## Biomedical and Biopsychosocial Aspects of Pain Management

## **MI-CCSI**

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## Disclosures

- Consultant to Community Health Focus Inc.
- Consultant to Swing Therapeutics, Inc.
- Funded for research by NIH
- Funded for research by PCORI

Why is Chronic Pain so difficult to treat?



Why is Chronic Pain so difficult to treat?



- > Multiple underlying causes
- > Individual differences
- > Neuroplasticity
- >Treatment limitations
- **▶**Biopsychosocial factors
- >A holistic approach is required

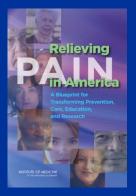
Why is Chronic Pain so difficult to treat?





## **Chronic Pain Numbers**





100 Million People
- US



150 Million
- 37 Countries

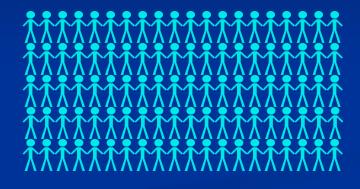
Eccleston, C., Wells, C. (2017). European Pain Management. Oxford University Press



# More people have Chronic Pain than Diabetes, Heart Disease, and Cancer Combined

Chronic Pain

100 Million



**Diabetes** 

29.1 Million

Heart Disease

27.6 Million

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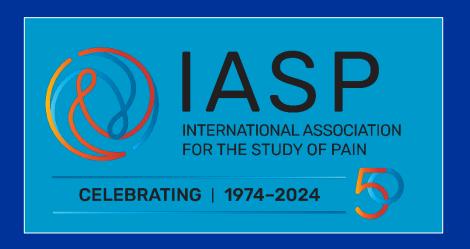
Cancer

13.7 Million



### What is Pain?

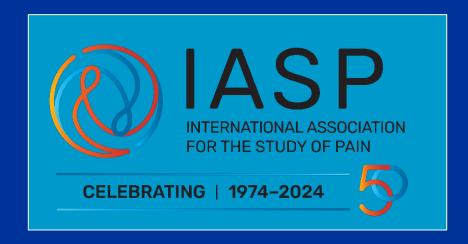
An unpleasant sensory and emotional experience associated with, or resembling that associated with, actual or potential tissue damage



### What is Pain?

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

- •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.
- •A person's report of an experience as pain should be respected.
- •Although pain usually serves an adaptive role, it may have adverse effects on function and social and psychological wellbeing.
- •Verbal description is only one of several behaviors to express pain; inability to communicate does not negate the possibility that a human or a nonhuman animal experiences pain.



## How Does Pain Happen?

## Which person has pain?



Normal



Bone on bone

## Which person has pain?

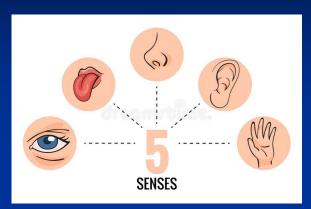


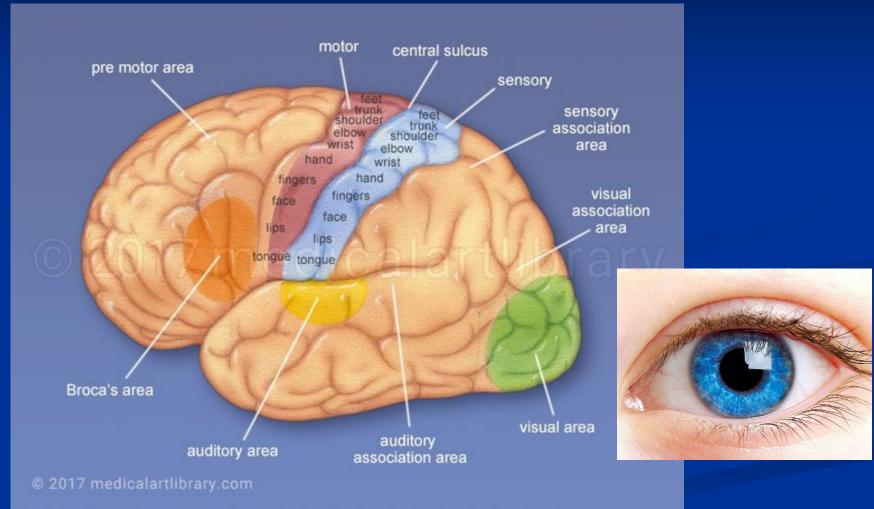


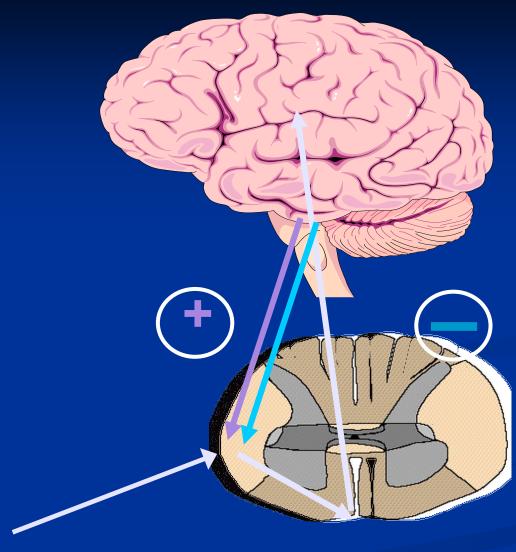
Bone on bone 30-40% Have no pain

"Normal"
10% have pain

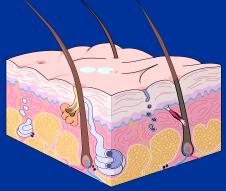
## Multi-Stage Sensory Processing







Nociception



## CNS Neurotransmitters Influencing Pain

#### **Facilitation**

Gabapentinoids, ketamine

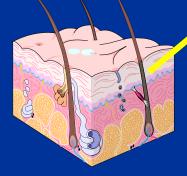
Glutamate and EAA

Substance P

Nerve growth factor

Serotonin (5HT<sub>2a, 3a</sub>)

Anti-migraine drugs (– triptans), cyclobenzaprine





Descending antinociceptive pathways

Norepinephrineserotonin (5HT<sub>1a.b</sub>), dopamine

tramadol

Opioids

Low dose naltrexone

Cannabinoids

**GABA** 

Gammahydroxybutyrate, moderate alcohol consumption No knowledge of endocannabinoid activity but this class of drugs is effective

- 1. Schmidt-Wilcke T, Clauw DJ. Nat Rev Rheumatol. Jul 19 2011.
- 2. Clauw DJ. JAMA. 2014.

## Neurobiological perspective

Brain regions associated with pain processing involve both sensory and affective/cognitive regions

- Sensory / discriminative dimension
  - Somatosensory cortices (S1, S2)
  - Dorsal posterior insula
- Affective / Cognitive dimensions
  - Anterior insula
  - Prefrontal cortex
  - Anterior cingulate cortex
  - Thalamus
  - Amygdala
  - Hippocampus



## Neurobiological perspective

Brain regions associated with pain processing involve both sensory and affective/cognitive regions

- Sensory discriminative dimension
  - Some Cortices (S1, S2)
  - Do poste insula
- Affective / Cognitive dimensions
  - Anterior insula
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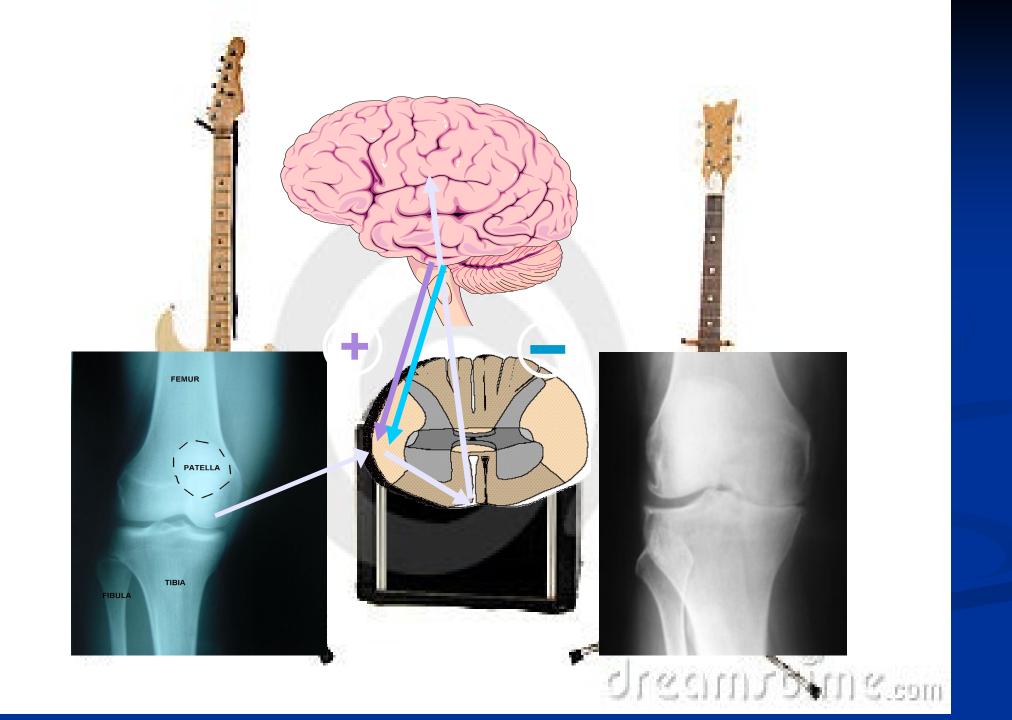


## Neurobiological perspective

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#### Norepinephrine

Concentration
Circadian rhythms
Attention
Stress
Energy

#### Norepinephrine |

Concentration
Circadian rhythms
Attention
Stress
Energy

#### Serotonin

Well-being
Sleep
Affect /Mood
Appetite

#### Norepinephrine |

Concentration
Circadian rhythms
Attention

Stress

Energy

#### <u>Serotonin</u>

Well-being Sleep

Affect /Mood

Appetite

#### **Dopamine**

Attention Pleasure Reward

#### Norepinephrine |

Concentration
Circadian rhythms

Attention

Stress

Energy

#### **Serotonin**

Well-being

Sleep

Affect /Mood

Appetite

#### **Dopamine**

Cognitive

Function

Attention

Pleasure

Reward

#### **Glutamate**

Major Exciter of CNS, Synaptogenesis and neurogenesis

#### Norepinephrine

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**Function** 

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Reward

#### **GABA**

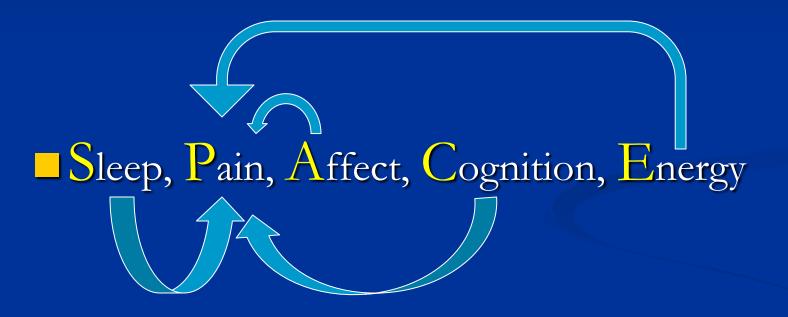
Major Inhibitor of CNS, Sleep/wake cycle

## Shared Neurotransmitters Explain

■ The complexity of chronic pain presentation

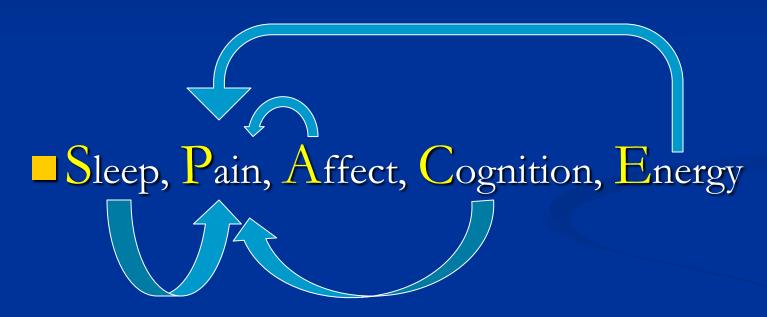
## Shared Neurotransmitters Explain

■ The complexity of chronic pain presentation



## Shared Neurotransmitters Explain

■ The complexity of chronic pain presentation



■ SPACE represents new targets for treating pain perception

## How is Pain Classified?

Time

**Body Location** 

**Suspected Etiology** 

Acute Vs Chronic

Head, Neck, Back, Pelvis

Cancer, Rheumatic, etc.

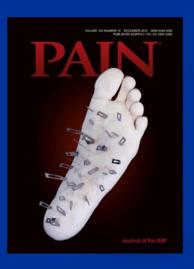
#### Newest Classification: Pain Mechanisms

Nociceptive Peripheral damage or inflammation

Neuropathic

Nociplastic







## Nociceptive Pain

Mechanical, Thermal, Chemical Inflammation or Damage



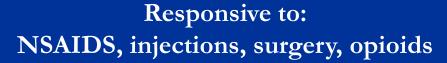




Pain is well localized Consistent with injury/damage



Classic Examples: OA, autoimmune disorders, Cancer pain, Injury







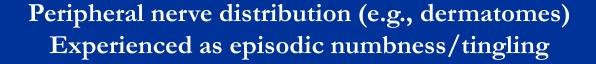






## Neuropathic Pain (Peripheral or Central)

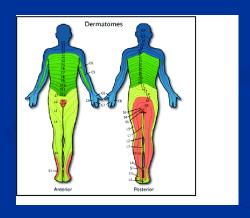
Nerve Damage or Entrapment



Responsive to:
Local treatments of nerves
Surgery, injections, CNS-acting drugs

Classic Examples:
Painful diabetic neuropathy, Sciatica,
Carpal Tunnel, Post-Stroke pain









## Nociplastic Pain

CNS augmentation of peripheral nociception or CNS generation

#### Pain is widespread

- Disproportionate to injury
- Accompanied by CNS-mediated symptoms

Responds to: CNS acting drugs and Non-pharm treatments

#### Classic Examples:

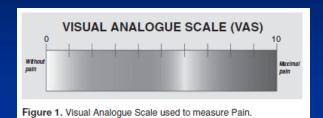
- Fibromyalgia
- Chronic overlapping pain conditions (COPCS)

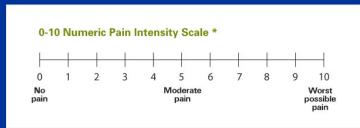


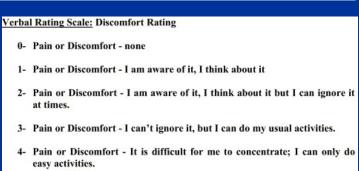
## How Do you Assess Pain?

- Intensity
- Multi-focal (widespread)

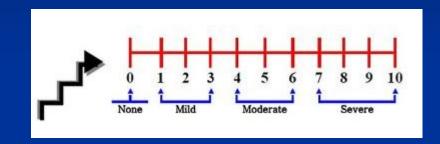
## Intensity







5- Pain or Discomfort - Such that I cant do anything.







#### **Chronic Pain 2**

## Nociplastic pain: towards an understanding of prevalent pain conditions

Mary-Ann Fitzcharles\*, Steven P Cohen\*, Daniel J Clauw, Geoffrey Littlejohn, Chie Usui, Winfried Häuser

Lancet 2021; 397: 2098-110

See Comment page 2029

This is the second in a Series of three papers about chronic pain

\*Contributed equally

Department of Rheumatology and Alan Edwards Pain Management Unit, McGill University, Montreal, QC, Canada (M-A Fitzcharles MBChB);

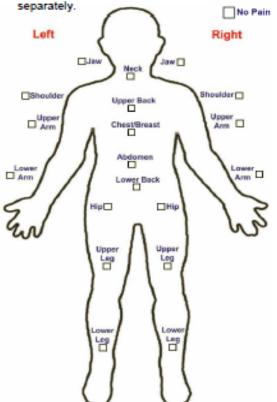
(M-A Fitzcharles MBChB); Department of Psychiatry and Behavioral Sciences and Department of Anesthesiology

Nociplastic pain is the semantic term suggested by the international community of pain researchers to describe a third category of pain that is mechanistically distinct from nociceptive pain, which is caused by ongoing inflammation and damage of tissues, and neuropathic pain, which is caused by nerve damage. The mechanisms that underlie this type of pain are not entirely understood, but it is thought that augmented CNS pain and sensory processing and altered pain modulation play prominent roles. The symptoms observed in nociplastic pain include multifocal pain that is more widespread or intense, or both, than would be expected given the amount of identifiable tissue or nerve damage, as well as other CNS-derived symptoms, such as fatigue, sleep, memory, and mood problems. This type of pain can occur in isolation, as often occurs in conditions such as fibromyalgia or tension-type headache, or as part of a mixed-pain state in combination with ongoing nociceptive or neuropathic pain, as might occur in chronic low back pain. It is important to recognise this type of pain, since it will respond to different therapies than nociceptive pain, with a decreased responsiveness to peripherally directed therapies such as anti-inflammatory drugs and opioids, surgery, or injections.

## 2010/11/16 ACR criteria for FM

#### Fibromyalgia Symptoms (Modified ACR 2010 Fibromyalgia Diagnostic Criteria)

1. Please indicate below if you have had pain or tenderness over the past 7 days in each of the areas listed below. Check the boxes in the diagram below for each area in which you have had pain or tenderness. Be sure to mark right and left sides



2. Using the following scale, indicate for each item your severity over the past week by checking the appropriate box.

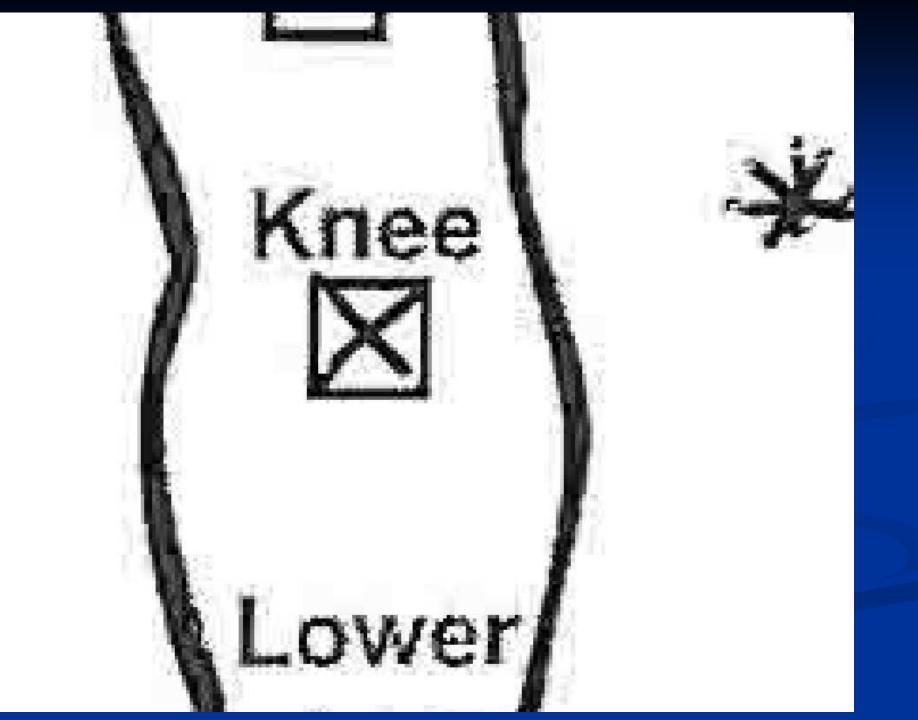
#### No problem

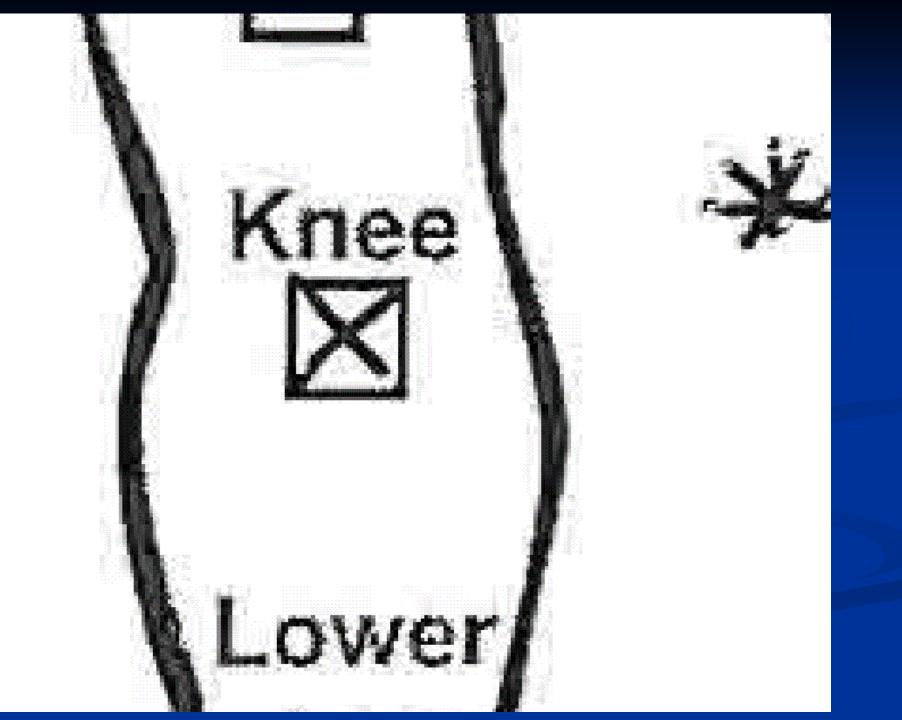
Slight or mild problems: generally mild or intermittent Moderate: considerable problems; often present and/or at a moderate level

Severe: continuous, life-disturbing problems

	No problem	Slight or mild	Moderate	Severe
a. Fatigue				
<ul> <li>b. Trouble thinking or remembering</li> </ul>				
<ul> <li>c. Waking up tired (unrefreshed)</li> </ul>				
During the past 6 months have you had any of the following symptoms?     No Yes				
a. Pain or cramps in lov	n 🗆			
b. Depression				
c. Headache				
4. Have the symptoms in questions 2-3 and pain been present at a similar level for <u>at least 3 months</u> ? No □ Yes □				
<ol> <li>Do you have a disorder that would otherwise explain the pain?</li> <li>No □ Yes □</li> </ol>				

- Wolfe et. al. *Arthritis Rheum.* Jun 15 2009;61(6):715-716. 2. Wolfe et. al.
- J Rheumatol. Feb 1 2011. 3. Clauw DJ. JAMA, 2014.



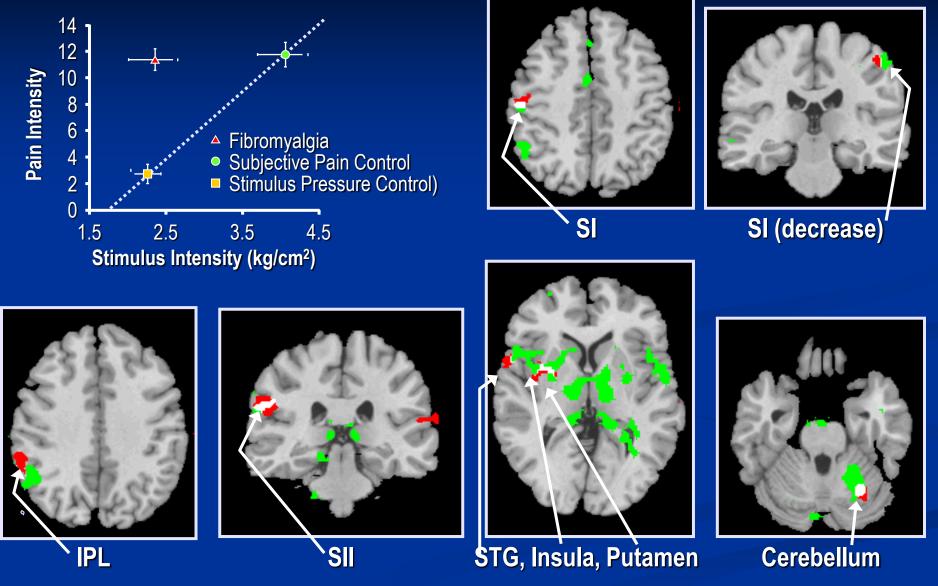


### What Do We Know About the Central Amplifier?

## Pathophysiology of centralized pain states— understading the "central amplifier"

- Most patients display augmented pain and sensory processing on quantitative sensory testing and functional neuroimaging<sup>1,3</sup>
- Manifest by increased connectivity to pro-nociceptive brain regions and decreased connectivity to anti-nociceptive regions<sup>2,3</sup>
- These abnormalities are being driven by imbalances in concentrations of CNS neurotransmitters that control sensory processing, sleep, alertness, affect, memory<sup>3,4</sup>
- Autonomic, HPA, and peripheral abnormalities likely play a prominent role in some individuals

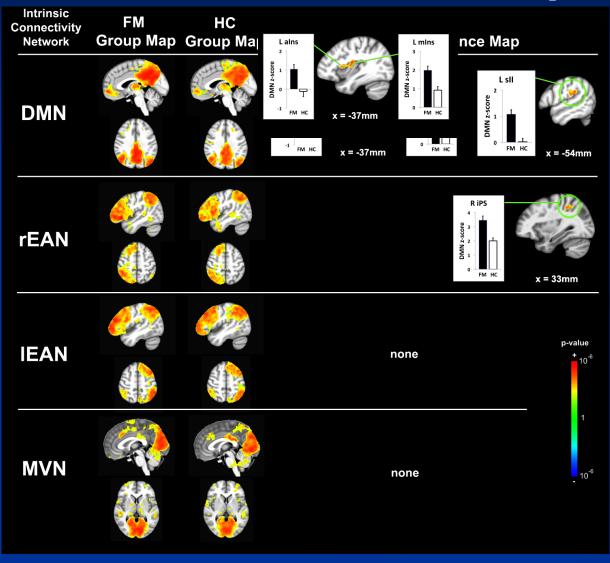
### fMRI in Fibromyalgia



STG=superior temporal gyri; SI=primary somatosensory cortex SII=secondary somatosensory cortex; IPL=inferior parietal lobule.

Gracely. Arthritis Rheum. 2002;46:1333-1343.

## Intrinsic Brain Connectivity is Altered in FM patients



• In FM, DMN and rEAN show greater intrinsic connectivity within component DMN (PCC), and rEAN (iPS) as well as limbic (insula), and sensorimotor (SII) regions outside conventional network boundaries.

•All FM vs. HC differences driven by greater connectivity for FM patients

## Changes in size and shape of brain regions indicate CNS neuroplasticity in chronic pain

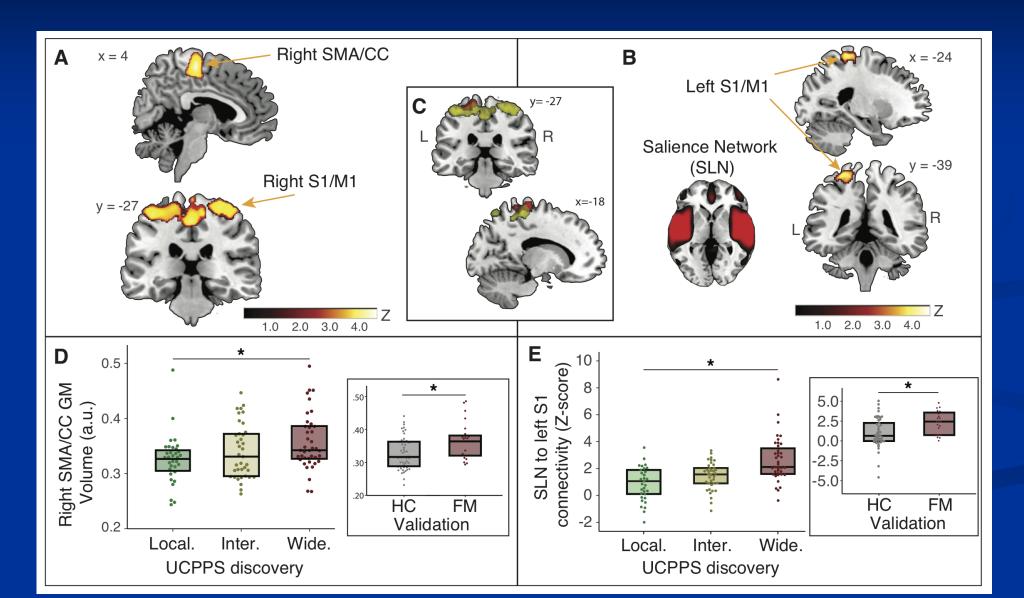
- Apkarian<sup>1</sup> was first to show that chronic pain may be associated with decrease of size of brain areas involved in pain processing
- More recently seen in virtually all other chronic pain states including headache, <sup>2</sup> IBS, <sup>3</sup> FM<sup>4</sup>
- May be partially due to co-morbid mood disturbances<sup>6</sup>
- Data from NIH MAPP network suggests *increase* in size of and connectivity to S1 may represent neural signature for widespreadedness of pain

<sup>1.</sup> Apkarian et al. *J Neurosci*. 2004;24:10410-5. 2. Schmidt-Wilcke et al. *Pain*. 2007;132 Suppl 1:S109-16.

<sup>3.</sup> Davis et al. Neurology. 2008;70:153-4. 4. Kuchinad et al. J Neurosci. 2007;27:4004-7.

<sup>5.</sup> Chen et al. Psychiatry Res. 2006;146:65-72. 6. Hsu et. al. Pain. Jun 2009;143(3):262-267. 7. Kutch et. al. IASP 2016

## Increased Gray Matter Volume in and Connectivity to Sensory Cortex In Widespread Pain





AN OFFICIAL JOURNAL OF THE AMERICAN COLLEGE OF RHEUMATOLOGY



Original Article

#### Heritability of the Fibromyalgia Phenotype Varies by Age

Diptavo Dutta, Chad M. Brummett, Stephanie E. Moser, Lars G. Fritsche, Alexander Tsodikov, Seunggeun Lee, Daniel J. Clauw, Laura J. Scott , ... See fewer authors ,

First published: 17 November 2019 | https://doi.org/10.1002/art.41171 | Citations: 3

#### Results

Overall, the FM score had an estimated heritability of 13.9% (SE 2.9%) ( $P = 1.6 \times 10^{-7}$ ). Estimated FM score heritability was highest in individuals  $\leq 50$  years of age (23.5%; SE 7.9%) ( $P = 3.0 \times 10^{-4}$ ) and lowest in individuals  $\geq 60$  years of age (7.5%; SE 8.1%) (P = 0.41). These patterns remained the same when we analyzed FM as a case–control phenotype. Even though women had an  $\sim 30\%$  higher average FM score than men across age categories, FM score heritability did not differ significantly by sex.

#### Conclusion

Younger individuals appear to have a much stronger genetic component to the FM score than older individuals. Older individuals may be more likely to have what was previously called "secondary FM." Regardless of the cause, these results have implications for future genetic studies of FM and associated conditions.



## Neurobiological antecedents of multisite pain in children

Chelsea M. Kaplan<sup>a,\*</sup>, Andrew Schrepf<sup>a</sup>, Ishtiaq Mawla<sup>b</sup>, Eric Ichesco<sup>a</sup>, Kevin F. Boehnke<sup>a</sup>, Adriene Beltz<sup>c</sup>, Emily Foxen-Craft<sup>d</sup>, Michael P. Puglia II<sup>a</sup>, Alexandre Tsodikov<sup>e</sup>, David A. Williams<sup>a,b,f,g</sup>, Afton L. Hassett<sup>a</sup>, Daniel J. Clauw<sup>a,f,g</sup>, Steven E. Harte<sup>a,b,g</sup>, Richard E. Harris<sup>a,b,g</sup>

#### **Abstract**

Altered brain structure and function is evident in adults with multisite chronic pain. Although many such adults trace their pain back to childhood, it has been difficult to disentangle whether central nervous system alterations precede or are consequences of chronic pain. If the former is true, aberrant brain activity may identify children vulnerable to developing chronic pain later in life. We examined structural and functional brain magnetic resonance imaging metrics in a subset of children from the first 2 assessments of the Adolescent Brain and Cognitive Development Study. Children (aged 9-10) who were pain free at baseline and then developed multisite pain 1 year later (n = 115) were matched to control children who were pain free at both timepoints (n = 230). We analyzed brain structure (cortical thickness and gray matter volume) and function (spontaneous neural activity and functional connectivity). Results were deemed significant at the cluster level P < 0.05 false discovery rate corrected for multiple comparisons. At baseline, children who subsequently developed multisite pain had increased neural activity in superior parietal /primary somatosensory and motor cortices and decreased activity in the medial prefrontal cortex. They also exhibited stronger functional connectivity between the salience network, somatosensory, and default mode network regions. No significant differences in the brain structure were observed. Increased neural activity and functional connectivity between brain regions, consistent to that seen in adults with chronic pain, exist in children before developing multisite pain. These findings may represent a neural vulnerability to developing future chronic pain.

Keywords: Multisite pain, Children, fMRI, Functional connectivity, Risk factors

# Sub-threshold FM is Highly Predictive of Surgery and Opioid Non-responsiveness in Patients Undergoing Arthroplasty and Hysterectomy

- Primary hypothesis of studies is the measures of centralized pain in OA (FMness) will predict failure to respond to arthroplasty and hysterectomy
- Extensive preoperative phenotype using validated selfreport measures of pain, mood, and function
- Two outcomes of interest:
  - Postoperative opioid consumption
  - Pain relief from procedure at 6 months

<sup>1.</sup> Brummett, C.M., et al., Anesthesiology, 2013. 119(6): p. 1434-43.

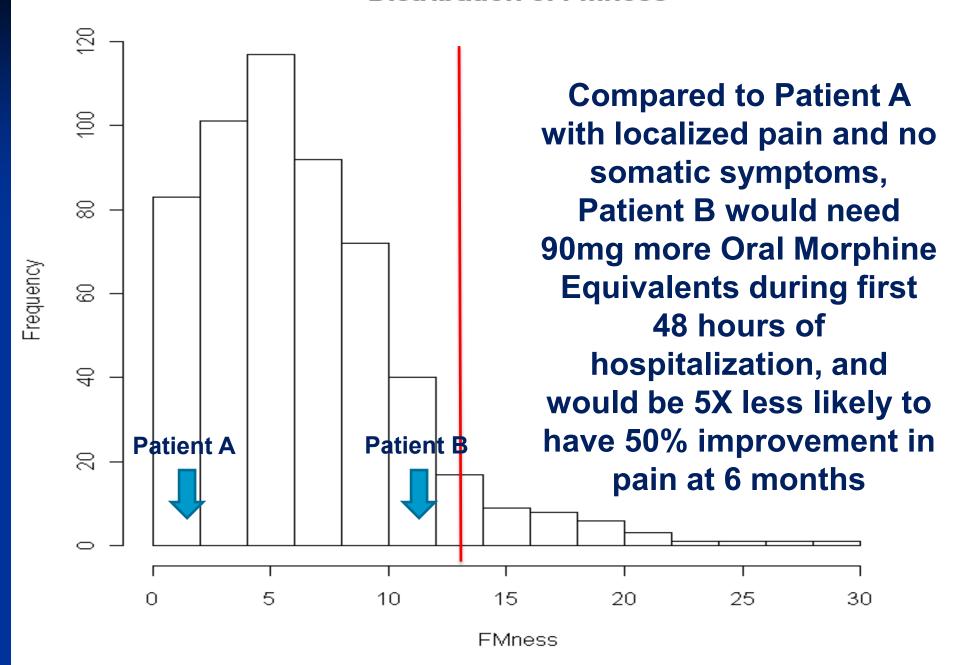
<sup>2.</sup> Brummett, C.M., et al., Arthritis Rheumatol, 2015. 67(5):1386-94.

<sup>3.</sup> Janda, A.M., et al., Anesthesiology, 2015. 122(5): p. 1103-11.

## Each one point increase in fibromyalgianess led to:

- 9 mg greater oral morphine requirements during acute hospitalization (8mg greater when all individuals taking opioids as outpatients excluded)
- 20 25% greater likelihood of failing to respond to knee or hip arthroplasty (judged by either 50% improvement in pain or much better or very much better on patient global)
- These phenomenon were linear across entire scale up to a score of approximately 18 and equally strong after individuals who met criteria for FM were excluded
- This phenomenon was much stronger than and largely independent of classic psychiatric factors

#### Distribution of FMness



## Incidence and predictors of persistent pelvic pain following hysterectomy in women with chronic pelvic pain

Sawsan As-Sanie, MD, MPH; Sara R. Till, MD, MPH; Andrew D. Schrepf, PhD; Kendall C. Griffith, MD; Alex Tsodikov, PhD; Stacey A. Missmer, ScD; Daniel J. Clauw, MD; Chad M. Brummett, MD

**BACKGROUND:** Chronic pelvic pain is a debilitating problem that afflicts 15% to 20% of women in the United States. Although more than 200,000 hysterectomies are performed annually for the treatment of chronic pelvic pain, previous studies indicate that 1 in 4 women undergo the discomfort and morbidity of hysterectomy without the relief of pain. The factors that predict treatment failure remain poorly characterized.

**OBJECTIVE:** To describe the incidence of persistent pelvic pain 6 months following hysterectomy in women with chronic pelvic pain and determine whether a simple, self-reported measure of central sensitization is associated with a greater risk of persistent pelvic pain following hysterectomy.

**STUDY DESIGN:** We conducted a prospective, observational cohort study of women undergoing hysterectomy at an academic tertiary care center for a benign indication. Patients with preoperative chronic pelvic pain, defined as average pelvic pain  $\geq 3$  on a 0 to 10 numeric rating scale for >3 months before hysterectomy, were included in this analysis. The patients completed validated assessments of pain, anxiety, depression, and centralized pain (using the 2011 Fibromyalgia Survey Criteria, 0–31 points) preoperatively and 6 months after hysterectomy. The demographic information, surgical history, intrapperative findings, and surgical pathology were abstracted from the

**RESULTS:** Among 176 participants with pelvic pain before hysterectomy, 126 (71.6%) were retained at 6 months, and 15 (11.9%) reported persistent pelvic pain. There was no difference in age (P=.46), race (P=.55), average pain severity during menses (P=.68), average overall pelvic pain (P=.10), or pain duration (P=.80) in those with and without persistent pelvic pain. Whereas intraoperative findings of endometriosis (P=.05) and uterine fibroids (P=.03) were associated with a higher incidence of persistent pain on univariate analysis, the surgical route (P=.46), pelvic adhesions (0.51), uterine weight (P=.66), and adenomyosis on histopathology (P=.27) were not related to the risk of persistent pain. Higher preoperative centralized pain scores (P=.01) but not depression (P=.64) or anxiety (P=.45) were more common in women with persistent pelvic pain. Multivariate logistic regression adjusting for age, preoperative pain severity, anxiety, depression, and operative findings of endometriosis and fibroids indicated that every 1-point increase in centralized pain before hysterectomy was associated with a 27% increase in the odds of persistent pelvic pain (odds ratio, 1.27; 95% confidence interval, 1.03—1.57) 6 months after surgery.

**CONCLUSION:** Although the majority of women with chronic pelvic pain report considerable improvement in pain following hysterectomy, higher degrees of centralized pain before hysterectomy is a robust pre-

### Chronic Overlapping Pain Conditions



RESEARCH EDUCATION TREATMENT ADVOCACY



The Journal of Pair, Vol 17, No 9 (September), Suppl. 2, 2016: pp T93-T107

Available online at www.ipain.org and www.sciencedirect.com

Overlapping Chronic Pain Conditions: Implications for Diagnosis and Classification



William Maixner, \*,† Roger B. Fillingim,‡ David A. Williams,§ Shad B. Smith, \*,† and Gary D. Slade \*,¶,||

-Term defined by the National Institutes of Health  $\sim 2013$ 

-Conditions likely to co-exist sharing neurobiological underpinnings

-Conditions predominantly (or solely) affecting women

-Any number and combination of conditions is possible

-Several conditions can develop at once or gradually over years

US COPCs<sup>1</sup> **Prevalence Irritable Bowel Syndrome** 44 Million Temporomandibular Disorder 35 Million **Chronic Low Back Pain** 20 Million **Interstitial Cystitis / BPS;** 8 Million chronic prostatitis Migraine Headache 7 Million **Tension Headache** 7 Million **Endometriosis (Painful)** 6 Million Vulvodynia 6 Million **Fibromyalgia** 6 Million Myalgic Encephalopathy / CFS 4 Million

<sup>1</sup>Veasley, C. et al (2015). White paper from the *Chronic Pain Research Alliance*.

<sup>\*</sup>Center for Pain Research and Innovation, \*Department of Dental Ecology, \*Department of Epidemiology, University of North Carolina at Chapel Hill, Chapel Hill, North Carolina.

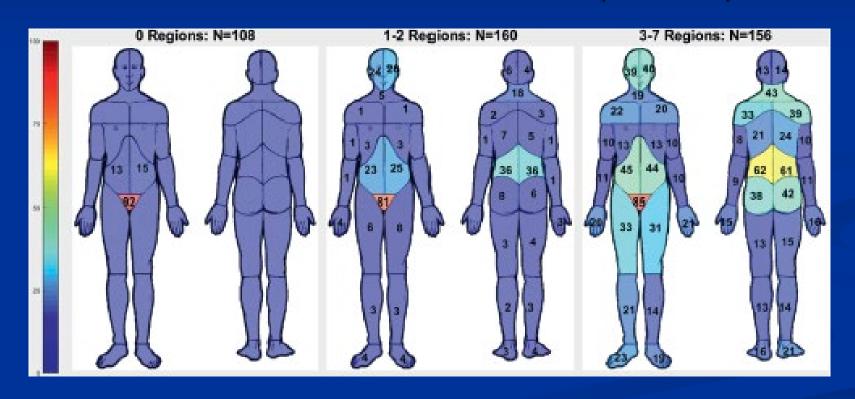
<sup>&</sup>lt;sup>1</sup>Center for Translational Pain Medicine, Department of Anesthesiology, Duke University, Durham, North Carolina.

<sup>&</sup>lt;sup>†</sup>Pain Research and Intervention Center of Excellence, University of Florida, Gainesville, Florida.

<sup>&</sup>lt;sup>§</sup>Chronic Pain and Fatigue Research Center, Department of Anesthesiology, University of Michigan, Ann Arbor, Michigan.

### Can you have just 1 COPC and still have it be a COPC?

#### Pain Distribution in IC (MAPP)



Tertiles of body pain distribution:

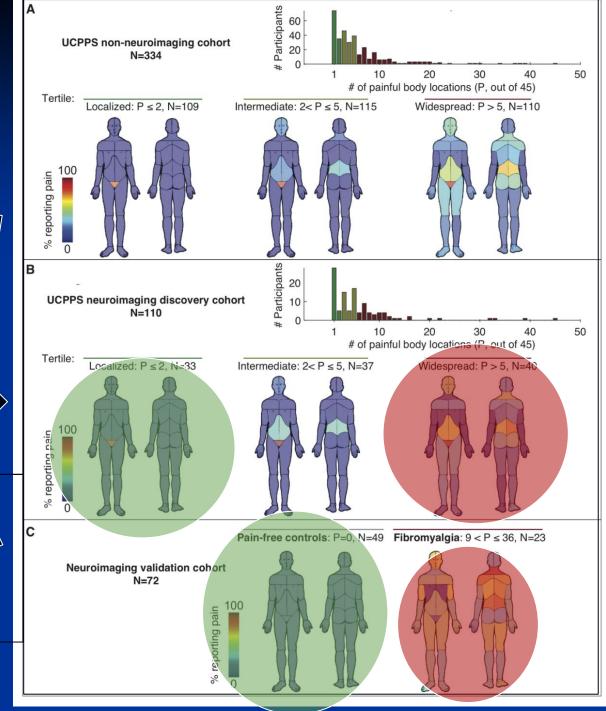
local, intermediate,



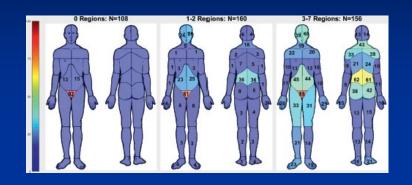
Non-Neuroimaging (N=334)

Neuroimaging (N=110)

Validation FM (N=23) HC (N=49)



### Similar Pattern in other COPCs



Condition	Index only	1-2 regions + index	>= 3 + index
IC (n=424)	25%	38%	37%
C. Migraine (n=1601)	29%	26%	45%
Mixed COPCs (n=9169)	25%	7	5%

## Two methods for assessing COPCS for clinical research

- ICD-10 Computable Phenotype
- The Chronic Overlapping Pain Conditions Screener (COPCS)





#### Original Reports

### ICD-10 Codes for the Study of Chronic Overlapping Pain Conditions in Administrative Databases



Andrew Schrepf, \* Vy Phan, † J. Quentin Clemens, † William Maixner, † David Hanauer, † and David A. Williams\*

\*Chronic Rain and Fatigue Research Center, Department of Anesthesiology, University of Michigan Medical School, Ann Arbor, Michigan, \*Trinity College-Hartford, Hartford, Connecticut, \*Department of Urology, University of Michigan Medical School, Ann Arbor, Michigan, \*Center for Translational Pain Medicine, Department of Anesthesiology, Duke University School of Medicine, Durham, North Carolina, \*Department of Pediatrics, University of Michigan Medical School, Ann Arbor, Michigan

FM	IBS	IC/BPS, CP	VVD	MI
• M79.7	• K58.[0,1,2,8,9]	<ul><li>N30.[1,3]</li><li>N41.1</li></ul>	• N94.8[10,18,19]	• G43.xx
ТТН	TMD	cLBP	ME/CFS	ENDO+pain
• G44.[201,209,211,2 19,221,229]	<ul> <li>M26.[60,62,63]</li> <li>S03.0xxA</li> </ul>	• M54.[5,40,41,42,89]	• R53.82	<ul> <li>N80.xxx +R10.2,</li> <li>N80.xxx +N94.[4,5,6,10,11, 12,19]</li> </ul>

	FM	IBS	TMD	UCPPS	ENDO	VVD	cLBP	cTTH	MI	CFS
FM										
IBS	10.18									
TMD	5.64	3.70								
UCPPS	9.91	9.10	4.75							
ENDO	4.06	5.05	1.87	18.62						
VVD	3.14	3.97	1.85	24.99	15.56					
cLBP	5.29	2.29	1.24	2.34	2.30	1.20				
сТТН	2.43	1.58	2.64	1.94	1.25	N/A	3.36			
MI	5.27	3.30	6.13	3.29	3.21	1.63	1.99	4.27		
CFS	6.07	2.90	1.48	2.78	1.86	1.19	1.75	1.82	2.67	
diab neurop	2.60	1.66	N/A	0.86	N/A	N/A	2.08	0.51	1.06	1.18
COPD	3.14	1.78	0.89	1.05	0.54	0.50	1.92	0.71	1.11	1.29
chronic viral										
hepatitis	2.20	1.48	0.56	1.19	N/A	N/A	1.22	0.56	0.68	1.01

#### Odds ratio =

## Chronic Overlapping Pain Conditions Screener (COPC-S)





The Journal of Pain, Vol 25, No. 1 (January), 2024: pp 265–272

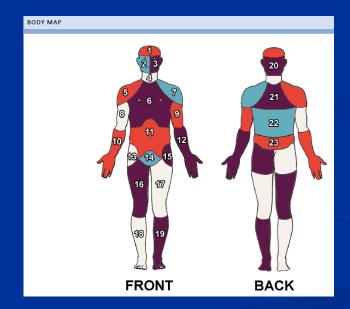
Available online at www.ipain.org and www.sciencedirect.com

The Chronic Overlapping Pain Condition Screener



Andrew Schrepf,\* William Maixner,<sup>†</sup> Roger Fillingim,<sup>‡</sup> Christin Veasley,<sup>§</sup> Richard Ohrbach,<sup>¶</sup> Shad Smith,<sup>†</sup> and David A. Williams\*

https://copcscreener.com



Rep	ort
Condition	Class
FM	Y/N
IBS	Y/N
TMD	Y/N
MI	Y/N
TTH	Y/N
IC/CP	Y/N
cLBP	Y/N
ME/CFS	Y/N
ENDO-pain	Y/N
VVD	Y/N
Total	x/10

### Stretch Break

### Music about Pain

- Pain Three Days Grace
- Pain Tupac Shakur
- Pain Pusha T
- Pain Jimmy Eats World
- Pain War on Drugs
- Pain Hollywood Undead
- Pain The Used

- Hurt Johnny Cash
- Everybody Hurts R.E.M.
- Hurt Nine Inch Nails

- King of Pain Police
- No More Pain 2Pac
- Feel No Pain Sade
- The Sweetest Pain Dexter Wansel
- I Feel Pain A Boogie wit Da Hoodie
- I Can't Stand the Pain James Brown
- Physical Pain Future Island

- Give Me Novocain Green Day
- Pain Killer Judas Priest
- The Drugs Don't Work The Verve

## Most Pain Care Visits occur within Primary Care



Peterson K, et al.. VA ESP Project #09-199; 2017.

## Primary Care Physicians Receive Little Training in Pain Management

- 80% of American Medical Schools have no formal pain education
- Those that do, report 5 or fewer hours
  - Emphasis of education is often cellular and subcellular rather than interpersonal or social in nature
- Only 34% of physicians reported feeling comfortable treating chronic pain
  - Only 1% found it a satisfying practice

#### Pain Medicine Versus Pain Management: Ethical Dilemmas Created by Contemporary Medicine and Business

John D. Loeser, MD\*† and Alex Cahana, MD, PhD\*†

Biomedical Model
Interventional
Pain Medicine

Biopsychosocial model
Interdisciplinary
Pain Management

- Procedure Driven
- Focus on curing/fixing

Patient is passive recipient

- Focus on multidisciplinary teams
- Focus on pain management

Patient is active participant

## How good is our black bag for treating chronic pain?

Treatment	Impact on Chronic Pain
Long term opioids	32% reduction
Pain drugs generally (across classes)	30% - 40% get 40% - 50% relief
Spinal fusion	75% still have pain
Repair herniated disk	70% still have pain
Repeat Surgery	66% still have pain
Spinal cord stimulators	61% still in pain after 4 yrs. average pain relief 18% across studies

## Are Invasive Procedures Effective for Chronic Pain? A Systematic Review

Wayne B. Jonas, MD,\* Cindy Crawford,<sup>†</sup> Luana Colloca, MD, PhD,<sup>‡</sup> Levente Kriston, PhD,<sup>§</sup> Klaus Linde, MD, PhD,<sup>¶</sup> Bruce Moseley, MD,<sup>∥</sup> and Karin Meissner<sup>|||</sup>\*\*

**Conclusions**. There is little evidence for the specific efficacy beyond sham for invasive procedures in chronic pain

Pain Medicine, 20(7), 2019, 1281–1293

doi: 10.1093/pm/pny154

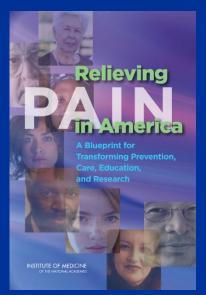
Advance Access Publication Date: 10 September 2018

Review Article



## Recommendations in Multiple Federal Documents

Self-Management, Evidence-Based, Patient-Centric, Multi-Modal Pain Care

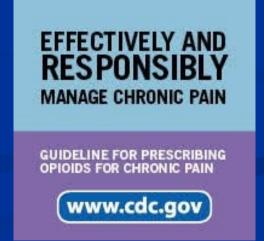












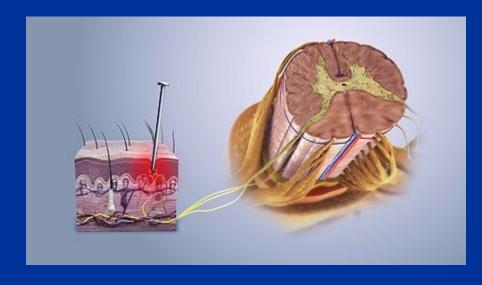
## What's a doctor to do?



## We Need to Approach Chronic Pain Differently

### Thinking Differently about Pain

- Damaged tissue and pain are not the same thing
- Sometimes they occur together, so they seem to be the causal
- Nociception provides bodily information that may or may not be interpreted as pain





Nociception

**PAIN** 

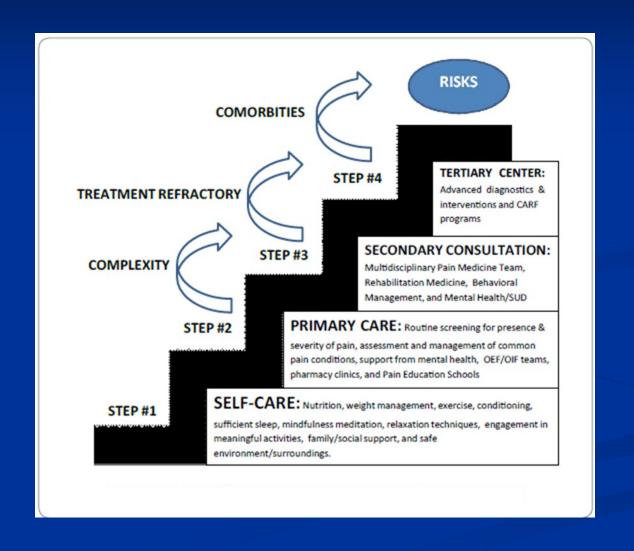
### Thinking Differently about Chronic Pain

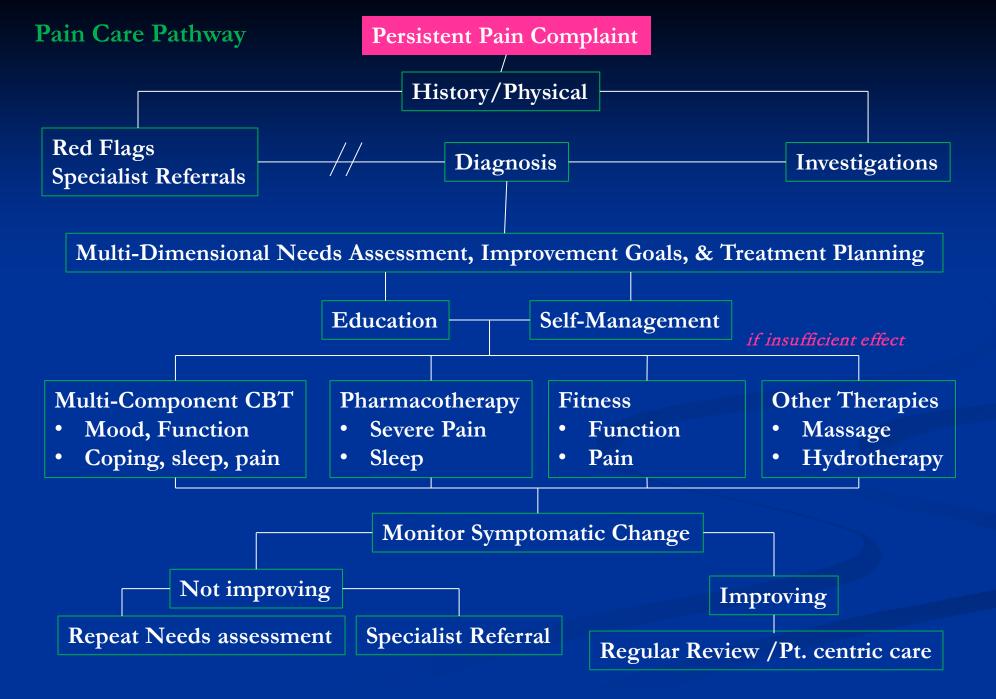
- Pain is a Perceptual Experience formed in the brain
  - Other perceptual experiences with flexible biological associations include the following:
    - hunger, itch, tickle, urinary urgency, orgasm

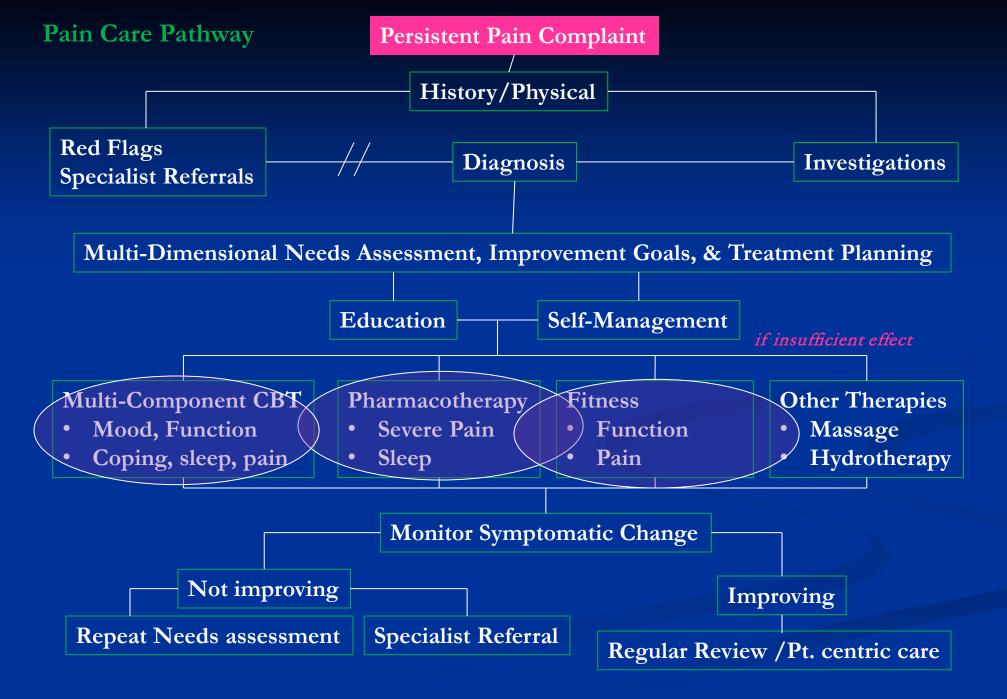
### Thinking Differently about Chronic Pain

■ Treating a perception requires different techniques than fixing damaged tissues

### VA's Stepped Care Model of Pain Management







### Treating Based on Mechanisms

Any combination may be present

1 11 y Convolvation in though to product			
	Peripheral (nociceptive)	Neuropathic	Centralized Pain
NSAIDs	+	-	_
Opioids	+	+	_
Surgery/ Injections	+	+	_
Tricyclics	+	+	+
SNRIs	+	+	+
Gabapentinoid	-	+	+
CBD	+	-	_
THC	_	+	+

Modified from Goldenberg et al. *JAMA*. 2004;292:2388-95.

### Nonpharmacological Therapies

## **Strong Evidence**

- Education
- Aerobic exercise
- Cognitive behavior therapy (ACT, CBT-i, empowered selfcare

## **Modest Evidence**

- Strength training
- Hypnotherapy, biofeedback, balneotherapy, yoga, Tai Chi
- Neuromodulation
- Acupuncture, chiropractic, manual and massage therapy

## Weak Evidence

Trigger point injections

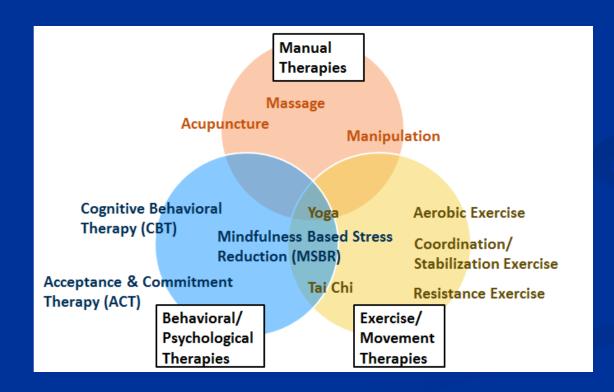
## No Evidence

Doing nothing

## Non-Pharmacological Pain Treatments in VHA

VA State of the Art Conference Nov. 2016: Evidence-based non-pharmacological approaches for MSK pain management

- Evidence to support CIH and conventional therapies.
- Provision of multi-modal therapies accessible from Primary Care.



### VHA Directive 1137: Advancing Complementary and Integrative Health (May 2017)

- List 1: Approaches with published evidence of promising or potential benefit.
  - Acupuncture
  - Massage Therapy
  - Tai Chi
  - Meditation
  - Yoga
  - Clinical Hypnosis
  - Biofeedback
  - Guided Imagery
- Chiropractic Care was approved as a covered benefit in VHA in 2004 and is part of VA whole health care.
- To be made available across the system, if recommended by the Veteran's health care team.

### Can we use diet/nutrition to treat chronic pain?

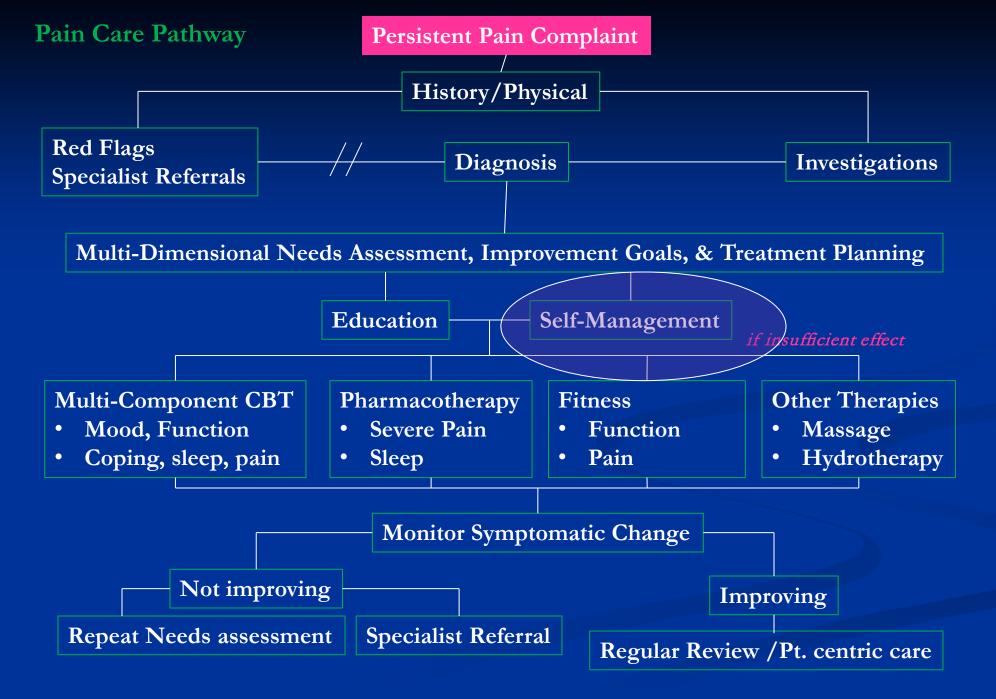


\*Chronic Pain and Fatigue Research Center, Department of Anesthesiology,†Department of Internal Medicine, ‡School of Public Health, University of Michigan, Ann Arbor, Michigan.

**Abstract:** Weight loss is known to improve pain localized to weight-bearing joints but it is not known how weight loss affects the spatial distribution of pain and associated somatic symptoms like fatigue. We sought to determine if weight loss using a low-calorie diet improves pain, affect, and somatic symptoms commonly associated with chronic pain conditions in an observational study. We also documented changes in inflammatory markers in serum before and after weight loss. Participants were 123 obese individuals undergoing a 12- to 16-week calorie restriction weight loss intervention. The spatial distribution of pain, symptom severity (eg, fatigue, sleep difficulties), depression, and total fibromyalgia scale scores were measured before and after weight loss. Pain (P = .022), symptom severity (P = .004), depression (P < .001), and fibromyalgia scores (P = .004) improved after weight loss; men showed greater improvement than women on somatic symptoms and fibromyalgia scores (both P < .01). Those who lost at least 10% of body weight showed greater improvement than those who lost <10%. Levels of the regulatory cytokine interleukin-10 increased after the intervention (P = .002). Weight loss may improve diffuse pain and comorbid symptoms commonly seen in chronic pain participants.

**Perspective:** This article presents the effect of a weight loss intervention on characteristics of chronic pain, including the spatial distribution of pain and comorbid somatic symptoms. Weight loss appeared to produce larger improvements in somatic symptoms for men.

© 2017 by the American Pain Society



### How to ERASE S.P.A.C.E.

**E**motions

Reflections

Actions

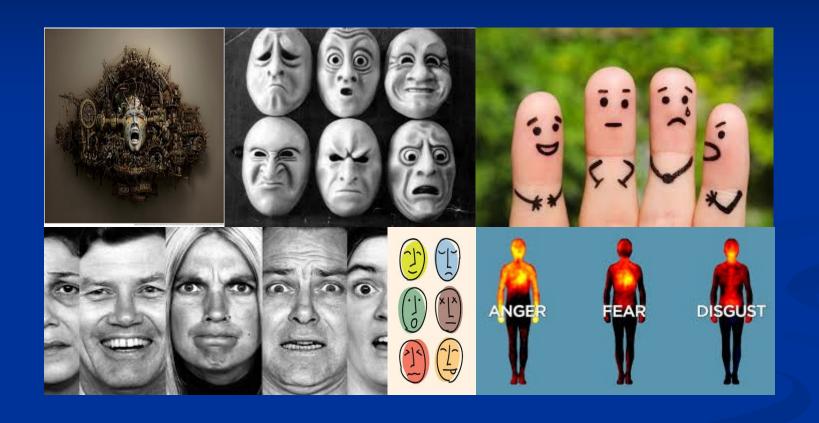
Sleep

**Environment** 

Sleep, Pain, Affect, Cognitive changes, Energy deficits

### ERASE

## **E**motions



Altering pain perception through Emotions

Patients do not need to be mentally ill to have chronic pain



# Approaches to Resolve Negative Affect Influencing Chronic Pain



Emotional Awareness and Expression Therapy (EAET)

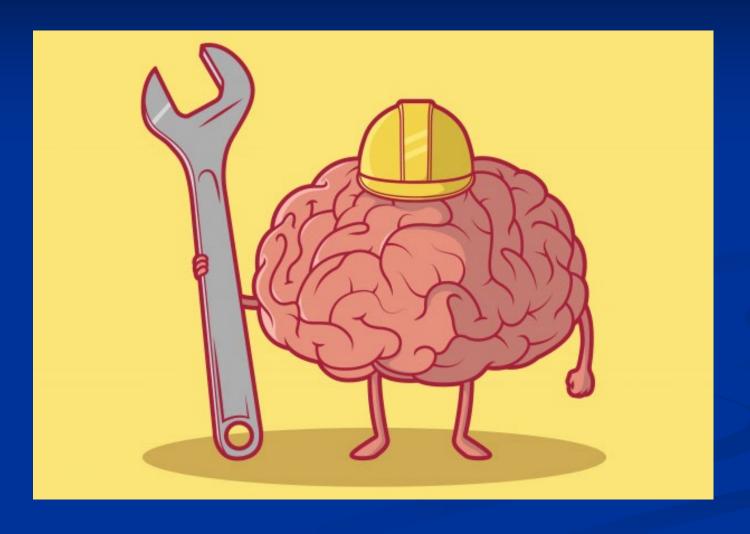


Pleasant Activity Scheduling



Traditional Psychotherapy

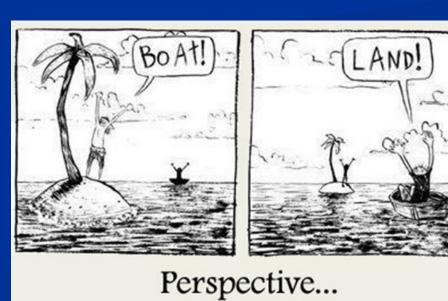
## Reflections



Using Cognition to alter pain perceptions

## Reframing





### The Relaxation Response







Visual Imagery



Meditation



Biofeedback

## Actions



Using <u>Behavior</u> to alter pain perceptions and provide a foundation of wellness

### Exercise

- Multiple reviews and metaanalyses, and professional society guidelines recommend exercise and physical activity for the treatment of chronic pain and fatigue
- Increase Fitness
- Increase Function





## **Lifestyle Physical Activity**





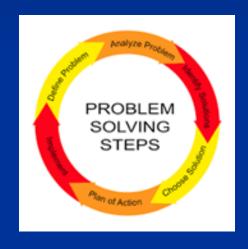


## **Pacing for Energy Efficiency**





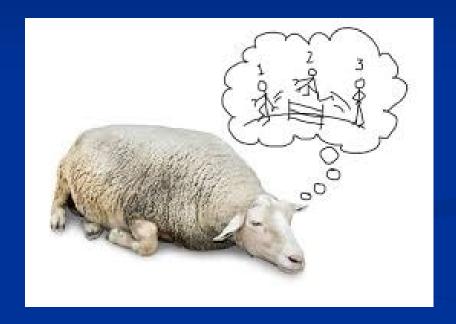
## Problem Solving / Goal Setting





### ERASE

## Sleep



Altering Pain via Sleep

## Behavioral and Sleep Hygiene Skills

### **Timing**

Regular bed time/wake time

### **Sleep Behavior**

Get in bed only when sleepy Use bed for sleep Get up after 15' if no sleep

### **Thermal Tips**

Decline in core temp signals sleep Exercise, warm bath before bed

### **Environment**

Steady room temperature Keep room dark

### **Ingestion**

Decrease nicotine
Decrease Caffeine
Alcohol interferes with sleep
Light snack is recommended

### **Mental Control**

Effort will not produce sleep Avoid mental stimulation Seek mental quiescence

### ERASE

## **Environment**



Using the Environment to alter pain perceptions and provide a foundation of wellness

## Social Challenges



Dr. -Patient



Family



Friends



Employer and co-workers

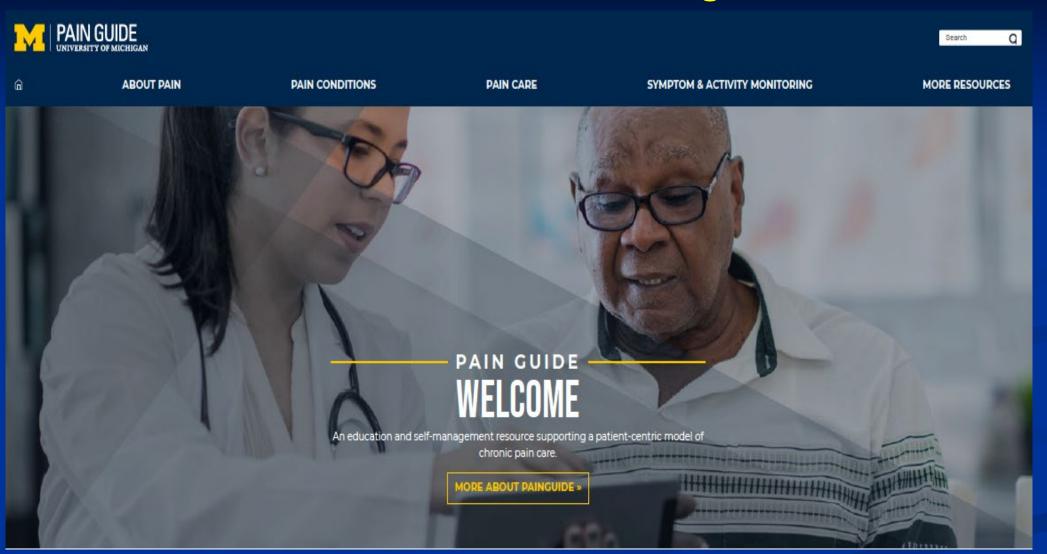
## **Physical Challenges**







### Web-based self-management



http://PainGuide.com

#### **Self Care**



#### Exercise

Exercise, when done safely, can benefit you physically and mentally It helps prevent deconditioning of muscles which is often associated with more pain. Studies find that exercise is one of the most beneficial approaches to managing pain.



#### Relaxation

Teaching the body to relax can both diminish muscle tension and decrease stress. To work properly, regular practice is needed so that the body learns a rhythm of relaxation and can relax on your command. Less tension and less stress can lead to decreased pain intensity.

Learn more >



#### Communication skills

Conflictual social relationships with family, friends, doctors, and employers can make pain worse. Alternatively, these same relationships can be used constructively to make pain better. Communication skills can help make social relationship work in your favor.

Learn more >



#### Sprituality

The belief in something "bigger," "more powerful," or "more knowledgeable" than oneself has been key to many individuals being able to successfully deal with pain. Spirituality may refer to a specific religious belief or it can be any belief that provides a source of strength and comfort to the individual with pain

Learn more >



#### Pacing

People with pain often "over do" resulting in pain flare ups. Pacing can allow activities to get accomplished safely, without flare-ups, and in a manner that conserves energy (i.e., with less fatigue).

Learn more >



#### Reframing

What we think influences how we feel and how much pain we experience. Sometimes negative thoughts become automatic and make us feel worse. Learning to reframe our thinking in realistic terms that challenge negative automatic thinking can help diminish pain intensity.

Learn more >



#### **Nutrition &** supplements

Eating a healthy diet has many benefits for everyone; however there may be some specific benefits for pain sufferers. The examination of pain and diet is an emerging literature.

Read nutrition & supplements tips >



#### **Managing Emotions**

Emotions are integral to the production of pain. You cannot have pain without emotions. Thus anything we can do to alter the emotional content of one's brain will influence pain. Better management of stress can influence pain as well as engaging in pleasant activities. The pleasant activities will help diminish pain intensity.

Learn more >



#### Sleep

Pain and Sleep are closely related such that poor sleep can make pain worse. These are a number of behavioral sleep strategies that can be used to get a more refreshing night's sleep.

Learn more >



#### Acupressure

Like acupuncture, which uses needles, acupressure is an ancient treatment that uses the pressure of one's own finger on the skin so as to help rebalance the flow of energy through the body as a means of reducing symptoms such as pain.

Learn more >



#### Ergonomics/Posture

How you sit, stand, transition and lift can either make pain worse or allow you to function even with pain. This section offers help in optimizing how you interact with your environment in ways that don't exacerbate pain.

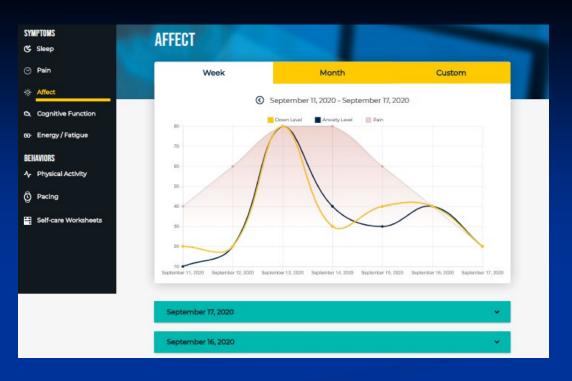
Learn more >



#### Resilience

We often focus on fixing what is broken but we can't lose sight of our personal strengths that help us get through challenging times. Finding our sources of resilience can be a valuable tool for reducing pain and living a quality-filled life.

Learn more >



### **PAIN CARE**

#### Self Care

### **Professional Care**

Medications

Therapies

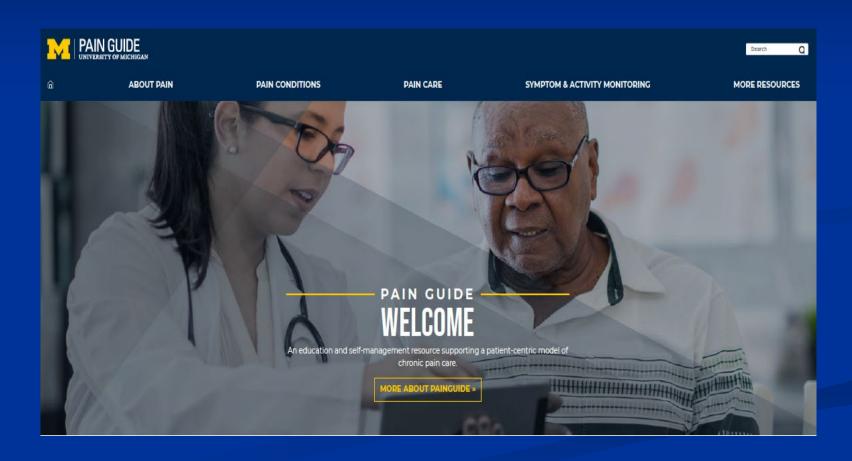
Devices

Procedures



# Pain Guide can serve as the foundation for guided interventions





### **Bottom Line**

- 1. Pain is not located in a body part. It is a perception and needs to be treated as a perception.
- 2. Taking time to just listen to the patient's story is a necessary part of pain treatment. You will be establishing rapport and identifying pain and non-pain treatment targets.
- 3. If you recommend self-management (exercise, relaxation, sleep hygiene etc.), ask about it with the same enthusiasm and regularity that you ask about drugs. Patients learn what you think is <u>really</u> important by what you ask about.

