

# UNRAVELING CHRONIC PAIN: DIAGNOSING AND TREATING FIBROMYALGIA



*Dana Harrell- Sanders  
"Do You See What I Feel?"*

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Northwell Health, Division of Rheumatology  
February 2019

## CME ACCREDITED UPDATES IN MEDICINE ELEARNING SERIES

**COURSE NAME:**

Medicine RSS eLearning Modules

**CME eLEARNING ACTIVITY NAME:**

Unraveling Chronic Pain: Diagnosing and treating Fibromyalgia

**PROGRAM DESCRIPTION, EDUCATIONAL GOAL AND RATIONALE:**

Evidence based guidelines are constantly changing and being updated for several core areas of Internal Medicine throughout the year. It is important for physicians to practice the most up-to-date standard of care in all specialties to promote patient health and well-being. Our series of lectures at the medicine regularly scheduled series promotes continuing education for the practicing internist and highlights important updates in medical practice in these core areas. Physicians in general practice often and do not have the time to keep themselves up-to-date with medical advances as they are busy seeing patients in the clinical setting. The Medicine Regularly Scheduled Series gives these physicians the opportunity to learn these advances in an academic setting.

## CME ACCREDITED UPDATES IN MEDICINE ELEARNING SERIES

### TARGET AUDIENCE:

Physician Partners and Premium Network  
community-based providers

### LEARNING OBJECTIVES:

- To recognize Fibromyalgia (FM): primary and secondary.
- To differentiate mechanism of acute vs. chronic pain.
- Identify and explain the evidence to support a pathophysiologic basis for FM.
- State the most significant recommendations for treatment.
- Identify resources that can be used to help patients that live with FM and recognize need to refer when necessary.

## CME ACCREDITED UPDATES IN MEDICINE ELEARNING SERIES

### FACULTY PRESENTER/AUTHOR:

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Long Island Jewish Medical Center

## CME ACCREDITED UPDATES IN MEDICINE ELEARNING SERIES

### ACCREDITATION:

Northwell Health is accredited by the Accreditation Council for Continuing Medical Education to provide Continuing Medical Education for physicians.

### CREDIT DESIGNATION:

Northwell Health designates this Continuing Medical Education activity for a maximum of **1** *AMA PRA Category I credits*<sup>TM</sup>. Physicians should only claim credit commensurate with the extent of their participation in the activity

### METHOD OF PHYSICIAN PARTICIPATION:

#### To receive credit the participants must:

Read/view the entire educational activity.

Input name and credentials to gain CME credit.

## **CME ACCREDITED UPDATES IN MEDICINE ELEARNING SERIES**

**COURSE HOST:**

Department of Medicine  
Northwell Health

**ESTIMATED TIME TO COMPLETE ACTIVITY:**

90 minutes

**ACKNOWLEDGEMENT OF COMMERCIAL SUPPORT:**

An announcement of program support will be made to all attendees at the beginning of each educational activity.

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### **FACULTY DISCLOSURES:**

Drs. Sandy Balwan, George Boutis, John Raimo and Sean LaVine have nothing to disclose. Dr. Stamatoukas disclosures are on the next slides.

**RELEASE DATE:** 2/28/19

**REVIEW DATE:** 2/28/19

**PROGRAM EXPIRATION:** 7/30/19

# Disclosures

- Pfizer Medical Outcome Specialist



- Recognize fibromyalgia (FM)
  - Primary: no other explanation for chronic pain
  - Secondary: outcomes impacted by presence of FM
- Differentiate mechanism of acute versus chronic pain
- Explain the evidence to support a pathophysiologic basis for FM
- State the most significant recommendations for treating FM
  - Non opioid management
  - Maintain work, even if needs to be adjusted-
- Identify resources that can be used to help patients live with FM and recognize need to refer when necessary

## Objectives

- Arnold, L. M., et al. (2016). Fibromyalgia and chronic pain syndromes: A white paper detailing current challenges in the field. *Clin J Pain*; 32(9): 737-746.
- *Atzeni, F. (2019). Review: One year in review 2019: Fibromyalgia. Clin & Exp Rheum; 37 (Suppl. 116): S3-S10. \**
- Clauw, D. (2015). Diagnosing and treating chronic musculoskeletal pain based on mechanisms. *Best Practice & Research Clinical Rheumatology*; 29(1): 6-19.
- Dowell, D., Haegerich, R., & Chou, R. (2016). CDC Guideline for prescribing opioids for chronic pain- United States, 2016. *JAMA*; 315(15): 1624-1645.
- Fitzcharles, M.A. (2013). 2012 Canadian guidelines for the diagnosis and management of fibromyalgia syndrome: Executive summary. *Pain Res Manag*; 18(3): 119-126.
- Hauser, W., & Fitzcharles, M. (2019). Facts and myths pertaining to fibromyalgia. *Dialogues Clin Neurosci*; 20(1): 53–56.
- *Harris, R. & Clauw, D. 2006. How do we know that the pain in fibromyalgia is real? Current Pain and Headache Reports; 10(6): 403-407. \**
- Lopez-Sola, M., et al. (2017). Towards a neurophysiological signature for Fibromyalgia. *Pain*; 158(1): 34-47.
- Macfarlane GJ et al. (2016). EULAR revised recommendations for the management of fibromyalgia. *Ann Rheum Dis*; 0: 1-11. doi:10.1136/annrheumdis-2016-209724.
- F. Wolfe et al. (2016). 2016 Revisions to the 2010/2011 fibromyalgia diagnostic criteria. *Seminars in Arthritis and Rheumatism* 46 (2016) 319–329

## Key Evidence Based Articles

- <https://www.youtube.com/watch?v=STmQm-nHS08>



*Art courtesy of PainExhibit.org*

“Crucified by Pain”

June 28th, 2013 by Dana Harrell-Sanders

This painting represents those that suffer from fibromyalgia and chronic fatigue. Sometimes it's hard to put into words what those of us with these syndromes feel. I hope my art expresses it to those that don't understand.

**WIDESPREAD PAIN**

**STIFFNESS**

**SLEEP DISTURBANCE**

**COGNITIVE DIFFICULTIES**

**MOOD DISTURBANCE**

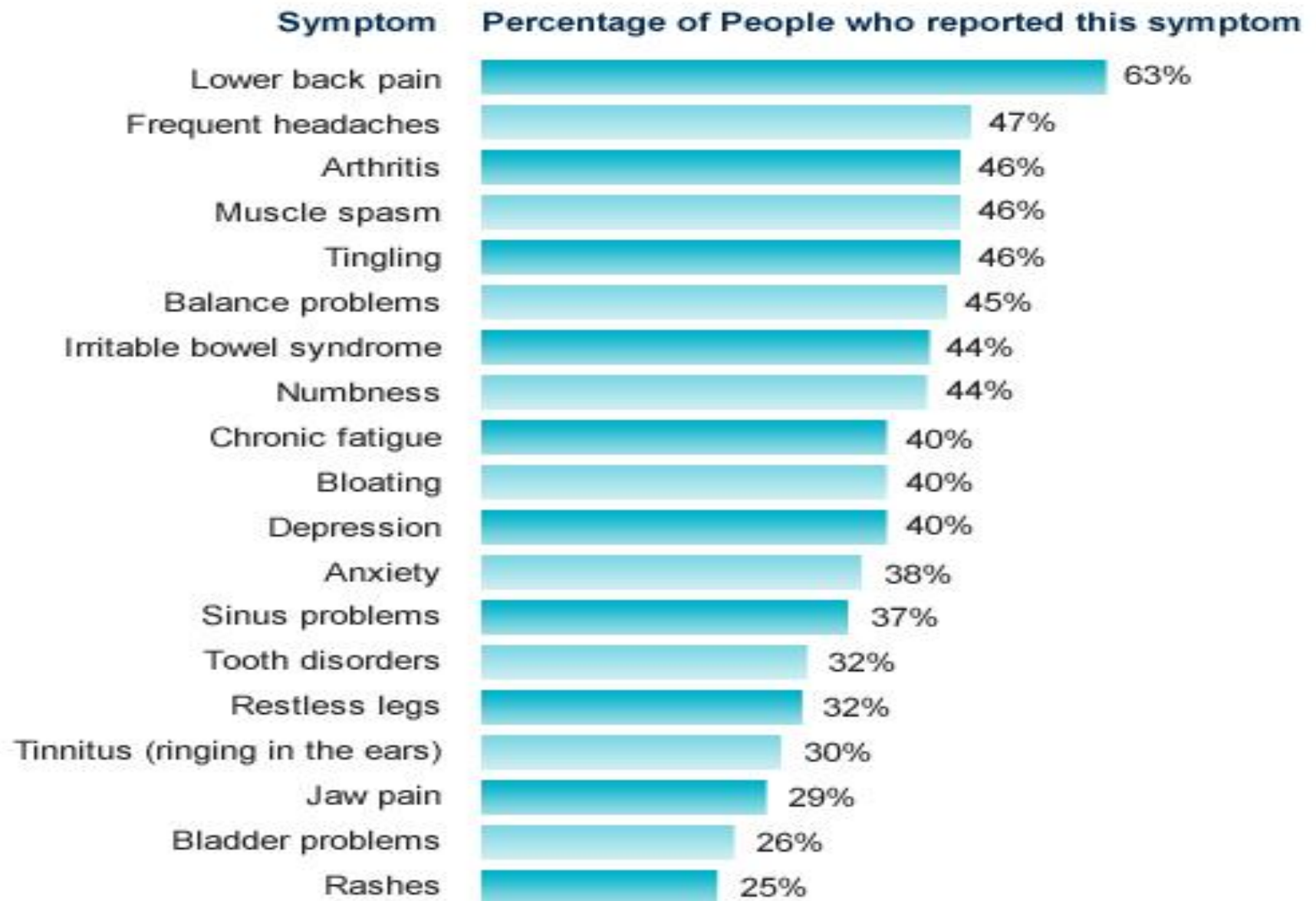
Here's what it feels like

# Fibromyalgia

- “There has to be something wrong... you can't feel this bad”
- “ I hurt all over”
- “It feels like I always have the flu”
- “I'm always so tired”
- “I can't fall asleep and when I do, I wake up all the time”
- “I have numbness and tingling sometimes here, sometimes there...”
- “ I feel like I can't think straight, trouble focusing, remembering”
- “All my tests are negative, why do I feel like this?”
- “I've been to neurology, gastroenterology, ortho, endocrinology, no one knows what's wrong”

What does it sound like?

# National Fibromyalgia Association “snapshot” 2005 3 Day survey



Condition	Description	%
Irritable Bowel Syndrome	Constipation, diarrhea, bloating	32-80%
Temporal Mandibular Dysfunction	Facial/ jaw pain, limited opening	75%
Chronic Fatigue Syndrome	Unexplained, fatigue	21-80%
Headaches	Lasting > 30 mins	10-80%
Multiple Chemical sensitivities	Cause unpleasant symptoms	33-55%
Interstitial Cystitis	At least 8 months pain, urgency, frequency	13-21%
Restless Leg Syndrome	Unpleasant urge to move legs	32%
Numbness	Migratory tingling, pins, needles	44%

*Bennett, R. (2007). Internet survey 2,569 people with FM. BMC MSK Disor; 8: 27.*

# Pain Epidemic in America

## **BY THE NUMBERS:**

- >116 Million people
- 20% chronic pain
- 8% High impact chronic pain:
- Costs \$ 560-635 Billion/ year
- More than DM, CAD, CA combined!
- 14% of all Federal HC \$\$ attributable to pain
- Nearly \$2,000 /person living in the US

- Prevalence:
  - 2-4% population est 5 million adults... 2<sup>nd</sup> only to OA
    - 16% OA
    - 21% RA
    - 36% SLE
    - Unknown: SS, AS, PsA-
      - 75% go undetected... imagine that!
  - 7-8:1 ratio female: male\* in Clinical settings
  - 1-2:1 Epidemiology studies using 2010/11-2016 criteria (survey criteria)
  - 17 missed days work vs. 6 without...
- Morbidity & Mortality
  - Not higher but morbidity is...
  - Death rates from suicide and injuries higher (OR 3.31 suicides/ 1.45 accidental deaths)
  - “will never cause destruction except to personal life”
  - Disability up from 20% in 1996 to 31% in 2015
  - Avg costs \$3,400-3,600 annually direct with total costs as high as \$7,945!! (ER, OV, procedures/ tests, hosp)
  - Indirect costs can be as high as \$8,285 (absenteeism, unemployment, and disability)

Fitzcharles, M. A., Perrot, S., & Hauser, W. (2018). Comorbid fibromyalgia. *Eur J Pain*; 22: 1565-1576.

Walitt, B., Nahin, R., Katz, R., Bergman, M., & Wolfe, F. (2016). *Plos one*; 10(9): doi 10.1371

Wolfe, F., et al. (2011). *A, C & R*; 63(1): 94 & Wolfe, F. et al (2013). *A, C & R*; 65(5): 777

Chandran, A., et al. (2012). *J Manag Care Pharm*; 18(6): 415

Jones, G.T., et al (2015). *A & R*; 67(2): 568



- Retrospective EMR review 2015-2016
- 41,739: 79% female
- 1,600 in Division of Rheumatology alone!
- Medications: most frequent Rx...
  - **Glucocorticoids**
  - **Opiates**
  - **NSAIDs**
  - Of Note: Less than 1%:
    - Anticonvulsants
    - SSRI
    - SNRI
    - TCA

# Prevalence at Northwell

# Prevalence of Myalgia/ FM dx in Rheumatology Nationally

## Rheumatology Informatics System Effectiveness: 55 sites, 239,302

49,345  
 + 34,768  
**84,113** Pain States

# Rise Registry

Table 2. Selected rheumatologic diagnoses captured in the RISE registry\*

Diagnosis at last encounter	ICD-9 codes	No.
Degenerative joint disease		
OA, generalized or localized	715.00, 715.04, 715.09–715.18, 715.20–715.38, 715.80, 715.89	76,381
Inflammatory rheumatic diseases		
RA	714.0, 714.1, 714.2, 714.81	60,102
Polymyalgia rheumatica	725	7,850
Sjögren's syndrome	710.2	15,800
SLE	710.0	13,940
Psoriatic arthritis	696.0	13,550
Spondyloarthritides	720.0–720.2, 720.8, 720.89, 720.9, 729.9	10,265
Vasculitis		
Temporal arteritis	446.5	1,596
Granulomatosis with polyangiitis	446.4	686
Behçet's disease	136.1	301
Henoch Schonlein	287.0	106
Takayasu disease	446.7	85
Goodpasture's syndrome	446.21	4
Scleroderma	710.1	2,754
JIA	714.3, 714.31–714.33	1,342
Dermatomyositis/polymyositis	710.3, 710.4	2,366
Sarcoidosis	135	1,548
Relapsing polychondritis	733.99	886
Crystalline arthropathies		
Gout	274.xx	9,887
CPDD	275.49, 712.1–712.3, 712.8	1,131
Pain syndromes		
Myalgia or myositis (fibromyalgia)	729.1	49,345
Low back pain	724.1	34,768
Infectious arthritis		
Lyme disease	88.81	913
Septic arthritis	711.xx	284

\* All data reflect values at end of last observed clinical encounter. Diagnoses are not mutually exclusive across diagnostic categories; for example, a patient may be captured twice in the Table if they have both rheumatoid arthritis (RA) and osteoarthritis (OA). RISE = Rheumatology Informatics System for Effectiveness; ICD-9 = International Classification of Diseases, Ninth Revision; SLE = systemic lupus erythematosus; JIA = juvenile idiopathic arthritis; CPDD = calcium pyrophosphate deposition disease.

# Fibromyalgia & Mood Disorders

## Risks and Consequence

- Depression 21-83%
- Anxiety 18-60%
- Depression & Anxiety 30-50%
- PTSD as much as 23%
- Bipolar as much as 21%
- Substance Abuse Disorder up to 25%
  
- Suicide rate 10 x HIGHER than General Public
- And 3 x more at risk than other chronic pain patients

- Certal, C., et al. (2018). *The impact of sleep in fibromyalgia, an exploratory study.* *J Psych & Clin Psychiatry*; 9(5): 456-459.
- Guglielmo, D., et al. (2018). *Sx of anxiety & depression adults with arthritis- US, 2015-2017.* *MMWR*; 67(39): 1081-1087.
- Hooten, W. M. (2018). *Chronic pain and mental health disorders: Shared neural mechanisms,* *Mayo Clin Proc*; 91(7): 955-970.
- Mckernan, L. Lenert, M., Crofford, L., & Walsh, C. (2018). *Outpatient engagement lowers predicted risk of suicide attempts in fibromyalgia.* *Arthritis Care & Research.* doi: 10.1002/acr.23748

# Good News

- Little change over time:
  - Hoskin 2018
- Face-Face time matters:
  - Those without SA spent up to 40% more face-face time than those that attempted Suicide.
  - Those without SI spent 3 x more face-face
  - McKernan 2018
- Both more likely with Younger age, Mental illness, Co-morbid illness & Frequent inpatient admissions...
  - Suicide Ideations: more polysomatic sx
  - Suicide Attempts: obesity & **drug dependence!**

*Mckernan et al. (2018). Arthritis Care & Research. doi: 10.1002/acr.23748*

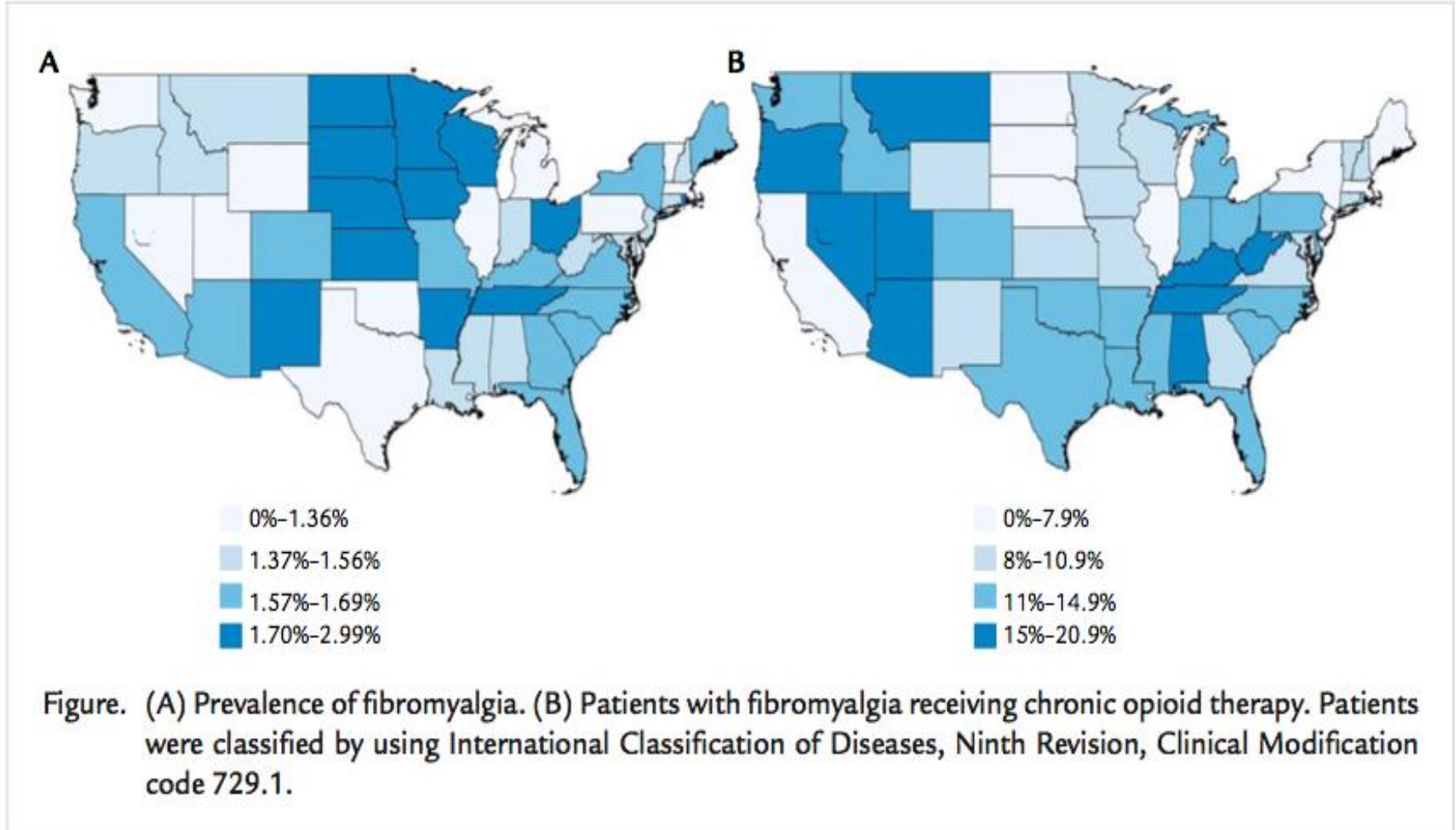
*Hoskin, T., Whipple, M., Nanda, S., & Vincent, A. (2018). Arthritis Research & Therapy; 20: 37.*

# Clusters of FM patients

Low D & A	High D & A	
<b>Cluster I</b> Low D & A Low Pain	<b>Cluster III</b> High D & A Low Pain	<b>Low Pain</b>
<b>Cluster II</b> Low D & A High Pain	<b>Cluster IV</b> High D & A High Pain	<b>High Pain</b>

- Vincent, A. et al. (2014). *Arthritis Research & Therapy*; 416: 463.
- Hoskin, T., Whipple, M., Nanda, S., & Vincent, A. (2018). *Arthritis Research & Therapy*; 20: 37.

# Prevalence of FM and Use of Opioids

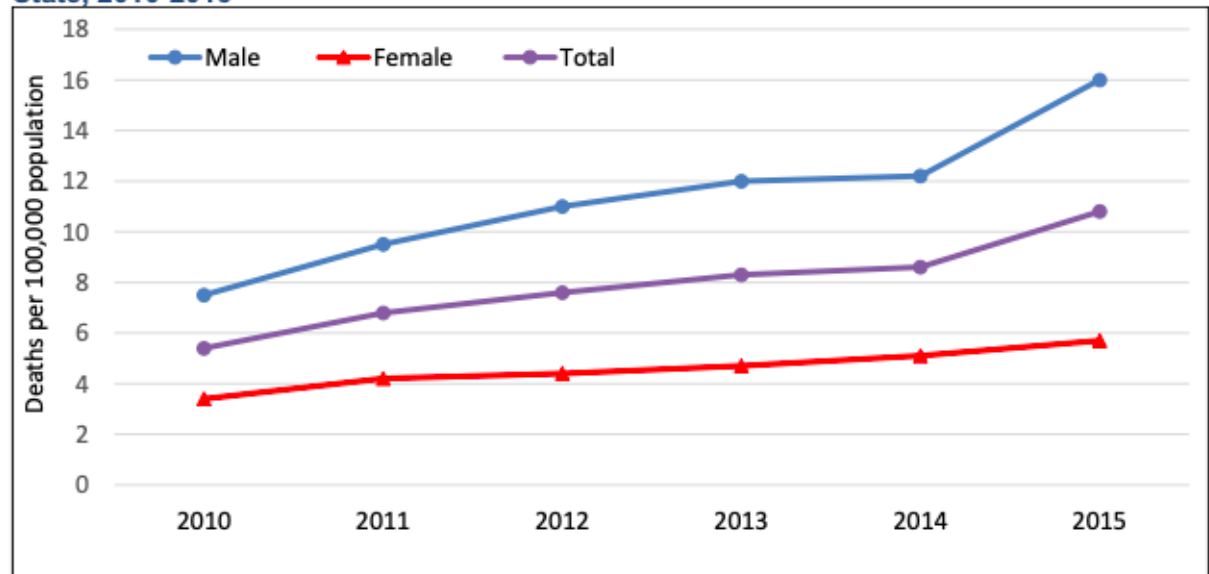


**Commercially Insured 2007-2009 10-30% patients on Chronic Opioids**



Figure 2 shows the annual trends from 2010 to 2015 for the age-adjusted rates of all opioid overdose deaths per 100,000 population by gender, as well as the total rate for NYS. The rate of all opioid overdose deaths per 100,000 population in New York was consistently higher for males than females, although the rate of all opioid overdose deaths steadily increased for each gender. In 2015, the rate for males (16.0) was almost three times higher than the rate for females (5.7).

**Figure 2. Age-adjusted Rates of Overdose Deaths Involving Opioids by Gender, New York State, 2010-2015**



**Source: CDC Wonder**

### The Numbers:

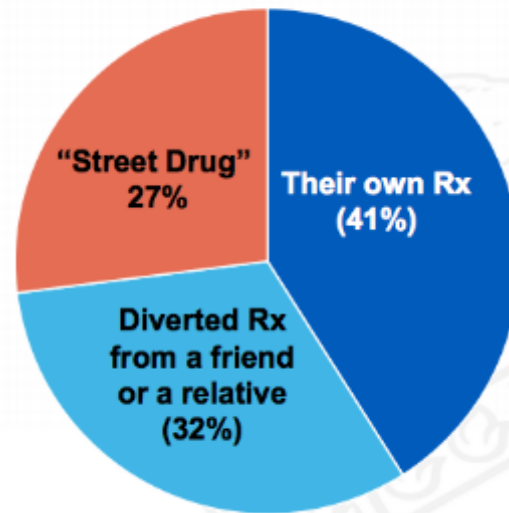
- 1 person every 19 mins killed
- 2012 NY 259 million Rx = one Rx for every American adult!
- 2 out of 5 teens in LI abused Rx drugs!

**Prescription  
drugs serve as  
“gateway drugs”  
to illicit drug use.**

Patients who are prescribed opiates for acute pain can develop an addiction.

Canfield et al. *J Addict Med* 2010;4:108-113

**Self-Reported Etiology of  
Opioid Addiction, n = 75**



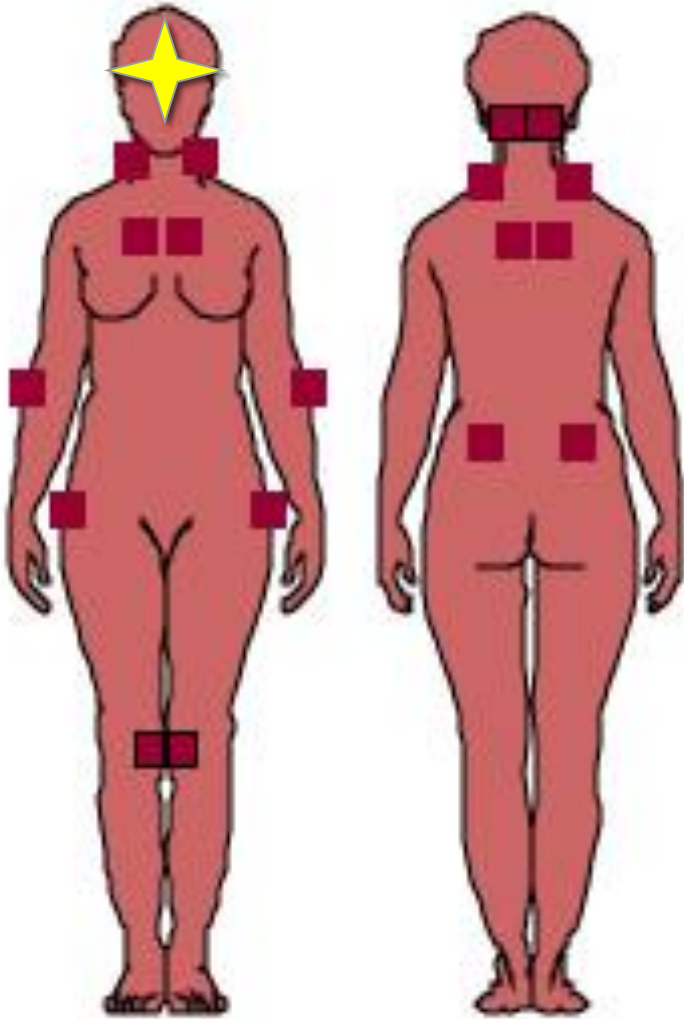
**73% came from Rx drugs!!!**



# Making the Diagnosis

## To Count or Not to Count

- 1990 ACR Criteria 85% sensitivity
  - Used in most FM trials
  - Widespread pain above/ below, R/L side
  - 11 of 18 tender points
  - 4 kg force- blanch fingernail
- Too subjective
- 25% FM did not meet criteria



Tender points  
are NOT  
Trigger Points!



Be sure to check control... forehead,

# History of Diagnostic Criteria:

- 1990 ACR Criteria: 85% sensitivity & 90% specificity
  - The **tender point exam 11/18 required**
  - Specificity of criteria was near perfect and considered “Gold Standard”
  - 25% patients were missed
- 2010 ACR Criteria: eliminated tender point exam but requires examiner
  - Widespread Pain Index
  - Symptom Severity Score- if your SS was high enough you could have regional pain and still +
- 2011 Modified 2010 ACR Criteria- eliminated the examiner
  - Questionnaire
  - Fibromyalgia Symptom Scale
  - Severity: “Fibromyalgianess”
  - Score: 0-31 (combination of WPI & SS)
  - > 13 sensitivity 96% and specificity of 91.8%
- 2016 Revisions to modified 2010/2011
  - Complete self report
  - Generalized pain defined as pain in 4 out of 5 regions
  - Dx of FM is valid irrespective of other: Primary vs. Secondary

FM= WPI  $\geq$  7 & SS  $\geq$  5

or WPI 3-6 & SS  $\geq$  9

### Widespread Pain Index (1 point per check box; score range: 0-19 points)

① Please indicate if you have had pain or tenderness during the past 7 days in the areas shown below. Check the boxes in the diagram for each area in which you have had pain or tenderness.

### Symptom Severity (score range: 0-12 points)

② For each symptom listed below, use the following scale to indicate the severity of the symptom during the past 7 days.

- No problem
- Slight or mild problem: generally mild or intermittent
- Moderate problem: considerable problems; often present and/or at a moderate level
- Severe problem: continuous, life-disturbing problems

	No problem	Slight or mild problem	Moderate problem	Severe problem
<b>Points</b>	0	1	2	3
A. Fatigue	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
B. Trouble thinking or remembering	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
C. Waking up tired (unrefreshed)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>

③ During the past 6 months have you had any of the following symptoms?

	0	1
A. Pain or cramps in lower abdomen	<input type="checkbox"/> No	<input type="checkbox"/> Yes
B. Depression	<input type="checkbox"/> No	<input type="checkbox"/> Yes
C. Headache	<input type="checkbox"/> No	<input type="checkbox"/> Yes

### Additional criteria (no score)

④ Have the symptoms in questions 2 and 3 and widespread pain been present at a similar level for at least 3 months?

No       Yes

⑤ Do you have a disorder that would otherwise explain the pain?

No       Yes

Total Score 0-31 > 12 = Fibromyalgia and KEY impact outcomes  
**2016** Revisions to 2010/2011 FM Diagnostic Criteria



# Challenges:

- *Conceptual change from primary pain condition to multisystem syndrome*
- *Understand that “Fibromyalgiansess” will impact outcomes*
- *Should be measured on all chronic pain patients... determines outcomes!*

Polysymptomatic Distress Scale (PSD) =  
Fibromyalgia Symptom Questionnaire (FSQ)  
“Fibromyalgiansess”  
Score 12-13

# Is FIBROMYALGIA real?



Cynthia Yolland  
Queen Creek, AZ "Today's Forecast"

*"This is an expression of the two worlds that is pain, especially in hidden syndromes and chronic pain. There is the outer world that is normal. "You look wonderful". Then the turmoil that is going on inside. I wanted to take the pain out of the hidden box and shake it, then show it to my doctors, and my family, and friends. In fact, a copy of this picture is part of my medical records."*

## Pathophysiology

# Is Fibromyalgia Real? Yes...

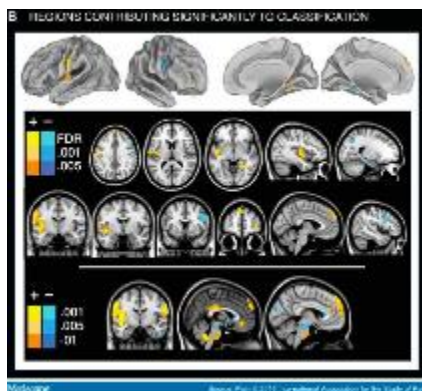
- Prototypical Central Amplification/ Sensitization
- Hyperalgesia
- Volume turned up way too high!
  - More sensitive to stimulation
  - Exaggerated response to that stimulation
  - Inability to modulate the pain



**Here's why...**

## Pathophysiology:

- Work of... seminal papers
  - Functional MRI's
  - Pain threshold
  - Central Sensitization & Pain Amplification
- Altered pain processing peripherally and centrally contribute to central amplification **facilitation** and dampened effect **inhibitory** mechanisms
  - Bidirectional neural signals:
    - Pain & Mood
    - Pain & Sleep
  - **Failure of endogenous opiate systems**



# Is Fibromyalgia real? No longer a debate

## Signature for Fibromyalgia

### How Do We Know That the Pain in Fibromyalgia Is “Real”?

Richard E. Harris, PhD, and Daniel J. Clauw, MD

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Current Pain and Headache Reports 2006, 10(4):403–407  
Current Science Inc. ISSN 1528-8325  
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Fibromyalgia is a common idiopathic pain condition often resulting in increased morbidity and disability in patients. The lack of peripheral abnormalities in this disease has led clinicians and researchers alike to question if this syndrome represents a valid entity. Recent genetic findings suggest that specific gene mutations may predispose individuals to develop fibromyalgia. In addition, neurobiological studies indicate that fibromyalgia patients have abnormalities within central brain structures that normally encode pain sensations in healthy pain-free controls. Future studies that focus on central neurobiological and/or genetic influences in fibromyalgia may bring insight into mechanisms of this problematic disease and ultimately result in improved treatments.

#### Introduction

Fibromyalgia is a common systemic disorder estimated to affect 2% to 4% of the population, second in prevalence among rheumatologic conditions to osteoarthritis [1]. There are legitimate controversies surrounding many aspects of fibromyalgia, including precisely how it should be defined, as well as whether individuals with this condition are disabled or deserve compensation [2]. However, at the same time, there have been rapid advances in our aggregate scientific knowledge about fibromyalgia. There are now multiple, converging lines of evidence confirming that the pain of fibromyalgia is “real” and that there are strong neurobiological underpinnings to this condition [3]. In fact, there has been a parallel recognition that many pain syndromes classically thought to be “idiopathic,” such as irritable bowel syndrome, tension headache, temporomandibular syndrome, and idiopathic low back pain,

share overlapping symptom expression and underlying mechanisms with fibromyalgia [4–6].

#### The American College of Rheumatology

Criteria for Fibromyalgia:  
The Good and the Bad

In 1990, the American College of Rheumatology (ACR) established classification criteria for fibromyalgia [7]. These criteria require that an individual must have both chronic widespread pain involving all four quadrants of the body as well as the axial skeleton, and the presence of 11 of 18 tender points on examination. Two positive aspects of the ACR criteria are that 1) as intended, these allowed investigators throughout the world to use a single set of classification criteria to standardize research studies, and 2) these criteria require that individuals display widespread tenderness in addition to pain. In fact, many of the mechanistic and physiological studies of fibromyalgia have explored the underlying mechanism for this tenderness (see following text). However, the ACR criteria have not been as uniformly positive in the societal sense. Two of the (unintended) consequences of these criteria are that they 1) are often used in clinical practice to diagnose individual patients in clinical settings (this was not the intended purpose and should not be rigidly adhered to); and 2) skew the diagnosis of fibromyalgia primarily toward women with high levels of distress (by assessing tenderness by “counting tender points”).

#### The Problems with Tender Points

When the ACR criteria were published, it was thought that tender points were somehow areas that were increasingly tender in individuals with fibromyalgia, and many therapies were even specifically aimed at reducing tender points (eg, injections with anesthetic or corticosteroids). Since then, research has definitively shown that patients with fibromyalgia have tenderness or decreased pain thresholds extending throughout their body, and not only at localized areas considered tender points [8,9]. Thus, we now know that tender points are merely areas where anyone is more tender and that overall tenderness in an individual can be successfully predicted by just assessing



## TIMELINE:

### 1858

- Florence Nightingale...Described FM sx - Died at age 90

### 1904

- Gower first description in medicine
- Fibrositis

### Early 1975-1980s

- Symthe & Yunus modern Fibromyalgia
- First clinical trials: FM vs HC
- Moldofsky- SLEEP disordered sleep and increased PAIN

### Disordered Sleep: Non restorative sleep and Chronic Fatigue

- Lower sleep quality: alpha intrusions and less slow wave deep sleep
- Lower efficiency, fragmented
- Diagnostic criteria for FM with worse sleep associated w/ increased symptoms
- Restorative sleep, decrease WPS

- Moldofsky, H., Scarisbrick, P., & Smythe, H. (1975). *MSK sx and Non-REM sleep disturbance ...Psychosomatic Medicine*; July/ August.
- Wu, Y.L., et al. (2017). *Sleep disturbances in fibromyalgia: A meta-analysis . J Psychosomatic Res*; 96: 89-97.
- Yunus, M., et al. (1981). *Primary fibromyalgia (fibrositis). Clinical study of 50 patients with matched normal controls. Seminars in Arthritis & Rheum*; 11(1): 151-171

#### Florence Nightingale

Florence Nightingale was a pioneer in health during the Victorian period. She also wrote about health conditions and dietary information based on notes about patients she'd treated during her time in the Ottoman Empire. Her first book, published in 1858, was called *Notes on Matters Affecting the Health, Efficiency and Hospital Administration of the British Army*. Her fibromyalgia was triggered by an infection, and she was finally bedridden in 1896. After years of struggling with this disorder, she died in her sleep at the age of 90.



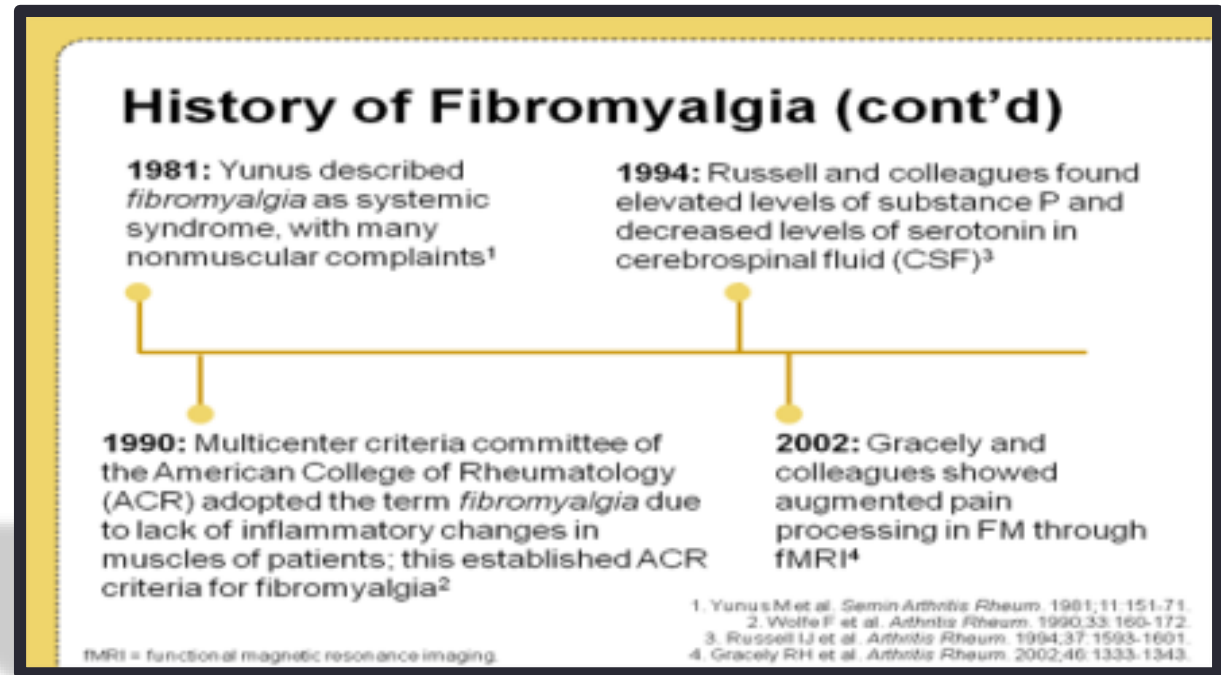
# History of Fibromyalgia

# History of Fibromyalgia

## 1990

### ACR Criteria for FM

- Tender Point Count  
11/18 > 3m
- Clinical trials



## 1990-2010

- Substance P 3 x higher **PAIN**
- Serotonin, Norepi, Dopa lower
- Endogenous opioids higher
- Gaba low, glutamate high
- NGF higher **PAIN**

(see reference list for details)

# fMRI- Fibromyalgia Signature

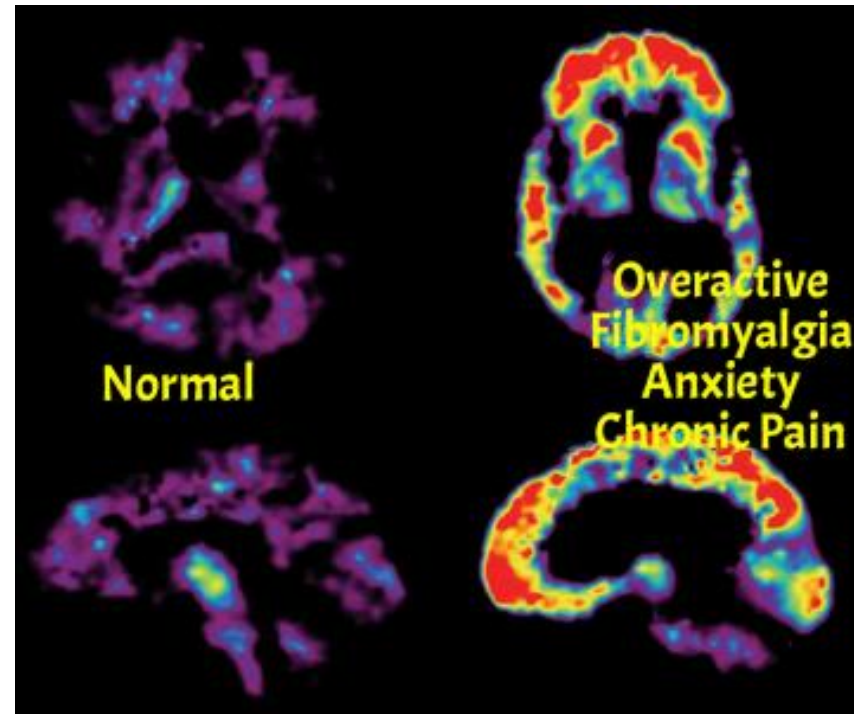
- **2002-** Gracely & Clauw fMRI
  - AC&R: 835-843 (N=16 FM vs. 16 matched)
- **2016-** Shi, H., et al.
  - Sem Arthritis & Rheum; 46: 330-337. Meta Analysis
- **2017-** Lopez et al.
  - Pain; 158(1): 34-47. (N=40 vs. 40 matched)

## *Fibromyalgia Signature*

ARTHRITIS & RHEUMATISM  
Vol. 46, No. 5, May 2002, pp 1330-1340  
DOI 10.1002/art.10225  
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### Functional Magnetic Resonance Imaging Evidence of Augmented Pain Processing in Fibromyalgia

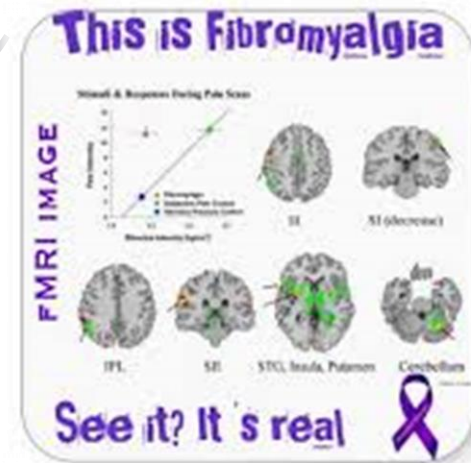
Richard H. Gracely,<sup>1</sup> Frank Petzke,<sup>2</sup> Julie M. Wolf,<sup>3</sup> and Daniel J. Clauw<sup>2</sup>



- f MRI findings:
  - Central Sensitization now referred to as AMPLIFICATION- occurs in **MOST CHRONIC PAIN STATES-**
- Functional MRI:
  - Noted altered activity in areas brain responsible for pain
  - Now evidence of “FM” signature
  - Studies suggest changes in brain activity following CBT for Depression/ Pain.
    - *deCharms, C., et al. (2005). Control over brain activation and pain learned by using real-time functional MRI. PNAS; 102(51). 18626-18631.*

## *Signature for Fibromyalgia*

## Functional MRI's Evidence



# TIMELINE:

## 2004

- Arnold et al. Genetic basis confirmed

## 2013

- Uceyler et al. SFN prevalent up to 60%

## 2015- 2018

- Brummet et al. (2015) **“Fibromyalgiansess”** > 13 FSQ

### Others:

- Janda, et al. (2015). *FM survey criteria are associated with increased postoperative opioid consumption in women undergoing hysterectomy. Anesthesiology; 122: 1103-11.*
- Ablin et.al. (2017). *Effect of FM symptoms on outcomes of spinal surgery. Pain Medicine; 18: 773-380*
- Kurien, et al. (2018). *Preoperative neuropathic pain-like symptoms and central pain mechanisms in knee OA predicts poor outcomes 6 m after TKR surgery. The J of Pain; 19 (11): 1329-1341.*



Consider calculating this score when determining outcomes in all conditions...

# “Fibromyalgianess”

- Large prospective observational study followed x 6 months postop N= 464
- Effects outcomes...
- Higher FS was predictive of less improvement in pain postop
- For every 1 degree in FSS, 9 mg increase in MSO4 equivalent
- 6 months after surgery determinant improvement in pain

Brummett, C., et al. (2015). Outcomes following TKR & THR. *Arthritis & Rheum*; 67(5): 1386

ARTHRITIS & RHEUMATISM  
Vol. 67, No. 5, May 2015, pp 1386-1394  
DOI 10.1002/art.21601  
© 2015 The Authors. *Arthritis & Rheumatism* is published by Wiley Periodicals, Inc. on behalf of the American College of Rheumatology. This is an open access article under the terms of the Creative Commons Attribution-NonCommercial-NoDerivs License, which permits use and distribution in any medium, provided the original work is properly cited, the use is non-commercial, and no modifications or adaptations are made.

## Characteristics of Fibromyalgia Independently Predict Poorer Long-Term Analgesic Outcomes Following Total Knee and Hip Arthroplasty

Clara M. Brummett,<sup>1</sup> Andrew G. Lizzit,<sup>1</sup> Adam L. Hassett,<sup>1</sup> Alex Fancher,<sup>2</sup> Brian R. Halstrom,<sup>1</sup> Nathan L. Wood,<sup>1</sup> David A. Williams,<sup>2</sup> and Daniel J. Clauw<sup>1</sup>

**Objective.** While psychosocial factors have been associated with poorer outcomes after knee and hip arthroplasty, we hypothesized that augmented pain perception, as occurs in conditions such as fibromyalgia, may account for decreased responsiveness to primary knee and hip arthroplasty.

**Methods.** A prospective, observational cohort study was conducted. Preoperative phenotyping was

The contents of this article are solely the responsibility of the authors and do not necessarily represent the views of the National Institute of Arthritis and Musculoskeletal and Skin Diseases or

conducted using validated questionnaires to assess pain, function, depression, anxiety, and satisfaction. Participants also completed the 2011 Fibromyalgia survey questionnaire, which addresses the widespread body pain and somatoid symptoms associated with characteristics of fibromyalgia.

**Results.** Of the 668 participants, 464 were evaluated 6 months after surgery. Since individuals who met criteria for being classified as having fibromyalgia were subjected to random loss favorable, all primary analyses excluded these individuals (69% of the cohort).

# Pathogenesis- Summary

- Pain **CATASTROPHIZING**, or characterizations of pain as awful, horrible, unbearable, is increasingly being recognized as an important factor in the experience of pain.
- Endorphin receptors Mech dysfunction- reason for failure to respond to opiates... Should NOT be used
  - Endogenous opioid levels are elevated
  - Opioid receptor binding is diminished as seen in depression and anxiety\*\*



## Summary:

# Mechanisms of Amplified Pain

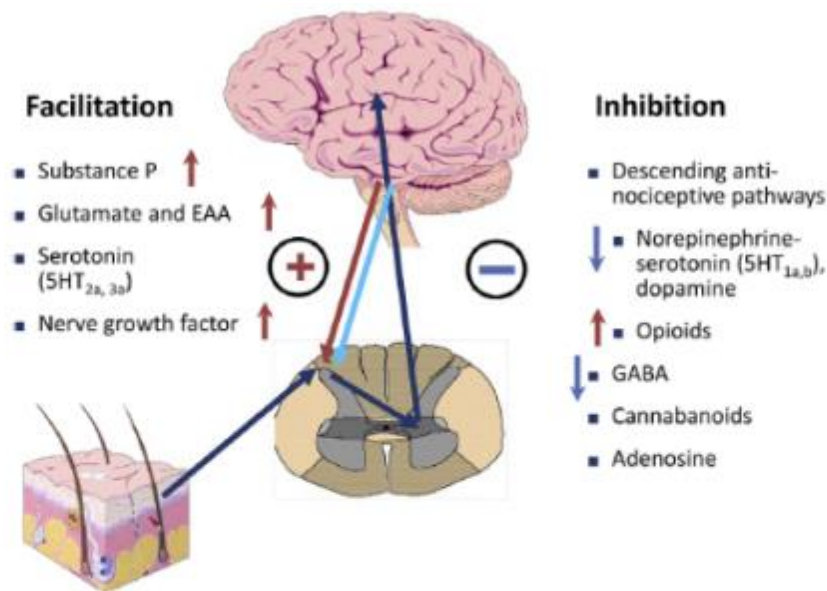


Fig 3. CNS influences on pain and sensory processing.

endogenous opioid system. Both CSF levels and brain activity by functional neuroimaging appear to be augmented, not reduced (as would be necessary to cause augmented pain processing), in FM, which may be the reason why opioidergic drugs do not work well to treat FM and related pain syndromes [60,61].

*Potential role of peripheral factors in centralized pain states*

Aberrant brain Transmission is the Hallmark of FM



# What causes it?

- Non Modifiable Causes:

- Trauma: physical or emotional/psychological often – sexual/ abuse
- Illness: viral infections, lyme or EBV, Hep C
- Genetic predisposition becoming more clear, but also learned: 50%



- Modifiable Causes:

- STRESS, chronic #1
- Sleep deprivation
- Mood: anxiety & depression
- Obesity!!!
- Personality characteristics, coping mechanisms-

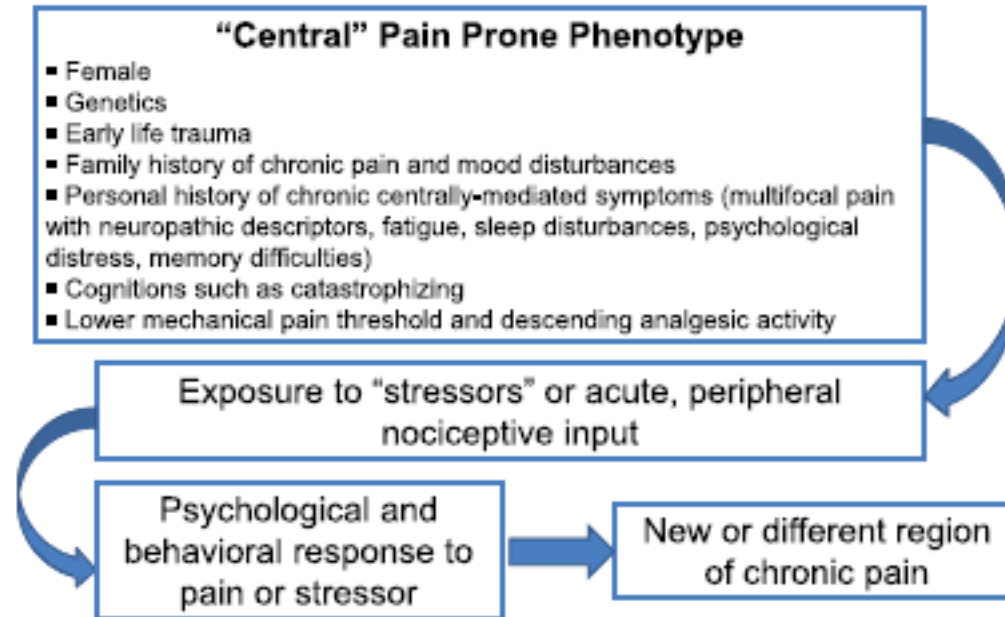


Arnold, L., et al. (2004). *Arthritis & Rheum*; 50(3): 944-952

Clauw, D. (2015) *Best Practice & Research Clinical Rheumatology*; 29(1): 6-19.

Nakamura, I., et al. (2014). *Artiritis & Rheum*; 66(7): 1093

# Pain Prone Phenotype:



**Fig. 2.** Pain-prone phenotype.

Clauw, D. (2015). *Dx and treating chronic musculoskeletal pain based on mechanisms. Best Practice & Research Clinical Rheumatology*; 29(1): 6-19.

## Standardize Care:

- Should avoid expensive dx studies
  - CBC & CMP
  - ESR
  - CRP
  - TSH
  - CPK
- All other studies should be driven by specifics
  - No evidence viral or other infectious cause
  - Vit D: not directly r/t but levels < 20, many chronic pain low D
  - ANA only with direction
- Sleep study when necessary
- Psychiatric screening: anxiety & depression



Overnight Sleep Study or Polysomnogram

# Diagnostic Studies

# Guidelines for FM and Chronic Pain

## Chronic Pain

H & P, Dx as indicated

Dx FM or other CP state

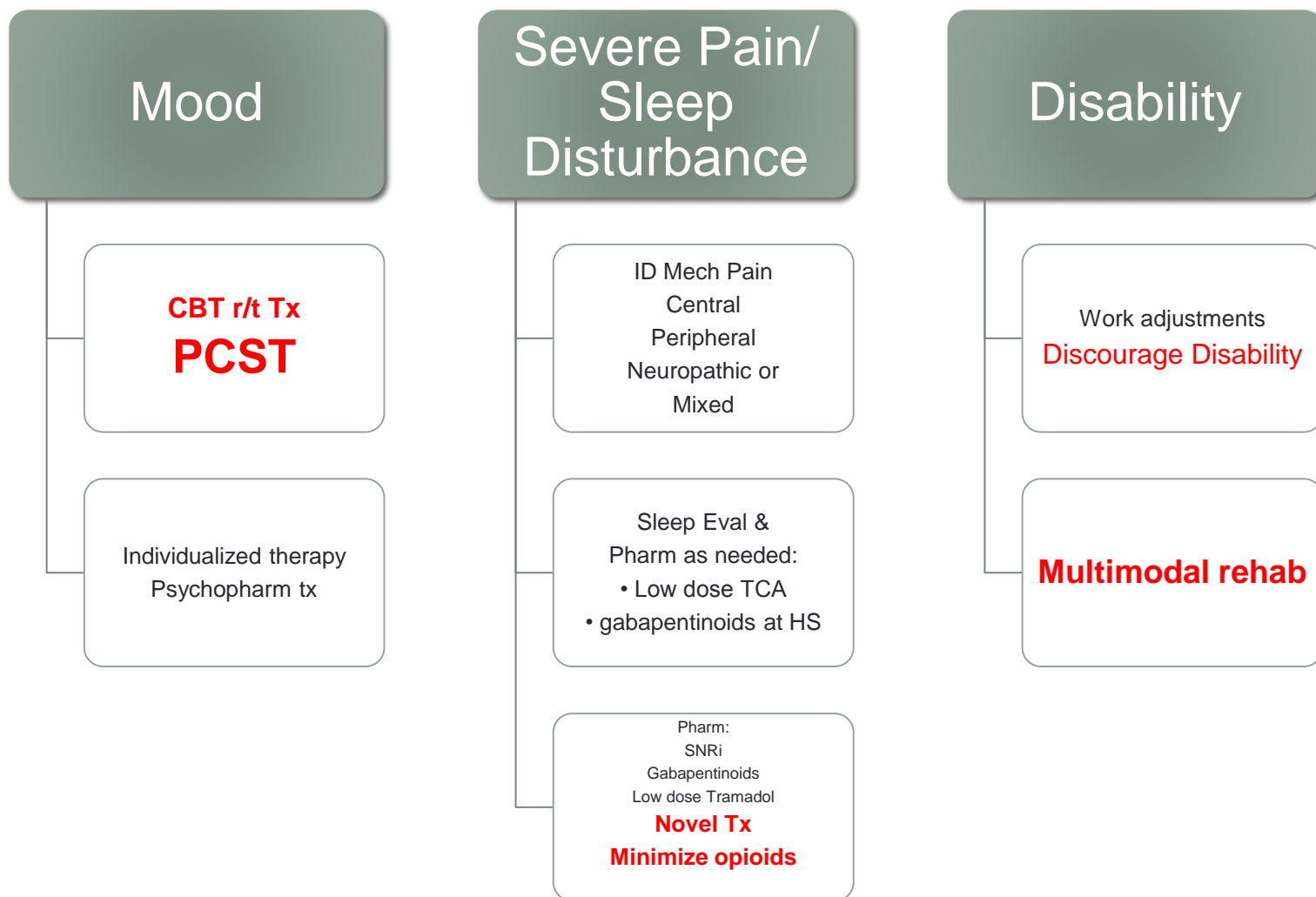
Patient / Provider Education

PT, graded exercise & other NP therapies

Reassessment & tailored Individualized Tx

- Macfarlane, G.J., et al. (2017). EULAR revised recommendations for the management of FM. *Ann Rheum Dis*; 76: 318-328.
- Tick, H., et al. (2017). A consortium pain task force white paper. [www.nonpharmpaincare.org](http://www.nonpharmpaincare.org).
- <https://www.cdc.gov/drugoverdose/prevention/index.html>

# Guidelines for FM and Chronic Pain



• Macfarlane, G.J., et al. (2017). EULAR revised recommendations for the management of FM. *Ann Rheum Dis*; 76: 318-328.

• Tick, H., et al. (2017). A consortium pain task force white paper. [www.nonpharmpaincare.org](http://www.nonpharmpaincare.org)

## Basic Tenements:

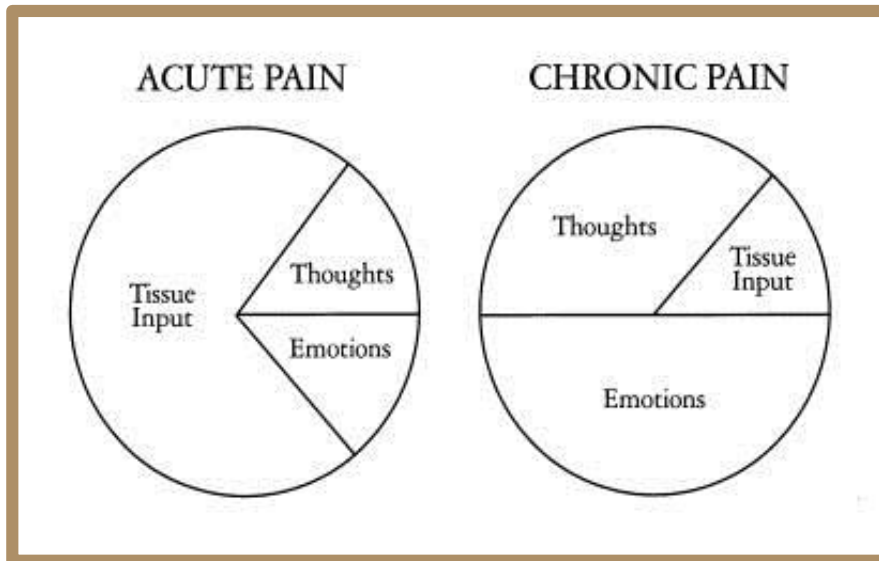
Fibromyalgia is a Disorder (constellation of symptoms) not a Disease

1. Symptoms are not caused by an organic disease
2. It is a legitimate condition
3. Symptoms are persistent in nearly all patients often turned on or worsened
  - Psychological stress
  - Emotional conflict
4. Total relief of symptoms is seldom achieved: **No magic pill!**
5. The symptoms do not lead to disability and **DO NOT** shorten life expectancy
6. Most patients learn to adapt to the symptoms over time
7. Self management strategies to modulate symptoms should be emphasized
8. Goals of treatment are:
  - Improvement in quality of life
  - Maintenance of function in everyday life
  - Reduction of symptom burden

# Treatment of FM: Education is KEY

# Understanding Pain: Patient Teaching

Acute pain **NOT** effectively treated- leads to CPS



Persistent Pain  
> 3 mnths  
Is  
Chronic Pain

This is how I explain it to patients...

- Make the Diagnosis
  - **Education at diagnosis sets expectations**
  - Identify resources
- Evaluate for co morbid conditions
  - Identify any underlying condition: SS, PsA
  - Sleep disorders
  - Identify other co morbid pain conditions
    - IBS; Migraines/ HA; IC/ chronic prostatitis; TMJ; LBP/ neck pain/ myofascial pain; chronic pelvic pain;
  - Evaluate for depression, anxiety, other co morbid psychiatric issues: PHQ, GAD-7
  - Social stressors
    - Family
    - Work/ leisure activity
  - Obesity
  - Physical strengthening: Strongest evidence
- Other Non Pharmacologic strategies
  - CBT vs. CST: **in person vs. online support- using [www.FibroGuide.com](http://www.FibroGuide.com) or [www.Knowfibro.com](http://www.Knowfibro.com)**
  - Positive Affective interventions: Teach Happiness
  - Complementary and Alternative Treatments: acupuncture
- Pharmacotherapy to control pain: weak evidence at best

## Standardize Care



# Physical Strengthening

- Start somewhere:
  - 50% of what you think you can do and slowly increase
- Physical therapy:
  - Cardio
  - Tai Chi...

# Self Management Strategies:

Response to CBT... change in sensitivity

- • *Ang et al. (2010). AC & R; 62(5): 618-23.*
- **(N=17 FM vs. 16 UC 6 wk telephone CST)**
  
- • *Emery, C.F. et al. (2006). J Pain Symp Mngmt; 31(3): 262-269.*
- **(N=65 for 45 mins CST- telephone)**
  
- *Dear, et. al. (2018). J of Pain; 19(12): 1491-1503.*
- *12-24 month f/u Internet CBT based program*
- *N=490 into 4 groups varying amounts clinician time*
  - **(n= 143 regular, n=141 optional contact, n= 131, n= 75 waitlist)**



## What's the Evidence?

- If you don't address SLEEP and PSYCH, **NOTHING** you do will work!
- Behavioral strategies first...
- Pharmacotherapy when necessary:
  - Address sleep
  - Mood
  - Modify Central Pain processing



**Standard of care: Treatment**

# Understanding Sleep...

## ASSESS SLEEP:

Describe the problem...

- Sleep: trouble falling, staying and/or Non restorative
  - What is INSOMNIA?
- Daytime fatigue
- This will determine next step: what is interfering w/ sleep
  - Pain
  - Anxiety
  - Behavioral issues
  - OSA: snoring, apneic episodes (incidence)
    - Disrupted sleep as a component of FM (scientific evidence)
    - Contributing to chronic fatigue and widespread pain
- Sleep study when necessary...
- Caution medications can impair sleep...

Address the problem...

Sleep

- Dark, cool, quiet room
- **No screen time 30-45 mins, if reading w/ electronic device- night mode- melatonin**
- **Bed is for sex and sleep only**
- Routine night time ritual – same time going & waking up
- Minimize distractions at night...
- Balance activity and rest throughout the day., every day
- No heavy exercise, food, ETOH, stress, smoking before bed... allow for transition
- **Get out of bed if not asleep after 20 minutes; something calm 30 minutes, then try again**
- Don't go to bed if not drowsy.
- **Naps: and not after 3:00 p.m or longer than 45-60 mins**
- Caution: antihistamines/ hypnotics to help with sleep, poor quality.
- **Don't let children or pets interfere with your sleep.**



## Sleep: Do's & Don'ts

# Targeted Therapies: Neurochemical

- Treat Sleep disorder
- Address Mood disorders...
- Avoid Opioids: High levels of endogenous opioids in CSF
  - Down regulation of opiate mu receptors
    - High risk for failure
    - Misuse and abuse
    - Potentiating hyperalgesia...
- Increase Serotonin, Norepi and Dopa
- Block substance P
- Increase Gaba & decrease glutamate
- Block NGF

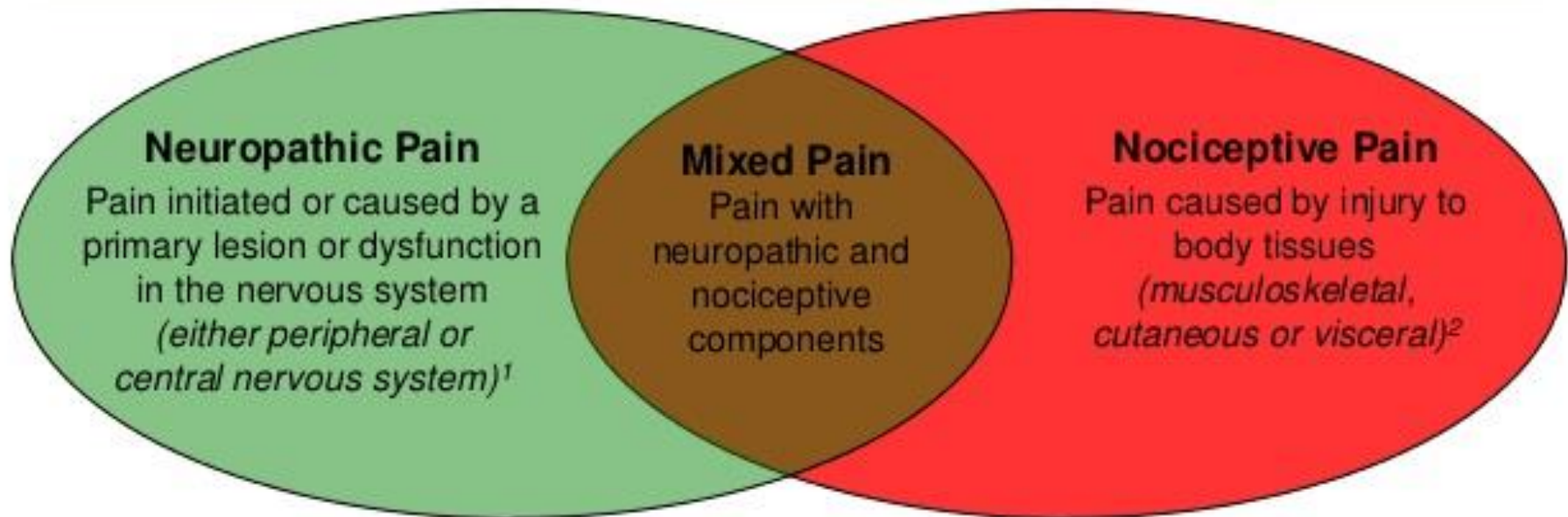
## • KEY RESOURCES:

- *Atzeni, F. (2019). Review: One year in review 2019: Fibromyalgia. Clin & Exp Rheum; 37 (Suppl. 116): S3-S10. \**
- Clauw, D. (2015). Diagnosing and treating chronic musculoskeletal pain based on mechanisms. *Best Practice & Research Clinical Rheumatology*; 29(1): 6-19.
- Fitzcharles, M.A. (2013). 2012 Canadian guidelines Pain Res Manag; 18(3): 119-126.
- Hauser, W., & Fitzcharles, M. (2019). Facts and myths pertaining to fibromyalgia. *Dialogues Clin Neurosci*; 20(1): 53–56.
- Macfarlane GJ et al. (2016). EULAR revised recommendations for the management of fibromyalgia. *Ann Rheum Dis*; 0: 1-11. doi:10.1136/annrheumdis-2016-209724

# Drug Therapies:

- Optional NOT Obligatory
- No 2 people are the same...
- Should be targeted to the underlying process: Essential
  - Sleep: need to clarify the issue...
  - Mood
  - GI/ GU issues
- Pain modulation

# Presentation Across Pain States Varies



## **Examples**

- Postherpetic neuralgia
- Trigeminal neuralgia
- Painful diabetic neuropathy
- **Postsurgical neuropathic pain**
- Posttraumatic neuropathy
- Central poststroke pain

## **Common descriptors<sup>2</sup>**

- Burning
- Tingling
- Hypersensitivity to touch or cold

## **Examples**

- Low back pain with radiculopathy
- Cervical radiculopathy
- Cancer pain
- Carpal tunnel syndrome

## **Examples**

- Pain due to inflammation
- Limb pain after a fracture
- Joint pain in osteoarthritis
- Postoperative visceral pain

## **Common descriptors<sup>2</sup>**

- Aching
- Sharp
- Throbbing

1. International Association for the Study of Pain. IASP Pain Terminology;

2. Raja et al. in Wall PD, Melzack R (Eds). Textbook of Pain. 4th Ed. 1999;11-57.



# Medications & SLEEP

- Benzodiazepines & ETOH
- Stimulants: caution
- Ambien, Lunesta...Gaba, benzo receptor
- Melatonin, Romelteon
- Marijuana: CBD vs THC
- Antidepressants
  - **Amitriptyline**, nortriptyline, trazodone
  - SSRI, SNRIs and NDRIs
- Muscle relaxants: several
  - **Cyclobenzaprine**: TCA properties
  - Tizanidine/ Baclofen: anti spastic agents
  - Carisprodol
- Psychotropic agents:
  - Quetiapine
  - Mirtazepine: TCA properties
- Gabapentinoids

- MEDICATIONS:
  - Alpha 2 delta ligands (anticonvulsants): excellent...
    - analgesic benefits through substance P reduction
    - anxiolytic centrally
    - Restores slow wave deep restorative sleep...
    - Mood stabilizer
    - Small fiber neuropathy
  - Gabapentin & Pregabalin ... others available
    - Gabapentin does not require PA, gentle titration easy
    - Start slow... only at night initially for tolerance
    - Titrate to lowest effective dose as tolerated

## Pharmacotherapy

## Treatments to Consider:

- Rozerem or Melatonin/ Magnesium & Zinc
- TCA- Amitriptyline/ Tetracyclic- Trazodone- Na channel calms excitable nerves
- Muscle relaxants
  - Cyclobenzaprine: TCA properties
- antispastic agents
  - Tizanidine
  - Baclofen
- Seroquel (quetiapine): excellent for insomnia in low doses
- Anti depressants depend on the presentation
  - Pain: SNRIs...Duloxetine & Milnacipran
  - Anxiety/ and Older: Sertraline, citalopram
  - Cognitive issues: Bupropion

# Pharmacotherapy

- Analgesics:
  - **Heat & Ice**
  - APAP
  - NSAIDs \* **Lowest dose for shortest term\*** tx regional problems
    - Naproxen/ Celecoxib has safest CV profile
    - Ibuprofen/ Celebrex / meloxicam better GI tolerance
    - Should always be taken with GI protection
  - Topical vs oral
  - Precautions... requires careful monitoring & Education
  - Anticonvulsants: Gabapentin & Pregabalin
  - Anti depressants: what is the problem?

### Controversial:

- LDN
- Marijuana
  - CBD
  - THC

# Medications

*Nissen, S., et al. (2016). Cardiovascular safety of celecoxib, naproxen, or ibuprofen for arthritis. NEJM; 375(26): 2519-2529.*

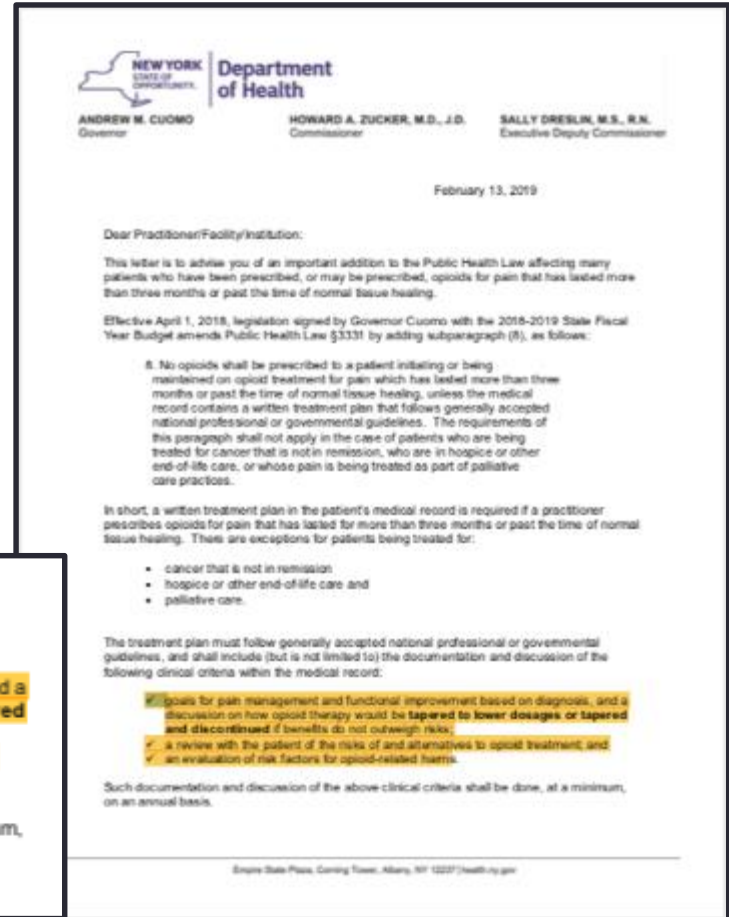
## • Opioid Use:

- Tramadol: mu and 5HT
- BUT... NYS requires documentation
- Lowest dose, shortest time
- Risk assessment
- Have an out
- Measure function!

The treatment plan must follow generally accepted national professional or governmental guidelines, and shall include (but is not limited to) the documentation and discussion of the following clinical criteria within the medical record:

- ✓ goals for pain management and functional improvement based on diagnosis, and a discussion on how opioid therapy would be tapered to lower dosages or tapered and discontinued if benefits do not outweigh risks;
- ✓ a review with the patient of the risks of and alternatives to opioid treatment; and
- ✓ an evaluation of risk factors for opioid-related harms.

Such documentation and discussion of the above clinical criteria shall be done, at a minimum, on an annual basis.



# Pharmacotherapy

# Risk Assessment prior to Opioid Rx...

## Scoring (Risk)

0-3: low

4-7: moderate

≥ 8: High

## Opioid Risk Tool

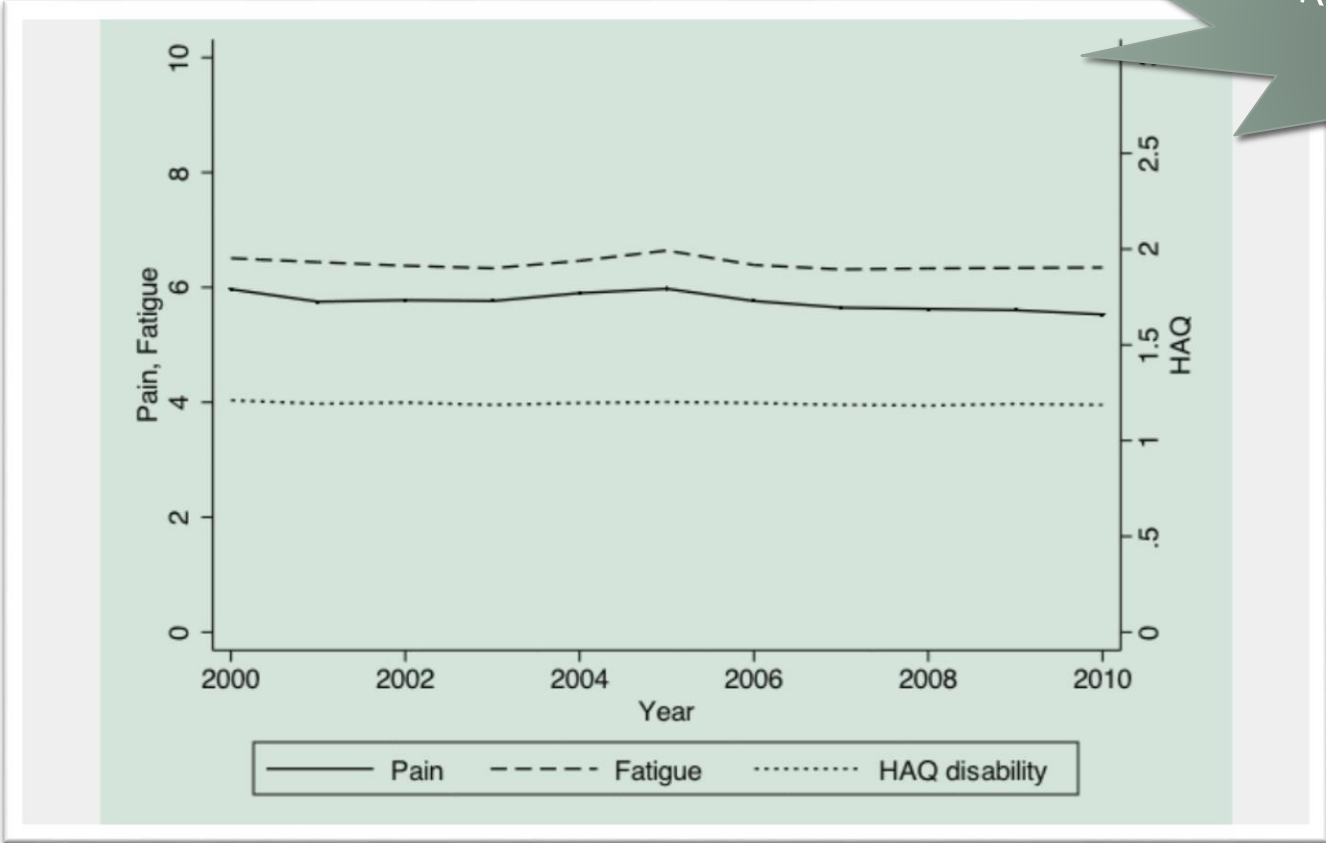
This tool should be administered to patients upon an initial visit prior to beginning opioid therapy for pain management. A score of 3 or lower indicates low risk for future opioid abuse, a score of 4 to 7 indicates moderate risk for opioid abuse, and a score of 8 or higher indicates a high risk for opioid abuse.

Mark each box that applies	Female	Male
<b>Family history of substance abuse</b>		
Alcohol	1	3
Illegal drugs	2	3
Rx drugs	4	4
<b>Personal history of substance abuse</b>		
Alcohol	3	3
Illegal drugs	4	4
Rx drugs	5	5
<b>Age between 16—45 years</b>	1	1
<b>History of preadolescent sexual abuse</b>	3	0
<b>Psychological disease</b>		
ADD, OCD, bipolar, schizophrenia	2	2
Depression	1	1
<b>Scoring totals</b>		



Remember-  
Opiates Don't work

Remember- Opiates  
Don't work



N= 3,123 pts from 2000-2011 Longitudinal study w/ 19,201 self reported assessment  
Wolfe et al (2011). *European J Pain*; 17: 581-586

## Limitations of Medications...

# Pharmacotherapy

- The Patients Role is Critical



Patient Materials





**Living with Chronic Pain is....  
like driving with a flat tire**

Need all **4** tires to “move” smoothly...

1. Medical treatments
1. Physical Strength/ endurance
3. Social Support
4. Coping strategies:
  - Relaxation
  - Distractions
  - Pacing/ Pleasant activities
  - Cognitive restructuring
  - Problem Solving/ Goal Setting

**Fill your tires with....**

*Adapted from **Penney Cowen**, founder American Chronic Pain Association.  
[www.theapca.org](http://www.theapca.org) Wonderful patient centered resource!!*

# Realistic Expectations

# Disability and Fibromyalgia

- Strong connection between work and HRQOL
  - Better overall function
  - HRQOL
  - Life satisfaction
- Applying for disability promotes “illness” behaviors
  - Difficulty controlling condition through this process
- Cross Sectional Internet Survey: **Need EARLY ID and Interventions**
- N= 198 (n=73 FT, n=29 PT, n=96 U)
  - Pain intensity: no difference
  - Functional disability & HRQOL : significant
  - Secondary health conditions
  - Depression: employed lowest
  - Social support: highest in employed
  - Life satisfaction: significant differences

# SHARED MEDICAL VISITS

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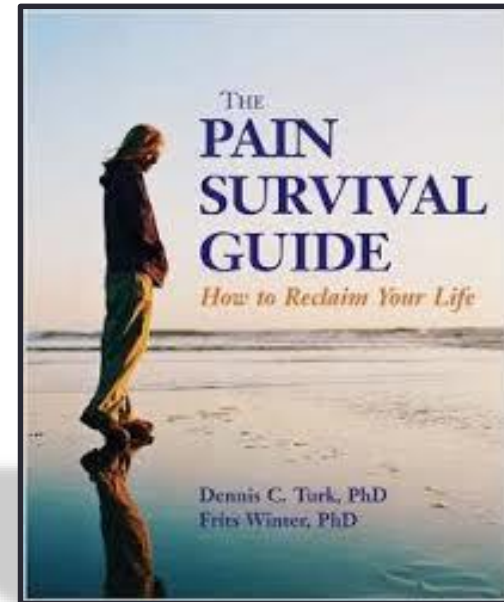
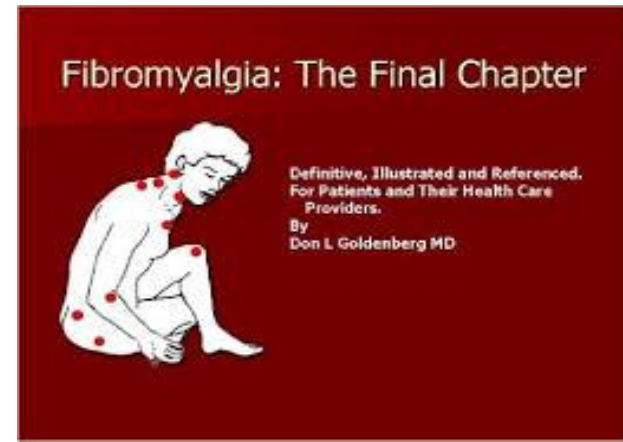
## Solutions for the Fibromyalgia Visit



- 1-2 x month
- Pre visit assessments
- H & P abbreviated to confirm dx
- Group educational session
- Support group
- Lecture Series

*Tara Shuey "Fairylnd of Pain"*  
*From Painexhibit.org*

- Self Management Skill: Running the car... COPING SKILLS
  - Pain Education: understanding mechanisms of control
  - Relaxation: PMR, mini sessions, guided imagery
  - Activity/ Pacing
    - Pleasant Activity scheduling “Homework- 40 Things”
    - Aerobic exercise critical
    - Stretching regimens
    - Learning to PACE activities: life in planned increments
  - Distraction techniques: learning how and when
  - Cognitive restructuring: more than just positive thinking
    - Problem solving
    - Recognizing and Challenging negative thinking patterns
- Resources:
  - Don Goldenberg: The Final Chapter
  - U Michigan’s fibroguide: <http://fibroguide.med.umich.edu/>



# Standard of Care Treatment



# Fibromyalgia Wellness Center Resources

Christine Stamatos DNP, ANP-C



### Websites:

- The Arthritis Foundation  
[www.arthritis.org](http://www.arthritis.org)
- The American College of Rheumatology  
[www.rheumatology.org](http://www.rheumatology.org)
- The American Chronic Pain Association  
[www.theacpa.org](http://www.theacpa.org)
- University of Michigan  
[www.fibroguide.com](http://www.fibroguide.com)
- Calm  
[www.calm.com](http://www.calm.com)
- Headspace  
[www.headspace.com](http://www.headspace.com)
- Pain Online  
<http://www.painonline.org/>
- NIH  
<https://www.nih.gov/health-information>
- Dance Therapeutics  
<http://www.dancetherapeutics.com/>

### Podcasts:

- The Nutrition Diva
- The Savvy Psychologist

### Books:

- Don Goldenberg's eBook
- Fibromyalgia: The Final Chapter*
- Dennis Turk & Fritz Winter's *Pain Survival Guide*
- Clair Davies & Amber Davies *The Trigger Point Therapy Workbook*

### SM Tools:

- Activity Tracking Devices
- Shiatsu Heated, Kneading Massage
- Massage stick
- The Back Knobber.



- Calm
- Headspace
- Relaxation
- Autogenic ~~Tag~~
- ~~Herzife~~
- Gratitude
- Sleep cycle alarm clock
- My fitness pal
- ACPA pain log
- Pain Diary

GEM



**Other Modifiable Factors:** Physical strength & Endurance

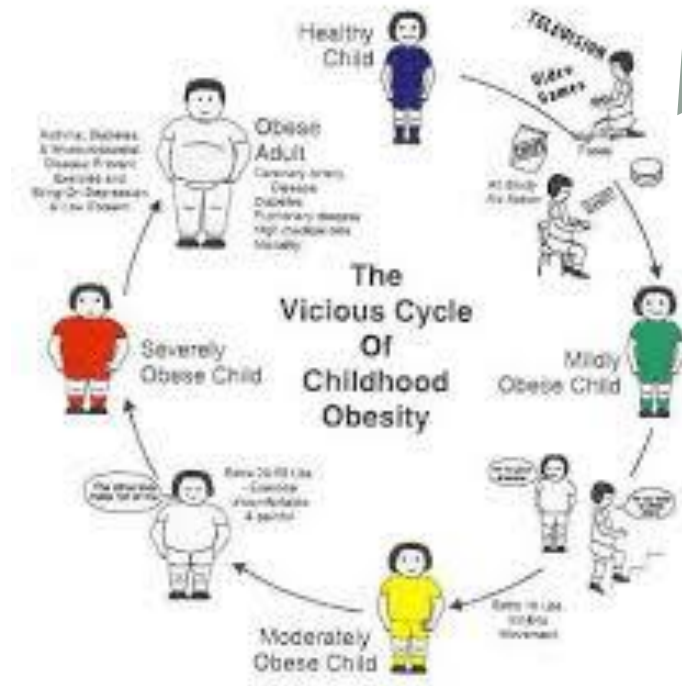


**Fibromyalgia & Obesity, Is there a link?**

**Risk of Fibromyalgia in Obesity**

## Fatigue, Widespread Pain:

- low self esteem
- inactivity
- loss muscle mass
- increased joint pain
- Chronic pain is different...
- Inflammatory state
- Depression
- Sleep disturbance
- Metabolic syndrome
- Cardiovascular challenges
- Unwell state...and worse....



**It starts early...**

**FIBROMYALGIA!**

# Realistic Expectations

- Important: How you THINK and FEEL dramatically impacts your pain.
- Stress/ Fatigue/ Anger/ Sadness/ GUILT
  - INCREASE PAIN!!!!!!
- Good Sleep/ Laughter/ Happiness/ Relaxation
  - DECREASE PAIN!!!!!!
- You have SOME control.
- May not be able to change reason for the PAIN... can learn to change your response to the pain.
- Acceptance: not give up... don't stop fighting... **JUST TAKE THE LEAD!**





# Realistic Expectations



**FEAR**  
is nothing but the anticipation of pain.  
Whether it's...  
physical, mental,  
spiritual or emotional.  
[WWW.LIVELIFEHAPPY.COM](http://WWW.LIVELIFEHAPPY.COM)



**BREAKING THE CYCLE...**

**Kinesiophobia**



## Brief Action Planning

### Provider.... 5 A's

1. **A**ssess- factors affecting health
2. **A**dvice- gives personalized, specific advice
3. **A**gree- collaboratively select
4. **A**ssist- provide assistance where necessary
5. **A**rrange- arrange follow up



Realistic Expectations: Goal Setting

- Ang, D., et al. (2010). CBT attenuates nociceptive responding in patients with fibromyalgia: A pilot study. *Arthritis Care & Research*; 62(5): 618-623.
- Arnold, L., et al. (2004). Family study of FM. *Arthritis & Rheum*; 50(3): 944-952.
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- Clauw, D., Arnold L., & McCarberg, B. (2011). The science of fibromyalgia. *Mayo Clin Proc*; 86(9): 907-911.
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- <http://painexhibit.org/en/galleries/pain-visualized/>

**Thank you!**

**Any Questions?**