

Rheumatoid Arthritis, Low Dose Naltrexone, and New Life

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April 27, 2018

Case Study Introduction

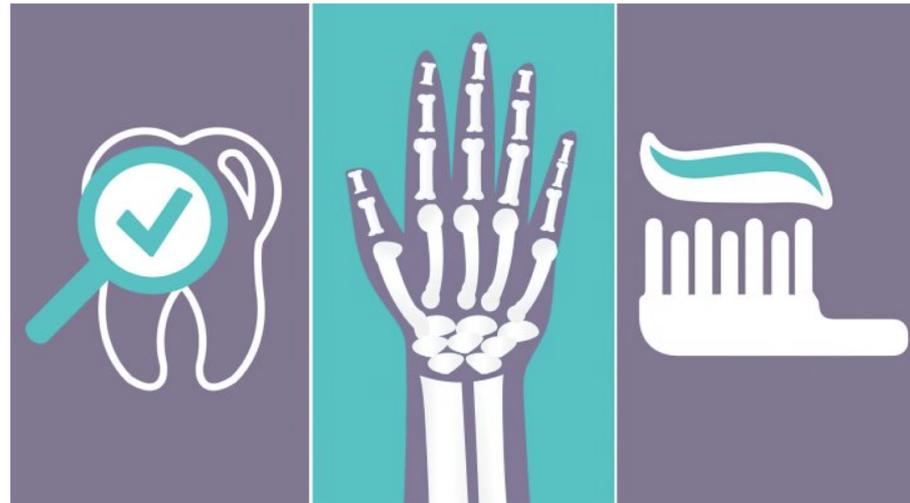
- ▣ 26y old F
- ▣ Rheumatoid Arthritis (RA), poorly controlled
- ▣ Chronic Methotrexate Therapy
- ▣ Presented with desire to wean off of pharmacological agents for RA

Pertinent Past Medical History

- ▣ Grew up eating diet of processed foods
- ▣ Child of divorce
- ▣ Long-term abusive relationship
 - ▣ Verbal, emotional, physical abuse
 - ▣ Lived in fear long-term
- ▣ Joint stiffness and fatigue began almost immediately after ending relationship
- ▣ Hx chronic yeast infections
- ▣ Father w/ hx of Hashimotos hypothyroidism

Rheumatoid Arthritis

- Autoimmune Condition
- 3 Requirements:
 - Genetic Predisposition
 - Environmental Trigger(s)
 - Intestinal Permeability
- Several theories for etiology of RA
 - Infectious mimics
 - Dysbiosis drivers
 - New associations: oral flora/gingival bacteria



Methotrexate (MTX)

- ❑ Contraindicated in pregnancy
 - ❑ Folic acid antagonist
- ❑ Mechanisms of action:
 - ❑ Competitive inhibitor of dihydrofolate reductase (DHFR)
 - ❑ Inhibition of enzymes involved in purine metabolism
 - ❑ Inhibit T cell activation
 - ❑ Selective down-regulation of B cells



Initial Findings

- ▣ Presented with severe pain in both hands, bilateral shoulder pain, and rest of joints “feel bruised”
- ▣ Initial labs unremarkable other than high titers of RF/CCP antibodies
 - ▣ CRP/ESR minimally elevated
 - ▣ Pain out of proportion to findings
- ▣ Elevated Candida antibodies
 - ▣ Tx: Diflucan/Nystatin/Natural Antifungals



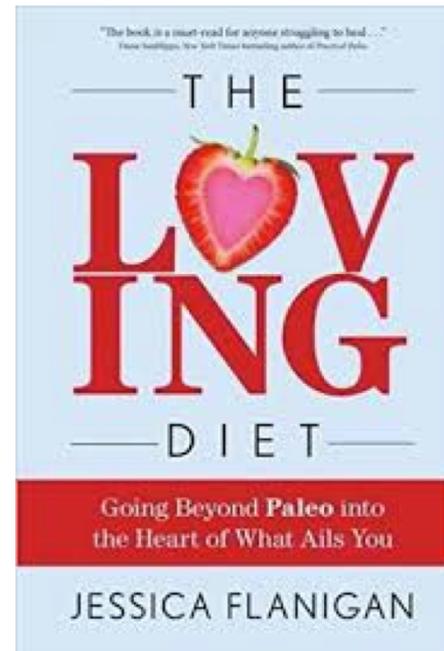
Diet and Lifestyle Changes: Paramount

- ▣ Do not immediately discontinue patient's medications!
- ▣ Must be gradual process
- ▣ Continued her low dose MTX, Folic Acid, work to optimize vitamin D levels
- ▣ Autoimmune dietary template
- ▣ First gluten eliminated
- ▣ Then strict elimination diet:
 - ▣ Autoimmune Paleo with low FODMAP approach
- ▣ Address trauma → referral for EMDR therapy



Short-Term Elimination Diet Stressed

- ❑ Do not wish to reduce oral tolerance
 - ❑ Reduction of microbial diversity
- ❑ Do not attempt long-term restriction diets in setting of chronic gut infections or presence of dysbiotic flora
 - ❑ E.g. “Commensals”
- ❑ Autoimmune Paleo template
 - ❑ “Loving Diet” by Jessica Flanigan
 - ❑ Helped with yeast overgrowth sx
 - ❑ Nightshade-Free diet may be important for RA patients



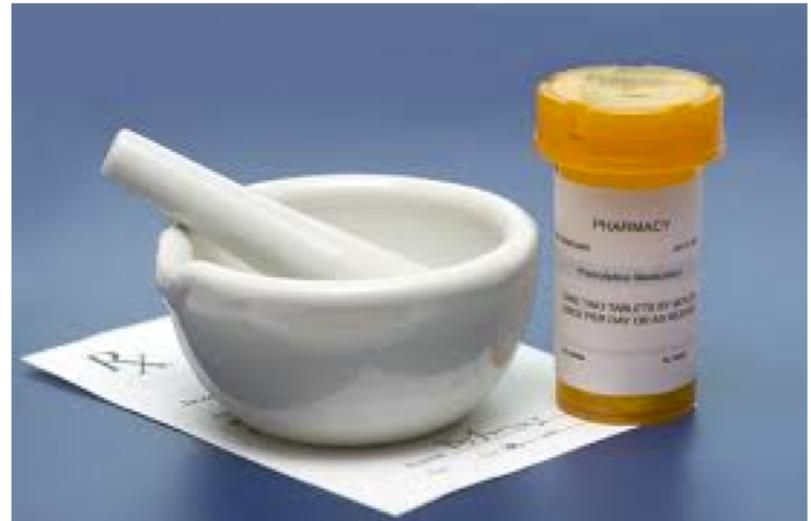
One Month Later...

- ▣ Joint pain much improved
 - ▣ Swelling confined to R middle finger, +synovitis
 - ▣ No further shoulder pain
 - ▣ Can clearly see knuckles on left thumb and fingers



Low Dose Naltrexone Prescribed

- ▣ Dosing: Bihari Protocol
- ▣ Compounded 1.5mg → 3.0mg → 4.5mg oral tablet
- ▣ Increased dose every 2 weeks
- ▣ Nightly dosing
- ▣ Side Effects: none



Symptoms Continued To Improve

- ▣ After 6 months on LDN
 - ▣ Weaned methotrexate
- ▣ Patient attributed residual hand stiffness to overuse
- ▣ Methylation testing, folate level testing



3 Months Later...

- ▣ Patient discovered she was unexpectedly pregnant
- ▣ Maintained on 4.5mg low dose naltrexone throughout pregnancy and post-partum period
 - ▣ Extensive research and data on pregnancy & LDN (Dr. Phil Boyle)
- ▣ Uneventful pregnancy and delivery of full-term, healthy female infant
- ▣ No complications



Flare 8 Weeks Post-Partum

- ▣ Post-partum autoimmune flares common secondary to immune system shifts
 - ▣ Th1/Th2 mediated
- ▣ Patient's RA symptoms quickly mitigated by targeted anti-inflammatory supplementation and immune system support
 - ▣ Omega-3 Fatty Acids
 - ▣ SPMs (Specialized Pro-Resolving Mediators)



Proposed MOAs

- ▣ LDN and Rheumatoid Arthritis
 - ▣ Opioid antagonism
 - ▣ Increased endogenous beta endorphins and met enkephalins
 - ▣ Novel microglial cell modulator
 - ▣ Suppression of activation
 - ▣ Prevent production of reactive oxygen species →
 - ▣ Reduction of neuro-excitotoxicity →
 - ▣ Reduction of pro-inflammatory signaling cascade
- ▣ LDN shown to profoundly benefit conditions associated with elevated inflammatory markers
 - ▣ ESR, CRP

Conclusion

- ▣ Low Dose Naltrexone is an important consideration for the reduction of chronic inflammation from autoimmune, and musculoskeletal/neurological conditions
- ▣ Must treat underlying co-morbidities (especially gut infections) and optimize nutrition to maximize efficacy

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LDN 2018 Conference Presentation
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