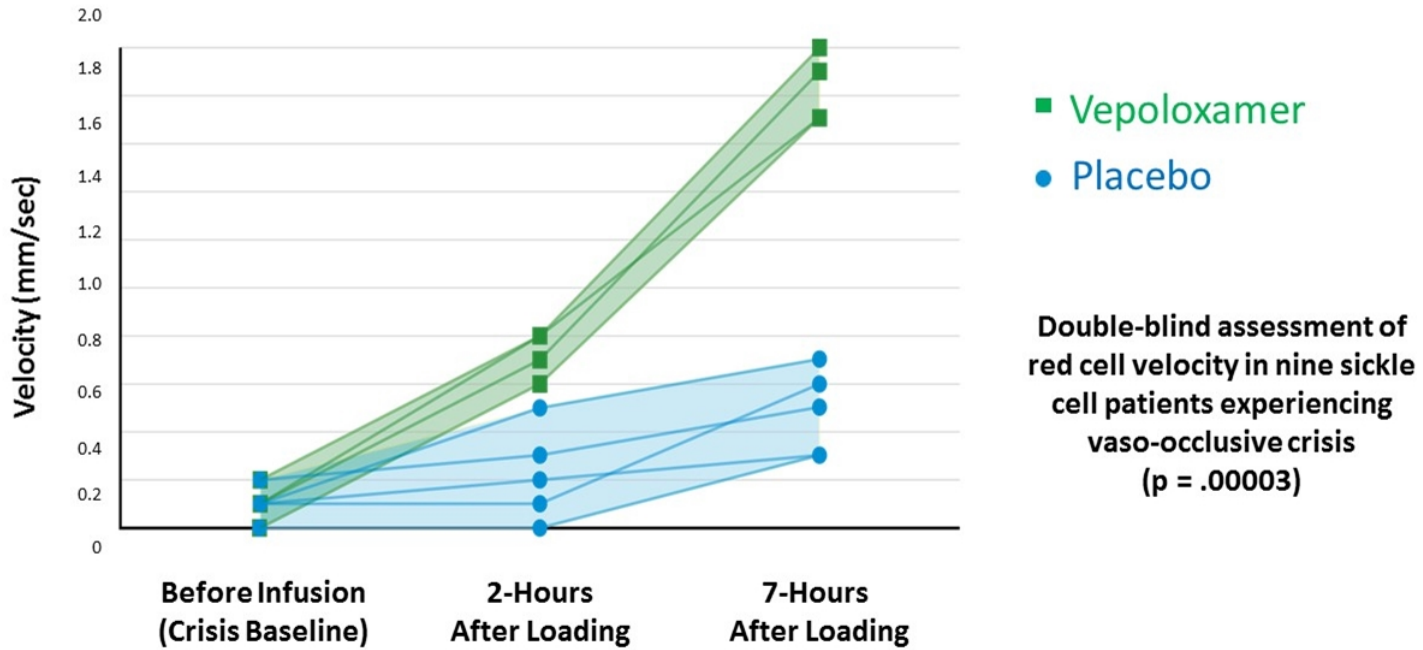
# Vepoloxamer in Sickle Cell Disease (SCD)

- **The underlying pathology of vaso-occlusive crisis is reduced blood flow**
- **Vepoloxamer improves blood flow by addressing multiple pathological mechanisms:**
  - Adhesion
  - Hemolysis
  - Aggregation
  - Viscosity

**Objective of treatment with vepoloxamer: Improve blood flow to prevent ischemic injury and shorten the duration of crisis**

# Vepoloxamer Improved Blood Flow in Sickle Cell Patients

Vepoloxamer significantly improved microvascular blood flow in sickle cell patients experiencing crisis (randomized sub-study)



# Vepoloxamer Development History

- **Over 100 nonclinical studies completed**
- **Phase 2 SCD – statistically significant shorter crisis and less opioid use**
- **Phase 2 ACS\* – ~50% shorter hospitalization stay vs. historical control**
- **Phase 3 SCD – shorter duration of crisis (p-value = 0.07)**

## **Lessons learned from prior sponsors and applied to Mast's Phase 3 study:**

1. Vepoloxamer has activity in SCD
2. Study design is key to success (prior endpoint poorly designed)
3. Key FDA feedback:
  - Utilize a clinically meaningful endpoint
  - Use as objective an endpoint as possible
  - Avoid use of pain scores due to variability
  - Provide a plan to minimize data loss

# The Phase 3 “EPIC” Study

## Evaluation of Purified Poloxamer 188 In Vaso-Occlusive Crisis

- **Largest placebo-controlled study in sickle cell disease ever completed**
- **Double-Blind, Placebo-Controlled, International (~75 sites)**
  - 388 patients, randomized 1:1 to standard of care +/- vepoloxamer
  - 1hr loading dose followed by 48h continuous infusion
- **Primary Endpoint**
  - Duration of crisis (time of randomization to last dose of parenteral opioid)
- **Secondary Endpoints**
  - Re-hospitalization for VOC within 14 days
  - Occurrence of acute chest syndrome within 120 hours of randomization
- **Other Assessments**
  - Safety
  - Duration of hospitalization
  - Biomarkers
  - Opioid utilization
  - Sub-study outcomes





# EPIC Study: Primary Endpoint

- **Assessment: Duration of vaso-occlusive crisis**
- **Definition: Time from randomization to last dose of parenteral opioid (LPC)**
- **Advantages:**
  - Aligns with FDA recommendations
    - Sensitive and specific data collection
    - Objective
    - Minimal data loss
  - Medical expert support
  - Clinically meaningful to experts in the field
- **Powering:**
  - 90% power to detect a 17% difference in treatment arms with a statistical significance level of  $p=0.05$

# EPIC Study: Safety

## ➤ **DSMB Evaluations**

- Independent, unblinded DSMB (4 clinicians and 1 statistician)
- Meetings at 25, 58, 145 and 250 patients

## ➤ **DSMB meeting held at 250 patients**

- No unexpected safety signals identified
- DSMB members deemed no additional meetings were necessary

# EPIC Study: Key Characteristics

- **U.S. patients = ~60%**
  - **Average patient age = 15 years (range 4-46, ~30% adults)**
  - **Hydroxyurea use = >60%**
  - **Blinded analysis at 250 patients:**
    - Average duration of crisis (pooled) & coefficient of variation consistent with statistical assumptions in study design (79h and ~50%)
    - Minimal regional variability (U.S. vs ex-U.S.)
- ✓ **Top-line data anticipated September 2016**

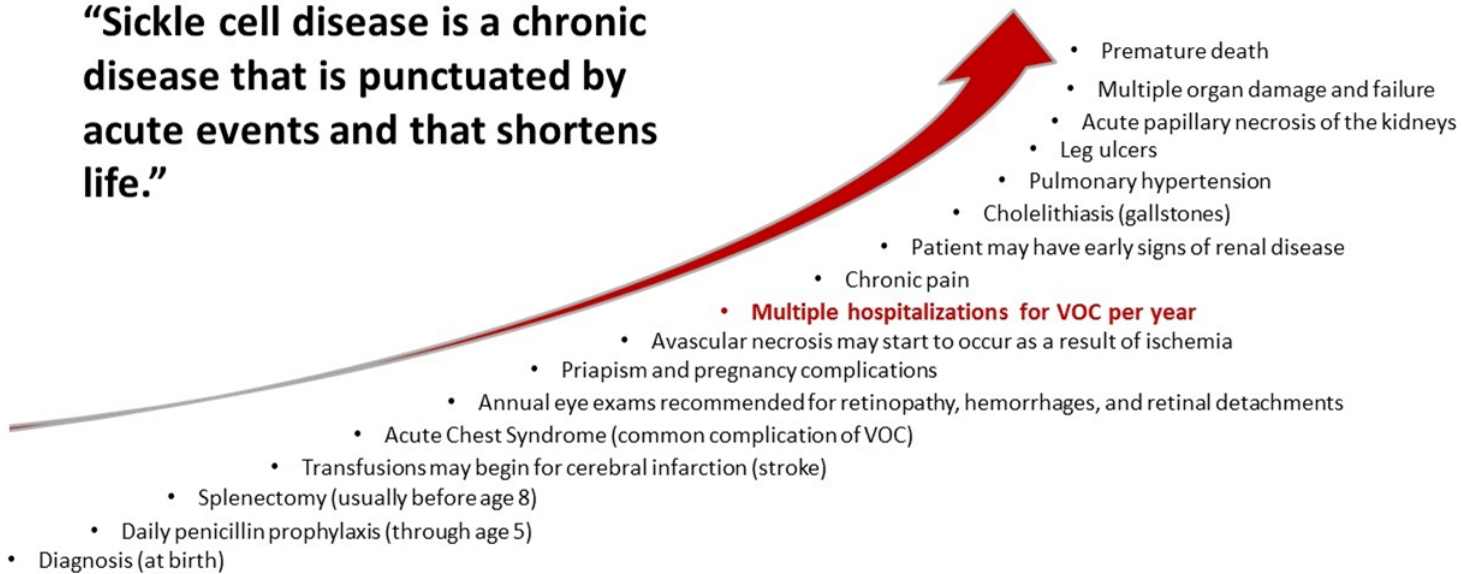
# Vepoloxamer in SCD: Regulatory Consideration

- Significant unmet need – no disease-modifying therapies for an ongoing crisis
- Support among medical / advocacy / patient communities
- Orphan Drug Designation
- Fast Track Designation
- SCD is part of FDA “Patient-Focused Drug Development Initiative”
- Healthcare disparity concerns
- NDA-supportive clinical studies:
  - Thorough QT study - *complete*
  - Special population study - *enrolling*
  - Repeat-administration study – *enrolling*
    - ✓ One patient who completed the EPIC Study has subsequently received three doses of vepoloxamer in the elective repeat-dose study

# Vepoloxamer: Commercial Opportunity in Sickle Cell Disease

# SCD: A Lifetime of Complications

**“Sickle cell disease is a chronic disease that is punctuated by acute events and that shortens life.”**

- 
- Diagnosis (at birth)
    - Daily penicillin prophylaxis (through age 5)
    - Splenectomy (usually before age 8)
  - Transfusions may begin for cerebral infarction (stroke)
  - Acute Chest Syndrome (common complication of VOC)
  - Annual eye exams recommended for retinopathy, hemorrhages, and retinal detachments
  - Priapism and pregnancy complications
  - Avascular necrosis may start to occur as a result of ischemia
  - **Multiple hospitalizations for VOC per year**
  - Chronic pain
  - Patient may have early signs of renal disease
  - Cholelithiasis (gallstones)
  - Pulmonary hypertension
  - Leg ulcers
  - Acute papillary necrosis of the kidneys
  - Multiple organ damage and failure
  - Premature death



Source: Martin H. Steinberg, M.D., Management of Sickle Cell Disease, New Eng J Med; 1999; Vol 340, No 13

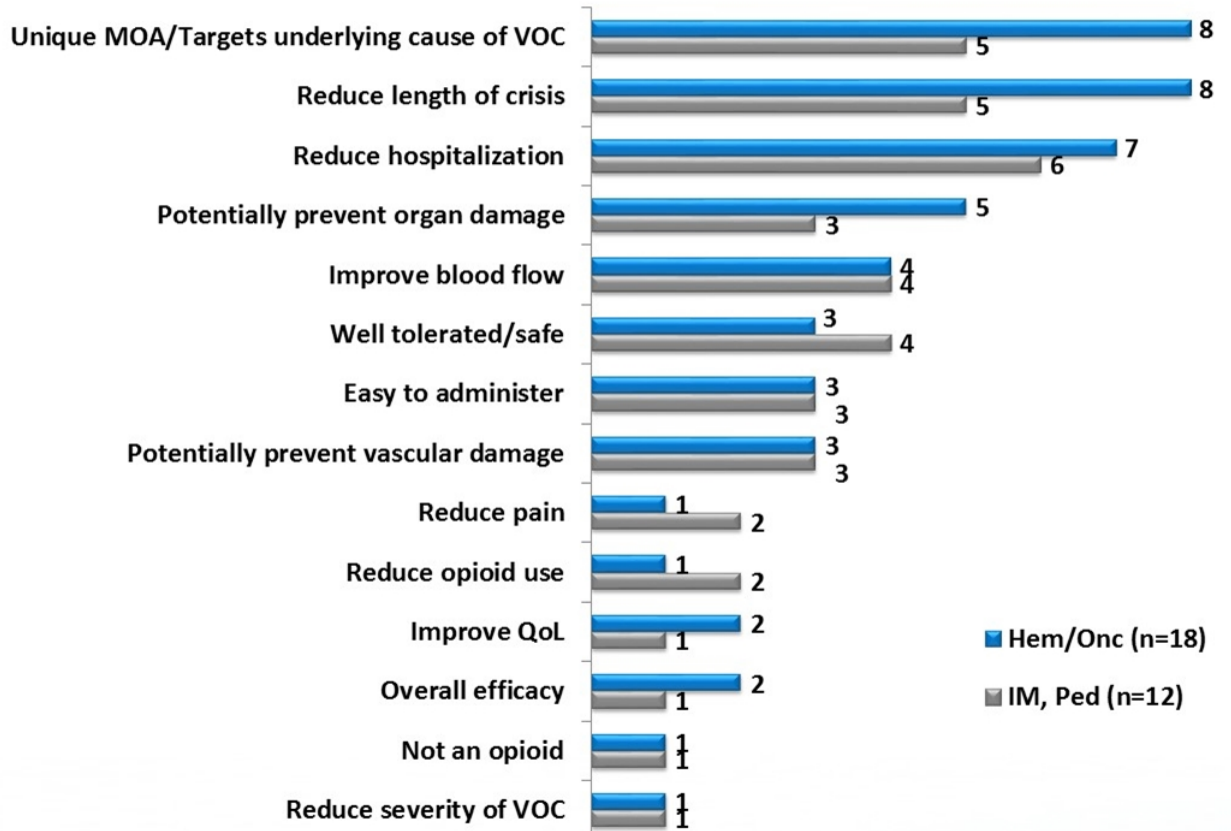
# Vaso-Occlusive Crisis is the Hallmark of SCD

- **80-100k hospitalizations annually in the U.S.**
- **Current treatment is palliative**
  - Consisting of hydration and IV opioids
  - Does not treat underlying pathophysiology of the disease
  - No VOC interventional therapy available
- **Hospitalized on average for approximately 4-5 days**
- **40% of patients will have acute chest syndrome complication**
- **12-41% are re-hospitalized within 30 days of hospital discharge**
- **Shorter life expectancy due to ischemic injury to organs (~45y)**
- **Organ failure is leading cause of death**

Sources: Life-expectancy: Hassell, K. Am J Prev Med 2010; 38(4S): S512-521; Utilization data calculated from data in Brousseau, D., et al., Acute Care Utilization and Rehospitalizations for Sickle Cell Disease JAMA Apr 7 2010 and HCUP 2010-2012; SCD prevalent pop: Brousseau DC, Panepinto JA, Nimmer M, Hoffmann RG. The number of people with sickle-cell disease in the United States: national and state estimates. Am J Hematol 2010;85:77-8; Steinberg, M.H., Management of Sickle Cell Disease, New Eng J Med; 1999; Vol 340, No 13; Platt, et al., Mortality in Sickle Cell Disease (N Engl J Med 1994;330:1639-44)

# Market Research: Expected Features and Potential Benefit Align with Unmet Needs for VOC Treatment

**Key Advantages/Features of Vepoloxamer**  
 (# of ranked mentions, unaided)

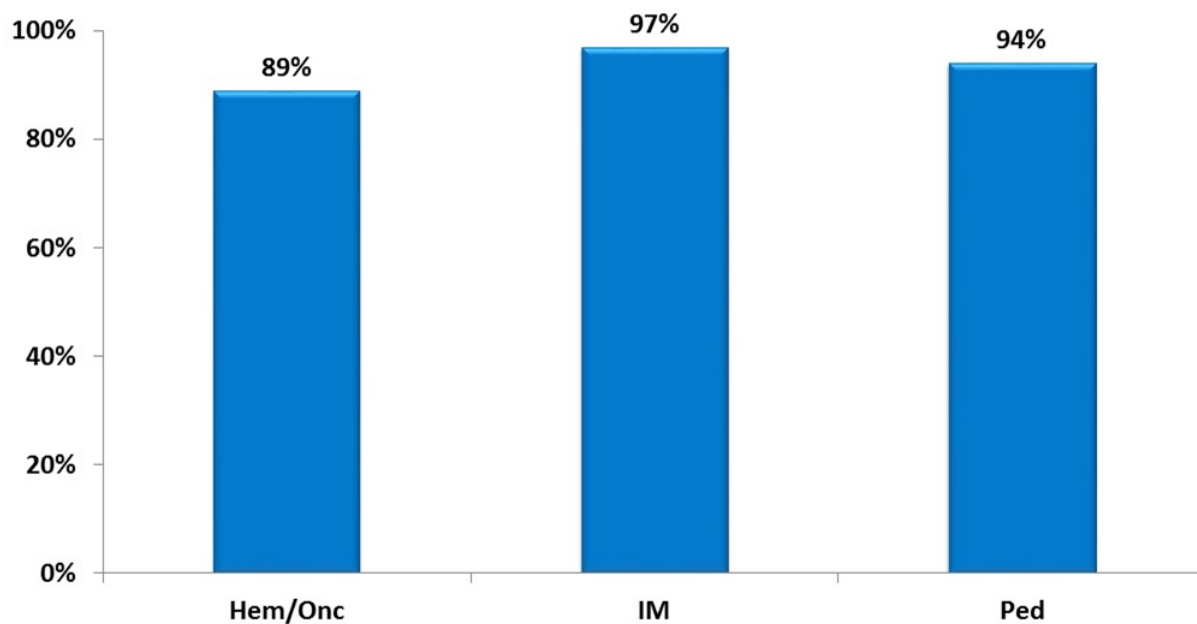




# Market Research: Physicians Report a High Percent of Patients to be Treated with Vepoloxamer at Peak

## Average % of Patients Treated with Vepoloxamer at Peak

n=30



Survey-reported time to peak adoption: approx. 6 months)

# Development Landscape in SCD

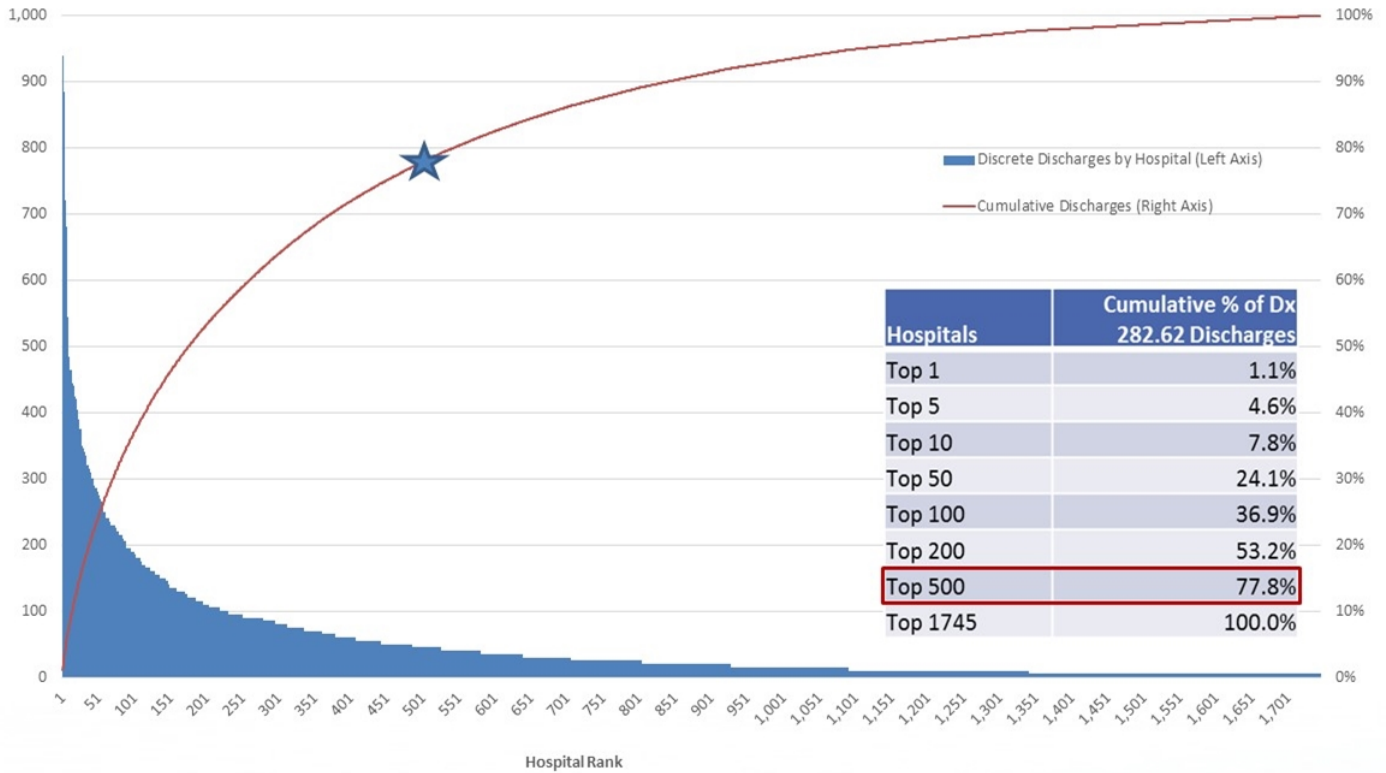
Vepoloxamer has the potential to be the first and only interventional treatment to reduce the duration and severity of an ongoing VOC

Stage	VOC Intervention	VOC Reduction	SCD Corrective Treatment
Marketed		hydroxyurea	Bone Marrow Transplant
Phase 3	<b>vepoloxamer</b> rivipansel	L-glutamine	
Phase 2		SC-411 Sanguinate Sevuparin SelG1 GBT440	
Phase 1		CXA-10 NiCord/CordLn NKTT120 PF-04447943 SCD-101	Gene therapy

# Concentration of SCD Treatment in the U.S. Offers an Attractive Commercial Opportunity

- Top 500 U.S. hospitals treat nearly 80% of SCD patients in crisis
- Effective field promotion with small hospital sales force (~30)

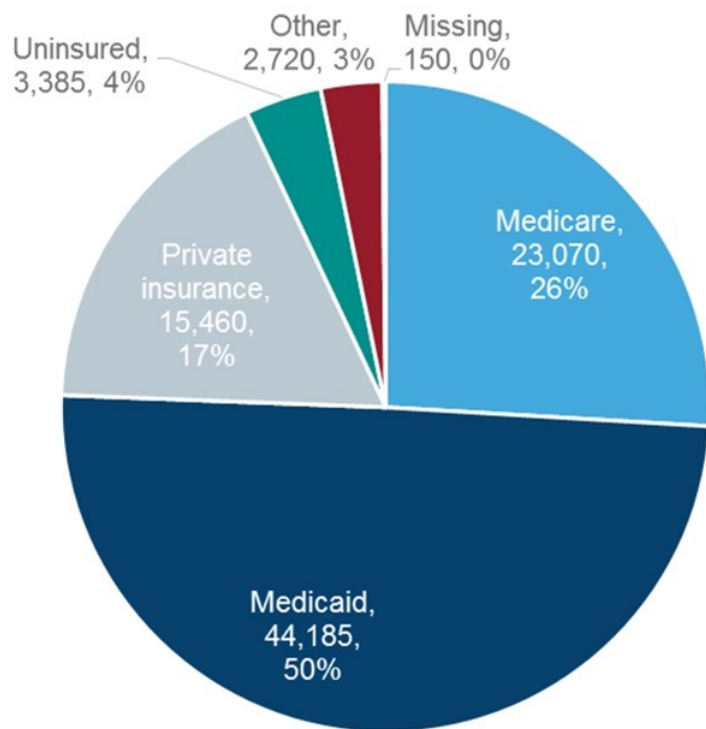
Distribution of Dx 282.62 (HbSS with VOC) Discharges by US Hospital



Source: Analysis of HCUP NIS 2013 data; Dx 282.62 is 89.7% of all VOCs.

# U.S. Sickle Cell Disease Hospital Payer Mix

Sickle Cell Disease, Payer Mix  
2012 HCUP data



- **Inpatient prospective payment system (IPPS)**
- **Patients with public insurance have minimal share of costs**
- **Medicaid plans may seek additional funding via CHIP, per diem rates, or modified FFS rates**
- **Vepoloxamer expected to meet criteria for additional Medicare payment (NTAP)**

# Commercial Potential Outside the U.S.

## ➤ Over 10 million patients worldwide

## ➤ Europe

- Approximately 40,000 patients
- >50% reside in two countries: UK and France
- Most patients concentrated in large metro areas: Paris and London



## ➤ MENA

- Over 850,000 with SCD
- Many treatment centers provide care on par with standard of the U.S. and E.U.

Sources: Hassell KL. Population estimates of sickle cell disease in the U.S. *Am J Prev Med* 2010;38(4 Suppl):S512-21; Data on file, Mast Therapeutics 2015; VOI Consulting analysis, 2015

# Vepoloxamer Positioned for Success in SCD

- **Novel therapy for a genetic disease with high unmet needs**
  - Unique and relevant mechanism of action
  - No approved disease-modifying therapies available for VOC intervention
- **Significant first-to-market advantage in multiple territories**
  - Clinical development >2 years ahead of other SCD programs
  - Orphan Drug Designation in U.S. and E.U.
- **Concentrated market**
  - Top 500 U.S. hospitals treat nearly 80% of SCD patients in crisis
  - 96% of SCD patients in the U.S. have insurance coverage
- **Research supports rapid adoption & significant market penetration**
  - Ranked 4.4 out of 5 as a “breakthrough medical innovation” by pharmacy directors at key SCD institutions
  - KOLs and community physicians express high intent to use

# Development of Vepoloxamer in Heart Failure

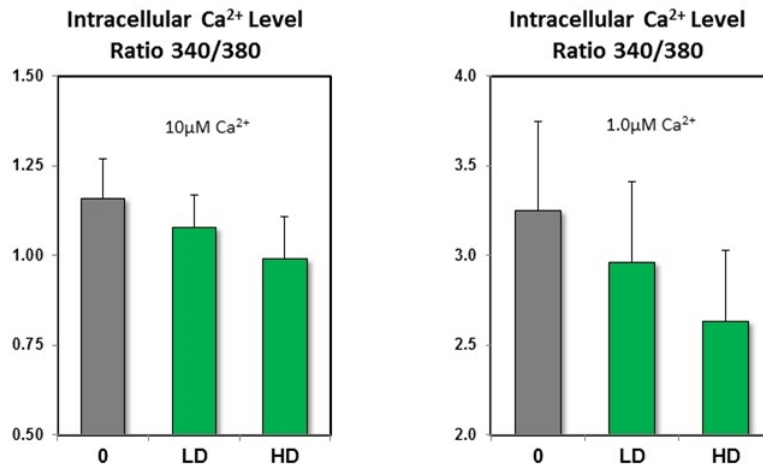
**Objective: Restore Membrane Integrity To Improve Cardiac Performance**

**Phase 2 Study - Enrolling**

# Development Rationale in Heart Failure

- Elevated wall tension in a dilated (e.g. failing) heart impairs normal membrane repair activity
- Permeabilized membranes allow *unregulated calcium influx* and cardiac troponin leak

## Cardiomyocytes Isolated from Animals with Advanced Heart Failure Exhibit Reduced Intracellular Calcium



LD = 1.5 mg/mL vepoloxamer  
HD = 4.5 mg/mL vepoloxamer



# Heart Failure Model Results

## ➤ **Functional improvement (single administration)**

- Vepoloxamer elicited improvements in Left Ventricle (LV) systolic and diastolic function that persisted for up to 2 weeks
- Ejection fraction (EF) and stroke volume (SV) increased

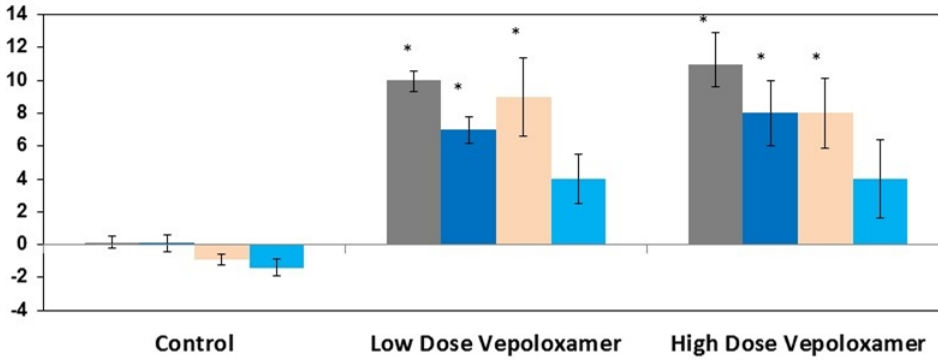
## ➤ **Biomarkers (single administration)**

- Functional improvements supported by significant reductions of NT-pro BNP for up to 2 weeks
- Membrane sealing supported by significant reductions in plasma troponin for up to 2 weeks
- Data suggests vepoloxamer can preserve cardiomyocytes by limiting calcium entry into the cell

## ➤ **Results support clinical development of vepoloxamer for the treatment of acute and chronic heart failure**

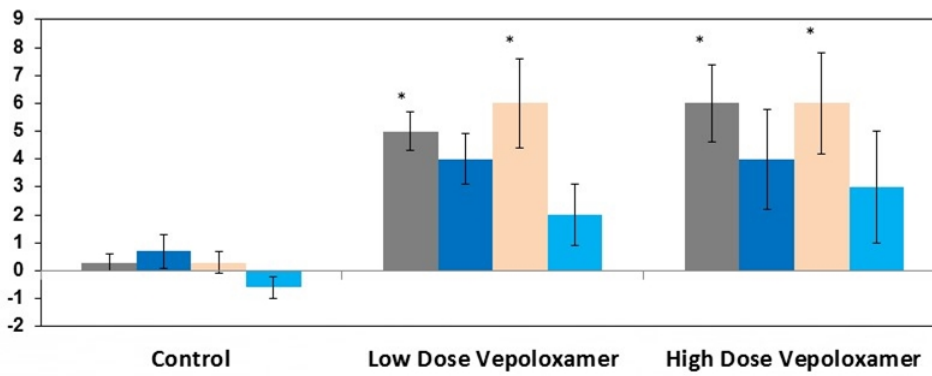
# Heart Failure Model – Functional Improvement (single administration)

**Δ LV Ejection Fraction (%)**



■ 2 Hours Post  
■ 24 Hours Post  
■ 1 Week Post  
■ 2 Weeks Post  
 \* p < 0.05 vs. Control

**Δ Stroke Volume (mL)**

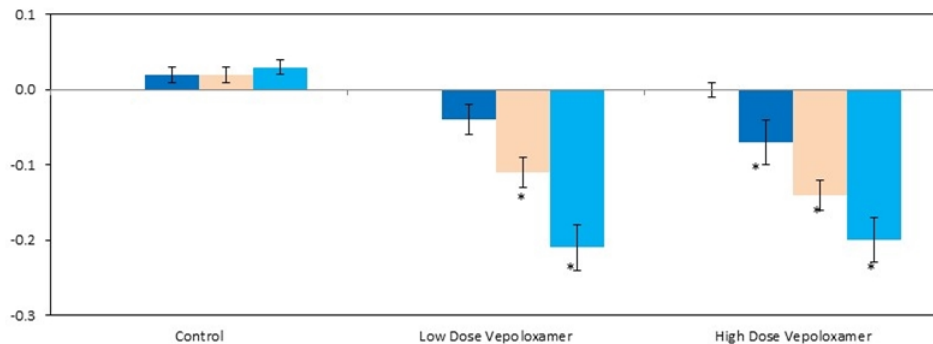


➤ **Vepoloxamer elicited improvements in LV systolic and diastolic function that lasted for up to 2 weeks**

Study conducted by: Hani N. Sabbah  
 Henry Ford Health System

# Heart Failure Model – Biomarkers (single administration)

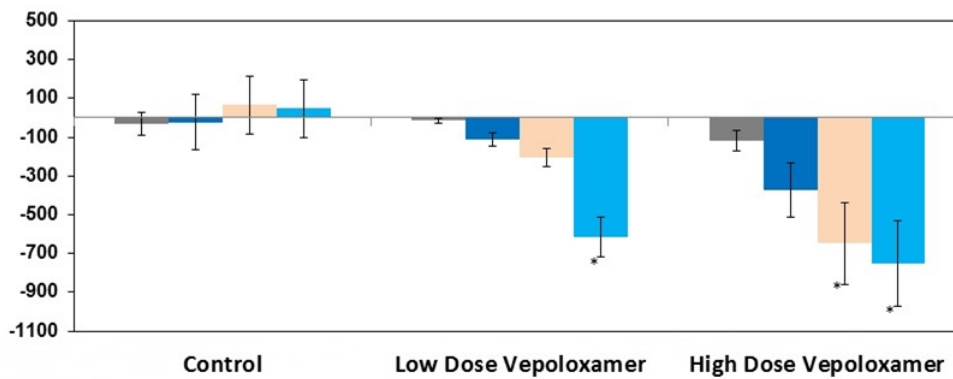
$\Delta$  Plasma Troponin-I (ng/mL)



- 2 Hours Post
- 24 Hours Post
- 1 Week Post
- 2 Weeks Post

\* p < 0.05 vs. Control

$\Delta$  nt-pro BNP (pg/mL)



- Troponin data suggests vepoloxamer can preserve cardiomyocyte by limiting calcium entry into the cell.
- Functional improvement supported by significant reductions of NT-proBNP for up to 2 weeks.

Study conducted by: Hani N. Sabbah  
Henry Ford Health System

# Heart Failure Phase 2 Study Design

- **Randomized, double-blind, placebo-controlled, multi-center Phase 2 study in chronic heart failure ongoing**
- **N=150 patients, 3 dose arms, single 3-hour administration**
- **Efficacy assessments:**
  - Cardiac function
  - Biomarkers
  - Exercise tolerance
- **Patient enrollment ongoing**

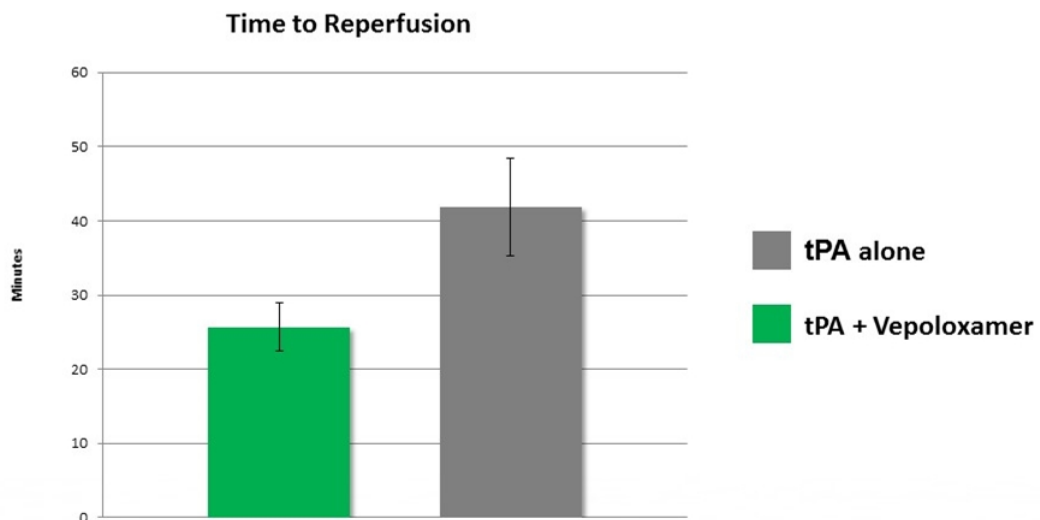
# Development of Vepoloxamer in Ischemic Stroke

**Objective: Accelerate reperfusion and reduce reperfusion injury**

**Phase 2-Ready**

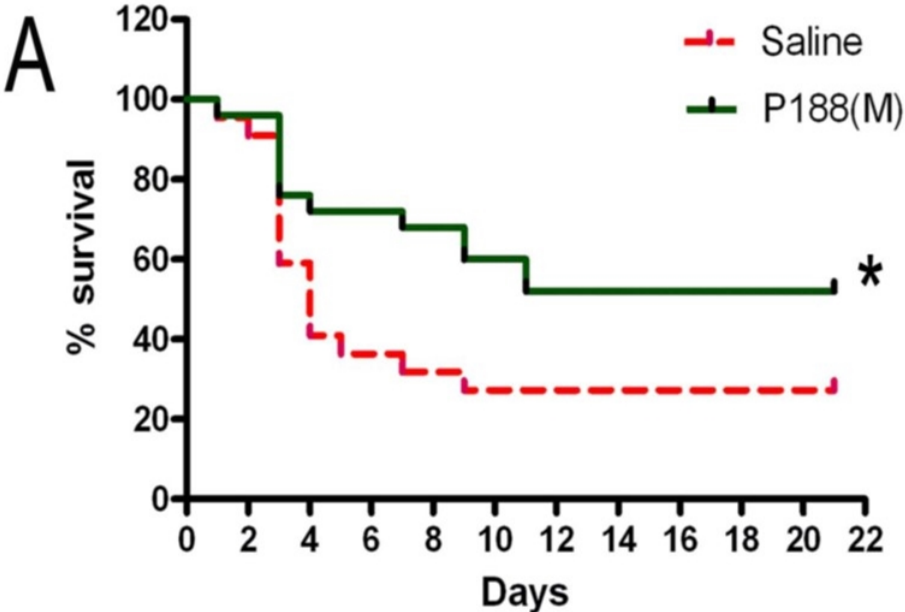
# Vepoloxamer in Ischemic Stroke

- **In stroke, restoring blood flow is critical (“time is tissue”)**
  - Vepoloxamer improves blood flow as a stand-alone agent
  - In combination with a thrombolytic, vepoloxamer shortens time to thrombolysis by up to 40%
  - Seals and protects ischemia-injured tissue
    - Neuronal tissue
    - Blood brain barrier integrity



# Vepoloxamer Alone or with tPA Improves Outcomes in Experimental Stroke Models

- Two hour occlusion of MCA with silicon coated nylon suture
- Only 27% of control mice survived vs. 52% of mice treated with poloxamer 188\* (n=51)



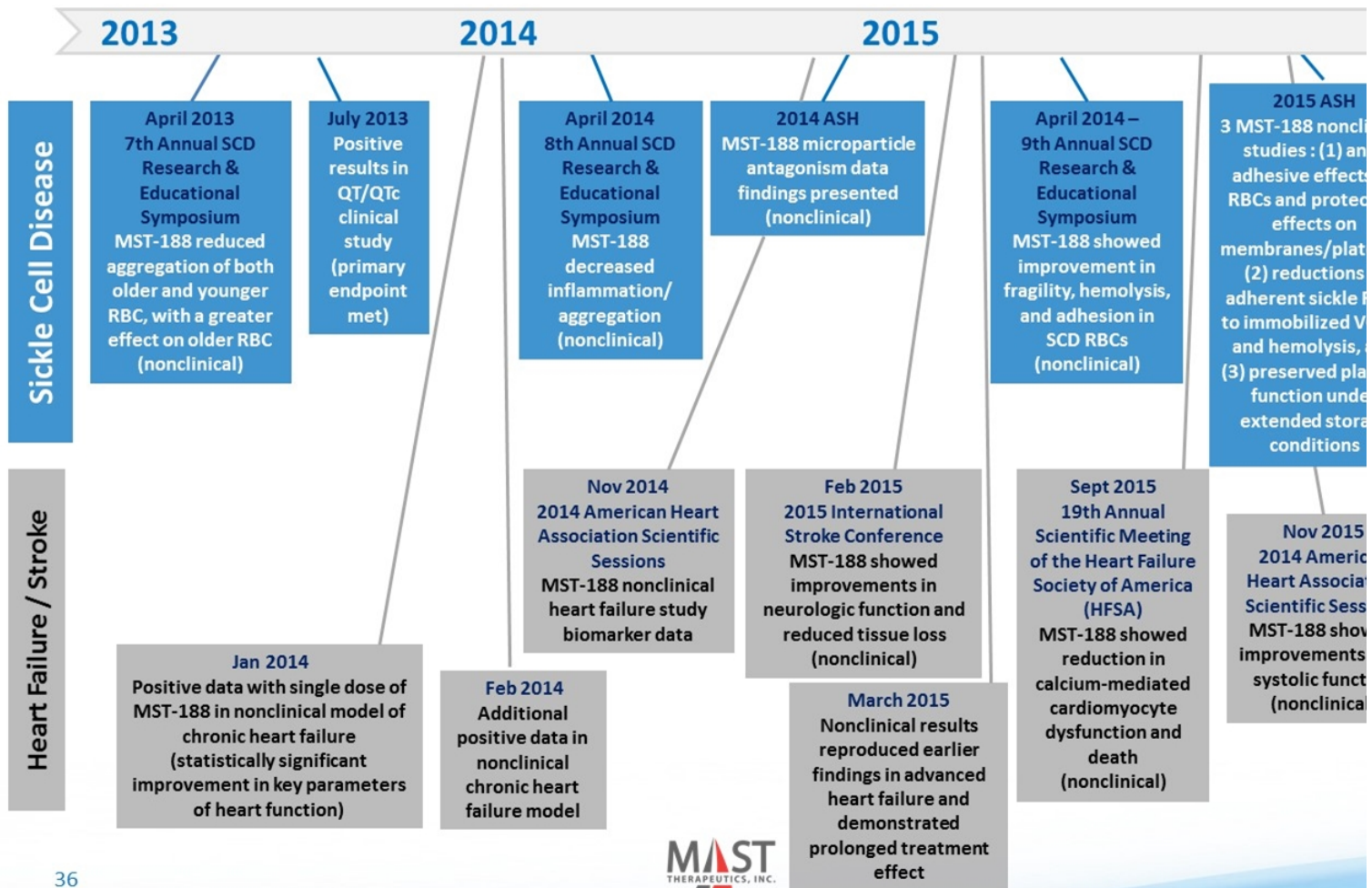
PLOS ONE 2013 (8) 4: e61641  
\* Vepoloxamer is purified poloxamer 188

# SBIR Grant-Funded Research

- **SBIR grant awarded by NIH in August 2016**
- **Phase 1 nonclinical study**
  - Evaluate the effect of a combination treatment with vepoloxamer and tissue plasminogen activator (tPA) on infarct volume and functional outcome in a nonclinical model of embolic stroke



# Key Data Generated by Mast Following Acquisition of Vepoloxamer



# Vepoloxamer Market Exclusivity

Form of Protection	Indication	Status
Orphan Drug Designation (Market Exclusivity)	SCD	Granted (US/EU)
Patents – Composition of matter	SCD, HF, IS	US 9,403,941 (to 2035), also pending w/w
Patents – New formulation	HF	Filed, pending w/w
Patents – Methods of use	SCD, HF, IS	Filed, pending w/w
Data Exclusivity	SCD, HF, IS	Eligible (US/EU)
Trade Secret & Know-How	SCD, HF, IS	Varies

SCD = sickle cell disease  
 HF = heart failure  
 IS = ischemic stroke

# Development of AIR001 in Heart Failure

**Objective: Improve exercise tolerance and hemodynamics in HFpEF patients**

## Multiple Phase 2 Studies

# AIR001 Overview

## ➤ AIR001 is nitrite\* for intermittent inhalation

- Delivered via proprietary handheld nebulizer
- Activity includes dilation of blood vessels and reduced inflammation
  - Not limited to role as nitric oxide donor as nitrite has direct mitochondrial oxygen-sparing activity
- Hemodynamic benefits include reductions in:
  - Pulmonary capillary wedge pressure
  - Right atrial pressure
  - Mean pulmonary arterial pressure
- Safety data available in approximately 140 subjects (well-tolerated)

\* Note: Nitrite is a different molecule and has separate activity compared to organonitrates or nitric oxide.

# AIR001 Clinical Data

## ➤ **Three Phase 1 studies**

- Established Maximum Tolerated Dose (MTD)
- Acute improvements in hypoxia-induced pulmonary hypertension
- No drug-drug interaction with sildenafil

## ➤ **Phase 2a study in PAH (n=29)**

- Well-tolerated; no treatment-related serious adverse events
- Improvements seen in median pulmonary vascular resistance (PVR) and median distances in 6-minute walk test
- Methemoglobin levels remained normal (<1.5%)

## ➤ **Phase 2a study in HFpEF (n=30)**

- Met primary endpoint; statistically significant decrease in pulmonary capillary wedge pressure during exercise compared to placebo
- Well-tolerated; no treatment-related serious adverse events

# AIR001 Clinical Development Plan

- **Heart Failure with Preserved Ejection Fraction (HFpEF)**
  - Responsible for ~50% of heart failure hospitalizations
  - No approved medications
  
- **Supporting ongoing institutional-sponsored Phase 2a study**
  - Evaluating hemodynamic effects – interim data expected mid-2016
  - Reported positive interim data in May 2016
  
- **AIR001 recently selected by the Heart Failure Clinical Research Network for Phase 2 study in HFpEF**
  - n=100
  - Multicenter, randomized, double-blind, placebo-controlled
  - First patient enrolled July 2016

# MSTX Financial Overview

- **Cash/investments at 6/30/2016: \$35.1 million**
- **Principal debt balance: \$14.1 million\***
  - \$10 million prepayment on 10/14/16 if EPIC data not positive or not announced
- **Market capitalization: ~\$72 million\***
- **Shares outstanding: ~ 212 million\***
- **Average daily volume (3 mo.): ~ 3.3 million\***

# Management Team and Board of Directors

## Management Team

**Brian Culley, CEO**

**Ed Parsley, CMO**

**Brandi Roberts, CFO**

**Martin Emanuele, SVP Development**

**Greg Gorgas, SVP Commercial**

**Mark Longer, VP Regulatory**

**Neurocrine, UC San Diego, The Scripps Research Inst.**

**Aires, Pfizer, CSL, Encysive, U. Texas Medical**

**Alphatec, Artes, Stratagene, Pfizer, PwC**

**DaVita, SynthRx, Kemia, Avanir, DuPont**

**Theragence, Biogen Idec, Chiron, Cetus, Upjohn Co.**

**AstraZeneca, Amylin, Pfizer**

## Board of Directors

**Matthew Pauls, CEO**

**Peter Greenleaf, CEO**

**Brian Culley, CEO**

**Howard Dittrich, EIR, CMO**

**David Ramsay, CFO (ret.)**

**StrongBridge Biopharma**

**Sucampo Pharmaceuticals**

**Mast Therapeutics**

**Frazier Healthcare Partners**

**Halozyme**



# Key Takeaways and Investment Highlights

## 1. Mast is the leader in sickle cell disease

- Potential first-in-class therapy for an orphan disease with high unmet need
- Enrollment completed in pivotal Phase 3; top-line data expected September
- More than 2 years ahead of nearest competitor
- Extensive patient-focused activity:
  - Created the leading SCD app, VOICE Crisis Alert (>3000 downloads)
  - Created the (5<sup>th</sup> Annual) SCD Drug Development Conference
  - Sponsor and volunteer at charity events, SCD radio show, etc.

## 2. Vepoloxamer has potential in other serious vascular diseases, including heart failure and stroke

## 3. Encouraging clinical data generated from AIR001 for heart failure

- Met primary endpoint in 30-patient randomized blinded trial
- Selected by HFN as subject of 100-patient Phase 2 study in HFpEF



