

SAVI Dual™ Migraine Therapy *from eNeura*

— for acute and preventive treatment

- FDA cleared, prescription non-drug therapy for patients 12 and older
- Similar efficacy to prescription medication
- Only non-invasive, Central Neuromodulation therapy for migraine
- No medication risks or side effects
- SAVI Smart Diary™ automatically records treatment history

www.eneura.com

SAVI Dual™ Migraine Therapy

Migraine relief is as easy as 1-2-3

- **Easy** patient-administered
- **Effective** non-drug option
- **Superior tolerability**
- **Quick** less than a minute
- **Personalized** treat as needed
- **SAVI Smart Diary™** automatically records treatment history



1

Power

Press center button to turn on device

2

Position

Cradle SAVI Dual to the back of the head

3

Pulse

Press treatment button to deliver a pulse

Repeat as Prescribed



Watch the SAVI Dual “How to” video at www.eneura.com

How SAVI Dual™ works

Quieting The Migraine Storm

Non-invasive, **Central Neuromodulation** therapy developed to **treat acute attacks** and **prevent future migraine days**



Source: Andreou AP, Holland PR, Akerman S, et al. Transcranial magnetic stimulation and potential cortical and trigeminothalamic mechanisms in migraine. Brain 2016; 139: 2002–2014.

- Electromagnetic fields pass painlessly through bone and tissue
- Non-Invasive pulses signal the brain to calm hyperactive nerves and return to a more natural state
- Safety well established –science used in MRI’s for decades
- Single pulse delivered in less than a second
- Often referred to as Single Pulse Transcranial Magnetic Stimulation (STMS)
- Pulse strength about half that of typical MRI



Watch the SAVI Dual “Quieting the Migraine Storm” video at www.eneura.com

SAVI Dual™ for acute treatment of migraine

Well-established efficacy and tolerability

Proven effective

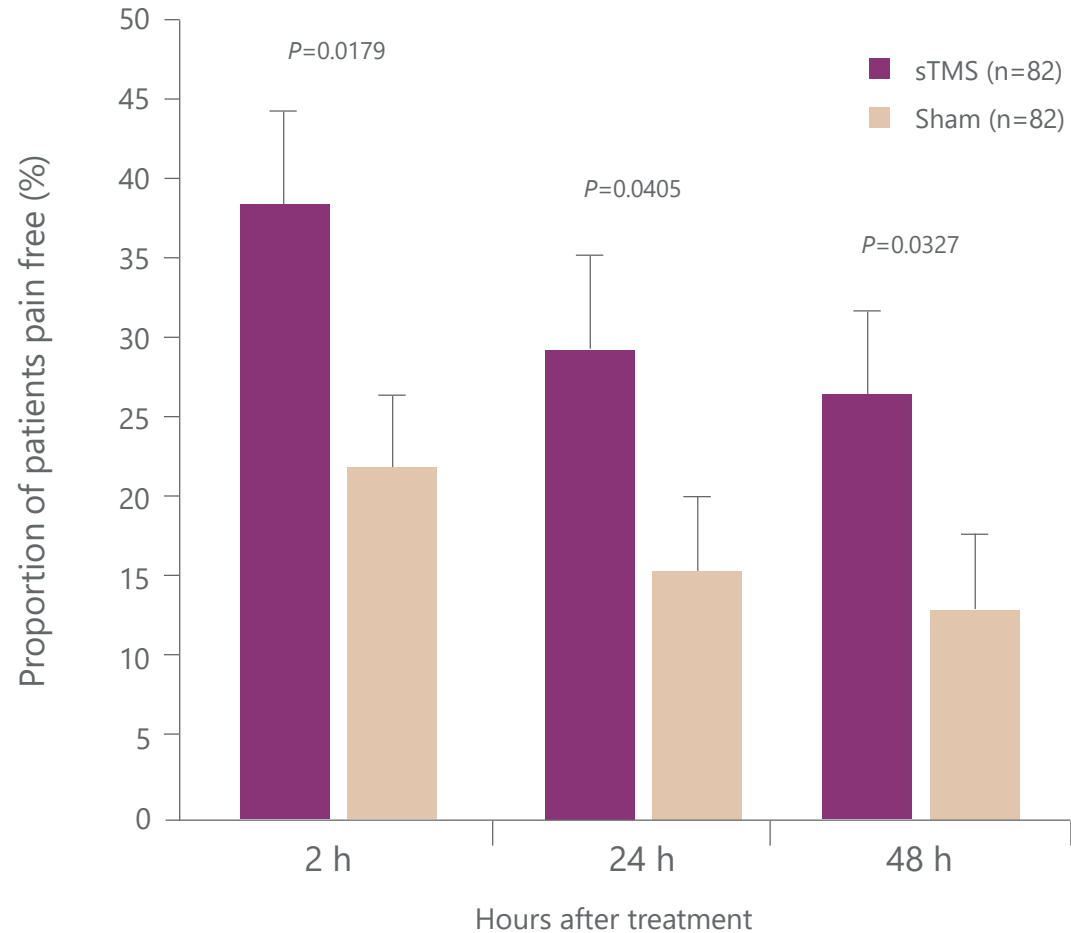
- Pain freedom 2 hours post treatment
- Sustained pain freedom at 24 and 48 hours

Demonstrated tolerability

- AEs were minimal and mild– 5% SAVI vs 2% sham

Source: Lipton RB et al. *Lancet Neurol.* 9(4):373-380.

Significant and sustained pain freedom

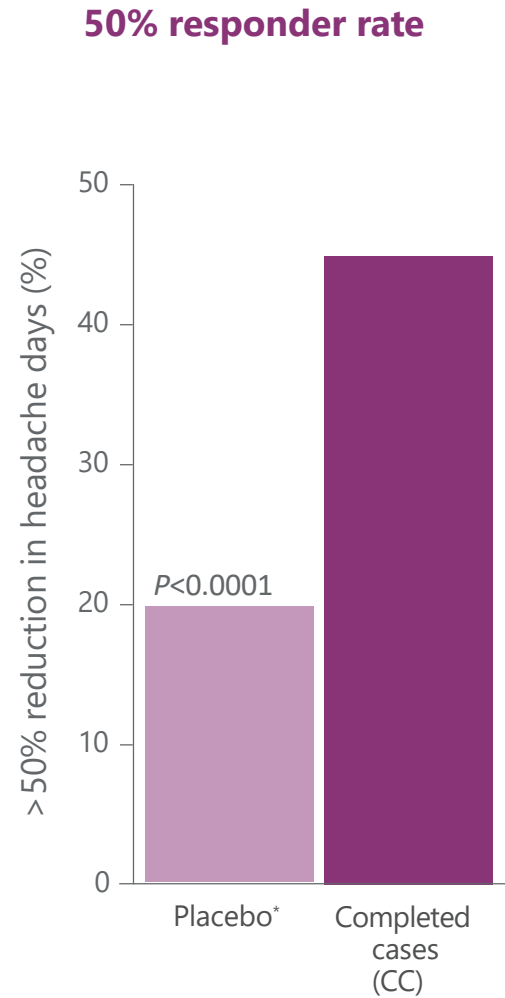


SAVI Dual™ for preventive treatment of migraine

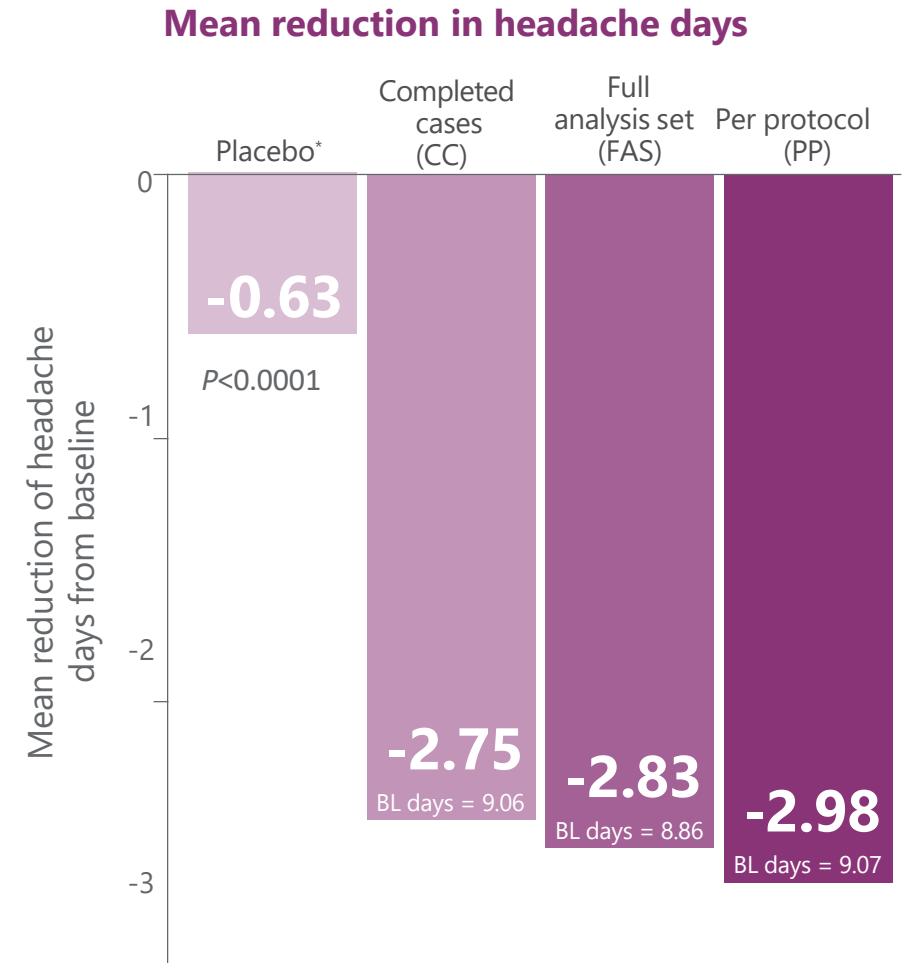
More patients with fewer headache days

-2.75 ± 0.4 less headache days with SAVI treatment vs placebo

Significantly reduced use of acute rescue medications by 3.18 days



*Statistically derived.



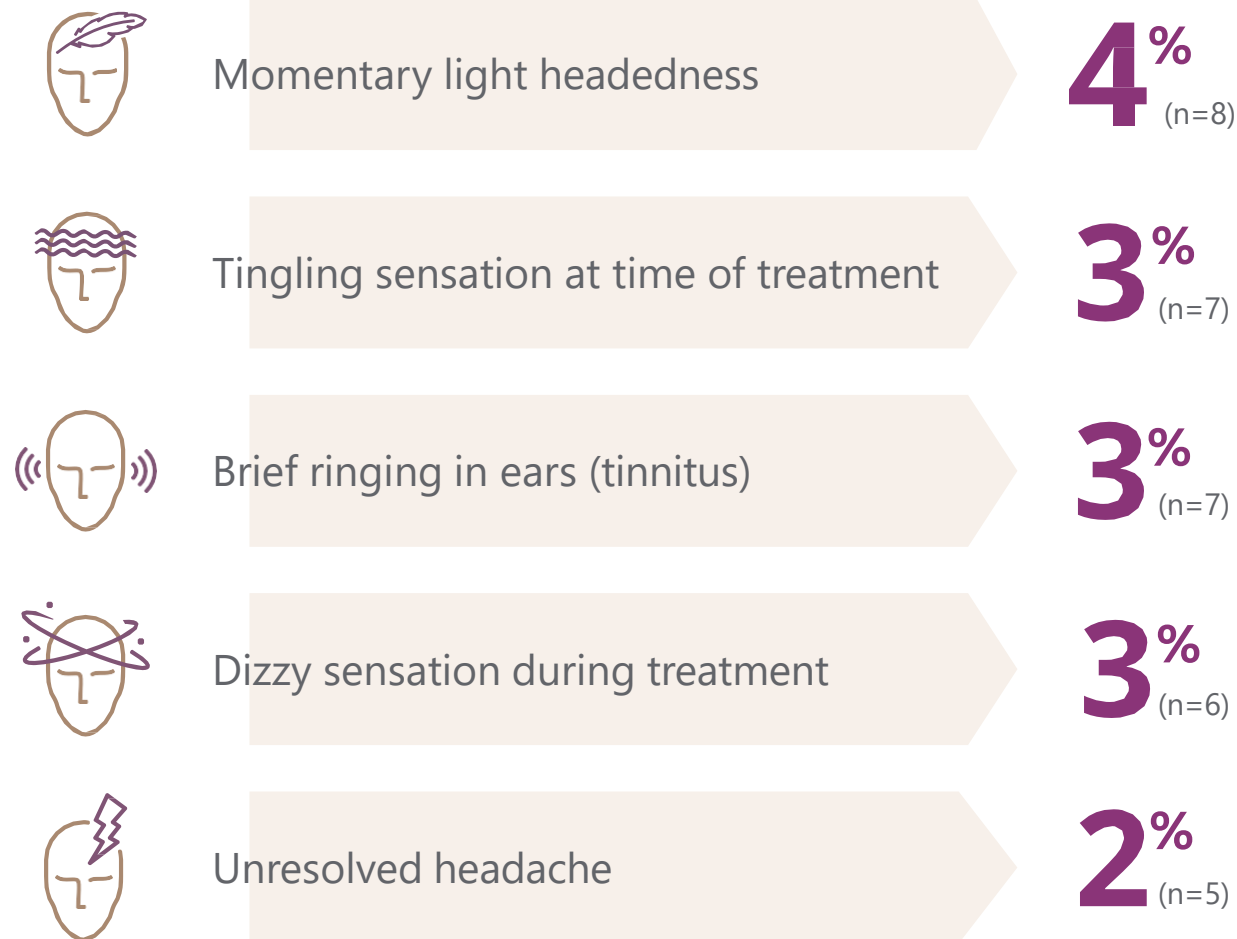
Well-tolerated with no serious adverse events

SAVI Dual is contraindicated for patients with conductive metal implants in the head and neck, excluding dental work.

MRI protocols may be a reasonable guide in assessing use of SAVI Dual in patients with metal implants in the head and neck.

Refer to the Prescriber IFU for a complete list of contraindications and warnings.

5 most common adverse events were mild and typically resolved immediately after treatment



N=217

Source: Starling AJ et al. *Cephalalgia*. 2018 [epub ahead of print].

Clinical Studies and RWE from tertiary headache clinics

- High frequency, refractory patients
- Failed numerous acute and preventive medications
- Well-tolerated with no serious adverse events

Sources:

- Bhola R, et al. Single-pulse transcranial magnetic stimulation (sTMS) for the acute treatment of migraine: Evaluation of outcome data for the UK post market pilot program; . J Headache Pain 2015 16: 535
- Bhola R, et al. Use of Single Pulse Transcranial Magnetic Stimulation (sTMS) to Treat Migraine with Medication Overuse: Response Over Time. Presented at: 2019 American Headache Society Annual Meeting; July 11-14, 2019; Philadelphia, PA. P222LB
- Lloyd JO, et al Single-Pulse Transcranial Magnetic Stimulation for the preventive treatment of difficult-to-treat migraine: a 12-month prospective analysis. J Headache Pain 2023

SAVI Dual™ Real-World Evidence – Mirrors Clinical Trial Results



- Reduction in Headache Days



- Reduction in Medication Use



- Reduction in Medication Overuse Headache



- Improved Disability Scores



- Sustained Response at 3, 6 and 12 months

American Headache Society (AHS) Prescribing Guidelines Dual Migraine Therapy with Neuromodulation

Diagnosis of ICHD-3 migraine with aura, migraine without aura, or chronic migraine

- Inability to tolerate migraine specific medications
- Limits to medication due to comorbid and coexisting illness or concomitant medication
- Adjunctive treatment in those experiencing frequent attacks who are at risk of developing medication-overuse headache and/or chronic migraine
- Patients who prefer to avoid systemic medications

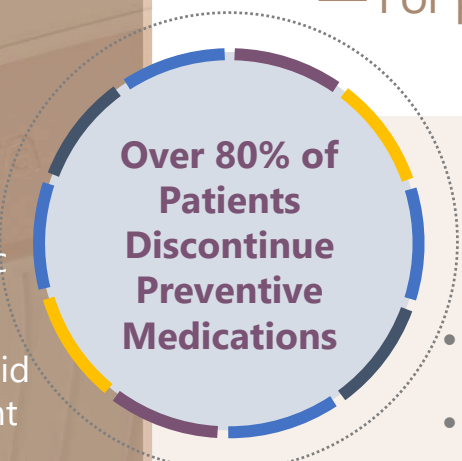
Source:

Ailani J, Burch RC, Robbins MS; Board of Directors of the American Headache Society. The American Headache Society Consensus Statement: Update on integrating new migraine treatments into clinical practice. *Headache*. 2021 Jun 23

Marcus SC, Shewale AR, Silberstein SD, et al. Comparison of healthcare resource utilization and costs among patients with migraine with potentially adequate and insufficient triptan response. *Cephalalgia*. 2020;40(7):639-649

Prescribing SAVI Dual™ Migraine Therapy

— For patient selection, AHS guidelines are a good place to start



Over 80% of Patients Discontinue Preventive Medications

Additional considerations in patient selection:

- Frequent migraine- need both acute and preventive therapy
- Women of childbearing age concerned about medication use
- History of treatment compliance
 - Patients who use SAVI Dual as prescribed are far more likely to see significant improvement in overall well being
- Reasonable expectations for preventive therapy – It may take a few months for preventive therapy to kick in

Comparing Neuromodulation Devices

	FDA Cleared for Acute + Preventive Treatment of Migraine			
	SAVI Dual™	CEFALY®	gammaCore™	Nerivio®
Mechanism	Central Neuromodulation	Peripheral Neuromodulation Transcutaneous Electrical Nerve Stimulation (TENS)*		
Met Primary Endpoint ^{1,2}				
Acute (Therapeutic Gain)	Yes (17%) ³	Yes** (7%) ⁴	No (-) ⁵	Yes (19%) ⁶
Preventive (Therapeutic Gain)	Yes (26%) ⁷	No (-) ⁸	No (-) ⁹	Yes (15.9%) ¹⁰
Availability	Rx	OTC	Rx	RX
Treatment Duration	10 sec	120 mins	4 mins	45 mins
Remote Therapeutic Monitoring (RTM) Automated compliance /Outcome	Yes	No	No	No
Pricing includes unlimited treatment	Yes	No	No	No

¹Guidelines of the International Headache Society for controlled trials of acute treatment of migraine attacks in adults: Fourth edition. *Cephalalgia*.

²Guidelines of the International Headache Society for controlled trials of preventive treatment of migraine attacks in episodic migraine in adults. *Cephalalgia*.

³Lipton RB, et al. Single-pulse transcranial magnetic stimulation for acute treatment of migraine with aura: A randomized, double-blind, parallel-group, sham-controlled trial. *Lancet Neurol* 2010; 9: 373–380

⁴Kuruvilla, D.E., Mann, J.I., Tepper, S.J. et al. Phase 3 randomized, double-blind, sham-controlled Trial of e-TNS for the Acute treatment of Migraine (TEAM). *Sci Rep* 12, 5110 (2022).

⁵Tassorelli C, Grazi L, de Tommaso et al. Noninvasive vagus nerve stimulation as acute therapy for migraine: The randomized PRESTO study. *Neurology*. 2018 Jul 24;91(4):e364-e373.

⁶Yarnitsky D, Dodick DW, Grosberg BM, et al. Remote Electrical Neuromodulation (REN) Relieves Acute Migraine: A Randomized, Double-Blind, Placebo-Controlled, Multicenter Trial. *Headache*.

⁷Diener HC, Goadsby PJ, Ashina M, et al. Non-invasive vagus nerve stimulation (nVNS) for the preventive treatment of episodic migraine: The multicentre, double-blind, randomised, sham-controlled PREMIUM trial. *Cephalalgia*. 2019 Oct;39(12):1475-1487.

⁸Tepper SJ, Rabany L, Cowan RP et al. Remote electrical neuromodulation for migraine prevention: A double-blind, randomized, placebo-controlled clinical trial. *Headache*. 2023; 63:377–389.

⁹Tepper SJ, Rabany L, Cowan RP et al. Remote electrical neuromodulation for migraine prevention: A double-blind, randomized, placebo-controlled clinical trial. *Headache*. 2023; 63:377–389.

Therapeutic Gain = Active response – Placebo response (sources available upon request)

Pricing sources: Federal Supply Schedule (2023), company websites

*TENS devices available over the counter for the treatment of pain in various product configurations, and price ranges

**Pain Status measured immediately after 2 hours of continuous treatment.

Well-tolerated:
treat as needed to
improve overall well-
being

Prescription includes plenty of pulses
for daily preventive treatment and
for treating migraine attacks as
prescribed

**Sample Treatment Plan based on SAVI Dual
clinical studies:**

Daily preventive treatment:

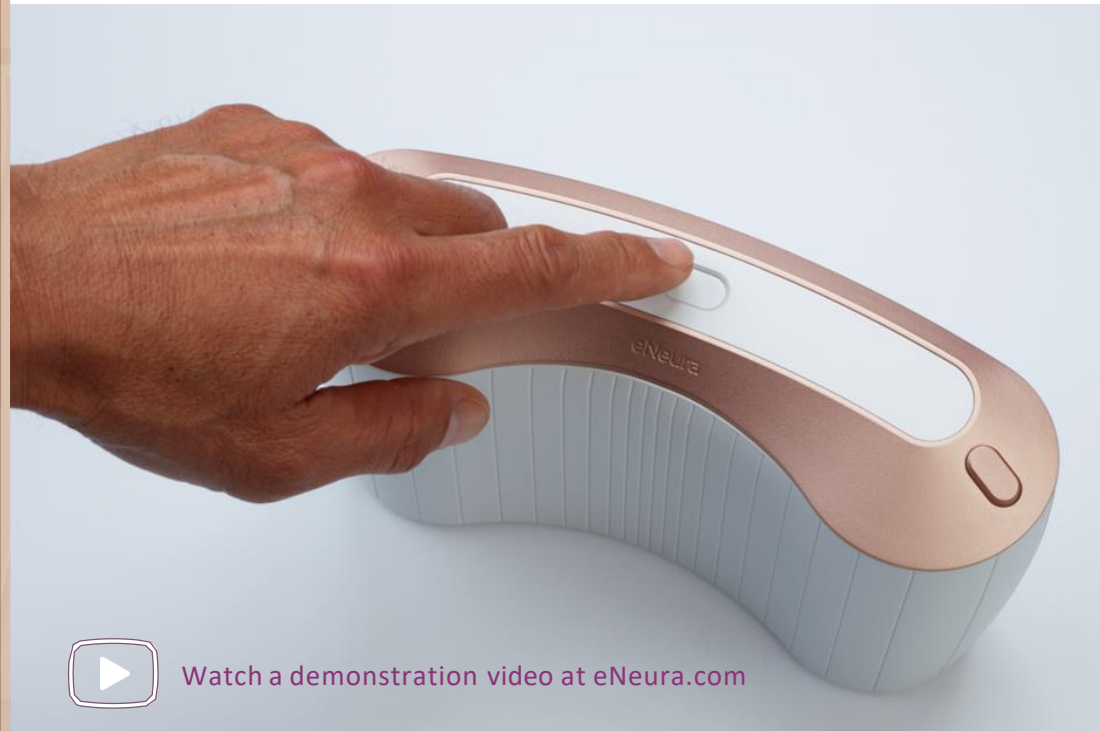
- 4 pulses twice daily

Acute Treatment:

- 4 pulses at the first sign of the migraine attack
- the earlier, the better
- If needed, treat again every 15-30 minutes
until migraine attack subsides

SAVI Dual™ Migraine Therapy Suggested Treatment Plan

Unlimited, Personalized treatment puts the patient in control



- Takes less than a minute
- Treat as early and often as needed
- Encourage patient to use SAVI Dual for 3 months before adjusting therapy

To learn more, review the SAVI Dual Patient Quick Guide at www.eneura.com/support

SAVI Smart Diary™ for your patients

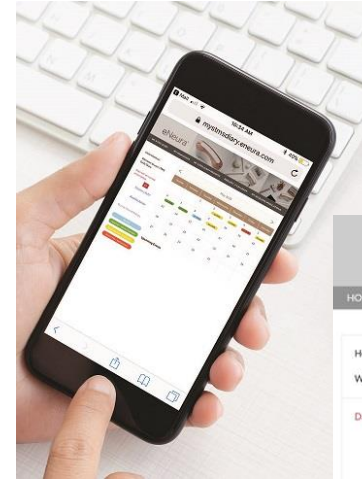
Secure, wireless headache diary made easy

Allows patient to quickly record key data in about 15 seconds

- Patient can work with HCP to personalize treatment programs and optimize overall well-being
- Compliant with PHI Standard
- Patients can login directly from eNeura website

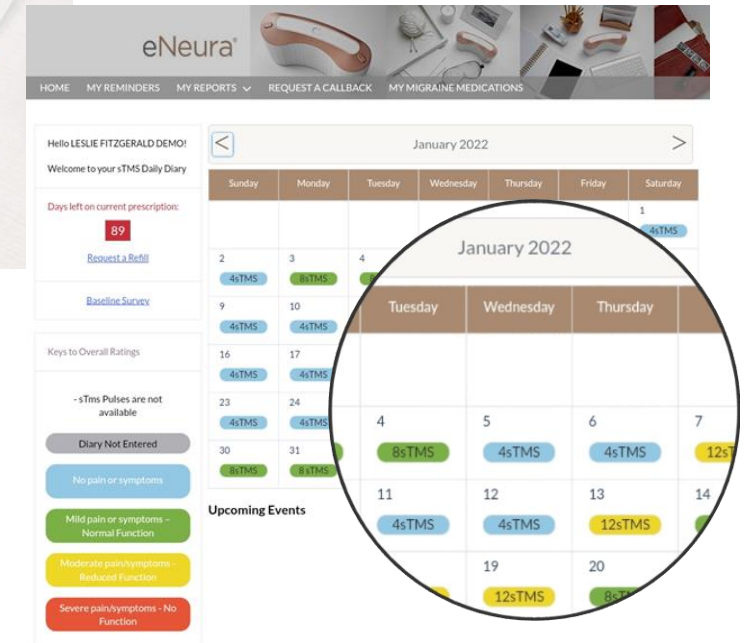


SAVI Dual automatically tracks and records treatment history



Patient records:

- Daily disability status
- Medication Use
- Headache Status

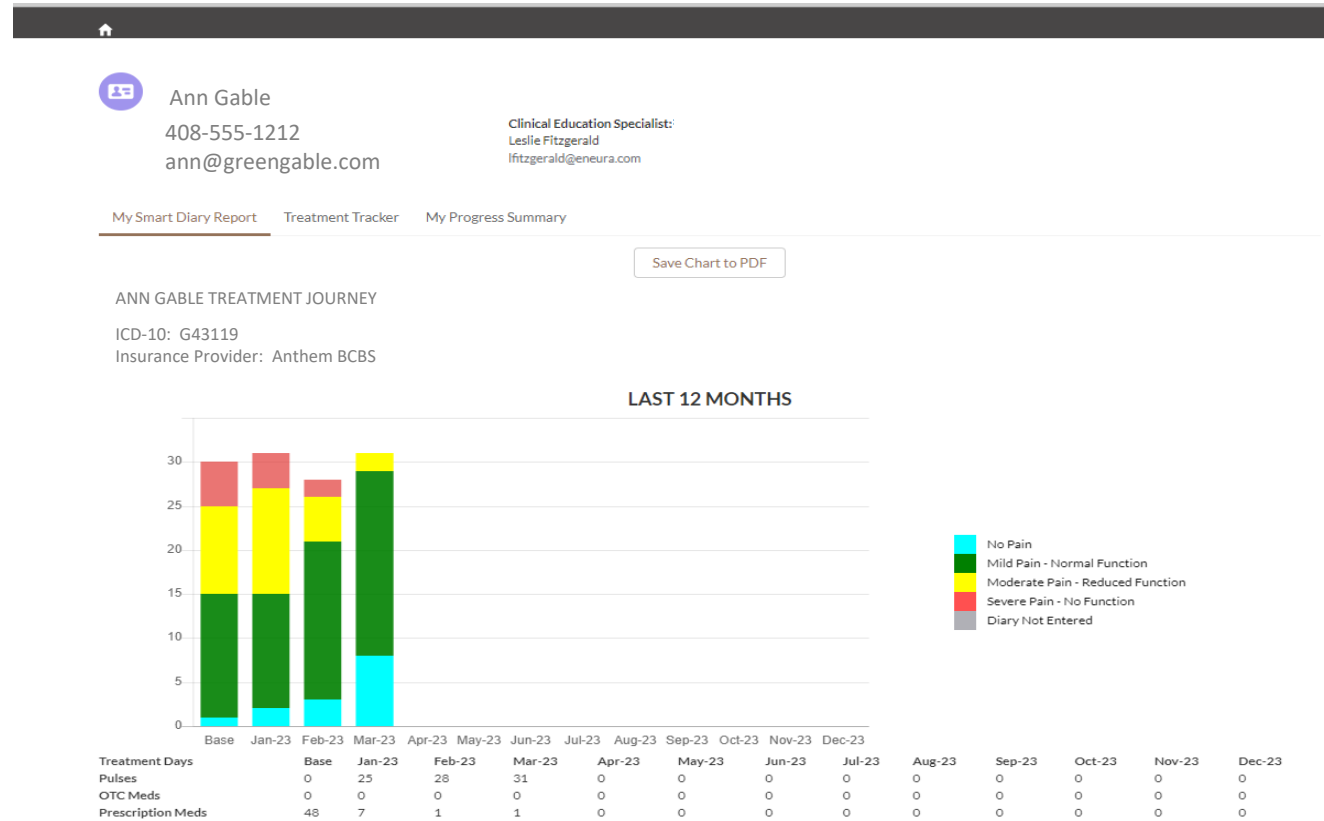


Clinicians and patients can easily track treatment progress

SAVI Dual™ Remote Therapeutic Monitoring (RTM) for Healthcare Providers

Remote Therapeutic Monitoring (RTM)

- Allows you and your patients to monitor migraine patterns and treatment progress
- Work with your patients to personalize treatment programs and optimize overall well-being
- Treatment Journey compares patient's baseline data to SAVI Dual treatment data



To access SAVI RTM, login at [eneura.com](https://www.eneura.com) – click **HCP login**

To learn more about SAVI RTM, review the SAVI Remote Therapeutic Monitoring (RTM) Quick Guide at <https://www.eneura.com/for-healthcare-professionals/>

Personalized Patient Support

eNeura provides your patient one-to-one support

eNeura's **Customer Care team** is here to answer any questions your patient may have, such as how to fill or renew their prescription or take care of their SAVI Dual.

The dedicated **eNeura Clinical Specialist** will walk your patient through their first treatment, step-by-step. They will find it remarkably easy.

Your eNeura Migraine Team



Customer Care



Clinical Education Specialists

Company Information

eNeura Inc. is a wholly-owned subsidiary of Aruene Corporation

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Cage Code: 8YSK9

Email: customercare@eneura.com

Fax: 877.264.1818

Phone: 833.499.9300, option 1

How to prescribe SAVI Dual™ Migraine Therapy

1



Submit a Consult Request to Prosthetics Dept.

2



Prosthetics: Fax PO to eNeura Customer Care

3



eNeura ships directly to your patient

Ordering Information

Item # 16140-0020-13-3 3-month activation of SAVI Dual Migraine Therapy

Item # 16140-0020-13-12 12-month activation of SAVI Dual Migraine Therapy

Activation purchase includes:

- Unlimited use of SAVI Dual for acute and preventive migraine therapy during the activation period
- Patient access to wireless SAVI Smart Diary
- HCP access to SAVI Remote Therapeutic Monitoring (RTM)

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SAVI Dual™ Migraine Therapy

Available Resources at www.eneura.com:

Click the HCP tab for:

- ✓ SAVI Dual Quick Guide – also ships with every device
- ✓ SAVI Remote Therapeutic Monitoring (RTM) Quick Guide
- ✓ How to prescribe

Click the Clinical Data and Publications tab for:

- ✓ Clinical trials and studies
- ✓ Real world evidence
- ✓ Safety data
- ✓ Mechanism of Action
- ✓ Professional Guidelines

