



**FUNCTIONAL
MEDICINE**

Continuing Education

The Autoimmune Puzzle: Interconnection Between Immune Disease & the Gut

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Disclosures

- No financial relationships with any commercial interests to disclose

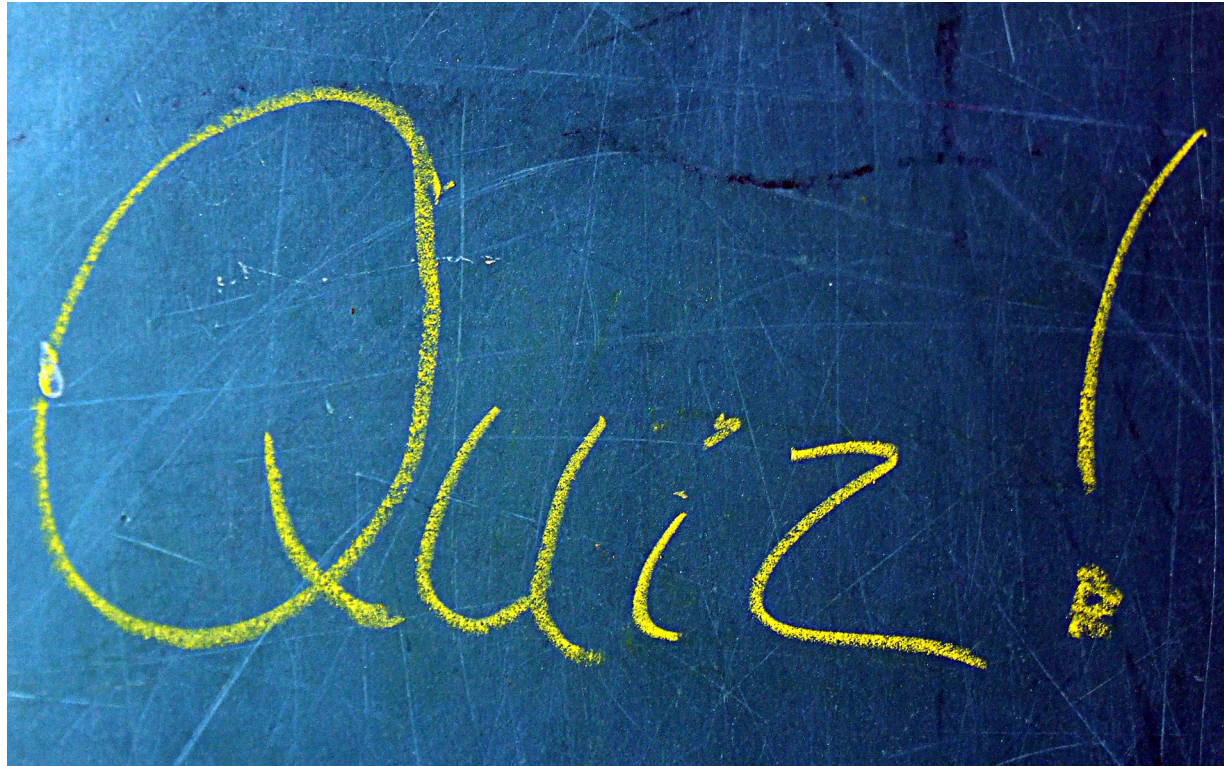
Today's Agenda

1. Describe the pathophysiological pathways that lead to allergic and autoimmune disease
2. Examine complex interaction of immune disease as it relates to GI integrity
3. Identify appropriate dietary modifications for patients with immune disorders.

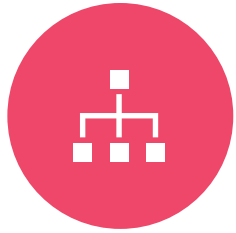
Autoimmune Disease vs Allergic Disease



Let's test our foundational knowledge



Autoimmunity: The Functional Medicine Window



SYSTEM'S
BASED
APPROACH



UPSTREAM
MEDICINE



VS



"SYMPTOM-
BASED"
APPROACH



DOWNSTREAM
MEDICINE

What is autoimmunity?

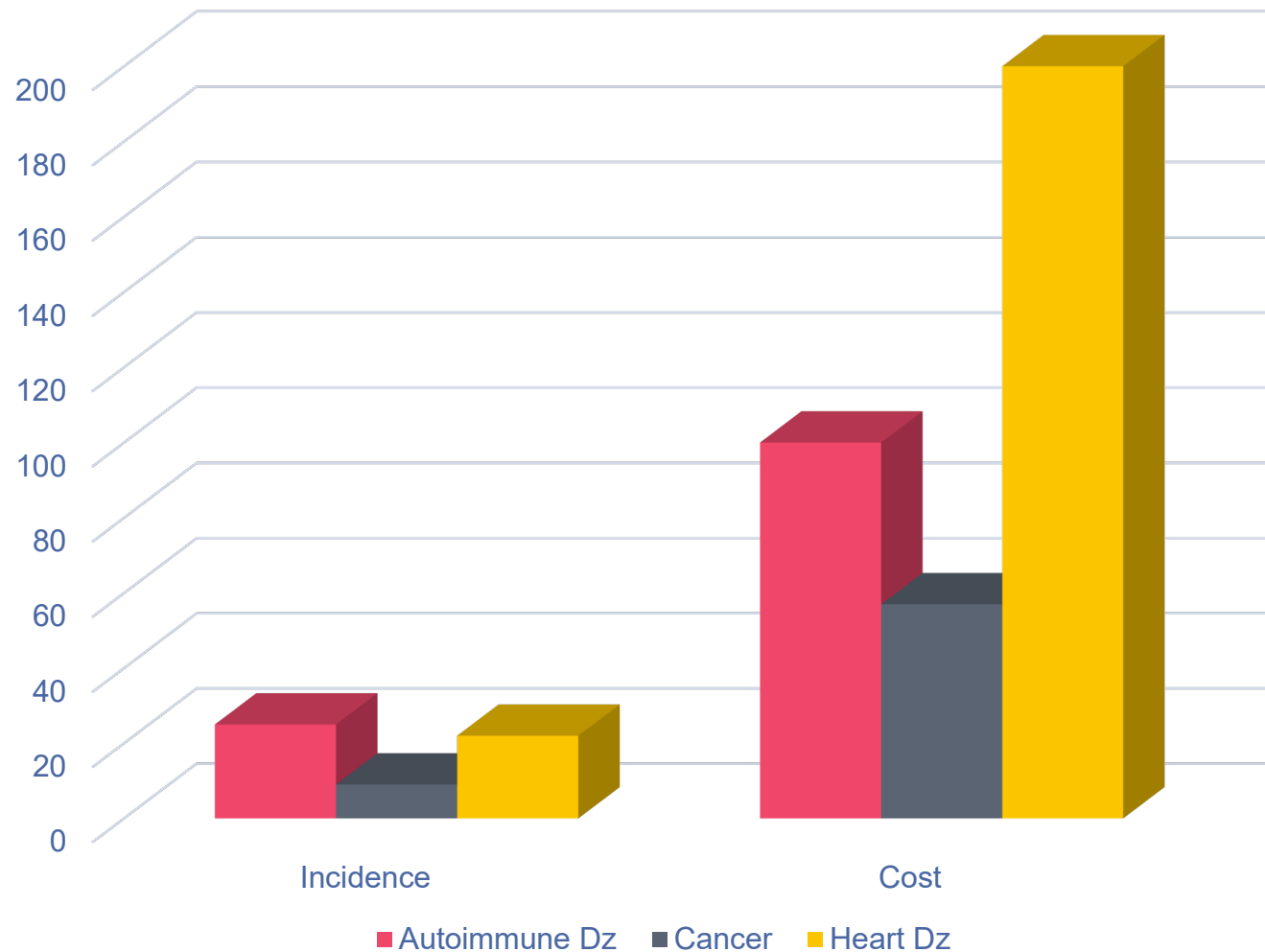
80 types of autoimmune disease as defined by the NIH	Defined as	Etiology	Prognosis/treatment
• Hoshimoto's, celiac, psoriasis, T1D, Lupus, MS, RA, etc...	• Genetically driven disease with some environmental factors • Autoantibodies, CRP, ANA, complement testing, ESR, immunoglobulin testing, genetic testing, electrophoresis	• 23.5 million Americans according to the NIH* • More common among women and white patients	• Difficult to diagnose • No cure, but can reduce the symptoms • Pharmacotherapy – DMARDs, anti-TNF biologics, other biologics, steroids, NSAIDS , opioids

Autoimmune Disease



**American
Autoimmune**

Related Diseases Association, Inc.



NIH incidence (US):

- AD 23.5 million*
- Cancer 9 million
- Heart disease up to 22 million

Cost (NIH, ACS, AHA):

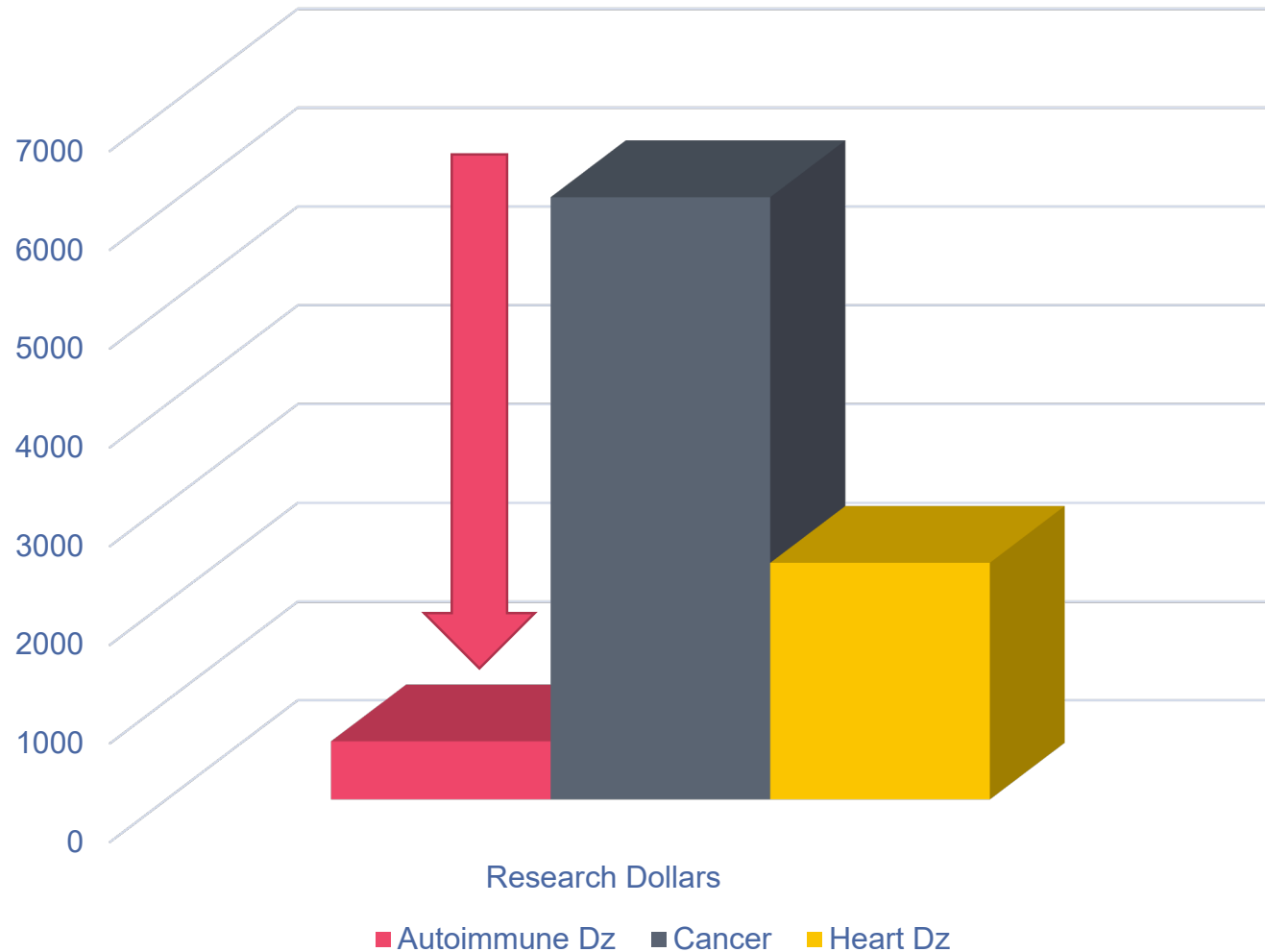
- AD ~\$100 **billion**
- Cancer ~\$57 **billion**
- Heart and stroke costs ~\$200 **billion**

Autoimmune Disease



**American
Autoimmune**

Related Diseases Association, Inc.



2003 NIH Research Funding:

- AD **\$591 million**
- Cancer **\$6.1 billion**
- Heart disease and stroke **\$2.4 billion**

The problem with our current treatment “options” for treating Autoimmune Disease

- Disproportionate funding vs cost and disease burden.
- Immunosuppressant treatments have high side effect burden and lead to significant long-term side effects.
- *“Specialists are generally unaware of interrelationships among the different autoimmune diseases or advances in treatment outside their own specialty area.”*
- Medications are often ineffective and offer symptom management, without addressing the root cause of disease.

Medications used in Autoimmune disease

NSAIDs

Glucocorticoids

Conventional Disease-Modifying Anti-Rheumatic Drugs (cDMARDs)

- Methotrexate
- Leflunomide
- Gold compounds, sulfasalazine, azathioprine, cyclophosphamide, antimalarials, d-penicillamine, cyclosporine

Anti-TNF Biologics:

- Infliximab
- Etanercept
- Adalimumab
- Golimumab
- Certolizumab Pegol

Cost and Effectiveness of Biologics for Rheumatoid Arthritis in a Commercially Insured Population

Jeffrey R. Curtis, MD, MS, MPH; Benjamin Chastek, MS; Laura Becker, MS; Caroleen Quach, MSPH;
David J. Harrison, PhD; Huifeng Yun, PhD; George J. Joseph, PhD, MS; and David H. Collier, MD

ABSTRACT

BACKGROUND: Administrative claims data, diagnosis, and procedure data, but the literature on rheumatoid arthritis (RA) historically has limited effectiveness research. A claims-based algorithm was developed to estimate effectiveness for RA from claims data and modifications.

OBJECTIVE: To implement the claims-based algorithm in the Optum Research Database to estimate biologic effectiveness for RA.

METHODS: The cohort included patients with RA in the Optum Research Database who were diagnosed between January 2007 and December 2010, and who had at least 12 months before through 12 months after the index date. Patients were categorized as effectively treated by the claims-based algorithm if they met all of the following criteria in the post-index period: (1) a medication claim for a biologic, or at least as many claims for intravenous biologics; (2) no increased use of biologics; (3) no new nonbiologic disease; (4) no new or increased oral glucocorticoid treatment; and (5) no more than 1 glucocorticoid injection. Drug costs (all biologics) and administration costs (intravenous biologics) were obtained from allowed amounts on claims. Biologic cost per effectively treated patient was defined as total 1-year biologic cost divided by the number of patients categorized by the algorithm as effectively treated with that index biologic. Sensitivity analysis was conducted.

“The index biologic was categorized as effective by the algorithm for **28.9% of patients overall**, including 30.6% for subcutaneous biologics and 22.1% for intravenous biologics. The index biologic was categorized as effective in the first year for **32.7% of etanercept (794/2,425)**, **32.3% of golimumab (40/124)**, **30.2% of abatacept (89/295)**, **27.7% of adalimumab (514/1,857)**, and **19.0% of infliximab (147/773)** patients.”

- This study used the validated algorithm to estimate the cost of effective biologic treatment from adjudicated claims in 5,474 patients with RA who received an approved biologic between 2007 and 2010.
- Using the claims-based algorithm, mean 1-year biologic cost per

Reasons for Drug Discontinuation

- Cost of Immune modulating Rx = \$1-3K/month
- Drug inefficacy varies in studies (as high as 84%) cited as the #1 reason cited for discontinuation of drug therapy
- They don't cure, they manage symptoms (downstream)
- Serious long-term risks
- Efficacy often short-lived (agent switching required)

It's not enough, we need a better solution

The Functional Medicine Perspective

1. Your genes play a role, but your environmental exposures (toxins, “bugs” etc), lifestyle (stress, sleep, exercise), and diet play a larger role in outcome

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2. **Autoimmune disease is a spectrum, it moved backwards and forwards (not just progressively worse)**

The Functional Medicine Perspective

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2. Autoimmune disease is a spectrum, it moved backwards and forwards (not just progressively worse)
3. **All autoimmune disease is rooted in the same primary disorder → the immune system. It’s not an “organ-specific” disease but rather a whole system.**

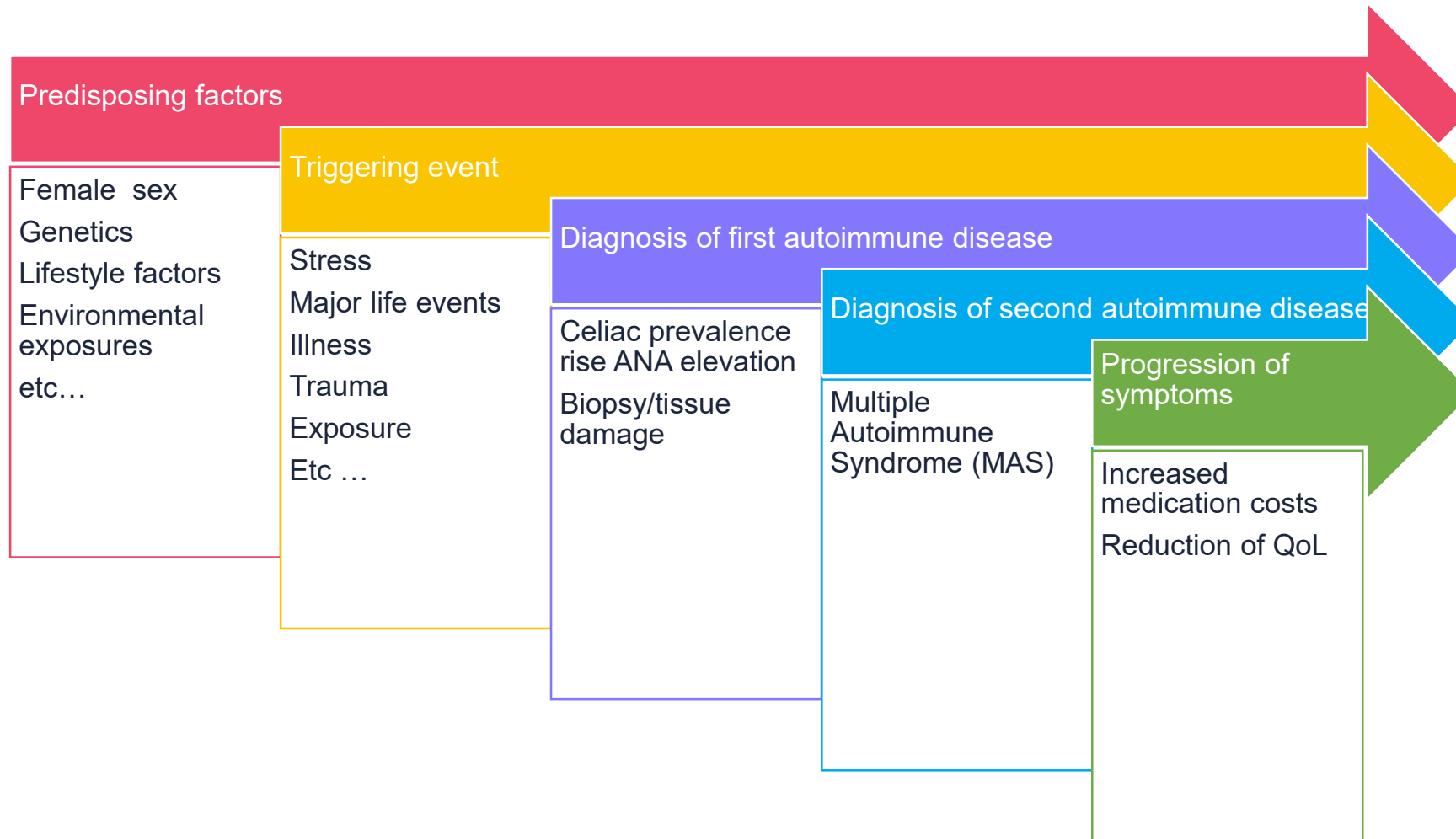
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4. **Symptom-relief with medication is an option, but it isn’t the only option**

The Functional Medicine Perspective

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3. All autoimmune disease is rooted in the same primary disorder → the immune system. It’s not an “organ-specific” disease but rather a whole system.
4. Symptom-relief with medication is an option, but it isn’t the only option
5. **Diet plays a key role in both the etiology and the remission of autoimmune disease. So does lifestyle modification.**

Autoimmune disease exists on a spectrum



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Harpreet S, et al. *Reumatologia*. 2016;54(6):326–329

<https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5241371/>

FUNCTIONAL MEDICINE MATRIX

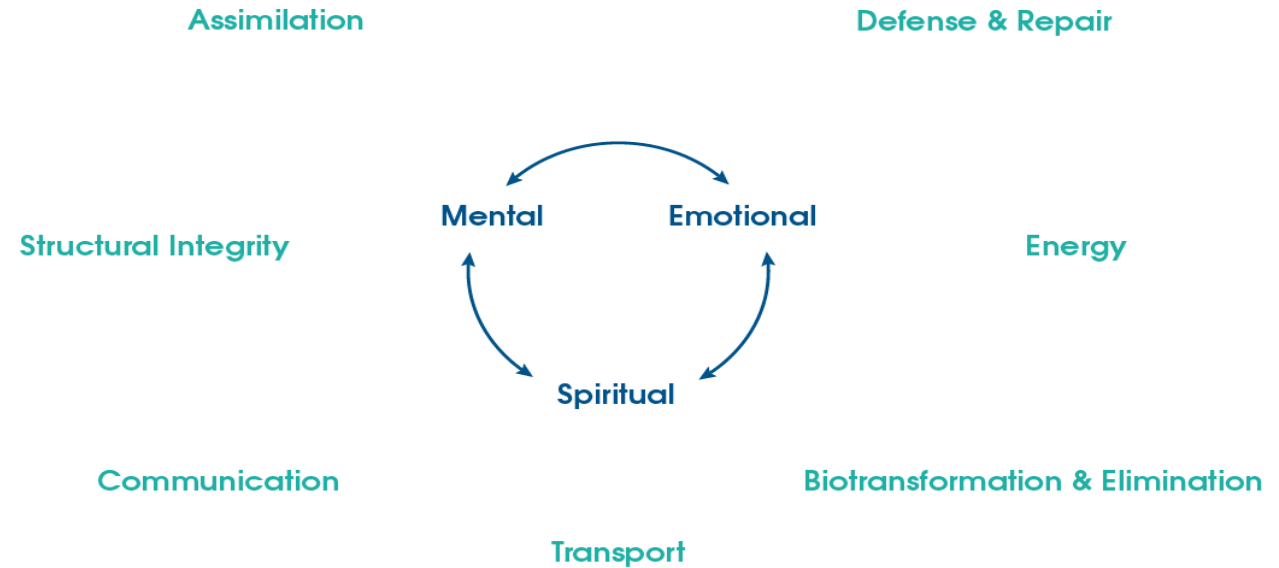
Retelling the Patient's Story

Antecedents

Triggering Events

Mediators/Perpetuators

Physiology and Function: Organizing the Patient's Clinical Imbalances



Modifiable Personal Lifestyle Factors

Sleep & Relaxation

Exercise & Movement

Nutrition

Stress

Relationships

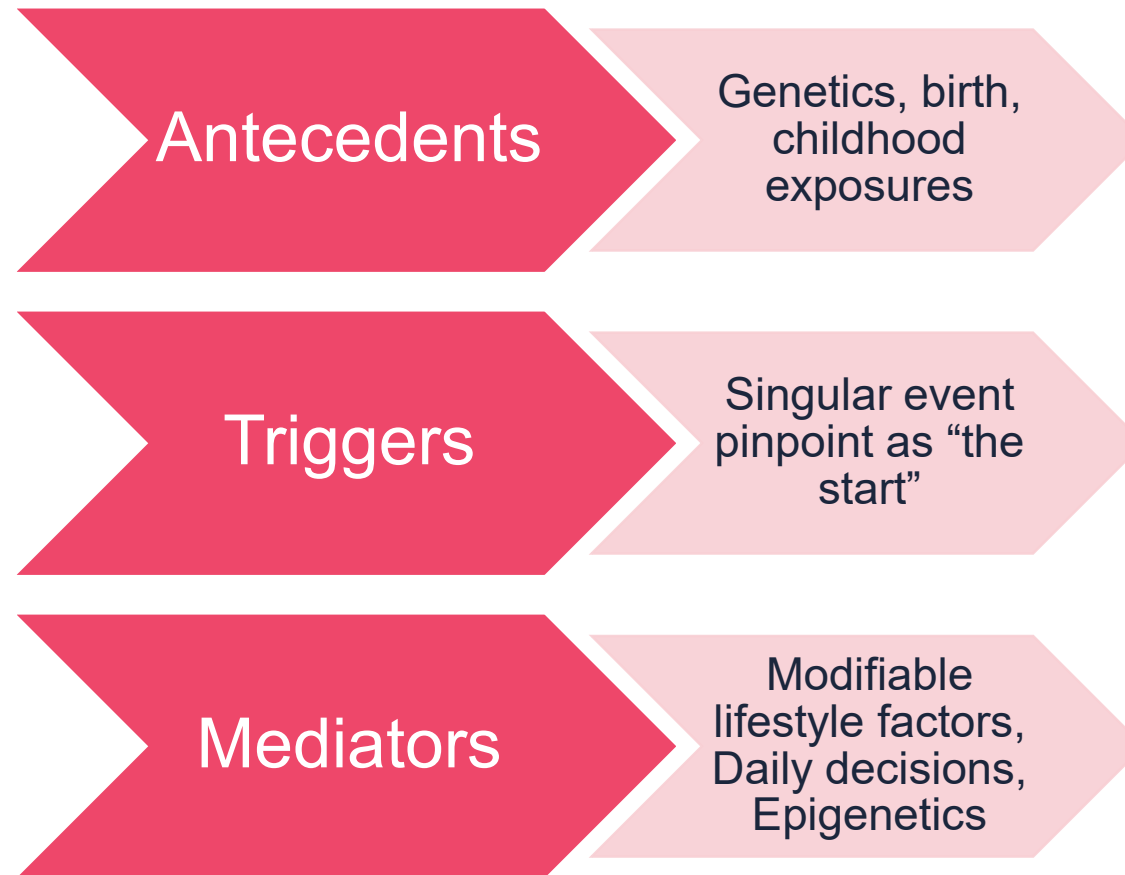
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Version 2



The *ATMs*



FUNCTIONAL MEDICINE TIMELINE

Mediators/Perpetuators

Antecedents

Triggers or Triggering Events

Preconception

Prenatal

Birth

Current Concerns

Signs, Symptoms or Diseases Reported

Name: _____

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Version 3



Pathophysiology Review

How do you feel about Immunology?



Defense & Offense

Physiological barriers

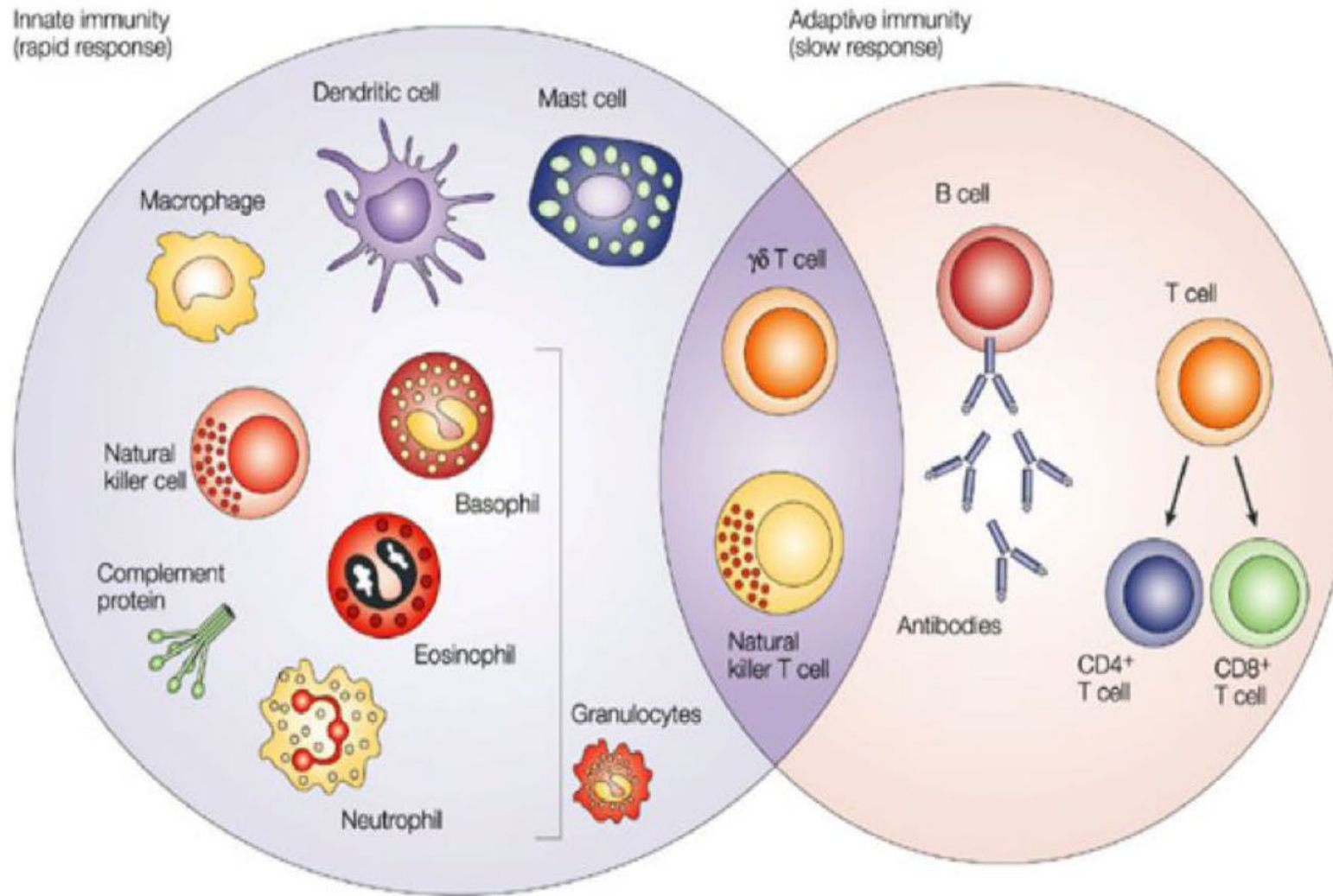
- VALT/GALT/MALT/BALT/NALT

Innate vs adaptive

- Cell mediated (T-cell)
- Humoral (immunoglobins)



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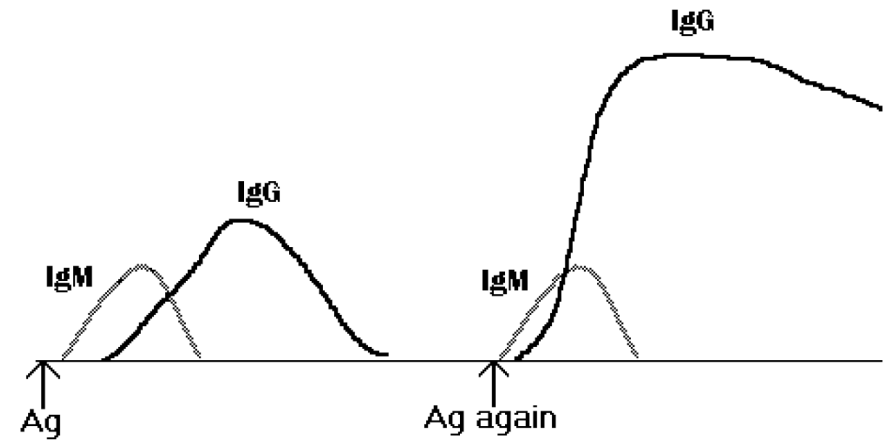


Innate & Adaptive Immunity

Dranoff G. Cytokines in cancer pathogenesis and cancer therapy. Nat Rev Cancer. 2004 Jan;4(1):11-22. Review.

Nature Reviews | Cancer

Innate Immunity



IgM

- Oldest antibody phylogenetically; 500 times more efficient than IgG at activating complement.

IgA

- Made by plasma cells in lymphoid tissues near mucous membrane
- **Secretory Component** - protective against invaders in the gut (first line of immunological defense)

IgG

- Most abundant in circulation
- Plasma half-life of IgG is about **3 weeks**
- **Opsonizing** - vital for clearance of most extracellular bacteria.

IgE

- Type I Immunopathology due to histamine-containing mast cells and basophils activation
- Protective for parasites.

Compliment Cascade

Compliment cascade – 3 Pathways to activation:

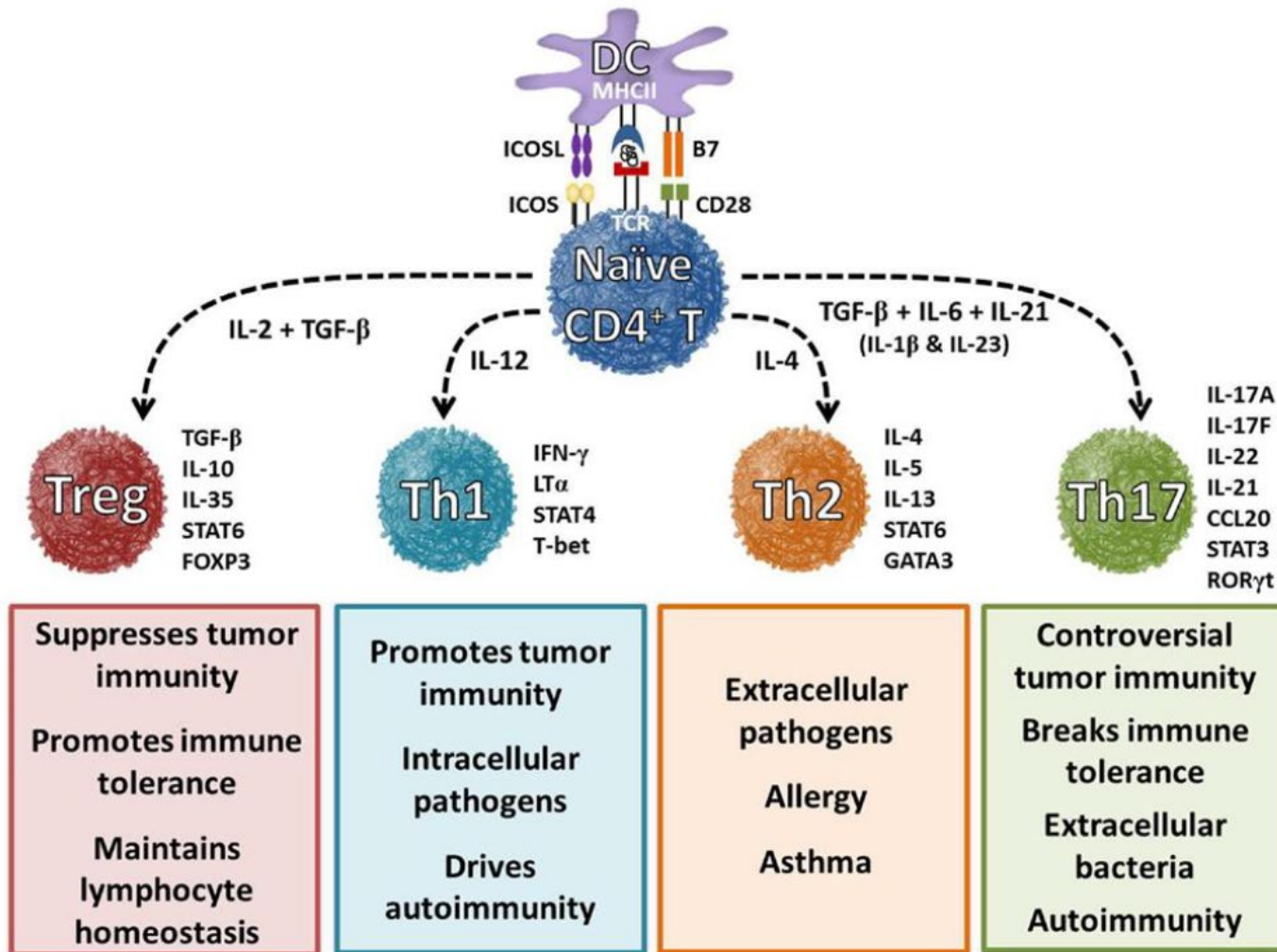
- Classic
- Alternative
- Lectin

Results in 4 actions:

- Lysing – cell breakdown
- Opsonizing - phagocytosis
- Chemotactic – chemokine release
- Anaphylactic – mast cells, basophile response

Th17 cells in cancer: the ultimate identity crisis

Stefanie R. Bailey^{1,2*}, Michelle H. Nelson^{1,2}, Richard A. Himes³, Zihai Li¹, Shikhar Mehrotra² and Chrystal M. Paulos^{1,2*}



Adaptive Immunity

What's the tipping point?



FUNCTIONAL MEDICINE MATRIX

Retelling the Patient's Story

Antecedents

Genetics/Family History
C-section birth
Frequent childhood antibiotic use

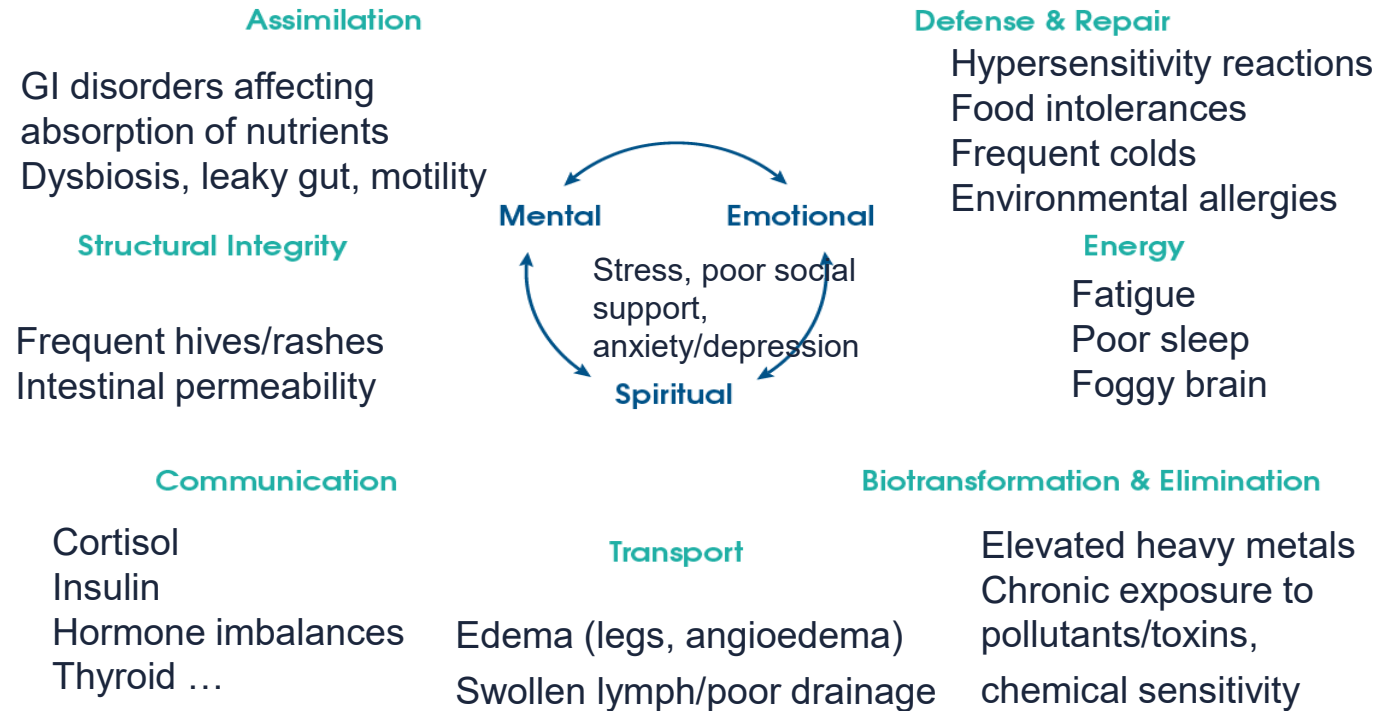
Triggering Events

Tick bite
Home renovation
Family stress, traumatic event
Physical injury or surgery

Mediators/Perpetuators

Standard American diet
Chronic infection (viral, bacterial, parasite)
Intestinal permeability/dysbiosis/SIBO

Physiology and Function: Organizing the Patient's Clinical Imbalances



Modifiable Personal Lifestyle Factors

Sleep & Relaxation

Sleep hygiene

Exercise & Movement

Regular exercise
Type, frequency

Nutrition

Glycemic load
Plant-based, variety
Color
Inflammatory foods

Stress

Amount of stress and stress management

Relationships

Social network
Friends and family
Stressful or supportive?



Pathogenesis and treatment of autoimmune rheumatic diseases

Eric Liu and Andras Perl

KEY POINTS

- Autoimmunity occurs in the setting of genetic predisposition and environmental influence.
- Both genetic and environmental factors modulate the innate and adaptive arms of the immune system towards self-reactivity and inflammation.
- Metabolic pathways serve as key sensors and effectors of genetic, epigenetic, and environmental cues that shape the development and function of the immune system.
- Activation of the mechanistic target of rapamycin has emerged as a central regulator of metabolic stress that underlies pro-inflammatory lineage development in the adaptive immune system.
- Blockade of mTOR reverses the depletion of Tregs and CD8 memory T cells and moderates the expansion of inflammatory Th17 cells in autoimmune diseases with promising therapeutic efficacy.



Prevention of allergy with diverse and healthy microbiota: an update

Diana Chernikova^a, Irene Yuan^b, and Marcus Shaker^{b,c,d}

Purpose of review

Microbiota consist of symbiotic microscopic neighbors that have been described, clear effective therapeutic interventions have been described, clear effective therapeutic interventions have been described, clear effective therapeutic interventions have been described.

Recent findings

The human microbiome is influenced by multiple factors (e.g., cesarean section), breastfeeding, diet, presence of medications (particularly antacids), lifestyle, and development of atopic responses and tolerance have been described. Metabolic pathways to promote the health of their immune system away from pro-allergic, Th2-dominated responses to more Th1-regulatory-influenced behaviors.

Summary

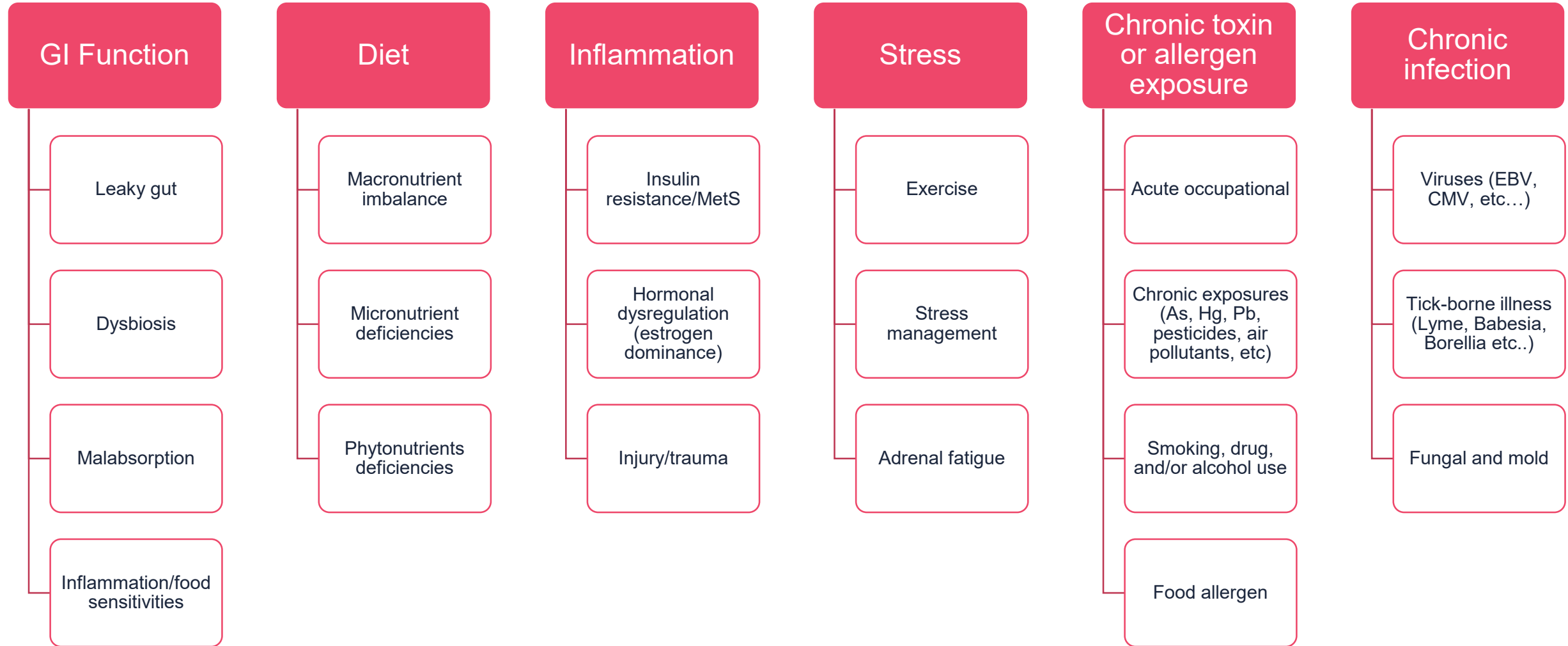
Evidence suggests that specific healthy infant microbiome signatures may influence development of some components of the allergic march of childhood by decreasing atopic dermatitis, asthma, and food allergy. Further understanding of factors that influence healthy microbiota may lead to specific strategies tailored for early intervention and disease prevention.

Keywords

allergy, asthma, atopic dermatitis, microbiome, microbiota, prevention

Evidence suggests that specific healthy infant microbiome signatures may influence development of some components of the allergic march of childhood by decreasing atopic dermatitis, asthma, and food allergy.

Immune Disease Drivers (ATMs)



Autoimmune Triggers

PAMPS

- Bugs (bacterial, viral, parasites, candida, mold)

DAMPS

- Trauma or surgery

Food reactivity or increased sensitivity

- Advanced glycation end products (AGEs)
- IgG/IgA/IgE development
- Mannose binding lectins
- FODMAP/Sulphur/histamine sensitivity

Toxins acute exposure or increased chemical sensitivity

- Mercury exposure (Amalgam removal)
- Daily household exposure or home renovations (new furniture, carpet removal, etc)



Contents lists available at ScienceDirect

Biochemical Pharmacology

journal homepage: www.elsevier.com/locate/biochempharm

Review

Adipokines: Linking metabolic syndrome, the immune system, and arthritic diseases

Vera Francisco^{a,1}, Clara Ruiz-Fernández^{a,1}, Jesús Pino^a, Antonio Mera^b, Miguel A. González-Gay^c, Rodolfo Gómez^d, Francisca Lago^e, Ali Mobasher^f, Oreste Gualillo^{a,*}

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^e Molecular and Cellular Cardiology Group, SERGAS (Servizo Galego de Saude) and IDIS (Instituto de Investigación Sanitaria de Galicia) (Inflammatory Diseases), Santiago University Clinical Hospital, Santiago de Compostela, Spain

^f Department of Regenerative Medicine, State Research Institute Centre for Inn

ARTICLE INFO

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Leptin
Lipocalin-2
Metabolic syndrome
Obesity
Osteoarthritis
Rheumatoid arthritis

ABSTRACT

Metabolic syndrome (MetS) is a cluster of obesity and visceral adiposity that increase the risk of cardiovascular diseases, such as type 2 diabetes and atherosclerosis. Advances in the understanding of the pathogenesis of MetS have identified as a possible link between

obesity has been established as a significant contributing factor to the increased prevalence of MetS. Adipokines, which play important physiological roles in metabolic activities contributing to the pathogenesis of MetS, are also involved in the regulation of autoimmune and/or inflammatory processes associated with arthritic diseases. Therefore, MetS and dysregulated secretion of pro-inflammatory adipokines have been recognized as a molecular link between CVD and arthritis diseases. In the present paper, we review recent evidence supporting the role played by adipokines, in particular leptin, adiponectin, and lipocalin-2, in the modulation of the immune system, MetS and arthritic diseases. The underlying cellular and molecular mechanisms are discussed, as well as potential new therapeutic strategies.

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Published in final edited form as:

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The gut microbiome shapes intestinal immune responses during health and disease

June L. Round* and Sarkis K. Mazmanian*

*Division of Biology, California Institute of Technology, Pasadena, California 91125, USA

Abstract

Immunologic dysregulation is the cause of autoimmunity, allergy and cancer. The gas between the host immune system and micro discuss findings which indicate that develop influenced by intestinal bacterial colonization mediate host–symbiont interactions that recent evidence to support an emerging cor result in immunological dysregulation that disease. Perhaps the mammalian immune sys fact, controlled by the microbes themselves.

“The gastrointestinal tract is the primary site of interaction between the host immune system and the microorganism, both symbiotic and pathogenic... **disturbances in the bacterial microbiota result in immunological dysregulation that underlie disorders such as inflammatory bowel disease.**”



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The gut microbiome shapes intestinal immune responses during health and disease

June L. Round* and Sarkis K. Mazmanian*

*Division of Biology, California Institute of Technology, Pasadena, California 91125, USA

Abstract

Immunologic dysregulation is the cause of many autoimmune diseases, including autoimmunity, allergy and cancer. The relationship between the host immune system and the gut microbiome is discussed, and we discuss findings which indicate that dysregulation is influenced by intestinal bacterial colonization. We discuss how these mediate host–symbiont interactions that lead to the recent evidence to support an emerging paradigm that the result in immunological dysregulation that may underlie disorders such as inflammatory bowel disease. Perhaps the mammalian immune system which appears designed to control microbes is, in fact, controlled by the microbes themselves.

“Perhaps the mammalian immune system which appears designed to control microbes is, in fact, controlled by the microbes themselves.”

Environmental exposures and mechanisms in allergy and asthma development

Liza Bronner Murrison, ... , Jocelyn Biagini Myers, Gurjit K. Khurana Hershey

J Clin Invest. 2019;129(4):1504-1515. <https://www.jci.org/articles/view/124612>

Review Series

Environmental exposures interplay with the development and progression of allergic diseases. This is as a result of complex gene-environment interactions and host response. Research shows an association between increasingly modern Westernized lifestyles, including urbanization, time spent indoors, and air pollution, with increased exposure to air and traffic pollution, and other early-life and lifelong risk factors for the development and exacerbation of allergic diseases. It is increasingly recognized that the timing, load, and route of allergen exposure affect allergic disease phenotypes and development.

Understanding the mechanisms and interaction of environmental, viral, and allergen exposures with immune pathways that impact disease development. This Review highlights epidemiologic and mechanistic evidence linking environmental exposures to the development and exacerbation of allergic airway responses.

Research shows an association between the rise of allergic diseases and increasingly modern Westernized lifestyles, which are characterized by increased urbanization, time spent indoors, and antibiotic usage. These environmental changes result in increased exposure to **air and traffic pollution, fungi, infectious agents, tobacco smoke, and other early-life and lifelong risk factors for the development and exacerbation of asthma and allergic diseases**. It is increasingly recognized that the timing, load, and route of allergen exposure affect allergic disease phenotypes and development.

So ...

- Immune dysregulation might be linked to general rise of chronic disease like cardiometabolic disease, which in turn, impacts inflammation.
- The GI houses the “cockpit” of the immune system - major site of immune regulation
- The microbiome is having a two-way conversation involving epigenetic expression and immune modulation and we can go as far as to say that the host is, “***in fact, controlled by the microbes themselves***”
- GI integrity (physiological, biochemical, and immunological barrier). Leaky gut when combined with genetic risk and contributing factors can increase risk of development of autoimmune disease

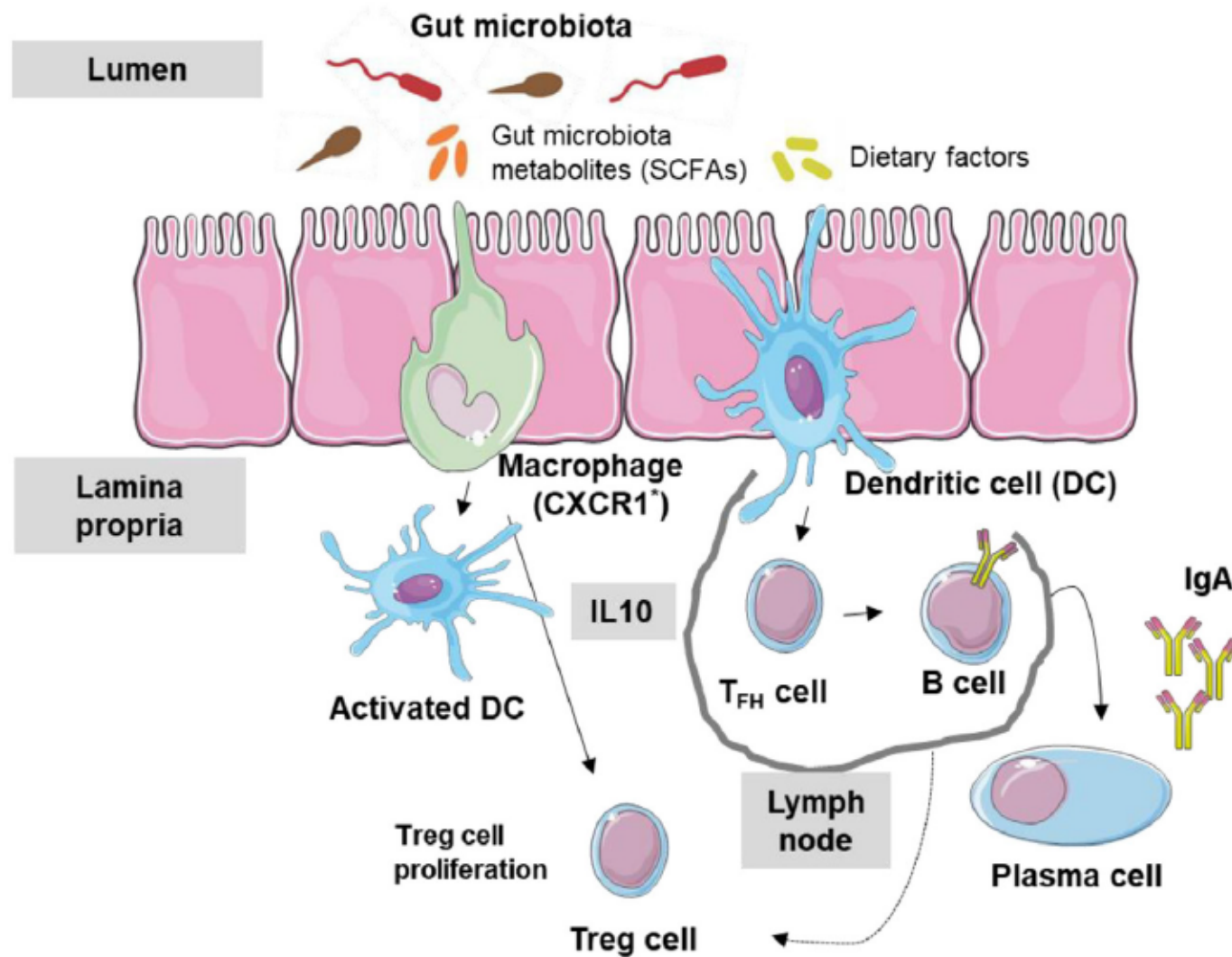


FIGURE 1 | Interaction between gut microbiota and immune system. Gut microbiota metabolites and dietary factors constitute the main antigen load of the gastrointestinal tract. Macrophages (CXCR1⁺) and dendritic cells (DCs) are stimulated and T regulatory (Treg) cells are activated by metabolic products such as short chain fatty acid (SCFA). Follicular T cells activate B cells inducing the production of IgA antibodies.

A few (more) comments on allergic disease and the gut

Allergy (atopic sensitization) is considered the strongest risk factor for development childhood asthma.

The microbiota is influenced by multiple environmental and dietary factors, and has a complex role in development of allergic diseases.

Environmental exposures and lifestyle have been implicated as primary drivers in the dramatic rise of childhood allergies.



Lynch SV. Gut Microbiota and Allergic Disease. New Insights. *Ann Am Thorac Soc*. 2016.

<https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5015732/>

Pascal M, et al. Microbiome and Allergic Diseases. *Front Immunol*. 2018;9:1584.

<https://www.ncbi.nlm.nih.gov/pmc/articles/PMC6056614/>

Modulating Risk: The connection to the gut

Altered gut microbial colonization in very early life has been linked with increased risk of allergies.

- Severe food- or aero-allergens have a significantly higher risk of developing asthma.

Key factor influencing gut microbiome diversity is infant feeding/breastfeeding

- Lactobacilli and *Bifidobacteria*, oligosaccharides, range of fatty acids which impact microbiome and capacity to produce metabolites that protect against allergies and asthma.
- *Lachnospiraceae* and *Ruminococcaceae* impact gut microbiome by producing SCFAs, acting as an essential energy source for gastrointestinal colonocytes and exerting anti-inflammatory effects that modulate allergic disease (MOA increasing epithelial barrier function and inducing Treg cells, DCs precursors, and IL-10 production).

Supplementing Probiotics and prebiotics show promise for the development of a preventive therapy.

- MOA likely due to restoring microbiome balance, reducing dysbiosis, and improving immune response.

More studies with larger sample sizes are needed to dissect the role of microbiome in allergic diseases.

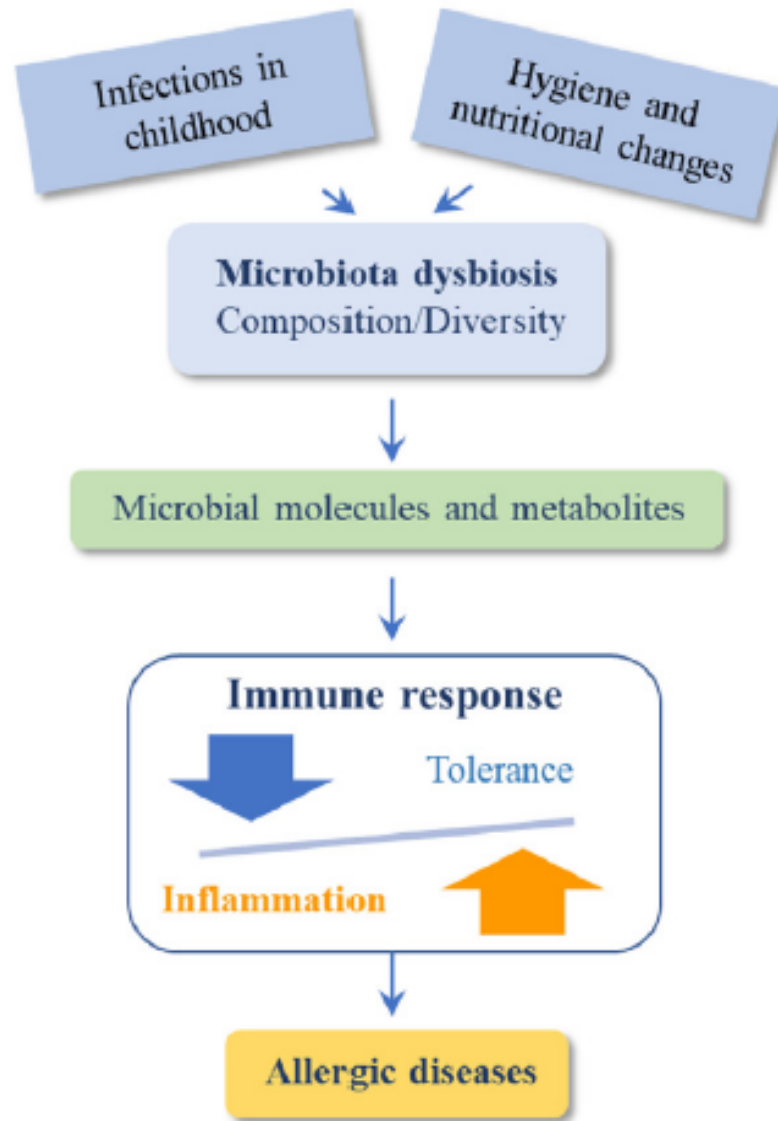
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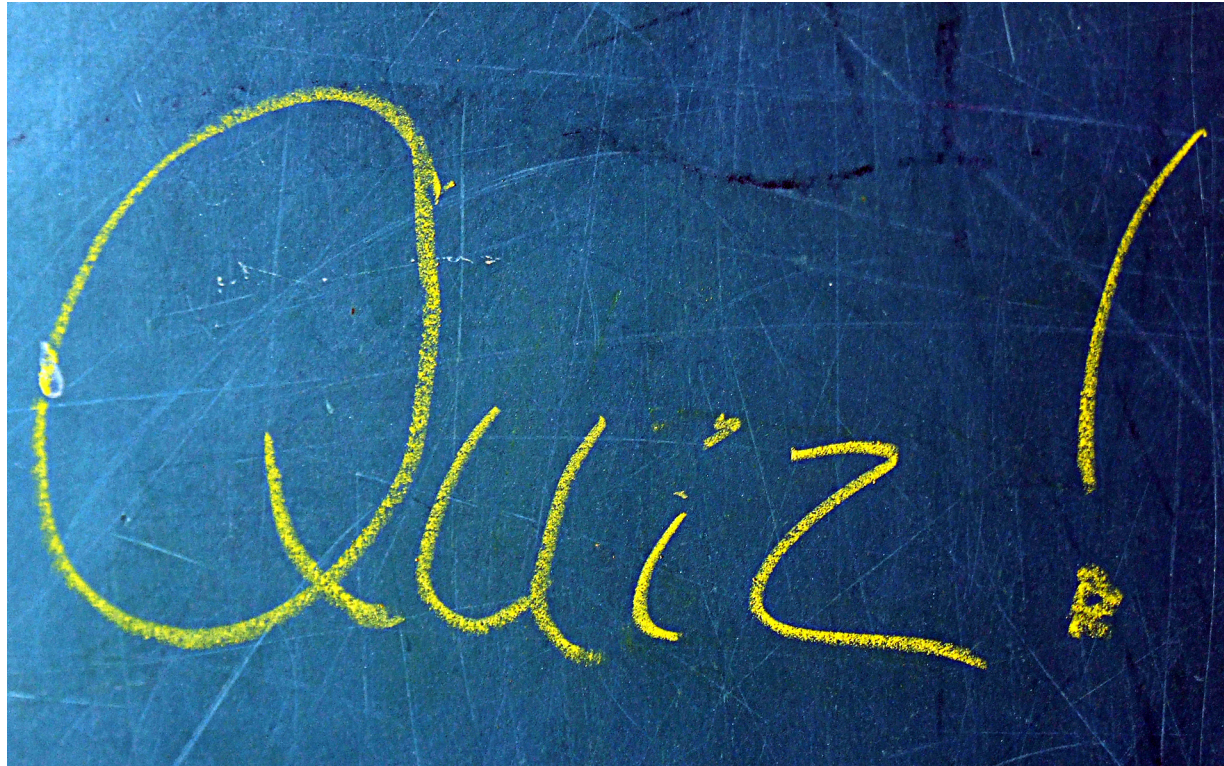
In Summary...



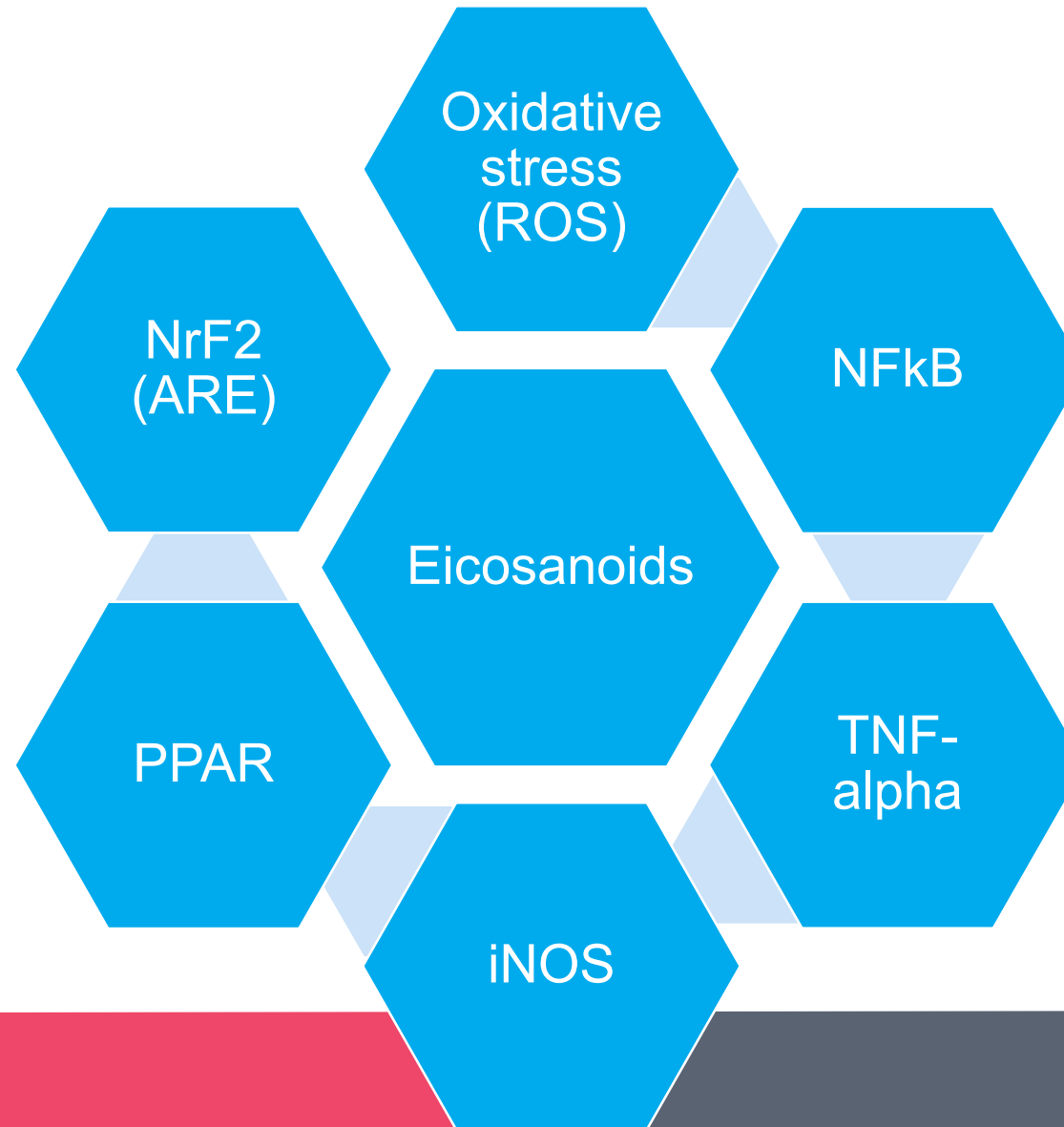
Pascal M, et al. Microbiome and Allergic Diseases. *Front Immunol.* 2018;9:1584.
<https://www.ncbi.nlm.nih.gov/pmc/articles/PMC6056614/>

FIGURE 2 | Dysbiosis induce qualitative and quantitative changes in the microbiota that directly affect immunological mechanisms leading to allergic diseases.

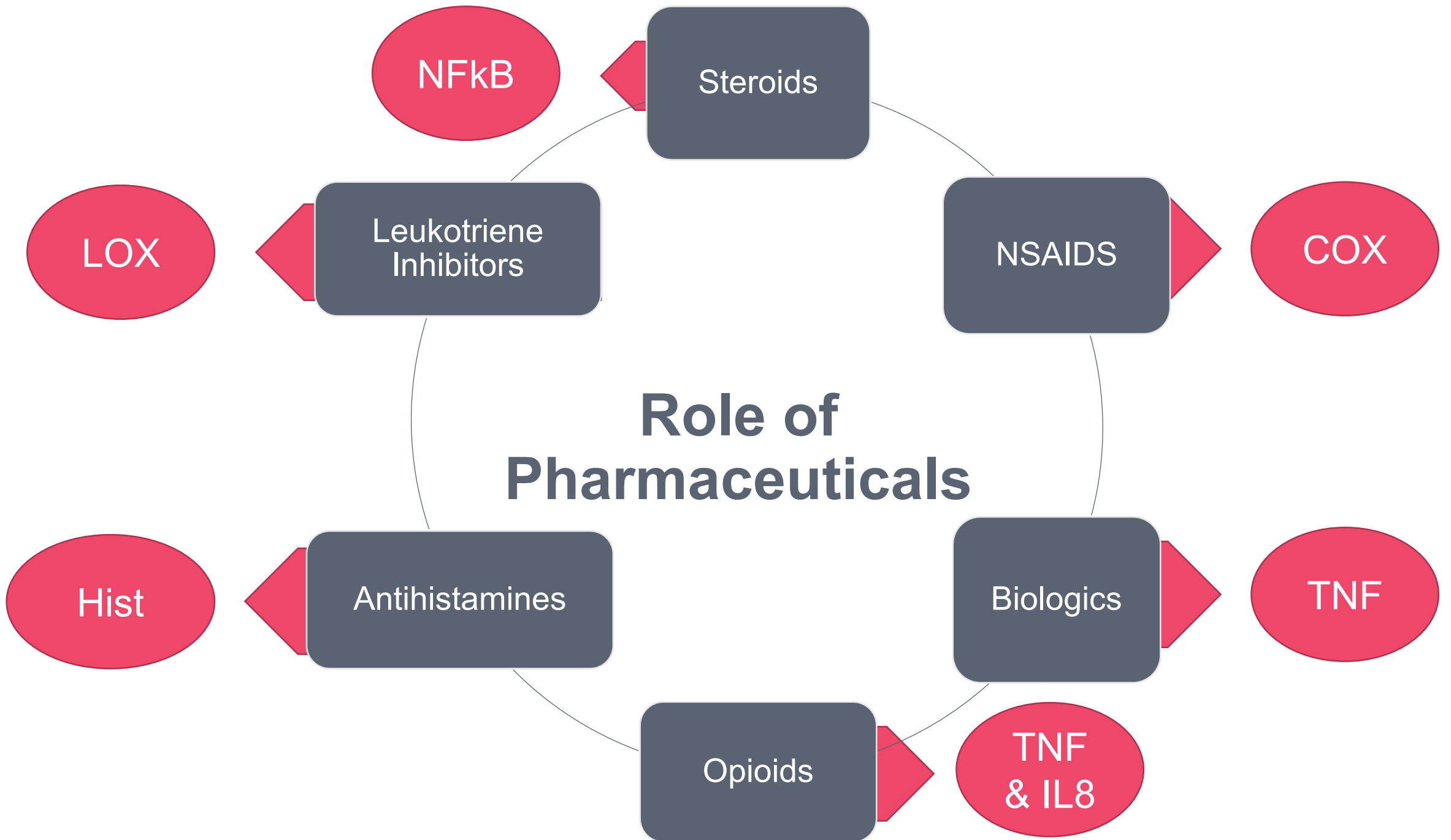
Let's test our foundational knowledge



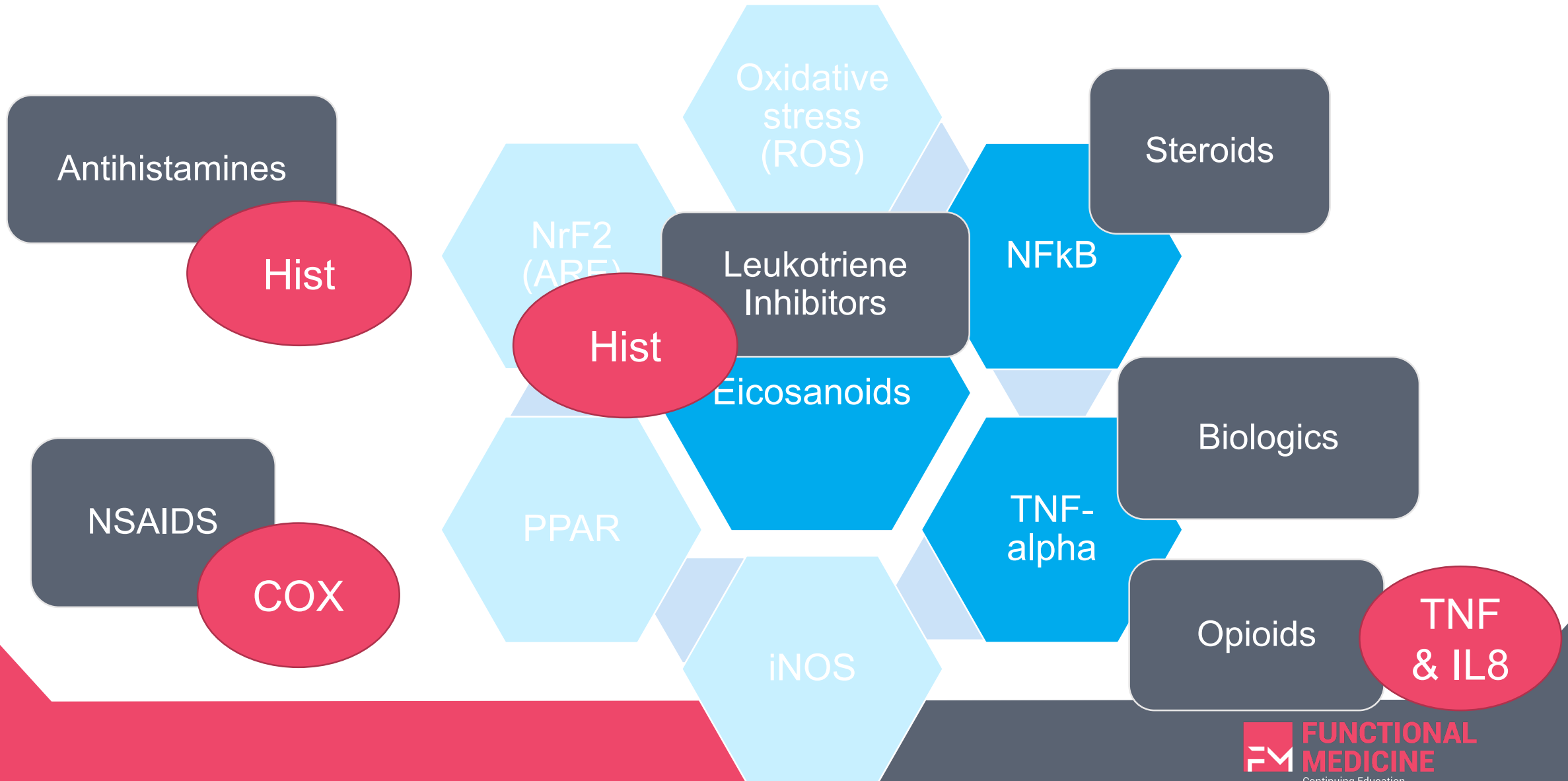
Mediating Inflammatory Response



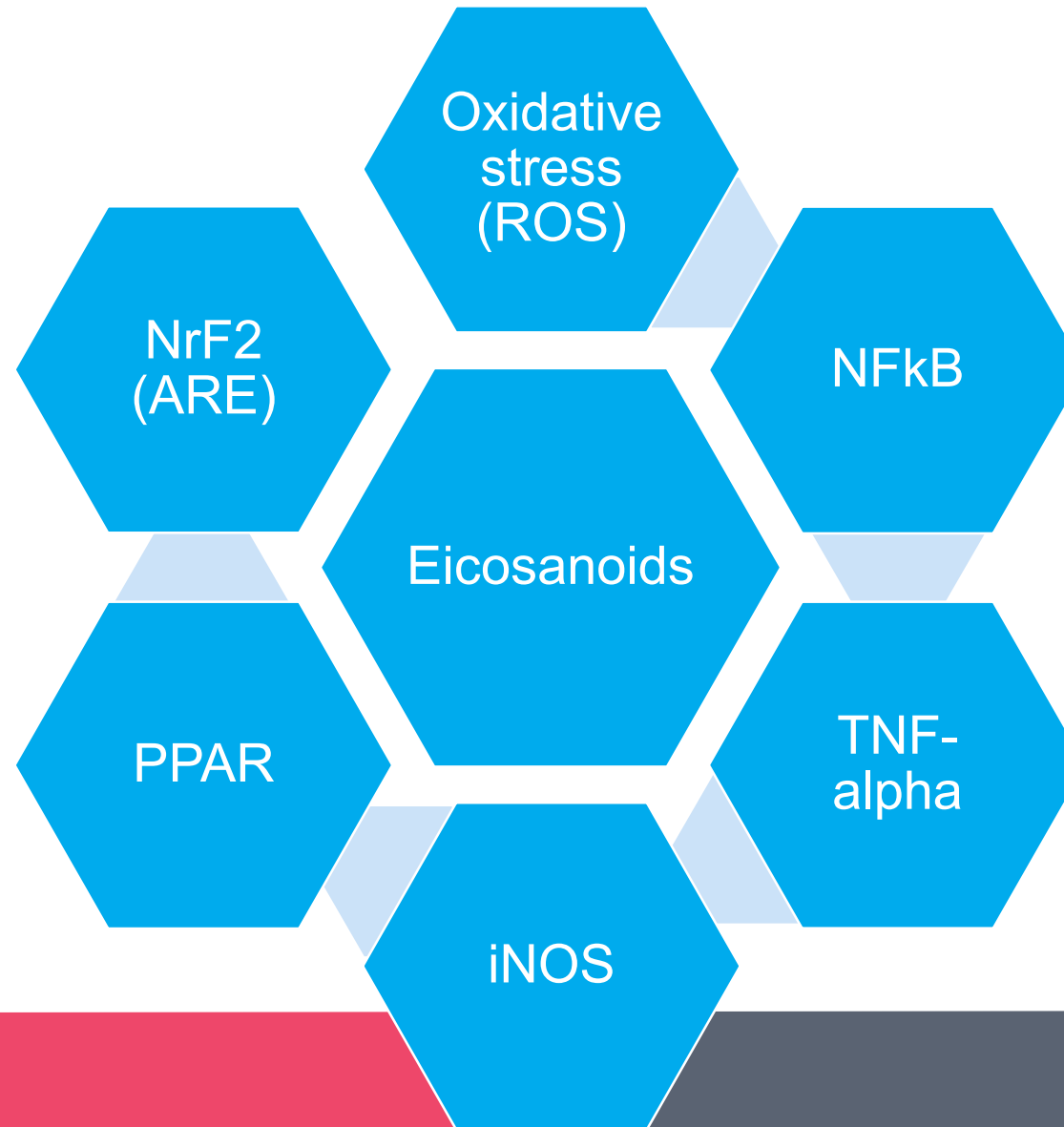
Role of Pharmaceuticals



Mediating Inflammatory Response



Mediating Inflammatory Response



NF-κB Inhibitors

- Resveratrol
- Curcumin
- EGCG
- Lycopene
- Allicin (garlic)
- Quercetin
- Vitamin C
- Vitamin D3
- Vitamin E
- Vitamin B6
- Folate
- B12
- B5
- Mg
- ALA
- CoQ10
- EPA/DHA
- Carnosine
- Dark chocolate
- Blueberries
- Ginger

Nrf2 inducers

(polyphenols, flavonoids, organosulfur)

- Sulforaphane
- Curcumin
- Ginger
- Trans-resveratrol
- Quercetin
- ALA
- Carnosol (rosemary)
- Garlic
- Coffee & EGCG
- Cinnamon
- Ginkgo biloba

PPAR activation

- PUFAs including
 - Omega 3
 - ALA
 - DHA
 - LA
- Along with cofactors:
 - Zn
 - Mg
 - Vitamin C
 - B3
 - B6
 - Saturated FA (some)

Nature's Immune Modulators

Arachidonic Cascade Modulators

Phospholipase A2

- Licorice
- Quercitin
- Vitamin E

COX1 + COX2 Inhibitors

- Bromelain
- Ginger*
- EPA/DHA*
- Quercitin*
- Garlic
- Turmeric
- Willow
- Barberry
- Licorice
- Wild yam
- Chamomile
- EGCG
- Rosmarinic acid
- Skullcap
- Thyme
- Clove

**both COX1+ COX2 activity*

5-LOX Inhibitors

- Quercitin
- Boswellia
- Turmeric
- Licorice
- Ginger
- EPA
- EGCG
- Skullcap
- Thyme

Vitamin D and autoimmunity

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Objectives: To review and evaluate the role of vitamin D in autoimmune diseases.

Method: We searched PubMed using keywords such as 'vitamin D' and 'autoimmunity'. We compiled and reviewed various studies including prospective studies, genetic studies, and experimental models that investigated the role of vitamin D in autoimmune diseases.

Results: There is evidence based on these various studies that several key autoimmune diseases are modulated by vitamin D. These diseases include, but are not limited to, multiple sclerosis (MS), scleroderma or systemic sclerosis (SSc), autoimmune thyroid diseases, rheumatoid arthritis (RA), and primary biliary cirrhosis (PBC).

Conclusions: Although there is evidence for vitamin D modulation of autoimmune diseases, the mechanism for this association has yet to be elucidated. Additional data are needed to corroborate these findings.

RESULTS: There is evidence based on these various studies that several key autoimmune diseases are modulated by vitamin D. These diseases include, but are not limited to, multiple sclerosis (MS), scleroderma or systemic sclerosis (SSc), autoimmune thyroid diseases, rheumatoid arthritis (RA), and primary biliary cirrhosis (PBC).

Diet & Lifestyle Interventions

Upstream Plan: Nutrition Overview

Overarching diet

- Elimination diet
- Autoimmune Diet/Paleo (AIP vs Whals)
- Fasting

Focus on: Cruciferous veggies, garlic, onion, rosemary, cherries, dark chocolate, blueberries, turmeric, raspberries and grapes, fish, egg*, flax, nuts/seeds, grass-fed beef and wild game, organ meats, green tea.

- Antioxidants and cofactors)
- Direct immune modulators (Vit C and D)
- FIBER

**Eggs re often removed during elimination phase and challenged*

SYSTEMATIC REVIEW



The effects of the Mediterranean diet on rheumatoid arthritis prevention and treatment: a systematic review of human prospective studies

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Abstract

Rheumatoid arthritis is a progressive autoimmune disease characterized by chronic inflammation. In the absence of effective pharmacotherapy, people living with rheumatoid arthritis may benefit from a healthy diet. The aim of the present systematic review is to discuss the effects of the Mediterranean diet on the prevention of rheumatoid arthritis in human prospective studies. We searched for intervention studies reporting improvement in the pain visual analogue scale and the questionnaire for rheumatoid arthritis score ($p < 0.05$) in the Mediterranean diet group compared with the control group in the 28 joint count disease activity score for rheumatoid arthritis. The Mediterranean diet has identified beneficial effects of the Mediterranean diet in reducing disease activity in people with rheumatoid arthritis. However, there is currently insufficient evidence to support widespread recommendation of the Mediterranean diet for prevention of rheumatoid arthritis.

In conclusion, presently, there is insufficient evidence to support the use of the MD for the prevention of RA. However, the results from this review suggest that the MD may provide benefit for PLRA in conjunction with medical treatment. **While the intervention studies presented in this report suggest a beneficial effect in using a MD for the management of arthritic pain and function, these trials have many shortcomings, and their results should be interpreted with caution.** The MD is a well-balanced, nutritionally adequate diet that encompasses all the food groups.

Paleolithic and Mediterranean Diet Pattern Scores Are Inversely Associated with Biomarkers of Inflammation and Oxidative Balance in Adults^{1–3}

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Abstract

Background: Chronic inflammation and oxidative balance are associated with poor health and chronic diseases. A diet–inflammation/oxidative balance association may relate to diet patterns.

Objective: We investigated associations between 2 diet pattern scores, the Mediterranean Diet Pattern Score (MDPS), and circulating concentrations of 2 related biomarkers, high-sensitivity C-reactive protein, and F₂-isoprostane, a reliable marker of in vivo lipid peroxidation.

Methods: In a pooled cross-sectional study of 30- to 74-y-old men and women in the Atherosclerosis Risk in Communities Study (n = 646), we created diet scores from responses on Willett food frequency questionnaire. We measured plasma hsCRP and F₂-isoprostane concentrations by ELISA and gas chromatography-mass spectrometry, respectively. Both diet scores were calculated and categorized into quintiles, and their associations with biomarkers were estimated with the use of general linear models to calculate and compare adjusted means and 95% CIs for each quintile using ordinal logistic regression.

Results: There were statistically significant trends for decreasing geometric mean concentrations with increasing quintiles of the Paleolithic and Mediterranean diet scores. Comparing those in the highest with those in the lowest quintiles of the Paleolithic and Mediterranean diet scores were 0.61 (95% CI: 0.36, 1.05; P-trend = 0.06) and 0.71 (95% CI: 0.42, 1.20; P-trend = 0.01), respectively, for a higher hsCRP concentration, and 0.51 (95% CI: 0.27, 0.95; P-trend 0.01) and 0.39 (95% CI: 0.21, 0.73; P-trend = 0.01), respectively, for a higher F₂-isoprostane concentration.

Conclusion: These findings suggest that diets that are more Paleolithic- or Mediterranean-like may be associated with lower levels of systemic inflammation and oxidative stress in humans. *J Nutr* 2016;146:1217–26.

CONCLUSION:

These findings suggest that diets that are more Paleolithic- or Mediterranean-like may be associated with lower levels of systemic inflammation and oxidative stress in humans.

Randomized control trial evaluation of a modified Paleolithic dietary intervention in the treatment of relapsing-remitting multiple sclerosis: a pilot study

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 Degenerative Neurological and Neuromuscular Disease
 4 January 2017
[Number of times this article has been viewed](#)

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Background/objective: A Paleolithic diet may be useful in the treatment and management of multiple sclerosis (MS) patients, but past research has been limited. The purpose of this pilot study was to evaluate a modified Paleolithic diet (MPDI) in combination with other treatments. **Methods:** We measured the effects of a MPDI in a randomized controlled trial. Subjects (one man) completed the control group and the MPDI group. **Results:** Significant improvements were seen in the Multiple Sclerosis Quality of Life-54 and time to complete tasks in the MPDI group compared to the control group. **Conclusion:** A Paleolithic diet may be useful in the treatment and management of MS, by reducing perceived fatigue, increasing mental and physical quality of life, increasing exercise capacity, and improving hand and leg function. By increasing vitamin K serum levels, the MPDI may also reduce inflammation. **Keywords:** diet therapy, nutrition therapy, gluten-free, quality of life, fatigue, complementary medicine, alternative medicine

CONCLUSION:

A Paleolithic diet may be useful in the treatment and management of MS, by reducing perceived fatigue, increasing mental and physical quality of life, increasing exercise capacity, and improving hand and leg function. By increasing vitamin K serum levels, the MPDI may also reduce inflammation.



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Nutrition and fasting mimicking diets in the prevention and treatment of autoimmune diseases and immunosenescence

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Abstract

Complex and coordinated signals are necessary to initiate and sustain the development, proliferation, and differentiation of lymphocytes. These signals, which are known to regulate immune function, also depend on the metabolic state of the organism. Recent studies have shown that the type and levels of nutrients can influence the generation, survival and function of lymphocytes and therefore can affect several autoimmune diseases. Here, we review the dysregulation of lymphocytes during autoimmunity and aging, the mechanisms associated with loss of immune function, and how fasting mimicking diets and other dietary interventions affect autoimmunity and immunosenescence.

...periodic IF/FMD combines a period of severe restriction sufficiently long to promote the death of a significant portion of damaged cells with a period of high nourishment re-feeding able to promote the opposite effect on growth and other factors leading to multi-system regeneration



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Diet mimicking fasting promotes regeneration and reduces autoimmunity and multiple sclerosis symptoms

In Young Choi^{1,†}, Laura Piccio^{2,†}, Patra Childress³, Bryan Bollman², Arko Ghosh⁴, Sebastian Brandhorst¹, Jorge Suarez¹, Andreas Michalsen⁵, Anne H. Cross², Todd E. Morgan¹, Min Wei¹, Friedemann Paul^{6,7}, Markus Bock^{6,7,*}, and Valter D. Longo^{1,4,8,9,*}

Summary

Dietary interventions have not been effective in relapsing-remitting multiple sclerosis (RRMS). We show that periodic 3 day cycles of a fasting mimicking diet (FMD) promote axonal regeneration and remyelination and reduce symptoms in a murine experimental model. The FMD reduced clinical severity in a murine model of RRMS. These improvements were associated with increased axon number, increased cell number, reduced levels of pro-inflammatory cytokines, TH1 and TH17 cells, and antigen-presenting cells (APCs). Moreover, the FMD promoted axonal regeneration and remyelination in axons in a murine model of RRMS. These findings support its effects on both suppression of inflammation and axonal regeneration. Preliminary data suggesting that a FMD or a chronic ketogenic diet are safe, feasible and potentially effective in the treatment of relapsing remitting multiple sclerosis (RRMS) patients (NCT01538355).

...associated with increased corticosterone levels and regulatory T (Treg) cell numbers and reduced levels of pro-inflammatory cytokines, TH1 and TH17 cells, and antigen-presenting cells (APCs)... **We also report preliminary data suggesting that an FMD or a chronic ketogenic diet are safe, feasible, and potentially effective in the treatment of relapsing-remitting multiple sclerosis (RRMS) patients**

Dietary recommendations



Summary of dietary Recommendations

Emphasis on:

- Fiber-dense foods and lower carbohydrate diet
- Anti-inflammatory fats (some)
- Apoptogenic and antioxidant properties of micronutrients and phytonutrients

Paleo

- Might be due to reduced carbohydrates, avoidance of processed foods, and possibly avoidance of lectin and phytic acid in grains and legumes.

Keto and Intermittent Fasting

- Benefit may be similar to paleo in addition to that from added anti-inflammatory effect of ketosis state

AIP/Wahls etc..

- Combine some of the above benefits into a protocol

Removal of IgG/IgA and cross-reactive foods (always with goal of safe expansion)

- May contribute to delayed food sensitivities of mast cell reactivity

Elimination diet and careful challenge (gold standard)

- Basic elimination may include gluten, dairy (casein), egg, soy, corn, peanuts, shellfish

5R Protocol

Remove	Potential triggers, including polypharmacy, pathogenic organisms, food intolerances, sensitivities and allergies, or toxic exposure
Replace	Digestive aid to support improved nutrient absorption and metabolism, including digestive enzymes, or agents that promote improved motility and regular bowel movements
Reinoculate*	Provide an environment where good bacteria can thrive and where bad ones cannot
Repair	Support of the cellular repair process through the above, as well as by providing specific nutritional support for the regeneration of the GI protective barrier
Rebalance	lifestyle factors that influence the gut bacteria such as stress, sleep, exercise and relationships and assure ongoing gut health

**Arguably, reinoculated isn't as simple as "reseeding" but we use the term to simplify*

Role of supplements



Useful supplements in allergic disease

Work horses:

- Quercetin
- Vit C
- Vit D3/K2
- Vit A

Anti-inflammatory support (fish oil, selective proresolving mediators, ginger, resveratrol, curcumin, etc...)

Adaptogens (caution in some patients)

GI protocol (antimicrobial herbs, glutamine, etc...)

Probiotics and prebiotics (caution in some patients)

Targeted approach: Consider nutrition evaluation and stool testing to assess assimilation and nutrient utilization

Putting it all together...

FUNCTIONAL MEDICINE MATRIX

Retelling the Patient's Story

Antecedents

Genetics/Family History
C-section
Frequent childhood antibiotic use

Triggering Events

Tick bite
Family stress, traumatic event

Mediators/Perpetuators

Standard American diet (SAD+)
SIBO methane dom
Intestinal permeability
Stress

Physiology and Function: Organizing the Patient's Clinical Imbalances

Assimilation

GI distress/bloating
Constipation.
Positive SIBO
Indication so leaky gut
Structural Integrity

Intestinal permeability
Heat rash

Communication

Cortisol blunted due to
chronic steroid use
Fasting Insulin elevated (7)

Defense & Repair

Frequent colds
Chemical sensitivity (perfume)
Heat sensitivity

Energy

Fatigue
Poor sleep
Foggy brain

Biotransformation & Elimination

Constipation
chemical sensitivity



Transport

Edema (legs)

Modifiable Personal Lifestyle Factors

Sleep & Relaxation

Sleep irregular,
Not enough

Exercise & Movement

Exercise
induces heat
rash

Nutrition

Diet is SAD+

Stress

Amount of stress
high, has some
good stress
management

Relationships

Strong social network
Friends and family

FUNCTIONAL MEDICINE TIMELINE

Mediators/Perpetuators

Family stress
Standard American diet

Chronic infection (viral, bacterial, parasite)
Intestinal permeability/dysbiosis/SIBO

Antecedents

Genetics/Family History
SAD and high sugar diet in childhood

Triggers or Triggering Events

Preconception

Prenatal

Birth

C-section birth

Frequent childhood antibiotic use

Grad school - stress

Mother's illness
Mother's death
Father's death shortly after

Tick bite

SLE Diagnosis
Home renovation

Physical injury (car accident)

Lupus diagnosis -initiated hydrocortisone and hydroxychloroquine

Current Concerns

Signs, Symptoms or Diseases Reported

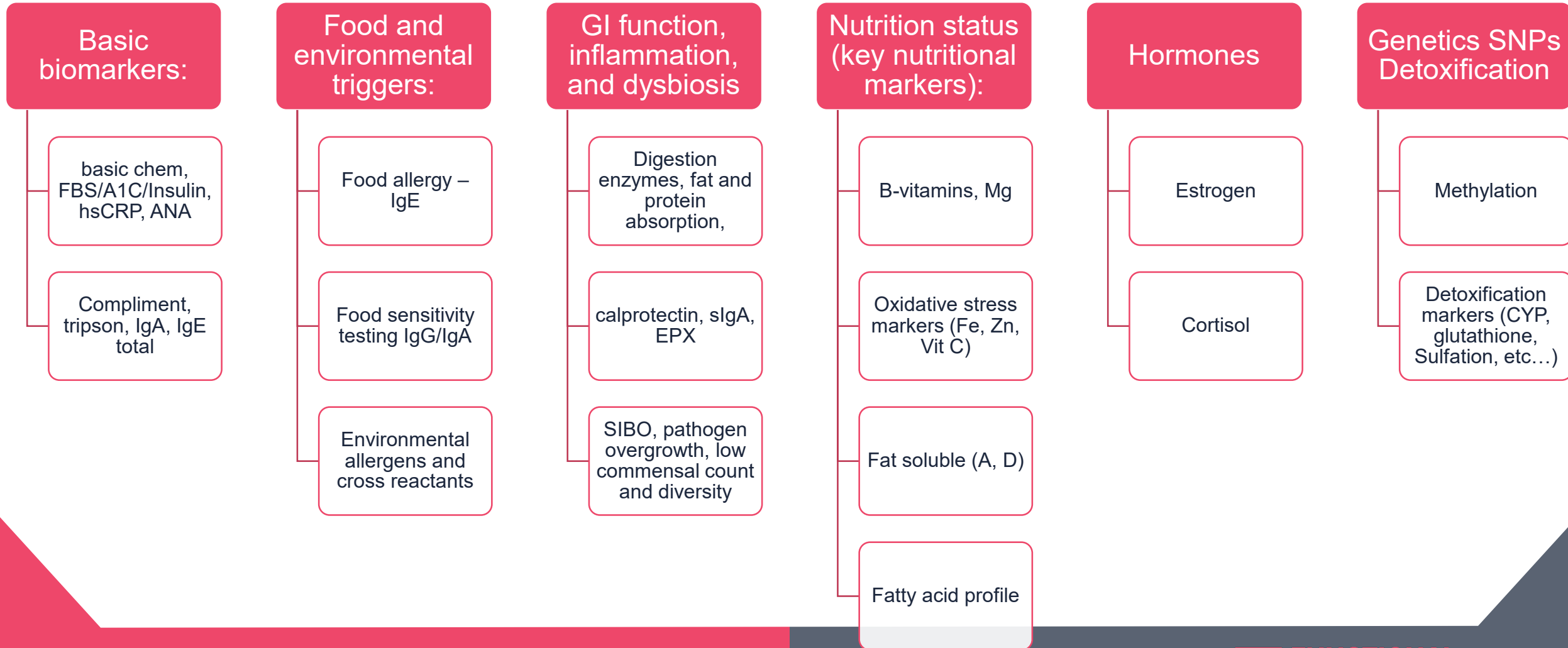
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Version 3



Autoimmune Assessment: Labs



Implications of dysbiosis

Opportunistic Bacteria			
Additional Dysbiotic/Overgrowth Bacteria	Result		Normal
<i>Bacillus</i> spp.	6.61e5	High	<1.50e5
<i>Enterococcus faecalis</i>	<dl		<1.00e4
<i>Enterococcus faecium</i>	1.02e4	High	<1.00e4
<i>Morganella</i> spp.	<dl		<1.00e3
<i>Pseudomonas</i> spp.	<dl		<1.00e4
<i>Pseudomonas aeruginosa</i>	<dl		<5.00e2
<i>Staphylococcus</i> spp.	<dl		<1.00e4
<i>Staphylococcus aureus</i>	7.85e2	High	<5.00e2
<i>Streptococcus</i> spp.	2.02e4	High	<1.00e3
<i>Methanobacteriaceae</i> (family)	1.14e8		<5.00e9
Potential Autoimmune Triggers	Result		Normal
<i>Citrobacter</i> spp.	<dl		<5.00e6
<i>Citrobacter freundii</i>	4.69e5		<5.00e5
<i>Klebsiella</i> spp.	1.02e6	High	<5.00e3
<i>Klebsiella pneumoniae</i>	1.66e6	High	<5.00e4
<i>M. avium</i> subsp. <i>paratuberculosis</i>	<dl		<5.00e3
<i>Prevotella</i> spp.	4.57e7		<1.00e8
<i>Proteus</i> spp.	<dl		<5.00e4
<i>Proteus mirabilis</i>	<dl		<1.00e3
<i>Fusobacterium</i> spp.	6.43e5		<1.00e8



Implications of dysbiosis

Intestinal Health		
Digestion	Result	Normal
Steatocrit	<dl	<15 %
Elastase-1	502	>200 ug/g
GI Markers	Result	Normal
b-Glucuronidase	4158	<2486 U/mL
Occult Blood - FIT	0	<10 ug/g
Immune Response	Result	Normal
Secretory IgA	2889	510 - 2010 ug/g
Anti-gliadin IgA	101	0 - 157 U/L
Inflammation	Result	Normal
Calprotectin	0	<173 ug/g
Add-on Test	Result	Normal
Zonulin	208.2	<107 ng/g

Implications of dysbiosis

Methodology: Culture/MALDI-TOF MS, Automated and Manual Biochemical Methods, Vitek® 2 System Microbial identification and Antibiotic susceptibility

Gastrointestinal Microbiome

Bacteriology (Culture)

Lactobacillus spp.

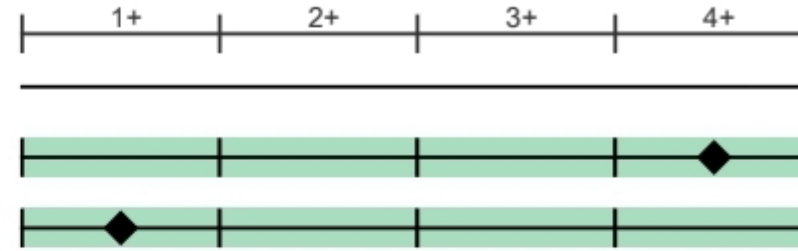
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Escherichia coli

4+ NP

Bifidobacterium

1+ NP



Additional Bacteria

alpha haemolytic Streptococcus

4+ NP

Klebsiella oxytoca

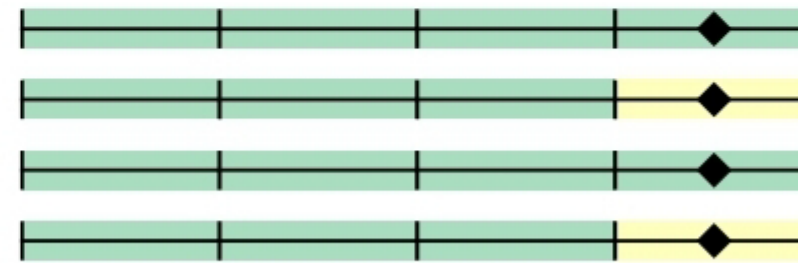
4+ PP

gamma haemolytic Streptococcus

4+ NP

Enterobacter cloacae

4+ PP



Mycology (Culture)

NG



Medical Symptoms Questionnaire (MSQ)

Patient Name _____

Rate each of the following symptoms based upon your typical history.

Point Scale 0 – *Never or almost never* have the symptom 3 – *I*
1 – *Occasionally* have it, effect is *not severe* 4 – *I*
2 – *Occasionally* have it, effect is *severe*

HEAD _____ Headaches
_____ Faintness
_____ Dizziness
_____ Insomnia

EYES _____ Watery or itchy eyes
_____ Swollen, reddened or sticky
_____ Bags or dark circles under eyes
_____ Blurred or tunnel vision
(Does not include near or far-sightedness)

EARS _____ Itchy ears
_____ Earaches, ear infections
_____ Drainage from ear
_____ Ringing in ears, hearing loss

NOSE _____ Stuffy nose
_____ Sinus problems
_____ Hay fever
_____ Sneezing attacks
_____ Excessive mucus formation

MOUTH/THROAT _____ Chronic coughing
_____ Gagging, frequent need to clear throat
_____ Sore throat, hoarseness, loss of voice
_____ Swollen or discolored tongue
_____ Canker sores

SKIN _____ Acne
_____ Hives, rashes, dry skin
_____ Hair loss
_____ Flushing, hot flashes
_____ Excessive sweating

HEART _____ Irregular or skipped heartbeats
_____ Rapid or pounding heartbeats
_____ Chest pain

Version 2

MEDICAL SYMPTOMS QUESTIONNAIRE (MSQ)

LUNGS _____ Chest congestion
_____ Asthma, bronchitis
_____ Shortness of breath
_____ Difficulty breathing
Total _____

DIGESTIVE TRACT _____ Nausea, vomiting
_____ Diarrhea
_____ Constipation
_____ Bloating, fullness
_____ Belching, passing gas
_____ Heartburn
_____ Intestinal/stomach pain
Total _____

JOINTS/MUSCLE _____ Pain or aches in joints
_____ Arthritis
_____ Stiffness or limitation of movement
_____ Pain or aches in muscles
_____ Feeling of weakness or tiredness
Total _____

WEIGHT _____ Binge eating/drinking
_____ Craving certain foods
_____ Excessive weight
_____ Compulsive eating
_____ Water retention
_____ Underweight
Total _____

ENERGY/ACTIVITY _____ Fatigue, sluggishness
_____ Apathy, lethargy
_____ Hyperactivity
_____ Restlessness
Total _____

MIND _____ Poor memory
_____ Confusion, poor comprehension
_____ Poor concentration
_____ Poor physical coordination
_____ Difficulty in making decisions
_____ Stuttering or stammering
_____ Slurred speech
_____ Learning disabilities
Total _____

EMOTIONS _____ Mood swings
_____ Anxiety, fear, nervousness
_____ Anger, irritability, aggressiveness
_____ Depression
Total _____

OTHER _____ Frequent illness
_____ Frequent or urgent urination
_____ Genital itch or discharge
Total _____

Grand Total _____

MSQ

- ✓ Quantifies subjective physical
- ✓ Highlights areas of priority



Diet, Nutrition, and Lifestyle Journal – 1 Day

Patient Name: _____ Date: _____

Food Plan Type: _____

Day 1

Day Event	Food & Drink Intake (include type, amount, brand)	Macronutrients (PFC) and Phytonutrients
Rising Time		
Breakfast Time		P_____F_____C ☐ R ☐ O ☐ Y ☐ G ☐ B/P/BL ☐ W/T/BR
Mid-AM Snack Time		P_____F_____C ☐ R ☐ O ☐ Y ☐ G ☐ B/P/BL ☐ W/T/BR
Lunch Time		P_____F_____C ☐ R ☐ O ☐ Y ☐ G ☐ B/P/BL ☐ W/T/BR
Mid-PM Snack Time		P_____F_____C ☐ R ☐ O ☐ Y ☐ G ☐ B/P/BL ☐ W/T/BR
Dinner Time		P_____F_____C ☐ R ☐ O ☐ Y ☐ G ☐ B/P/BL ☐ W/T/BR
PM Snack Time		P_____F_____C ☐ R ☐ O ☐ Y ☐ G ☐ B/P/BL ☐ W/T/BR
Bed Time		

P: Proteins; F: Fats; C: Carbohydrates; R: Red; O: Orange; Y: Yellow; G: Green; B/P/BL: Blue/Purple/Black; W/T/BR: White/Tan/Brown

Sleep & Relaxation	Exercise & Movement	Stress	Relationships
Sleep Quantity: _____ (hours) Quality: ☐ Poor ☐ Fair ☐ Good Relaxation ☐ Yes ☐ No Type/Amount:	Type, Duration, & Intensity ☐ Aerobic: ☐ Strength: ☐ Flexibility:	Stress Reduction Practices: Stressors:	Supporting: Non-supporting:

Mental	Emotional	Spiritual



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Version 2

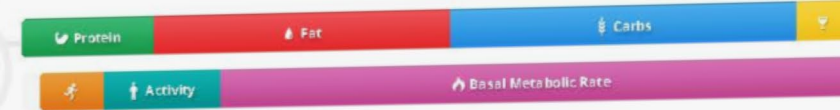
Diet Diary

- ✓ Meal timing
- ✓ Color and variety
- ✓ Track macronutrients (later)
- ✓ Foundational aspects:
 - Sleep & Relaxation
 - Exercise
 - Stress
 - Relationships
 - Mental, emotional and Spiritual factors/practices

Description	Amount	Unit
 Oatmeal, Regular or Quick, Dry	1	FDA Serving Size
 Raisins, Uncooked	1	miniature box each 1/2
 Flax Seeds, Fortified	1	tbsp. whole pieces
 Coconut Milk, Plain or Original, Fortified, Unsweetened	4	tbsp
 Tap Water	1	cup
 Salt	1	dash
 Pecans, Raw	1	oz
 Green Tea, Brewed, Unsweetened	1	cup
 vacuuming, general, moderate effort	10	minutes
 Mild stretching	15	minutes
 Cantaloupe, Fresh	1	wedge 1/8 of medium
 Seaweed, agar, dried	5	g
 Arugula, Raw	3	cup, cut pieces
 Spinach, Raw	1	cup, cut pieces
 Apple, Fresh, With Skin	1	very small 2 1/2" diamet
 Carrots, Raw	1	medium 6" to 7" long
 Black Olives	6	large
 Avocado, Black Skin, California Type	1	Serving (3 slice)
 Olive Oil	1	tbsp
 Lemon Juice, Fresh	1	wedge juice from one w
 Tap Water	2	cup
 Dates	1	each
 Rice Cake	2	regular 4" diameter
 Cashew Butter, Salted	3	tbsp
 swimming laps, freestyle, front crawl, slow, light or moderate effort	30	minutes
 Soup, Stock, Chicken, Home Prepared	1	cup
 Mushrooms, Cooked from Fresh	1	FDA Serving Size
 Chicken Breast, Skin Removed Before Cooking	1	medium split
 Brown Rice, Cooked in Salted Water	1	FDA Serving Size
 Spinach, Cooked from Fresh	1	FDA Serving Size
 Tap Water	2	cup
 White Wine, Other Types	1	× 1/2 cup
 Tap Water	1	cup
 Rice Cake	2	regular 4" diameter

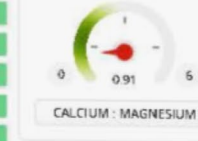
Daily Average Nutrition Report 2018 February 17 to 2018 February 24

Calories Summary



Carbohydrates			Vitamins		
Carbs	162.8 g	125%	B1 (Thiamine)	1.2 mg	111%
Fiber	25.2 g	120%	B2 (Riboflavin)	1.2 mg	113%
Net Carbs	137.6 g	106%	B3 (Niacin)	23.7 mg	170%
Starch	61.4 g	No Target	B5 (Pantothenic Acid)	5.7 mg	114%
Sugars	56.0 g	No Target	B6 (Pyridoxine)	2.4 mg	159%
Lipids			B12 (Cobalamin)	4.9 µg	205%
Fat	65.5 g	101%	Folate	369.0 µg	92%
Monounsaturated	28.8 g	No Target	Vitamin A	11189.8 IU	480%
Polyunsaturated	16.7 g	No Target	Vitamin C	130.3 mg	174%
Omega 3	2.8 g	255%	Vitamin D	77.9 IU	13%
Omega 6	13.8 g	125%	Vitamin E	7.5 mg	59%
Saturated	14.0 g	n/a	Vitamin K	211.4 µg	235%
Trans Fats	0.2 g	0.242 g	Minerals		
Cholesterol	83.9 mg	No Target	Calcium	417.9 mg	35%
Protein			Copper	2.3 mg	254%
Protein	66.4 g	144%	Iron	13.7 mg	172%
Cystine	0.9 g	165%	Magnesium	459.3 mg	144%
Histidine	1.9 g	234%	Manganese	6.6 mg	368%
Isoleucine	2.8 g	263%	Phosphorus	1183.1 mg	169%
Leucine	4.8 g	202%	Potassium	3332.1 mg	71%
Lysine	4.5 g	210%	Selenium	76.6 µg	139%
Methionine	1.5 g	269%	Sodium	1463.5 mg	113%
Phenylalanine	2.7 g	288%	Zinc	12.2 mg	152%
Threonine	2.6 g	228%	General		
Tryptophan	0.7 g	264%	Energy	1539.7 kcal	133%
Tyrosine	2.1 g	223%	Alcohol	13.6 g	No Target
Valine	3.3 g	239%	Caffeine	7.7 mg	No Target
			Water	2393.1 g	88%

Nutrient Balances



Reminder: *Eat the Rainbow*

- Color and variety
- 8 or more servings of veggies a day
- High fiber, low glycemic fruits & veggies
- LOTS of water
- Minimize caffeine and alcohol
- Avoid cigarettes
- Exercise
- Meditate



Phytonutrient Spectrum Foods

RED

Foods				Benefits	
Apples	Cranberries	Pomegranate	Shrubub	Anti-cancer	Gastrointestinal health
Beets (adults, kidney ref)	Cherries	Potatoes	Roobor tea	Anti-inflammatory	Heart health
Beets	Grapefruit (pink)	Radicchio	Tomato	Cell protection	Hormone health
Bell peppers	Goji berries	Radicchio	Watermelon		Liver health
Blood oranges	Grapes	Raspberries			
	Onions	Strawberries			
	Mint	Sweet and peppers			

ORANGE

Foods				Benefits	
Apricots	Mango	Pumpkin	Tangerines	Anti-cancer	Reduced mortality
Bell peppers	Nectarine	Squash (Acorn, butternut, kabocha, winter)	Turmeric root	Anti-bacterial	Reproductive health
Cantaloupe	Orange	Sweet potato	Yams	Immune health	Skin health
Carrots	Papaya			Cell protection	Source of vitamin A
	Pineapples				

YELLOW

Foods				Benefits	
Apples	Bell peppers	Lemons	Starfruit	Anti-cancer	Eye health
Asian pears	Corn	Millet	Succotash	Heart health	Heart health
Bananas	Corn-on-the-cob	Pineapple	Summer squash	Anti-inflammatory	Skin health
	Ginger root			Cell protection	Vascular health
				Cognition	

GREEN

Foods				Benefits	
Apples	Bok choy	Green peas	Okra	Anti-inflammatory	Skin health
Artichoke	Broccoli	Green tea	Olives	Brain health	Hormone balance
Asparagus	Broccoli	Greens (arugula, basil, chard, dandelion, kale, lettuce, mustard, spinach, turnip)	Peanut	Cell protection	Heart health
Avocado	Brussels sprouts		Snow peas		Liver health
Bamboo shoots	Cabbage		Watermelon		
Bean sprouts	Celery		Zucchini		
Bell peppers	Cucumbers				
Bitter melon	Edamame/Soy beans	Limes			
	Green beans				

BLUE/PURPLE/BLACK

Foods				Benefits	
Bell peppers	Cabbage	Grapes	Prunes	Anti-cancer	Cognitive health
Berries (blue, black, boysenberries, blackberries, mulberries)	Carrots	Kale	Raspberries	Anti-inflammatory	Heart health
	Cauliflower	Olives	Pineapple	Cell protection	Liver health
	Eggplant	Peanut			
	Figs	Potatoes			

WHITE/TAN/BROWN

Foods				Benefits	
Apples	Dates	Mushrooms	Shallots	Anti-cancer	Heart health
Applesauce	Garlic	Nuts (almonds, cashews, pecans, walnuts)	Soy	Anti-microbial	Hormone health
Bean dips	Ginger	Onions	Tahini	Cell protection	Liver health
Cauliflower	Jicama	Peanut	Tea (black, white)	Gastrointestinal health	
Cocoa	Legumes (chickpeas, dried beans or peas, lentils, mung beans, pinto beans, refried beans / taco mix)	Sweetener	Whole grains (barley, brown rice, oat, quinoa, rye, spelt, wheat)		
Coconut		Seeds (flax, hemp, pumpkin, sesame, sunflower)			
Coffee					



Elimination Diet Food Plan

PROTEINS

Proteins

Servings/day _____

Lean, free-range, grass-fed, organically grown animal protein; non-GMO, organic plant protein; and wild-caught, low-mercury fish preferred.

Animal Proteins:

- ☐ Fish: Halibut, herring, mackerel, salmon, sardines, tuna, etc.—1 oz
- ☐ Meat: All wild game, buffalo, elk, lamb, venison—1 oz
- ☐ Poultry (skinless): Chicken, Cornish hen, turkey—1 oz

1 serving as listed = 35–75 calories, 5–7 g protein, 3–5 g fat, 0–4 g carbs

Average protein serving is 3–4 oz (size of palm of hand).

Eliminate

Beef/veal, canned meats, cold cuts, eggs, frankfurters, pork, shellfish, whey, soy (miso, natto, tempeh, tofu, textured vegetable protein)

Plant Protein:

- ☐ Spirulina—2 T
- ☐ Protein Powder: Check label for # grams/scoop (1 protein serving=7 g) Hemp, pea, rice

LEGUMES

Proteins/Carbs

Servings/day _____

Organic, non-GMO preferred

- ☐ Bean soups—¾ c
- ☐ Dried beans, peas, or lentils (cooked)—½ c
- ☐ Flour, legume—¼ c
- ☐ Green peas (cooked)—½ c
- ☐ Hummus or other bean dip—½ c
- ☐ Refried beans, vegetarian—½ c

1 serving = 90–110 calories, 3–7 g protein, 0 fat, 15 g carbs

Eliminate

Soybean products (edamame, miso, soy sauce, tamari, tempeh, tofu, soy milk, soy yogurt, textured vegetable protein)

DAIRY ALTERNATIVES

Proteins/Carbs

Servings/day _____

Unsweetened, organic preferred

- ☐ Kefir: Coconut (plain) 4–6 oz
- ☐ Milk: Almond, coconut, flaxseed, hazelnut, hemp, rice—8 oz
- ☐ Yogurt: Coconut (cultured) 4–6 oz

1 serving = 25–90 calories, 1–9 g protein, 1–4 g carbs (nutritional values vary)

Eliminate

Butter, cheese, cottage cheese, cream, frozen yogurt, ice cream, milk, non-dairy creamers, soy milk, yogurt (dairy and soy), whey

NUTS & SEEDS

Proteins/Fats

Servings/day _____

Unsweetened, unsalted, organic preferred

- ☐ Almonds—6
- ☐ Brazil nuts—2
- ☐ Cashews—6
- ☐ Chia seeds—1 T
- ☐ Coconut (dried)—3 T
- ☐ Flaxseed (ground)—2 T
- ☐ Hazelnuts—5
- ☐ Hemp seeds—1 T
- ☐ Macadamias—2–3
- ☐ Nut and seed butters—½ T
- ☐ Pecan Halves—4
- ☐ Pine nuts—1 T
- ☐ Pistachios—16
- ☐ Pumpkin seeds—1 T
- ☐ Sesame seeds—1 T
- ☐ Sunflower seeds—1 T
- ☐ Walnut halves—4

1 serving = 45 calories, 5 g fat

Eliminate

Mixed nuts (with peanuts), peanuts, peanut butter

FATS & OILS

Fats

Servings/day _____

Minimally refined, cold-pressed, organic, non-GMO preferred

- ☐ Avocado—2 T or ⅛ whole
- ☐ Coconut milk, regular (canned)—1½ T
- ☐ Coconut milk, light (canned)—3 T
- ☐ Ghee/clarified butter (grass-fed)—1 t
- ☐ Olives: Black, green, kalamata—8
- ☐ Oils, cooking: Avocado, coconut, grapeseed, olive (extra virgin), rice bran, sesame—1 t
- ☐ Oils, salad: Almond, avocado, flaxseed, grapeseed, hempseed, olive (extra virgin), pumpkin, safflower (high-oleic), sunflower (high-oleic), sesame, walnut—1 t
- ☐ Prepared salad dressing with acceptable oils—2 T

1 serving = 45 calories, 5g fat

Eliminate

Butter, corn oil, cottonseed oil, margarine/spreads, mayonnaise, peanut oil, shortening, soybean oil

KEY

● High Histamine ■ Nightshades ▲ Fermented Foods

Notes: Nutritional amounts are based on average values for the variety of foods within each food category.

Dietary prescription is subject to the discretion of the health practitioner.

Eliminates:

- Gluten
- Dairy
- Eggs
- Soy
- Peanuts
- Shellfish
- Pork

Considerations:

- Paleo, Wahl's
- Low carbohydrate/higher fat
- Intermittent Fasting or FMD
- Nightshades
- Histamine
- Fermented foods*
- Food sensitivity testing

**Stomach pain or bloating after eating certain high-fiber food might be a symptom of bacterial overgrowth*



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VEGETABLES Non-starchy Carbs

Servings/day _____

- | | |
|---|--|
| ☐ Artichoke | ☐ Horseradish |
| ☐ Arugula | ☐ Jicama |
| ☐ Asparagus | ☐ Kohlrabi |
| ☐ Bamboo shoots | ☐ Leeks |
| ☐ Beets (cubed) | ☐ Lettuce, all |
| ☐ Bok choy | ☐ Microgreens |
| ☐ Broccoflower | ☐ Mushrooms● |
| ☐ Broccoli | ☐ Okra |
| ☐ Brussels sprouts | ☐ Onions |
| ☐ Cabbage | ☐ Parsley |
| ☐ Carrots | ☐ Peppers, all■ |
| ☐ Cauliflower | ☐ Radicchio |
| ☐ Celeriac root | ☐ Radishes |
| ☐ Celery | ☐ Salsa●■ |
| ☐ Chard/Swiss chard | ☐ Sea vegetables |
| ☐ Chervil | ☐ Scallions |
| ☐ Chives | ☐ Shallots |
| ☐ Cilantro | ☐ Snap peas/snow peas |
| ☐ Cucumbers | ☐ Spinach● |
| ☐ Daikon radishes | ☐ Sprouts, all |
| ☐ Eggplant●■ | ☐ Squash: Delicata, |
| ☐ Endive | pumpkin●, spaghetti, |
| ☐ Escarole | yellow, zucchini, etc. |
| ☐ Fennel | ☐ Tomato●■ |
| ☐ Fermented | ☐ Tomato juice●■—¾ |
| vegetables:●▲ | c |
| Kimchi, pickles, | ☐ Turnips |
| sauerkraut, etc. | ☐ Vegetable juice—¼ c |
| ☐ Garlic | ☐ Water chestnuts |
| ☐ Green beans | ☐ Watercress |
| ☐ Greens: Beet, collard, | |
| dandelion, kale, | |
| mustard, turnip, etc. | |

1 serving = ½ c, 1 c raw greens = 25 calories, 5 g carbs

Organic, non-GMO fruits, vegetables, herbs and spices preferred

VEGETABLES Starchy Carbs

Servings/day _____

- | | |
|---|--|
| ☐ Acorn squash | ☐ Potatoes (mashed, |
| (cubed)—1 c | made with non- |
| ☐ Butternut squash | dairy milk)—½ c |
| (cubed)—1 c | ☐ Root vegetables: |
| ☐ Plantain—⅓ c or | Parsnip, rutabaga—½ c |
| ½ whole | ☐ Yam—½ med |
| ☐ Potato: Purple, | |
| red, sweet, white, | |
| yellow—½ med | |

1 serving = 80 calories, 15 g carbs

Eliminate

Corn, Potato (if avoiding nightshades)

FRUITS Carbs

Servings/day _____

Unsweetened, no sugar added

- | | |
|---|--|
| ☐ Apple—1 sm | ☐ Melon, all—1 c |
| ☐ Applesauce—½ c | ☐ Mango—½ sm |
| ☐ Apricots—4 | ☐ Nectarine—1 sm |
| ☐ Banana—½ med | ☐ Orange—1 med |
| ☐ Blackberries—¾ c | ☐ Papaya—1 c |
| ☐ Blueberries—¾ c | ☐ Peach—1 sm |
| ☐ Dried fruit● | ☐ Pear—1 sm |
| (no sulfites)—2 T | ☐ Persimmon—½ |
| ☐ Figs—3 | ☐ Pineapple—¼ c |
| ☐ Grapes—15 | ☐ Plums—2 sm |
| ☐ Grapefruit—½ med | ☐ Pomegranate seeds |
| ☐ Juices (diluted)—½ c | —½ c |
| ☐ Kiwi—1 med | ☐ Prunes—3 med |
| ☐ Kumquats—4 | ☐ Raisins—2 T |
| ☐ Lemon—1 | ☐ Raspberries●—1 c |
| ☐ Lime—1 | ☐ Tangerines—2 sm |

1 serving = 60 calories, 15 g carbs

Eliminate

Citrus fruits (if directed by your healthcare provider)

GLUTEN-FREE GRAINS Carbs

Servings/day _____

Unsweetened, sprouted, organic preferred

- | | |
|--|--|
| ☐ Amaranth—¾ c | ☐ Millet—½ c |
| ☐ Brown rice cakes—2 | ☐ Oats: Rolled, |
| ☐ Buckwheat/ | steel-cut—½ c |
| kasha—½ c | ☐ Quinoa—½ c |
| ☐ Crackers: (nut, seed, | ☐ Rice—⅓ c |
| rice)—3-4 | ☐ Teff—¼ c |
| ☐ Flours for baking: | All grain servings are |
| Arrowroot, sorghum, | for cooked amounts. |
| tapioca—3 T | |

1 serving = 75-110 calories, 15 g carbs

Eliminate

Barley, corn, emmer, farro, kamut, rye, spelt, triticale, wheat

BEVERAGES, SPICES & CONDIMENTS

Unsweetened, no sugar added

- | | |
|---|--|
| ☐ Filtered water | ☐ Herbs and Spices, all |
| ☐ Sparkling/mineral | ☐ Condiments: |
| water | Mustard●, vinegars●▲ |
| ☐ Unsweetened | —use sparingly, suggest |
| coconut water | 1 T or less per serving |
| ☐ Green tea | |
| ☐ Fresh juiced fruits/ | |
| vegetables | |

KEY

● High Histamine ■ Nightshades ▲ Fermented Foods

Gut restoration:

First step: Diet

Herbals & Supplements:

- Turmeric, ginger, resveratrol
- Vitamins (D,A, C, B's)
- Minerals (Mg, Zn,Se, etc)
- Quercitin
- Boswellia
- Digestive enzymes & Betaine HCL
 - Swedish bitters, ACV
- L-glutamine
 - Bone broth & collagen
- Marshmallow, slippery elm, aloe, DGL
- Butyric acid & other prebiotic fibers (Arabigalactan, pectin, etc)
- Probiotics



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ReNew Food Plan

PROTEINS

Proteins

Servings/day _____

Lean, free-range, grass-fed, organically grown animal protein; non-GMO, organic plant protein; and wild-caught, low-mercury fish preferred.

Animal Protein:

- ☐ Egg ●—1
- ☐ Fish: Anchovies, cod, flounder/sole, herring, halibut, salmon, sardines, trout, etc.—1 oz
- ☐ Meat: Beef, buffalo, elk, lamb, venison, ostrich, etc.—1 oz
- ☐ Poultry (skinless): Chicken, Cornish hen, duck, pheasant, turkey, etc.—1 oz
- ☐ Mung bean/Edamame pasta ●—½ oz
- ☐ Natto ●▲—1 oz
- ☐ Spirulina—2 T
- ☐ Tofu (firm/extra firm) ●—1½-2 oz
- ☐ Tofu (soft/silken) ●—3 oz
- ☐ Tempeh ●▲—1 oz

Protein Powder:

- ☐ Check label for # grams/scoop (1 protein serving = 7g protein)
- ☐ Bovine collagen, egg, hemp, pea

Plant Protein:

- ☐ Black soybeans ●—¼ c
- ☐ Edamame ●—¼ c
- ☐ Hemp tofu—1½ oz

1 serving as listed = 35-75 calories, 5-7 g protein, 3-5 g fat, 0-4 g carbs

Average protein serving is 3-4 oz (size of palm of hand).

DAIRY ALTERNATIVES *Proteins/Carbs*

Servings/day _____

Unsweetened, organic preferred

- ☐ Yogurt: Coconut (plain) ●▲—4-6 oz
- ☐ Kefir: Coconut (plain) ●▲—4-6 oz
- ☐ Nut/seed milk: Almond, cashew, coconut, flaxseed, hazelnut, hemp—8 oz

1 serving = 25-90 calories, 1-9 g protein, 1-4 g carbs

NO DAIRY ALLOWED

NUTS & SEEDS

Proteins/Fats

Servings/day _____

Unsweetened, unsalted, organic preferred

- ☐ Almonds—6
- ☐ Brazil nuts—2
- ☐ Cashews ●—6
- ☐ Chia seeds—1 T
- ☐ Coconut (dried)—3 T
- ☐ Coconut wraps (raw, vegan)—1 wrap
- ☐ Flaxseed (ground)—2 T
- ☐ Hazelnuts—5
- ☐ Hemp seeds—1
- ☐ Macadamias—2-3
- ☐ Nut and seed butters: Almond, cashew, macadamia, pecan, sunflower, tahini, walnut ●—½ T
- ☐ Pecan halves—4
- ☐ Pine nuts—1 T
- ☐ Pistachios—16
- ☐ Pumpkin seeds—1 T
- ☐ Sesame seeds—1 T
- ☐ Sunflower seeds ●—1 T
- ☐ Walnut halves ●—4

1 serving = 45 calories, 5 g fat

FATS & OILS

Fats

Servings/day _____

Minimally refined, cold pressed, organic, non-GMO preferred

- ☐ Avocado ●—2 T or ½ whole
- ☐ Coconut butter (raw)—1 t
- ☐ Coconut milk, regular (BPA-free canned or boxed)—1½ T
- ☐ Ghee/clarified butter (grass-fed)—1 t
- ☐ Oils, cooking: Avocado, coconut, ghee/clarified butter, olive (extra virgin), sesame—1 t
- ☐ Oils, salad: Almond, avocado, flaxseed, hempseed, olive oil (extra virgin), sesame, walnut—1 t
- ☐ Olives: ● Black, green, kalamata—8
- ☐ Pesto (olive oil)—1 t

1 serving = 45 calories, 5 g fat

KEY

● High Histamine ■ Nightshades ▲ Fermented Foods

Notes: Nutritional amounts are based on average values for the variety of foods within each food category.

Dietary prescription is subject to the discretion of the health practitioner.

Paleo (lectins)

- With vegetarian option (soy*)

Eliminates:

- Gluten
- Dairy
- Peanuts
- Shellfish
- Pork

**Soy can be problematic in certain individuals, so this might take a deeper conversation about their choice to be vegetarian – always approached with respect for their autonomy balanced with informed decision making.*

NO LEGUMES (Except those specifically listed) and NO GRAINS (Bread, pasta, cereal, oats, etc.)

VEGETABLES Non-starchy Carbs

Servings/day_____

- Brassicales**
(i.e. Cruciferous)

 - ☐ Arugula
 - ☐ Broccoflower
 - ☐ Broccoli
 - ☐ Broccoli sprouts
 - ☐ Brussels sprouts
 - ☐ Cabbage
 - ☐ Cauliflower
 - ☐ Horseradish
 - ☐ Kohlrabi
 - ☐ Radishes

Detoxifying Leafy Greens

 - ☐ Bok choy
 - ☐ Chard/Swiss chard
 - ☐ Chervil
 - ☐ Cilantro
 - ☐ Endive
 - ☐ Escarole
 - ☐ Greens: Beet, collard, dandelion, kale, mustard, turnip, etc.
 - ☐ Microgreens
 - ☐ Parsley
 - ☐ Radicchio

Thiols

 - ☐ Chives
 - ☐ Daikon radishes
 - ☐ Garlic
 - ☐ Leeks
 - ☐ Onion
 - ☐ Scallions
 - ☐ Shallots
- Liver & Kidney Support**

 - ☐ Artichokes
 - ☐ Asparagus
 - ☐ Celery
 - ☐ Sprouts, all

Other Non-Starchy Vegetables

 - ☐ Bamboo shoots
 - ☐ Bean sprouts
 - ☐ Beets (not canned)
 - ☐ Carrots
 - ☐ Cucumbers
 - ☐ Eggplant
 - ☐ Fennel
 - ☐ Green beans
 - ☐ Jicama
 - ☐ Kimchi
 - ☐ Lettuce, all
 - ☐ Mushrooms
 - ☐ Okra
 - ☐ Peppers, all
 - ☐ Salsa
 - ☐ Sauerkraut
 - ☐ Sea vegetables
 - ☐ Shirataki noodles
 - ☐ Snap peas/snow peas
 - ☐ Spinach
 - ☐ Squash: Delicata, pumpkin, spaghetti, yellow, zucchini, etc.
 - ☐ Tomato
 - ☐ Turnip
 - ☐ Watercress

1 serving = ½ c, 1 c raw greens = 25 calories, 5 g carbs

NO STARCHY VEGETABLES (root vegetables)

Organic, non-GMO fruits, vegetables, herbs and spices preferred

FRUITS Carbs

Servings/day_____

- Unsweetened, no sugar added
- ☐ Blackberries—¾ c
 - ☐ Blueberries—¾ c
 - ☐ Cherries—12
 - ☐ Cranberries—¾ c
 - ☐ Kiwi—1 med
 - ☐ Pomegranate seeds—½ c
 - ☐ Raspberries—1 c
 - ☐ Strawberries—1¼ c
- 1 serving = 60 calories, 15 g carbs
- NO OTHER FRUITS ALLOWED

HERBS & SPICES

- ☐ Basil
- ☐ Bay leaf
- ☐ Black pepper
- ☐ Cayenne pepper
- ☐ Chili powder
- ☐ Cilantro
- ☐ Cinnamon
- ☐ Cloves
- ☐ Cacao powder (100% raw)
- ☐ Coriander seed
- ☐ Cumin
- ☐ Curry powder
- ☐ Dill
- ☐ Fenugreek
- ☐ Garlic powder
- ☐ Ginger
- ☐ Himalayan salt
- ☐ Nutmeg
- ☐ Onion powder
- ☐ Oregano
- ☐ Parsley
- ☐ Paprika
- ☐ Pumpkin spice
- ☐ Red curry paste
- ☐ Rosemary
- ☐ Sage
- ☐ Sea salt
- ☐ Thyme
- ☐ Turmeric
- ☐ Vanilla bean (whole)

BEVERAGES

- Unsweetened, no sugar added
- ☐ Broth (organic): Bone, meat, vegetable
 - ☐ Coconut water kefir
 - ☐ Filtered water
 - ☐ Seltzer water
 - ☐ Tea: Green, herbal
 - ☐ Vegetable juice (fresh, raw, cold pressed)
- NO COFFEE, ALCOHOL, CAFFEINE, SODA

CONDIMENTS

- ☐ Coconut aminos
 - ☐ Lemon/lime juice (fresh)
 - ☐ Miso
 - ☐ Mustard: Dijon, stone ground
 - ☐ Tamari
 - ☐ Vinegars: Apple cider, balsamic, white, etc.
- Use sparingly, suggest 1 T or less per serving.
- NO SUGARS, NATURAL SWEETENERS, OR ARTIFICIAL SWEETENERS, INCLUDING (BUT IS NOT LIMITED TO) ASPARTAME, SLENDA, STEVIA, AND SUGAR ALCOHOLS.

KEY
● High Histamine ■ Nightshades ▲ Fermented Foods

- Considerations:
- Low carbohydrate/higher fat
 - Ketogenic, remove egg and soy
 - Intermittent Fasting or FMD
 - Nightshades
 - Histamine
 - Fermented foods*
 - Food sensitivity testing

Key Take Aways

1. **Remember the T in ATMs.** In a **SYSTems-Based approach**, when we think of autoimmunity, we want to consider the underlying triggers within the context of the *perfect storm* : Bugs, environment, food, stress/trauma

Key Take Aways

1. **Remember the T in ATMs.** In a **SYSTems-Based approach**, when we think of autoimmunity, we want to consider the underlying triggers within the context of the *perfect storm*: Bugs, environment, food, stress/trauma
2. **When in doubt, start with the gut.** Autoimmune disease exists on a spectrum with **interplay of GI dysfunction and immune dysregulation**, with symptoms developing/manifesting as a consequence of dysbiosis, intestinal permeability, and issues with nutrient absorption and digestion.

Key Take Aways

1. **Remember the T in ATMs.** In a **SYSTems-Based approach**, when we think of autoimmunity, we want to consider the underlying triggers within the context of the *perfect storm*: Bugs, environment, food, stress/trauma
2. **When in doubt, start with the gut.** Autoimmune disease exists on a spectrum with **interplay of GI dysfunction and immune dysregulation**, with symptoms developing/manifesting as a consequence of dysbiosis, intestinal permeability, and issues with nutrient absorption and digestion.
3. **Food as Medicine.** Diet and lifestyle factors are an essential foundation for mitigating autoimmune disease and putting our patients in **remission**.

Today's Agenda

1. Describe the pathophysiological pathways that lead to allergic and autoimmune disease
2. Examine complex interaction of immune disease as it relates to GI integrity
3. Identify appropriate dietary modifications for patients with immune disorders.

How did we do?



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Questions

