



Steve Davis, CEO

**38th Annual
J.P. Morgan
Healthcare
Conference**

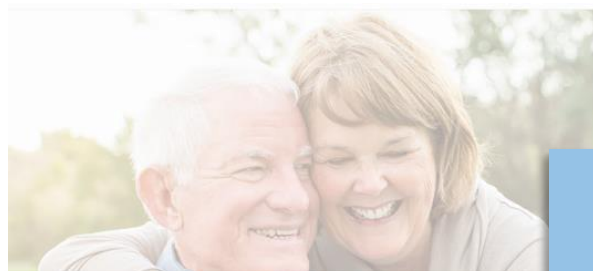
JANUARY 14, 2020

Forward-Looking Statement

This presentation contains forward-looking statements. These statements relate to future events and involve known and unknown risks, uncertainties and other factors which may cause our actual results, performance or achievements to be materially different from any future results, performances or achievements expressed in or implied by such forward-looking statements. Each of these statements is based only on current information, assumptions and expectations that are inherently subject to change and involve a number of risks and uncertainties. Forward-looking statements include, but are not limited to, statements about (i) plans for, including timing and progress of commercialization of, NUPLAZID® or for the clinical development of our product candidates, including pimavanserin and trofinetide; (ii) benefits to be derived from and efficacy of our product candidates, including the use of pimavanserin in dementia-related psychosis, schizophrenia, depression or other neurological or psychiatric indications, potential advantages of NUPLAZID versus existing antipsychotics or antidepressants, and expansion opportunities for NUPLAZID; (iii) estimates regarding the prevalence of PD, PD Psychosis, dementia-related psychosis, schizophrenia or depression and the potential use of trofinetide in Rett syndrome; (iv) potential markets for any of our products, including NUPLAZID and trofinetide; and (v) our estimates regarding our future financial performance, cash position or capital requirements.

In some cases, you can identify forward-looking statements by terms such as “may,” “will,” “should,” “could,” “would,” “expects,” “plans,” “anticipates,” “believes,” “estimates,” “projects,” “predicts,” “potential” and similar expressions (including the negative thereof) intended to identify forward-looking statements. Given the risks and uncertainties, you should not place undue reliance on these forward-looking statements. For a discussion of the risks and other factors that may cause our actual results, performance or achievements to differ, please refer to our annual report on Form 10-K for the year ended December 31, 2018 as well as our subsequent filings with the SEC. The forward-looking statements contained herein are made as of the date hereof, and we undertake no obligation to update them for future events.

ACADIA in 2020 – Building a Leading CNS Platform



TRANSFORMING STANDARD
OF CARE FOR PDP PATIENTS

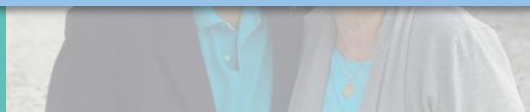
50% NUPLAZID®
NET SALES
GROWTH YOY¹



INNOVATIVE PIPELINE

4 LATE-STAGE
PIPELINE PROGRAMS

3 POSITIVE PIVOTAL
STUDIES



PIMAVANSERIN FOCUSED ON
SIGNIFICANT PATIENT NEED

35X POTENTIAL INCREASE
IN ADDRESSABLE
MARKET BEYOND PDP²



Dedicated to Improving Lives of Patients, Families, and Caregivers

¹2019 net sales guidance of \$330-340M, represents a 50% increase in revenue and 32% volume growth year-over-year at the mid-point of the range.

²ACADIA Market Research based on estimated U.S. treated populations for patients with dementia-related psychosis (DRP), adjunctive treatment for major depressive disorder (MDD), and the negative symptoms of schizophrenia (NSS).

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2020 Strategic Pillars

Drive
NUPLAZID® Growth
in PDP



Deliver
DRP Opportunity
to the Market



Develop
Innovative Treatments
For Unmet Needs



Successful Execution and Strong Balance Sheet
Position ACADIA for Significant Long-term Growth

The Potential of Pimavanserin

A Novel Selective Serotonin Inverse Agonist

Current – NUPLAZID®

- First and only FDA-approved treatment for PDP
- FDA Breakthrough Therapy
- Patent protection into 2030¹



Future – Pimavanserin

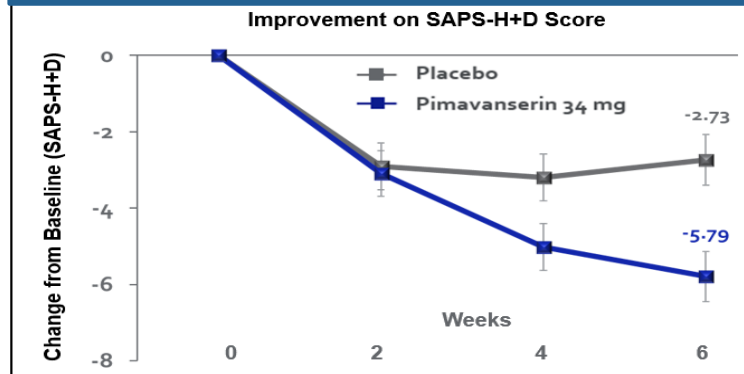
- Robust efficacy in pivotal studies across 3 additional CNS indications:
 - DRP (FDA Breakthrough Therapy)
 - MDD (adjunctive treatment)
 - Negative symptoms of schizophrenia

Safety in Late-Stage Clinical Trials

- DRP - No negative impact on cognition or impairment of motor function
- MDD - Improved symptoms of sexual dysfunction with no increased sedation or weight gain
- Schizophrenia - No effect on vital signs, weight, and metabolic syndrome

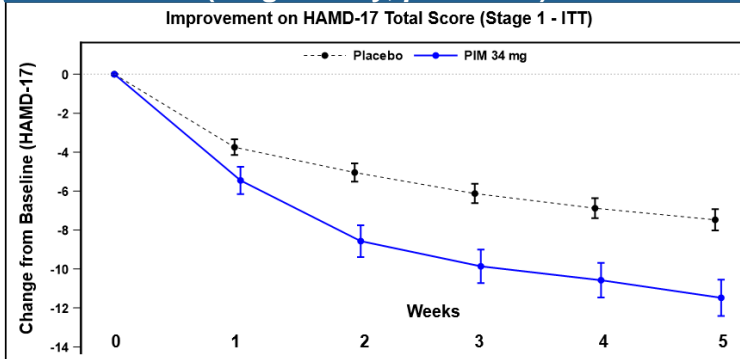
Pimavanserin – Robust and Consistent Efficacy Across Four Disease Areas

PDP: Pivotal -020 Study, $p=0.0014$

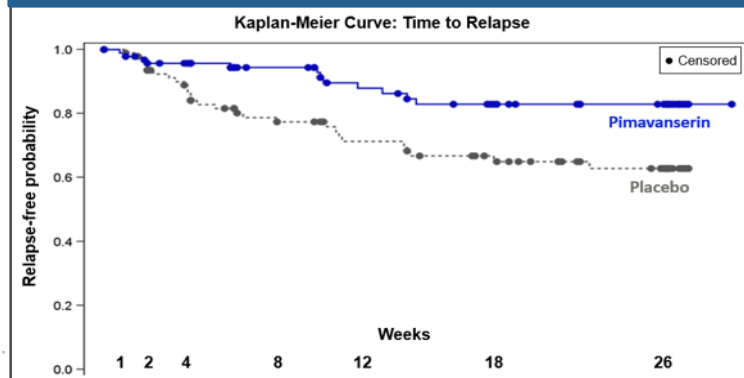


MDD: Pivotal CLARITY Study

(Stage 1 only, $p=0.0003^{1}$)

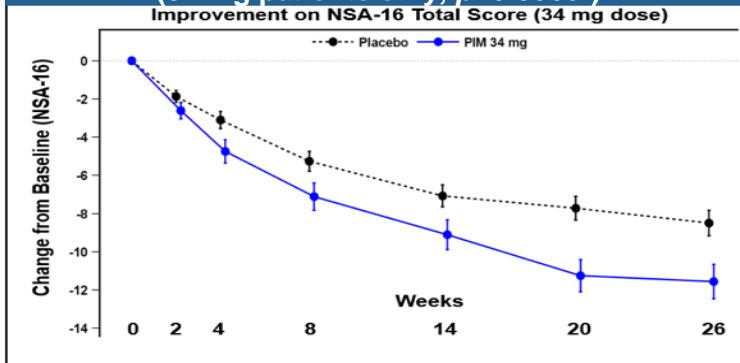


DRP: Pivotal HARMONY Study, $p=0.0023$



NSS: Pivotal ADVANCE Study

(34 mg patients only, $p=0.0065^{2}$)

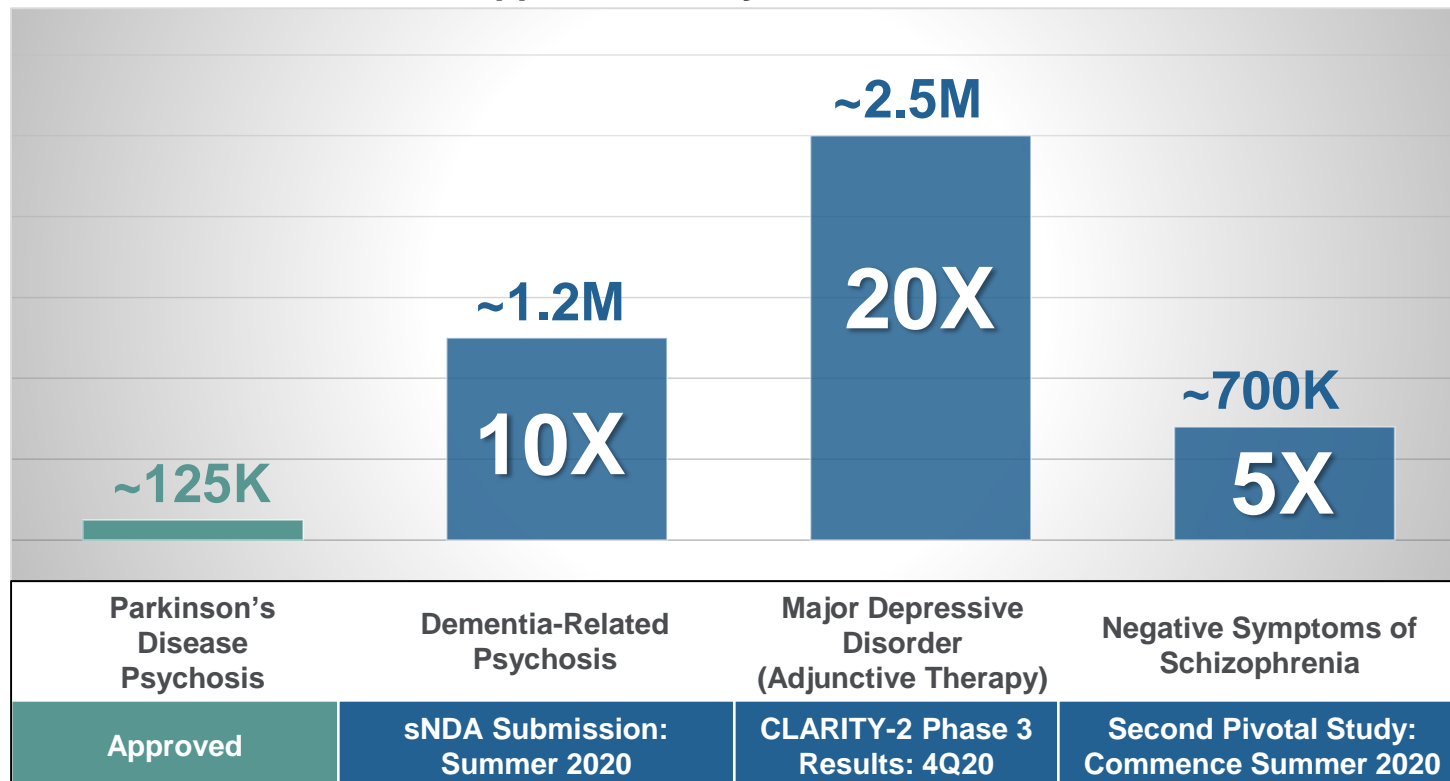


¹Primary endpoint was HAMD-17 total score compared to placebo (wgt. avg. of Stage 1 + 2) in a sequential parallel comparison design ($p=0.039$). Graph shows results from Stage 1.

²Primary endpoint was NSA-16 total score at 26-weeks in patients that received either 20 mg or 34 mg pimavanserin + background antipsychotic vs placebo + background antipsychotic ($p=0.043$). Graph shows results for patients that received the higher 34 mg dose. Provided January 14, 2020 as part of an oral presentation and is qualified by such; contains forward-looking statements; actual results may vary materially; ACADIA disclaims any duty to update.

Pimavanserin – Potential to Provide Meaningful Advances for Patients

U.S. Addressable Market Opportunities by Indication¹



2020 Strategic Pillars

Drive **NUPLAZID® Growth** **in PDP**



Deliver **DRP Opportunity** **to the Market**

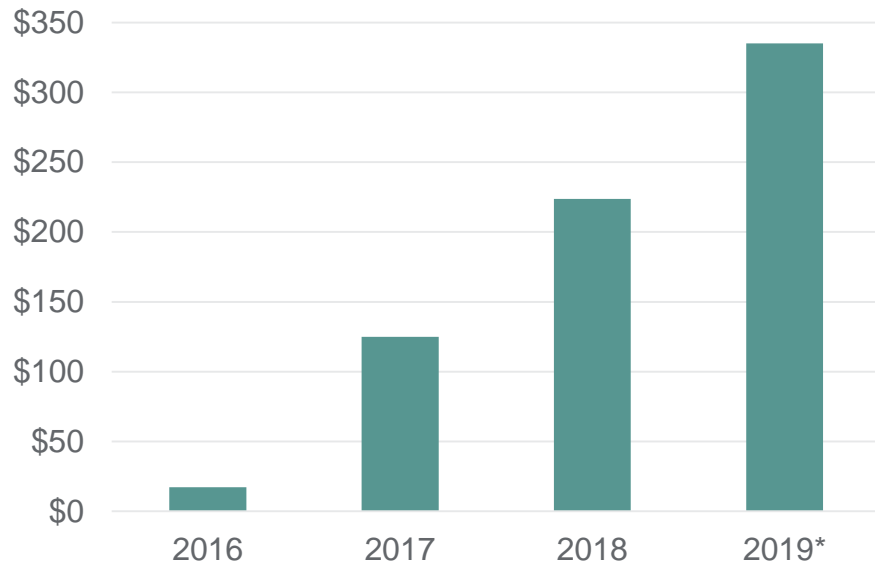


Develop **Innovative Treatments** **For Unmet Needs**



Drive NUPLAZID® Growth in Parkinson's Disease Psychosis

Net Sales (in millions)



- **2019 net sales guidance: \$330-340M¹**
+50% revenue / +32% volume YoY
- **High teens market penetration exiting 2019**
- **Continued growth leveraging:**
 - MDS Evidence based guidelines²
 - *NUPLAZID only therapy recognized as clinically useful and acceptable level of safety risk without specialized monitoring*
 - New caregiver burden and long-term clinical safety data
 - Digital and patient/caregiver campaigns

Significant Future Growth Opportunity in PDP

2020 Strategic Pillars

Drive
NUPLAZID® Growth
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Deliver
DRP Opportunity
to the Market



Develop
Innovative Treatments
For Unmet Needs



Deliver New Opportunity for Dementia-Related Psychosis

No FDA-approved treatments for DRP

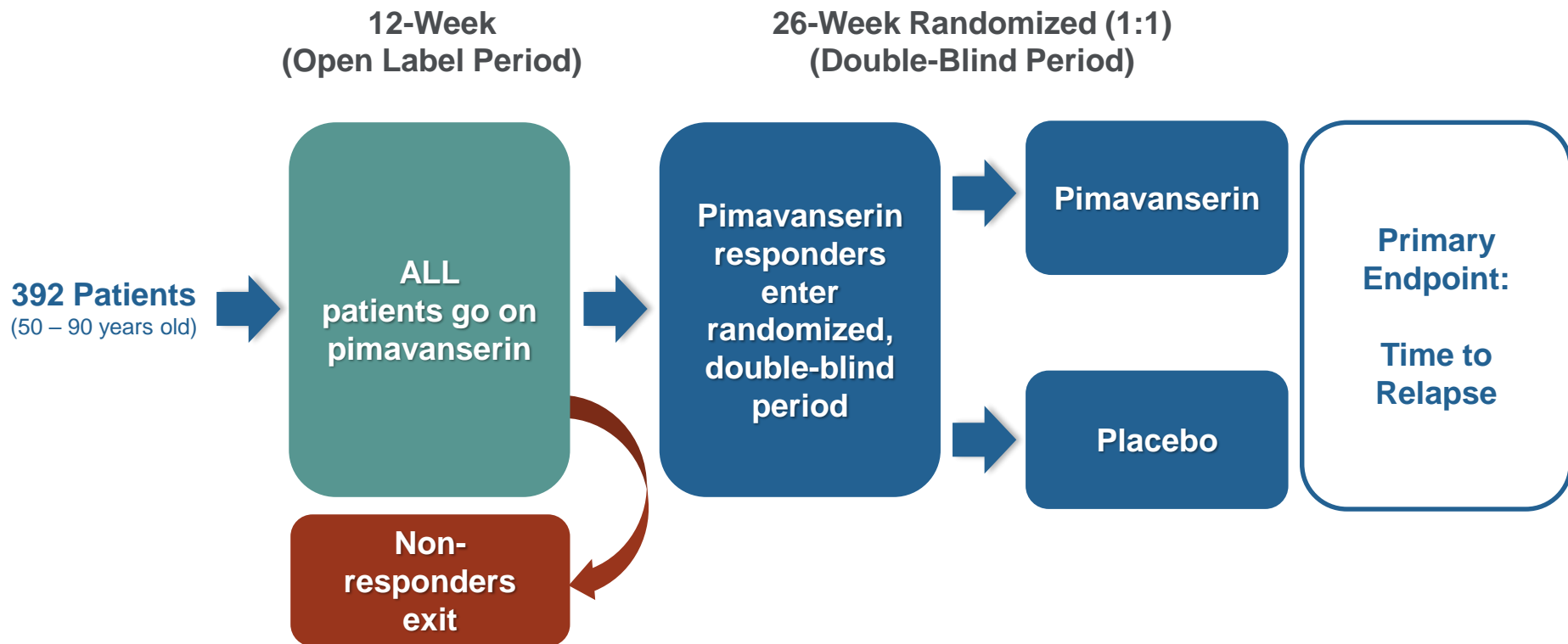


Pimavanserin has Breakthrough Therapy Designation for the treatment of DRP

Today, antipsychotics used off-label¹:

- Accelerate cognitive decline
- Impair motor function
- Cause extrapyramidal symptoms
- Increase sedation
- Cause orthostatic hypotension

Phase 3 HARMONY Relapse Prevention Study in DRP



Robust Positive Phase 3 HARMONY Results

1

Achieved Meaningful Primary Endpoint

- *Pimavanserin significantly reduced the risk of relapse of psychosis by **2.8 fold***
- ***Hazard Ratio = 0.353***
- ***One-sided p-value = 0.0023***

2

Strong Open-Label Efficacy Results

- ***61.8%** of eligible patients met the pre-specified response criteria at weeks 8 and 12*
- ***75.2%** improvement from baseline on SAPS-H+D¹ score at week 12*

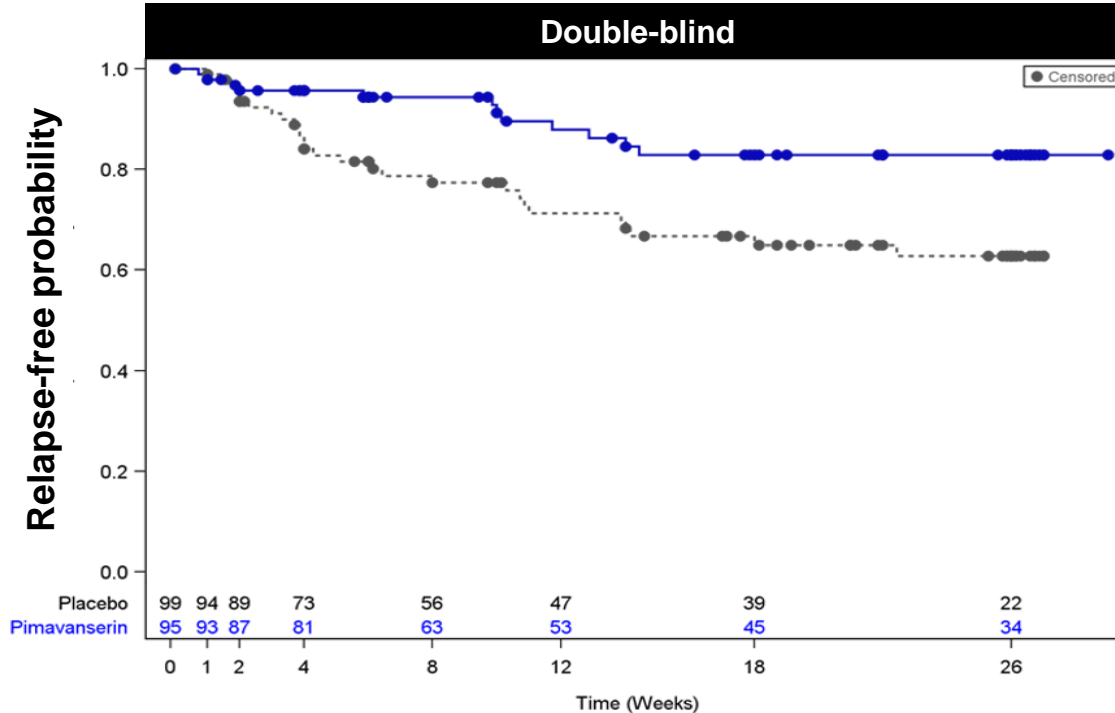
3

9-Month Safety and Tolerability Results

- *Well-tolerated in chronic treatment of frail and elderly patients with significant comorbidities*
- *No worsening of cognition²*
- *No worsening of motor function³*

2.8 Fold Reduction in Risk of Relapse of Psychosis

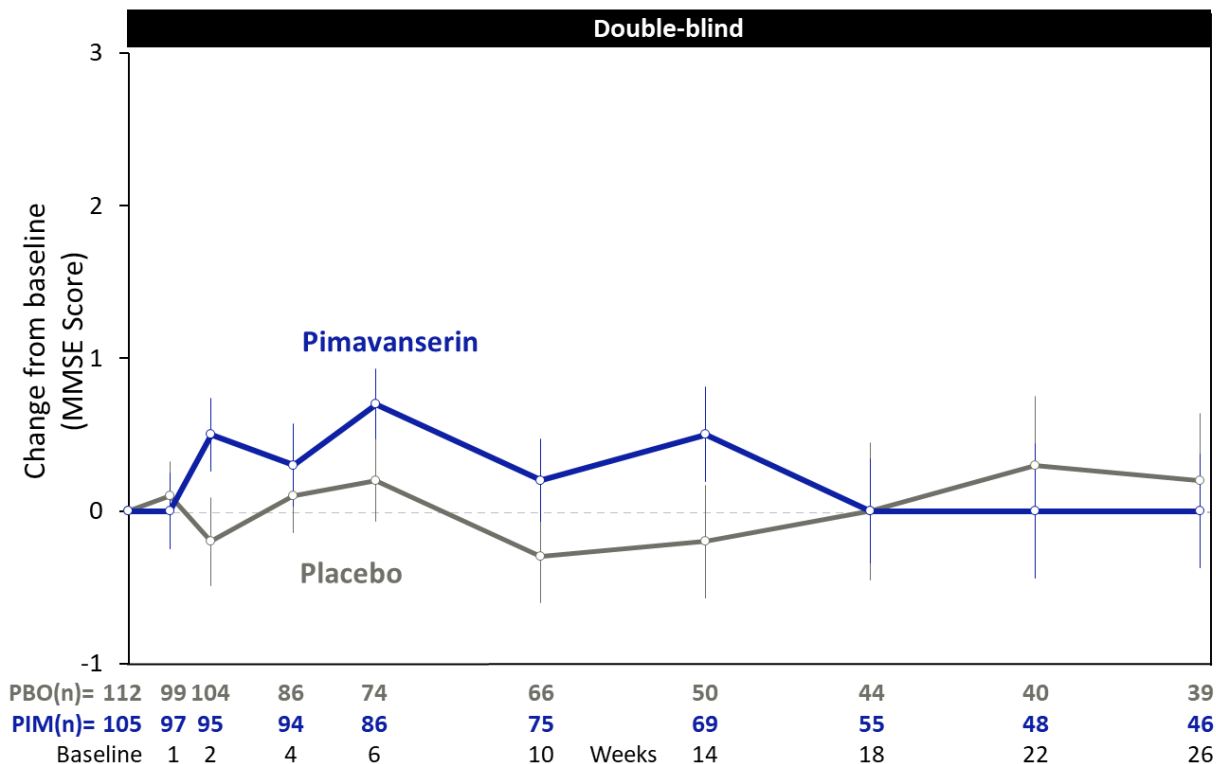
Primary Endpoint: Time to Relapse



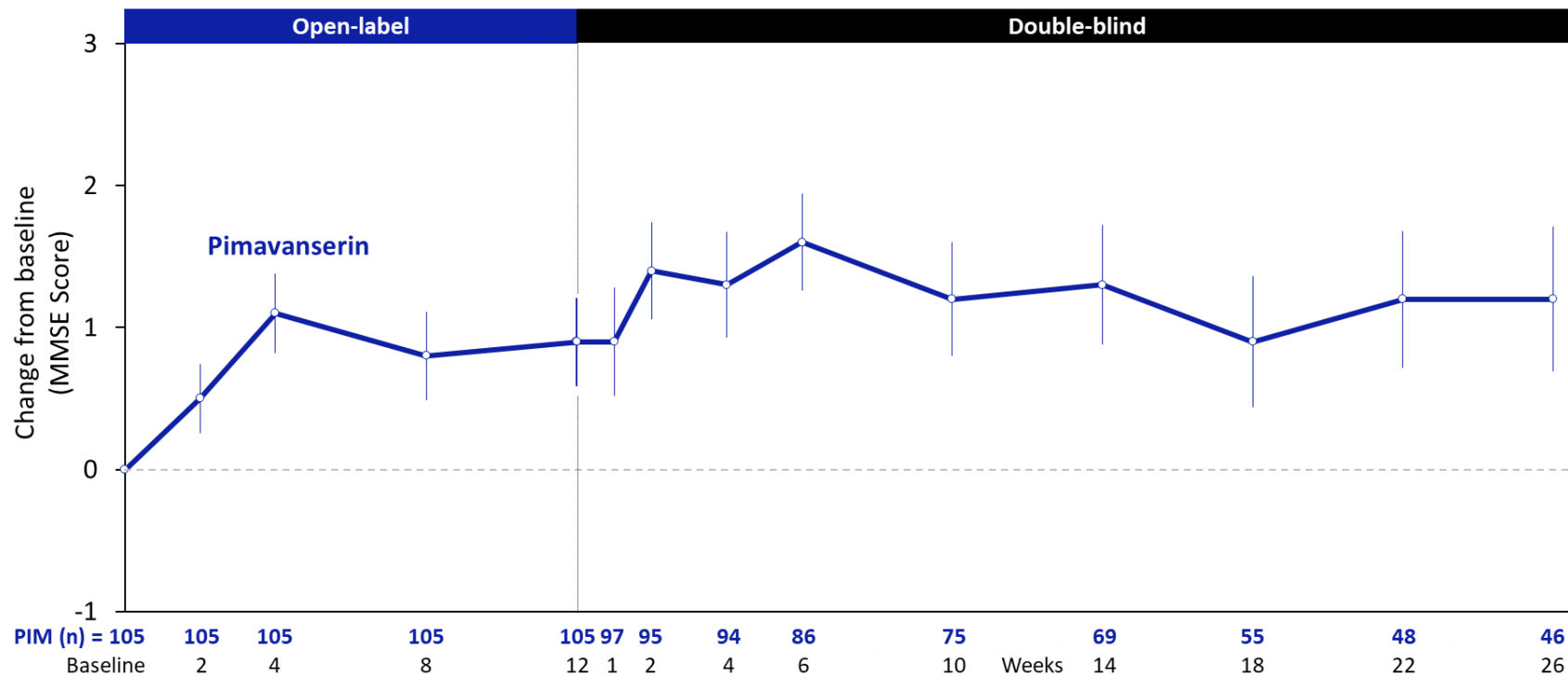
Hazard Ratio = 0.353

One-sided p-value = 0.0023

No Negative Impact on Cognition (MMSE) Over 6 Months Compared to Placebo¹



No Negative Impact on Cognition (MMSE) Over 9 Months of Treatment¹



DRP Next Steps

1. Pre-sNDA meeting request submitted ✓
2. Plan to submit sNDA in summer 2020

sNDA will include the following:

Pivotal Efficacy

Positive Phase 3
HARMONY Study

Supportive Efficacy

Positive Phase 2 (019)
Alzheimer's Disease
Psychosis Study¹
&
Positive data in PDP (020)
patients with dementia²

Large Safety Database

Safety and
Tolerability Data
from Completed
& Ongoing Studies

Pimavanserin has Breakthrough Therapy Designation for the Treatment of DRP

¹Ballard C, et al. Lancet. 2018;17:213-222.

²NUPLAZID Prescribing Information; Cummings J, et al. Lancet. 2014;383:533-540.

NUPLAZID (pimavanserin) is only approved in the U.S by the FDA for the treatment of hallucinations and delusions associated with Parkinson's disease psychosis.

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2020 Strategic Pillars

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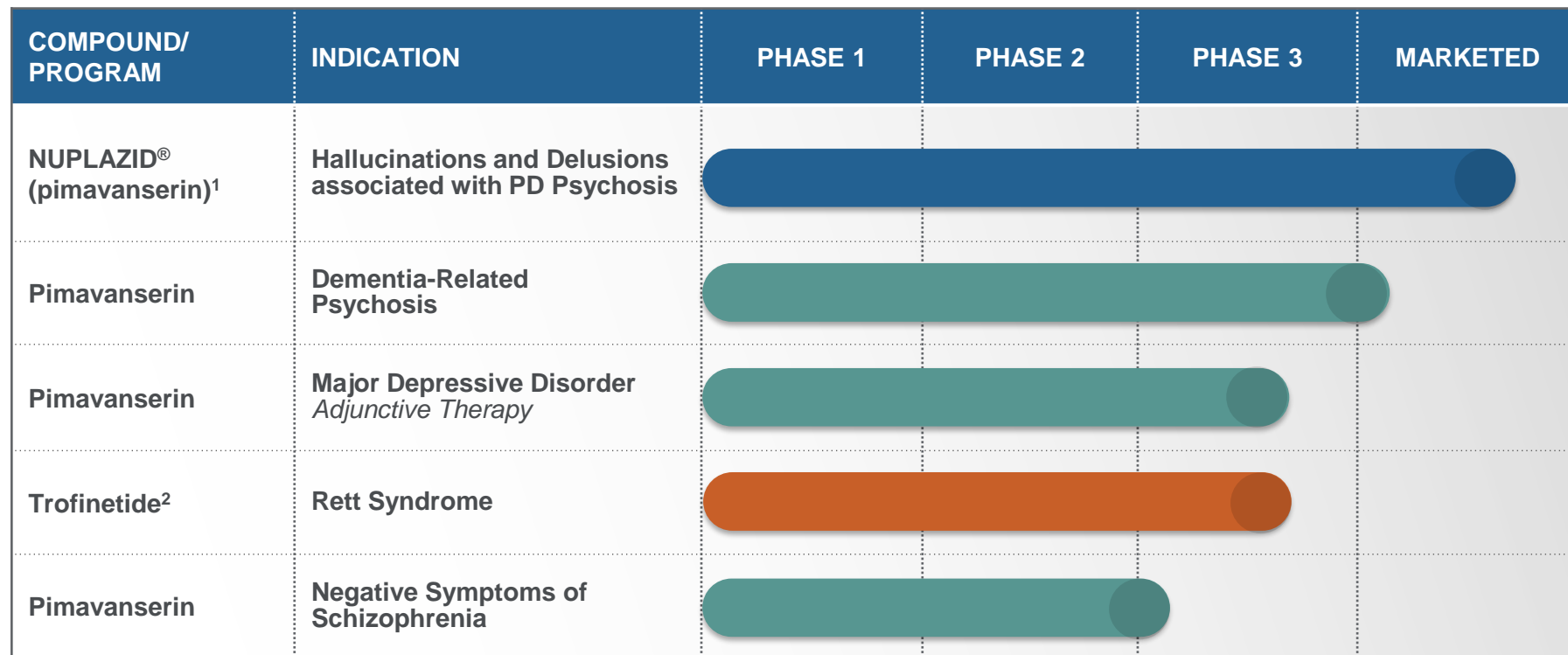
Deliver
DRP Opportunity
to the Market



Develop
Innovative Treatments
For Unmet Needs



Innovative Late-Stage Pipeline



Major Depressive Disorder – Adjunctive Therapy

High unmet need for differentiated adjunctive therapy



- **~17M patients in the U.S. have MDD¹**
 - Majority of patients with MDD do not respond to initial antidepressant therapy
- **~2.5M treated with adjunctive therapy²**
- **Adjunctive use of existing antipsychotics can lead to significant side effects:**
 - Sexual dysfunction
 - Sedation
 - Weight gain
 - Cognitive impairment
 - Extrapyrimalidal symptoms
 - Rare but serious tardive dyskinesia

¹National Institute of Mental Health. (2017). Major Depression. Retrieved from <http://www.nimh.nih.gov/health/statistics/major-depression.shtml>.

²IMS NSP, NPA, NDTI MAT-24 month data through Aug-2017; PLOS One, Characterization of Treatment Resistant Depression Episodes in a Cohort of Patients from a US Commercial Claims Database, Oct 2013, Vol 8, Issue 10.

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Advancing Adjunctive Treatment for MDD

CLARITY Results

Meaningful Efficacy:

Primary endpoint achieved – Depression¹

- **HAMD-17 (*p-value*=0.039)**

Robust effect in the parallel design Stage 1

- **HAMD-17 (*p-value* = 0.0003;
Effect size = 0.63)**

Key secondary endpoint achieved - Disability¹

- **SDS (*p-value*=0.004)**

Secondary Outcome Findings:

- Early and sustained antidepressant treatment effect²
- Improvement in sexual dysfunction symptoms
- Improvement in daytime sleepiness
- No meaningful weight gain
- No cognitive side effects observed
- No extrapyramidal symptoms observed
- No tardive dyskinesia observed

CLARITY-2 Phase 3 Study Results Expected 4Q 2020

Two ongoing Phase 3 studies with only one additional positive study necessary for sNDA

¹HAMD-17: 17-item Hamilton Depression Rating Scale; SDS = Sheehan Disability Scale.

²Week 1 separation from placebo observed in Stage 1 (n=207) and Week 10 separation from placebo observed in Stage 1 patients who were not re-randomized in Stage 2 (n=174).

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Negative Symptoms of Schizophrenia

No FDA-approved treatment for the negative symptoms of schizophrenia



- ~40 - 50% of schizophrenia patients experience predominant negative symptoms¹
- **Negative symptoms include** apathy, lack of emotion, social withdrawal, restricted speech, and blunted affect and can lead to:
 - Low social functioning
 - Long-term disability
 - Significant caregiver burden

Summary of Top-line ADVANCE Results

- 1 The study achieved statistical significance on the primary endpoint
 - ▶ Improvement in NSA-16¹ total score compared to placebo at 26 weeks
p-value = 0.043
 - ▶ Greater improvement on NSA-16 was observed in patients on the higher 34 mg dose (n=107) vs. placebo ***unadjusted p-value = 0.0065***
 - ▶ Second pivotal study will evaluate 34 mg vs. placebo
 - ***Study to commence in summer 2020***
- 2 Pimavanserin was well-tolerated when added to background antipsychotic therapy with low rates of AEs, SAEs, and discontinuations

Trofinetide for the Treatment of Rett Syndrome

No FDA-approved treatment for Rett syndrome



- Debilitating neurologic rare disease
- 6,000 to 9,000 patients in the U.S.¹
- Symptoms manifest primarily in young females:
 - Cognitive, sensory, emotional, and motor impairment
 - Loss of independence
 - Loss of purposeful hand use
 - Loss of spoken communication

Trofinetide Clinical Program

Phase 2 Study

Phase 2 study:

- Statistically significant improvements in **RSBQ** and **CGI-I**
- Positive Phase 2 study results published in *Neurology*^{®1}

Clinical Program

LAVENDER Phase 3 study ongoing:

- ~180 females (ages 5 – 20) with Rett syndrome
- Double-blind, placebo-controlled
- Co-primary endpoints: RSBQ and CGI-I
- 12-week study duration

LILAC 9-month extension study:

- To evaluate LT tolerability and safety of trofinetide

LAVENDER Results Expected in 2021

Upcoming Clinical and Regulatory Milestones

COMPOUND	INDICATION	MILESTONE	EXPECTED TIMING
Pimavanserin	Dementia-Related Psychosis	Pre-sNDA Meeting Request Submitted	1Q20 ✓
		sNDA Submission	Summer 2020
Pimavanserin	Major Depressive Disorder <i>Adjunctive Therapy</i>	CLARITY-2 Results Expected	4Q20
		CLARITY-3 Results Expected	1H21
Pimavanserin	Negative Symptoms of Schizophrenia	Initiate ADVANCE-2	Summer 2020
Trofinetide	Rett Syndrome	LAVENDER Results Expected	2021



Improving lives
for patients, caregivers
and families

