

Relieve pain. Restore health.

Corporate Presentation | September 2026



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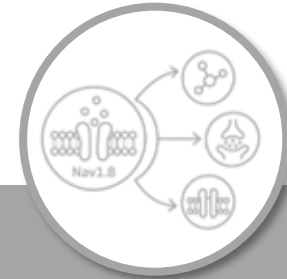
Reimagining Pain Management Away From Opioids: Building The Innovative Pain Company



Targeting pain, one of the largest areas of unmet need, where innovation has been limited



Late clinical stage portfolio of differentiated $\text{Na}_v1.8$ inhibitors across acute (Onzotrigine¹) and chronic pain (LTG-321)



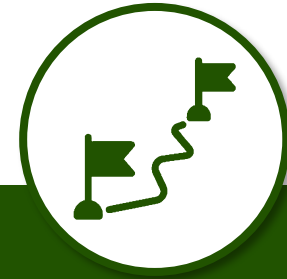
Opportunity to expand to complementary mechanism of actions additive to $\text{Na}_v1.8$



Deep capabilities across R&D to pioneer innovation in pain



Well capitalized with ~\$397MM raised in upsized IPO and expected cash runway into 2029



Multiple clinical milestones expected in 2H 2027

¹ Previously referred to as LTG-001

Experienced Leadership Team with Deep Expertise in Pain



Nima Farzan
CEO

- 3X CEO, IPO and ~\$700m capital raised as CEO
- Private & public company exits



Neil Singla
CMO

- Founder/CEO of preeminent pain CRO/site network
- Along with team, conducted >250 pain studies over 20+ years



Bryan Moyer
SVP, Discovery

- 20+ years of pain and CNS research
- Discovered multiple compounds in clinic



Neha Krishnamohan
CFO / CBO

- 3X CFO, 15+ years of experience in healthcare & finance
- Executed \$100B+ financing and M&A transactions



Naomi Lowy
SVP, Global
Regulatory Affairs

- 18+ years in FDA & regulatory advisory
- Former deputy director pain division of FDA (DAAP)¹
- Co-authored recent FDA acute/chronic pain guidances



¹ DAAP: Division of Anesthesia, Addiction Medicine, and Pain Medicine



Latigo Built for the Emerging Inflection Point in Pain Innovation

Committed to Building a Best-in-Class, Multi-modal, Pain Focused Company



Na_v1.8
Pioneering
Innovation in
Pain



New Emerging
Pain Targets



Scarcity of Pain
Pipelines



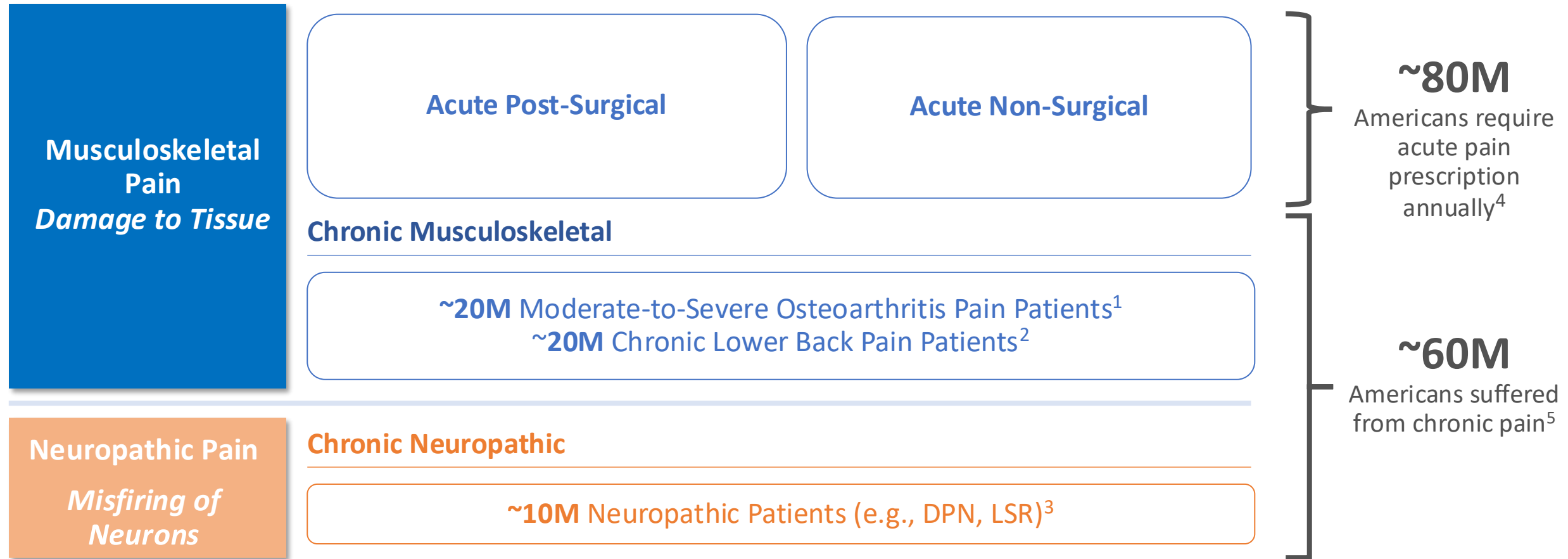
Differentiated
Pain Capabilities
at Latigo

Pain represents a significant market with substantial unmet need



Pain Remains a Debilitating Condition, Despite 250M Pain Prescription Medications Every Year in the US

Pain represents one of the largest and most pervasive therapeutic markets in the US



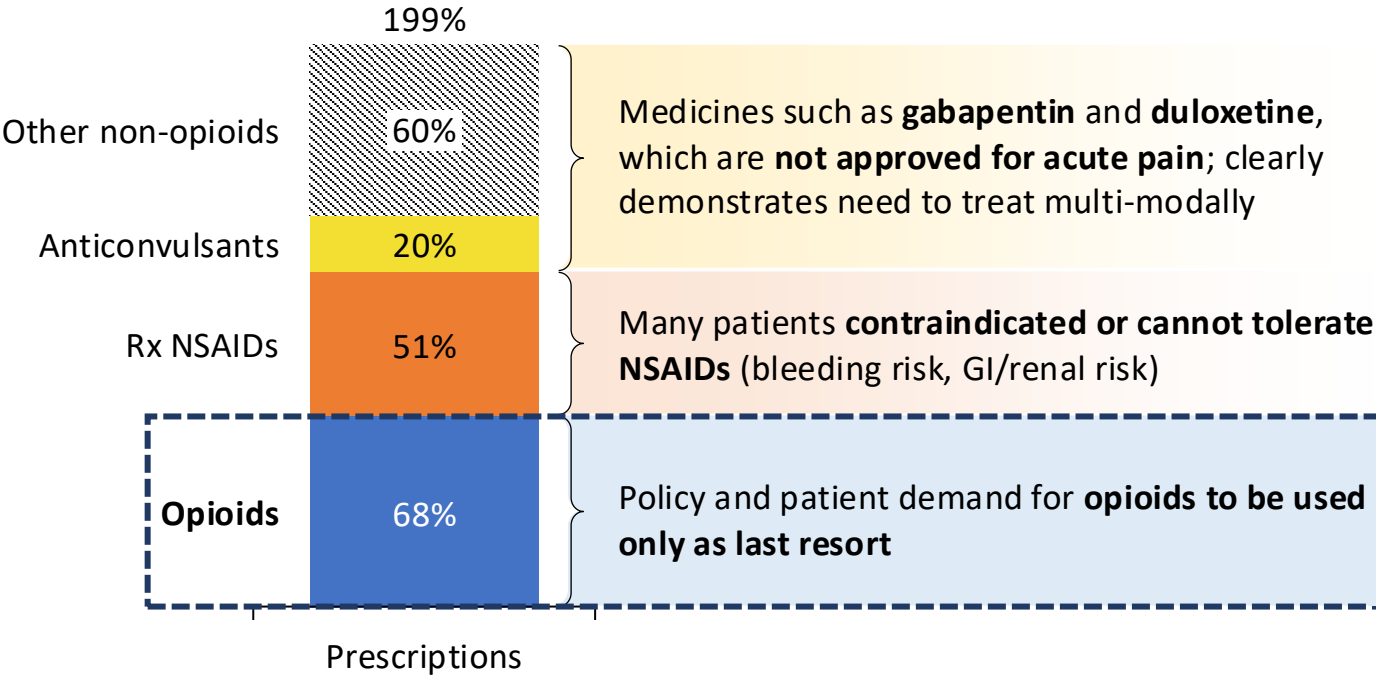
¹ 2020 CDC data and Schepman et al., JHEOR. 2022;9(1):58-67; ² Feldman and Nahin, The Journal of Pain, 2022; 23, 2144-2154; ³ Lopez et al., Pain Reports. 2026 Jan; 11; ⁴ AARP 2025 ⁵ Zajacova et al., Pain. 2026 Jan 1;167(1):142-149.
Note: DPN: Diabetic Peripheral Neuropathy; LSR: Lumbosacral Radiculopathy



Substantial Unmet Need in Pain Demonstrated by Significant Off-Label Use and Continued Reliance on Opioids, Despite Clear Risks

Average Acute Pain Patient Receives ~2 Pain Management Prescriptions¹

Percent of patients receiving each type of prescription pain medicine per episode
(sums to more than 100% due to multi-modal treatment)



Opioid Challenges

- **Addiction Risk:** ~11% of patients persistently use opioids after minor surgical procedures²
- **Dispersion Risk:**
 - Unused opioids can be dispersed into community
 - ~67-92% of patients have excess opioids after surgery³
- **Side Effects:** Constipation, nausea, sedation etc.

Note: Other includes Na_v1.8 blockers, neuropathic adjuvants (e.g., Gabapentin, Duloxetine), muscle relaxants, topical / local anesthetics; steroid injections and any combination therapy with Opioid / NSAIDs
¹2025 Komodo claims data analysis. ² Lawal OD, et al. JAMA Netw Open. 2020;3(6):e207367; ³ Bicket MC, et al. JAMA Surg. 2017;152(11):1066-1071

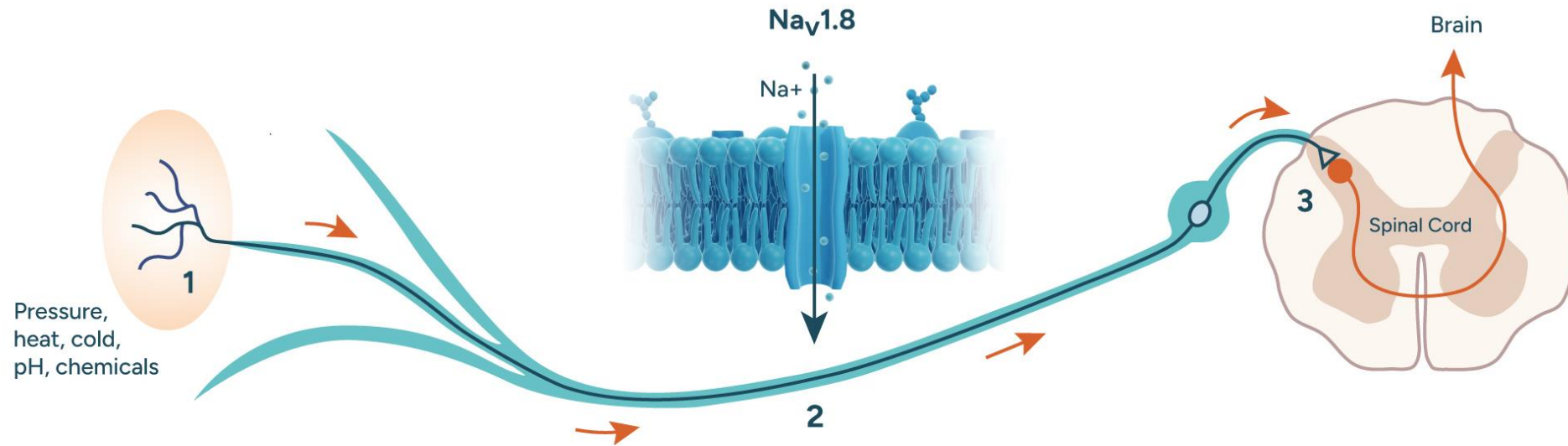


Na_v1.8 Transmits Pain Signals Independent of Source, Peripherally Blocking Pain Sensation

Transduction of pain signal

Transmission of pain signal

Perception of pain signal



Na_v1.8 transmits diverse pain signals

Modified from Waxman and Zamponi Nat Neurosci 17(2): 153-63, 2014 and Chung et al Nat Rev Disease Primers 8:45, 2022

Latigo Pipeline of Pain Therapies, Focused on Na_v1.8

Molecule	Discovery	Preclinical	Phase 1	Phase 2	Phase 3	Recent Progress / Anticipated Milestones
Onzotrigine (LTG-001) Acute Pain	Demonstrated highest analgesic effect seen in abdominoplasty trial*; results published in NEJM ¹					Abdominoplasty Trial Completed ² Phase 3 Bunionectomy & Safety Trial Initiation 2H 2026; Topline Results 2H 2027
LTG-321 Chronic Pain	Potential First-in-Class in OA					Phase 2 Initiated; Topline Results 2H 2027
Onzotrigine IV Acute Pain	Designed to allow IV to oral therapy transition					Bioavailability Study Completed
LTG-418	Potential for improved potency, novel formulations					Drug Candidate Selected

Undisclosed targets in discovery for chronic pain

*Based on management's assessment of currently available information.

¹ The New England Journal of Medicine

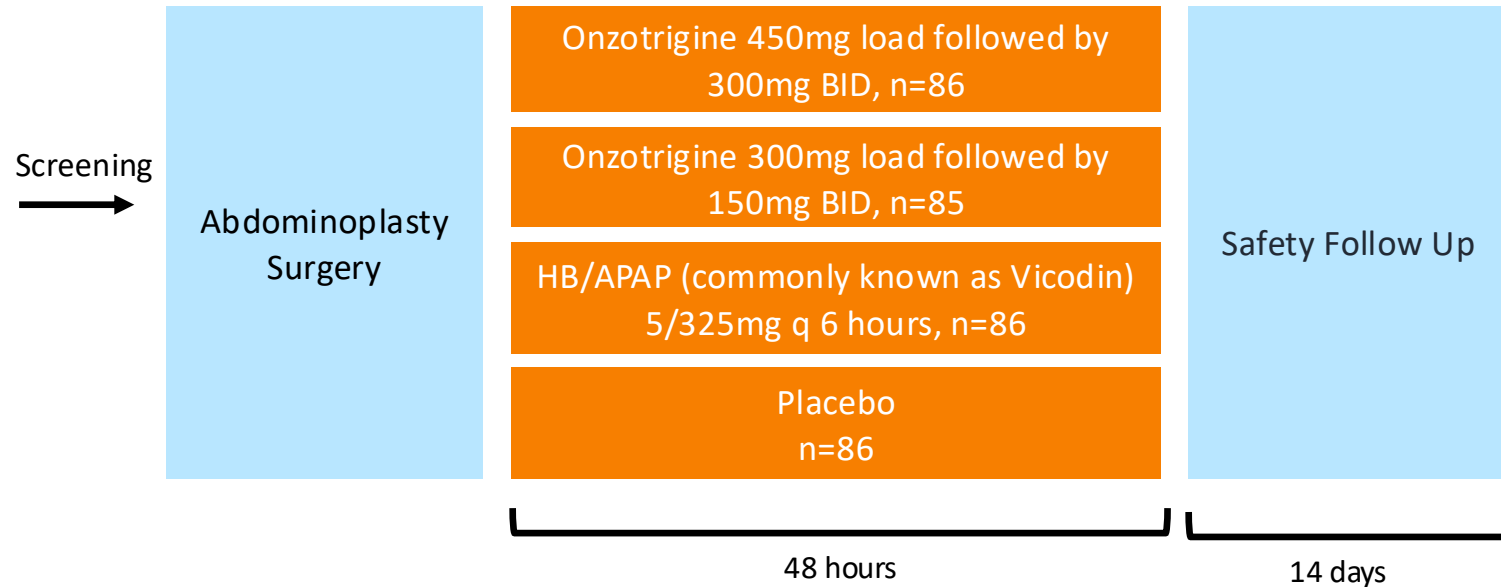
² Based on our discussions with the FDA, this trial may serve as one of the two pivotal adequate well-controlled trials required to support approval of onzotrigine for the treatment of moderate to severe acute pain, including postoperative pain.



Abdominoplasty Trial Exploring Onzotrigine – One of Two Pivotal Adequate Well-Controlled Trials to Support Acute Pain Approval¹

Onzotrigine
Acute Pain

Dose-ranging trial of Onzotrigine in pain after abdominoplasty, n=343



Enrollment initiated – July 2025

Enrollment completed – December 2025

Primary Endpoint

- Summed Pain Intensity Difference (SPID) from 0-48 hours, high dose onzotrigine vs. placebo

Design (matched to suzetrigine for benchmarking)

- Same active opioid comparator as used with suzetrigine Phase 3
- Same time-to-onset measures used in suzetrigine Phase 3
- Added opioid rescue medication (oxycodone 5mg) to enable opioid-sparing endpoints vs. ibuprofen 400mg for suzetrigine

Disposition and Demographics

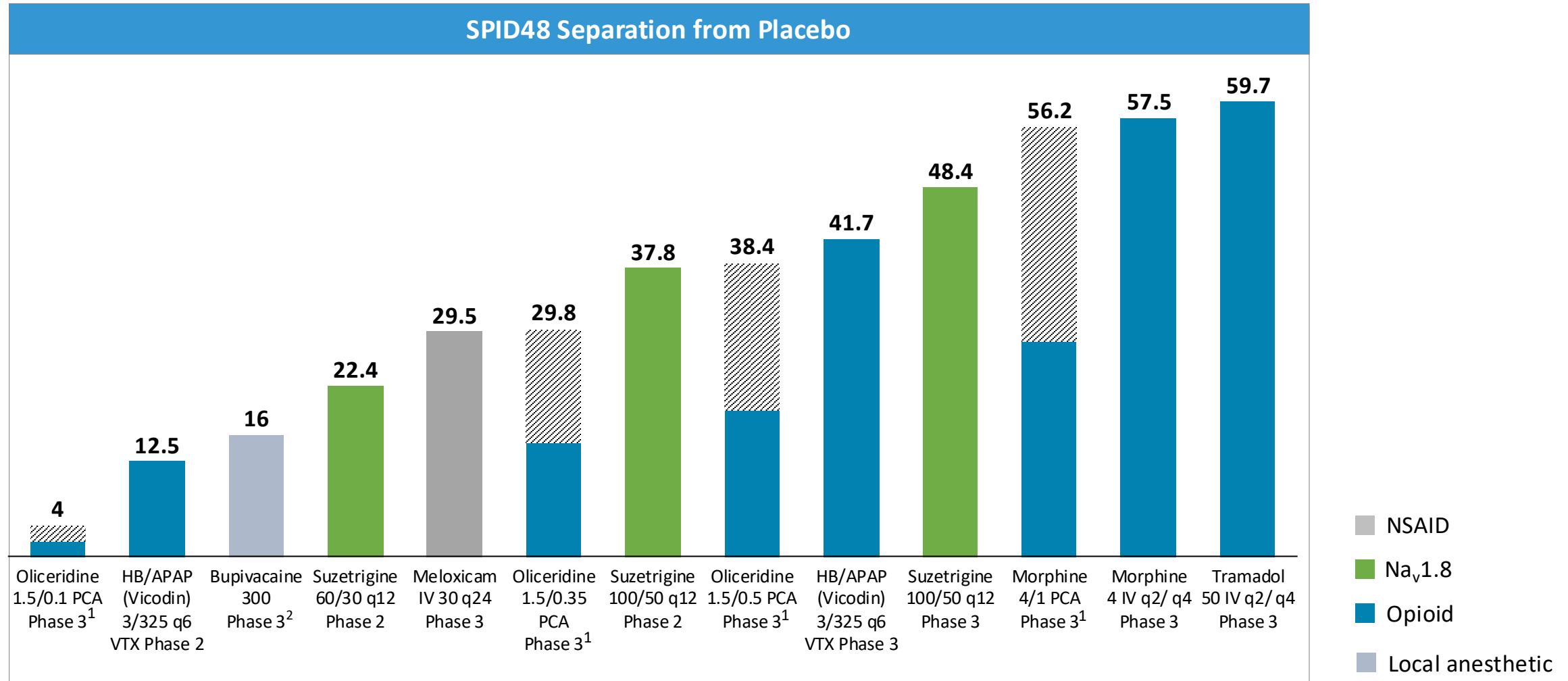
- 4 sites in the US
- 334 (97%) of subjects completed study treatment
- 328 (96%) of subjects completed study
- Average age 41 years, 68% of patients were white
- Baseline NPRS: 7.4

¹Based on our discussions with the FDA, this trial may serve as one of the two pivotal adequate well-controlled trials required to support approval of onzotrigine for the treatment of moderate to severe acute pain, including postoperative pain.



Analgesic Effect Seen in Separate Previously Reported Abdominoplasty Trials (6 Trials, 13 Dose Arms)

Onzotrigine
Acute Pain



Note: All dosing in mg unless otherwise noted; ¹Estimated by doubling of SPID24 data for oliceridine dose arms and morphine 4mg IV bolus + 1mg IV PCA from oliceridine label ²bupivacaine hydrochloride implant. Reflects AUC48

Note: These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.



Onzotrigine High Dose: 50% Greater Analgesic Effects than Vicodin Comparator; 2x Faster than Third-Party Phase 3 Trial of Suzetrigine

Onzotrigine
Acute Pain

Onzotrigine Observed Differentiation

~30% greater analgesic effect vs. suzetrigine¹

~50% greater analgesic effect vs. Vicodin

2.3x faster pain relief

Opioid Sparing

	Onzotrigine abdominoplasty High Dose ¹	Suzetrigine Phase 3 abdominoplasty ²
Analgesic Effect: SPID48 placebo subtracted		
Study Drug	62	48
HB/APAP (Vicodin)	41	42
Onset: Time to Meaningful Pain Relief	52 minutes	119 minutes
Opioid Sparing: Participants Not Requiring Opioid Rescue	52%	Not studied
		19% didn't use rescue (ibuprofen)

¹ Onzotrigine High Dose: 450mg loading dose followed by 300mg BID

² Suzetrigine phase 3 abdominoplasty trial. These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.



Abdominoplasty Trial Results Published in NEJM on July 30th 2026¹

Onzotrigine
Acute Pain

Only the second original research publication in NEJM reporting clinical results for a novel drug for acute pain in the last 15 years

ORIGINAL ARTICLE

Phase 2b Trial of a Na_v1.8 Inhibitor for Acute Pain

Neil Singla, M.D.,¹ Nathaniel P. Katz, M.D.,^{2,3} Ben Vaughn, M.S.,¹ Timothy Rogier,¹ Todd Bertoch, M.D.,⁴ Harold Minkowitz, M.D.,^{3,6} and Mallory Loflin, Ph.D.¹

ABSTRACT

Includes comparative data of the analgesic effects observed with onzotrigine (LTG-001) relative to other agents, including IV opioids in abdominoplasty

DISCUSSION

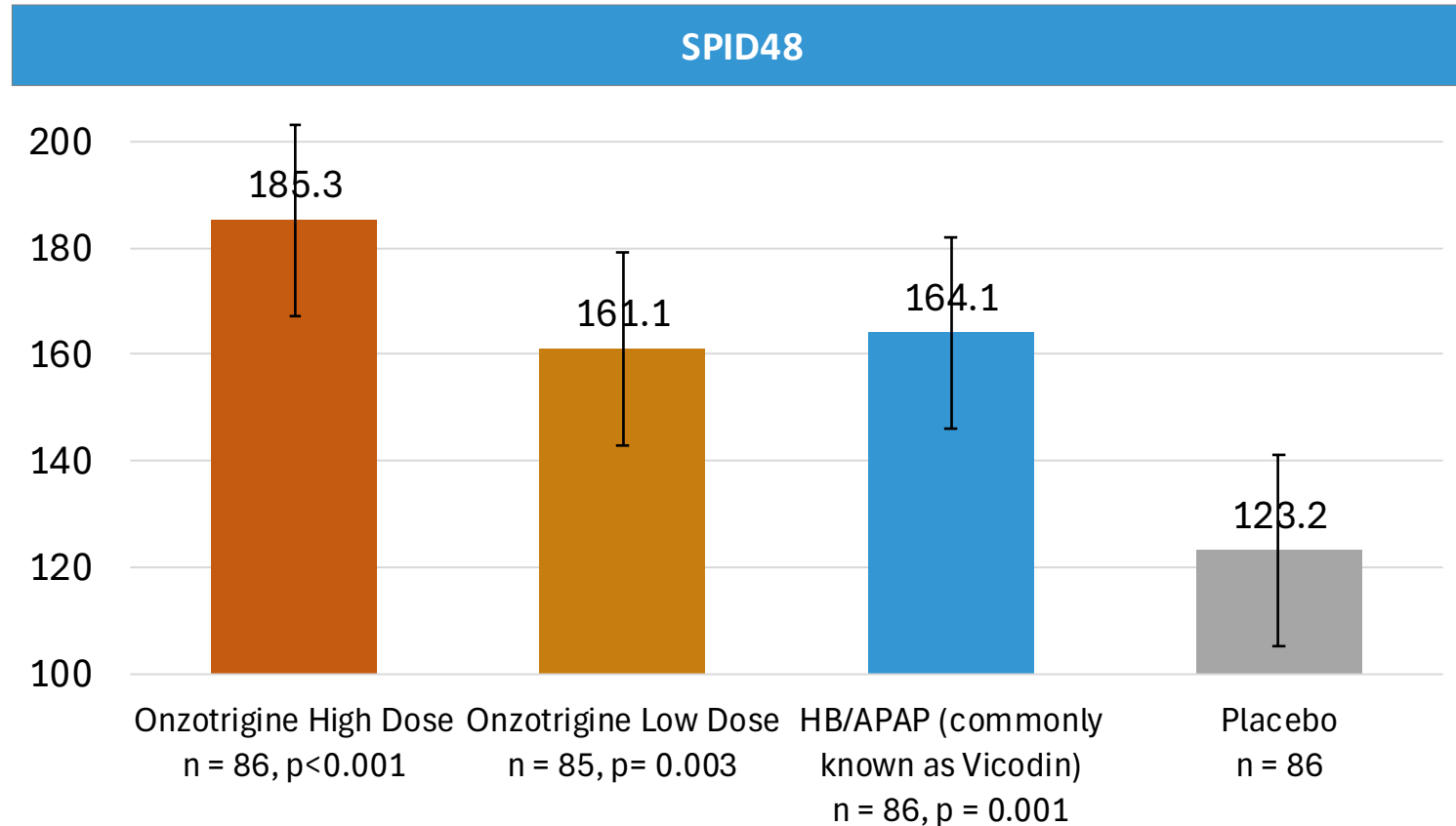
For contextualization of the results, the magnitude of the effect²⁴ was compared with the published results from other placebo-controlled trials that were conducted under a sponsor-initiated investigational new drug application and had the SPID48 as an outcome in patients undergoing abdominoplasty.^{13,25,26} In a comparison of the least-squares mean difference in the SPID48 from placebo across these trials, the value for high-dose LTG-001 (62.1) was similar to those for opioids that were administered intravenously (57.5 and 59.7),^{18,25} and the value for low-dose LTG-001 (37.8) was similar to those for the approved Na_v1.8 inhibitor suzetrigine (48.4) and for oral hydrocodone bitartrate–acetaminophen (40.9 and 41.7) and higher than the value for meloxicam (an NSAID) administered intravenously (29.5) (Fig.

¹ NEJM: New England Journal of Medicine; N Engl J Med 2026;395:454-64.



Onzotrigine Demonstrated Significantly More Pain Reduction than Placebo (Primary Endpoint of SPID48)

Onzotrigine
Acute Pain



- All active arms showed statistically significant separation from placebo ($p = <0.001-0.003$)
- Onzotrigine high dose Standardized Effect Size (SES) of 0.74 compared to 0.53 for suzetrigine¹ (onzotrigine 40% higher)

SPID48 placebo
subtracted

62.1

37.8

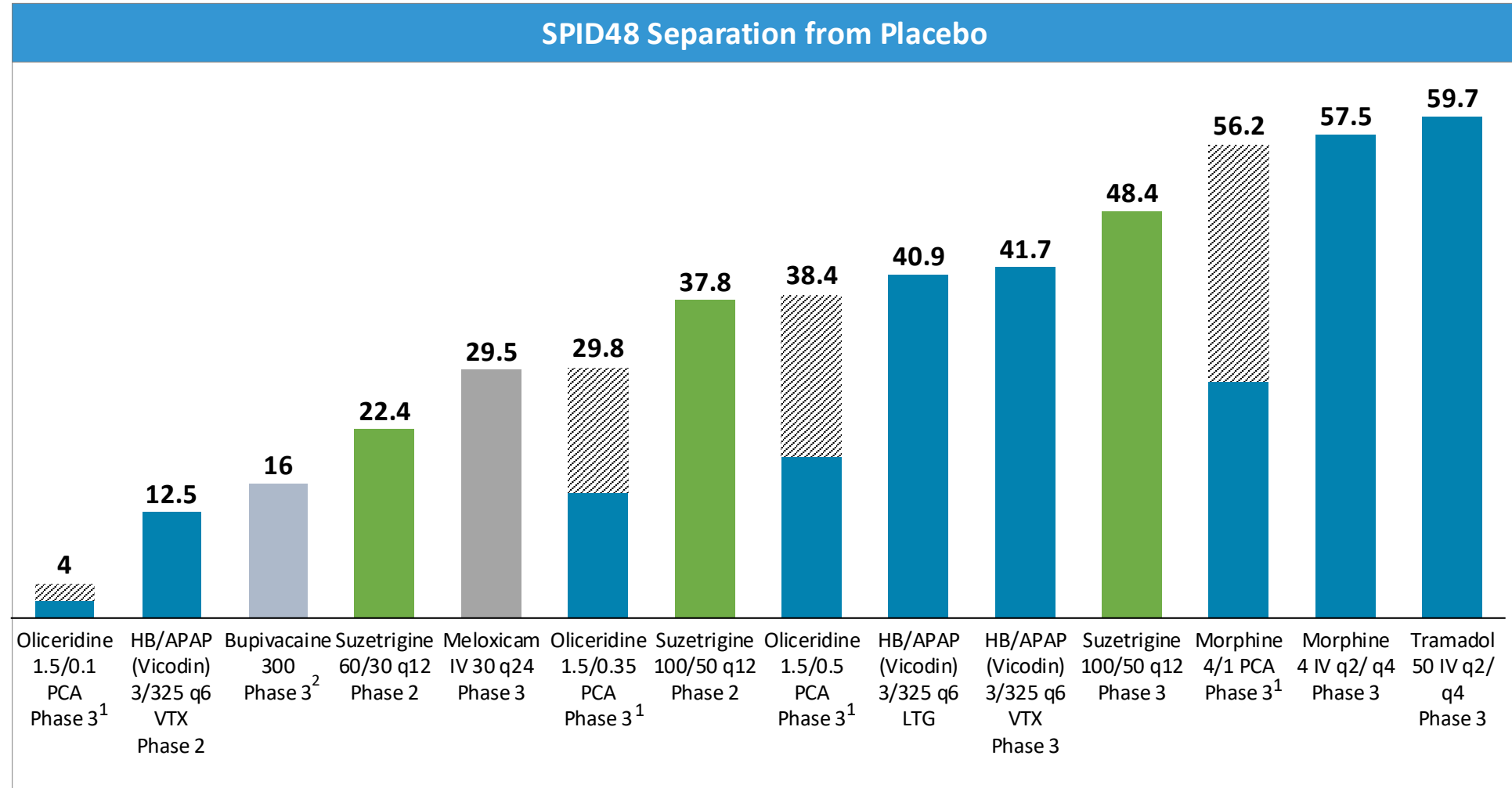
40.9

¹These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.



Onzotrigine had the Highest Analgesic Effect Seen Relative to Dose Arms in Separate Previously Reported Abdominoplasty Trials (7 Trials, 14 Dose Arms)

Onzotrigine
Acute Pain



- Opioid-level analgesia – Onzotrigine high dose: highest placebo separation over 48 hours
- Surpassed two historical studies using 4mg IV morphine
- Surpassed Vicodin comparator by 50%
- More than doubled NSAID
- Average placebo response: 123³

- NSAID
- Na_v1.8
- Opioid
- Local anesthetic

Note: All dosing in mg unless otherwise noted; ¹Estimated by doubling of SPID24 data for oliceridine dose arms and morphine 4mg IV bolus + 1mg IV PCA from oliceridine label ²bupivacaine hydrochloride implant: Reflects AUC48 ³The average pain reduction seen from the placebo arms from previously reported abdominoplasty trials that were conducted over 48 hours excluding onzotrigine.

Note: These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.



Onset Matters for Acute Pain and Onzotrigine’s Abdominoplasty Trial Data Suggest Rapid Onset for Na_v1.8

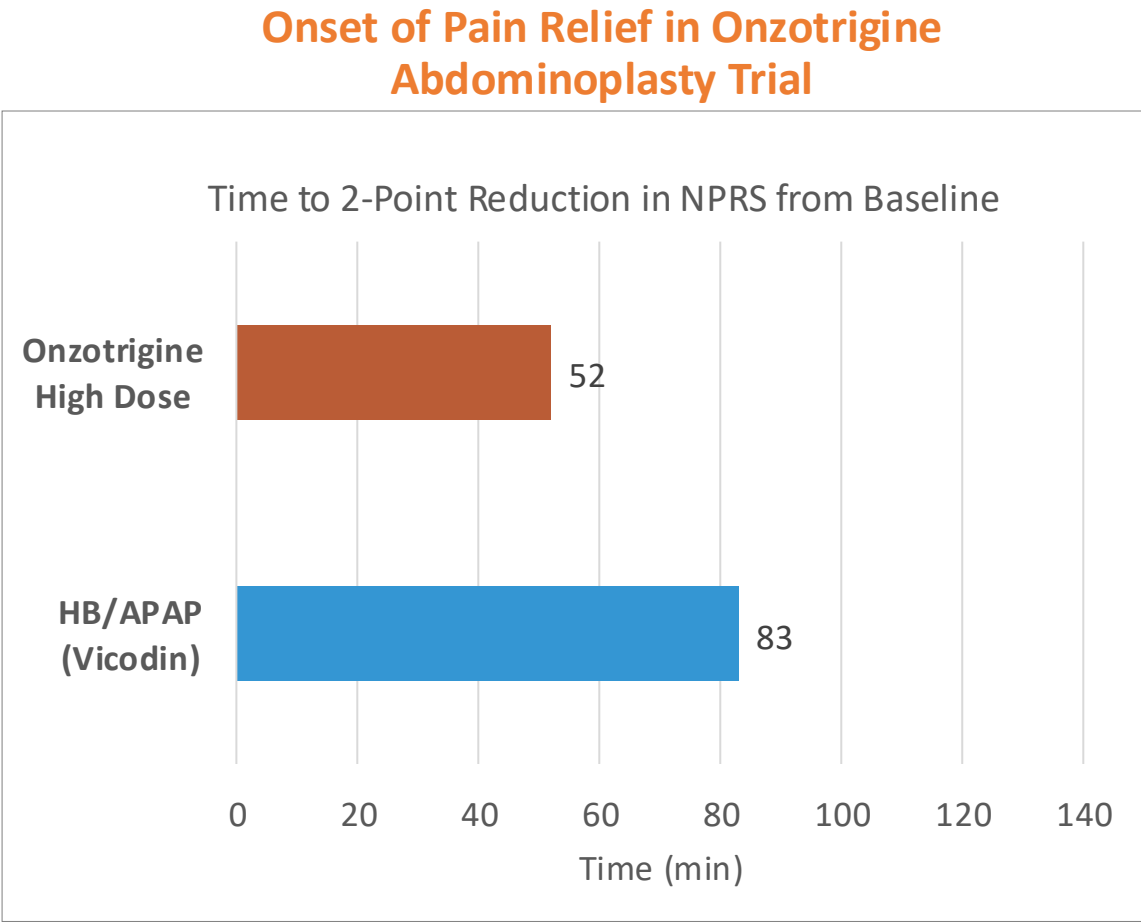
Why Onset Matters for Acute Pain

- Patients want immediate pain relief
- Clinicians and regulators worry about “drug stacking” – where patients take extra doses or other MoAs if they don’t feel early relief

Onset of Pain Relief Across Acute Pain Labels

Acute Pain Medicine (oral)	Onset in Label
Dsuvia (sufentanil)	54 min
Celebrex (celecoxib)	60 min
Ultracet (tramadol/APAP)	<1 hour
Zipsor (diclofenac)	<1 hour
Journavx (suzetrigine)	2-4 hours

Note: These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.



Onzotrigine Met Opioid-Free Endpoint FDA Has Used for Opioid Sparing Labels in Third-Party Drugs

Onzotrigine
Acute Pain

% of Opioid-Free Patients		P-value v. placebo
Onzotrigine High Dose	52.3	<0.001
Onzotrigine Low Dose	49.4	<0.001
Placebo	22.1	

There are only three therapies (infiltration analgesics) which include opioid free claims in their labels

Exparel (Bupivacaine Nerve Block or Instillation)	
Hemorrhoidectomy (72 hours)	
Active	28%
Placebo	10%
Shoulder Surgery (48 hours)	
Active	13%
Placebo	1%

Xaracoll (Bupivacaine Implant)	
Inguinal Hernia Repair 1 (72 hours)	
Active	36%
Placebo	22%
Inguinal Hernia Repair 2 (72 hours)	
Active	28%
Placebo	12%

Zynrelef (Bupivacaine + Meloxicam Instillation)	
Bunionectomy (72 hours)	
Active	29%
Placebo	2%
Inguinal Hernia (72 hours)	
Active	51%
Placebo	22%



Onzotrigine Was Generally Well Tolerated With Overall AE Levels Below Placebo

Onzotrigine
Acute Pain

- AEs were mostly mild to moderate
- Overall AEs lower in active than in placebo
- No individual AE was meaningfully more common in any active arm than in placebo, except pyrexia and presyncope, which were mild to moderate
 - Pyrexia: 5 of the 6 cases in the high dose arm were deemed unrelated to treatment
 - Presyncope: All cases were deemed unrelated to treatment
- Serious AEs for onzotrigine deemed unrelated to treatment

	High Dose	Low Dose	HB/APAP (commonly k/a Vicodin)	Placebo
Participants with any TEAEs, n (%)	43 (50.0)	53 (62.4)	54 (62.8)	56 (65.1)
Participants with any severe TEAE (Grade 3+), n (%)	3 (3.5)	3 (3.5)	6 (7.0)	3 (3.5)
Participants with serious AEs, n (%)	1 (1.2)	1 (1.2)	1 (1.2)	1 (1.2)
Participants with AEs leading to treatment discontinuation, n (%)	0	0	4 (4.7)	0
AEs reported in ≥3%* and at greater rate than placebo, n (%)				
Pyrexia	6 (7.0)	2 (2.4)	1 (1.2)	2 (2.3)
Presyncope	5 (5.8)	0	1 (1.2)	1 (1.2)

AE: adverse event; TEAE: treatment-emergent adverse event

*Note. 3% threshold represents occurrence in >2 patients



Higher Level of Na_v1.8 Target Inhibition in Nerve Supports Onzotrigine's Clinical Results

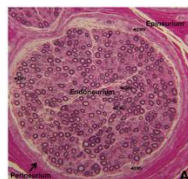
Onzotrigine
Acute Pain

Latigo Differentiated Capability to Look at Target Inhibition at Site of Action

1) Systemic Dosing of Na_v1.8 Inhibitors



2) Dissection of Sciatic Nerve at Various Times



3) Quantitate Drug in Nerve by Mass Spec



Onzotrigine Engaged More Na_v1.8 Channels at the Site of Action vs. Suzetrigine

	Onzotrigine Target Dose: 450 / 300 mg BID	Suzetrigine Target Dose: 100/50 mg BID
Drug at Nerve (Site of Action)	1790 nM	18 nM ¹
Free Fraction	21.6%	1%
In Vitro Potency Human DRG Neurons	104 nM	0.68 nM / 2.52 nM ²
Potency Shift w/assay to Mimic Nerve Environment [^]	2.7x	9.3x / 5.5x ²
Potency at Site of Action	281	10.2 ³
Predicted Target Inhibition % (Range for 2x Variation in Potency or Free Fraction)	86% (76-93)	64% (47-78) ¹

Note: These data are derived from different preclinical studies at different points in time, with differences in design. As a result, cross-study comparisons cannot be made, and no head to head preclinical studies were performed.

[^]Electrophysiology assay to determine compound potency conducted with 2% albumin to mimic protein level in endoneurial fluid surrounding axons expressing Na_v1.8 channels in vivo; Synthesized version of suzetrigine used;

¹ Incorporates parent & active metabolite. Drug at nerve reflects sum of parent & active metabolite; ² Parent and active metabolite respectively; ³ average potency shift of parent & metabolite



Planned Designs for Onzotrigine Phase 3 Bunionectomy and Open-Label Safety Trials

Onzotrigine
Acute Pain

	Phase 3 Trial for Onzotrigine in Moderate to Severe Acute Pain after Bunionectomy	Phase 3 Onzotrigine Safety Trial
Onzotrigine Regimen	Onzotrigine 450mg load, then 300mg BID (both trials)	
Study Population	Bunionectomy surgery (n = ~400)	Surgical & non-surgical (n = ~400)
Design	1:1 randomization, onzotrigine vs. placebo, double-blind, placebo-controlled, parallel group study	Open-label, onzotrigine as part of multi-modal regimen
Rescue	Oxycodone (same as prior abdominoplasty)	N/A
Primary Endpoint	SPID48: onzotrigine vs. placebo	Safety-related findings
Key Secondary Endpoints	Onset of action, opioid-sparing effect	% Opioid free, Patient & Clinician Global Assessment
Duration	48h treatment period	Up to 30 days treatment period

Initiation Planned for 2H 2026



Data Generated for Onzotrigine to Date Support Differentiated Target Product Profile to Approved Na_v 1.8 Inhibitor Suzetrigine

Onzotrigine
Acute Pain

	Suzetrigine's Approved Label	Onzotrigine Target Product Profile
Indication	Broad acute pain	Broad acute pain
Magnitude of Pain Relief	SPID48 Δ vs. placebo = 48 (Abdominoplasty)	SPID48 Δ vs. placebo = 62 (Abdominoplasty)
Onset of Action	119 mins (Abdominoplasty)	52 mins (Abdominoplasty)
Opioid Sparing	-	Potential for Opioid-free claim with 52% Opioid-free data in Abdominoplasty
Safety Profile	No SAEs Pruritus, muscle spasms, CPK elevation, rash	Comparable or lower AE incidence vs. suzetrigine (Abdominoplasty)
Duration of use	No duration restriction Drug not studied for >14 days	No duration restriction Drug not studied for >30 days*
Contraindications	- Avoid use with strong and moderate CYP3A inducers - Contraindicated with strong CYP3A4 inhibitors Dose reduction with moderate - Additional contraceptive measures	None based on data generated to date

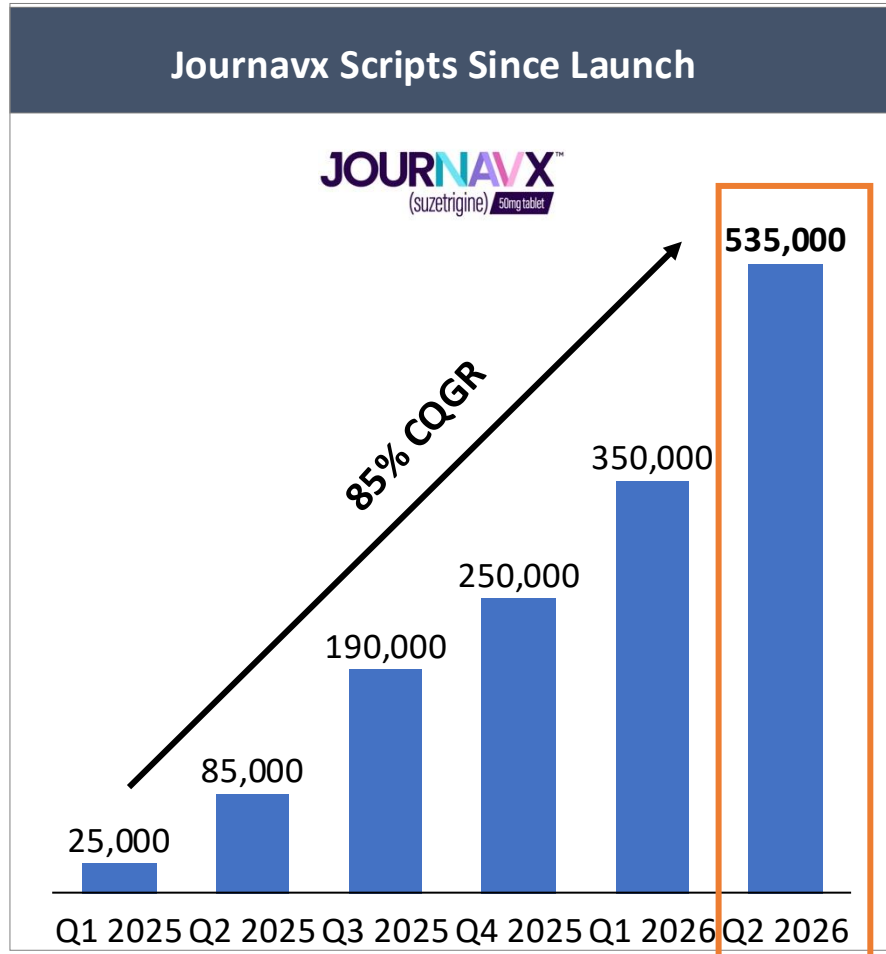
**possible based on planned safety study design*

Note: These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.



Strong Early Launch Momentum with Rapid Patient Demand for Journavx

Onzotrigine
Acute Pain



Clear Patient Demand for Journavx with Accelerating Growth

- Recent quarter scripts of 535K nearly matched all script volume from first year of launch
- Q2 '26 volume equates to ~>\$600MM¹ in gross revenue on a run rate

Market access has been improving

- 260M lives covered, formally contracted with all 3 national PBMs
- 22 state Medicaid programs provide coverage
- 3 of 4 Medicare Part D plans recently covered
- Recently eligible for separate, non-DRG payment for Medicare outpatient procedures (NOPAIN Act)

High GTN limiting initial revenues due to Patient Support Program (PSP)

- \$79M revenue in 1H 2026 vs. \$60M for full year 2025
 - 71% QoQ sequential growth
- PSP should be discontinued by 2027 as access improves
- Vertex guiding to >3x prescription and revenue growth in 2026 vs. 2025 and GTN normalizing in the 1H2027

Source: Vertex Pharmaceuticals Q4 2024, Q3 & Q4 2025, Q1 & Q2 2026 earnings presentations and earnings call transcript. 2026 JPM press release.

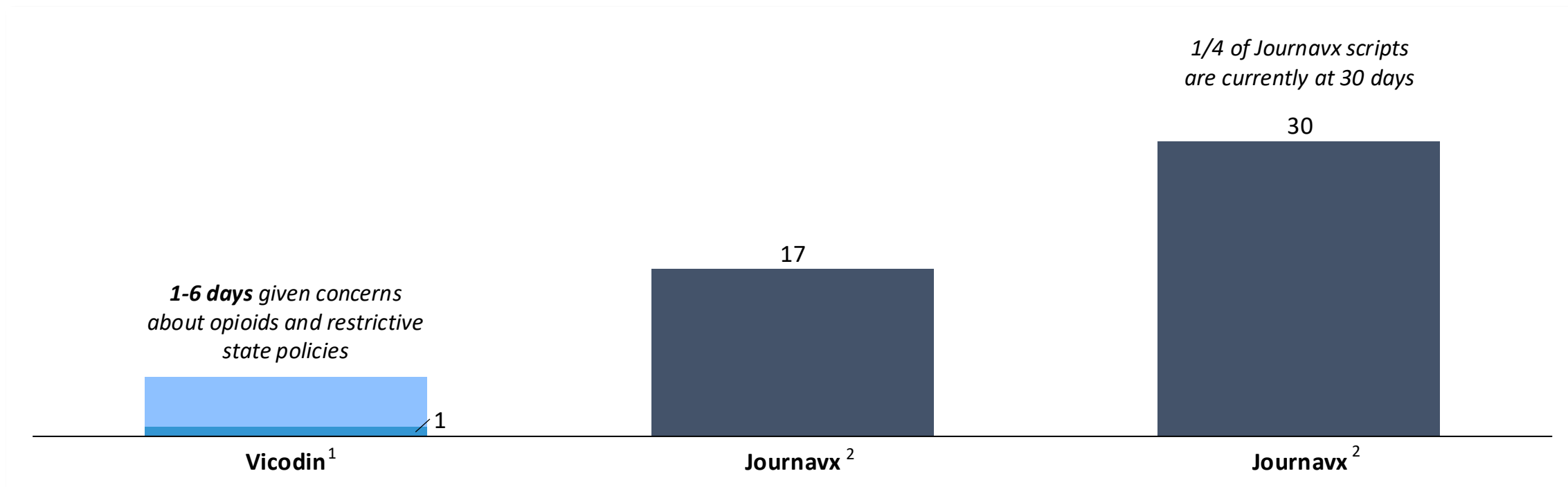
Assumes 535,000 scripts per quarter, average prescription length of 10 days at \$31 / day



Journavx's Retail Scripts Are at an Average of 17 Days with 1/4 at 30 Days

Onzotrigine
Acute Pain

Retail Days of Therapy (Average Treatment Duration in Days)



Average script length longest amongst anesthesiologists/pain specialists & rheumatologists –specialties typically managing chronic pain

¹ Vicodin prescription days: Tran et al., J Manag Care Spec Pharm. 2017 May;23(5):532-539; ² IQVIA Claims data of retail scripts from February 2025 to January 2026. Reflects volume weighted average.



Market Research Shows Substantial Majority of HCPs Prefer Onzotrigine Target Product Profile (TPP) – Driven By Onset

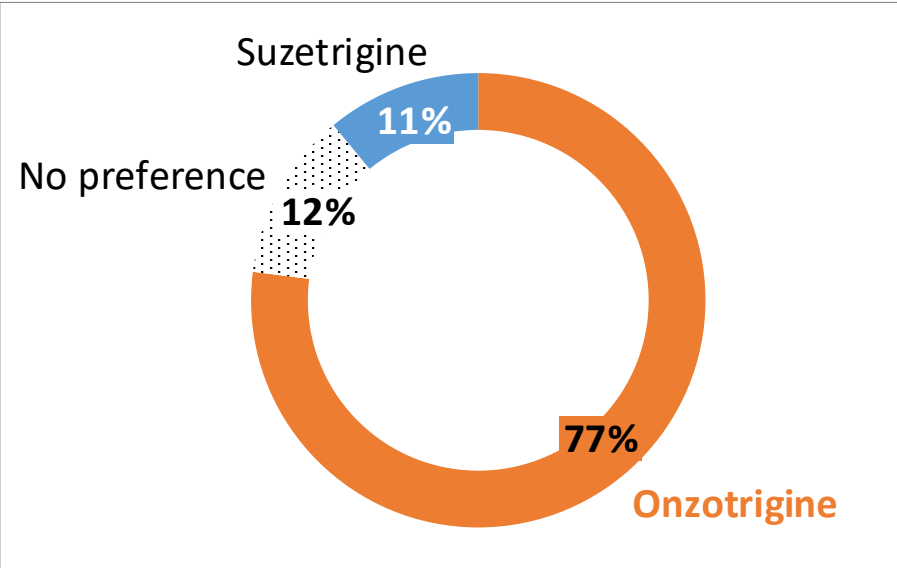
Onzotrigine
Acute Pain

Blinded Target Product Profiles Tested

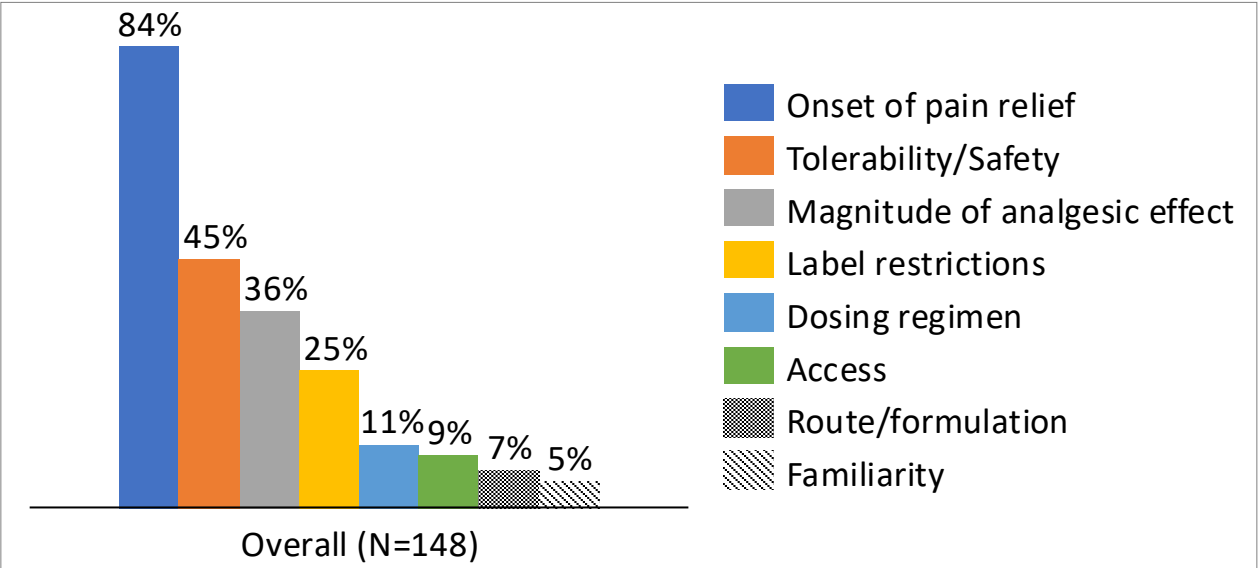
		Suzetrigine (on market)	Onzotrigine TPP (estimated profile prior to abdominoplasty trial results)
Analgesic Effect	SPID48 vs. PBO	29-48	36-57
	Onset	2-4 hours	1 hour

Actual results in abdominoplasty were *better* than what we tested

Product Preference



Factors Driving Preference for Onzotrigine



Q15.Based on the profiles provided, which product would you be more likely to prescribe overall?

Q16.What are the most important factors influencing your preference between Product A and Product B? (Select all that apply) | HCPs that stated preference for Product B. N=148

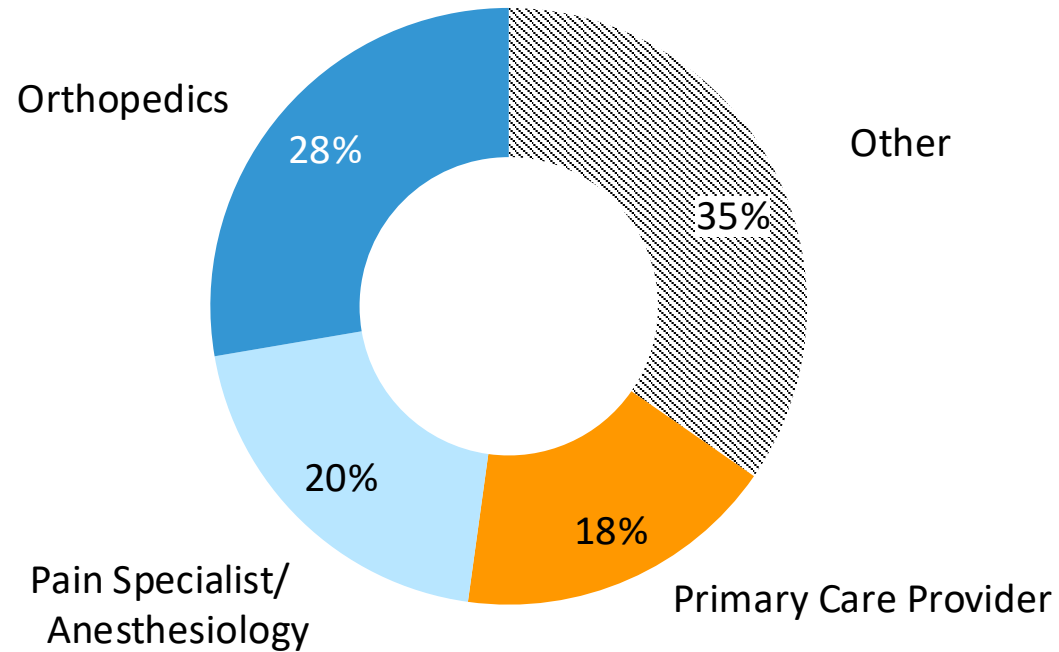
Source: Market survey conducted by Latigo in 2025



Orthopedics and Pain Specialists Drive Primary Retail Script Volume for Journavx

Onzotrigine
Acute Pain

Share of Rx by Specialty¹ (% of Total Prescriptions)



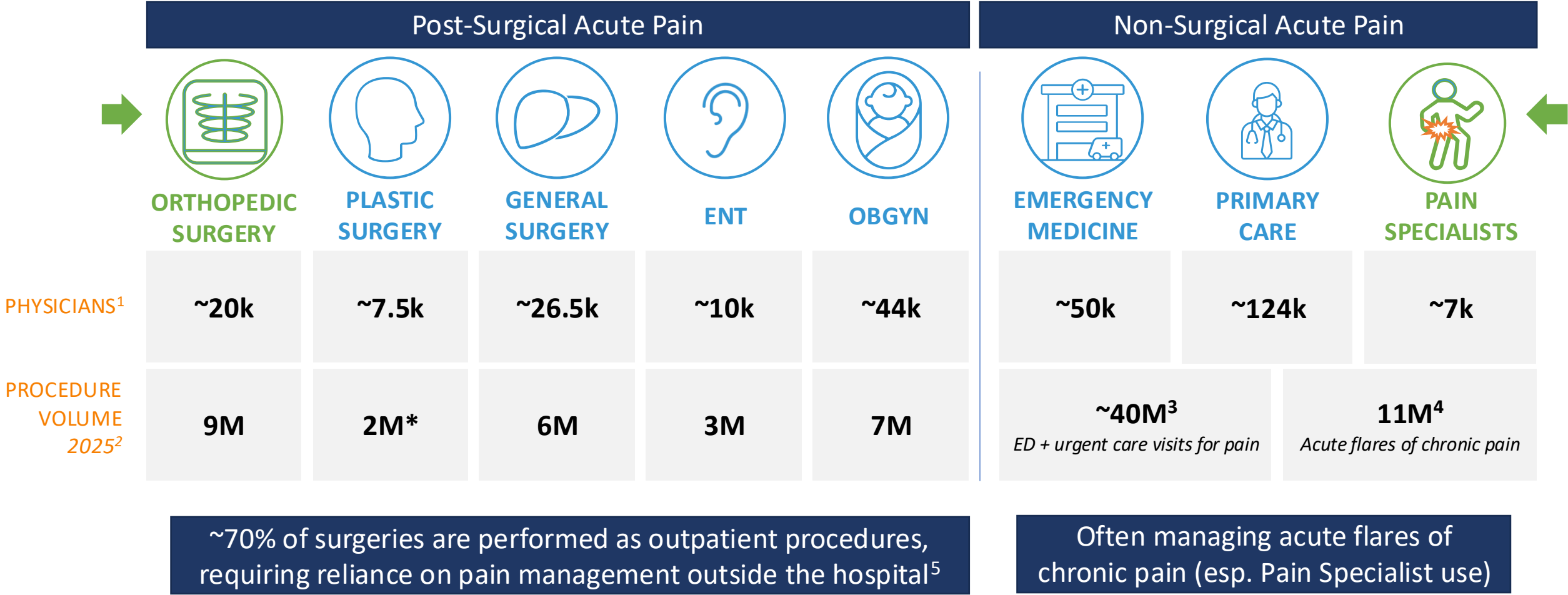
Significant primary care volume indicates organic growth driven by unmet need, outside of Vertex Pharmaceutical's promotional focus on institutional accounts

Note: excludes RN/NP/PA. Other includes physical therapy, plastic surgery, podiatry, neurology, general surgery, dentist, etc. ¹ IQVIA Claims data of retail scripts from February 2025 to January 2026. Reflects volume weighted average.



Orthopedic Surgeons & Pain Specialists Are Key Call Points (And Largest Volume of Journavx Scripts Today Along with PCPs)

Onzotrigine
Acute Pain

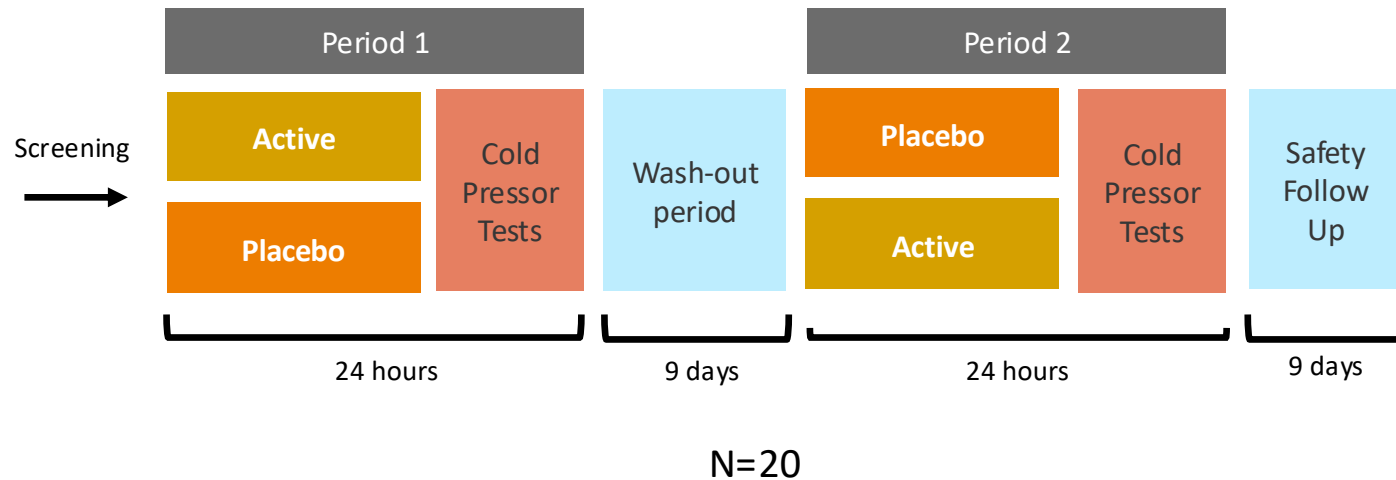


¹AMA physician workforce database. ²Life science intelligence 2024; ³CDC 2022 and FAIR Health White Paper, 2025; ⁴Todd et al., West J Emerg Med. 2010 Dec;11(5):408-15.; ⁵ Ambulatory Surgery Center Association, 2023. *Excludes “skin lesion procedures” which require minimal post-procedure pain management



Latigo Data Support Cold Pressor Test (CPT) as a Reliable Pharmacodynamic Biomarker for Drug Development Decision Making

Randomized, Double-blind, Placebo-controlled, Cross-Over Study Cold-Pressor Test



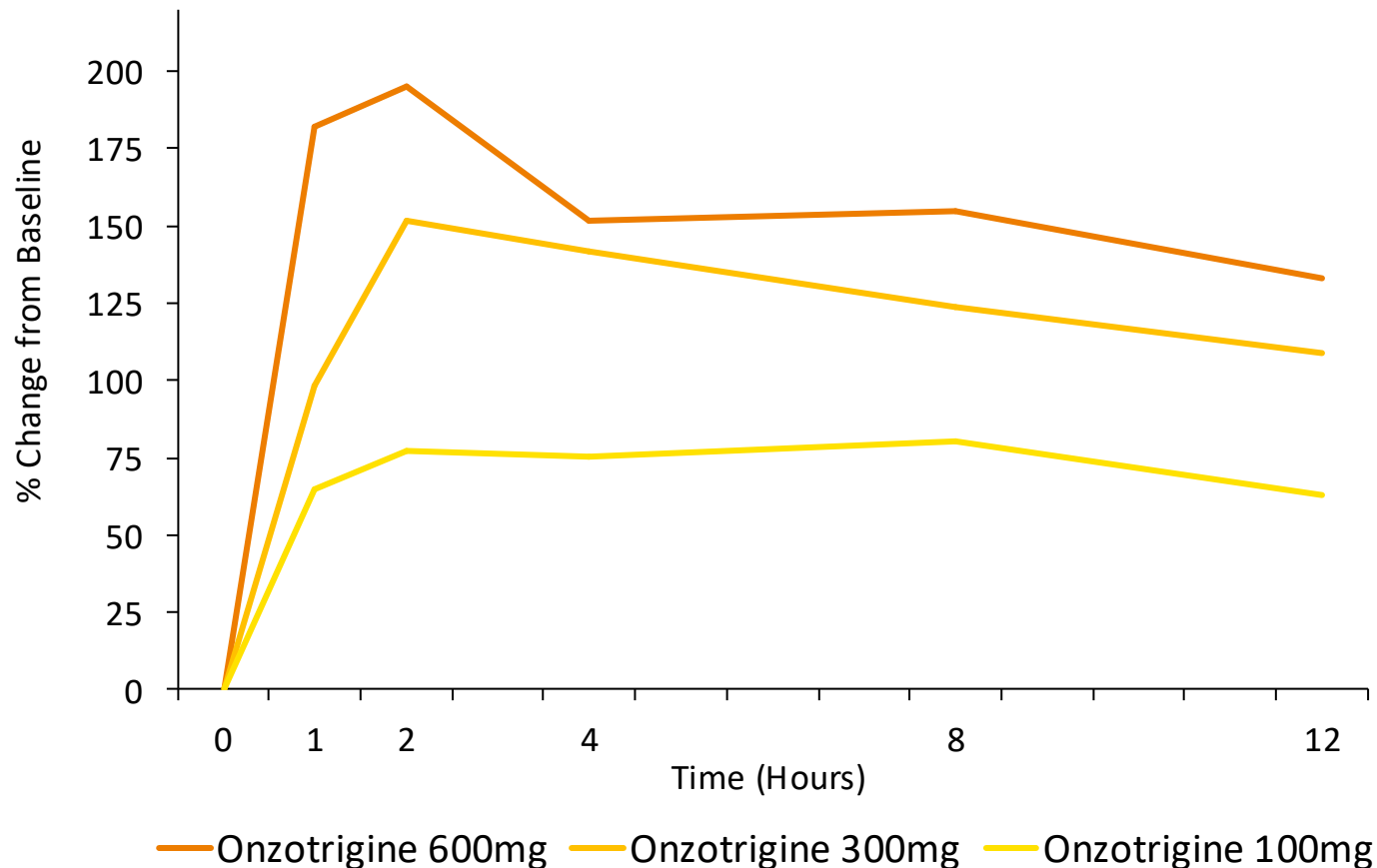
Study has been run with onzotrigine (100mg, 300mg, 600mg), LTG-305 (480mg 1 - and 5- day dosing), LTG-321 (90mg, 270mg), and suzetrigine (100mg)

Cold Pressor Test



CPT Has Predicted Analgesic Effects in a Later-Stage Clinical Trial

Cold Pressor Mean PTT % Change from Baseline, Placebo-Subtracted



PTT: Pain tolerance threshold

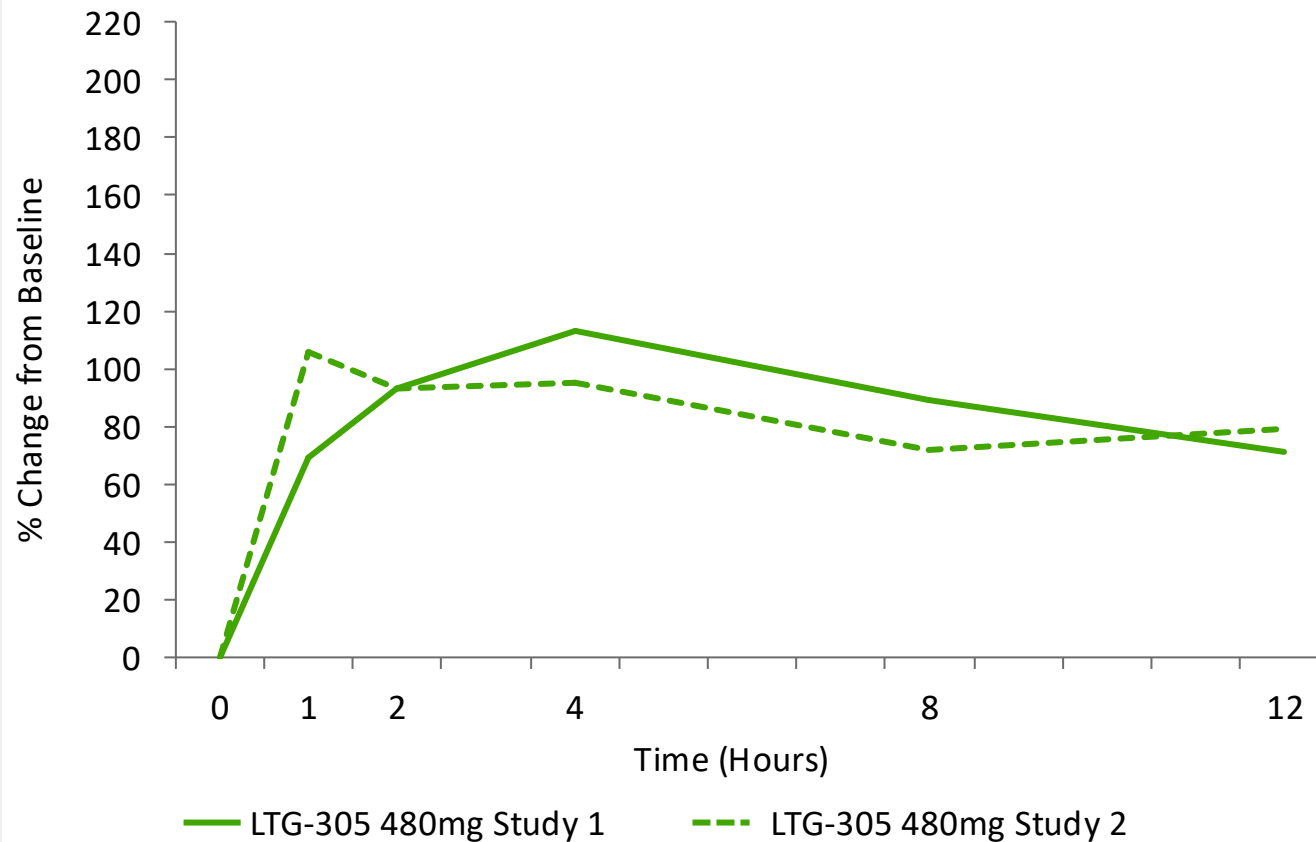
CPT as an Archetypal Translational Model

- **Dose Ranging:** The two drugs we tested with multiple doses onzotrigine (shown on left) and LTG-321 both showed dose ranging with separation across each dose level



CPT Has Predicted Analgesic Effects in a Later-Stage Clinical Trial (Cont.'d)

Mean PTT % Change from Baseline, Placebo-Subtracted



PTT: Pain tolerance threshold

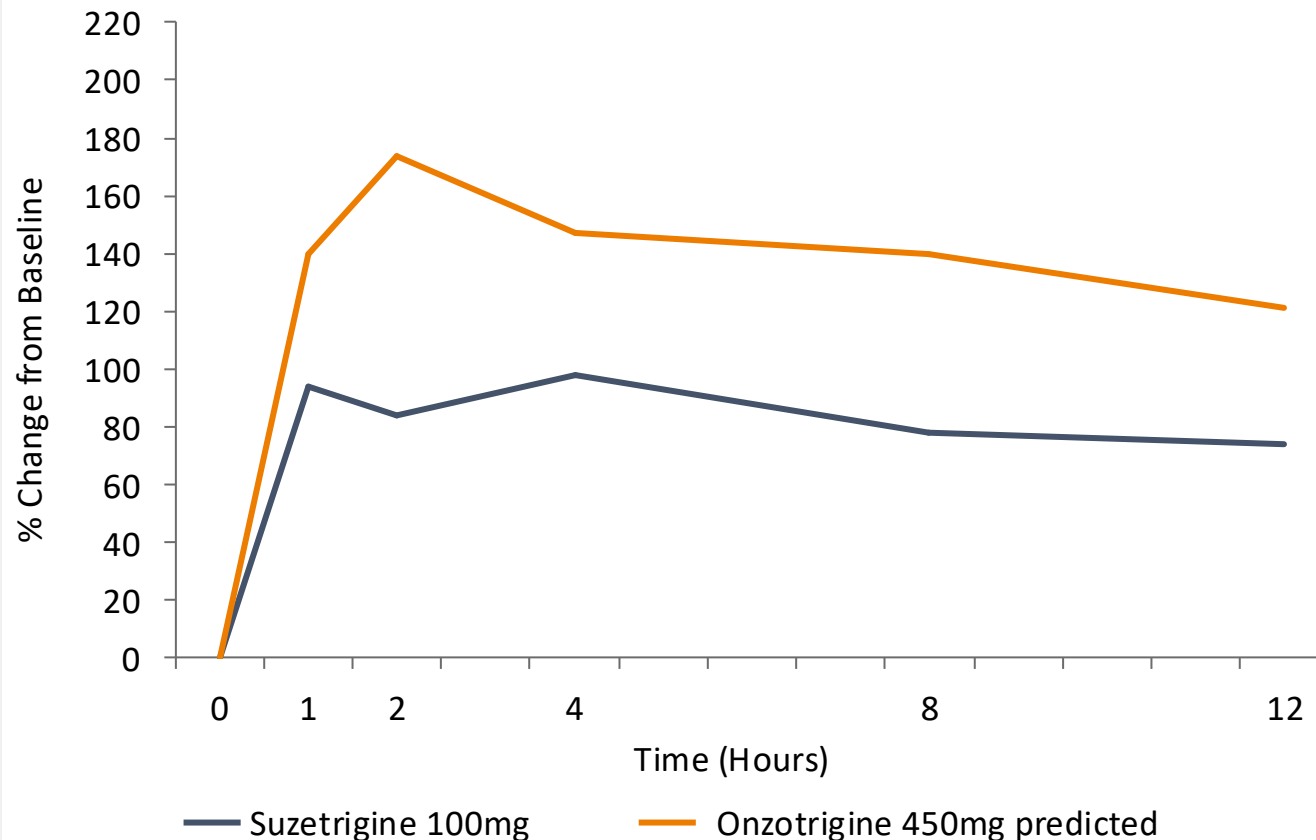
CPT as an Archetypal Translational Model

- **Repeatable:** We repeated the CPT with LTG-305 and got nearly consistent results



CPT Has Predicted Analgesic Effects in a Later-Stage Clinical Trial (Cont.'d)

Mean PTT % Change from Baseline, Placebo-Subtracted



CPT as an Archetypal Translational Model

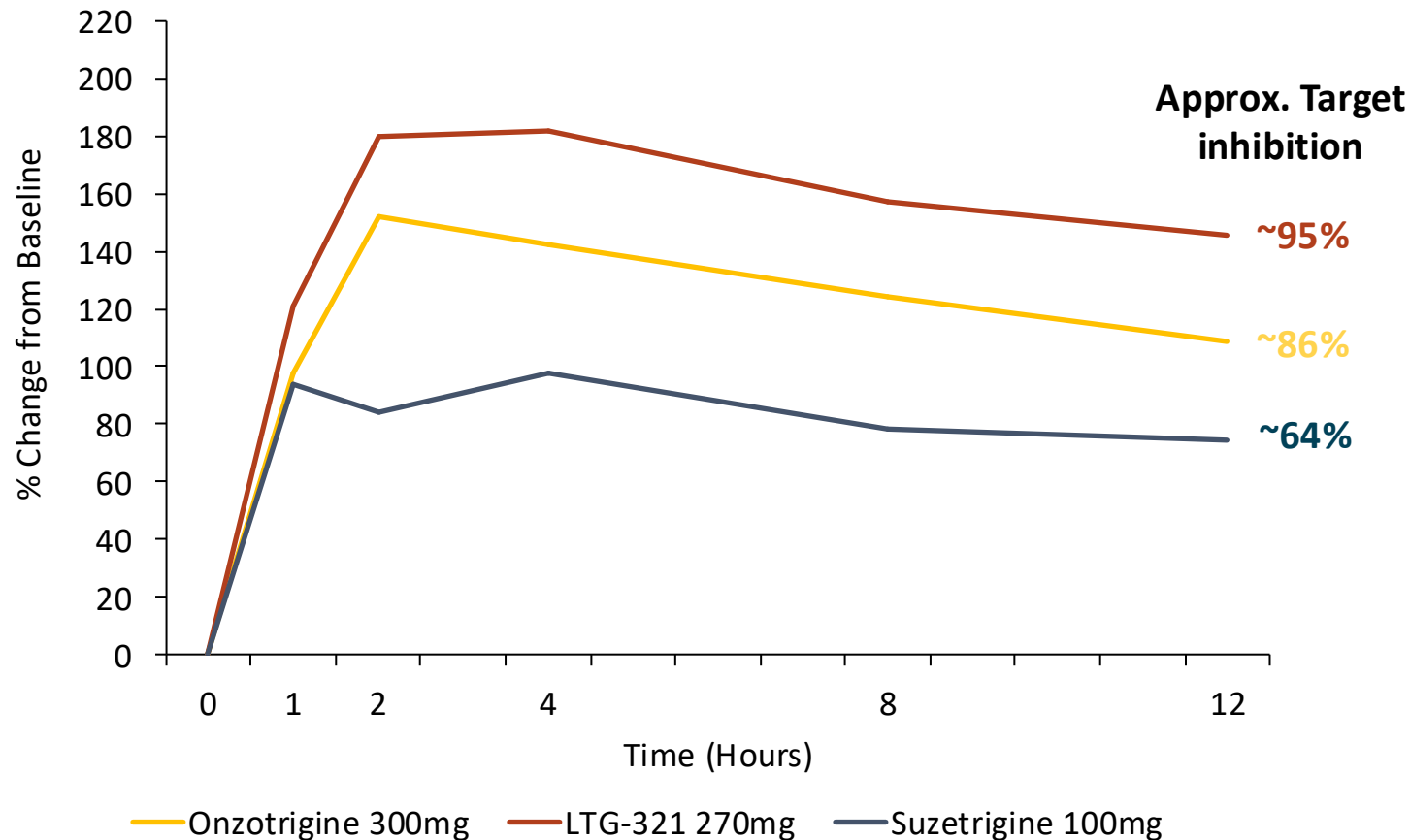
- **Predictive:** CPT predicted analgesic effect for a single dose in abdominoplasty
 - CPT data – Ran 100mg commercially-available suzetrigine in same CPT trial (450mg of onzotrigine (predicted) demonstrated ~70% higher AUC than 100mg of suzetrigine)
 - SPID12 of abdominoplasty (measuring impact of only the first dose) – 450mg of onzotrigine was 80% higher than 100mg of suzetrigine

Note: These data are derived from different clinical trials at different points in time, with differences in patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.



LTG-321: Highest Known CPT Data for Any Na_v1.8

Mean PTT % Change from Baseline, Placebo-Subtracted



Results

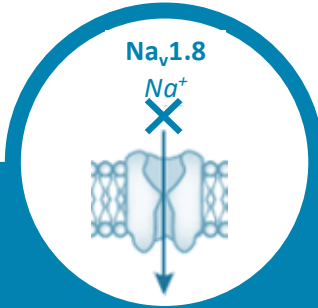
LTG-321 analgesic effect at 270 mg dose, surpassed onzotrigine 300 mg by 26%¹

Note: These data are derived from different clinical trials at different points in time, with differences in trial design and patient populations. As a result, cross-trial comparisons cannot be made, and no head-to-head clinical trials have been conducted.

¹ Reflects total aggregate effect or AUC over 12 hours



Why Proof of Concept in Osteoarthritis (OA) for Chronic Pain?



Preclinical Evidence of Potential Mechanism
of Action in Musculoskeletal Pain



Mechanism of Action Validated in OA



Most Reliable Chronic Pain Model

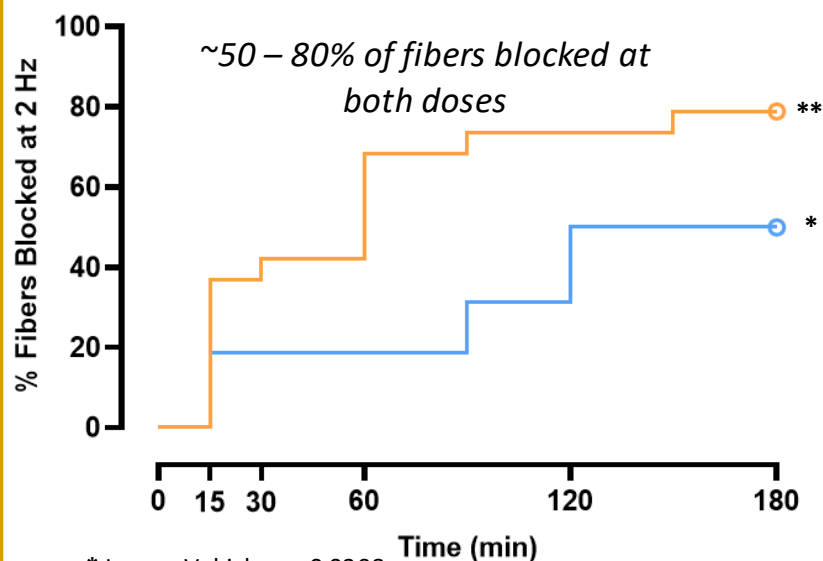


Latigo's Expertise

Lower Na_v 1.8 Target Inhibition Needed for Action Potential (AP) Block in Preclinical Musculoskeletal Pain Models

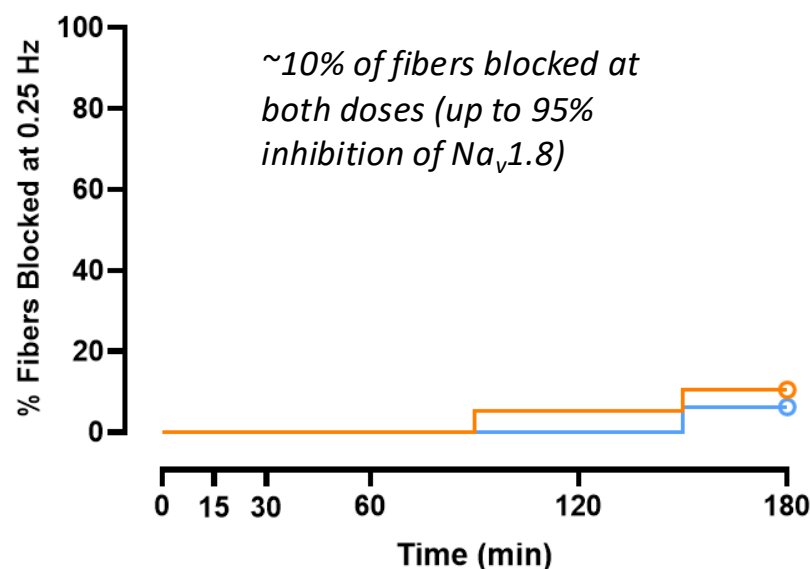
LTG-321
Chronic Pain

High Frequency – 2 Hz



Firing Frequency Mimicking
Musculoskeletal Pain

Low Frequency – 0.2 Hz



Firing Frequency Mimicking
Neuropathic Pain

>99% inhibition of Na_v 1.8 required to meaningfully reduce low frequency AP firing in neuropathic pain model vs. **>70% inhibition** to reduce higher frequency AP firing musculoskeletal pain model

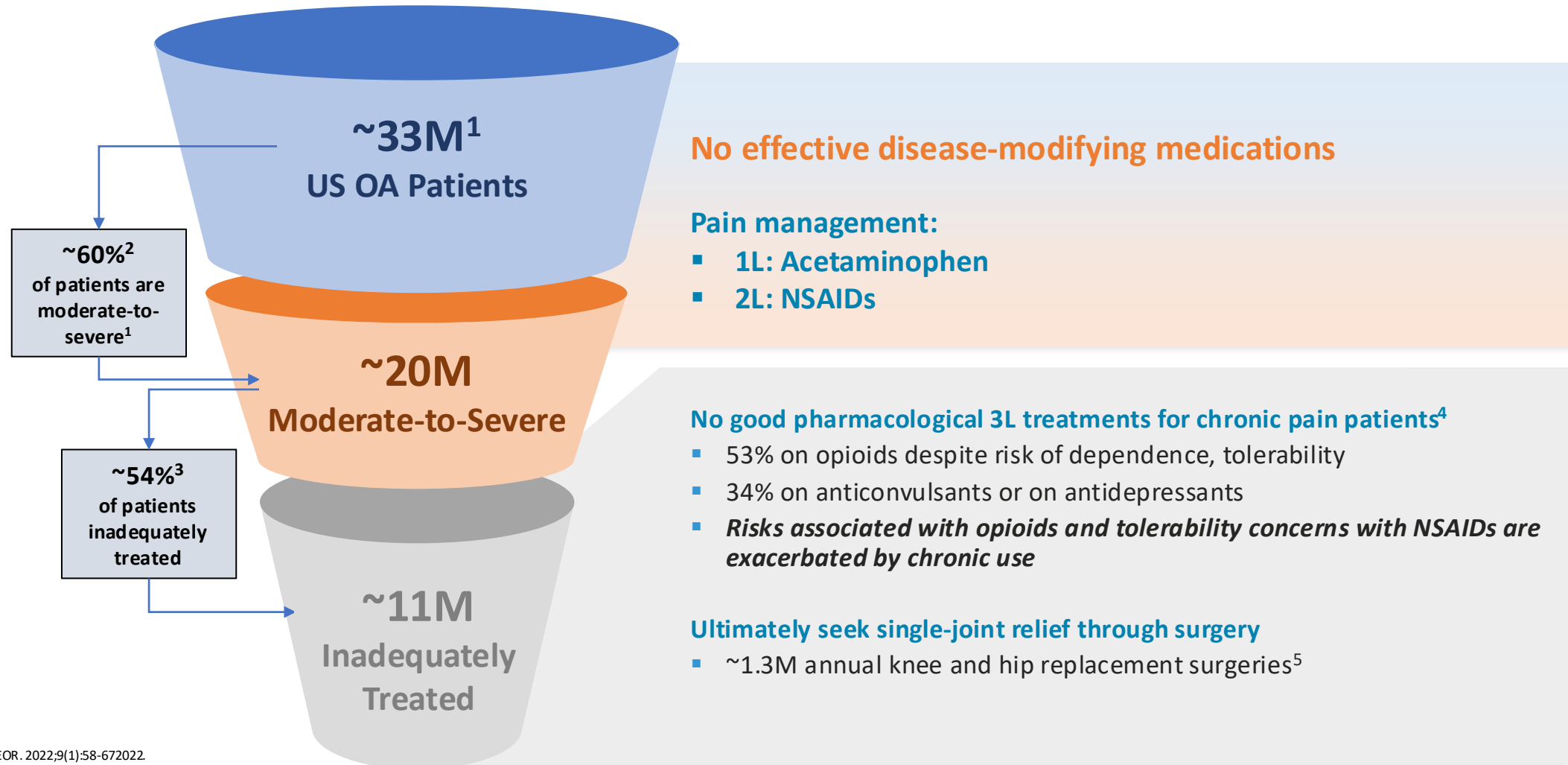
Target Inhibition Achieved with Na_v 1.8 Alone
Informs Musculoskeletal Pain as Initial
Prioritization for Chronic Pain Development

- Onzotrigine (70-80% Na_v1.8 Inhibition, n=16 fibers)
2.8 mg/kg bolus + 0.67 mg/kg/hr infusion i.v.
- Onzotrigine (90-95% Na_v1.8 Inhibition, n=19 fibers)
8.4 mg/kg bolus + 2 mg/kg/hr infusion i.v.



Limited Treatment Options for Patients with Osteoarthritis Pain Today, Driving Patients Towards Chronic Opioid Use Despite Risks

LTG-321
Chronic Pain



¹2020 CDC data

²Schepman et al., JHEOR. 2022;9(1):58-672022.

³Conaghan et al., Rheumatology. 2015;54(2):270-277.

⁴2025 Komodo claims data analysis

⁵American College of Rheumatology, March 2025

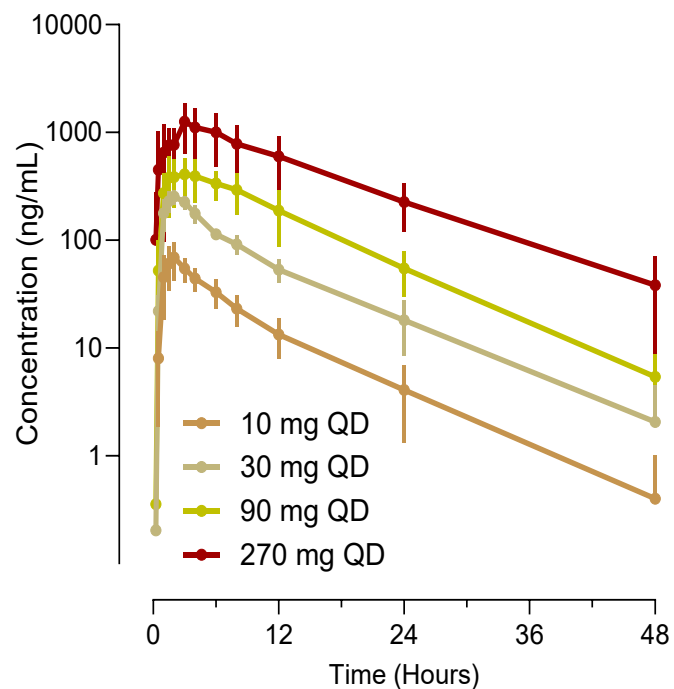


LTG-321: Dose Proportional Exposure Across All Doses in Phase 1 SAD / MAD

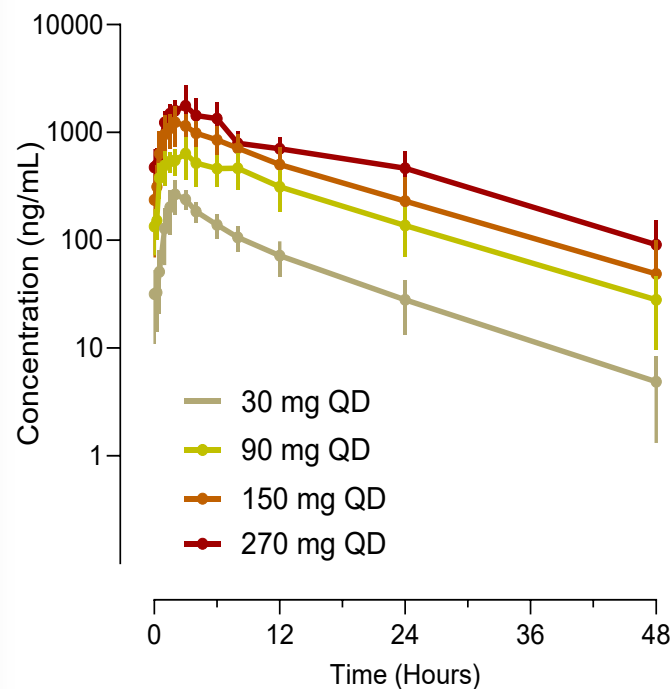
LTG-321
Chronic Pain

Dose-proportional Exposure Observed Across All Doses Tested in Phase 1 SAD / MAD Trial¹

LTG-321 Single Ascending Dose (SAD)
Log Scale



LTG-321 Multiple Ascending Dose (MAD)
Day 14 - Log Scale



Results

- Phase 1 SAD/MAD¹: LTG-321 well tolerated in healthy participants across cohorts
- Explored in SAD cohorts 10mg up to 270mg
- MAD cohorts explored 30mg up to 270mg once daily for 14 days
- Dose proportional exposure observed across all doses tested in participants across all cohorts
- ~33x 28-day NHP tox safety margin at 150mg QD dose

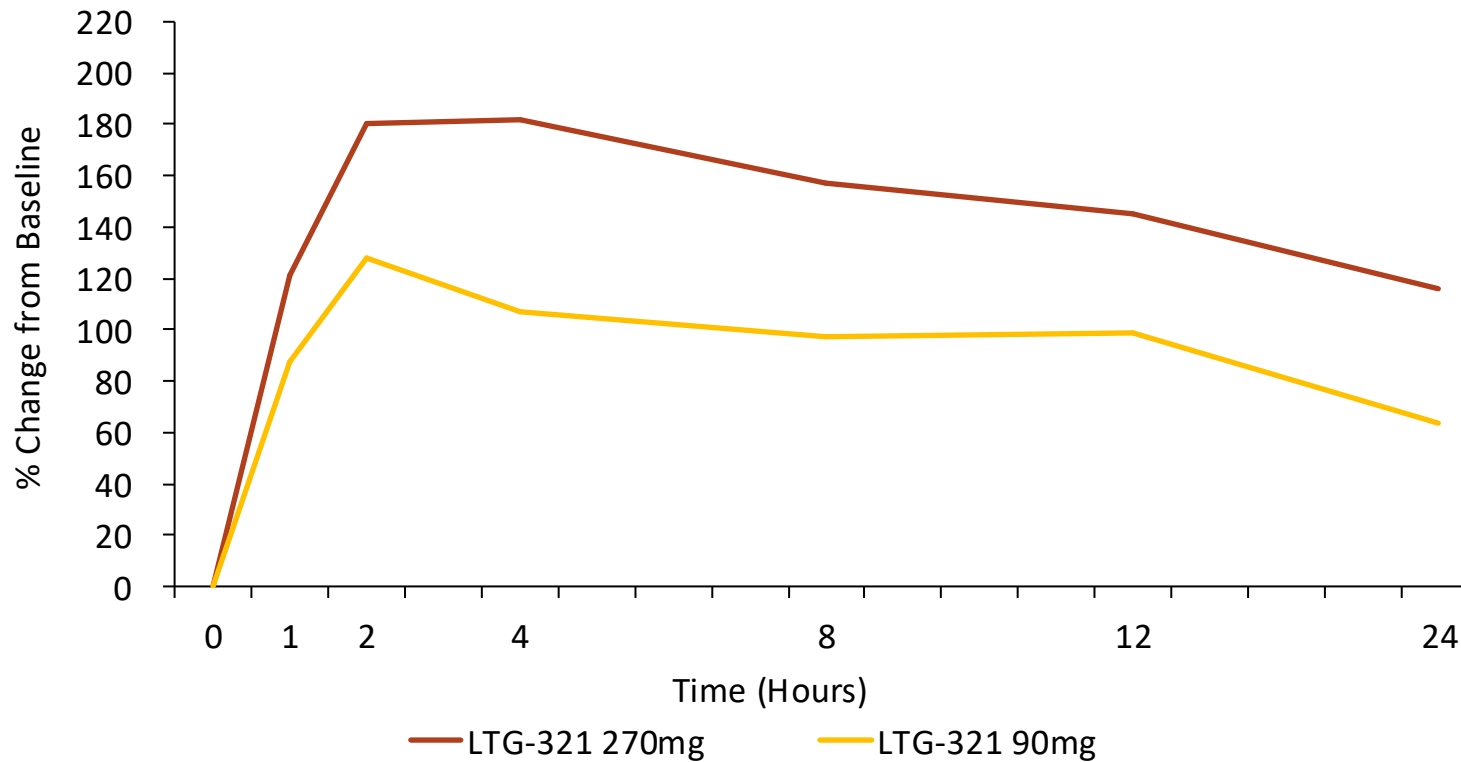
¹ As of May 2026 data cut off

Note: SAD: Single ascending dose; MAD: Multiple ascending dose; PK: Pharmacokinetic



LTG-321: Highest Known CPT data for Any Na_v1.8

Cold Pressor Mean PTT % Change from Baseline, Placebo-Subtracted



Results / Phase 2 Trial Dose Selection

- Both doses tested showed continued activity out to 24 hours
- LTG-321 analgesic effect at 270 mg dose, surpassed onzotrigine 300 mg by 26%¹
- 150 mg repeat dosing exposure was comparable to 270 mg single dose
- Will target **150mg QD for Phase 2 OA trial**

LTG-321 Dose / Cohort	C _{max} (ng/mL)	AUC ₀₋₂₄ (h*ng/mL)
270 mg CPT	1050	12400
150 mg MAD Day 14	1330	14100

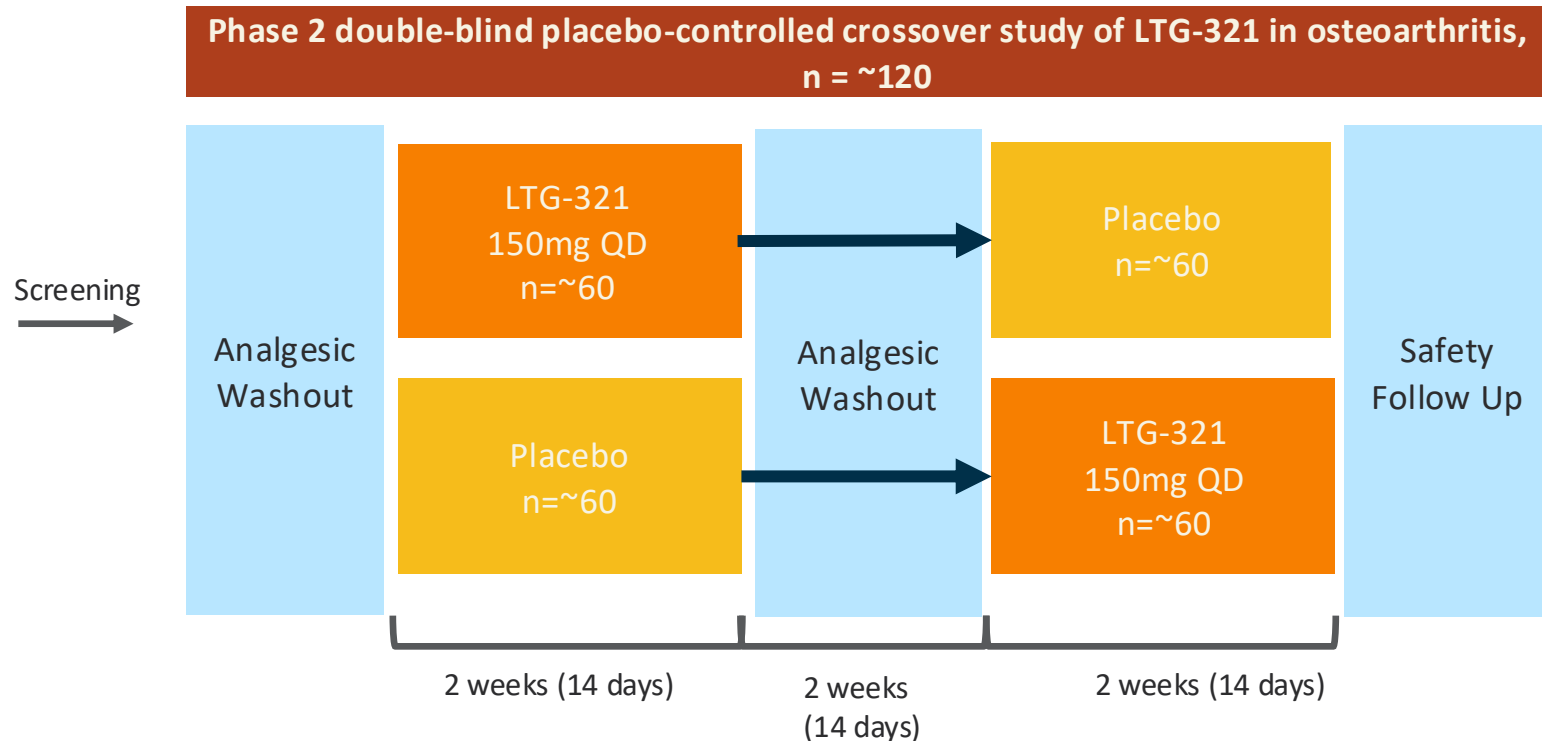
As of May 2026 data cut off

¹Reflects total aggregate effect or AUC over 12 hours



LTG-321 Chronic: Initiated Phase 2 Within-Subject Crossover Trial in Osteoarthritis (OA) of the Knee

LTG-321
Chronic Pain



Design

- Crossover design intended to reduce variability

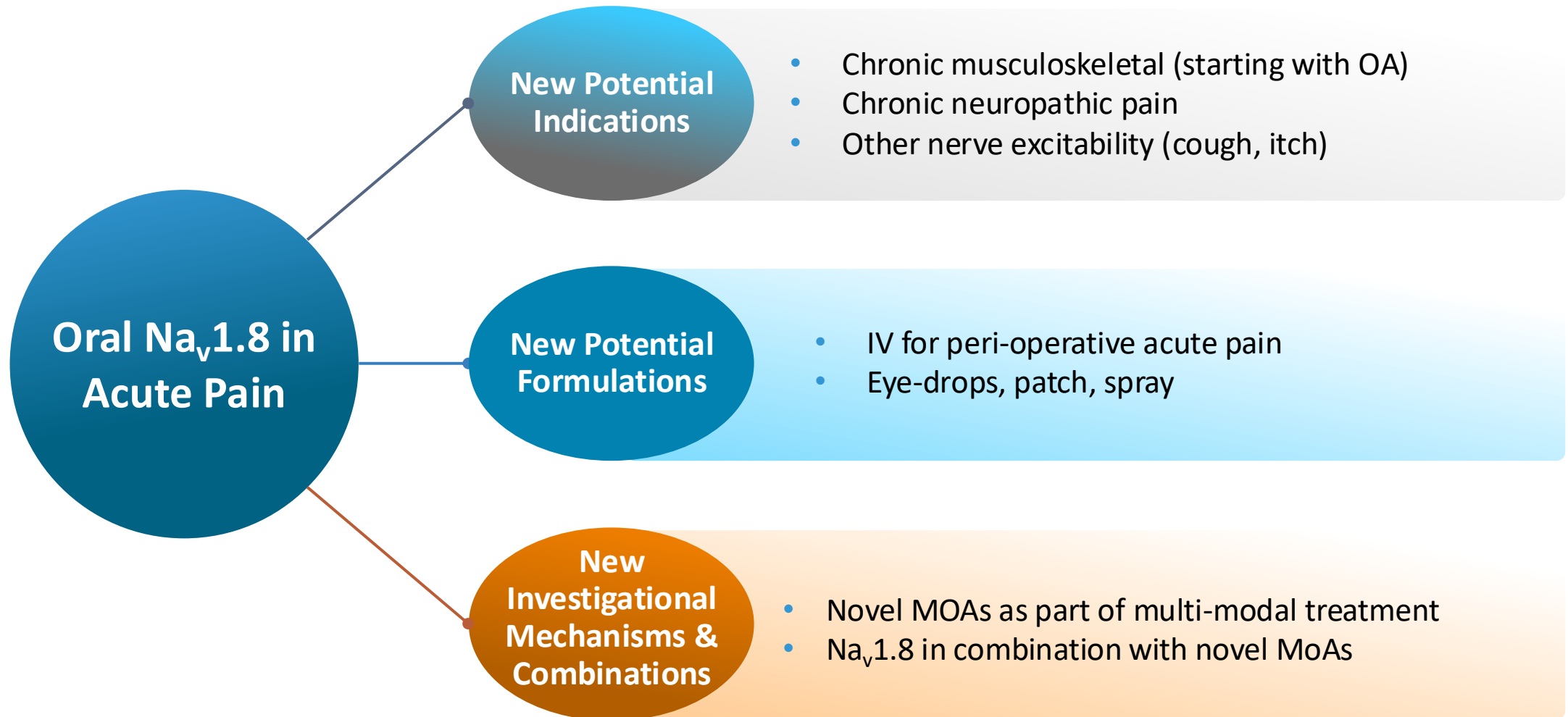
Primary Endpoint

- Change in WOMAC pain subscale from Baseline to 2 Weeks, LTG-321 vs. placebo

Rescue

- Acetaminophen (500mg every 4 to 6 hours up to max 3,000mg per day)

Oral Na_v1.8 in Acute Pain is the First Step in Life Cycle Management to Expand Portfolio



Na_v1.8 is Potentially the Ideal Foundational Therapy Target for a Multi-modal Pain Franchise

- Well tolerated for broad patient population based on suzetrigine approved label and data from clinical trials of other Na_v1.8 product candidates
- Can potentially be used in combination with other mechanisms

Novel MOAs involved in peripheral transmission

+ K_v7.2/7.3

+ Na_v1.9

+ Na_v1.6

+ Ca_v3.2

+ Na_v1.7

+ Other MOA

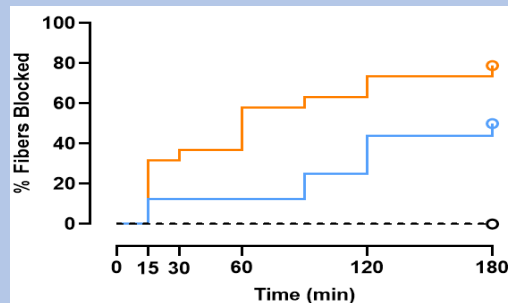
Na_v1.8

Execution of Best-in-Class Na_v1.8 Multi-Modal Pain Strategy – Latigo's Differentiation

Differentiated Pain Capabilities at Latigo

Pain Biology Expertise

- Internal e-phys models to validate targets and assess compounds
- Microneurography *in vivo* model – models drug activity without relying on behavior



in vivo & ex vivo E-phys

Clinical Translation

- Rapid, inexpensive proof of concept and early dose-finding in healthy volunteers with induced pain studies
- Accurate and has predicted analgesic effect in a late-stage trial



Cold Pressor Test

Clinical Execution

- Team has run over 250 pain studies
- Conducted full dose range finding abdominoplasty trial that read out in 6 months with potential best-in-class execution metrics



Abdominoplasty Trial

Anticipated Milestones for Onzotrigine and LTG-321

**Onzotrigine
(Acute Pain)**

2H2026: Initiate Phase 3
Bunionectomy & Safety
Trials

2H2027: Phase 3
Bunionectomy & Safety
Trials Topline Results

2026

2027

**LTG-321
(Chronic Pain)**

2H2027: Phase 2 OA Proof
of Concept
Topline Results



Thank You



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