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**UNITED STATES  
SECURITIES AND EXCHANGE COMMISSION**

Washington, D.C. 20549

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**FORM 8-K**

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

Date of Report (Date of earliest event reported):  
**December 17, 2019**

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**MILESTONE PHARMACEUTICALS INC.**

(Exact name of registrant as specified in its charter)

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**Québec**  
(state or other jurisdiction of incorporation)

**001-38899**  
(Commission File Number)

**Not applicable**  
(I.R.S. Employer Identification No.)

**1111 Dr. Frederik-Philips  
Boulevard, Suite 420  
Montréal, Québec CA**  
(Address of principal executive offices)

**H4M 2X6**  
(Zip Code)

Registrant's telephone number, including area code: **(514) 336-0444**

(Former name or former address, if changed since last report.)

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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 Instruction 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))

Securities registered pursuant to Section 12(b) of the Act:

**Title of each class**  
Common Shares

**Trading Symbol(s)**  
MIST

**Name of each exchange on which registered**  
The Nasdaq Stock Market LLC

Indicate by check mark whether the registrant is an emerging growth company as defined in as defined in Rule 405 of the Securities Act of 1933 (§ 230.405 of this chapter) or Rule 12b-2 of the Securities Exchange Act of 1934 (§ 240.12b-2 of this chapter).

Emerging growth company

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act.

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**Item 8.01. Other Events.**

On December 17, 2019, Milestone Pharmaceuticals, Inc. (the "Company") updated its corporate presentation that it intends to use in connection with presentations at conferences and meetings. The full text of the Company's corporate presentation is filed as Exhibit 99.1 hereto and incorporated herein by reference.

**Item 9.01. Financial Statements and Exhibits.**

**(d) Exhibits**

**Exhibit**

**No.**

**Description**

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<a href="#">99.1</a>	<a href="#">Corporate Presentation dated December 17, 2019.</a>
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**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.

**MILESTONE PHARMACEUTICALS INC.**

By: /s/ Amit Hasija  
Amit Hasija  
Chief Financial Officer

Dated: December 17, 2019

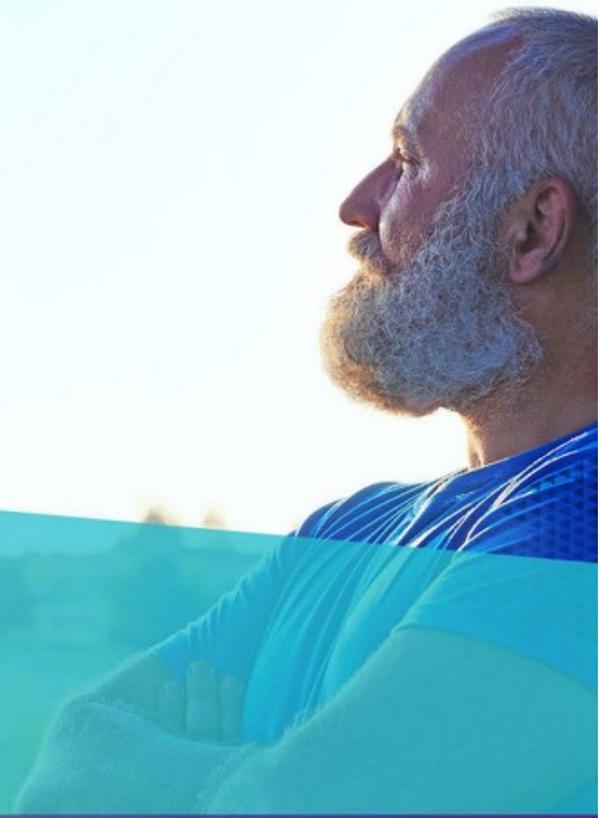
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**Milestone**  
PHARMACEUTICALS

**Corporate Overview**  
**December 2019**

**Joseph Oliveto**  
**Chief Executive Officer**



# Disclaimers



This Presentation contains forward-looking statements within the meaning of the safe harbor provisions of the Private Securities Litigation Reform Act of 1995, as amended. In some cases, you can identify forward-looking statements by terminology such as “aim,” “anticipate,” “assume,” “believe,” “contemplate,” “continue,” “could,” “design,” “due,” “estimate,” “expect,” “goal,” “intend,” “may,” “objective,” “plan,” “predict,” “positioned,” “potential,” “seek,” “should,” “target,” “will,” “would” and other similar expressions that are predictions of or indicate future events and future trends, or the negative of these terms or other comparable terminology. These forward-looking statements include, but are not limited to, statements about the initiation, timing, progress and results of our current and future clinical trials of etripamil, including our Phase 3 clinical trials of etripamil for the treatment of paroxysmal supraventricular tachycardia (“PSVT”), and of our research and development programs and clinical pipeline; our plans to develop and commercialize etripamil and any future product candidates; the expected benefits of using etripamil to treat PSVT; our expectations regarding the potential market size and the rate and degree of market acceptance of etripamil and any future product candidates and the implementation of our business model and strategic plans for our business, etripamil and any future product candidates. Such forward-looking statements are subject to inherent uncertainties, risks and assumptions that are difficult to predict. Factors that could cause actual results to differ include, but are not limited to, our dependence on the success of our Phase 3 clinical trials of etripamil for PSVT, the risks inherent in biopharmaceutical product development and clinical trials, including the lengthy and uncertain regulatory approval process, uncertainties related to the timing of initiation, enrollment and completion of clinical trials, and whether the clinical trials will validate the safety and efficacy of etripamil for PSVT or other indications, among others. These and other risks and uncertainties are described more fully in our annual and other periodic filings with the Securities and Exchange Commission (the “SEC”), including under the heading “Risk Factors” in our Quarterly Report on Form 10-Q filed with the SEC on November 13, 2019. We may not actually achieve the plans, intentions or expectations disclosed in our forward-looking statements, and you should not place undue reliance on our forward-looking statements. Actual results or events could differ materially from the plans, intentions and expectations disclosed in the forward-looking statements we make. We undertake no obligation to update or revise any forward-looking statements, whether as a result of new information, the occurrence of certain events or otherwise.

This Presentation contains trademarks, trade names and service marks of other companies, which are the property of their respective owners. Certain information contained in this Presentation and statements made orally during this Presentation relate to or is based on studies, publications, surveys and other data obtained from third-party sources and the Company’s own internal estimates and research. While the Company believes these third-party studies, publications, surveys and other data to be reliable as of the date of the Presentation, it has not independently verified, and makes no representation as to the adequacy, fairness, accuracy or completeness of, any information obtained from third-party sources. In addition, no independent sources has evaluated the reasonableness or accuracy of the Company’s internal estimates or research and no reliance should be made on any information or statements made in this Presentation relating to or based on such internal estimates and research.

## Milestone (Nasdaq: MIST) - Corporate Highlights



- Phase 3 Cardiovascular Company with data read out anticipated in middle 1H, 2020
- PSVT is a robust market represented by ~2M patients in US
- Paradigm-changing approach enabling patient self-management
- Potentially first new drug therapy in PSVT in > 25 years
- New Chemical Entity with proprietary IP protection until 2036
- Pipeline opportunities beyond the lead indication
- \$95M Initial Public Offering - May 13, 2019
- Cash & equivalents of \$136.5M (Sept. 30, 2019) – expected runway into Q3, 2021

PSVT = Paroxysmal Supraventricular Tachycardia

# Management Team



**Joseph Oliveto**  
Chief Executive Officer



**Amit Hasija**  
Chief Financial Officer



**Francis Plat, MD**  
Chief Medical Officer



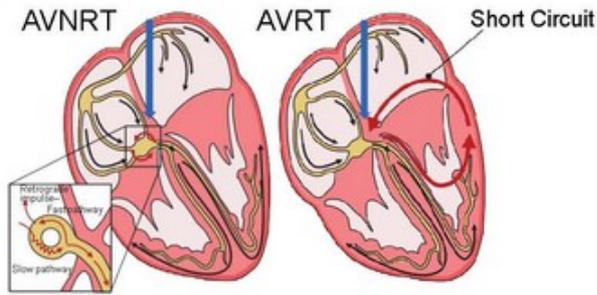
**Lorenz Muller**  
Chief Commercial Officer



**Philippe Douville, PhD**  
Chief Scientific Officer / Founder

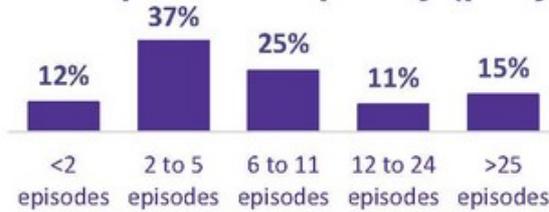


# Paroxysmal Supraventricular Tachycardia (PSVT)



- PSVT is a rapid heart rate condition that starts and stops without warning
- Heart rates >200 bpm are not uncommon
- Symptoms include
  - ✓ palpitations
  - ✓ sweating
  - ✓ chest pressure or pain, shortness of breath
  - ✓ sudden onset of fatigue
  - ✓ lightheadedness or dizziness
  - ✓ fainting or anxiety

## PSVT episode frequency (per yr.)



AVNRT = Atrioventricular Nodal Re-entrant Tachycardia AVRT = Atrioventricular Re-entrant Tachycardia bpm = beats per minute

Sources: Internal estimates based on market research

# Current Standard of Care for PSVT



**Current acute treatment options are invasive, inconvenient, anxiety-provoking and/or costly**

<b>Chronic / preventive</b>		<ul style="list-style-type: none"><li>• Chronic oral medication with modest efficacy and unpleasant side effects</li><li>• 4-7 episodes/year despite preventive medications</li></ul>
<b>Chronic / preventive</b>		<ul style="list-style-type: none"><li>• Catheter ablation</li><li>• ~80K ablations/year</li><li>• Only ~10% of patients opt for ablation</li></ul>
<b>Acute</b>		<ul style="list-style-type: none"><li>• IV adenosine or DC cardioversion in the ED</li><li>• &gt;150K ED visits/hospital admissions per year</li><li>• Many patients endure episodes when they occur</li></ul>

PSVT = Paroxysmal Supraventricular Tachycardia DC = Direct Current ED = Emergency Department

Sources: Internal estimates based on market research and longitudinal analysis of Truven/Marketscan and Medicare claims data; Page RL et al. 2015 ACC/AHA/HRS guideline for the management of adult patients with supraventricular tachycardia: executive summary: a report of the ACC/AHA Task Force on Clinical Practice Guidelines and the Heart Rhythm Society, *Circulation*. 2016;133:e471–e505

# A Paradigm-Changing Approach



**Opportunity to develop the first approved treatment to be used by patients whenever and wherever an episode of PSVT occurs**

Non-invasive

Convenient

Empowering

- Avoidance of ED visits/ hospital admissions
- Less need for chronic medications
- Alternative or bridge to ablation procedure



PSVT = Paroxysmal Supraventricular Tachycardia

# Etripamil



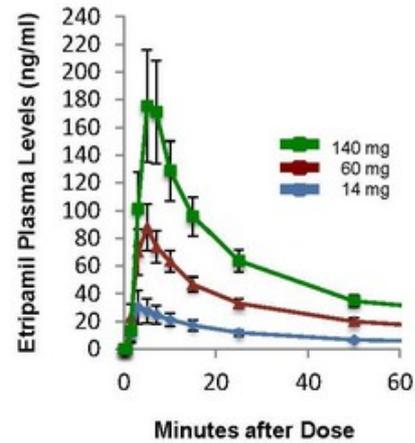
## A paradigm-changing approach for treating PSVT

Etripamil	
Class	Novel CCB
Potency (IC <sub>50</sub> )	11 nM
Metabolism	Rapid: Esterase-mediated

- **Clinically-validated mechanism**
  - Etripamil, Calcium Channel Blockers (CCBs), terminate PSVT through AV node modulation
- **Rapid onset of action**
- **Convenient patient self-administered nasal spray**
- **Short half-life**

AV = Atrio-ventricular

- **Rapid onset ( $T_{max} < 5 \text{ min}$ )**
- **Transient plasma levels**

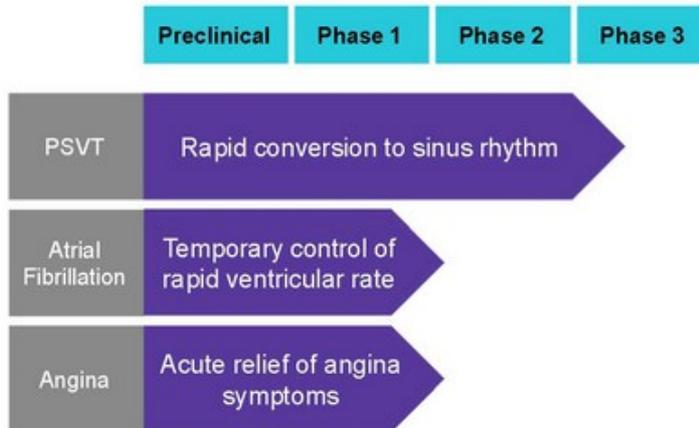


Error bars indicate standard error of the mean.

# Etripamil Clinical Pipeline



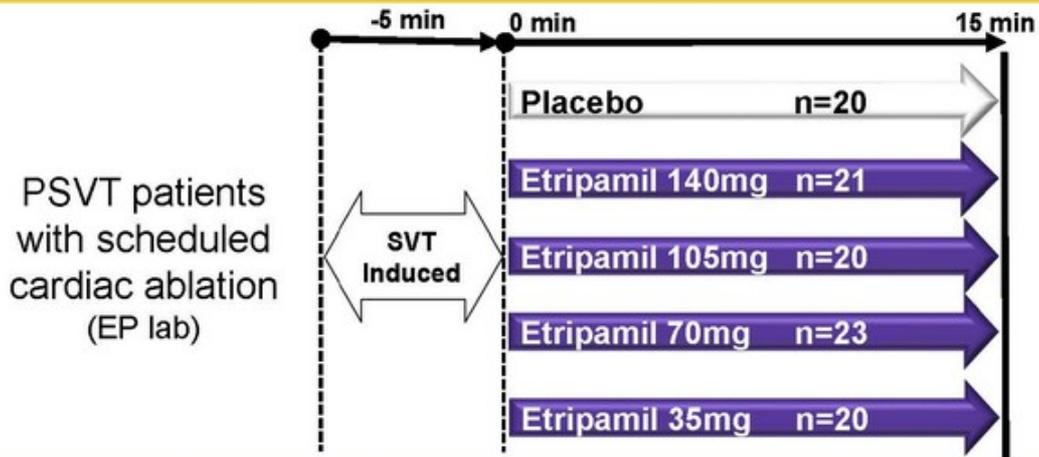
Pharmacology of L-type calcium channel blockers drives broad clinical utility



# Phase 2a/b Study Design



**Objectives: Demonstrate superiority of etripamil over placebo in terminating SVT and dose ranging trend analysis**



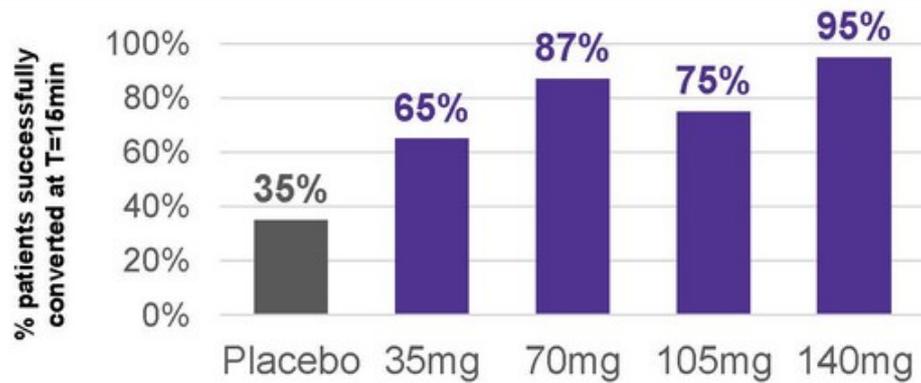
**Endpoint: conversion to sinus rhythm within 15 minutes**  
**>80% power to show a 50% absolute difference vs. placebo**

EP = electrophysiology, SVT = supraventricular tachycardia, PSVT = Paroxysmal Supraventricular Tachycardia

## Phase 2 Primary Endpoint



Etripamil three highest doses demonstrated 75-95% conversion rates which are statistically significant compared to placebo



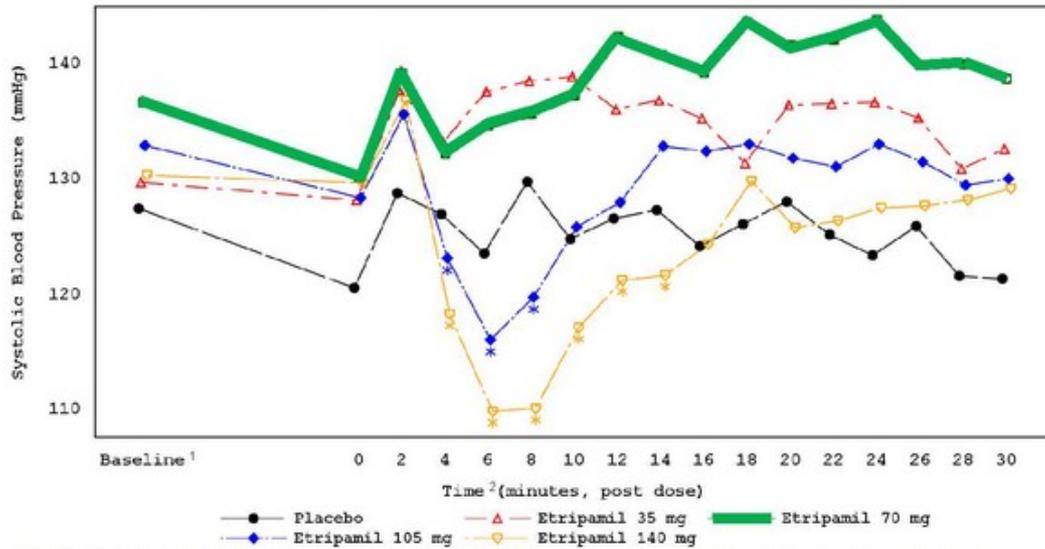
# patients converted at 15 min	7/20	13/20	20/23	15/20	20/21
p-value		0.1128	0.0006	0.0248	<.0001

Source: Stambler, B.S. et al.; Etripamil Nasal Spray for Rapid Conversion of Supraventricular Tachycardia to Sinus Rhythm; J Am Coll Cardiol. 2018;72(5):489-97

# Phase 2 Mean Systolic Blood Pressure Effects



**Etripamil 70 mg showed no drop in blood pressure**



<sup>1</sup> Baseline is defined as the average of the 20-min and 10-min pre-dose measurements. <sup>2</sup> Time 0 is defined as the average of the measurements during supraventricular tachycardia between 5 and 0 min before study drug administration. \*p < 0.05 versus baseline.

Source: Stambler, B.S. et al.; Etripamil Nasal Spray for Rapid Conversion of Supraventricular Tachycardia to Sinus Rhythm; J Am Coll Cardiol. 2018;72(5):489-97

## Phase 2a/b Clinical Conclusions



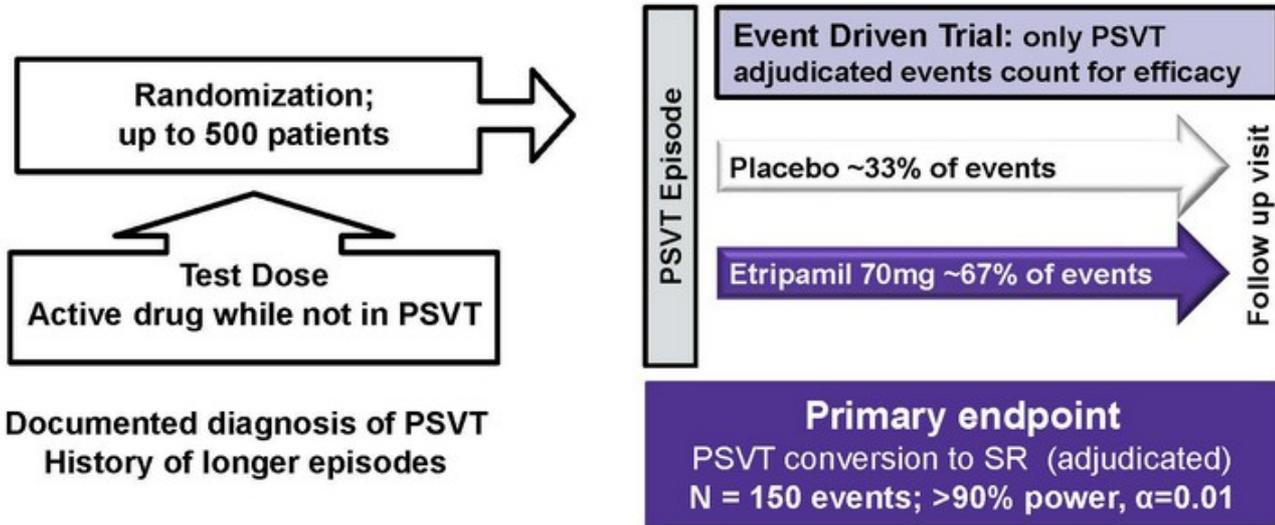
- Etripamil at 70, 105 and 140 mg is significantly better than placebo in terminating PSVT
- Median time to conversion <3 min with etripamil 70 mg
- 70 mg dose showed no mean blood pressure (BP) drop
- Most frequent side effect was nasal irritation or nasal congestion; however these were transient
- Etripamil 70 mg demonstrated the best efficacy/safety profile to take into Phase 3

Source: Adapted from Stambler, B.S. et al.; Etripamil Nasal Spray for Rapid Conversion of Supraventricular Tachycardia to Sinus Rhythm; J Am Coll Cardiol. 2018;72(5):489-97

# Pivotal Phase 3 Study Design



**Objective: Superiority of etripamil over placebo in terminating PSVT events in the outpatient setting**



SR = Sinus Rhythm; PSVT = Paroxysmal Supraventricular Tachycardia; Study randomization scheme 2:1 etripamil : placebo



## NODE-301

### Single pivotal efficacy study to support NDA submission

- Once target of 150 adjudicated events reached, collection of blinded data from patients who have not experienced an event to continue as separate dataset called "NODE-301B"

## NODE-303

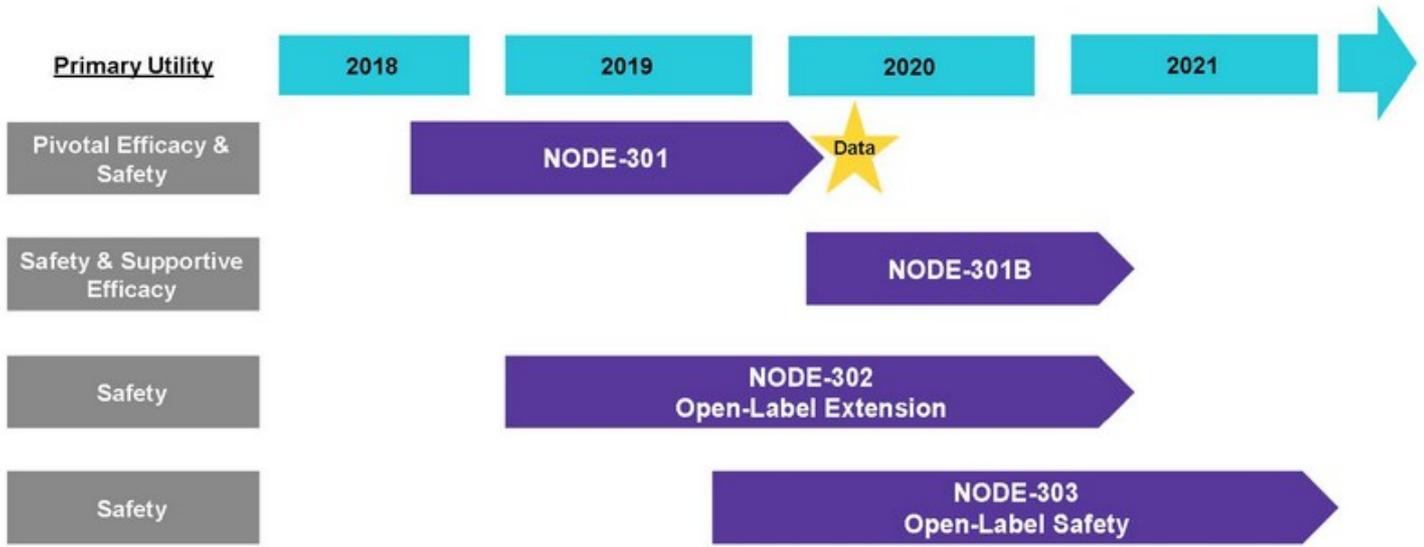
### Open-label global safety trial

- Removed the in-office safety test dose that is currently required in the NODE-301 study

## Population and Safety Database

- Program enrolling broad patient population including elderly and those on concomitant medications (e.g. calcium channel blockers and beta blockers)
- Total NDA safety data set of  $\leq 1,500$  unique patient events

# Etripamil PSVT Phase 3 Development Plan



PSVT = Paroxysmal Supraventricular Tachycardia

# PSVT Patient Characteristics



- Age: teens to elderly
- Gender: majority are female
- Episode frequency and duration varies widely
  - Median 4-7 per year despite chronic medications
  - Almost 40% of patients have at least 2 episodes/year >10 min\*
- Cardiovascular comorbidities in about half of patients
- 40% of patients have  $\geq 1$  ED visit per year\*



## Unmet Need

- Strongly negative experience associated with **adenosine in ED**
- Significant anxiety/fear of **ablation**
- Patients indicate **“significant impact”** on QOL

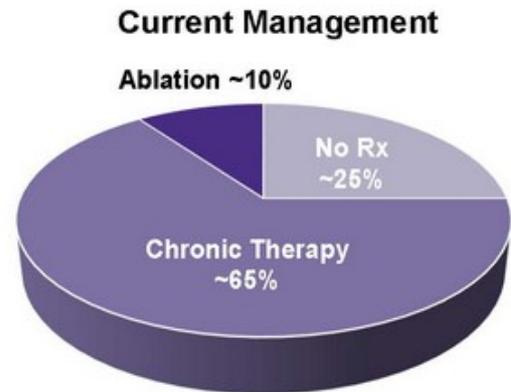
\*Estimates are for patients in year after initial diagnosis; rates drop by 13-29% in years following their initial diagnosis  
Sources: Internal estimates based on market research and longitudinal analysis of Truven/Marketscan and Medicare claims data

# Current US PSVT Market



Total annual US healthcare expenditures of ~\$3B

- Prevalence ~2M diagnosed PSVT patients
- ~300K newly diagnosed per year
- ~600K patients treated per year
- >150K ED/hospital visits per year
- ~80K ablations per year



Source: Sacks, N.C. et al; Prevalence of Paroxysmal Supraventricular Tachycardia (PSVT) in the US in Patients Under 65 Years of Age; Abstract and Oral Presentation at the International Academy of Cardiology Annual Scientific Sessions 2018, 23rd World Congress on Heart Disease; Precision Xtract, Boston, MA, USA; and data-on-file from IBM MarketScan® Commercial Research Database (<65y) and the Medicare Limited Dataset (≥65y), with demographic, enrollment and claims data for commercially insured (Truven) and Medicare covered patients using PSVT code 427.0 or I47.1 for up to a 9-year interval between 2008 and 2016 inclusive.

# Estimating Prevalence, Incidence, and Annually Treated Patients Using Longitudinal Claims Data



- Analyzed commercial and Medicare claims data over a 9-year period, where patients were required to have 5 years of continuous enrollment
  - ✓ 1+ PSVT code required in the ED or inpatient setting (unique patients managed acutely)
  - ✓ 2+ PSVT codes required in the outpatient setting (additional unique patients managed chronically)

Age Group	Year 1	Year 2	Year 3	Year 4	Year 5	Total
Age < 65	271,024	196,653	169,988	155,966	145,485	939,116
Age 65+	377,493	220,596	209,358	188,925	166,286	1,162,658
All Ages	648,518	417,249	379,346	344,891	311,771	2,101,775

↑

Annually Treated  
PSVT Patients

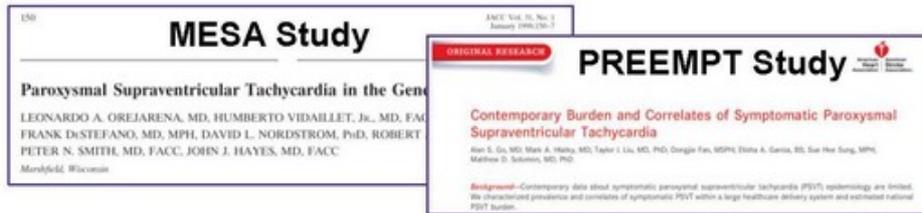
↑

Incident PSVT  
Patients

↑

Prevalent  
PSVT Patients

Source: Data on file from IBM MarketScan® Commercial Research Database (<65y) and the Medicare Limited Dataset (≥65y), with demographic, enrollment and claims data for commercially insured (Truven) and Medicare covered patients using PSVT code 427.0 or I47.1 for up to a 9-year interval between 2008 and 2016 inclusive.



## Strengths

- Provides important demographic and clinical characteristic data on patients with PSVT
- Positive Predictive Values from PREEMPT useful
- Less than 40% of incident cases in MESA would have been detected by PSVT ICD-9 Code 427.0

## Weaknesses

- Data only from patients presenting to healthcare settings acutely, with the episode confirmed on ECG during the encounter
- PSVT episodes were only adjudicated during the first healthcare encounter with a PSVT or PSVT-related code in PREEMPT
- Non-representative, small, and non-contemporary population (MESA)

Source: Orejarena LA, Vidaillet H Jr, DeStefano F, Nordstrom DL, Vierkant RA, Smith, PN, Hayes JJ. Paroxysmal supraventricular tachycardia in the general population. *J Am Coll Cardiol.* 1998;31:150–157. Alan S. Go, MD; Mark A. Hlatky, MD; Taylor I. Liu, MD, PhD; Dongjie Fan, MSPH; Elisha A. Garcia, BS; Sue Hee Sung, MPH; Matthew D. Solomon, MD, PhD. Contemporary Burden and Correlates of Symptomatic Paroxysmal Supraventricular Tachycardia. *J Am Heart Assoc.* 2018;7:e008759. DOI: 10.1161/JAHA.118.008759.

# Healthcare Cost and Utilization for PSVT



## Prevalence of Healthcare Utilization for PSVT

2014	Inpatient and Outpatient ablations*	Non cardiac ablation PSVT encounter					Total
		Hospitalizations with PSVT	Emergency Department Visits with PSVT	Outpatient Hospital Visits with PSVT	Outpatient Other	Office visits with PSVT	
HRU	79,347	103,865	51,048	260,207	503,269	1,174,087	
Cost (\$)	<u>\$24,071</u>	<u>\$8,860</u>	<u>\$1,003</u>	<u>\$853</u>	<u>\$195</u>	<u>\$123</u>	
Total (\$)	\$1.9B	\$920M	\$51M	\$221M	\$98M	\$144M	\$3.3B

\*Ablations are for patients with a PSVT diagnosis in the same year. Mean costs for inpatient and outpatient ablations reflect claims with ablation procedure code and inpatient and outpatient hospital settings.

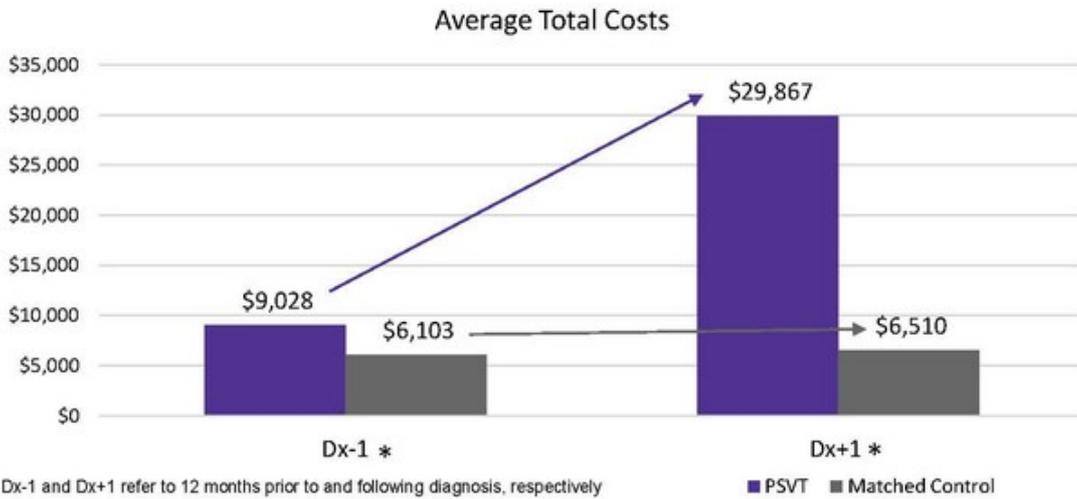
Source: Data on file from IBM MarketScan® Commercial Research Database (<65y) and the Medicare Limited Dataset (≥65y), with demographic, enrollment and claims data for commercially insured (Truven) and Medicare covered patients using PSVT code 427.0 or I47.1

HRU = Healthcare Resource Utilization

# Post-Diagnosis Total Healthcare Spending for Newly-Diagnosed PSVT Patients <65 Years Old Relative to Matched Controls

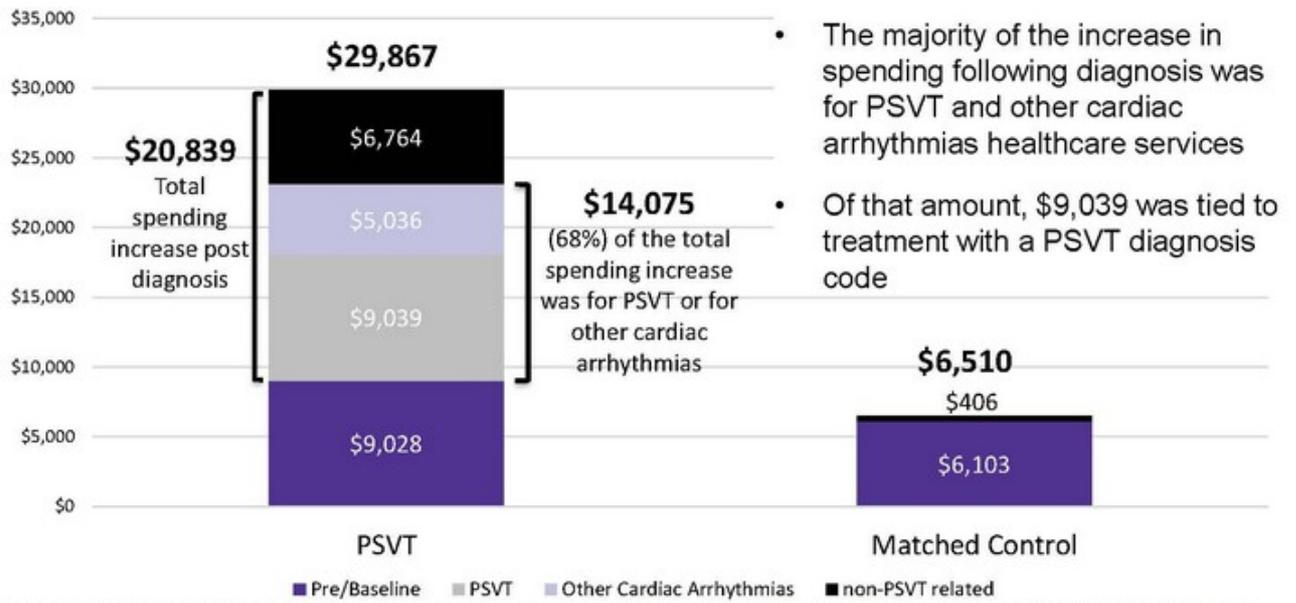


**In the year following diagnosis, total spending more than tripled for PSVT patients**



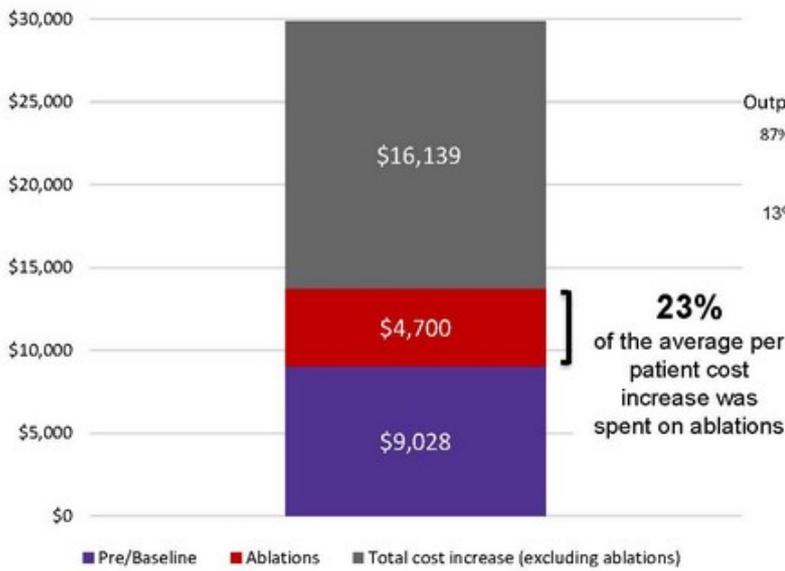
Source: Sacks, N et al.; "Healthcare Resource Use and Expenditures in Patients under 65 Years of Age and Newly Diagnosed with Paroxysmal Supraventricular Tachycardia (PSVT) in the United States"; Podium presentation at the International Academy of Cardiology, Annual Scientific Sessions 2018; 23rd World Congress on Heart Disease; July 2018

# Components of Spending in Newly-Diagnosed Patients <65 for PSVT and Other Cardiac Arrhythmias Healthcare Services



Source: Sacks, N et al.; "Healthcare Resource Use and Expenditures in Patients under 65 Years of Age and Newly Diagnosed with Paroxysmal Supraventricular Tachycardia (PSVT) in the United States"; Podium presentation at the International Academy of Cardiology, Annual Scientific Sessions 2018; 23rd World Congress on Heart Disease; July 2018

# Mean Cost per PSVT Ablation and Contribution to Average per Patient Cost Increase in Year after Diagnosis in Patients <65 Years Old



All settings*	0.14
Outpatient Hospital	0.12
Inpatient	0.02

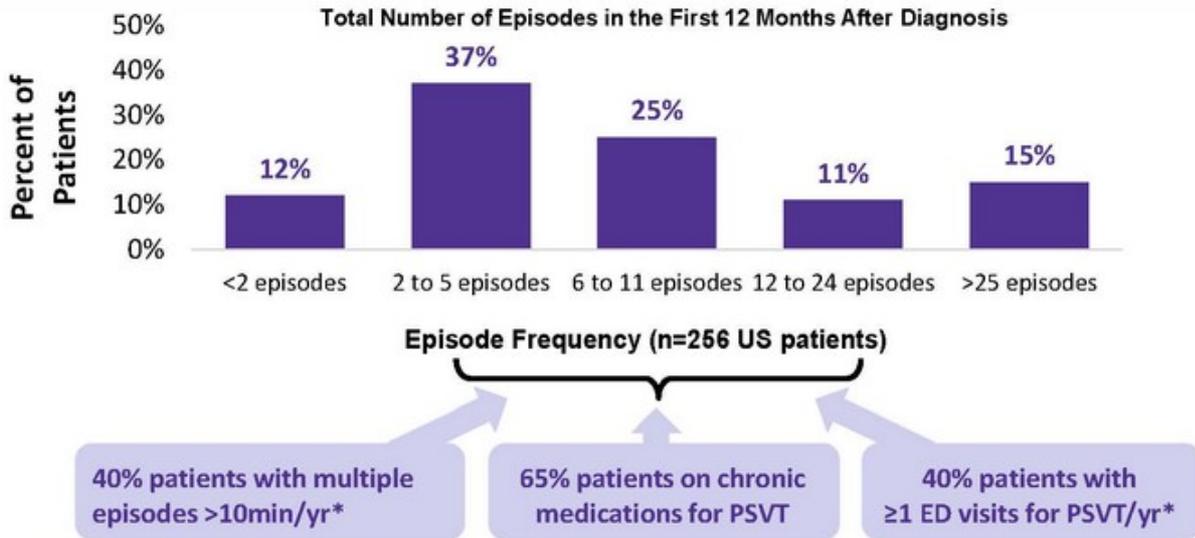
\* Includes ablations in unspecified outpatient settings

Source: Sacks, N et al.; "Healthcare Resource Use and Expenditures in Patients under 65 Years of Age and Newly Diagnosed with Paroxysmal Supraventricular Tachycardia (PSVT) in the United States"; Podium presentation at the International Academy of Cardiology, Annual Scientific Sessions 2018; 23rd World Congress on Heart Disease; July 2018

# Target Addressable Market for PSVT



Market research suggests TAM for PSVT of >800k patients



TAM – Target Addressable Market

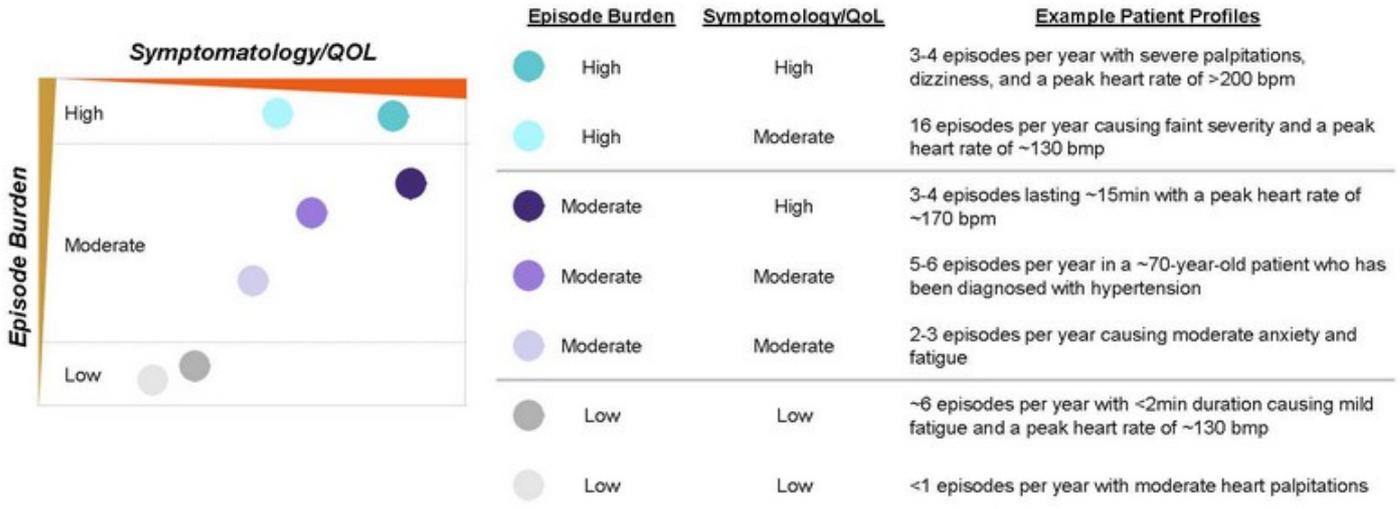
\*Estimates are for patients in year after initial diagnosis; rates drop by 13-29% in years following their initial diagnosis

Sources: Internal estimates based on market research

# Examples of PSVT Patient Profiles



Patients with PSVT present with a large range of episode and quality of life burdens

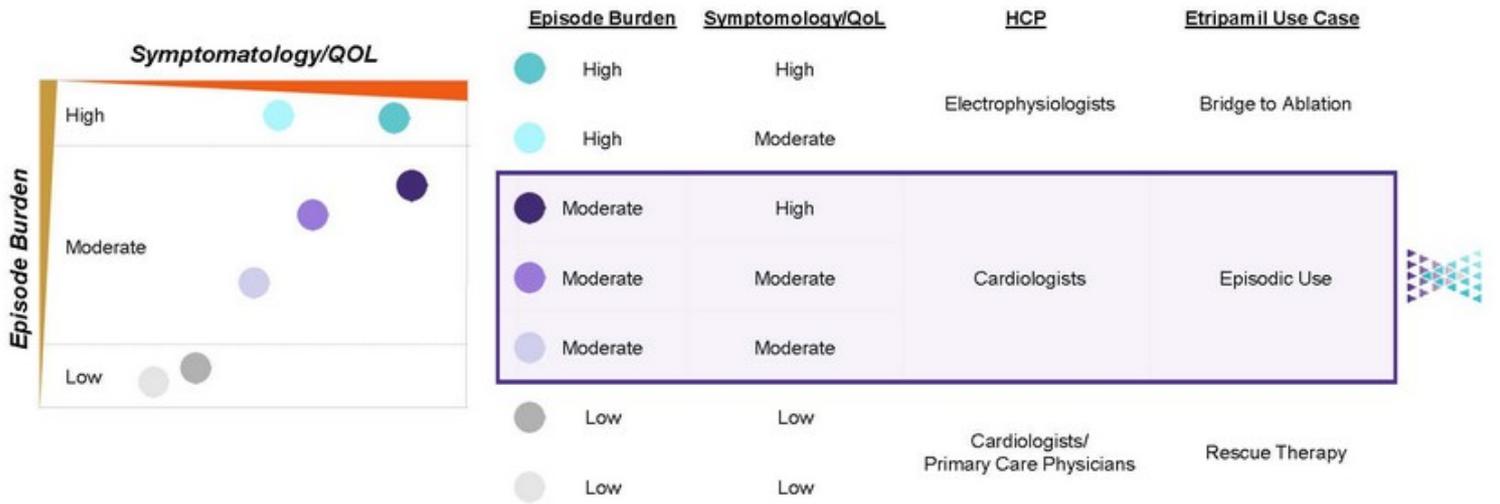


Sources: Internal market research

# Etripamil Use Case and Target Prescriber



Patients experiencing moderate episode burden are the anticipated target user of etripamil



Sources: Internal market research

# Etripamil – Addressing Market Needs



## Potential for high receptivity to the etripamil profile across stakeholders

<b>Patients</b> 	<b>Physicians (Cards, EPs, PCPs)</b> 	<b>Payors</b> 
<b>Future with Etripamil– a Better Treatment Option</b>		
<ul style="list-style-type: none"> <li>• Self-management of acute episodes</li> <li>• Less need for chronic medications</li> <li>• Avoidance of 50-75% of ED visits/hospital admissions</li> </ul>	<ul style="list-style-type: none"> <li>• Better risk/reward profile</li> <li>• Expected to have significant adoption in unablated patients</li> <li>• Alternative to ablation</li> <li>• Bridge to ablation</li> </ul>	<ul style="list-style-type: none"> <li>• Reduction in ED/hospital admissions</li> <li>• Deferral of ablation</li> <li>• Improvement in patient satisfaction</li> </ul>

Cards = Cardiologists, EPs = Electrophysiologists, PCPs = Primary Care Physicians

Sources: Internal market research

# MIST Healthcare Practitioner (HCP) Survey Results



Physician Reaction to Phase 2 Target Product Profile (TPP) for Etripamil							
 <b>Overall Receptivity from MIST Quantitative Survey (n=353)</b>	<p><i>"On a scale of 1-7, where 1 = not at all favorable and 7 = extremely favorable, please rate your receptivity to Product X."</i></p>  <p>Avg (mean): 5.6      All HCPs (n=353)</p>						
<b>Reaction to TPP from MIST Qualitative Survey (n=30)</b>	<ul style="list-style-type: none"> <li>• Potential for a fast onset of action and high conversion rates within 30 to 60 minutes of administration noted to be significantly better than current approaches (vagal maneuvers, pill-in-the-pocket)</li> <li>• HCPs familiar with and comfortable prescribing CCBs to the PSVT population</li> <li>• May require additional tests to rule out potential contraindications</li> </ul>						
<b>Potential Utilization from MIST Quantitative Survey (n=353)</b>	<p><i>"Of PVST patients not contraindicated to Product X, please estimate the share that would be prescribed Product X."</i></p>  <table border="1"> <tr> <td><b>Cards (n=253)</b></td> <td>54%</td> </tr> <tr> <td><b>EPs (n=50)</b></td> <td>47%</td> </tr> <tr> <td><b>PCPs (n=50)</b></td> <td>58%</td> </tr> </table>	<b>Cards (n=253)</b>	54%	<b>EPs (n=50)</b>	47%	<b>PCPs (n=50)</b>	58%
<b>Cards (n=253)</b>	54%						
<b>EPs (n=50)</b>	47%						
<b>PCPs (n=50)</b>	58%						

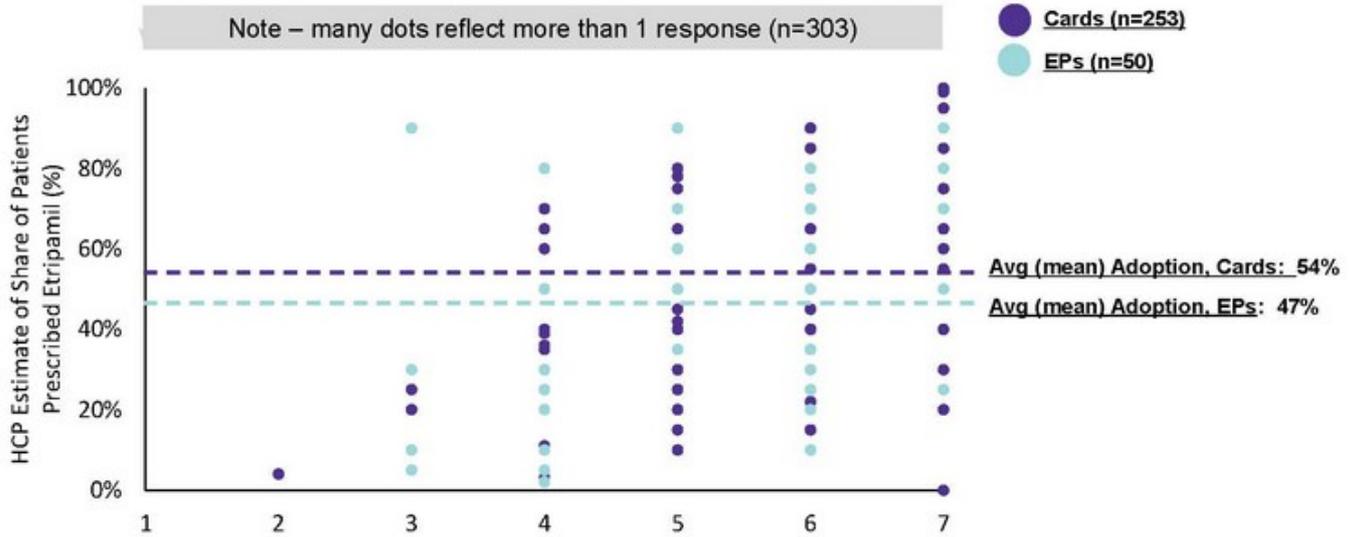
Source: Triangle Insights Group Qualitative (n=30) and Quantitative (n=353) Market Research with General Cardiologists, Electrophysiologists, and Primary Care Physicians, November 2018 through April 2019. MIST = Milestone Pharmaceuticals, Inc.

# Variability in Response for Cardiologist Stated Adoption by Favorability Score



## MIST Quantitative Demand Survey – Cards and EPs Physician Favorability Score v. Estimated Adoption Share

Note – many dots reflect more than 1 response (n=303)

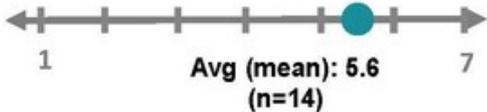
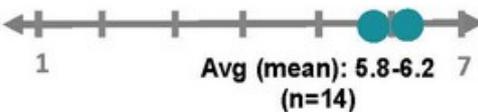


"On a scale of 1-7, where 1 = not at all favorable and 7 = extremely favorable, please rate your receptivity to Product X."

Source: Triangle Insights Group Qualitative (n=30) and Quantitative (n=353) Market Research with General Cardiologists, Electrophysiologists, and Primary Care Physicians, November 2018 through April 2019. MIST = Milestone Pharmaceuticals, Inc.

# MIST Payer Survey Results



 <b>Payer Reaction to Phase 2 Target Product Profile (TPP) for Etripamil</b>	
<b>Overall Receptivity from MIST Qualitative Survey (n=14)*</b>	<p><i>"On a scale of 1-7, where 1 = not at all favorable and 7 = extremely favorable, please rate your receptivity to Product X."</i></p>  <p>A horizontal scale from 1 to 7 with tick marks. A teal dot is placed at 5.6. Below the scale, it reads "Avg (mean): 5.6 (n=14)".</p>
<b>Likelihood of Coverage from MIST Qualitative Survey (n=14)*</b>	<p><i>"On a scale of 1-7, where 1 = not at likely and 7 = extremely likely, please rate your plan's likelihood to cover Product X (WAC \$500-\$1000)."</i></p>  <p>A horizontal scale from 1 to 7 with tick marks. Two teal dots are placed between 5 and 6, representing a range of 5.8-6.2. Below the scale, it reads "Avg (mean): 5.8-6.2 (n=14)".</p> <p><b>Payer rationale for reported likelihood of coverage</b></p> <ul style="list-style-type: none"> <li>✓ No other approved treatment options</li> <li>✓ Potential for fast onset of action and high conversion</li> <li>✓ Potential strong health economic message (cost offset of Emergency Department / inpatient admissions)</li> <li>✓ May want to see real world evidence in own population</li> </ul>

\*assuming demonstrated efficacy and no significant safety concerns

Source: Triangle Insights Group Qualitative (n=14) Market Research with Medical and Pharmacy Directors from both National and Regional Health Plans, April-June 2017  
 MIST = Milestone Pharmaceuticals, Inc.

# MIST Patient Survey Results



PSVT Patient Reaction to Phase 2 Target Product Profile (TPP) for Etripamil										
 <p><b>Overall Receptivity from MIST Qualitative Survey (n=20)</b></p>	<p><i>"On a scale of 1-7, where 1 = not at all favorable and 7 = extremely favorable, please rate your receptivity to Product X."</i></p> <div style="display: flex; align-items: center;">  <div style="margin-left: 20px;"> <ul style="list-style-type: none"> <li>• Patients had highest receptivity to etripamil of all stakeholders surveyed by MIST</li> <li>• Important because MIST anticipates patient-driven market dynamic</li> </ul> </div> </div>									
<p><b>Reaction to TPP from MIST Qualitative Survey (n=20)</b></p>	<ul style="list-style-type: none"> <li>• Estimated that patients would potentially have avoided ~50-75% of all Emergency Department visits</li> <li>• Expect etripamil to provide 'peace of mind' in between episodes and a sense of control over disease (reduces anxiety of the next event, allowing patients to perform activities that are limited without a reliable at-home therapy)</li> <li>• Minority of patients indicated an aversion to administering medications intranasally</li> </ul>									
<p><b>Expected Usage Rate by Patients from MIST Qualitative Survey (n=20)</b></p>	<p><i>"Please estimate the share of your PSVT episodes for which you would use Product X."</i></p> <div style="display: flex; align-items: center;"> <table border="0" style="margin-right: 20px;"> <tr> <td><b>\$60 OOP</b></td> <td><b>40%</b></td> <td><div style="width: 40%; height: 10px; background-color: #4a7ebb; border: 1px solid #ccc;"></div></td> </tr> <tr> <td><b>\$30 OOP</b></td> <td><b>60%</b></td> <td><div style="width: 60%; height: 10px; background-color: #999999; border: 1px solid #ccc;"></div></td> </tr> <tr> <td><b>\$10 OOP</b></td> <td><b>85%</b></td> <td><div style="width: 85%; height: 10px; background-color: #008080; border: 1px solid #ccc;"></div></td> </tr> </table> <div> <p>MIST would expect approximately 3-4 annual etripamil doses per patient based on patient stated usage rates at potential Tier 2 or Tier 3 out of pocket (OOP) costs</p> </div> </div>	<b>\$60 OOP</b>	<b>40%</b>	<div style="width: 40%; height: 10px; background-color: #4a7ebb; border: 1px solid #ccc;"></div>	<b>\$30 OOP</b>	<b>60%</b>	<div style="width: 60%; height: 10px; background-color: #999999; border: 1px solid #ccc;"></div>	<b>\$10 OOP</b>	<b>85%</b>	<div style="width: 85%; height: 10px; background-color: #008080; border: 1px solid #ccc;"></div>
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Source: Triangle Insights Group Qualitative (n=20) Market Research with PSVT Patients, April – June 2017  
MIST = Milestone Pharmaceuticals, Inc.

# PSVT Patient Management and Call Point Targeting



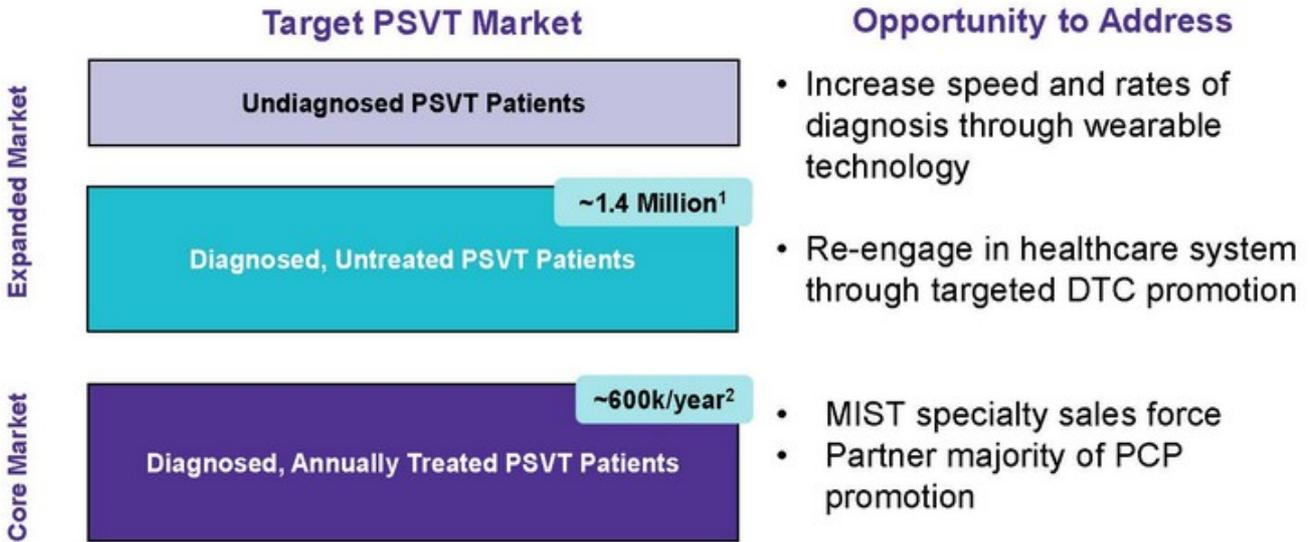
Majority of PSVT patients managed by CV specialists, leading to commercial efficiencies

		Clinical Cardiologists	Primary Care Physicians	Electro-physiologists
<b>% of PSVT patients managed</b>		<b>~60%</b>	<b>~30%</b>	<b>~10%</b>
<b>Long-term Use</b>	<i>Add to or Replace Chronic Medications</i>	<b>Primary Target</b>		
<b>Medium-term Use</b>	<i>Defer Ablation</i>			<b>Secondary Target</b>
<b>Short-term Use</b>	<i>Bridge to Ablation</i>			

- Targeted sales force to reach majority of available opportunity
- Significant overlap with most common CV portfolio call points

Source: Internal market research

# Core Market for PSVT with Potential for Expansion



Source: 1) Calculated as the difference between PSVT prevalence of 2M and annual treatment rate of ~600k from IBM MarketScan® Commercial Research Database (<65y) and the Medicare Limited Dataset (≥65y), 2008-2016 analyzed by Precision Xtract, 2019 2) Estimated number of unique patients with annual claims for PSVT from IBM MarketScan® Commercial Research Database (<65y) and the Medicare Limited Dataset (≥65y), 2008-2016 analyzed by Precision Xtract, 2019.

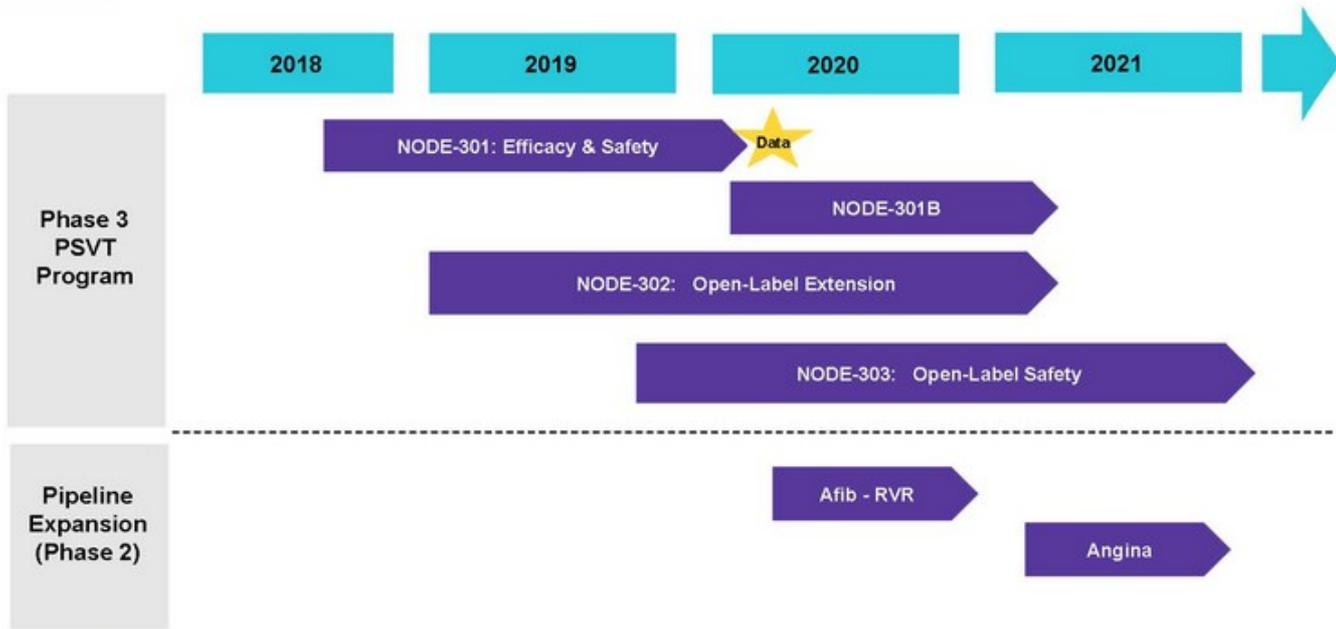
## Finances

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- Cash and equivalents of \$136.5M (as of September 30, 2019)
  - IPO (May 2019) net proceeds of approx. \$86M
- Runway expected into Q3, 2021
  - Phase 3 pivotal efficacy trial (NODE-301) data
  - Significant progression of Phase 3 safety study (NODE-303)
  - Continued PSVT market development via publications, patient education and Medical Affairs initiatives
  - Phase 2 proof of concept endpoint in atrial fibrillation
- 24.5M shares outstanding

# Etripamil Development Plan



PSVT = Paroxysmal Supraventricular Tachycardia  
Afib-RVR = Atrial Fibrillation with Rapid Ventricular Rate

## Milestone (Nasdaq: MIST) - Corporate Highlights



- Phase 3 Cardiovascular Company with data read out anticipated in middle 1H, 2020
- PSVT is a robust market represented by ~2M patients in US
- Paradigm-changing approach enabling patient self-management
- Potentially first new drug therapy in PSVT in > 25 years
- New Chemical Entity with proprietary IP protection until 2036
- Pipeline opportunities beyond the lead indication
- \$95M Initial Public Offering - May 13, 2019
- Cash & equivalents of \$136.5M (Sept. 30, 2019) – expected runway into Q3, 2021

PSVT = Paroxysmal Supraventricular Tachycardia



**Milestone**  
PHARMACEUTICALS

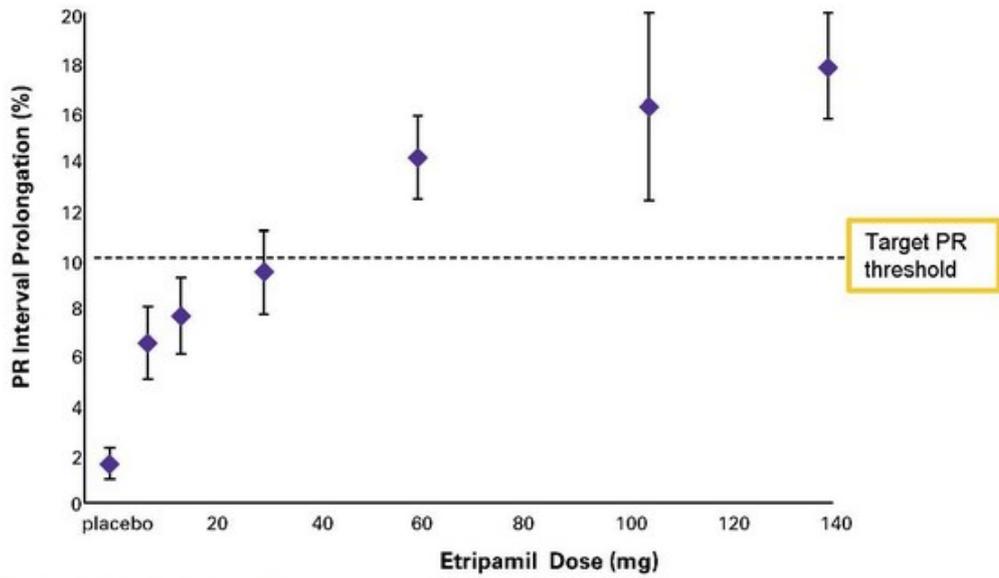
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**Thank you**

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# Etripamil Phase 1 Pharmacology

## PR Prolongation Used to Select Doses for Phase 2

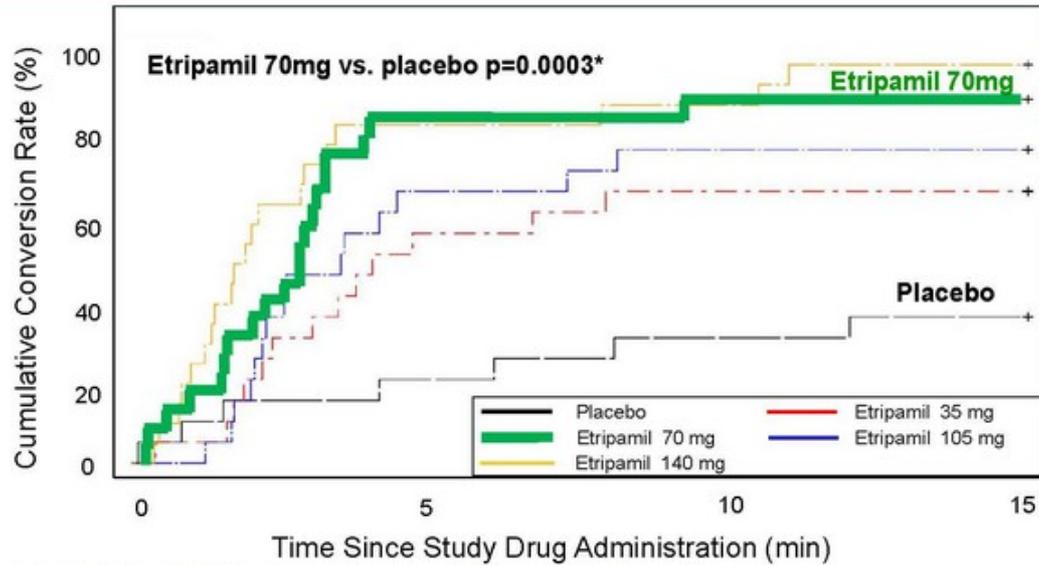


Error bars indicate standard error of the mean

## Phase 2 Time to Conversion



70mg etripamil dose showed rapid time to conversion (median < 3 min)



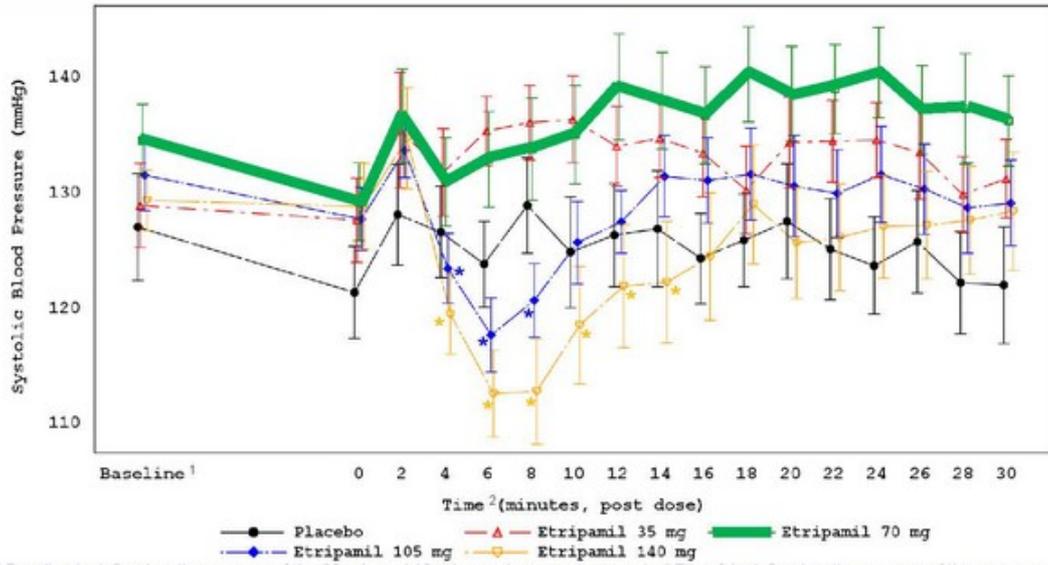
\*Hazard Ratio and 95% Confidence Intervals etripamil 70mg vs. placebo; 4.99 (2.09, 11.93)

Source: Stambler, B.S. et al.; Etripamil Nasal Spray for Rapid Conversion of Supraventricular Tachycardia to Sinus Rhythm; J Am Coll Cardiol. 2018;72(5):489-97

# Phase 2 Mean Systolic Blood Pressure Effects



Etripamil 70 mg showed no drop in blood pressure



<sup>1</sup> Baseline is defined as the average of the 20-min and 10-min pre-dose measurements. <sup>2</sup> Time 0 is defined as the average of the measurements during supraventricular tachycardia between 5 and 0 min before study drug administration. \*p < 0.05 versus baseline.

Source: Stambler, B.S. et al., Etripamil Nasal Spray for Rapid Conversion of Supraventricular Tachycardia to Sinus Rhythm; J Am Coll Cardiol. 2018;72(5):489-97

# Potential Commercial Opportunity for Etripamil in PSVT



<b>Number of annual PSVT ablations</b>	<b>80,000</b>
Ratio of etripamil-treated patients : ablation	x <u>3.5</u>
Total expected etripamil patients/year	280,000
Etripamil expected doses/patient/year	x <u>3</u>
Etripamil doses/year	840,000

TAM – Target Addressable Market

Sources: Internal estimates based on market research and longitudinal analysis of Truven/Marketscan and Medicare claims data