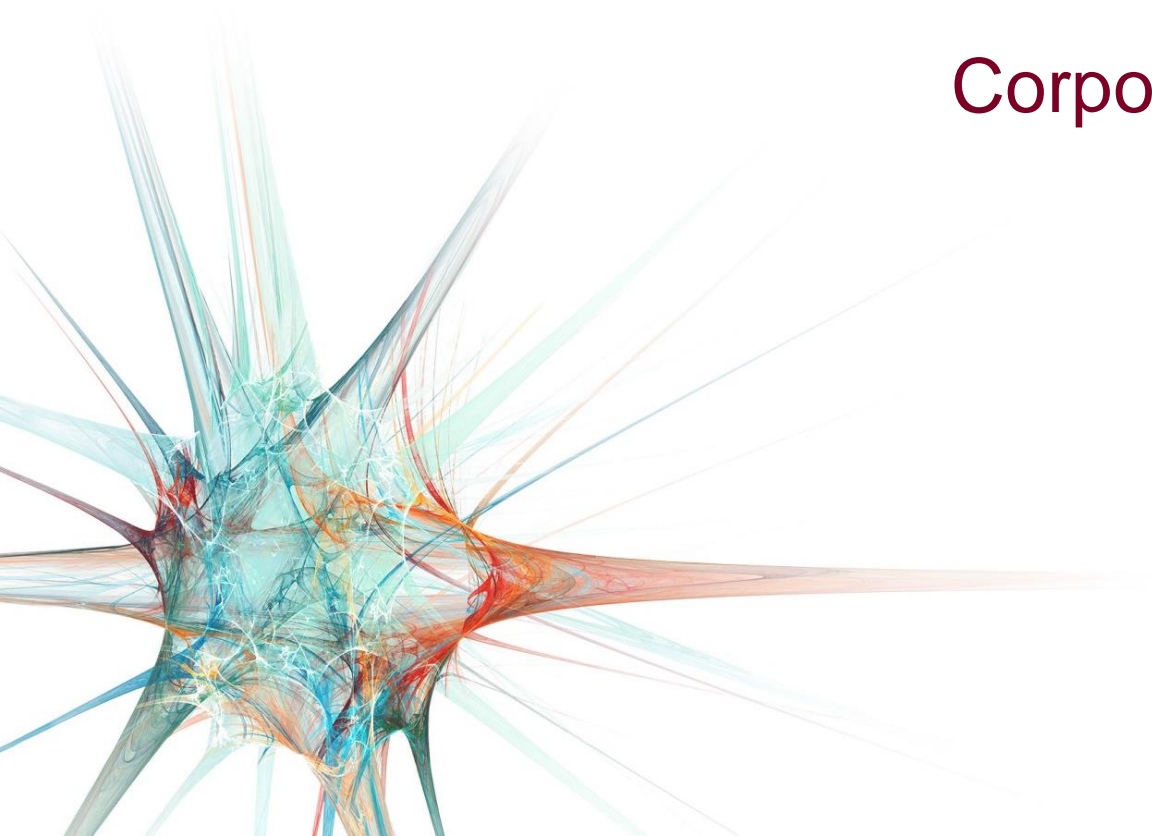




Corporate Presentation

January 2019



Safe Harbor Statement

This presentation contains “forward-looking statements” within the meaning of the Private Securities Litigation Reform Act of 1995. These forward-looking statements concern our product candidates, our development efforts, our collaborations, our technology, our intellectual property, our financial condition, our plans and our development programs. These statements involve risks, uncertainties and assumptions, and are based on the current estimates and assumptions of the management of Intra-Cellular Therapies, Inc. (the “Company” or “ITCI”) as of the date of this presentation and are subject to uncertainty and changes. Given these uncertainties, you should not place undue reliance on these forward-looking statements. Important factors that could cause actual results to differ materially from those indicated by such forward-looking statements include, among others, those set forth in our Annual Report on Form 10-K for the year ended December 31, 2017 filed with the Securities and Exchange Commission, as well as any updates to those risk factors filed from time to time in our periodic and current reports. All statements contained in this presentation are made only as of the date of this presentation, and the Company undertakes no duty to update this information unless required by law.

Intra-Cellular Therapies, Inc. (ITCI)

- Founded in 2002, leveraging technology to study intracellular signaling from the lab of Nobel Laureate Dr. Paul Greengard
- Focus on advancements in the treatment of CNS disorders
 - Lead program: Lumateperone (ITI-007)
 - For the treatment of schizophrenia
 - NDA under FDA review, PDUFA target action date Sept. 27, 2019
 - For the treatment of bipolar depression
 - Leader in the field of PDE1 inhibitors
 - Parkinson's' disease
 - Heart failure
- Well-capitalized
 - \$376 million in cash, cash equivalents and investment securities at 09/30/2018



ITCI Therapeutic Pipeline



Schizophrenia Is A Common Disabling Neuropsychiatric Disease



Schizophrenia affects ~1% of the global population¹

2.4 million adults in the United States have schizophrenia²

Economic Burden of schizophrenia exceeds **\$154 Billion** in the U.S.³

- Onset in early adulthood leads to life-long disability
- It is estimated that only about one-third of patients with schizophrenia can work regularly.⁴



1. National Alliance on Mental Illness. Available at <https://www.nami.org/Learn-More/Mental-Health-Conditions/Schizophrenia>. Accessed Dec 28, 2018.

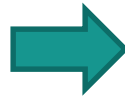
2. Schizophrenia. National Institutes of Health. Available at: <https://report.nih.gov/nihfactsheets/ViewFactSheet.aspx?csid=67> Accessed: Dec 28, 2018.

3. Cloutier M, et al. *J Clin Psychiatry*. 2016;77(6):764-771. 4. Bellack AS, et al. *Schizophr Bull*. 2007;33(3):805-822.

Lumateperone is Designed to Address Unmet Needs in Schizophrenia



Existing therapies are associated with significant adverse events including extrapyramidal symptoms, akathisia, metabolic disturbances, weight gain and cardiovascular dysfunction

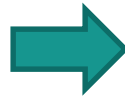


- Impact quality of life
- Lead to poor adherence with 74% of US patients discontinuing medication within 18 months¹

Weight loss and improvement in key cardiometabolic parameters were observed with long term treatment with lumateperone vs Standard of Care (SOC)

Existing drugs are:

- Only effective for positive symptoms; negative symptoms and depression are not effectively treated
- Social function is not improved



The unique pharmacology of lumateperone may translate into an advancement in the treatment of schizophrenia as a single, stand-alone drug therapy

Lumateperone: Novel, First-in-Class Molecule With an MOA That Predicts Clinical Benefits Across CNS Disorders



	<u>nM Ki</u>
5HT-2A	0.5
D2	32
D1	52
SERT	62

5-HT_{2A} Receptor Antagonist

- Improves sleep quality
- Reduces anxiety and hostility
- Enhances antipsychotic and antidepressant activity

Glutamatergic Phosphoprotein Modulator

NMDA & AMPA Enhancement via D1

- Efficacy for negative and positive symptoms
- Improved cognition
- Rapid-acting antidepressant activity

Dopamine Phosphoprotein D₂ Modulator (DPPM)

D₂ Pre-synaptic partial agonist and post-synaptic antagonist

- Antipsychotic efficacy for positive symptoms
- Reduced agitation

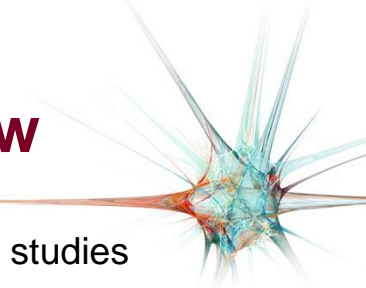
PHARMACOLOGY PREDICTS ROBUST EFFICACY ACROSS A BROAD RANGE OF SYMPTOM DOMAINS

AND PREDICTS HIGHLY FAVORABLE SAFETY/TOLERABILITY PROFILE

Serotonin Reuptake Inhibitor

- Antidepressant activity

Lumateperone Schizophrenia Program Overview



- Lumateperone (ITI-007 60 mg) met primary efficacy endpoint in 2 entirely U.S.-based studies
- Well-tolerated with a safety profile similar to placebo in all studies
- Once daily oral dosing with no need for titration to 60 mg
- Two studies with risperidone as active control showed statistically significant safety & tolerability benefits for lumateperone over risperidone
- NDA under FDA review, PDUFA target action date Sept. 27, 2019

3 Large Randomized, Double-Blind Trials

Primary outcome measure: change from baseline on Positive and Negative Syndrome Scale (PANSS) Total Score

ITI-007-005¹

*4-week treatment period
N=335*

- ITI-007 (60 mg or 120 mg)
- Risperidone 4 mg or
- Placebo

ITI-007-301

*4-week treatment period
N=450*

- ITI-007 (60 mg or 40 mg) or
- Placebo

ITI-007-302

*6-week treatment period
N=696*

- ITI-007 (60 mg or 20 mg)
- Risperidone 4 mg or
- Placebo

Open-Label Safety Switching Study - ITI-007 60 mg

First Part

*6-week treatment duration
N=302*

Second Part

Long-term treatment

ICH guidelines met
300 pts for 6 mos., 100 for 1 yr

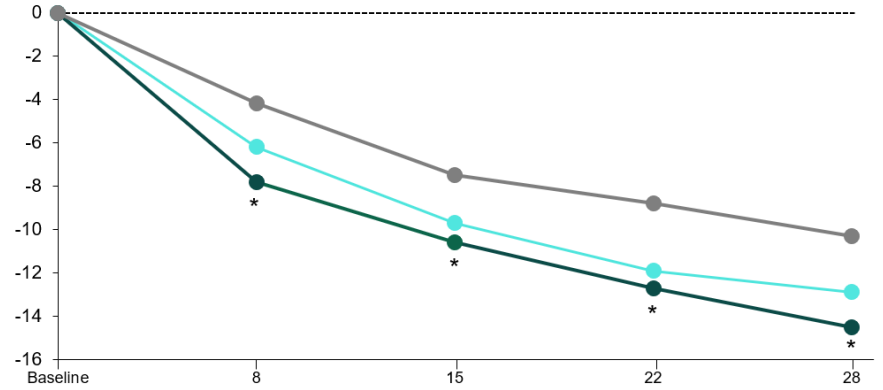
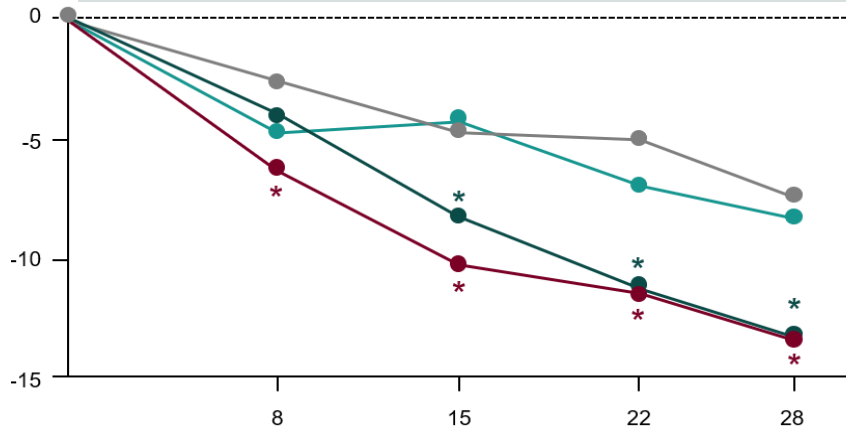
ITI-007 60 mg Met Primary Endpoint in 2 Positive Large Schizophrenia Studies



ITI-007-005

ITI-007-301

Mean Change From Baseline in PANSS Total Score



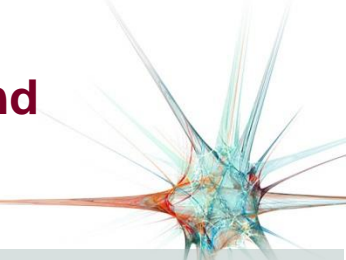
- Lumateperone 60 mg (n=76)
- Lumateperone 120 mg (n=80)
- Risperidone 4 mg (n=75)
- Placebo (n=80)

- Lumateperone 40 mg (n=123)
- Lumateperone 60 mg (n=130)
- Placebo (n=109)

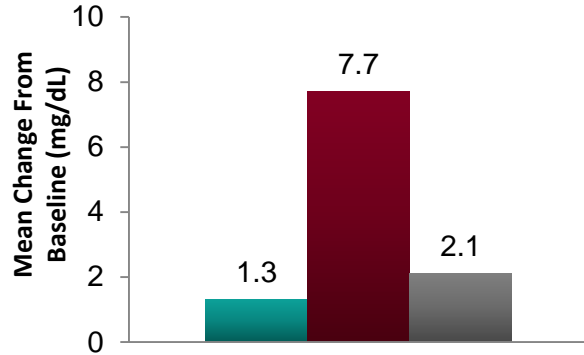
*P<.05 vs placebo.

PANSS=Positive and Negative Syndrome Scale.

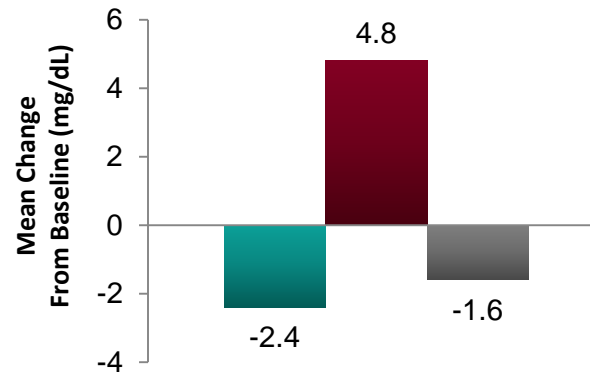
Pooled Data from 3 Acute Controlled Studies Demonstrated the Favorable Safety Profile of Lumateperone: Similar to Placebo And Superior to Risperidone on Key Safety Parameters



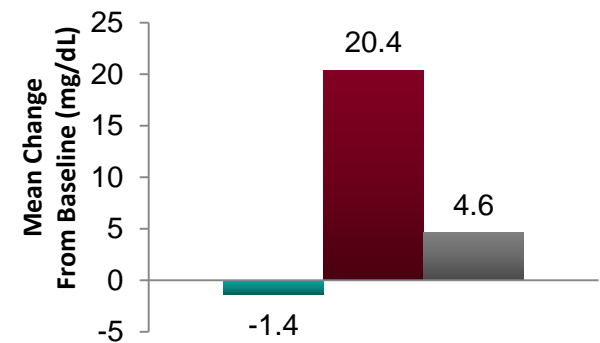
Fasting Glucose



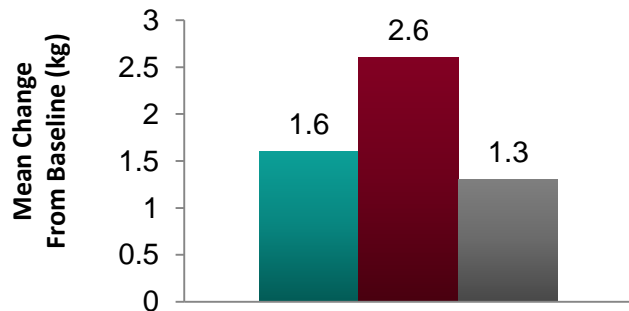
Total Cholesterol



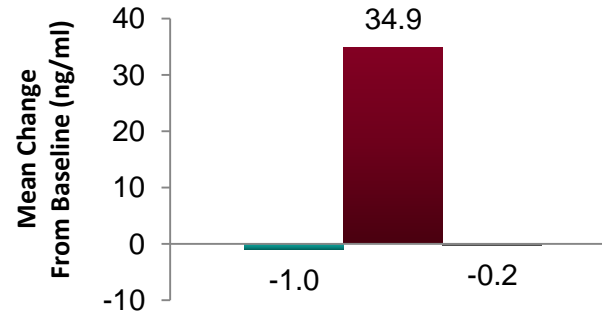
Triglycerides



Body Weight

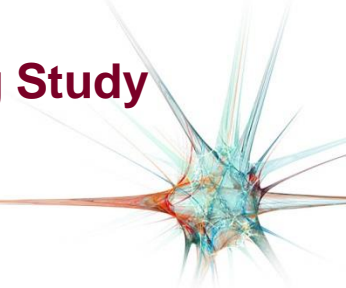


Prolactin



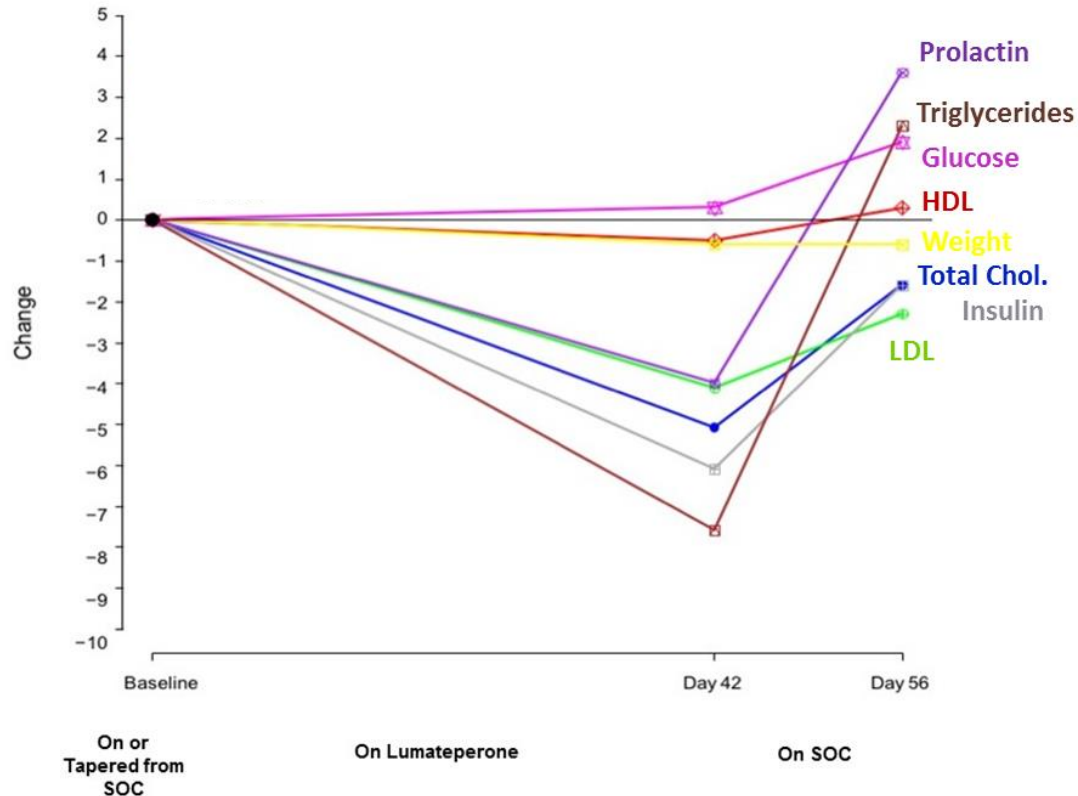
■ Lumateperone 20-120 mg (N=811) ■ Risperidone 4 mg (N=255) ■ Placebo (N=412)

Lumateperone Profile Confirmed in Open-label Safety Switching Study (Study 303) in Patients with Stable Schizophrenia Symptoms



- The study evaluated the safety of lumateperone 60mg in stable patients with schizophrenia in an outpatient setting similar to common clinical practice
- Part 1: 6-week treatment duration assessed:
 - the impact of switching to lumateperone from SOC antipsychotics (with no dose titration necessary for lumateperone)
 - the impact of switching back to SOC from lumateperone
- Part 2: up to 1-year treatment duration assessed:
 - the long-term impact of switching to lumateperone from SOC antipsychotics (with no dose titration necessary for lumateperone)

PART 1: 6-week Open Label Safety Switching Study, Key Cardiometabolic Parameters Improved on Lumateperone and Worsened Again After Switch Back to SOC (N=302)



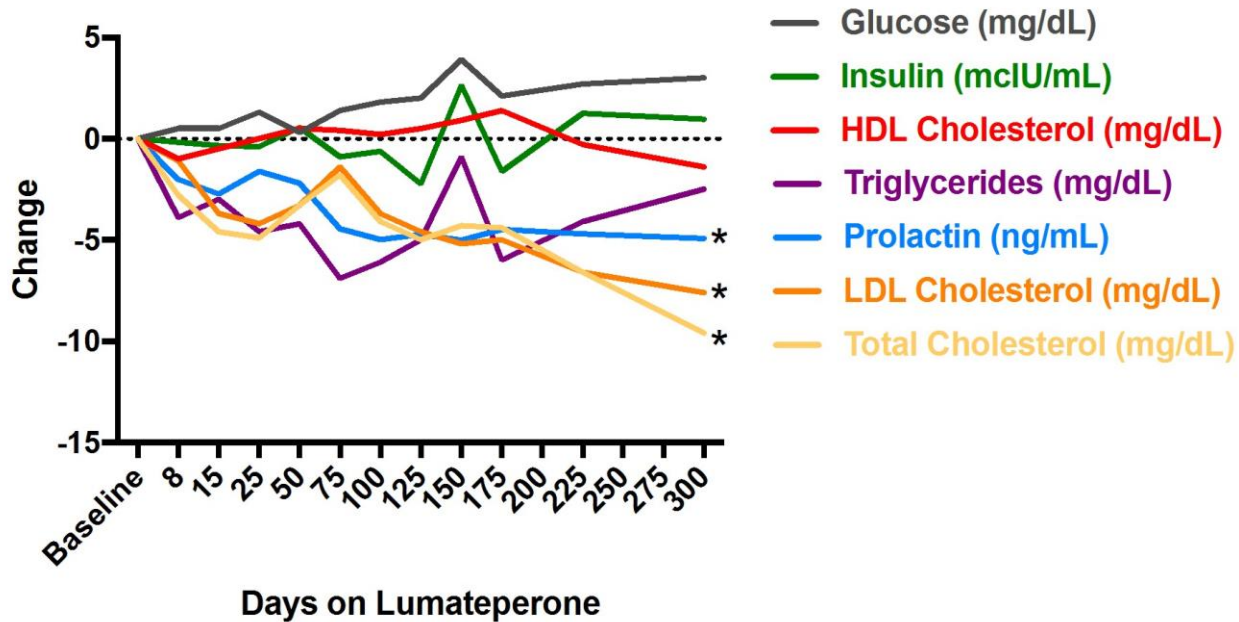
Lumateperone was generally well tolerated with a favorable safety profile

Statistically significant improvements from SOC baseline were observed in body weight, cardiometabolic and endocrine parameters over 6 weeks of treatment with lumateperone and worsened again when switched back to SOC medication

PART 2: Open Label Long-Term Study, Key Cardiometabolic Parameters Improved on Lumateperone



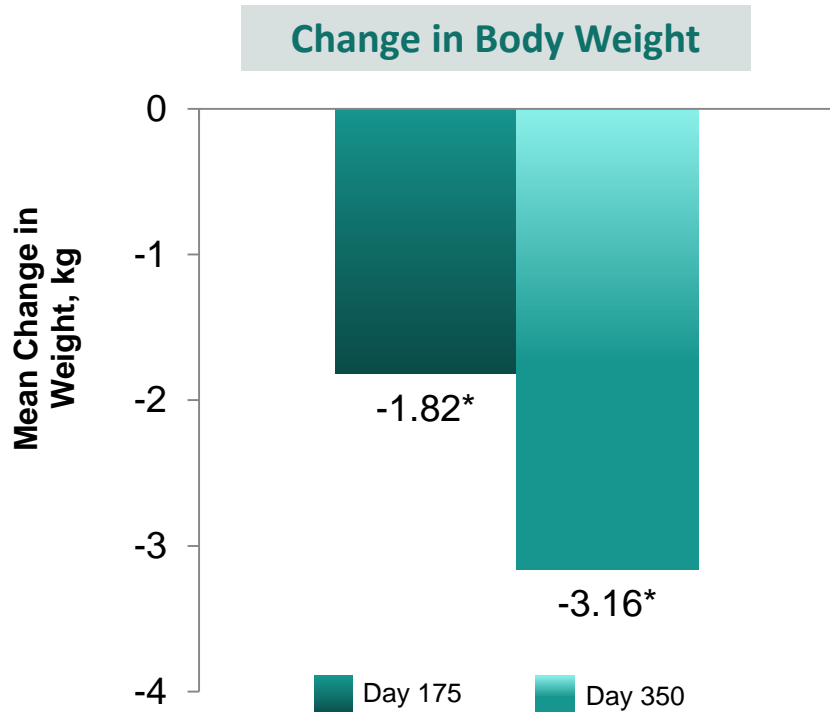
Lumateperone Exhibited a Favorable Cardiometabolic Profile with Long-Term Administration¹



1. Results meeting ICH guidelines for long-term safety presented at ACNP 2018 on 329 patients treated for at least 6 months and 108 patients treated for one year

* p<0.001 vs baseline

Open-Label Long-Term Study, Weight Decreased on Lumateperone¹



Patients With Significant Change in Weight ($\geq 7\%$) At Any Time During the Study

Weight decrease	24%
Weight increase	8%

* $P < 0.001$ compared to SOC baseline.

1. Results meeting ICH guidelines for long-term safety presented at ACNP 2018 on 329 patients treated for at least 6 months and 108 patients treated for one year. At baseline, the mean body weight was 92.5 kg,

Summary: Lumateperone Schizophrenia Program



- Program includes two large, well-controlled positive studies and supportive data from a third study
- In all three studies, lumateperone was well-tolerated with a safety profile similar to placebo
- In the 2 studies with risperidone as the active control, lumateperone was statistically significantly better than risperidone on key safety and tolerability parameters
- Long-term safety study confirmed the favorable cardiometabolic safety profile demonstrated in short-term studies. Weight loss seen in short-term studies continued over long-term treatment

Bipolar Depression is a Common Psychiatric Condition



- **Bipolar disorder has a 2.8% 12-month prevalence in US adults¹**
 - Affects ~6 million Americans²
 - Bipolar depression is the predominant presentation
 - Depressive episodes are longer and recur more often than manic/hypomanic episodes
- **Unmet Need**
 - Few Approved Treatments available
 - Safety and tolerability tradeoff limit use of existing agents

16 1. Bipolar Disorder. National Institute of Mental Health. Available at: https://www.nimh.nih.gov/health/statistics/bipolar-disorder.shtml#part_155458 Dec 28, 2018.
2. Depression and Bipolar Support Alliance Available at: https://secure2.convio.net/dabsa/site/SPageServer/?pagename=education_bipolar: Jan 3, 2019.

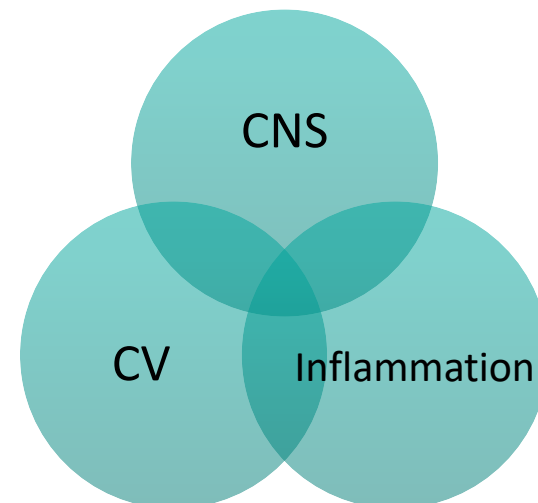
Lumateperone for the Treatment of Bipolar Depression and Other Depressive Disorders



- Potent 5-HT_{2A} receptor antagonism and D₂ receptor modulation coupled with inhibition of serotonin transporters and enhancement of glutamatergic neurotransmission suggest antidepressant activity
 - In previous studies depressed schizophrenia patients experienced robust improvements in depressive symptoms
 - Recent preclinical data support potential for rapid-acting antidepressant effects
- Bipolar depression Phase 3 clinical program includes
- Two monotherapy studies
 - One adjunctive study (lithium or valproate)

PDE1 Inhibition: Novel Approach to Modulation of Intracellular Signaling with Broad Therapeutic Potential

- PDE1 enzymes are highly active across a variety of neurological and cardiovascular diseases
- Our PDE1 inhibitors are designed to reestablish normal function in these disease states
 - Inhibitors of PDE1 block the breakdown of second messenger cyclic nucleotides (cAMP, cGMP) potentiating downstream intracellular signaling
- ITI-214, our lead PDE1 molecule, is being developed for the treatment of Parkinson's Disease and Heart Failure



ITI-214 for the Treatment of Parkinson's Disease (PD)



- **Over 2.2 million patients in the US and Europe^{1,2}**
- **Unmet Need**
 - Progressive neurodegenerative disorder with motor and non-motor symptoms
 - Dopamine replacement therapies (L-DOPA as gold standard) address motor symptoms, but are insufficient and have limiting side effects
 - Effects of dopamine replacement therapies wear off over time as disease progresses
- **ITI-214 is a Multi-Pronged Approach to the treatment of PD**
 - PDE-1 inhibition/ITI-214 enhances intracellular dopamine signaling pathways in the brain
 - ITI-214 may be neuroprotective and has potential disease modifying effects

1. Parkinson's Foundation. Available at <https://www.parkinson.org/Understanding-Parkinsons/Statistics>. Accessed Jan 3, 2019

2. European Parkinson's Disease Association. Available at http://ec.europa.eu/research/horizon2020/pdf/contributions/post/european_organisations/european_parkinson%27s_disease_association_-_epda.pdf. Accessed Jan 3, 2019

ITI-214 Phase 1/2 Results in PD



● Study ITI-214-105

- A Phase 1/2, randomized, double-blind, placebo-controlled, multiple ascending dose cohort study in patients with mild to moderate PD maintained on stable (concomitant) PD medication
- Patients randomly assigned to receive placebo or ITI-214 (1, 3, 10, 30, and 90 mg) administered orally once daily for 7 days

● ITI-214 was shown to have a favorable safety profile and was generally well tolerated

- across a broad range of doses (1 mg to 90 mg)
- without promoting or worsening motor complications

● Clinical signs consistent with reductions in motor symptoms and motor complications

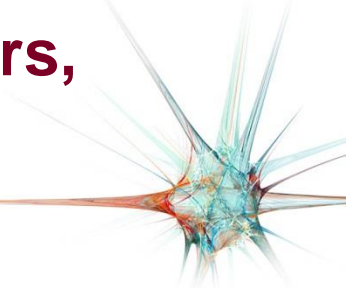
- signals of motor improvement seen in the MDS-UPDRS (and Subscales Part III and Part IV)
- reduced dyskinesia symptoms seen in UDysRS and increase in “ON” time without dyskinesia (Hauser Patient Motor Diary) in patients with dyskinesia at baseline
- several patients experienced profound improvements

ITI-214: Represents A Novel Mechanism of Action for The Treatment of Heart Failure



- **Heart failure (HF) affects approximately 5.8 million people and has a mortality rate of ~50% within 5 years¹**
 - Need for approaches that improve heart function acutely and with longer exposure, also counter maladaptive remodeling of the ventricle
- **Preclinical Data Has Shown Beneficial Effects on Cardiac Function**
 - PDE-1 inhibition increases cardiac contractility and cardiac output
 - Potential for attenuation or reversal of cardiac remodeling
 - ITI-214 acts by a novel mechanism of action that involves modulation of adenosine A_{2B} receptor signaling pathways
- **Ongoing translational study:** a randomized, double-blind, placebo-controlled study of escalating single doses of ITI-214 in patients with systolic HF to evaluate contractility and pharmacodynamic parameters

ITI-333 For Treatment of Substance Use Disorders, Pain, and Mood Disorders



- **Opioid crisis is a public health emergency**

- Nearly 12 million people in the United States misused opioids in the previous year¹
- 130+ people a day die from opioid-related drug overdoses²

- **ITI-333 Profile**

Receptor or Transporter	Ki (nM)
Serotonin 5-HT _{2A}	8.3
Dopamine D ₁	50
Mu opioid (MOP)	11

- High affinity at serotonin 5-HT_{2A}, dopamine D₁ and mu opiate receptors (MOP), acting as a partial agonist at MOP receptors
- Exhibits potent morphine-like analgesia in animal models, yet attenuates several morphine mediated behaviors (e.g., does not cause respiratory depression or GI loss of motility)
- No safety concerns have been noted with ITI-333 in animal models

1. Substance Abuse and Mental Health Services Administration. (2017). *Key substance use and mental health indicators in the United States: Results from the 2016 National Survey on Drug Use and Health* (HHS Publication No. SMA 17-5044, NSDUH Series H-52). Available at <https://www.samhsa.gov/data/sites/default/files/NSDUH-FFR1-2016/NSDUH-FFR1-2016.pdf>. Accessed Jan 3, 2019

2. U.S. Department of Health and Human Services. Available at <https://www.hhs.gov/opioids/>. Accessed Jan 3, 2019

Management Team



Sharon Mates, PhD, Founder, Chairman, President & Chief Executive Officer

Robert Davis, PhD, Senior Vice President, Chief Scientific Officer

Suresh Durgam, MD, Senior Vice President, Late Stage Clinical Development and Medical Affairs

Michael I. Halstead, Executive Vice President and General Counsel

Larry Hine, Senior Vice President Finance & Chief Financial Officer

Mark Neumann, Executive Vice President & Chief Commercial Officer

Juan Sanchez, MD, Vice President, Corporate Communications and Investor Relations

Andrew Satlin, MD, Executive Vice President & Chief Medical Officer

Kimberly Vanover, PhD, Senior Vice President, Early Stage Clinical Development and Translational Medicine

Key Financial Information



KEY METRICS

Total Cash, Cash Equivalents, and Investments¹	\$376.0 million
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Total Debt¹	\$0.0 million
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Common Shares Outstanding¹	54,713,831
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Stock Options / Restricted Stock Units Outstanding⁽¹⁾	5,478,117
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Intra-Cellular
T H E R A P I E S

