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*San Diego!*

# Pain Reprocessing Therapy: Retraining the Brain



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Speaker and all moderators and planners have no relevant financial relationships with ineligible companies to disclose.

# Learning Outcomes

- Explain the pathophysiology of nociplastic pain and distinguish it from nociceptive and neuropathic pain
- Describe the principles of Pain Reprocessing Therapy (PRT) and discuss the application in the management of nociplastic pain conditions

# Mechanistic Types of Chronic Pain

Pain Type	Defining Characteristics	Examples	Treatment
<b>Nociceptive</b>	Pain due to tissue injury, inflammation, damage or degeneration	Osteoarthritis, rheumatoid arthritis, fracture, burns	Topical analgesics, nonsteroidal anti-inflammatory drugs, acetaminophen, opioids, steroids
<b>Neuropathic</b>	Pain due to nerve injury or damage	Radiculopathy, diabetic neuropathy, chemotherapy-induced neuropathy	Topical or local therapy; systemic neuropathic medications such as gabapentin, pregabalin, and tricyclic antidepressants
<b>Nociplastic</b>	Pain arising from a sensitized nervous system (amplified processing of pain signals, decreased inhibition of pain or both)	Fibromyalgia, chronic back pain, chronic temporomandibular pain disorders	Multimodal management approach

# Nociplastic Pain

- Term proposed by the IASP in 2016
- A mechanistic descriptor for chronic primary pain defined as:  
***“Pain arising from the altered function of pain-related sensory pathways in the periphery and CNS causing increased sensitivity”***
- Due to changes in **neurophysiology** rooted in **neuroplasticity**
- Involves amplified processing and/or decreased inhibition of pain stimuli at multiple levels
  - Supraspinal
  - Spinal
  - Peripheral

Fitzcharles MA, et al. *Lancet*. 2021;397(10289):2098-2110.



# Localized Nociplastic Pain Conditions



Thapa R. *Cleve Clin J Med.*  
2025;92(4):236-247

## **Chronic Primary Headache and Orofacial Pain**

Chronic migraine

Chronic tension-type headache

Trigeminal autonomic cephalalgias

Chronic temporomandibular pain disorders without anatomic abnormality or explanation

Chronic burning mouth

Chronic primary orofacial pain

## **Chronic Visceral Pain Syndrome**

Chronic primary bladder pain syndrome or interstitial cystitis

Chronic pelvic pain syndrome

Irritable bowel syndrome

Chronic chest pain

Chronic abdominal pain

## **Chronic Primary Musculoskeletal Pain**

Primary cervical, thoracic, lower back and limb pain; extent of pain and suffering is greater than expected based on the underlying pathology

Complex regional pain syndrome

# 2020 Revised IASP Definition of Pain

- **IASP definition of pain:**

***“An unpleasant sensory and emotional experience associated with, or resembling that associated with, actual or potential tissue damage”***

- **Key notes:**

- Pain is always a personal experience that is influenced to varying degrees by biological, psychological, and social factors.
- **Pain and nociception are different phenomena.** Pain cannot be inferred solely from activity in sensory neurons.
- Through their life experiences, individuals *learn* the concept of pain.



Raja SN, et al. *Pain*. 2020;161(9):1976-1982.

# What is *pain*?

- Pain is a danger signal generated by the **brain**
- An internal reinforcement signal used for learning to guide behavior
- The brain is not passively waiting for nociceptive stimuli to impinge on it, but is actively making inferences based on prior experience and expectations
- Sometimes the brain infers incorrectly and generates a *false alarm*

Seymour B. *Neuron*. 2019;101(6):1029-1041  
Kaptchuk TJ, et al. *BMJ*. 2020;370:m1668.  
Büchel C, et al. *Neuron*. 2014;81(6):1223-1239.



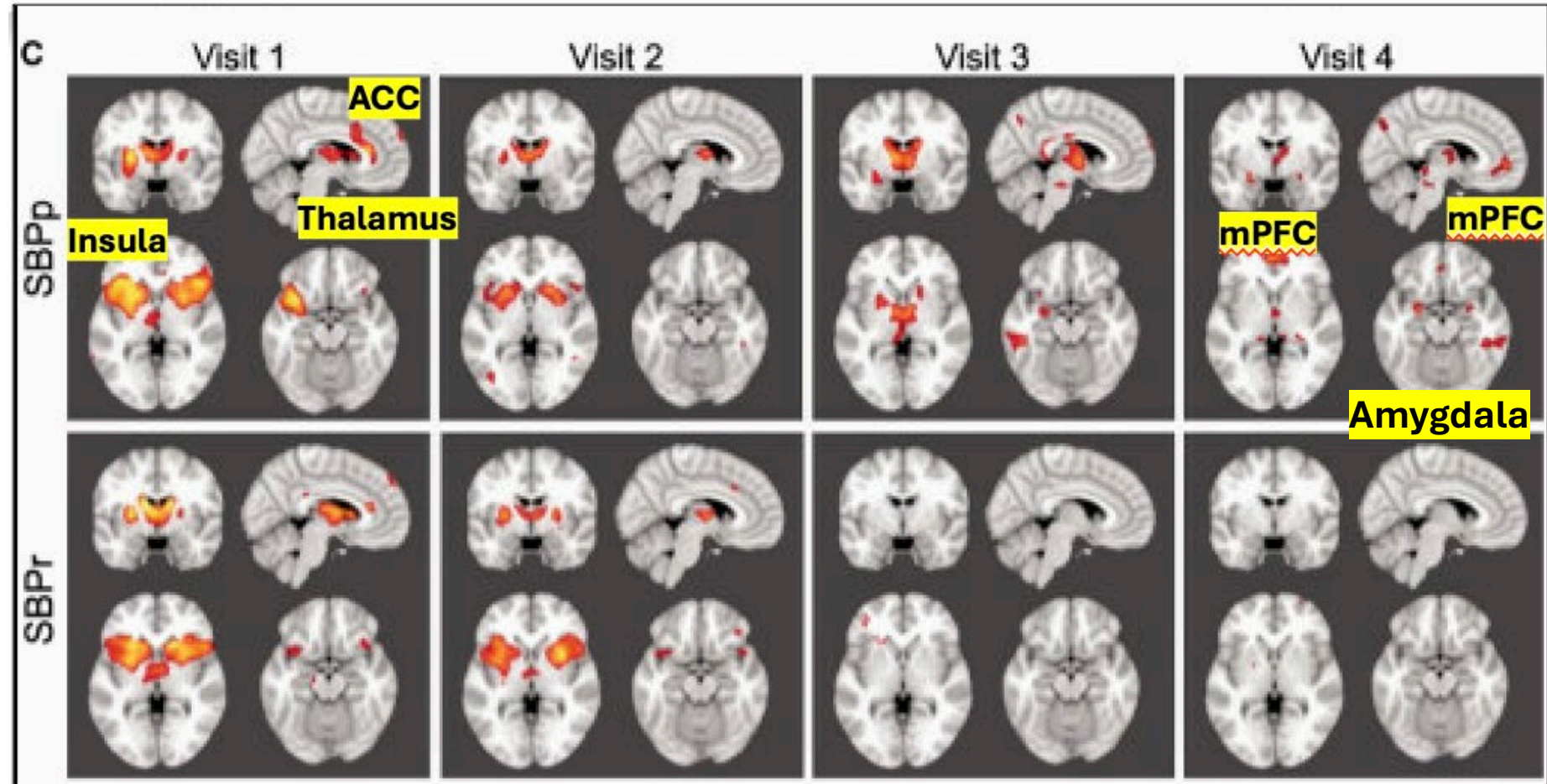
# Chronic Pain and the Brain

- As pain transitions from acute to chronic we see the following changes in brain activity
- Pain becomes:
  - Less closely tied to systems encoding nociceptive input
  - Increasingly associated with activity in the emotional/motivational systems associated with avoidance
- Brain regions serving allostasis and predictive control have been implicated in animal and human studies of chronic pain modulation

Ashar YK, et al. *JAMA Psychiatry*. 2022;79(1):13-23. || Kuner R, et al. *Nat Rev Neurosci*. 2016;18(1):20-30. || Hashmi JA, et al. *Brain*. 2013;136(Pt 9):2751-2768. || Roy M, et al. *Nat Neurosci*. 2014;17(11):1607-1612

# Shape Shifting Back Pain

- Chronification of back pain shifts brain representation from nociceptive to emotional circuits



Hashmi JA, et al. *Brain*. 2013;136(Pt 9):2751-2768.

If the brain can learn pain,  
can the brain ***unlearn*** the pain?

# Pain Reprocessing Therapy (PRT)

- Developed based on the neuroscience of chronic primary pain/nociplastic pain
- A mind-body intervention aimed to help patients reconceptualize their pain as due to **nondangerous brain activity** rather than *peripheral tissue injury*
- Retrains the brain to interpret & respond to signals from the body properly

Ashar YK, et al. *JAMA Psychiatry*. 2022;79(1):13-23.

# What does PRT involve?

1. **Education** about the brain origins and reversibility of nociplastic pain/primary pain
2. Gathering and reinforcing personalized **evidence** for the brain origins and reversibility of pain
3. Attending to and appraising pain sensations through a lens of **safety**
  - Somatic tracking
4. Addressing other emotional **threats**
5. Gravitating towards **positive** feelings and sensations



Ashar YK, et al. *JAMA Psychiatry*. 2022;79(1):13-23.



# PRT for Chronic Back Pain

“The Boulder Back Pain Study”

Research

JAMA Psychiatry | [Original Investigation](#)

## Effect of Pain Reprocessing Therapy vs Placebo and Usual Care for Patients With Chronic Back Pain A Randomized Clinical Trial

Yoni K. Ashar, PhD; Alan Gordon, LCSW; Howard Schubiner, MD; Christie Uipi, LCSW; Karen Knight, MD; Zachary Anderson, BS; Judith Carlisle, MA; Laurie Polisky, BA; Stephan Geuter, PhD; Thomas F. Flood, MD, PhD; Philip A. Kragel, PhD; Sona Dimidjian, PhD; Mark A. Lumley, PhD; Tor D. Wager, PhD



Ashar YK, et al. *JAMA Psychiatry*. 2022;79(1):13-23.



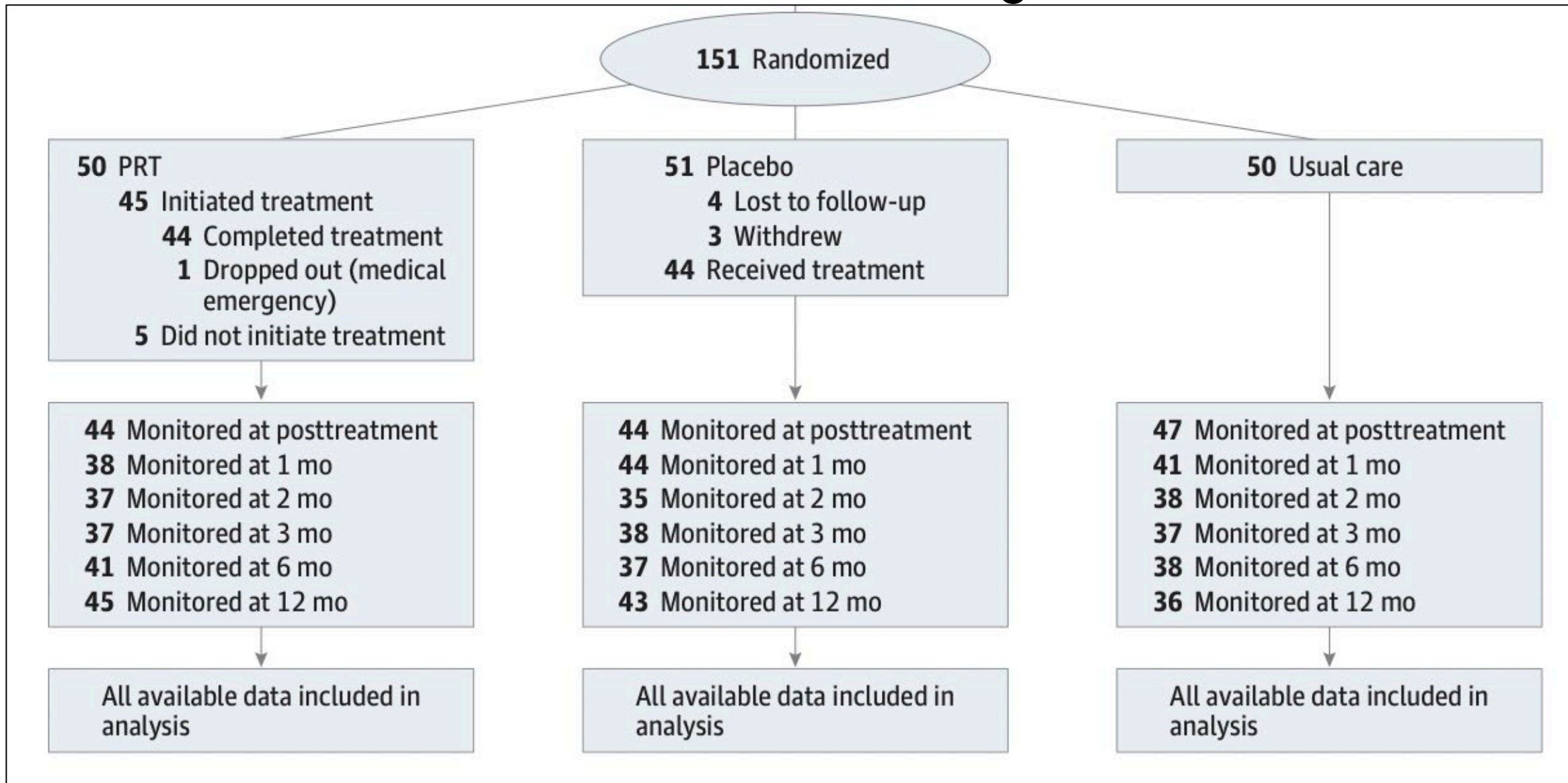
# Boulder Back Pain Study

- $N=151$  pts with chronic back pain
- Inclusion criteria:
  - Pt 21-70 yo
  - Back pain for at least 1/2 the days of the last 6 months
  - 1-week average pain intensity score of 4/10 or greater at screening
  - Excluded pts with leg pain worse than back pain or bowel/bladder incontinence



Ashar YK, et al. *JAMA Psychiatry*. 2022;79(1):13-23.

# Boulder Back Pain Study



# Boulder Back Pain Study

- Spinal anomalies among participants randomized to PRT with prior spinal imaging ( $N=20$ )
- All had at least 1 spinal anomaly, median of 4 per participant

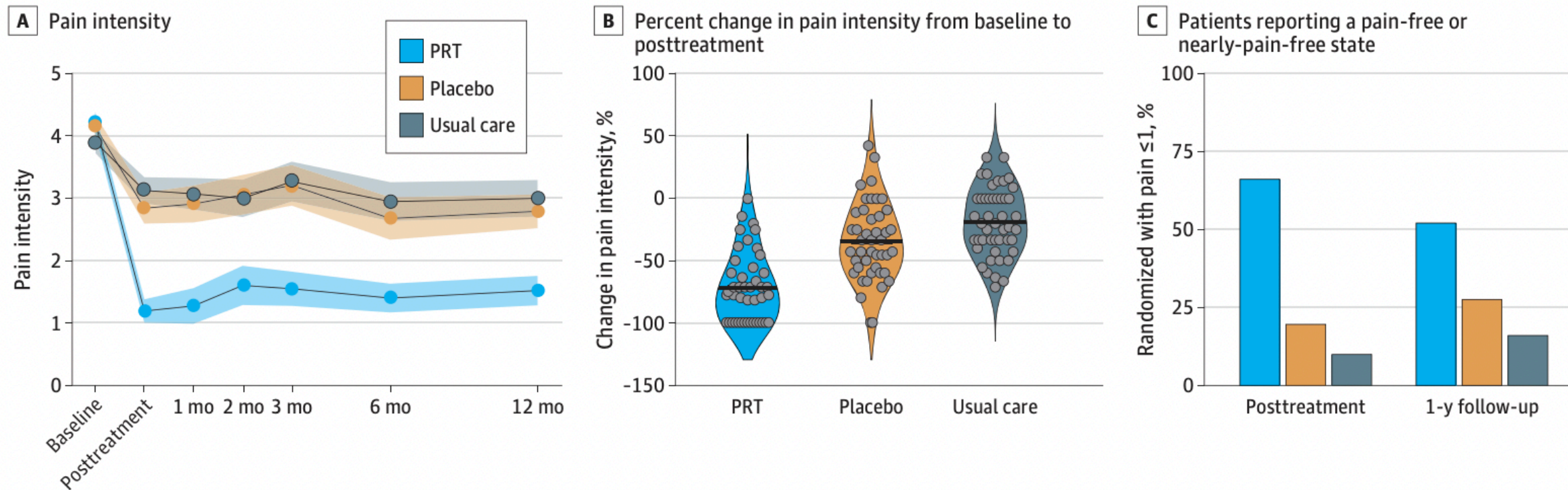
Radiological finding	<i>N</i>	%
Disc degenerative changes	15	75%
Disc herniation or rupture	7	35%
Spinal misalignment	14	70%
Osteoarthritic changes	13	65%
Neuroforaminal narrowing	9	45%
Central canal stenosis	9	45%

Ashar YK, et al. *JAMA Psychiatry*. 2022;79(1):13-23.



# Results

**Figure 2. Clinical Outcomes**



A, Shading indicates standard error. B, Dots represent individual participants; thick lines represent the group mean. C, Percentage of patients reporting pain scores of 0 or 1 of 10 (ie, pain-free or nearly pain-free) at posttreatment and at 1-year follow-up. PRT indicates pain reprocessing therapy.

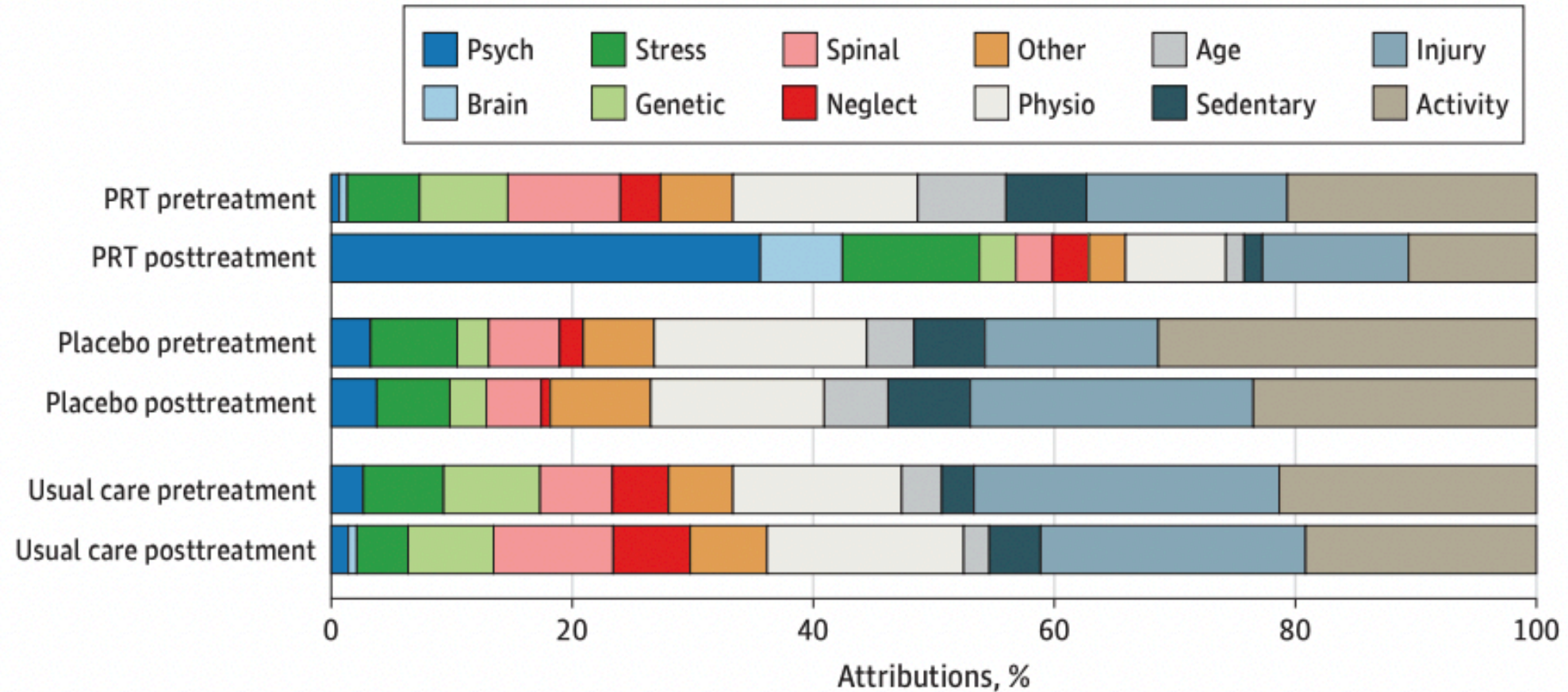
# Changes in Brain Connectivity

- Longitudinal fMRI outcomes:
  - Reduced responses to evoked back pain in the anterior midcingulate and the anterior prefrontal cortex for PRT vs placebo
  - Reduced responses in the anterior insula for PRT vs usual care
  - Increased resting connectivity from the anterior prefrontal cortex and the anterior insula to the primary somatosensory cortex for PRT vs both control groups
  - Increased connectivity from the anterior midcingulate to the precuneus for PRT vs usual care

Ashar YK, et al. *JAMA Netw Open*. 2023;6(9):e2333846.

# Reattribution of Pain

Figure 2. Pain Attribution Category Prevalence



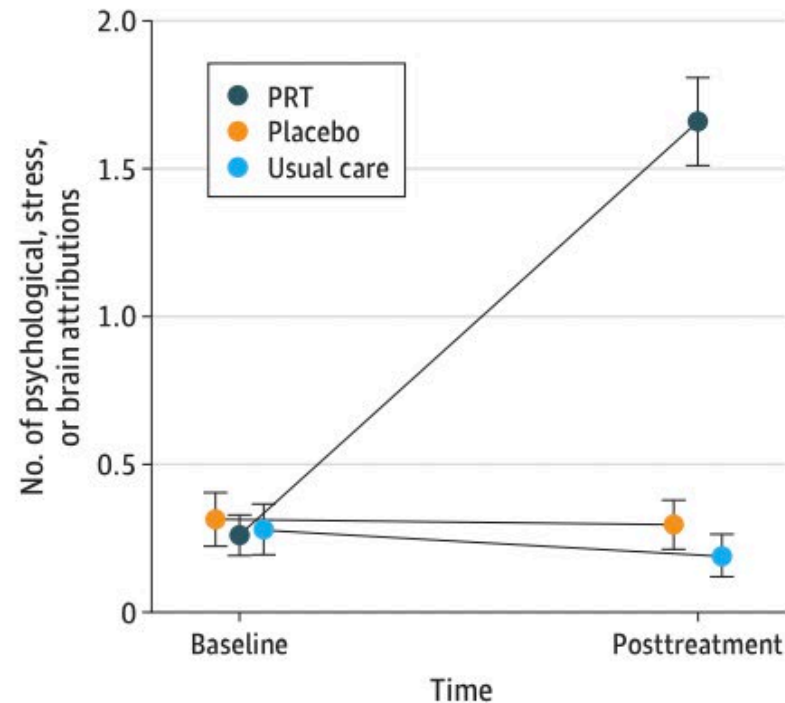


# Reattribution of Pain

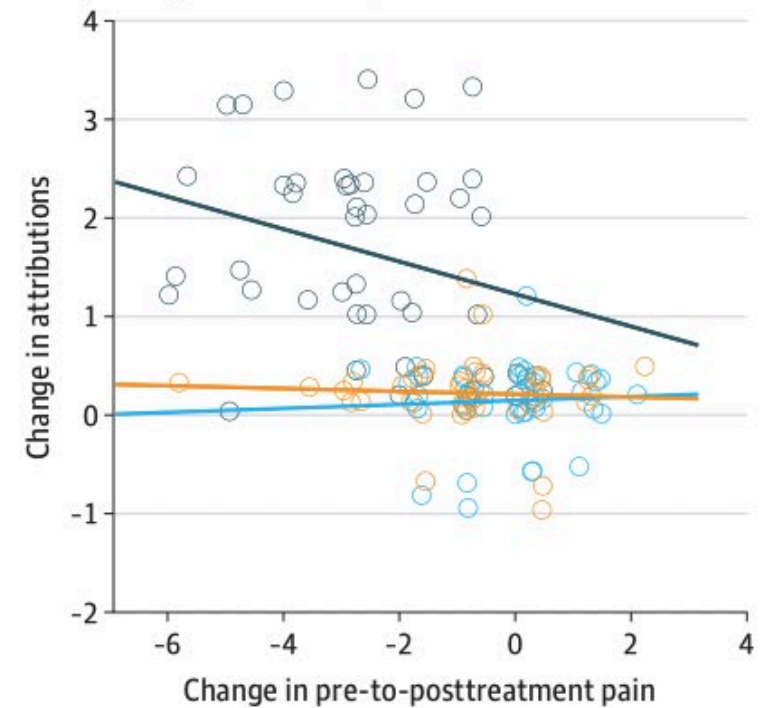
- PRT produced large increases in mind-brain attribution scores compared to control
- Increased mind/brain attributions within PRT significantly associated with decreases in pain intensity
- Reattribution alone not necessarily enough to change pain

**Figure 3. Effects of Pain Reprocessing Therapy (PRT) on Patients' Attributions Regarding the Underlying Causes of Chronic Back Pain**

**A** Psychological, stress, or brain attributions

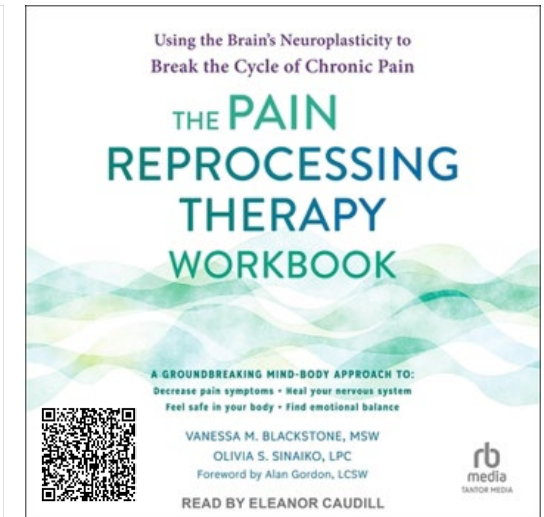
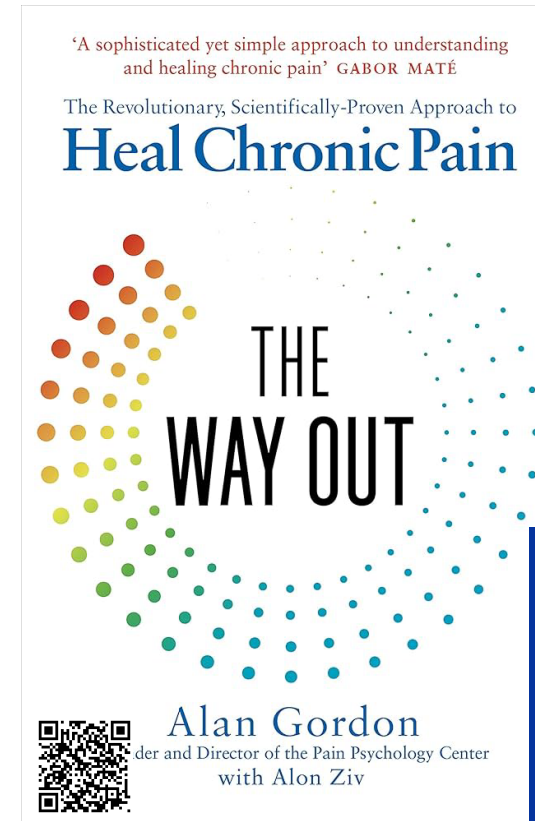


**B** Association of attributions with changes in pre-to-posttreatment pain



# Patient Education Resources

- Association for the Treatment of Neuroplastic Symptoms (ATNS)
  - Patient education videos on *Symptomatic.me*
- Book: *The Way Out* by Alan Gordon
- Pain Reprocessing Therapy Workbook
- Curable App
- Insight Timer (somatic tracking)
- Podcasts
  - Pain Reprocessing Therapy Podcast
  - The Story Behind The Symptoms (ATNS- David Clarke MD)



# Key Points

- Nociplastic pain is due to maladaptive rewiring of the nervous system that leads to sensitization of the peripheral and central nervous system
- Pain reprocessing therapy is a promising treatment for chronic low back pain

# Thank You!