



# Mastering fibromyalgia in the medical home: what we know in 2025

Lori Montgomery MD CCFP FCFP CHE
Clinical Associate Professor, Cumming School of Medicine
Department of Family Medicine,
Department of Anesthesiology, Perioperative and Pain Medicine

# **Disclosures: Montgomery**





- no industry affiliations
- teaching honoraria from ACFP, UCalgary, UBC, U of A, CBT Canada
- grant from Hotchkiss Brain Institute (U of C) and Alberta Innovates for a pilot clinical trial of a medication for opioid withdrawal; CIHR Transforming Health with Integrated Care (THINC) grant to study ECHO Pain
- medical leadership roles, PCA and AMA
- some slides from Dr Jessica Kirkwood

#### **Criteria for Success**





- build confidence diagnosing fibromyalgia
- explain the implications of a diagnosis to a patient
- outline an evidence-based treatment approach for fibromyalgia
- access a clinical pathway for FM and a toolkit of useful resources





# **POLLING QUESTIONS**





Your patient, Robin, has been presenting with frequent but variable pain complaints over the last six months. You suspect fibromyalgia — how do you confirm that?





# **Another patient, Alex:**

diagnosed with fibromyalgia 4 years ago, and was coping very well with it. After a recent COVID infection, pain and fatigue are significantly worse, so now Alex is visiting you to talk about medications. What are your options?

# Why are we here?





- prevalence ranges from 3-8% around the world
- 40% of patients referred to a tertiary pain clinic in one study
- increasingly, specialty services are declining referrals







# **Myths**

- women to men 7:1
- a relentlessly progressive disease
- a rheumatologic disease

Creed, Francis. A review of the incidence and risk factors for fibromyalgia and chronic widespread pain in population-based stud





# **Myths**

- women to men 7:1
- a relentlessly progressive disease
- a rheumatologic disease

# **Truth**

- likely closer to 3:1
- remitting/relapsing course, possibly with gradual improvement
- a nociplastic pain condition





#### **NOCIPLASTIC PAIN:**

"pain that arises from <u>altered nociception</u> despite no clear evidence of actual or threatened tissue damage causing the activation of peripheral nociceptors or evidence for disease or lesion of the somatosensory system causing the pain."

Raja SN, Carr DB, Cohen M, Finnerup NB, Flor H, Gibson S, Keefe FJ, Mogil JS, Ringkamp M, Sluka KA, Song XJ, Stevens B, Sullivan N





#### **NOCIPLASTIC PAIN:**

"pain that arises from <u>altered nociception</u> despite no clear evidence of actual or threatened tissue damage causing the activation of peripheral nociceptors or evidence for disease or lesion of the somatosensory system causing the pain."

i.e. this is a disorder of the central nervous system

Raja SN, Carr DB, Cohen M, Finnerup NB, Flor H, Gibson S, Keefe FJ, Mogil JS, Ringkamp M, Sluka KA, Song XJ, Stevens B, Sullivan N

# Fibromyalgia is NOT



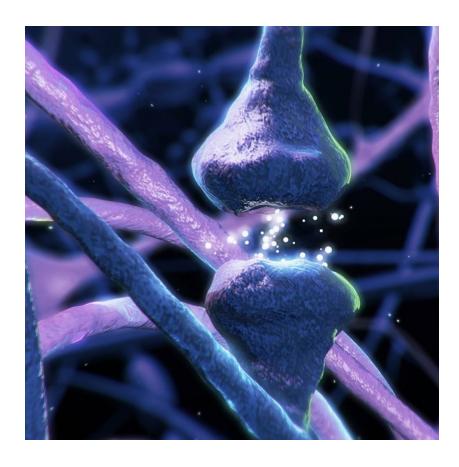


- a musculoskeletal disorder
- a psychiatric disorder
- a maladaptive coping mechanism
- a result of physical deconditioning



### **Small fibre pathology**





- about half of women with FM have reduced skin sensitivity
- small fibre quantity and function appear to be affected - the greater degree of pathology, the higher the symptom load
- this is also associated with CNS changes
- unclear which comes first

#### **Genetic Factors**







- strong familial associations noted for many years
- COMT gene
- serotonin transporter gene
- dopamine receptor gene

#### **Environmental Factors**



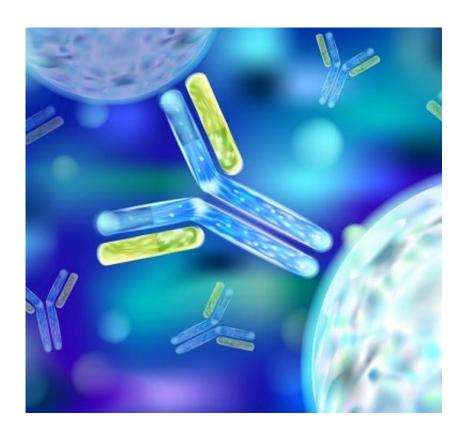




- strong association with trauma noted for many years
- stress, diet, social determinants of health are all noted to be associated with onset and progression of symptoms
- epigenetic factors are being investigated with promising results

#### **Auto-immune Factors**





- autoimmune factors have been postulated since the 1980s, often based on response to steroid therapy
- people with existing autoimmune conditions are at substantially increased risk of developing FM
- recently, there is evidence that FM presentation can be transferred to rodents through IgG infusion

#### Microbiome







- the gut microbiome is different in people with FM (and it isn't just IBS)
- transferring fecal microbiota from people with FM into mice induces pain hypersensitivity
- hypersensitivity in mice is eliminated by transferring fecal microbiota from healthy individuals
- via cross-talk between the gut microbiome and the immune system?

Minerbi, Amir; Khoutorsky, Arkady; Shir, Yoram; Decoding the connection: unraveling the role of gut microbiome in fibromyalgia. PAIN Reports 10(1):p e1224, February 2025. | DOI: 10.1097/PR9.000000000001224

# Not a diagnosis of exclusion

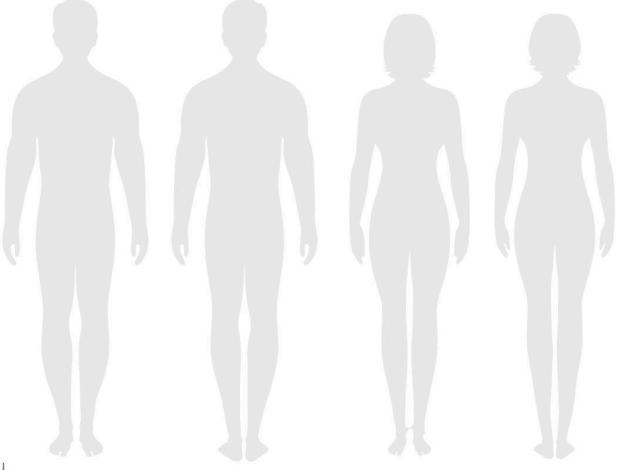




- many things on the differential can be ruled out by history and physical exam
- it has a typical clinical picture

#### Body map

Use the figures to record where pain occurs in detail. Shade the areas of your body where you have felt persistent or recurrent pain for the past 3 months or longer (chronic pain).



#### Calculating the WPI score

Use this checklist to calculate the widespread pain index (WPI) score. Tick the areas where you have had chronic pain for 3 months or longer.

# Region 1: left upper L jaw R jaw R shoulder girdle R shoulder girdle R upper arm R lower arm and/or L wrist/hand, L elbow R wrist/hand, R elbow

#### Region 3: left lower Region 4: right lower Region 5: axial Neck L hip and/or L buttock R hip and/or R buttock Lupper leg and/or L groin R upper leg and/or R groin Upper back L lower leg and/or R lower leg and/or Lower back L ankle/foot, L knee Chest (L and/or R) R ankle/foot, R knee Abdomen

Symptom severity scale (SSS)
------------------------------

Have your problems with the symptoms below been present for 3 months or more?

Yes

No

If yes, using the following scale, indicate the severity of each symptom over the past week by circling the appropriate number.

	No problem	Mild	Moderate	Severe
Fatigue	0	1	2	3
Trouble thinking or remembering	0	1	2	3
Waking up tired (unrefreshed)	0	1	2	3

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

Pain or cramps in lower abdomen

Depression

Headache

Total score\* for the SSS

Yes Yes

No No

 Also associated: TMJ dysfunction, painful bladder syndrome, IBS, sensitivity to light, sound, temperature, and medication side effects

• add 0 (none), 1 (a few), 2 (moderate amount), or 3 (lots)





WPI > 6 and SSS > 4
OR
WPI 3-6 and SSS > 8

 the higher the score, the more likely that FM is the sole explanation for symptoms

# Differential Diagnosis (or comorbid...)





- Inflammatory/rheumatologic conditions (eg SLE, Rheumatoid Arthritis, Polymyalgia Rheumatica)
- Hypermobility spectrum disorders
- Multiple Sclerosis
- Neuropathies/myopathies
- Obstructive sleep apnea

- Hypothyroidism
- Depression
- Post-viral syndromes
- Chronic fatigue syndrome/myalgic encephalitis
- Drug side effects (aromatase inhibitors, lipid lowering agents, high dose opioids)

#### **Treatment**

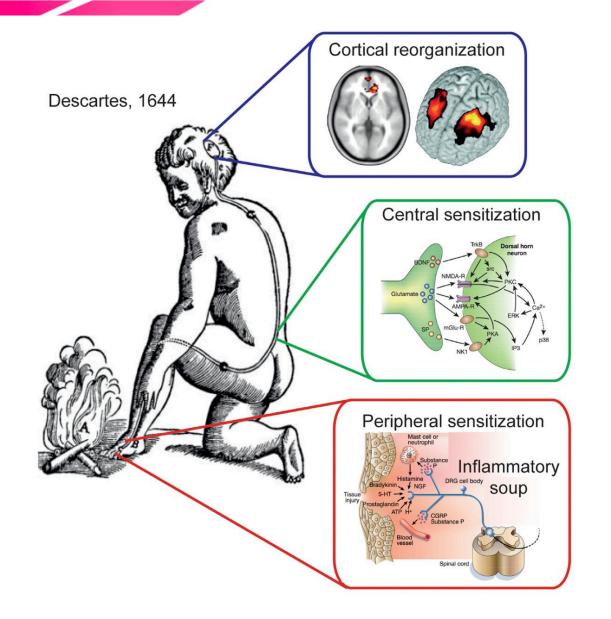






- education about the disorder is the first priority
- evidence is clear that pain science education (PSE) is a treatment that should be started early

Marris, D., Theophanous, K., Cabezon, P., Dunlap, Z., & Donaldson, M. (2019). The impact of combining pain education strategies with physical therapy interventions for patients with chronic pain: A systematic review and meta-analysis of randomized controlled trials. Physiotherapy Theory and Practice, 37(4), 461–472.

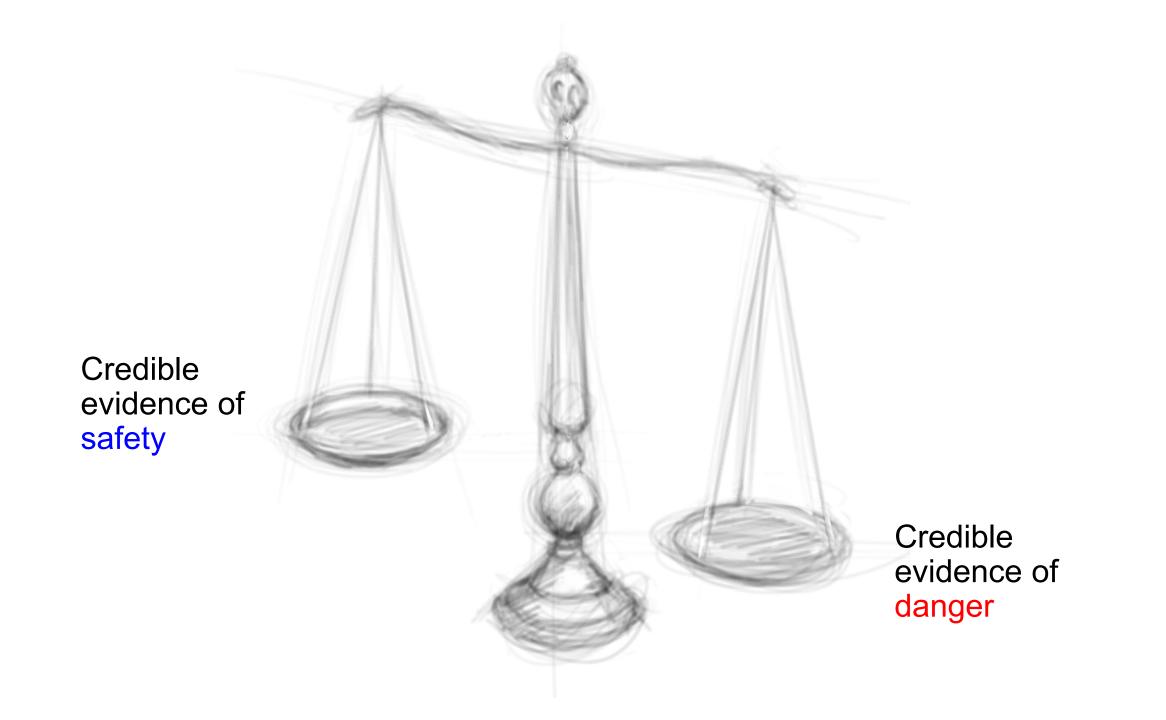




Marwan N. Baliki, A. Vania Apkarian, Nociception, Pain, Negative Moods, and Behavior Selection, Neuron, Volume 87(3): 474-491, 2015

# 1. Pain is protective





Pain is protective
 Persistent pain is overprotective



THE PAIN EXPERIENCE IS

DRIVEN BY

THE BRAIN'S

PERCEIVED NEED

TO PROTECT



1. Pain is protective

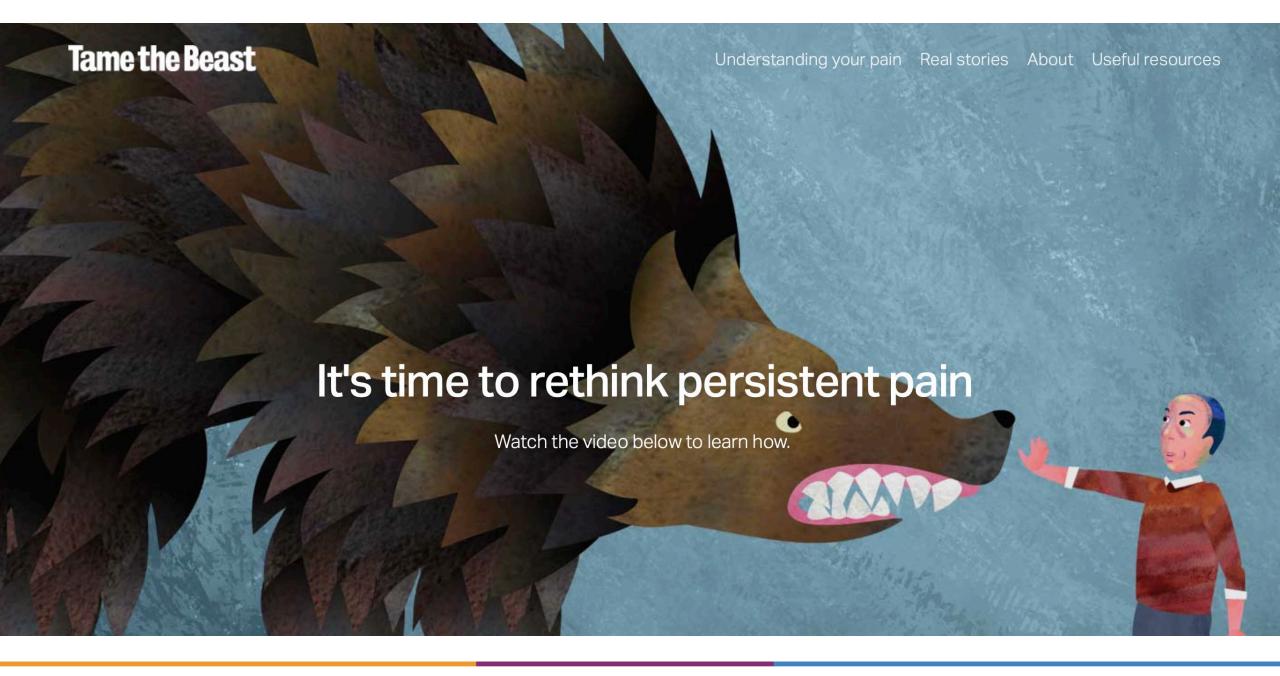
2. Persistent pain is overprotective

3. Many things impact pain, so many things can help





- current environment
- beliefs
- behaviours
- past experiences
- general health



evidence based medical management plan



#### **Tools for Practice**

Tools for Practice articles have been produced by the PEER team in collaboration with the ACFP since 2009.

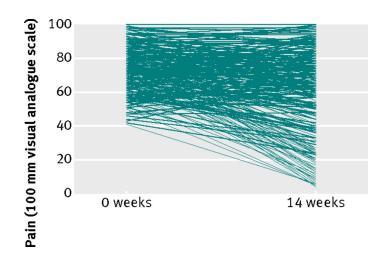
Click here for the entire collection

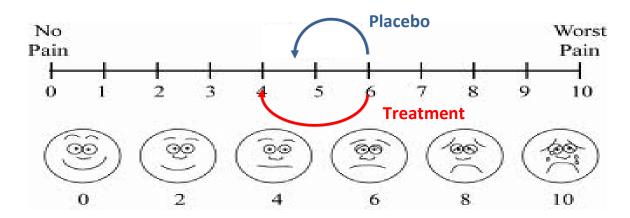


https://peerevidence.ca/tools-for-practice/

#### Pain Outcomes: Change in Scale

- On a 0-10 point scale: Baseline ~6/10.
  - Placebo reduces Pain: ~1.4
    - Treatment: ~2.0





# **Antidepressants**



TFP 375. October 2024.

## **Duloxetine**

- 4 systematic reviews
  - 6-7 RCTs, 2249-2642 patients, 8-27 weeks



- Proportion of patients with ≥30% reduction in pain:
  - 48% vs 36%, NNT=9
  - 30mg versus placebo: no difference
  - 60-120mg: ~50% versus 35% (placebo)
- Withdrawal due to adverse events: no difference
  - Exception: 120mg; 21% versus 11% (placebo), NNH=10.
  - Nausea (26%), constipation (15%), headache (14%) versus 4-8% placebo; hyperhidrosis 8% versus 1% placebo.

# Mirtazepine

### Systematic review

• 3 RCTs, 591 patients, average 30mg daily 7-14 weeks



- >30% pain reduction: 47% versus 34% (placebo), NNT=8
- ≥ 50% pain reduction: no difference

#### Adverse events:

- Adverse event withdrawals: no difference
- Somnolence 41% versus 14% (placebo); reporting any weight gain 19% versus 1% (placebo)



# other antidepressants

## SSRIs (fluoxetine, citalopram, paroxetine):

- Systematic Review (7 RCTs, 383 patients, 6-16 weeks).
- ≥30% reduction in pain: 33% versus 23% (placebo), NNT=10.
- Adverse event withdrawals: no difference

## Amitriptyline:

- Systematic Review (4 RCTs, 275 patients, 25-50mg daily, 8-24 weeks).
- >50% pain reduction: 36% versus 11%, NNT=5.
- Adverse event withdrawals and adverse events: no difference.



## **Antidepressants - Bottom line:**

- 1. In patients with fibromyalgia, meaningful pain reduction (~30% reduction) occurs in ~50% of patients on duloxetine or mirtazapine versus 35% on placebo over 7-14 weeks.
- 2. SSRIs and amitriptyline are likely effective over 6-24 weeks, based on limited evidence.
- 3. Versus placebo, antidepressants do not increase the risk of withdrawing treatment due to adverse effects, with the exception of high-dose duloxetine (120mg).



# Pregabalin

- 150-600mg daily
- systematic review (5 RCTs, 3283 patients, versus placebo, 8-14 weeks).
- ≥30% reduction in pain:
  - 150mg (1 RCT, 263 patients): no difference.
  - 300mg (39%); 450mg (43%); 600mg (39%) versus ~29% placebo.
     NNT=8-10.

# Pregabalin

Adverse events increase with higher daily doses:

Adverse Event	placebo	150mg dose	600mg dose	Number Needed to Harm (NNH)
Somnolence	~5%	16%	23%	6-9
Dizziness	~10%	23%	46%	3-8
Peripheral Edema	~2%	5%	11%	9-33
Withdrawal due to Adverse Events	~10%	10%	28%	6

# Gabapentin

- 1 Systematic Review<sup>3</sup>
- 1 publicly funded RCT<sup>4</sup>, 150 patients, titrated to average daily dose 1800mg); 12 weeks
- ≥30% reduction in pain:
  - 51% versus 31%, NNT=5
- Global improvement "better":
  - 68% versus 35%, NNT=3

#### Adverse events:

- Sedation 24% versus 4%, dizziness 25% versus 9%, NNH=5-7
- Withdrawal due to adverse events: no difference.

## **Bottom line:**

- 1. In patients with fibromyalgia, a meaningful pain reduction (30%) occurs in ~40% with **pregabalin** (300-600mg) compared to 30% with placebo. Somnolence (16-23%) dizziness (23-46%) and peripheral edema (5-11%) are increased compared to placebo.
- 2. **Gabapentin** (~1800mg) may improve pain in 50% compared with 30% placebo based on one small RCT with more dizziness and sedation (16-20%) than placebo.

90-day costs: pregabalin 300mg ~\$95; gabapentin 1800mg ~\$90.

# Less evidence...

- low dose naltrexone 4.5mg daily
  - rationale: elevated levels of circulating endorphins in FM
  - positive effects on spontaneous pain, hyperalgesia
  - very well-tolerated (most common side effect=vivid dreams)
  - small, short studies, often no control
  - no measurement of function
- magnesium supplementation
  - rationale: Mg is typically tissue bound and not measured in serum, so deficiency can be masked
  - studies are small, short, no controls
  - most studies examine Mg in combination with other interventions

Yang, J., Shin, K. M., Do, A., Bierle, D. M., Abu Dabrh, A. M., Yin, Z., ... Mohabbat, A. B. (2023). The Safety and Efficacy of Low-Dose Naltrexone in Patients with Fibromya

# **Exercise**

- almost any kind of exercise demonstrates improvement over baseline



- clinically meaningful changes compared to control are rare
- interventions and controls vary widely
- generally no responder analysis
- difficult to blind these studies
- usually small (not industry funded) studies



#### LivePlanBe+

We know how pain can affect your life. LivePlanBe+ is a program that helps us learn to make small changes that add up to big improvements in our well-being.

Go to Resource

Learn more



#### My SleepWell

Sleepwell has two main goals: 1) to help people with insomnia get their sleep back without medications; and 2) to help people stop taking sleeping pills safely and effectively.

Go to Resource

Learn more



# JNIVERSITY OF CALGARY

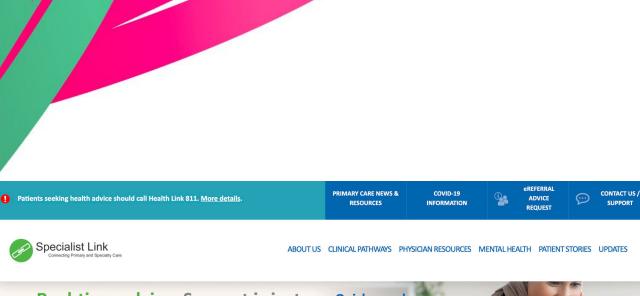
#### **Gentle Movement @ Home**

Guided movement and relaxation videos for pain

Go to Resource

Learn more

https://poweroverpain.ca



#### Real-time advice: Support is just **Quick search** a click away Specialist Link offers a provider-only tele-advice line. Need help? Contact us clinical care pathways and other resources that support Calgary-area

family physicians and nurse practitioners to care for their patients



# specialistlink.ca

#### Primary care pathway: Fibromyalgia

Pathway primer

Expanded details

Provider resources

Patient resources



■ Diagnosis is based on clinical judgement; It is not a diagnosis of exclusion; There is no diagnostic 'gold standard' best made and managed in the Medical Home with the support of a multi-disciplinary team. 

Earlier diagnosis and disclosure are likely associated with lower symptom severity, reduced healthcare costs and improved quality of life.

#### 2. Confirmatory history 🕥

Must be present for FM diagnosis. Core symptoms present for >3 months: ■ Widespread musculoskeletal pain (in four body quadrants plus axial region) 

Fatigue – intrusive – physical, cognitive, emotional ■ Sleep disturbance/non-restorative sleep ■ Symptoms cannot be explained by any other condition



#### 4. Commonly associated symptoms and diagnosis 👂

The more of these symptoms present, the more likely the diagnosis of FM is accurate. ■ Difficulty concentrating/cognitive disturbance

- Depression +/- anxiety may commonly present at time of diagnosis
- Sensitivity to temperature, weather change, light, sound.
- +/- significant sensitivity to medication Migraine and muscular type headache ■ Sleep disorder ■ TMJ disorder ■ Painful bladder syndrome/pelvic pain syndrome ■ Irritable Bowel Syndrome

#### 3. Differential/Co-existing diagnosis More than one may be present, and diagnosis

of FM may still be made: ■ Endocrinology: Hypothyroidism, hyperparathyroidism/hypercalcemia, abnormalities in cortisol

- Rheumatology: Hypermobility spectrum disorders, osteoarthritis, polymyalgia rheumatica, certain myopathic syndromes
- Neurology: Myalgic encephalomyelitis/chronic fatique syndrome, multiple sclerosis
- Respiratory: Obstructive sleep apnea, post-COVID/long-COVID
- Psychiatry: Depression
- Gastroenterology: Celiac disease, irritable bowel svndrome
- Hematology: Iron deficiency anemia, hemochromatosis

#### 5. Review lifestyle and medications 🕥

- Medications Rx and OTC Sleep history
- Movement/exercise history Social history

#### 6. Physical exam

Should be entirely normal unless co-morbidities ■ Rule out differential diagnosis +/- associated symptoms

#### 7. Investigations 🕥

Consider existing co-morbid conditions and the potential for other co-morbidities to occur

Basic screening lab work is a CBC, CRP, TSH, electrolytes and calcium, celiac screen, liver function tests and glucose. 

Other testing done based on clinical suspicion to exclude differential diagnosis and/or associated illness

#### 8. Disclosing the diagnosis 👂

FM is a diagnosis of nervous system processing (known as nociplastic RCP - FM pain). Although it may exist with any of these conditions. FM is not:

■ a musculoskeletal condition;
■ a psychiatric disorder;

■ a maladaptive coping; mechanism; ■ or physical deconditioning

#### information for patients

9. Management 👂 ■ Movement/exercise (strongest evidence) ■ Sleep hygiene/management

- Cognitive Behavioural Therapy for pain Medication (limited evidence)
- Follow up/Chronic disease management plan (in medical home w/team where available). ■ When/where to refer: Community resources: AHS

Alberta Healthy Living Program (online patient education resources) Clinical resources: PCN Multidisciplinary Team/resources, Specialty Care (e.g. FibroFOCUSTM treatment program through the Calgary Chronic Pain Centre, Alberta Virtual Chronic Pain Program)

Community (1) resources

Clinical resources At any point along pathway, consider the following to support clinical decision making:

#### 1. Non-urgent clinical advice:

Call Specialist Link (specialty) specific), incl. chronic pain advice

#### Specialist Link

Submit eReferral Advice Request (chronic pain. endocrinology, gastroenterology, neurology, or long COVID-19)

#### 2. Relevant Specialist Link pathways

3. PCN/Multidisciplinary Team collaboration





Your patient, Robin, has been presenting with frequent but variable pain complaints over the last six months. You suspect fibromyalgia — how do you confirm that?





# **Another patient, Alex:**

diagnosed with fibromyalgia 4 years ago, and was coping very well with it. After a recent COVID infection, pain and fatigue are significantly worse, so now Alex is visiting you to talk about medications. What are your options?





