

# Corporate Presentation

APRIL 15, 2024

# Forward-Looking Statements and Other Legal Notices

This presentation contains forward-looking statements about Longboard Pharmaceuticals, Inc. ("we," "Longboard" or the "Company"), including statements regarding: our vision; commercial opportunities and analogs; anticipated milestones and timing; the prevalence of, unmet need associated with, and market opportunity for, DEEs; the potential of a broad DEE indication and a broad-spectrum ASM; the potential of bexicaserin (LP352) (including to be best-in-class, to satisfy unmet need, to be a safer, efficacious, and less burdensome therapy, to have differentiated selectivity and specificity, to expand the market to a broader population, to limit adverse events, including those associated with activity at certain receptors, to be indicated across a range of DEEs, to avoid drug-drug interactions, including through optimized dosing, to be desired or preferred by physicians, patients and caregivers; to change the DEE landscape, to provide the cornerstone to build a world-class epilepsy franchise, and to expand, broaden or capture market share); plans regarding a global Phase 3 program for bexicaserin (including the approach, characteristics and timing for such a program); the product profile sampled with HCPs and caregivers; the potential of LP659 (including to be best-in-class or a market leader, to address multiple neurological disorders, to have strong scientific rationale, to be commercially attractive, to have greater selectivity and internalization-based signaling, and to limit off-target effects); expectations and objectives regarding the Phase 1 SAD study for LP659 (including regarding the timing of topline data, the number of participants, and key study objectives); LP659 indication assessment; our intellectual property; our ability to obtain regulatory approval and commercialize our drug candidates (in the manner we may propose or at all); and other statements that are not historical facts, including statements that may include words such as "will", "may", "can", "would", "plan", "anticipate", "expect", "beli

For such statements, we claim the protection of the Private Securities Litigation Reform Act of 1995. These forward-looking statements are subject to a number of risks, uncertainties and assumptions, including, but not limited to: topline data may not reflect the complete or final results of a particular study or trial, and are subject to change; nonclinical and clinical data are voluminous and detailed, and regulatory agencies may interpret or weigh the importance of data differently and reach different conclusions than us or others, request additional information, have additional recommendations or change their guidance or requirements before or after approval; our limited operating history; our history of incurring net losses and expectation that we will continue to incur net losses for the foreseeable future, and that we may never be profitable; our need for additional funding and related risks for our business, product development programs and future commercialization activities; the timing and success of clinical trials and preclinical studies we conduct; our ability to obtain and maintain regulatory approval to conduct our clinical trials (in the manner we propose or at all) and, ultimately, to market our product candidates; our ability to compete in the marketplace; risks regarding our license and dependencies on others; our ability to obtain and freedom to operate for our product candidates; our ability to manage our growth; and other risks and factors disclosed in our filings with the U.S. Securities and Exchange Commission (the "SEC"). We operate in a very competitive and rapidly changing environment. New risks emerge from time to time. It is not possible for our management to predict all risks, nor can we assess the impact of all factors on our business or the extent to which any factor, or combination of factors, may cause actual results to differ materially from those contained in any forward-looking statements we may make. The forward-looking events and circumstances discussed in this presentatio

Certain information contained in or that may orally accompany this presentation relate to or are based on studies, research, publications, surveys and other data obtained from third-party sources and our own internal estimates and research. While we believe these third-party studies, research, publications, surveys and other data to be reliable as of the date of this presentation, they have not been independently verified, and we make no representations as to the adequacy, fairness, accuracy or completeness of any information obtained from third-party sources.

This presentation discusses product candidates (bexicaserin and LP659) that have not yet been approved for marketing by the U.S. Food and Drug Administration (the "FDA") or any other regulatory authority.



Differentiated & innovative clinical approaches

### 

CNS programs with significant commercial opportunities

#### LONGBOARD THESIS

Our Vision is Backed by 20+ Years of World Class **GPCR** Research

A world where **devastating** neurological conditions are no longer devastating



Well-understood targets

### 

Bold & experienced leadership with expertise in CNS and rare disorders

Relevant M&A analogs

JAZZ - GW \$7.2B

PFE - ARNA \$6.7B

UCB - ZGNX \$1.9B



Pipeline with differentiated PK / PD and target engagement

# Longboard's Potentially Best-in-Class Product Candidates\*

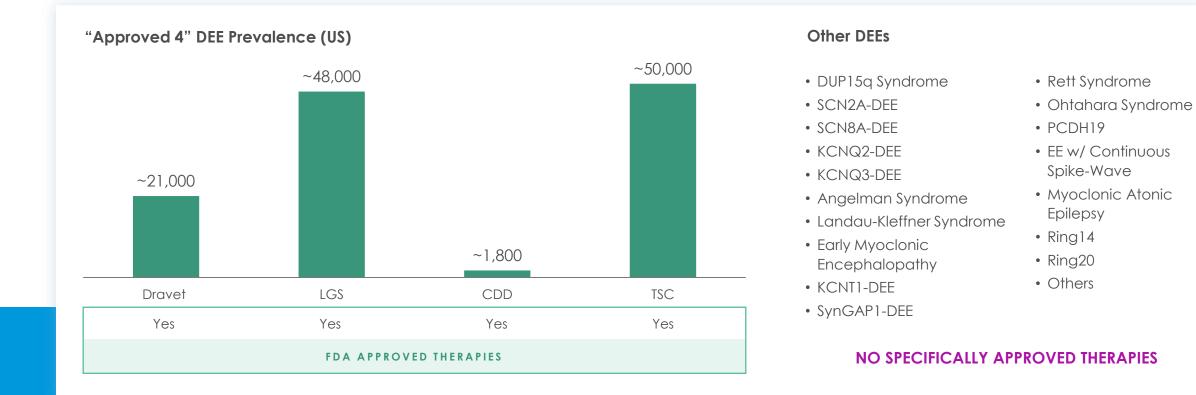
Program	MOA	Therapeutic Area	Preclinical	Ph I	Ph II	Ph III	Anticipated Milestones
							<ul> <li>PACIFIC Study Topline Data – Q1 2024</li> </ul>
Bexicaserin		DEEs and other refractory					<ul> <li>PACIFIC Data at medical meetings – Q2 2024</li> </ul>
(LP352)	(LP352) Superagonist	epilepsies					• PACIFIC OLE Data – H2 2024
							<ul> <li>Global Ph 3 Program Initiation – by YE 2024</li> </ul>
LP659	S1P Receptor Modulator	Multiple neurological diseases					<ul> <li>Ph 1 Initiation – Q4 2023</li> <li>Ph 1 Single-Ascending Dose Data – Q2 2024</li> </ul>

- We hold rights to other product candidates\*
- We are eligible to receive royalties of 9.5% 18.5% on sales of lorcaserin if approved for commercialization\*\*
- \* Through a License Agreement with Arena
- \*\* Through a Royalty Purchase Agreement with Arena

Definitions: DEEs=developmental and epileptic encephalopathies; \$1P=sphingosine 1-phosphate; PK=pharmacokinetics; PD=pharmacodynamics; EEG=electroencephalogram; OLE=Open-Label Extension

Developmental & Epileptic Encephalopathies (DEE) Landscape

# 4 DEE Syndromes Have Approved Therapies; 20+ Have None



#### The prevalence of all "Other DEEs" could exceed the total of the "Approved 4" combined

Sources: Dravet Syndrome Foundation, LGS Foundation

Definitions: LGS = Lennox-Gastaut Syndrome, CDD = CDKL5 Deficiency Disorder, TSC = Tuberous Sclerosis Complex; DEE = Developmental and Epileptic Encephalopathy

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# Surveyed HCPs Report a Need for More Effective and Safer Anti-Seizure Medications for Other DEEs

	Mean # of Seizures Per Week	Mean # of ASMs Per Patient
Dravet	12	3.4
LGS	19	3.5
TSC	6	3.0
CDD	13	2.9
Other DEEs	13	3.2

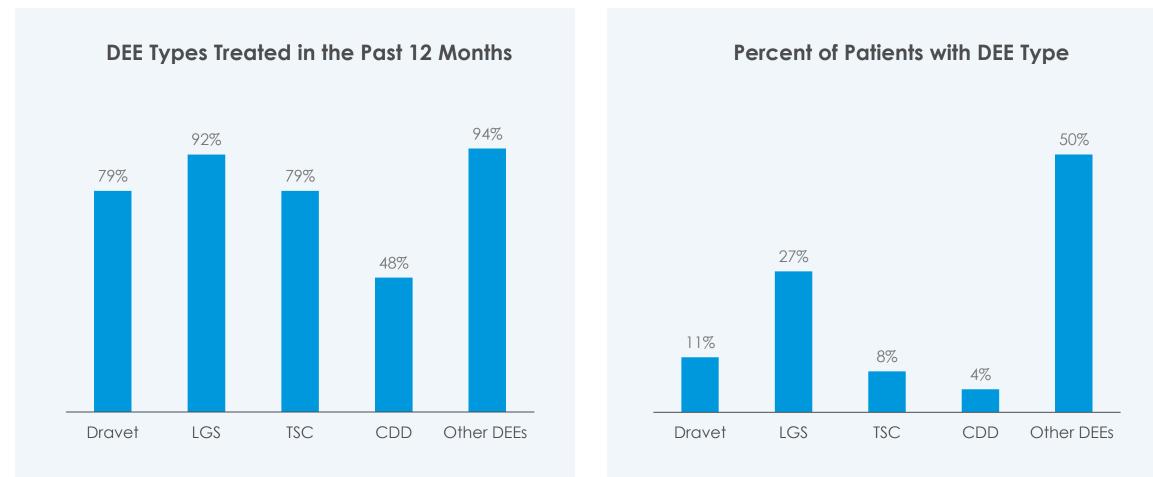
**Unmet Needs for Patients with Other DEEs** Among those who treat patients with other DEEs (not "Approved 4")



LONGBOARD PHARMACEUTICALS

Based on Longboard sponsored third-party qualitative and qualitative market research studies involving physicians specializing in neurology or epileptology. See slide 30 for more information about the studies.

# Nearly All Surveyed HCPs Treat Patients with All DEE Diagnoses; Collectively, the Number of "Other DEEs" is Significant



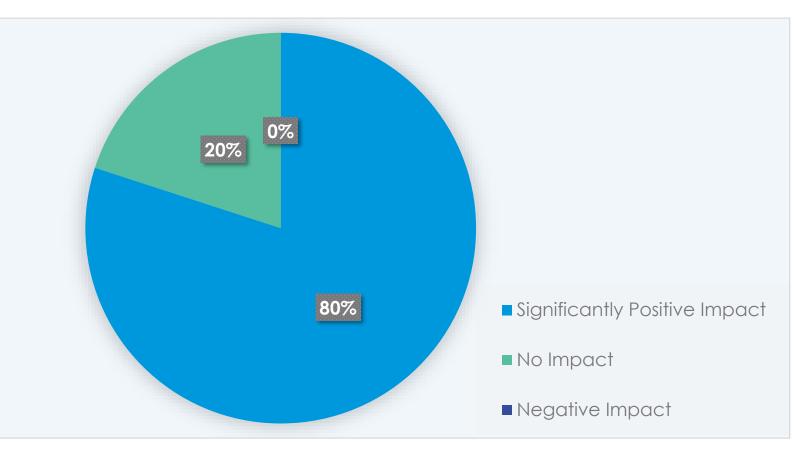
Based on Longboard sponsored third-party qualitative and qualitative market research studies involving physicians specializing in neurology or epileptology. See slide 30 for more information about the studies. Surveyed HCPs Prefer a Highly Selective 5-HT2C Agonist\* with a Broad DEE Indication, and Anticipate This Will Positively Impact DEE Patients' Treatment Options

80%

Anticipate a Broad DEE Indication will have a positive impact on treatment options for patients

**0%** Believe it would have a negative impact

LONGBOARD PHARMACEUTICALS



Based on Longboard sponsored third-party qualitative and qualitative market research studies involving physicians specializing in neurology or epileptology. See slide 30 for more information about the studies.

\*Survey sampled product profile for the 5-HT2C agonist that included an efficacy of 37-44% reduction in countable motor seizure frequency and generally well tolerated with BID dosing

# DEE Indications Represent a \$6B Total US + EU Market Opportunity<sup>1</sup>

A vast majority of the treatment options currently used are generic.

1) Longboard and third-party research 2022 – projections for 2040 for US + EU4/UK, 2) UCB FY 2023 Earnings Release and presentation dated February 28, 2024, 3) Jazz Pharmaceuticals FY 2023 Earnings Release dated February 28, 2024, 4) Jazz Corporate Overview August 2023

**Fintepla Epidiolex** Peak Sales Estimate Sales Estimate<sup>4</sup> (2027)(2025)≥€800M >S1B**Fintepla Epidiolex** 2023 Sales<sup>3</sup> 2023 Sales<sup>2</sup> \$846M **\$226M** 



# Bexicaserin (LP352)

Potential Best-in-Class 5-HT2C Superagonist - Entering a Ph 3 Program with the Goal of Treating a Broad Range of DEEs



# The Potential of Bexicaserin (LP352)

#### Greater Selectivity and Specificity

- 5-HT2 agonist designed to only bind to the 5-HT2C receptor\*
- 5-HT2 agonist no detected activity at receptors associated with significant adverse side effects: 5-HT2B (valvular heart disease and PAH) & 5-HT2A (psychiatric: insomnia, hallucinations, euphoria)

#### Preclinical Validation

- Reduced seizure activity in model of neuronal hyperexcitability in zebrafish
- Reduced epileptiform activity in fish and rodent models of disinhibition
- Reduced duration and number of epileptiform events in zebrafish model of Dravet Syndrome

#### Clinical Validation SAD/MAD in Healthy Volunteers

- In general, favorable safety & tolerability observed. AEs generally consistent with expected effects of serotonergic meds
- No observed food effect
- Potential prolactin biomarker which increased in a dose dependent and transient manner

#### Clinical Validation CSF/EEG \*\* in Healthy Volunteers

- Favorable safety & tolerability results observed, AEs generally consistent with previous studies
- Plasma & CSF PK concentration increased in a dose dependent & consistent manner
- Effects on qEEG activity within first few dose(s)
- Sustained dose-dependent effects on qEEG activity after continuous dosing, thus indicating receptor engagement

# IP protection on Composition of Matter up to 2041\*\*\* provides the opportunity to maximize the full potential of bexicaserin

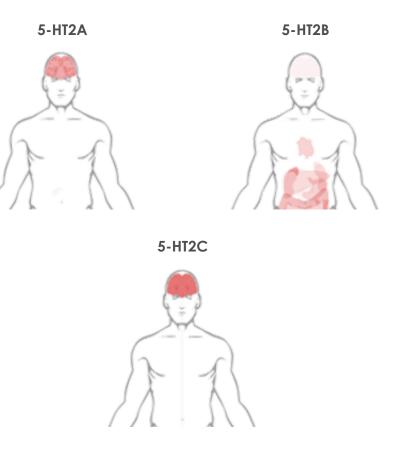
- \*Radioligand binding assays assessing >150 targets showed significant affinity only to 5-HT2C receptors
- \*\* Based on first two cohorts from the 102 study
- LONGBOARD PHARMACEUTICALS

\*\*\*Composition of matter through 2036 with potential for PTE / PTA (2041) Definitions: PAH = pulmonary arterial hypertension; AEs = adverse events; <u>CSF = cerebrospinal fluid</u>



# Bexicaserin (LP352) Designed to be a Next-Generation 5-HT2C with Greater Selectivity and Specificity

Dexicdselin (LP352) 5-HT2C SuperagonistSolution 5-HT2BNot detectableNot detectablen/aSuperagonist5-HT2ANot detectableNot detectablen/aNordexfenfluramine (an active metabolite of fenfluramine)15-HT2C72.410.4CNS, G Pulmond5-HT2A5-HT2B25.79.8Cardiad Pulmond5-HT2A1778120.2Psychiat		Serotonin Receptor Subtype	EC <sub>50,</sub> nM	Ki, nM	Potential Adverse Events Per Receptor Subtype
(LP352) 5-HT2C Superagonist5-HT2BNot detectableNot detectablen/aSuperagonist5-HT2ANot detectableNot detectablen/aNordexfenfluramine (an active metabolite of fenfluramine)15-HT2C72.410.4CNS, G Pulmond Pulmond Dimond T72ASuperagonist5-HT2C72.410.4CNS, G Pulmond Pulmond Pulmond	Revicaserin	5-HT2C	~120	~50	CNS, GI
Superagonist5-HT2ANot detectableNot n/aNordexfenfluramine (an active metabolite of fenfluramine)15-HT2C72.410.4CNS, G5-HT2B25.79.8Cardiaa Pulmond Pulmond5-HT2A1778120.2Psychiat	(LP352)	5-HT2B	-	-	n/a
Nordexfenfluramine (an active metabolite of fenfluramine)15-HT2B25.79.8Cardia Pulmond Dimond Cardia Pulmond5-HT2A1778120.2Psychiat		5-HT2A	_	-	n/a
(an active metabolite of fenfluramine)15-H12B25.79.8Cardia Pulmond5-H12B5-H12B1778120.2Psychiat		5-HT2C	72.4	10.4	CNS, GI
5-HT2A 1778 120.2 Psychiat	(an active metabolite	5-HT2B	25.7	9.8	Cardiac, Pulmonary
5-HT2C 39 13 CNS, G	or remoranine)	5-HT2A	1778	120.2	Psychiatric
		5-HT2C	39	13	CNS, GI
Lorcaserin <sup>2</sup> 5-HT2B 2380 147 n/a	Lorcaserin <sup>2</sup>	5-HT2B	2380	147	n/a
5-HT2A 553 92 Psychiat		5-HT2A	553	92	Psychiatric



1 Third party study previously commissioned by Arena, 2 BELVIQ FDA approved prescribing information 06/2012; Note: The above table is for illustrative purposes only and is not a head-to-head comparison. Differences exist between in vitro study designs and methodologies, and caution should be exercised when comparing data across studies Definitions: CNS= Central nervous system ; GI = Gastrointestinal; ASM = Anti-seizure medication Graphic source: Human Protein Atlas

# PACIFIC Study Topline Data

# Bexicaserin (LP352) Ph 1b/2a PACIFIC Study in Participants with DEEs

Screening Period	Randomization & Up-Titration	Maintenance*	Down- Titration	Follow-up Period	
5 Wks	Days 1-15	Days 16-75	Days 76- 80/90**	30 Days	
		<b>LP352</b> (n=43)			
	$6 \text{ mg} \rightarrow 9 \text{ mg} \rightarrow 12 \text{ mg}$	Participant remains on 6, 9 or 12 mg based on tolerability during up-titration			Open- Label Extension
		Placebo (n=9)			

#### Key Inclusion Criteria:

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- DEEs with average of ≥ 4 motor seizures per 4-week period during the 12 weeks prior to screening and ≥ 4 motor seizures in the 4-week period of screening
- DEEs (multiple syndromes) including LGS, Dravet syndrome, SCN2A-related epilepsies, CDD, among others

#### Key Exclusion Criteria:

Use of fenfluramine & lorcaserin

#### **Basic Information:**

- Sites: 34 sites
- Ages:  $\geq$  12 to  $\leq$  65 yrs old

#### No Echocardiograms Required in PACIFIC



Double-blind, placebocontrolled study to assess the safety, tolerability, pharmacokinetics and efficacy of bexicaserin

#### Study Objectives:

**Evaluate** reduction in countable motor seizures across a broad group of epilepsies

Identify potential indications for pivotal studies

Analyze concentration response to understand dosing in different seizure types and disorders

\* Maintenance Dose of bexicaserin (TID): 6 mg, 9 mg, 12 mg or placebo TID \*\* Up to a 15-day down-titration/taper period (reducing dose every 5 days) depending on

the last maintenance dose

Definitions: LGS = Lennox-Gastaut Syndrome, CDD = CDKL5 Deficiency Disorder

# PACIFIC Study Enrollment Summary





\*The diagnosis was at time of screening for PACIFIC and is subject to further refinement

# Diagnostic Eligibility Criteria: Dravet Syndrome, Lennox-Gastaut Syndrome (LGS) and Other DEEs



All participants: Treatment-resistant countable motor seizures with average of  $\geq$  4 observed/countable motor seizures per 4week period during 12 weeks before screening while on stable ASM treatment

	Dravet Syndrome	LGS	DEE Other
Onset	Between 3–19 months of age	Before 8 years of age	Unprovoked seizures before 5 years of age
Seizure Type	Generalized tonic-clonic, unilateral clonic or bilateral clonic seizures	Tonic or tonic/atonic seizures & more than 1 type of generalized seizure (tonic-clonic, tonic-atonic, atonic, tonic, myoclonic or drop)	Combined focal and generalized seizure types, or multiple generalized seizure types
Developmental History	Initially normal, then delayed	Delayed	Delayed
EEG		Consistent with LGS diagnosis*	Slow or disorganized
Additional Criteria	One of the following: • Emergence of another seizure type after the first • Induced by warm temperatures, fevers, or visual stimuli • Genetic test consistent with Dravet	More than 1 type of generalized seizure for ≥6 months before screening	No history of idiopathic generalized seizures

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# Bexicaserin Has the Potential to Change the DEE Landscape

In the Phase 1b/2a PACIFIC Study

L 59.8%

Median Reduction in Seizures\*

- 74.6% 
   Dravet
- 50.8% 📕 LGS
- 65.5% 🖊 DEE Other

- Bexicaserin provides the cornerstone to potentially build a world-class epilepsy franchise
- Studies to date highlight bexicaserin as potentially best-in-class
- Composition of matter IP protection up to 2041\*\* provides the opportunity to maximize the full potential of LP352
- Moving forward into a global Phase 3 program by YE 2024

Topline Participant Disposition & Safety Results Summary

# Demographics, Baseline Characteristics & Concomitant Medications

Parameter n(%)	Statistics	Bexicaserin (n=43)	Placebo (n=9)	Overall (N=52)
	Mean	23.8	26.7	24.3
Age (Years)	Standard Deviation	9.62	7.73	9.31
	Median	23.0	23.0	23.0
	Min, Max	12, 55	19, 41	12, 55
Sex	Male	21 (48.8)	7 (77.8)	28 (53.8)
JEX	Female	22 (51.2)	2 (22.2)	24 (46.2)
Weight (kg)	Median	55.20	72.76	59.36
weigin (kg)	Min, Max	29.2, 96.0	45.8, 110.7	29.2, 110.7
BMI (kg/m²)	Median	22.4	28.1	23.0
	Min, Max	17, 35	19, 34	17, 35
Baseline Countable Motor Seizures (Median)*	per 28-day period	40.0	24.1	38.2
	Clobazam	21 (48.8)	2 (22.2)	23 (44.2)
Concomitant Medications**	Cannabidiol	14 (32.6)	3 (33.3)	17 (32.7)
	Lamotrigine	13 (30.2)	4 (44.4)	17 (32.7)
	Levetiracetam	16 (37.2)	1 (11.1)	17 (32.7)

## Participant Disposition

n(%)	Ove	rall	Dravet Sy	ndrome	Lennox-Gasta (LG		DEE C	Other
	Bexicaserin	Placebo	Bexicaserin	Placebo	Bexicaserin	Placebo	Bexicaserin	Placebo
Safety Set	43 (100)	9 (100)	4 (9.3)	0	24 (55.8)	5 (55.6)	15 (34.9)	4 (44.4)
Full Analysis Set	35 (81.4)	9 (100)	3 (7.0)	0	17 (39.5)	5 (55.6)	15 (34.9)	4 (44.4)
Participants Completed	32 (74.4)	9 (100)	3 (7.0)	0	15 (34.9)	5 (55.6)	14 (32.6)	4 (44.4)
Participants Discontinued	11 (25.6)	0	1 (2.3)	0	9 (20.9)	0	1 (2.3)	0
Adverse Event	9 (20.9)	0	1 (2.3)	0	7 (16.3)	0	1 (2.3)	0
Consent withdrawn	1 (2.3)	0	N/A	N/A	1 (2.3)	0	N/A	N/A
Lost to follow-up	1 (2.3)	0	N/A	N/A	1 (2.3)	0	N/A	N/A

Note: Percentages are based on the number of participants in the Enrolled (Safety) Set

Safety Set includes all participants who signed informed consent or those who had their legally authorized representative sign for them

Full Analysis Set includes all participants in the Safety Set who complete Part 1 (titration) and have at least 1 post-baseline seizure measurement during Part 2 (maintenance)

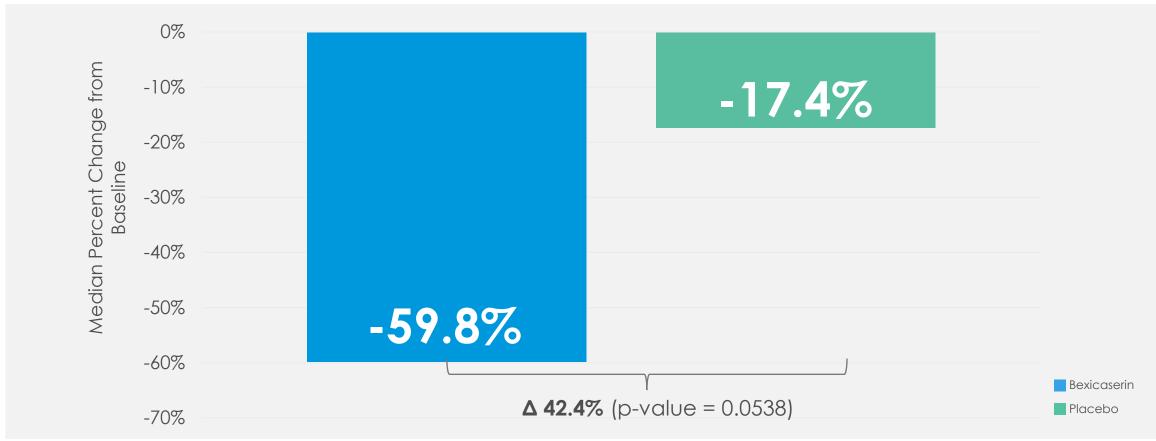
Efficacy analysis was on the Full Analysis Set includes all participants in the Safety Set who complete Part 1 (titration) and have at least 1 post-baseline seizure measurement during Part 2 (maintenance)

# Safety Results Summary

n(%)	Bexicaserin (LP352) (n=43)	Placebo (n=9)	Overall (N=52)
Participants with any TEAEs	35 (81.4)	8 (88.9)	43 (82.7)
Drug-Related TEAEs	28 (65.1)	3 (33.3)	31 (59.6)
TEAEs Leading to Discontinuation	9 (20.9)	0	9 (17.3)
TEAEs Leading to Discontinuation (Titration)	7 (16.3)	0	7 (13.5)
TEAEs Leading to Discontinuation (Maintenance)	2 (4.7)	0	2 (3.8)
Participants with any SAEs	3 (7.0)	0	3 (5.8)
Number of Deaths	0	0	0

- The most common AEs observed were somnolence, decreased appetite, constipation, diarrhea and lethargy
- SAEs were ankle fracture, constipation and increased seizures
- Vast majority of participants stayed on bexicaserin once they achieved the maintenance phase
- Favorable safety and tolerability results

Topline Efficacy Results Bexicaserin Achieved Median Observed Countable Motor Seizure Reduction of 59.8% vs. 17.4% Placebo Across the DEE Study Population



Bexicaserin Achieved Placebo Adjusted Mean Seizure Reduction of 51.9% (p-value = 0.0206, post-hoc exploratory analysis)

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## Bexicaserin Achieved Median Seizure\* Reduction Across Dravet, LGS, DEE Other Cohorts



## PACIFIC Results Pave the Way for Global Phase 3 Program

Bexicaserin (LP352) achieved a median percent reduction from baseline in seizure frequency during the treatment period of:

59.8% in broad DEE population (42.4% placebo-adjusted)

- 74.6% in Dravet cohort
- **50.8%** in LGS cohort (33.4% placebo-adjusted)
- 65.5% in DEE Other cohort (33.3% placebo-adjusted)

Results were shown on top of a contemporary polytherapy background with multiple ASMs including cannabidiol (32.7% of participants were receiving cannabidiol)

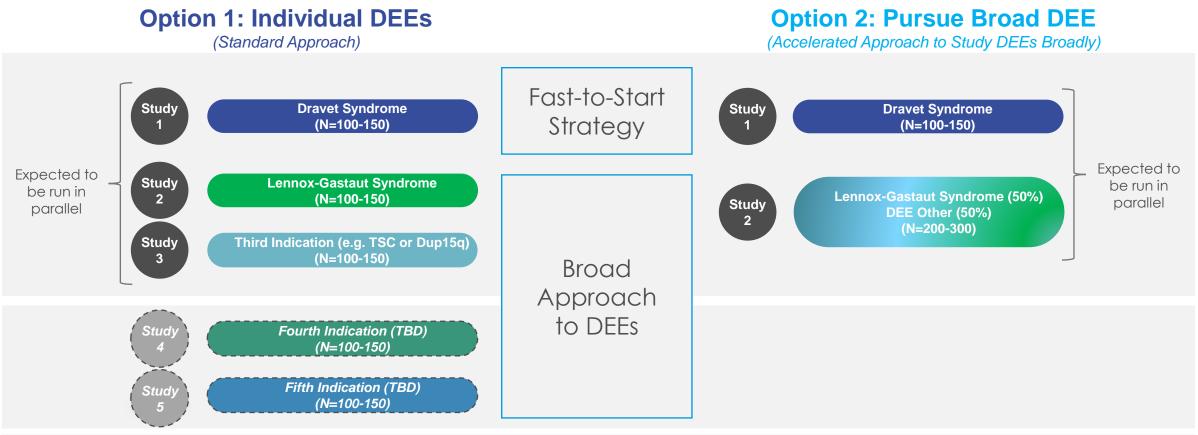
Favorable safety and tolerability results

- No echocardiograms required in PACIFIC study
- Metabolized via UGT pathway potentially reduces risk of Drug-Drug Interactions
- 86% of participants achieved the highest dose of 12 mg of bexicaserin in the maintenance period

100% of PACIFIC participants who completed the study entered the Open Label Extension Study

# Bexicaserin Phase 3 Global Program - Potential Paths Forward

Subject to Discussion with Regulatory Agencies



#### **Planned Study Parameters:**

Primary Endpoint: Reduction in Countable Motor Seizures

Ages:  $\geq 2$  to  $\leq 65$  yrs old (weight-based dosing for pts of lower weight/age)

Sites: ~80 sites across the US, AUS, EU, other potential regions

Open-Label Extension (OLE): Participants who complete any of Ph 3 studies are eligible to enter a 52-week OLE

# Commercial Opportunity for Bexicaserin (LP352)

## Potential Best-in-Class Profile Provides Multiple "Ways to Win"



Broaden market in "Approved 4" DEEs with preferred safety and burden profile



# Validating Continued Unmet Need in DEEs and Potential Of Bexicaserin (LP352)

#### **Background & Methodology**

#### **Quantitative HCP Research 100** Physicians

**Objective:** Validating Unmet Needs And **IP352** Potential

#### Criteria:

- Board Certified HCPs specializing in Neurology or **Epileptology**
- Treat at least 20 patients with DEEs in the past 12 months
- Familiar with Fintepla and Epidiolex
- Note: Most participants have some clinical experience with Epidiolex (92%) & Fintepla (68%)

#### **Qualitative HCP Research** 20 Physicians

**Objective:** Deeper Understanding Of Quantitative Findings (How & Why)

#### Criteria:

- Board Certified HCPs specializing in Neurology or Epileptology
- Treat at least **25 patients with DEEs** in the past 12 months
- Familiar with Fintepla and Epidiolex

Epileptologists = 5 (4 peds, 1 adult) Neurologists = 15 (13 peds, 3 adult)

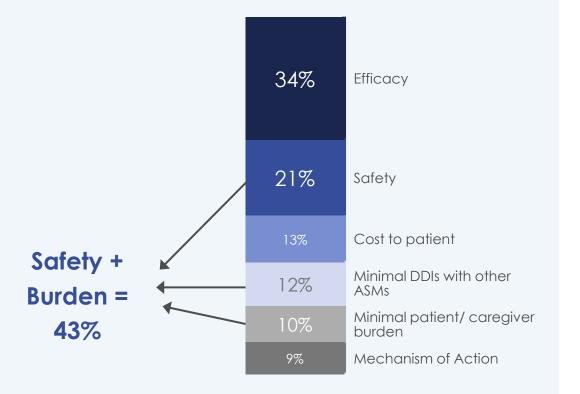
HCP = Health care providers

Based on Longboard sponsored third-party qualitative and qualitative market research studies involving physicians specializing in neurology or epileptology. Survey sampled product profile for LP352 case reviewed in this presentation: efficacy of 37-44% reduction in countable motor seizure frequency and generally well tolerated with BID dosing

# Surveyed HCPs Evaluate ASMs by Balancing Efficacy, Safety & Burden

#### **Influence on ASM Decisions**

When allocating 100 points across factors



# "

If I see a patient with epilepsy and give them enough valium, **they'll be seizure free**, **but then they'll be sleeping all day. That's not quality of life.** So, we must find the cocktail that gives them the **best seizure control with the least amount of side effects.**"

- Epileptologist, Primarily Pediatric

Based on Longboard sponsored third-party qualitative and qualitative market research studies involving physicians specializing in neurology or epileptology. See slide 30 for more information about the studies.

LONGBOARD PHARMACEUTICALS See side so for more information about the studies. \*Survey sampled product profile for the 5-HT2C agonist that included an efficacy of 37-44% reduction in countable motor seizure frequency and generally well tolerated with BID dosing

# LP659

Centrally Acting, Highly Selective Sphingosine-1-Phosphate (S1P) Receptor Modulator Targeting Multiple Neurological Diseases

# LP659 Potential Best-in-Class S1P Receptor Modulator with Broad Applicability

#### **STRONG SCIENTIFIC RATIONALE**

- Centrally acting \$1P receptor modulator
- Rapid onset & offset of action
- ✓ Highly selective to S1PR1,5
- ✓ No impact on \$1PR2,3 in preclinical models
- High oral bioavailability with direct impact on CNS glial cell \$1P receptors

#### **COMMERCIALLY ATTRACTIVE**

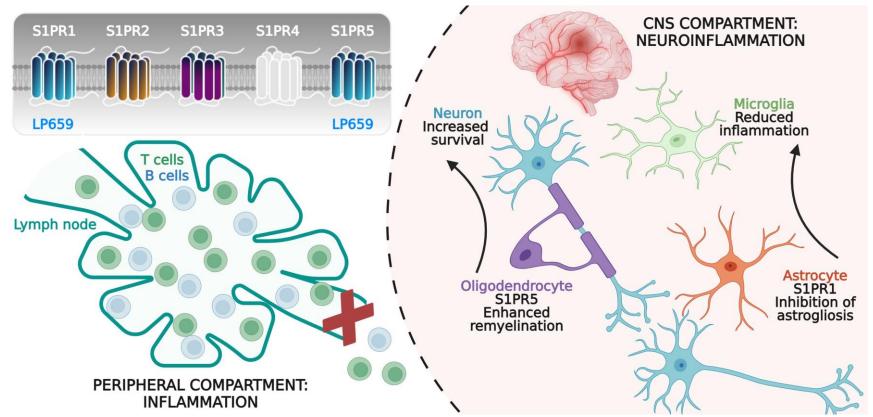
- S1P receptor modulators have generated billions of dollars of revenues in CNS indications
- Designed to potentially address multiple neurological disorders
- Opportunity for market leadership in \$1P receptor modulation in CNS

# S1PR1 Modulation Selectively Reduces Migration of Lymphocytes From Lymph Nodes

#### Treatment with S1P Receptor Modulator

- Functionally antagonizes S1PR1 by inducing receptor internalization and degradation, disrupting normal lymphocyte subset egress
- Decreases release of inflammatory cytokines and reduce organ/tissue damage
- Maintains immune surveillance
- Functional antagonism of S1PR1 receptor in astrocytes expected to attenuate neuroinflammation

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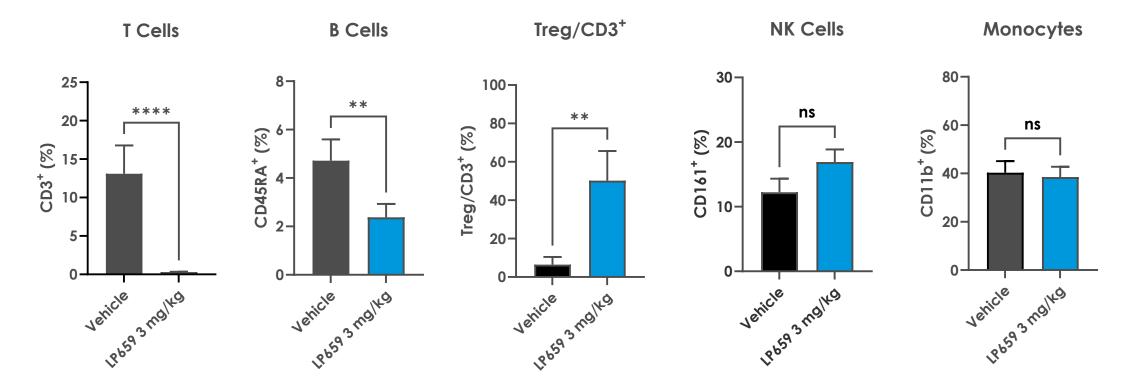
LP659 Designed to be a Potentially Next Generation Centrally-Acting S1PR1 Agonist with Greater Selectivity and Internalization-Biased Signaling

Most potent at S1PR1 internalization Greatest selectivity towards S1PR1 over S1PR5 50 25 S1PR1/5 **S1PR1** S1PR1 & 5 selective S1PR1 & 5 selective **40**· 20. **Relative Potency** Fold-selectivity 30. 15-**20** 10-10-5-0 Ω siponimod Fingolimod Ozonimod Ponesimod 18629 siponimod Oronimod Fingolimod

#### LP659 selectivity may limit off-target effects associated with currently approved S1P receptor modulators

# Modulation of Immune Tolerance Drives Efficacy by LP659

- LP659 potency in vivo parallels T and B cell lowering potential
- Proportion of Tregs over total CD3<sup>+</sup> cells is significantly increased by LP659
- No significant effects on NK and monocyte frequencies



# LP659 Ameliorated Disease Phenotypes in Multiple Preclinical Models

Disease / MoA	Autoimmune, CNS involvement	Autoimmune, CNS involvement	Autoimmune, PNS involvement	Autoimmune, PNS involvement	Neuro Degenerative
Model	Induced	Induced	Induced	Induced	Genetic
Species	Rat	Mouse	Rat	Rat	Human iPSC
Dosing	Prophylactic	Prophylactic	Prophylactic	Therapeutic	Therapeutic
Activity	+	+	+	+	+
Results	Dose-dependent amelioration of disease severity with parallel reduction of circulating T lymphocytes	Dose-dependent amelioration of disease severity with reduction of T and B cell infiltration, inflammatory markers, and loss of myelin in the spinal cord	Dose-dependent halting of disease progression with reduction of inflammatory cell infiltration and loss of myelin in the sciatic nerve	Blunting of disease severity with corresponding reduction of inflammatory cell infiltration in the sciatic nerve	Dose-dependent rescue of hyperexcitability in control neurons co- cultured with diseased astrocytes

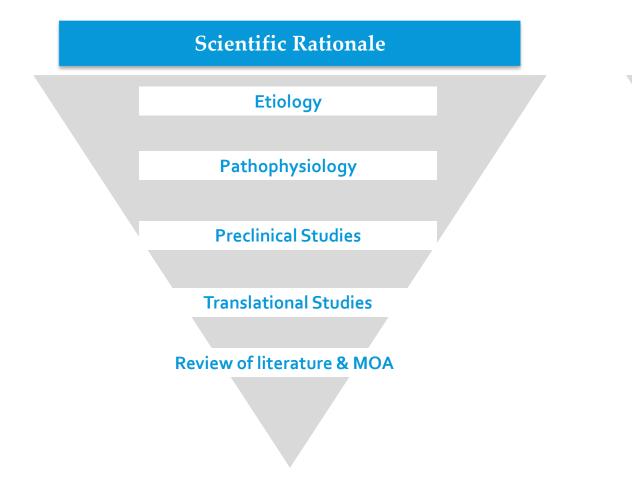
# LP659: Phase 1 Single-Ascending Dose (SAD) Study Objectives

First-In-Human, Randomized, Double Blind, Placebo Controlled, SAD Study to Assess the Safety, Tolerability, Pharmacokinetics & Pharmacodynamics of LP659 in up to 48 Adult Healthy Volunteers

#### Key Study Objectives:

- Assess the safety and tolerability of single ascending doses of LP659
- Determine the PK profile of LP659, and its metabolite(s), in single ascending doses
- Determine PD profile of single ascending doses of LP659

### Longboard Indication Assessment Process



#### **Commercial & Development Criteria**

Degree of Unmet Need

Patient Advocacy & Recruitment

**Clinical Trial Feasibility** 

Speed of Clinical Development

Approved SoC

**Competitive Headroom** 

**Disease Prevalence** 

**Pricing Potential** 

**Strategic Indication Selection** 

# Financial Summary & Upcoming Milestones

Cash, Cash Equivalents
 & Investments

~\$272.4 million As of January 31, 2024

Private Placement \$59.9 million Completed in March 2024

Shares Outstanding 38.9 million

As of March 28, 2024

#### Full-Year 2023 Operating Expenses

#### \$56.8 million

- R&D \$43.8 million
- G&A \$13.0 million As of December 31, 2023

	Key Milestones	Anticipated Timing
	PACIFIC Ph 1b/2a Topline Data in DEE Study	🕑 Q1 2024
Bexicaserin	PACIFIC Data at medical meetings	Q2 2024
(LP352)	PACIFIC Open-Label Extension Data	H2 2024
	Global Ph 3 Program Initiation	YE 2024
LP659	Ph 1 Initiation	🖌 Q4 2023
	Topline SAD Data	Q2 2024

Multiple clinical and preclinical studies in process to further support the development of bexicaserin & LP659

# Thank you

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