



PACIFIC Study Topline Data

JANUARY 2, 2024

Forward-Looking Statements

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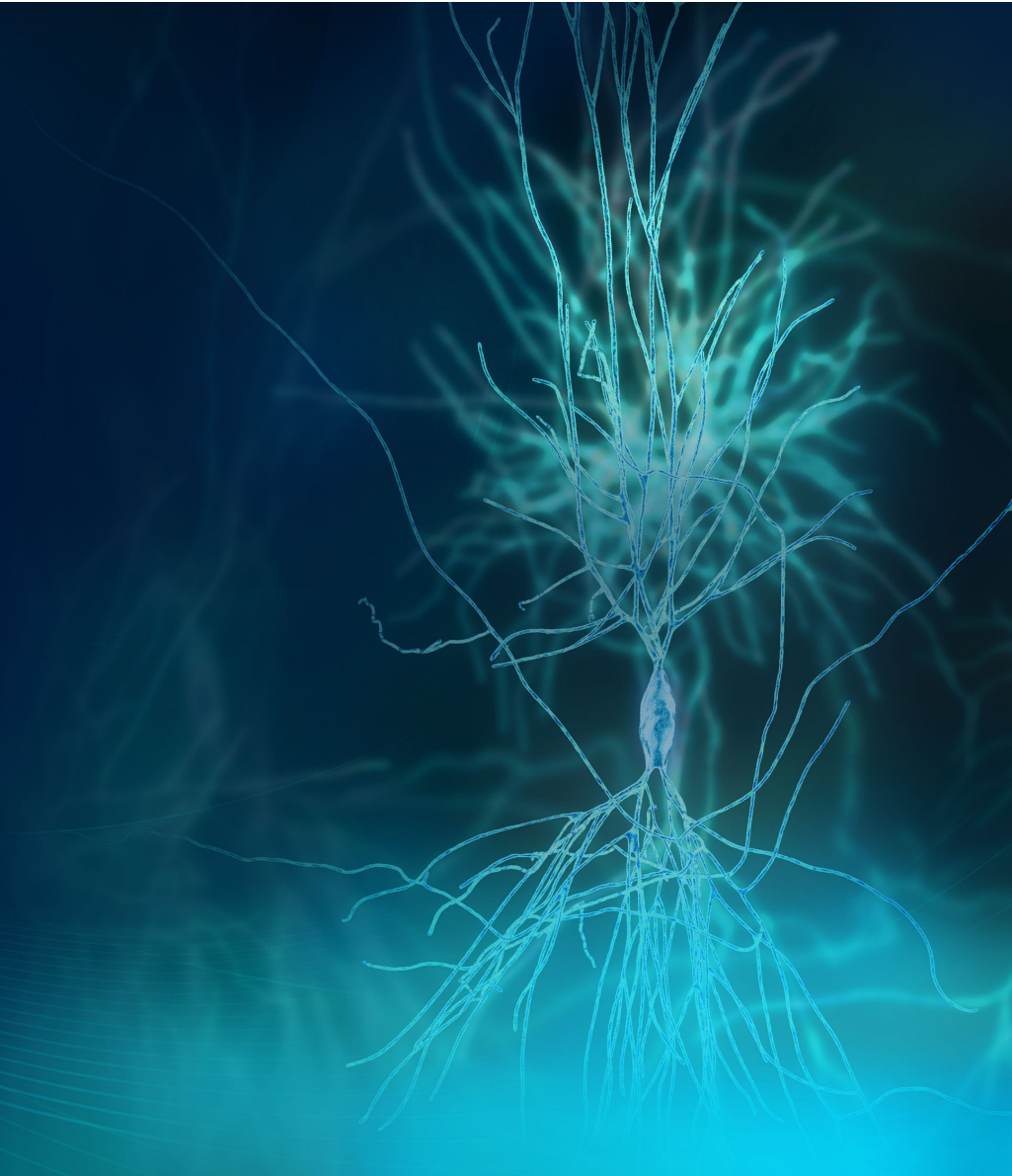
This presentation discusses product candidates that have not yet been approved for marketing by the U.S. Food and Drug Administration (the "FDA").





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Bexicaserin Has the Potential to Change the DEE Landscape

 **53.3%**

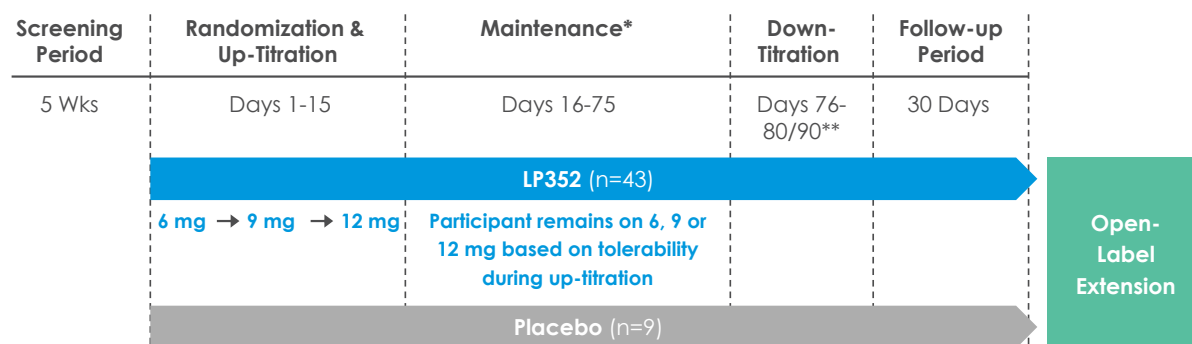
Median Reduction in Seizures*

- **72.1%**  **Dravet**
- **48.1%**  **LGS**
- **61.2%**  **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 through 2041**** provides the opportunity to maximize the full potential of LP352
- Rapidly moving forward with preparations for a **global Phase 3 program**



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



Key Inclusion Criteria:

- 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:** ≥ 12 to ≤ 65 yrs old

No Echocardiograms Required in PACIFIC



Double-blind, placebo-controlled 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

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



All participants: Treatment-resistant countable motor seizures with average of ≥ 4 observed/countable motor seizures per 4-week 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: <ul style="list-style-type: none"> • 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



Topline Participant Disposition & Safety Results Summary



Demographics, Baseline Characteristics & Concomitant Medications

Parameter	n(%)	Statistics	Bexicaserin (N=43)	Placebo (N=9)	Overall (N=52)
Age (Years)		Mean	23.8	26.7	24.3
		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)
		Female	22 (51.2)	2 (22.2)	24 (46.2)
Weight (kg)		Median	55.20	72.76	59.36
		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
Concomitant Medications		Clobazam	21 (48.8)	2 (22.2)	23 (44.2)
		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(%)	Overall		Dravet Syndrome		LGS		DEE 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



86% of Bexicaserin Treated Participants Achieved 12 mg Dose

Highest tolerated dose achieved for the Maintenance Period

	n(%)	Bexicaserin (LP352)	Placebo	Overall
All Participants		35	9	44
6 mg		1 (2.9)	0	1 (2.3)
9 mg		4 (11.4)	0	4 (9.1)
12 mg		30 (85.7)	9 (100)	39 (88.6)
Dravet Syndrome		3	0	3
6 mg		0	0	0
9 mg		1 (33.3)	0	1 (33.3)
12 mg		2 (66.7)	0	2 (66.7)
LGS		17	5	22
6 mg		0	0	0
9 mg		1 (5.9)	0	1 (4.5)
12 mg		16 (94.1)	5 (100)	21 (95.5)
DEE Other		15	4	19
6 mg		1 (6.7)	0	1 (5.3)
9 mg		2 (13.3)	0	2 (10.5)
12 mg		12 (80.0)	4 (100)	16 (84.2)



Safety Results Summary

n(%)	Bexicaserin (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

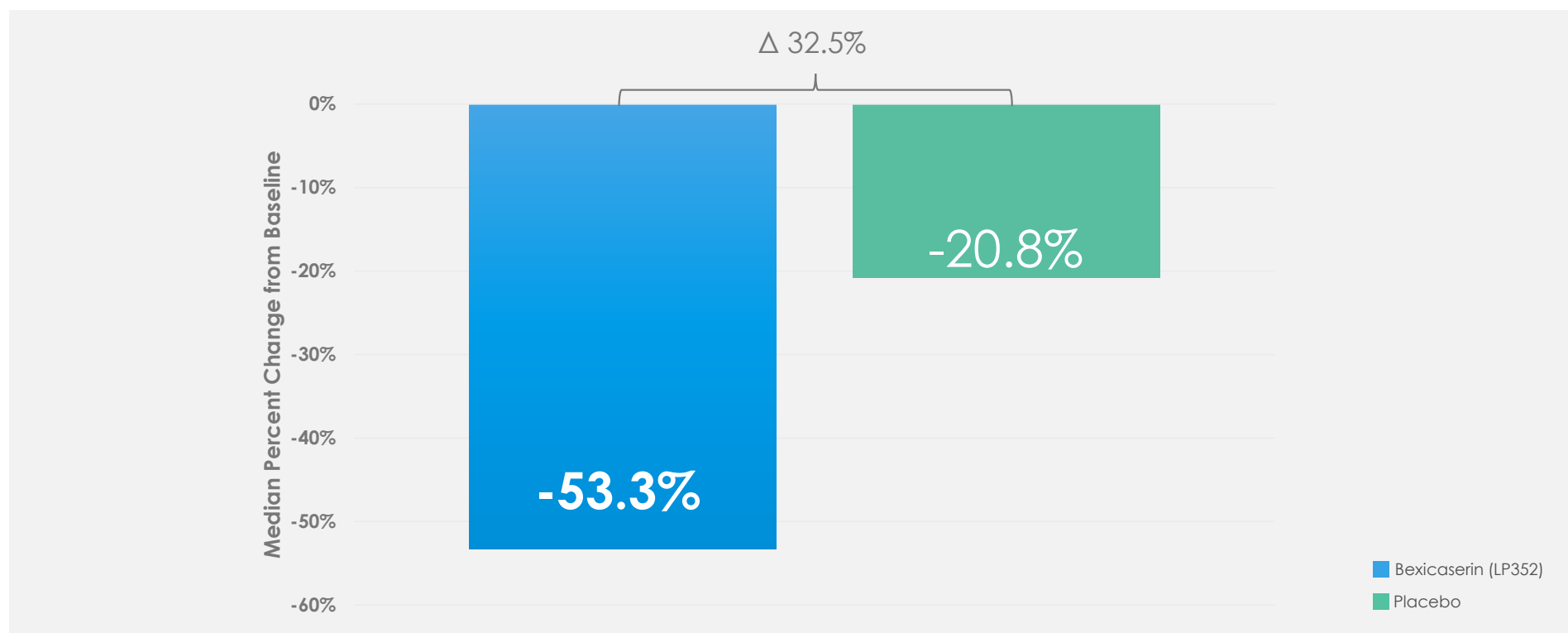
- The most common AEs* observed were somnolence, decreased appetite, constipation, diarrhea, lethargy, tremor, urinary tract infection, fatigue, pyrexia and agitation
- SAEs were ankle fracture, constipation and increased seizures
- Vast majority of participants stayed on bexicaserin once they achieved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



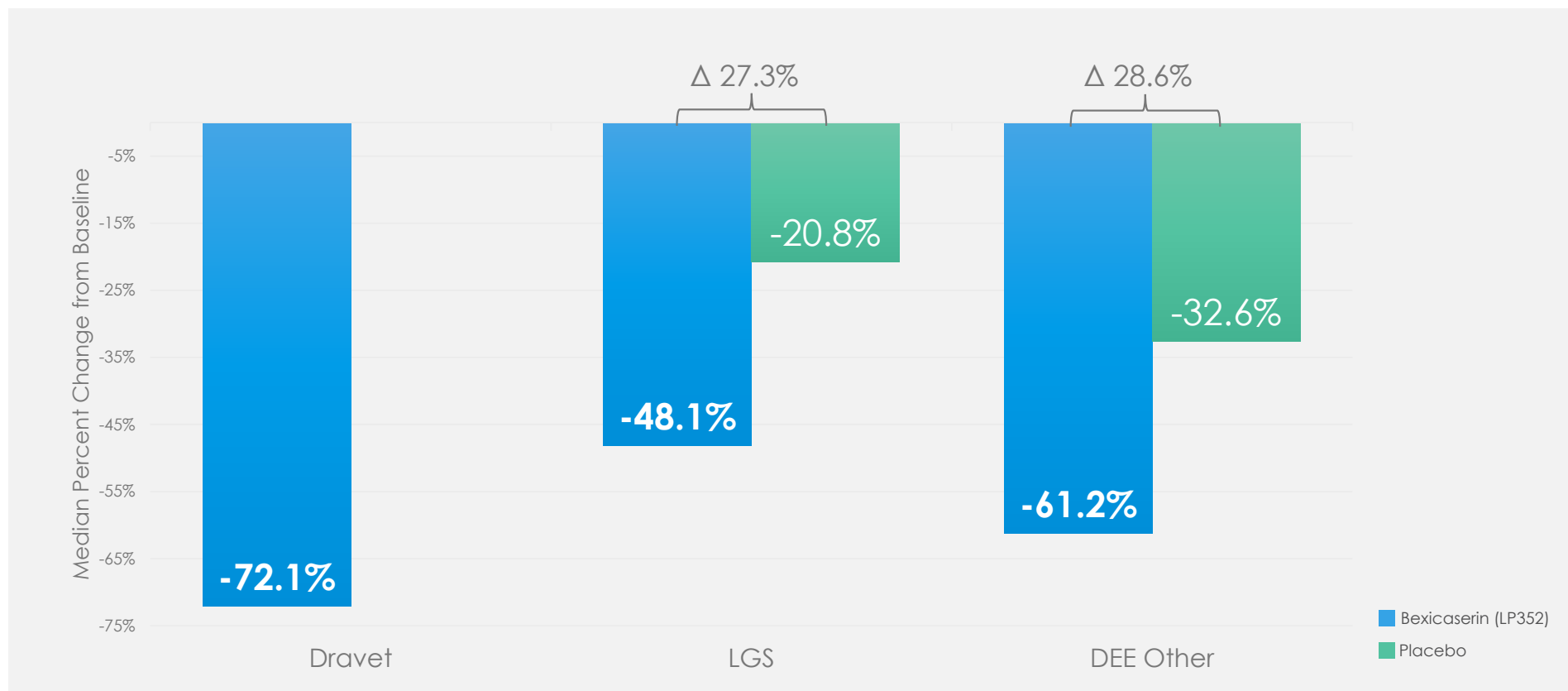
Topline Efficacy Results



Bexicaserin Achieved Median Seizure Reduction of 53.3% in Countable Motor Seizures Compared to 20.8% for Placebo Across the DEE Study Population



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



PACIFIC Results Pave the Way for Global Phase 3 Program

PACIFIC demonstrated meaningful efficacy results across a broad DEE population as well as in individuals with LGS and Dravet

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

53.3% in broad DEE population (32.5% placebo-adjusted)

72.1% in Dravet cohort

48.1% in LGS cohort (27.3% placebo-adjusted)

61.2% in DEE Other cohort (28.6% placebo-adjusted)

Results were demonstrated 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

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

- Awaiting analysis of the full PACIFIC dataset
- Utilizing key learnings for incorporation into the global Phase 3 program

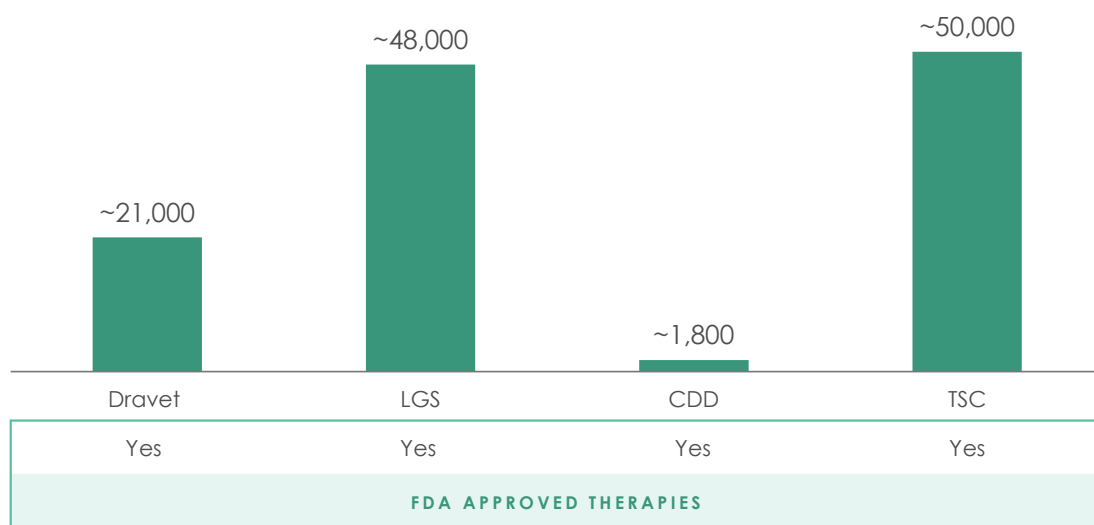


Summary & Next Steps



4 DEE Syndromes Have Approved Therapies; 20+ Have None

"Approved 4" DEE Prevalence (US)



Other DEEs

- DUP15q
- SCN2A related epilepsies
- SCN8A related epilepsies
- KCNQ2 related epilepsies
- KCNQ3 related epilepsies
- Angelman syndrome
- Landau-Kleffner Syndrome
- Early Myoclonic Encephalopathy
- KCNT1 related epilepsies
- SynGAP1 related epilepsies
- Rett Syndrome
- Ohtahara Syndrome
- PCDH19
- EE w/ Continuous Spike-Wave
- West Syndrome
- Myoclonic Atonic Epilepsy
- Ring14
- Ring20
- Others

NO SPECIFICALLY APPROVED THERAPIES

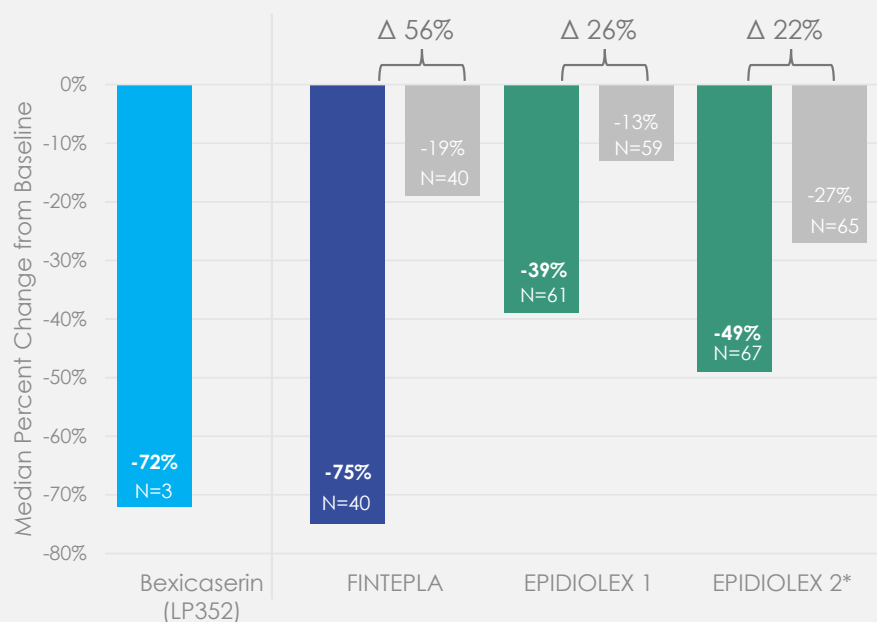
The prevalence of all "Other DEEs" could exceed the total of the "Approved 4" combined

Sources: Dravet Syndrome Foundation, LGS Foundation

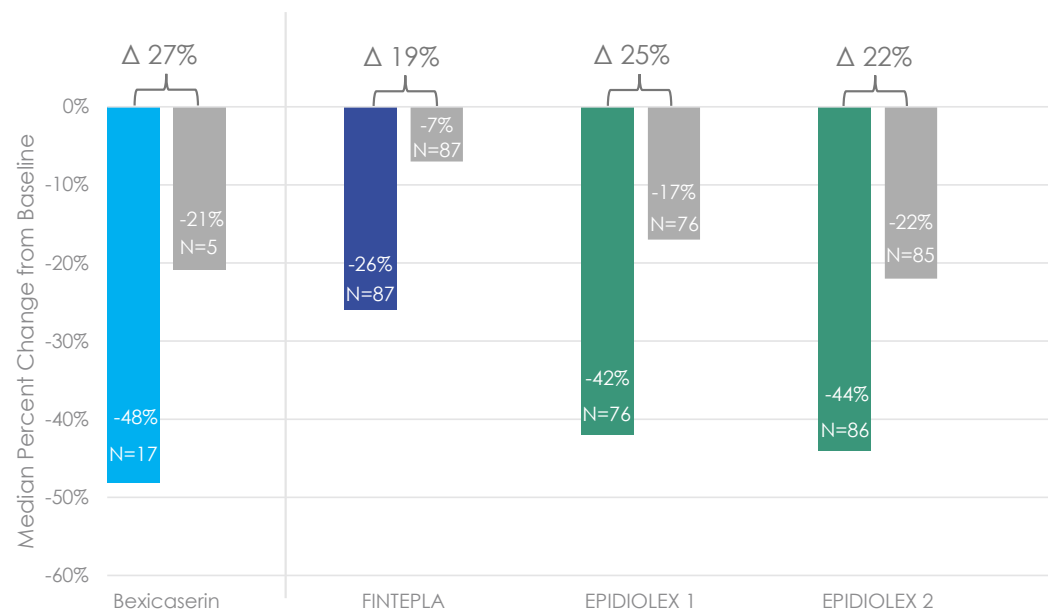
Definitions: LGS = Lennox-Gastaut Syndrome, CDD = CDKL5 Deficiency Disorder, TSC = Tuberous Sclerosis Complex; DEE = Developmental and Epileptic Encephalopathy

Competitive Landscape: Median Seizure Reduction for Bexicaserin and Approved Compounds in Dravet and LGS

Dravet Syndrome^{1,3,4}



Lennox-Gastaut Syndrome^{2,3,4}

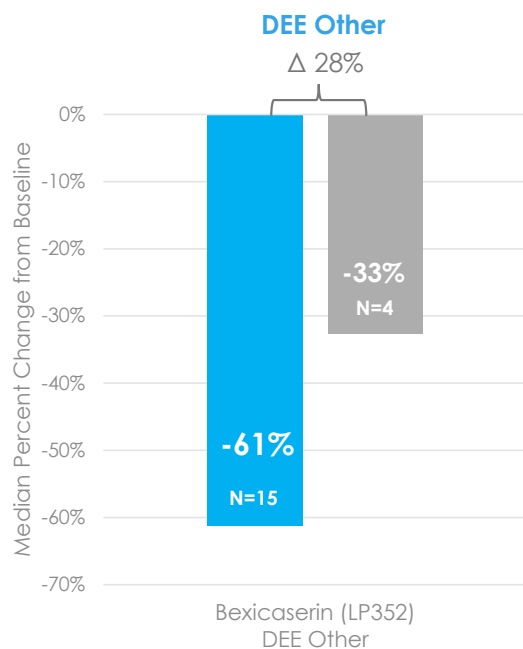


FOR ILLUSTRATIVE PURPOSES ONLY: Not a head-to-head comparison. Differences exist between trial designs and subject characteristics, and caution should be exercised when comparing data across studies.

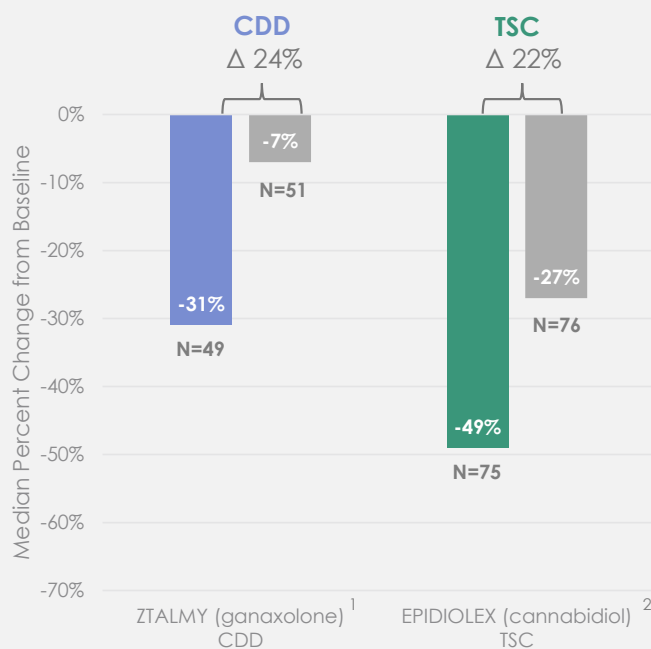
■ Bexicaserin (LP352)
■ FINTEPLA® (fenfluramine)
■ Epidiolex (cannabidiol)
■ Placebo

Significant Unmet Need in DEEs (Beyond Dravet and LGS)

“DEE Other” in PACIFIC



“Other DEEs” with Specifically Approved Therapies



“Other DEEs” With No Specifically Approved Therapies




- DUP15q
- SCN2A related epilepsies
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- EE w/ Continuous Spike-Wave
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- Ring14
- Ring20
- Others

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Thank you

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