



# ***Introduction to Functional Medicine***



*Dr. Jess Armine*

SOLVING THE HEALTH PUZZLE

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- Applied Kinesiology, Live Blood Cell Analysis, Nutritional Counseling

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***“The Neurotransmitter Whisperer”***

***“The Sherlock Holmes of Chronic Illness”***

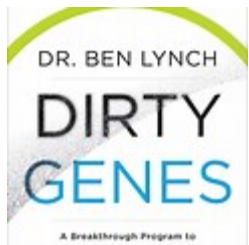
## Financial & Competing Interests disclosure

I am a self-employed, independent Health Care Practitioner in the United States.

I am not compensated for this webinar.

I have no Financial or Competing Interests with the LDN Research Trust or any other person or entity mentioned herein.

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## Dr Ben Lynch

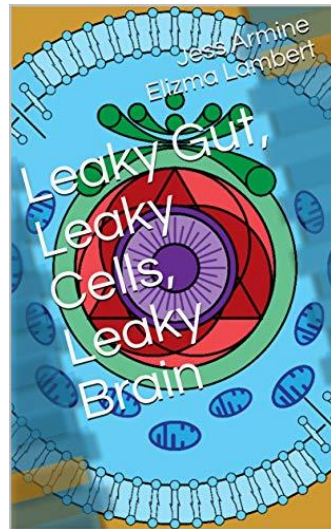
For the use of  
Strategene and concepts  
from his book, “Dirty  
Genes”

drbenlynch.com  
seekinghealth.com



Elizma Lambert, ND  
(elizmalambert.com)

For use of the concepts in her book  
coauthored with myself “*Leaky Gut,  
Leaky Cells, Leaky Brain*”



**Gilian Crowther,**  
NT/ND, Fellow of BANT,  
mNNA, CNHC reg.

**aonm.org**

**For the use of her  
CDR Slides**





# Today's Webinar is Unique

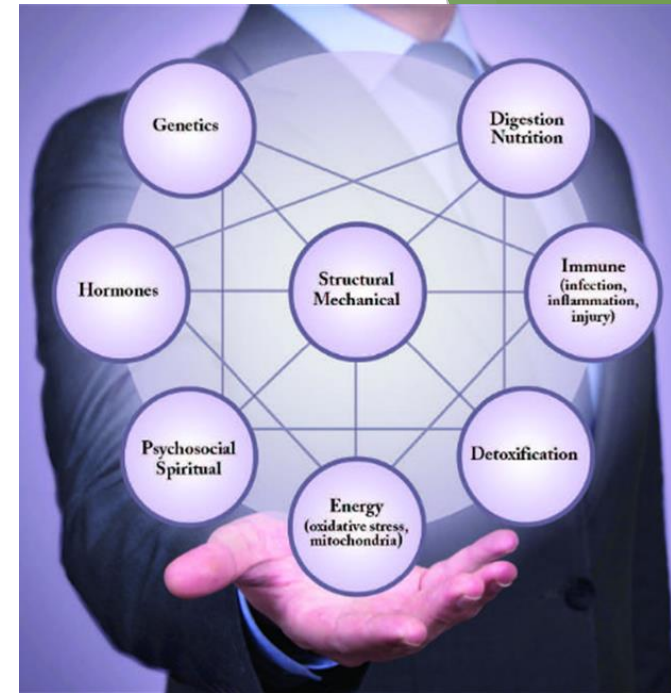
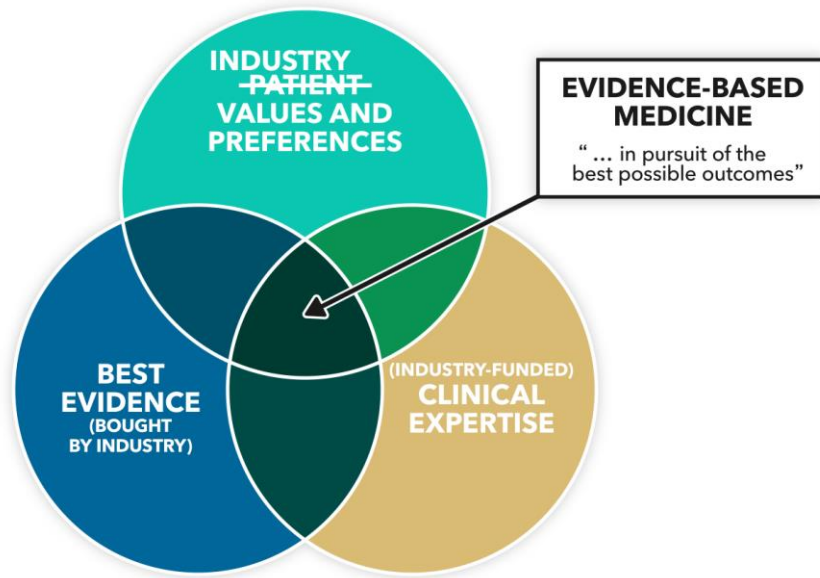
# What you can expect today...

## For Everyone

- ▶ Define what a Functional Medicine Practitioner is and does.
- ▶ Discuss the difference between classical and functional medicine.
- ▶ Show you how you benefit from true integrative approach
- ▶ Put it all together with Case Studies.
- ▶ Tips on how to pick a practitioner

## For the professionals

- ▶ The purpose of today's lecture is to be cohesive not divisive.
- ▶ For far too long our professions have been in divergence to the detriment of those who suffer.
- ▶ Please do not take anything that I say amiss as I am not being adversarial but will point out simple truths.
- ▶ We need to work together for the highest good of those we serve.



## Classical (Traditional) vs Functional Medicine\*

\*Please Note: I will use the term Functional Medicine Practitioner (FMP) to represent any healthcare professional that practices within the parameters that I am about to share with you. These practitioners have varied backgrounds and degrees

# ***Functional Medicine, the view of Classical Medicine***

- ▶ **Functional medicine** is a form of alternative medicine that encompasses a number of unproven and disproven methods and treatments.
- ▶ Its proponents claim that it focuses on the "root causes" of diseases based on interactions between the environment and the gastrointestinal, endocrine, and immune systems to develop "individualized treatment plans."
- ▶ It has been described as pseudoscience, quackery, and at its essence a rebranding of complementary and alternative medicine.



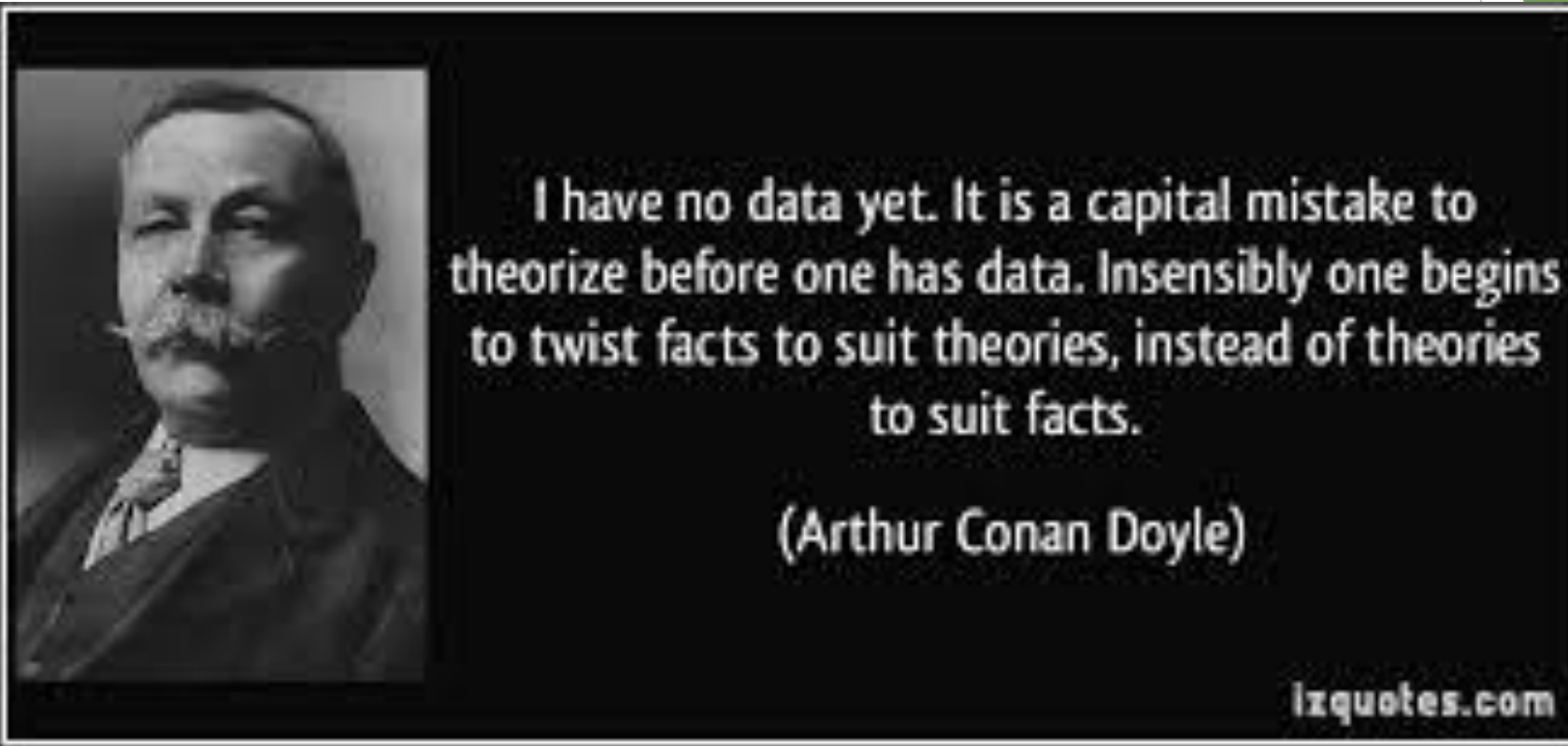
# *This begs the question...why would*

*The assessment of root causes.*

*The proven relationship between environment, endocrine,  
and the immune system (ex: psychoneuroendocrinology,  
Neuroendoimmunology, Gut-Brain Axis, etc.)*

*The development of individualized treatment plans*

***Be considered quackery?***



## WORDS TO LIVE BY



## Classical (Traditional) Medicine

- ▶ Takes sets of symptoms and labels them as diseases for the purpose of identifying a matching pharmaceutical protocol.
- ▶ Essentially, treating the result of an illness, not the illness.
- ▶ How did it get this way?

## 1960's and before

- ▶ The GP (General Practitioner) was king.
- ▶ The GP was the practitioner that knew you, your family, and was your advocate.
- ▶ Any specialists would report directly to your GP and he/she would take it from there,

## 1970's and beyond

- ▶ Transition to specialists and the GP was snubbed by the medical community.
- ▶ Medical training was done by referring to algorithms with suggested treatment protocols (Little and Brown “spirals”).
- ▶ Medicine became corporatized.
  - ▶ “Suggested Protocols” become standards of care.
  - ▶ Diagnoses must be “proven” by tests or treatment not supported.
  - ▶ Physicians are given less and less time to be with their patients.
  - ▶ Doctors have barely enough time to address the chief complaint and become “hemmed in” to their specialties.

**Doctors can't be physicians any longer resulting in treatment according to the Acute Care model. Those with chronic conditions suffer the most.**



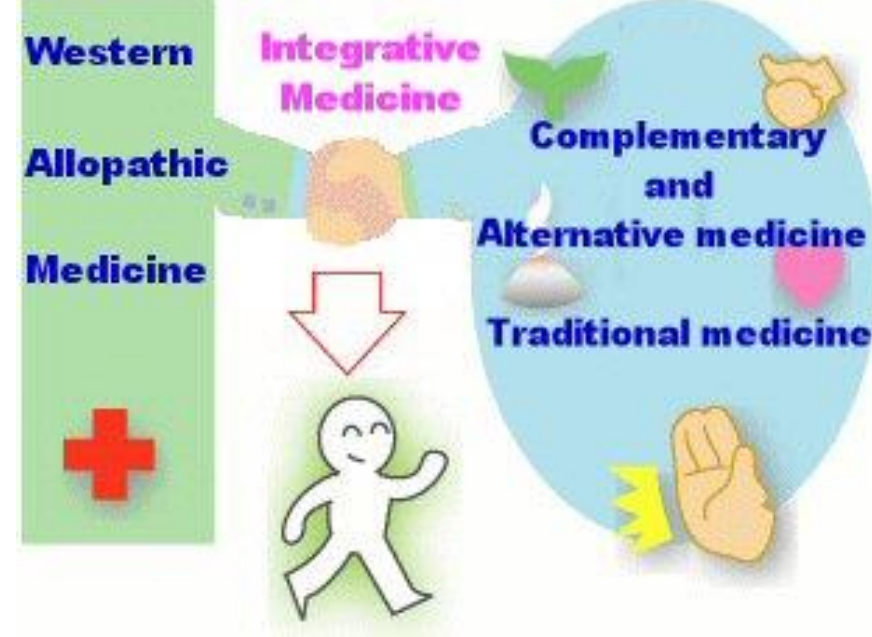
# Classical vs Functional Thought Patterns

## Acute Care Model

- ▶ **Premise:** Eradicate the root cause and the body will heal itself
- ▶ Results in chronic conditions:
  - ▶ Confusion as to why someone will not heal
  - ▶ Blaming the patient (histrionic, sx magnification, malingering)
  - ▶ Conclusion that certain pathologies cannot be healed.

## The FMP Model

- ▶ **Premise:** There are root causes and downstream effects (Symptoms created by the root causes)
- ▶ Both must be identified and treated
- ▶ In chronic conditions, the homeostatic(healing) mechanisms won't "re-boot" w/o intervention.
- ▶ Both foundational treatment and targeted root cause treatment are administered.
- ▶ In addition, the FMP will delve into the effect of the person's belief systems and coach them into a healthy mindset



Neither Allopathic nor Functional Medicine have all the answers!

Healthcare needs **TRUE** Integrative Medicine...  
A blending of allopathic and Functional.

# Collaborating is the best way. What holds us back? The way we think about things.

Basing	Ignoring	Ignoring	Forgetting
Basing our treatment parameters solely on “scientific proof” as demonstrated by placebo-controlled, double-blind studies	Ignoring observational or anecdotal evidence that may lack enough “scientific” studies	Ignoring intuitive insight	Forgetting the wisdom of Albert Einstein: <ul style="list-style-type: none"><li>• The intuitive mind is a sacred gift and the rational mind is a faithful servant. We have created a society that honors the servant and has forgotten the gift.</li></ul>

# Today, Let Agree to Think Differently

- ▶ Today let's:
  - ▶ To not depend on a single source of data
  - ▶ Consider data that was heretofore considered unusable because it was “unproven”, “alternative”, “woo woo”, or simply unfamiliar to us
- ▶ How will we do this?
  - ▶ Accept a combination of scientific and clinical data utilizing intuitive insight
  - ▶ Never, ever say (or think), “That can’t happen”.
  - ▶ Open our minds to what works but saying/thinking, “I wonder how that happened”?



# Why has Functional Medicine Flourished?

- ▶ When a desperate mother has a suffering child that no one can diagnose or treat successfully.
- ▶ It is the FMP that takes up the gauntlet.
- ▶ We will think outside the box.
- ▶ In other words, *“Who you gonna call?”*



▶ <https://www.imdb.com/title/tt0087332/atch!>

# FMP shines with Chronic Illnesses

*Autoimmune disorders, Fibromyalgia, MS, Parkinson's, Bipolar Disorder, ADD, ADHD, OCD, Depression, Dementia, Alzheimer's, Dysautonomia, Multiple Chemical Sensitivities, etc.*

## What they are

- ▶ Chronic illness have root causes that have resulted in expressions specific to the diagnosis.
- ▶ Chronic illnesses are pathophysiologic processes. As such, they can be resolved.
- ▶ Chronic illnesses often Require a multidisciplinary approach but most of all:
  - ▶ Chronic illnesses require a different point of view on the part of the practitioner.

## What they are not

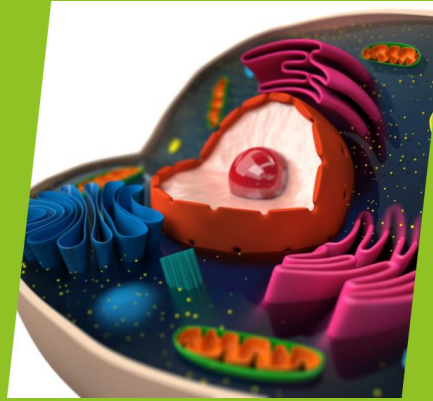
- ▶ You are not born with a chronic illness
- ▶ Chronic illnesses are not the fault of the patient
- ▶ Chronic illnesses are not chance occurrences or "rolls of the cosmic dice".
- ▶ Autoimmune disorders are NOT unrecoverable as they have precipitating factors that initiate the pathologic processes

# Functional Practitioners Thought Pattern

What does it mean to think “Outside the Box”?

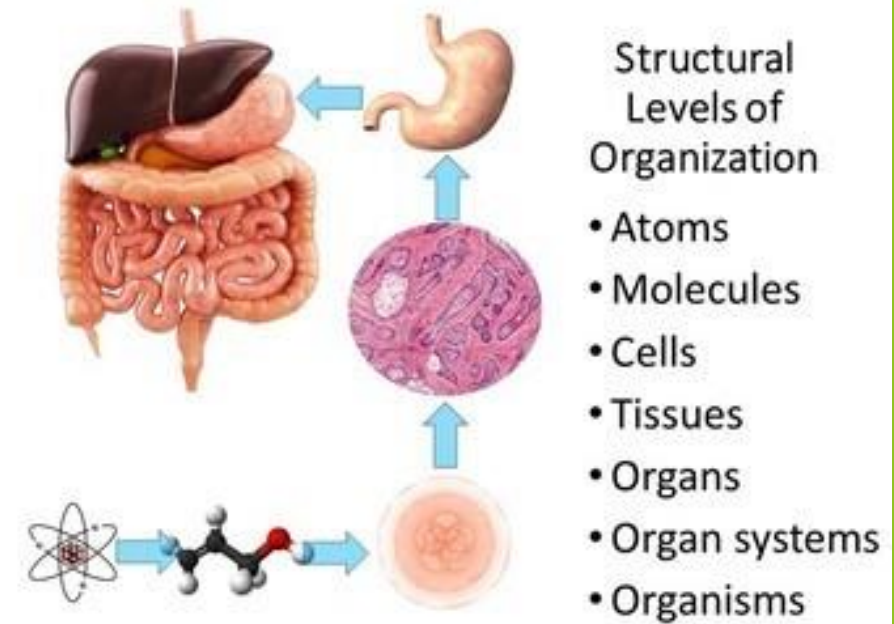
## Basic Premise:

All of life happens within the cell and is protected and supported by the cell membrane



► ***Heal the cells and you heal the body!***

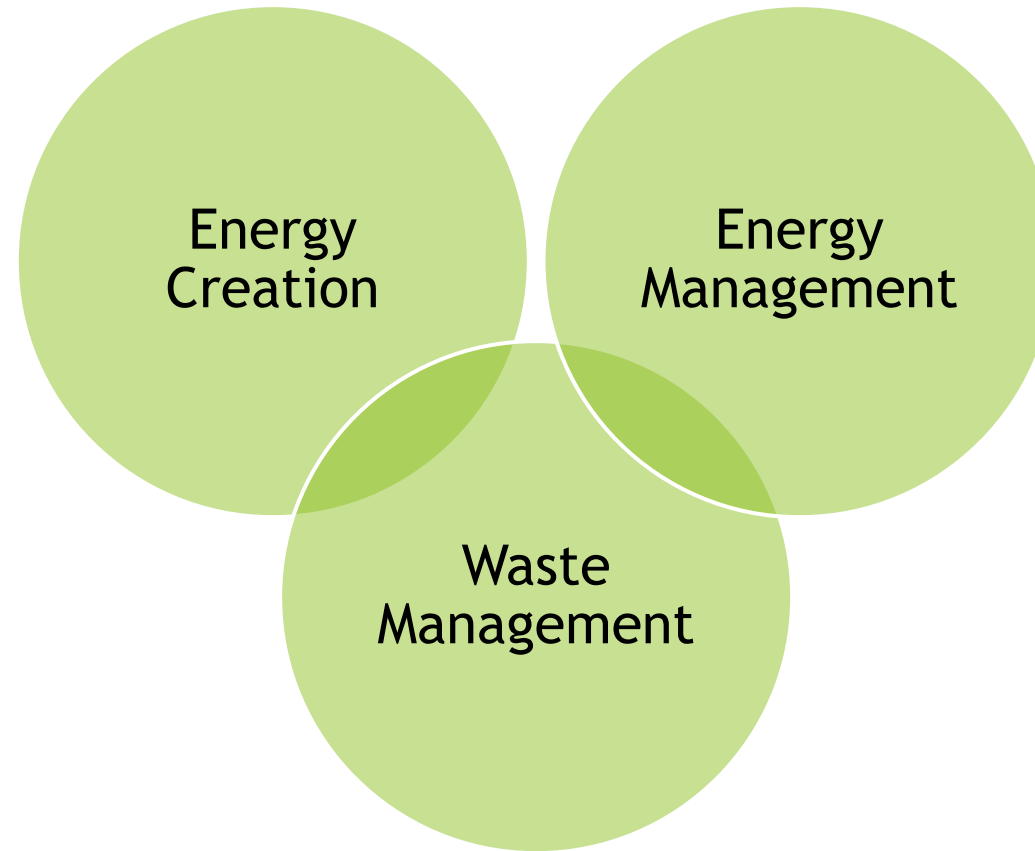
## Levels of Organization



& Life Functions



# Basis of Cellular Function...



# Bio-Individualized Medicine™



Bio-Individualized  
Medicine™ is a  
*Thought Paradigm*  
that includes  
consideration of  
ALL the following  
parameters



## *Ever Notice that Many Diseases Have Common Symptoms... Perhaps There is Common Causation?*

Symptom	Chronic Lyme	Fibromyalgia	ME/CFS	Dysautonomia
Fatigue	X	X	X	X
Chronic Pain	X	X	X	X
Mood Changes	X	X	X	X
Confusion/ Brain Fog	X	X	X	X
Numbness Tingling	X	X	X	X
Sensitivity to light	X	X	X	X
Inflammation	X	X	X	X



## The Cell Danger Response (CDR)

Naviaux, R.K., Metabolic features of the cell danger response, Mitochondrion (2013), <http://dx.doi.org/10.1016/j.mito.2013.08.006>

# What is the Cell Danger Response?

Mitochondrion 16 (2014) 7–17



Contents lists available at ScienceDirect

Mitochondrion

journal homepage: [www.elsevier.com/locate/mito](http://www.elsevier.com/locate/mito)



## Metabolic features of the cell danger response



Robert K. Naviaux\*

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Veterans Affairs Center for Excellence in Stress and Mental Health (CESAMH), La Jolla, CA, USA*

### ARTICLE INFO

Available online 24 August 2013

#### Keywords:

Oxidative stress  
Oxidative shielding  
Innate immunity  
Inflammation  
Purinergic signaling  
Mitochondria

### ABSTRACT

The cell danger response (CDR) is the evolutionarily conserved metabolic response that protects cells and hosts from harm. It is triggered by encounters with chemical, physical, or biological threats that exceed the cellular capacity for homeostasis. The resulting metabolic mismatch between available resources and functional capacity produces a cascade of changes in cellular electron flow, oxygen consumption, redox, membrane fluidity, lipid dynamics, bioenergetics, carbon and sulfur resource allocation, protein folding and aggregation, vitamin availability, metal homeostasis, indole, pterin, 1-carbon and polyamine metabolism, and polymer formation. The first wave of danger signals consists of the release of metabolic intermediates like ATP and ADP, Krebs cycle intermediates, oxygen, and reactive oxygen species (ROS), and is sustained by purinergic signaling. After the danger has been eliminated or neutralized, a choreographed sequence of anti-inflammatory and regenerative pathways is activated to reverse the CDR and to heal. When the CDR persists abnormally, whole body metabolism and the gut microbiome are disturbed, the collective performance of multiple organ systems is impaired, behavior is changed, and chronic disease results. Metabolic memory of past stress encounters is stored in the form of altered mitochondrial and cellular macromolecule content, resulting in an increase in functional reserve capacity through a process known as mitocellular hormesis. The systemic form of the CDR, and its magnified form, the purinergic life-threat response (PLTR), are under direct control by ancient pathways in the brain that are ultimately coordinated by centers in the brainstem. Chemosensory integration of whole body metabolism occurs in the brainstem and is a prerequisite for normal brain, motor, vestibular, sensory, social, and speech development. An under-

“An evolutionarily conserved response activated when a cell encounters a threat that could injure or kill it”



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## Metabolic features of the cell danger response



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### ABSTRACT

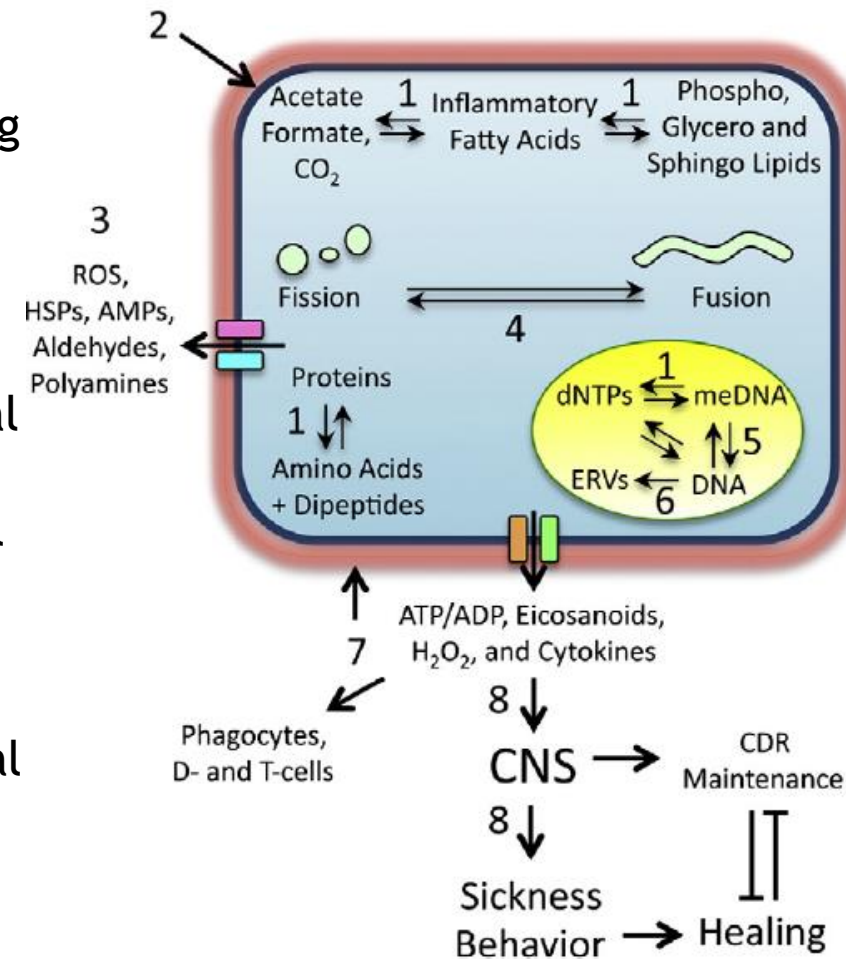
The cell danger response (CDR) is the evolutionarily conserved metabolic response that protects cells and hosts from harm. It is triggered by encounters with chemical, physical, or biological threats that exceed the cellular capacity for homeostasis. The resulting metabolic mismatch between available resources and functional capacity produces a cascade of changes in cellular electron flow, oxygen consumption, redox, membrane fluidity, lipid dy-

... Our mitochondria downregulate as a protective mechanism



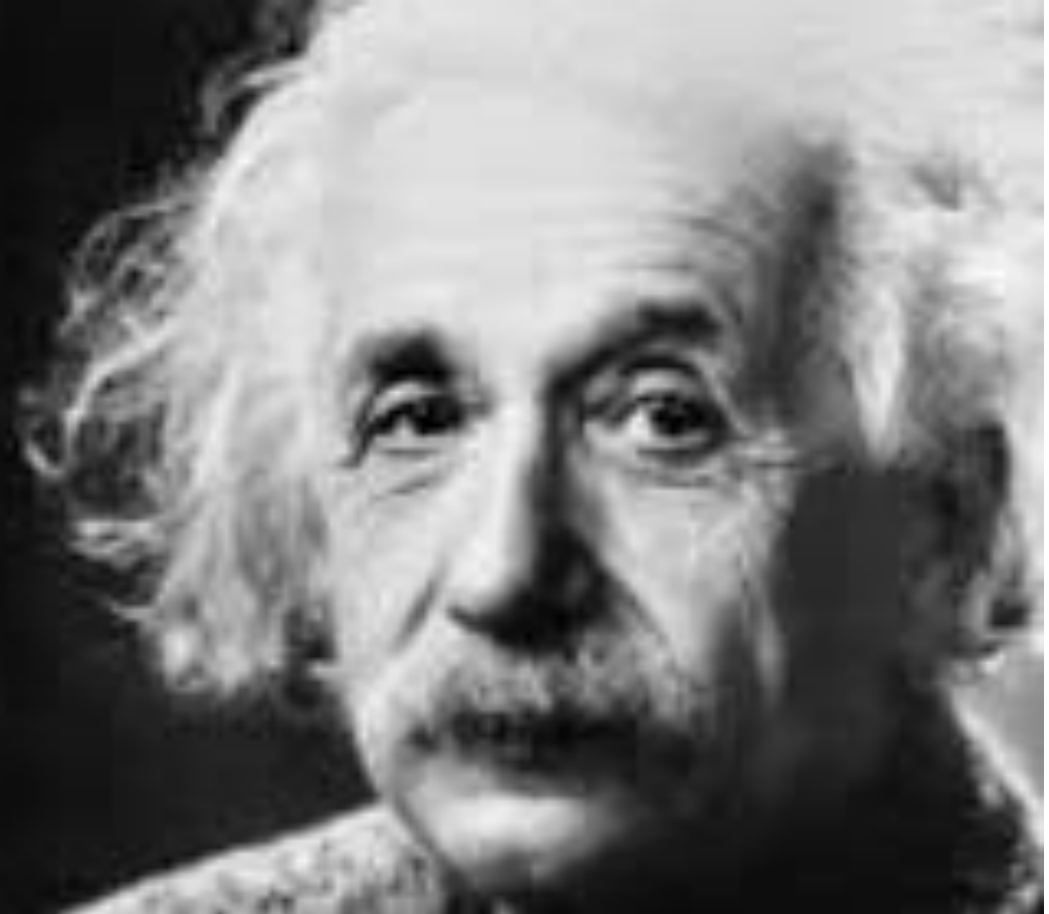
# What are the immediate results of acute CDR?

- 1) Mitochondria decrease oxygen consumption to oxidise the cellular environment, inhibiting assembly of monomeric building blocks into polymers, thus decreasing efficiency of RNA, protein, and DNA synthesis by the infecting pathogen
- 2) Stiffen cell membranes to limit pathogen egress
- 3) Release of antiviral and antimicrobial chemicals (HOCL)
- 4) Increase in autophagy/mitochondrial fission/mitophagy
- 5) Changes in DNA methylation: SAM is directed to polyamine synthesis to assist ROS and antiviral/antimicrobial polyamine aldehyde synthesis and release, lowering the SAM/SAH ratio



"If you can't explain it  
simply, you don't  
understand it well  
enough."

**- Albert Einstein**



# *The Cell Danger Response ?*

## *Put Simply:*

- ▶ Metabolic response of the cell to protect itself (and thereby you) from harm
- ▶ The basis of re-establishing homeostasis (normal function)
- ▶ It all occurs in the ***Mitochondria***





# ***MITO: THE MOVIE***

***THE MEDICAL COMMUNITY'S BEST KEPT SECRET - NOW REVEALED!***

YES!  
IT MAY BE  
ALL IN  
YOUR HEAD!

BUT,  
IT COULD BE  
AFFECTING  
OTHER  
ORGANS TOO!



GOT MITO?  
YOU  
ARE NOT  
ALONE!

DON'T KNOW  
A THING  
ABOUT  
MITO?

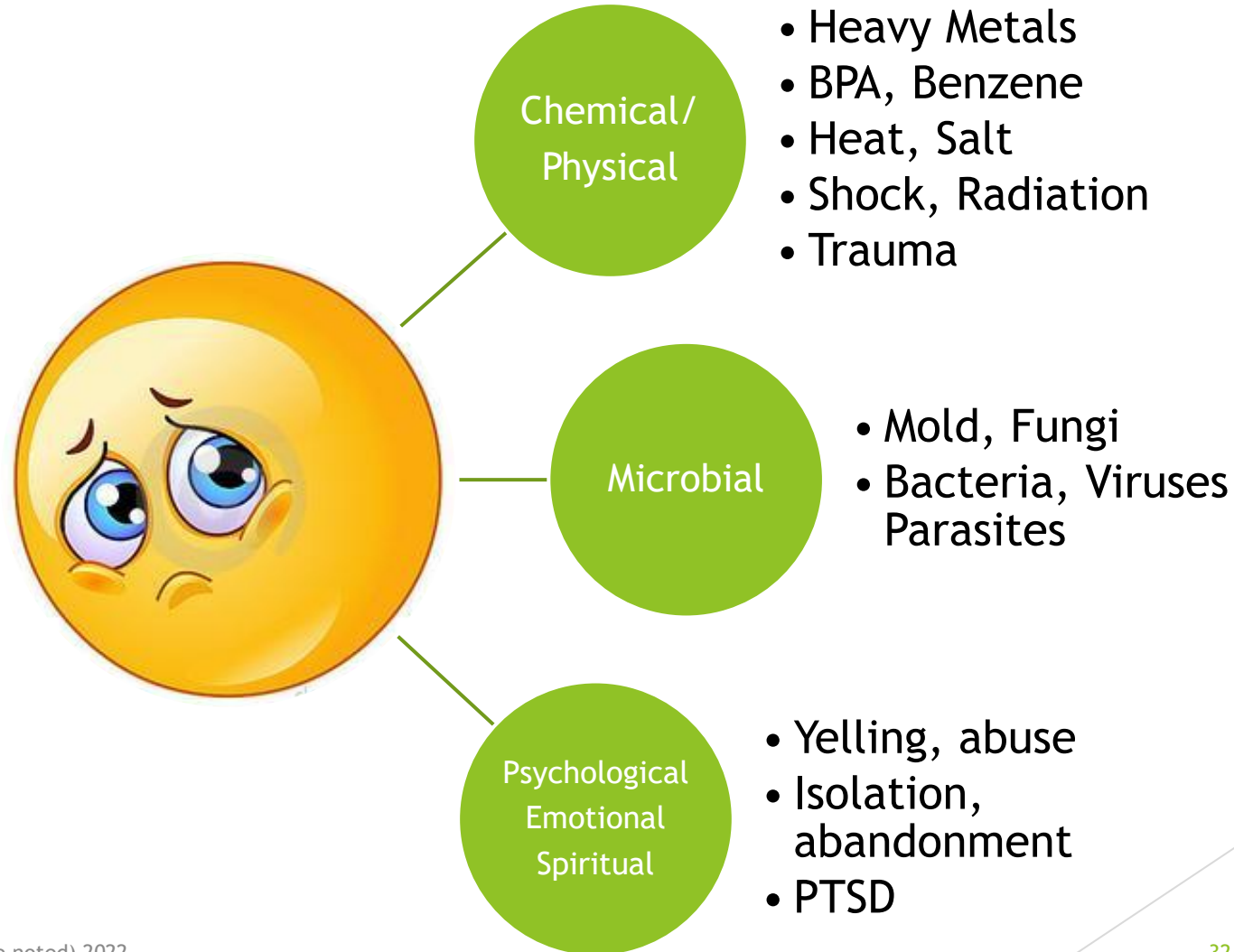
YOU TOO  
ARE NOT  
ALONE!

## **COMING SOON!**



# What activates the Cell Danger Response?

(and thereby damages the Mitochondria)





It's all in your head!

## The Relevance of Emotions on Health



# The Heart as a Psychoneuroendocrine and Immunoregulatory Organ

Carlo Dal Lin 1, Francesco Tona 1, Elena Osto 2 3

## Abstract

The heart can be viewed not just as muscle pump but also as an important checkpoint for a complex network of nervous, endocrine, and immune signals. The heart is able to process neurological signals independently from the brain and to crosstalk with the endocrine and immune systems. The heart communicates with the psyche through the neuro-endocrine-immune system in a highly integrated way, in order to maintain the homeostasis of the whole body with peculiarities specific to males and females.

The heart is able to process neurological signals independently from the brain

The heart communicates with the psyche through the neuro-endocrine-immune system in a highly integrated way

*Dal Lin C, Tona F, Osto E. The Heart as a Psychoneuroendocrine and Immunoregulatory Organ. Adv Exp Med Biol. 2018;1065:225-239. doi: 10.1007/978-3-319-77932-4\_15. PMID: 30051388.*

34



The **CDR** results in a Cascade of Changes...

# ...temporary interference in:

Lipid Dynamics

Cellular Electron  
Flow  
(Mitochondria-  
Energy)

O2 Consumption  
(Krebs-Energy)

Cellular fluidity  
(cell wall integrity)

Vitamin Availability  
(Biochemical  
Pathways)

Metal Homeostasis  
(how we get heavy  
metal burden just  
by breathing)

aggregation



# When the Danger has Passed...

- ▶ Sequence of anti-inflammatory and regenerative pathways are activated to:
  - ▶ Reverse CDR
  - ▶ Promote Healing
  - ▶ The interference is removed, and homeostasis (normal cell function) restored (Think of it as RE-BOOTING).
- ▶ BUT...

**IN CHRONIC OR MULTIPLE CDR  
THE INTERFERENCE REMAINS AND WORSE,  
SYNERGIZES!**



Chronic /Multiple CDR

Numerous Downstream effects (symptoms)

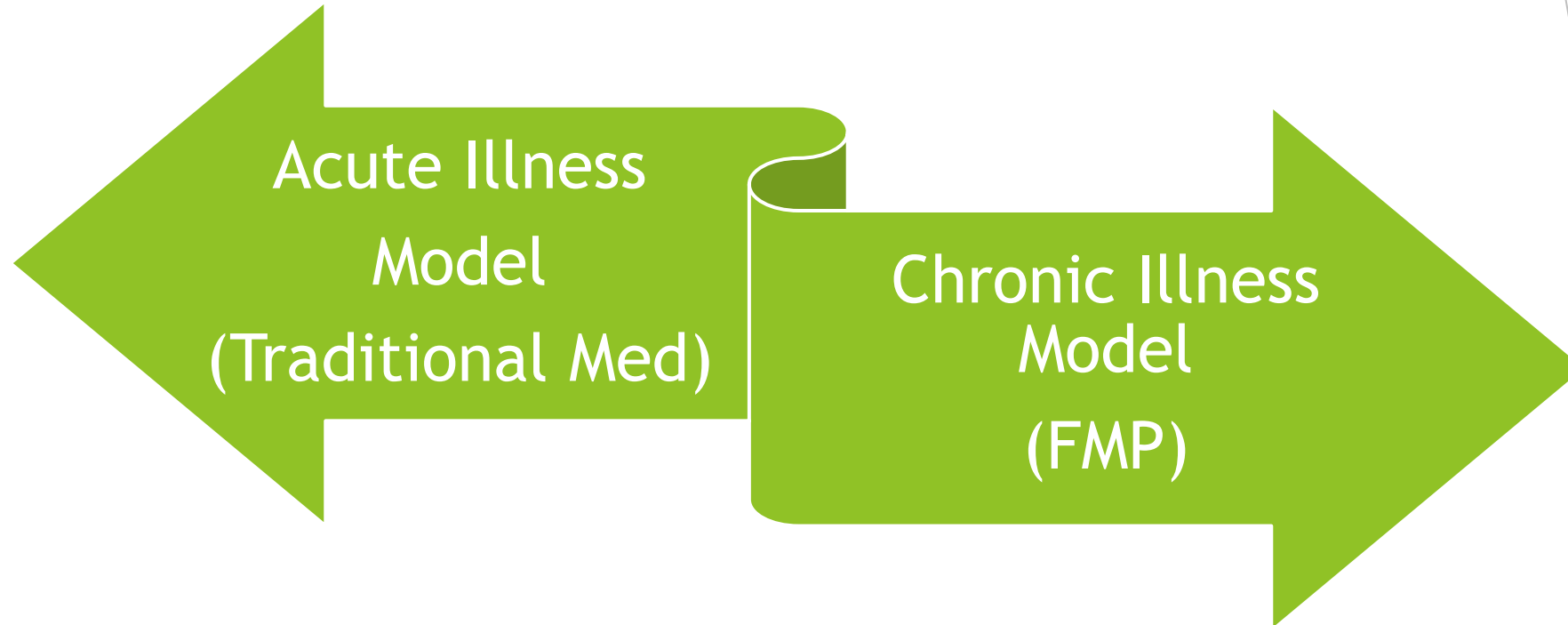
Healing Mechanisms will not re-boot.

The Negative Effects on the healing mechanisms synergize

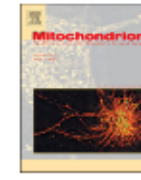
**Healing Becomes Impossible Unless Treating  
the Root Causes AND Downstream Effects**

**HERE'S WHERE THE DIVERGENCE BETWEEN ALLOPATHIC AND FMP THINKING  
OCCURS**

There is a significant difference in treating acute vs chronic illness.







## Metabolic features and regulation of the healing cycle—A new model for chronic disease pathogenesis and treatment

Robert K. Naviaux

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## ARTICLE INFO

**Keywords:**

Cell danger response  
Healing cycle  
Mitochondrial nexus  
Metabolic addiction  
Metabolic memory  
Purinergic signaling  
Metabokines  
Antipurinergic therapy  
M0, M1 and M2 mitochondria  
Ecoalleles  
Ecogenetics  
Allostasis  
Allostatic load  
Integrated stress response

## ABSTRACT

Without healing, multicellular life on Earth would not exist. Without healing, one injury predisposes to another, leading to disability, chronic disease, accelerated aging, and death. Over 60% of adults and 30% of children and teens in the United States now live with a chronic illness. Advances in mass spectrometry and metabolomics have given scientists a new lens for studying health and disease. This new lens provides a new perspective on chronic illness in terms and reframes the pathophysiology of chronic illness. Chronic illness is not a block healing and cause the normal stages of the healing cycle. When an injury occurs, active progress through the stages of healing, energetics and the disposition of oxygen and carbon dioxide is required for recovery. > 100 chronic illnesses can be organized in a hierarchical manner based on targetable chemosensory G-protein coupled and ion channel receptors. Healing. Metabokines are signaling molecules derived from the pathogenesis of chronic illness in this way, as a result of a remote trigger(s) that *caused* the initial injury, perpetuates the injury, *unlock* the healing cycle, and restore health when the injury is resolved. When caring for acute health, the careful identification of the trigger, or cause of the injury, is the anatomical local

## 1. Introduction

Much of modern Western medicine is based on the principles of acute interventions for poisoning, physical injury, or infection. These principles trace to historical figures like Paracelsus (1493–1541), Ambroise Paré (1510–1590), and Louis Pasteur (1822–1895). These acute care interventions are now widely used in the modern fields of pharmacology, toxicology, urgent care, emergency medicine, and surgery. When caring for acute disruptions in health, the careful identification of the trigger, or cause of the problem, and the anatomical location of the defect, is an important part of good medical care. However, when dealing with chronic illness, treatments based on the rules of acute care medicine have proven less helpful, and can even cause harm by producing unwanted side-effects (Qato et al., 2018).

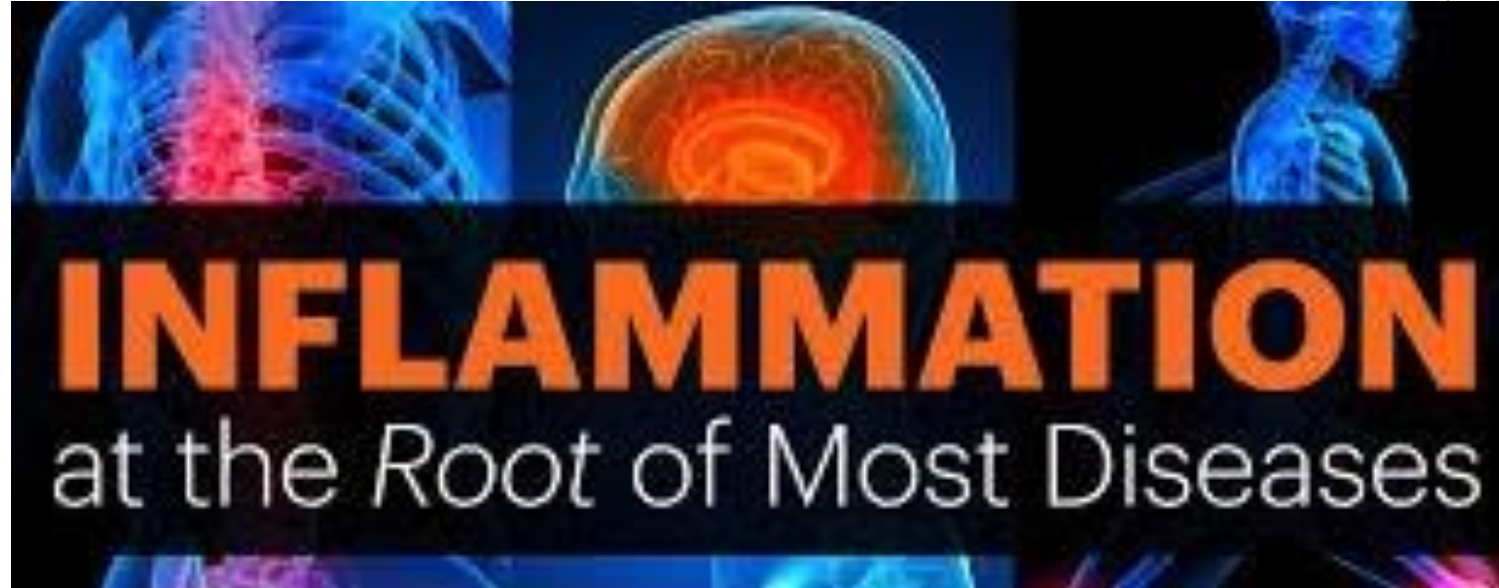
In chronic illness, the original triggering event is often remote, and may no longer be present. Emerging evidence shows that most chronic illness is caused by the biological reaction to an injury, and not the initial injury, or the agent of injury itself. For example, melanoma can

be caused by such traumatic stress. A bullet wound is more severe disease, therefore complete healing of the second injury is even when the disease is progressive dysfunction occurs in all organs results when cells are injured and re-injury, unlike every chronic illness recurrent infections, diabetic heart and pulmonary disease, fatigue syndromes, diseases, Alzheimer's.

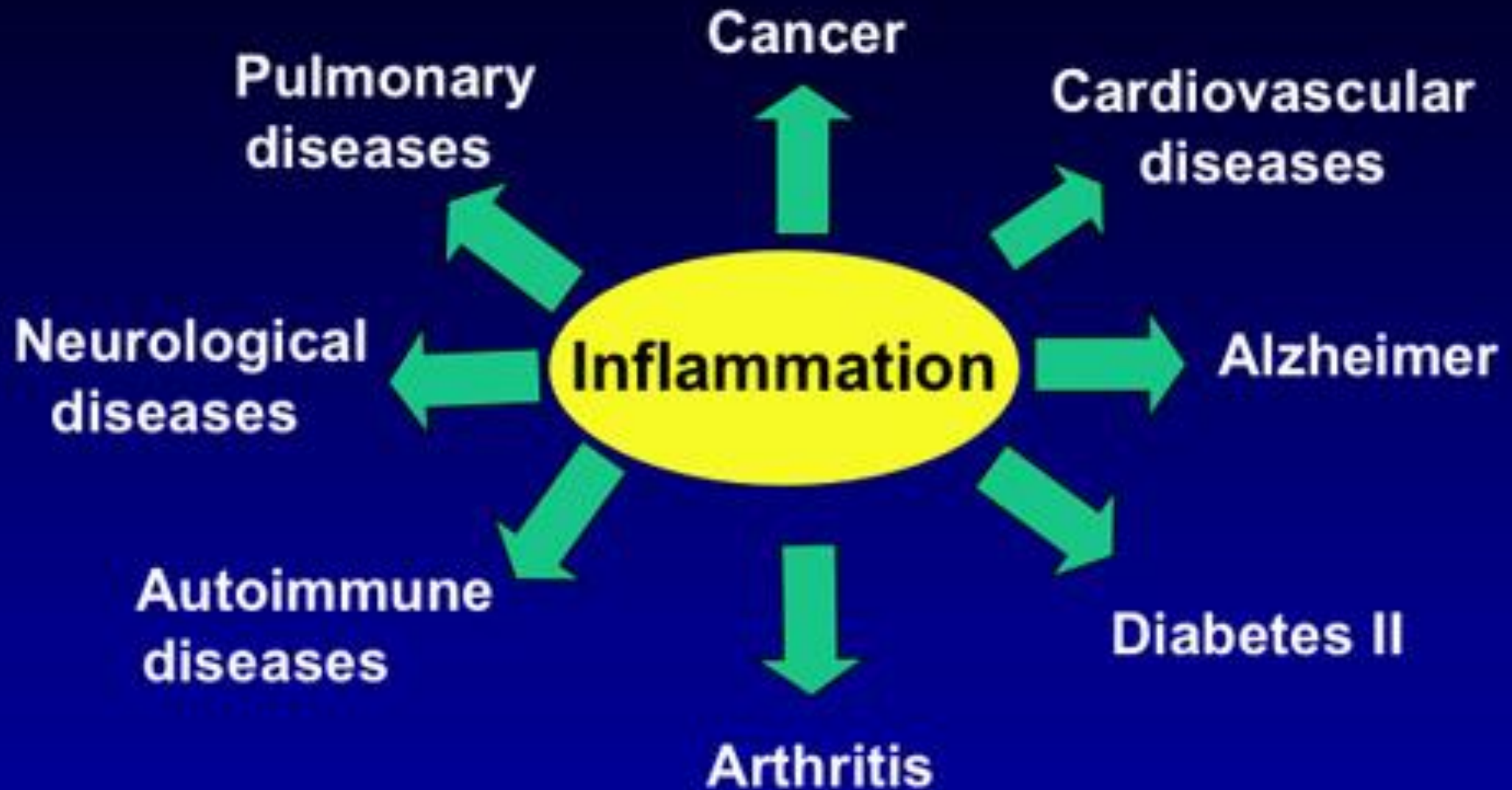
Great strides

When caring for **acute** disruptions in health, the careful identification of the trigger, or cause of the problem, and the anatomical location of the defect, is an important part of good medical care. However, when dealing with **chronic illness**, treatments based on the rules of acute care medicine have proven less helpful, *and can even cause harm by producing unwanted side-effects (emphasis added)*

Back to CDR... A “STUCK” Cell Danger Response creates



And all the suffering we face...





# HOW INFLAMMATION AFFECTS THE BODY

BROUGHT TO YOU BY



WWW.LIVELOVEFRUIT.COM

"Inflammation is at the root of practically all known chronic health conditions"  
Find out how to prevent it at [www.livelovefruit.com](http://www.livelovefruit.com)

## BRAIN

Pro-inflammatory cytokines cause autoimmune reactions in the brain, which can lead to depression, autism, poor memory, Alzheimer's disease and MS.



## SKIN

Chronic inflammation compromises the liver & kidneys, resulting in rashes, dermatitis, eczema, acne, psoriasis, wrinkles & fine lines.



## CARDIOVASCULAR

Inflammation in the heart & arterial & venous walls contributes to heart disease, strokes, high blood sugar (diabetes) and anemia.



## KIDNEYS

Inflammatory cytokines restrict blood flow to the kidneys. Complications like edema, hypertension, nephritis & kidney failure can result.



## BONES

Inflammation interferes with the body's natural ability to repair bone mass, increasing the number of fractures & leading to conditions like osteoporosis.



## LIVER

Build-up of inflammation leads to an enlarged liver or fatty liver disease. Increased toxic load build-up in the body.



## THYROID

Autoimmunity as a result of inflammation can reduce total thyroid receptor count & disrupts thyroid hormone function.



## LUNGS

Inflammation induces autoimmune reactions against the linings of airways. Can result in allergies or asthma.



## GI TRACT

Chronic inflammation damages our intestinal lining and can result in issues like GERD, Chron's disease and Celiac disease.



## MUSCLE

Inflammatory cytokines can cause muscle pain & weakness. Can manifest as carpal tunnel syndrome, or polymyalgia rheumatica, to name a few.

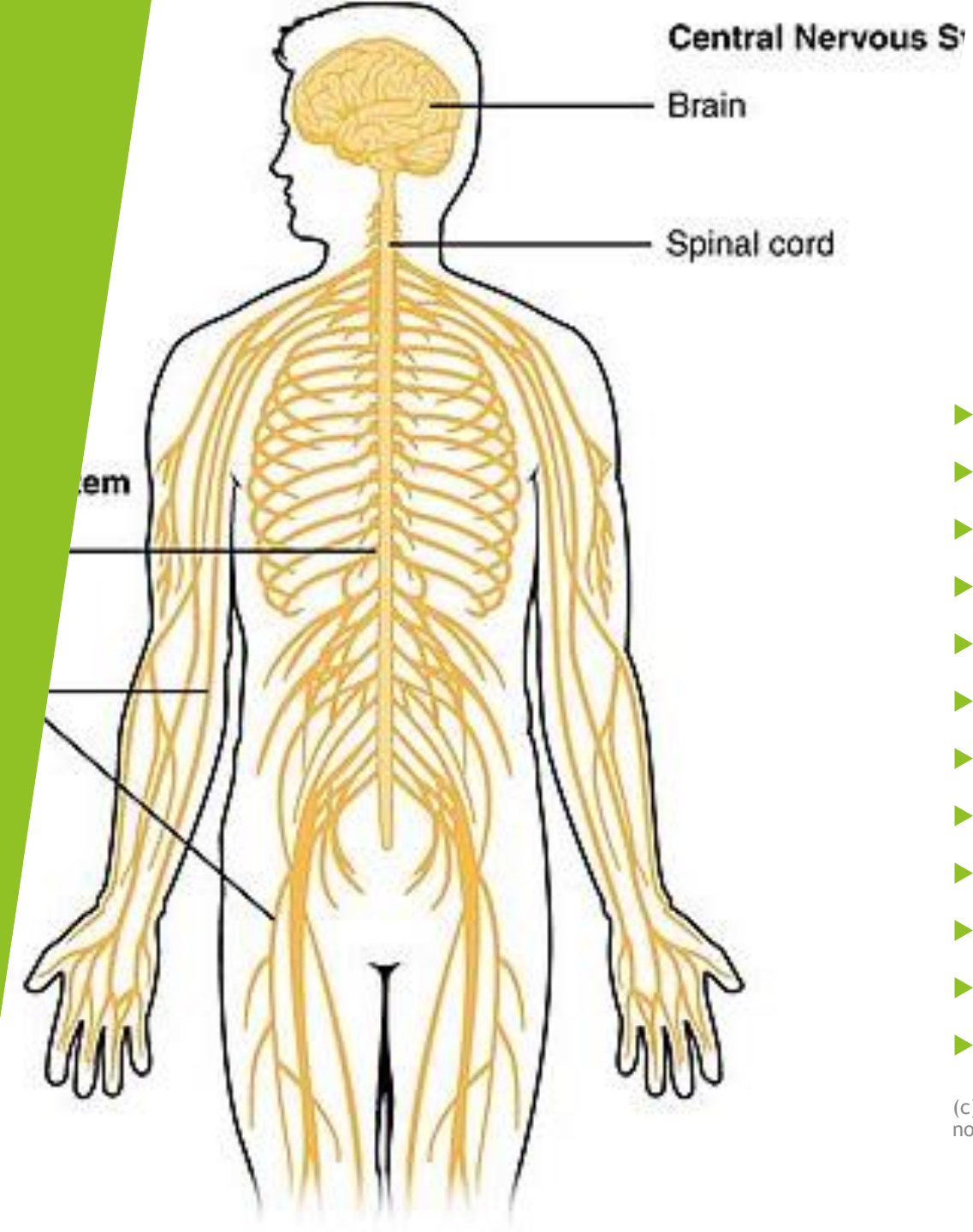


(c) Dr. Jess P. Armline 2018



# Chronic Inflammation Has Caused a Rise In Health Issues

- ▶ Cardiovascular Issues
  - ▶ GI (stomach) issues
  - ▶ Diabetes
  - ▶ Metabolic Disorders
  - ▶ “Adrenal Fatigue”
- 
- ▶ But The #1 Target For CIRS (Chronic Inflammatory Response Syndrome AKA Chronic Inflammation) Is In The.....



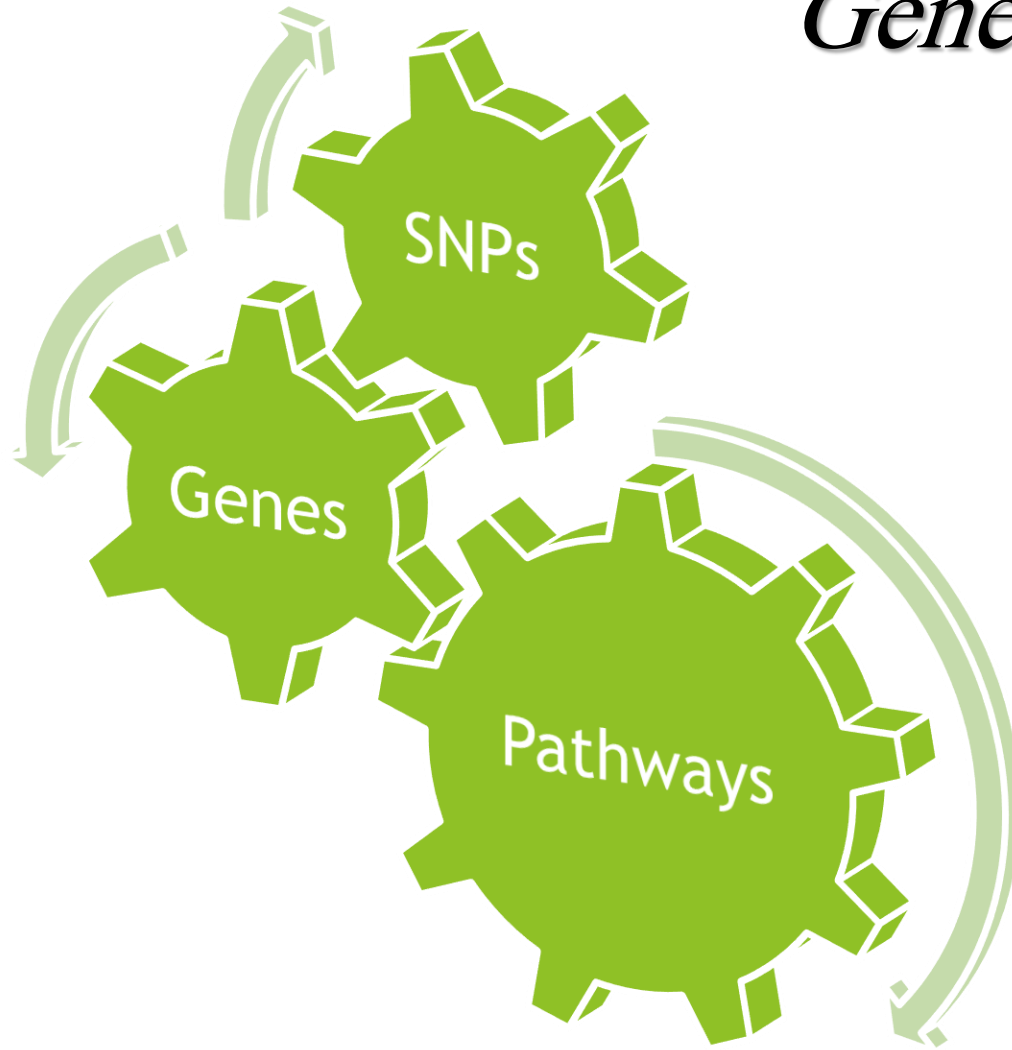
# Nervous System

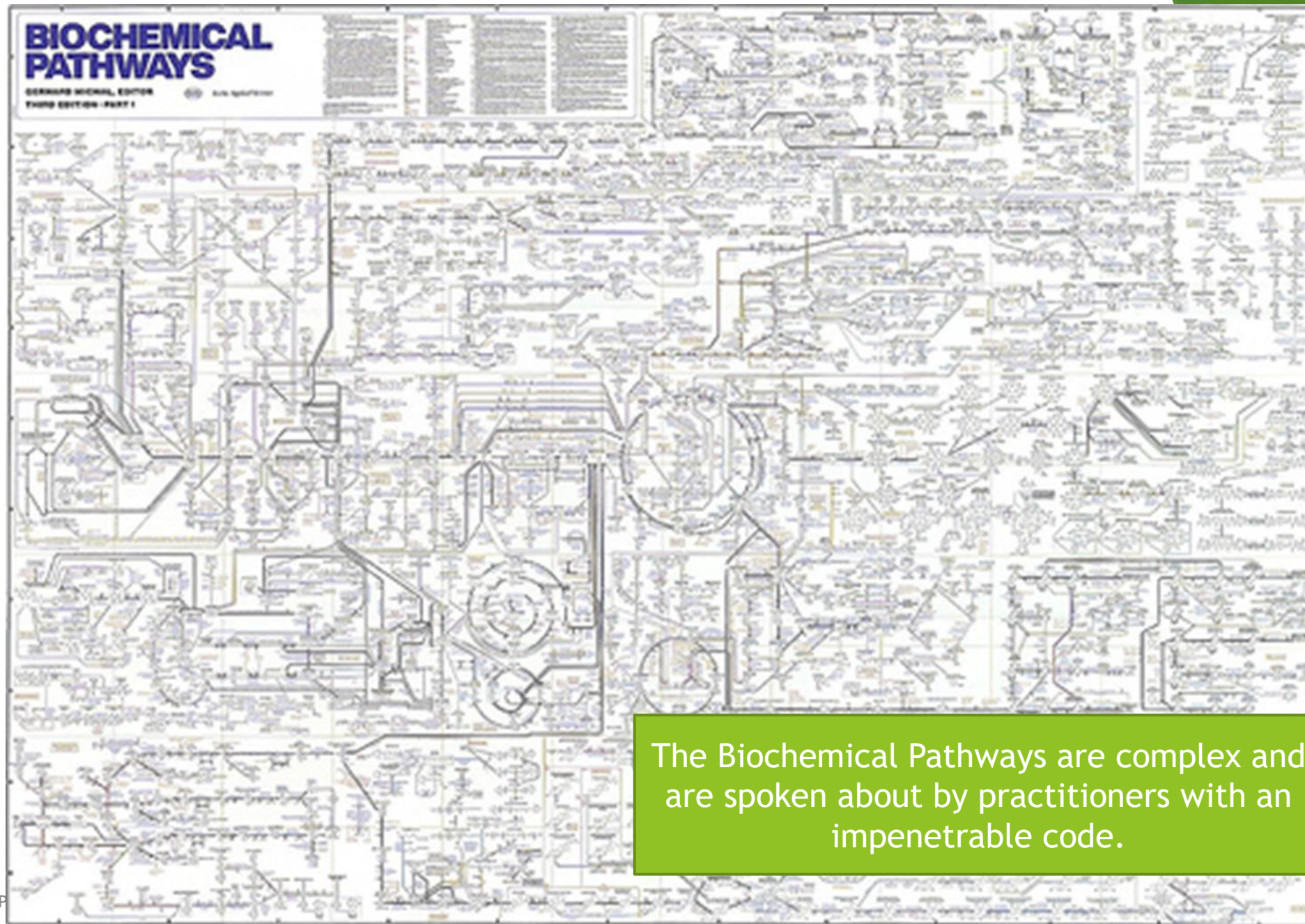
- ▶ ASD
- ▶ Anxiety
- ▶ OCD
- ▶ Migraines/Headaches
- ▶ Addictions/Cravings
- ▶ Behavioral Issues
- ▶ Dysautonomia (POTS, etc.)
- ▶ Neuropathies
- ▶ PMS/Menopausal disorders
- ▶ ADD/ADHD
- ▶ Depression
- ▶ Many More...

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# *The Role and Relevance of Genetics*





The Biochemical Pathways are complex and are spoken about by practitioners with an impenetrable code.

(c) Dr Jess P

<http://biochemical-pathways.com/#/map/1>



# SNP's (AKA Polymorphisms)...What Do They Mean?

*They are an Estimate of the Enzyme's Function*

*Think of highways of differing widths*

- / -



**Normal**  
Usual Enzyme  
Function

+ / -



**Heterozygous**  
60% Enzyme Function

+ / +



**Homozygous**  
20% Enzyme Function

# *Traffic\* Will Slow Down the Pathway's Function*

+/-



+/+



\*Traffic = bacteria, heavy metals, viruses, parasites, food allergens, candida, Leaky Gut Syndrome, lack of substrate, lack of cofactors, and coenzymes, presence of factors that will speed up or slow down enzymatic activity, etc.

# Timeless Wisdom

- ▶ THE PRESENCE OF A POLYMORPHISM DOES NOT MEAN YOU'RE ILL
- ▶ THE LACK OF A POLYMORPHISM DOES NOT MEAN YOU'RE WELL
- ▶ DR. JESS

The lack of a polymorphism (SNP) is not a guarantee that the pathway will work adequately.

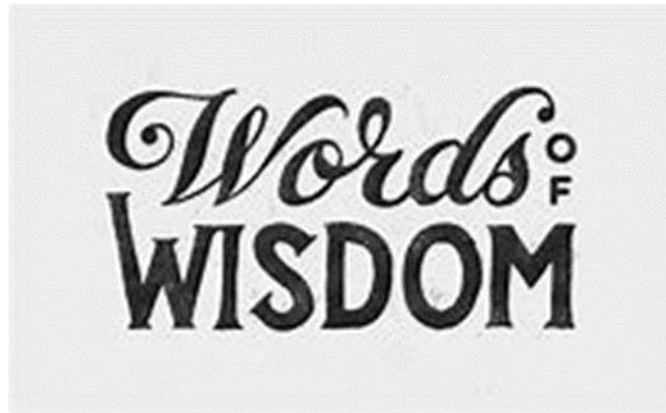
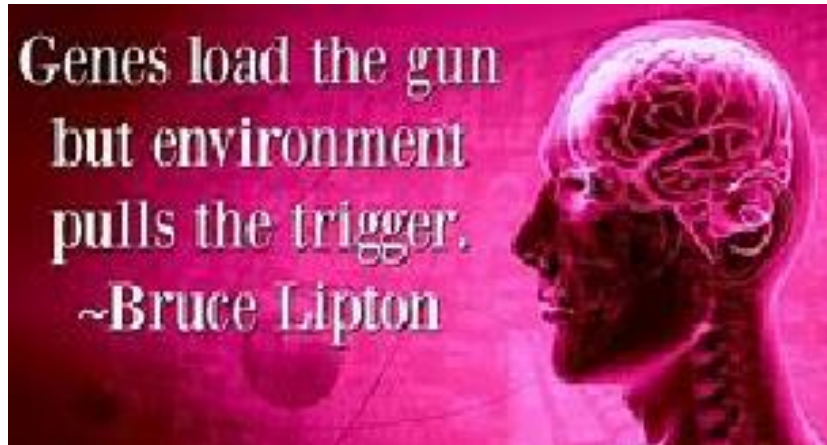
- / -

Think about it. Can you put enough traffic into an 8-lane highway to slow it down?

**YA THINK?**







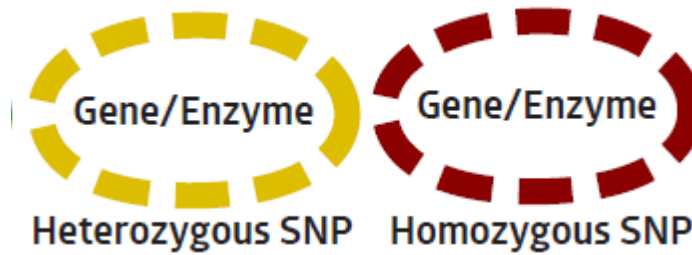
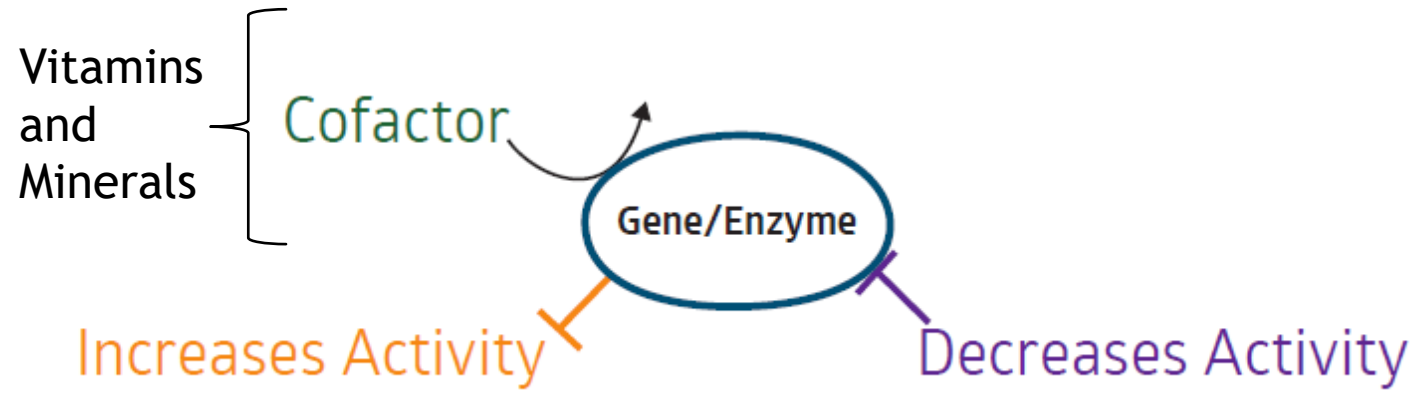
GENETICS

loads the gun, but

LIFESTYLE

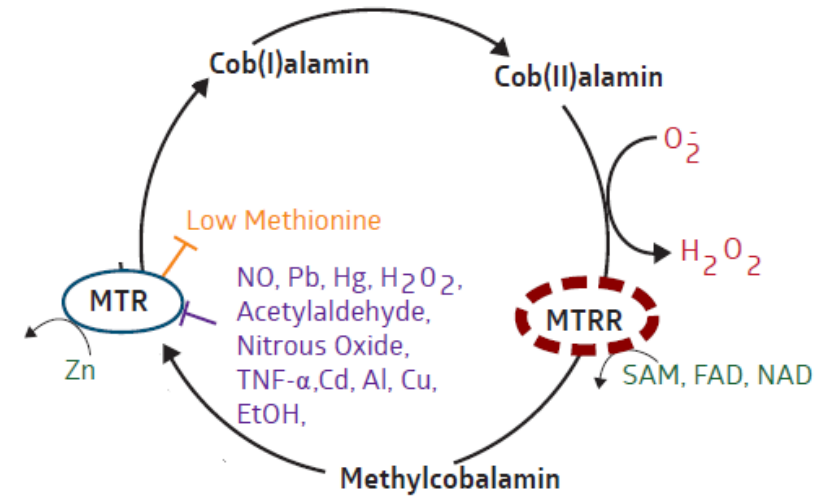
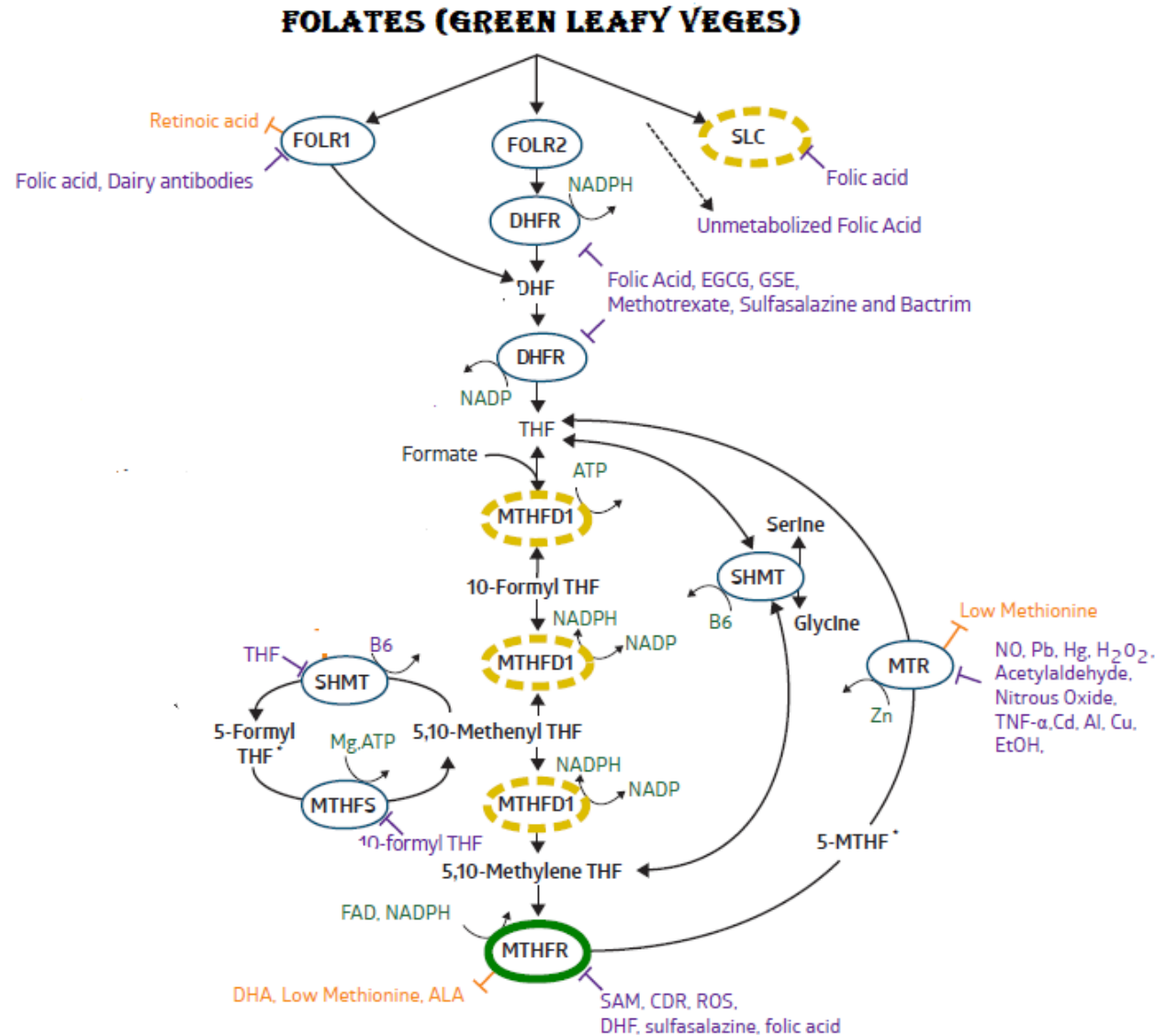
pulls the trigger.

The role of nutrition In optimizing pathway *function*



Courtesy of Dr Ben Lynch. Used with permission

# The role of nutrition and supplementation



Absorbable Folate and Cobalamin (B12)

NAD (B3), FAD (B2)

B6

Mg, Zn

SAMe

Most Important: ATP (your energy)

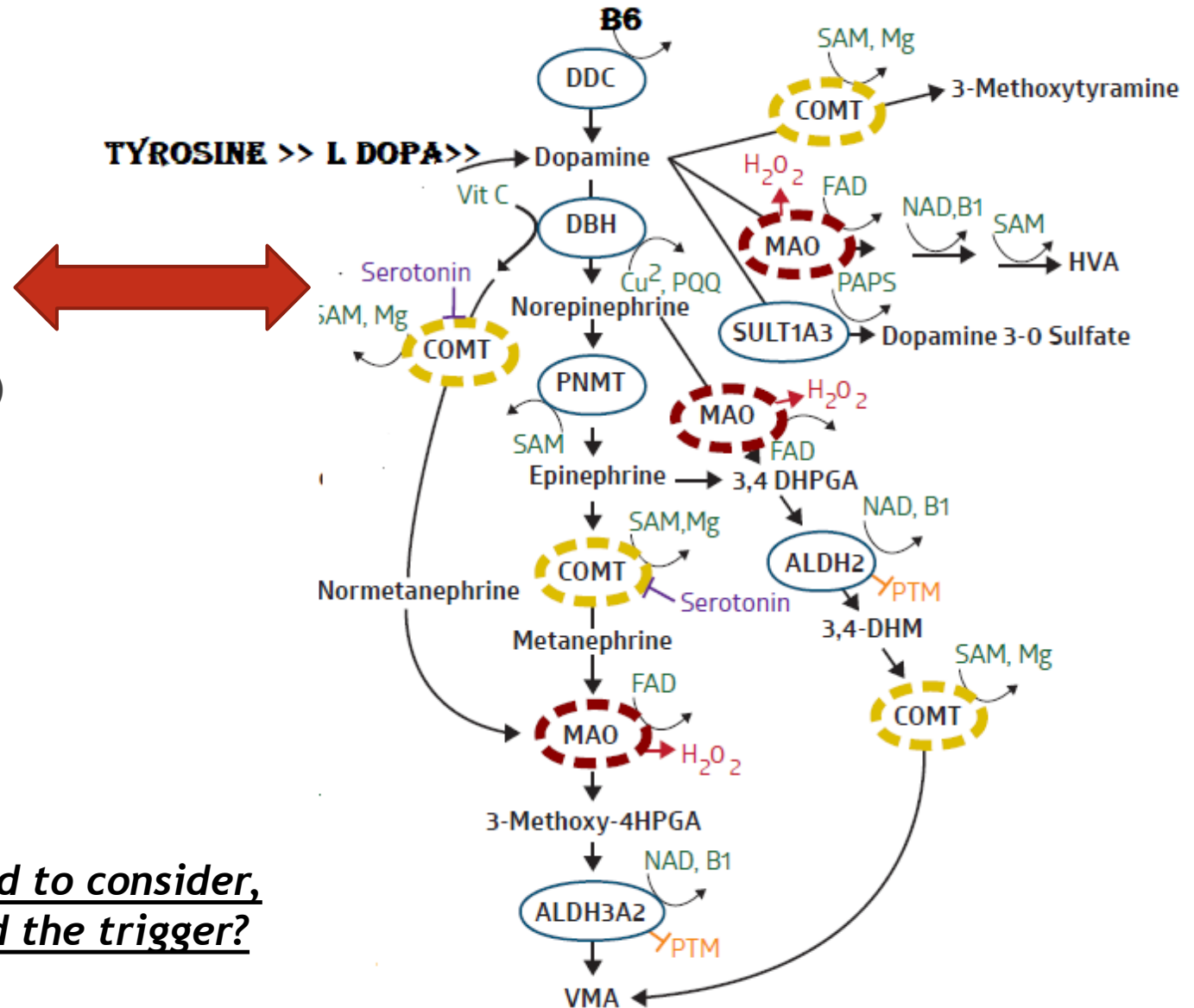
54

# What Predispositions?

- Excitation as expressed by Anxiety, OCD, ADD, etc.
- Pathway needs:
- Inhibitory support (Serotonin, GABA)
  - SAM
  - B6, Vit C
  - B1, B2 B3
  - Mg, Cu
  - PQQ



**Always need to consider, what pulled the trigger?**



rs1049748	ABP1 P574P	T	CT	+/-
rs1049793	ABP1/DAO H864A	G	CG	+/-
rs1049742	ABP1/DAO S332P	T	CC	-/-
rs10156191	ABP1/DAO T16M	T	CT	+/-
rs8911472	CNR1 A88853143C	C	AA	-/-
rs8928813	CNR1 A88861698G	G	AA	-/-
rs806380	CNR1 A88864653G	G	AG	+/-
rs806381	CNR1 A88865601G	C	AG	-/-
rs7752758	CNR1 A88866376G	G	AA	-/-
rs12528858	CNR1 A88867488G	G	AA	-/-
rs806378	CNR1 C88859551T	T	CT	+/-
rs9450898	CNR1 C88864063T	T	CC	-/-
rs4707436	CNR1 G88851751A	A	GG	-/-
rs6454673	CNR1 G88871049A	G	GG	+/+
rs1049353	CNR1 T453T	T	CC	-/-
rs806368	CNR1 T88850100C	C	TT	-/-
rs12720071	CNR1 T88851181C	C	TT	-/-
rs806369	CNR1 T88856178C	T	TT	+/+
rs806374	CNR1 T88857320C	T	TT	+/+
rs806376	CNR1 T88858648C	C	CT	+/-
rs806377	CNR1 T88858723C	T	CT	+/-
rs6454672	CNR1 T88861570C	C	TT	-/-
rs12205430	CNR1 T88867925C	C	TT	-/-
rs6454674	CNR1 T88872930G	T	TT	+/+
rs2502993	CNR2 A282A	A	AG	+/-
rs2501431	CNR2 G155G	A	AG	+/-
rs16828926	CNR2 G24215130A	A	AG	+/-
rs2501432	CNR2 G63A	T	CT	+/-
rs2229579	CNR2 H316T	A	GG	-/-
rs4649124	CNR2 L251L	A	AG	+/-
rs9424398	CNR2 T24221834G	G	GT	+/-
rs3741775	DAO A14747C	C	AC	+/-
rs18347	DAO A24464G	G	AA	-/-
rs14098	DAO C108286399T	T	CC	-/-
rs1000000	DAO G109284478T	G	GG	+/+
rs2070586	DAO G8864A	A	GG	-/-
rs7980427	DAO S93S	A	GG	-/-
rs2070587	DAO T887G	T	TT	+/+
rs2111902	DAO T9891G	T	TT	+/+

rs2073440	HDC A1932C	G	TT	-/-
rs17740807	HDC C92T	A	GG	-/-
rs854158	HDC T10086C	G	AG	+/-
rs16063498	HDC T1657C	G	AA	-/-
rs1800708	HFE 10795T>C	C	TT	-/-
rs2071302	HFE 11622T>C	C	TT	-/-
rs2794719	HFE 6382T>G	G	GT	+/-
rs9388837	HFE 6590C>T	T	CC	-/-
rs2071303	HFE 8828T>C	C	TT	-/-
rs1800562	HFE C282Y	A	GG	-/-
rs1799945	HFE H83D	G	CC	-/-
rs1050900	HNMT A'218T	T	AA	-/-
rs1050921	HNMT A47507G	G	AA	-/-
rs1050922	HNMT C29232A	A	CC	-/-
rs8430764	HNMT C3616T	T	TT	+/+
rs1050891	HNMT T939C	G	AA	-/-
rs347591	HRH1 G11290122T	G	GT	+/-
rs2087488	HRH1 G57C	C	GG	-/-
rs7851620	HRH1 G809A	A	GG	-/-
rs348070	HRH1 T'1687C	T	CT	+/-
rs901885	HRH1 T-17C	T	CT	+/-
rs11662595	HRH4 A817G	G	AG	+/-
rs11665084	HRH4 C413T	T	CT	+/-
rs4800573	HRH4 G'2144A	A	GG	-/-
rs1421125	HRH4 G'385T	T	GG	-/-
rs16940765	HRH4 T3537649C	C	TT	-/-
rs7997012	HTR2A T84185C	G	AG	+/-

Histamine

Extracellular

Intracellular

**WHY IS HISTAMINE IMPORTANT?**



[Adv Exp Med Biol. 2010;709:95-107.](#)

## Histamine in neurotransmission and brain diseases.

[Nuutinen S<sup>1</sup>](#), [Panula P.](#)

### ⊕ Author information

#### Abstract

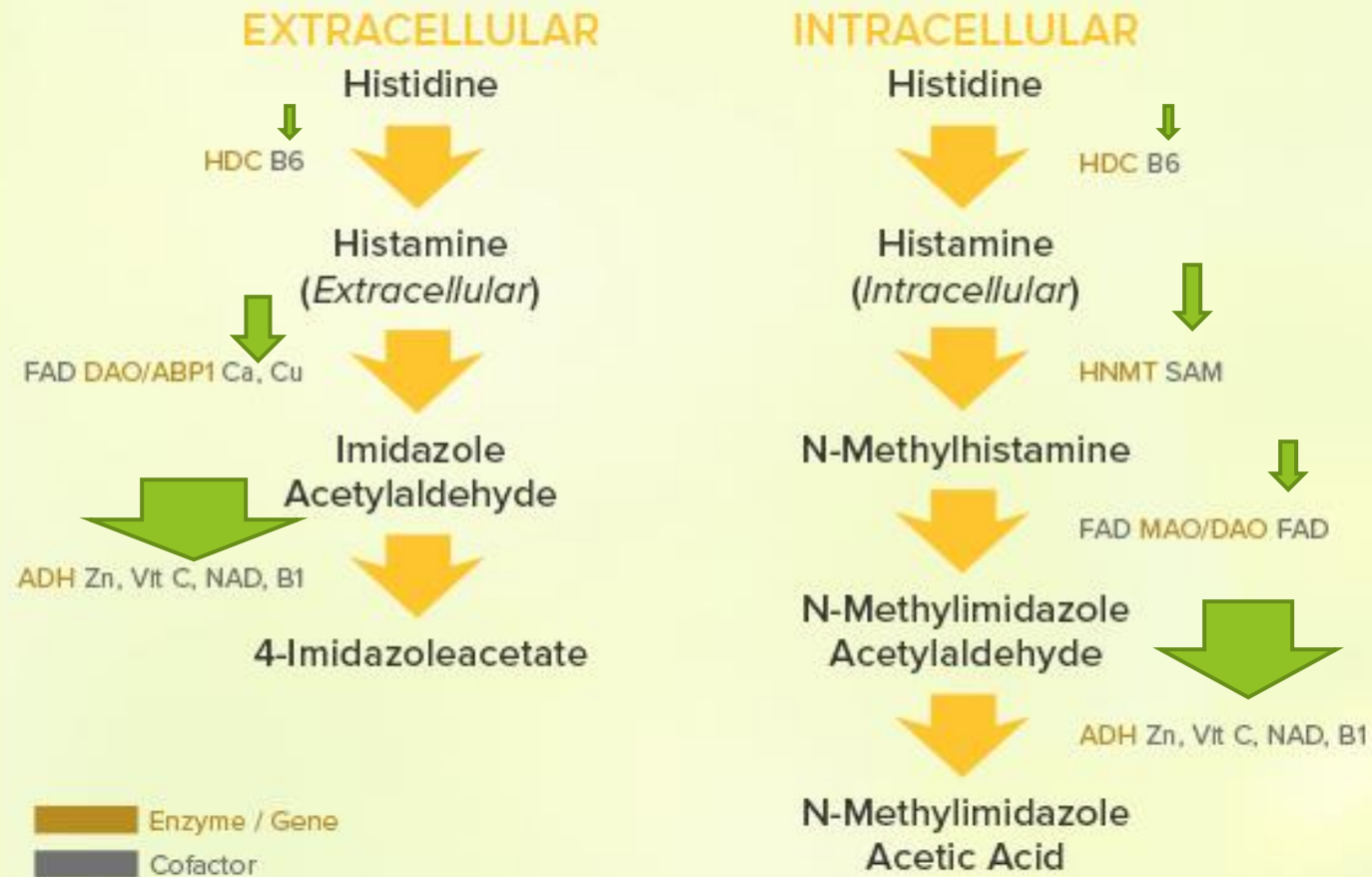
Apart from its central role in the mediation of allergic reactions, gastric acid secretion and inflammation, histamine serves an important function as a neurotransmitter in the central nervous system. The histaminergic neurons originate from the tuberomammillary nucleus of the posterior hypothalamus and send projections to most parts of the brain. The central histamine system is involved in many brain functions such as arousal, control of pituitary hormone secretion, suppression of eating and cognitive functions. The effects of neuronal histamine are mediated via G-protein-coupled H1-H4 receptors. The prominent role of histamine as a wake-promoting substance has drawn interest to treat sleep-wake disorders, especially narcolepsy, via modulation of H3 receptor function. Post mortem studies have revealed alterations in histaminergic system in neurological and psychiatric diseases. Brain histamine levels are decreased in Alzheimer's disease patients whereas abnormally high histamine concentrations are found in the brains of Parkinson's disease and schizophrenic patients. Low histamine levels are associated with convulsions and seizures. The release of histamine is altered in response to different types of brain injury: e.g. increased release of histamine in an ischemic brain trauma might have a role in the recovery from neuronal damage. Neuronal histamine is also involved in the pain perception. Drugs that increase brain and spinal histamine concentrations have antinociceptive properties. Histaminergic drugs, most importantly histamine H3 receptors ligands, have shown efficacy in many animal models of the above-mentioned disorders. Ongoing clinical trials will reveal the efficacy and safety of these drugs in the treatment of human patients.

PMID: 21618891

The central histamine system is involved in many brain functions such as arousal, control of pituitary hormone secretion, suppression of eating and cognitive functions.

The prominent role of histamine as a wake-promoting substance has drawn interest to treat sleep-wake disorders, especially narcolepsy, via modulation of H3 receptor function.

# HISTAMINE PATHWAY

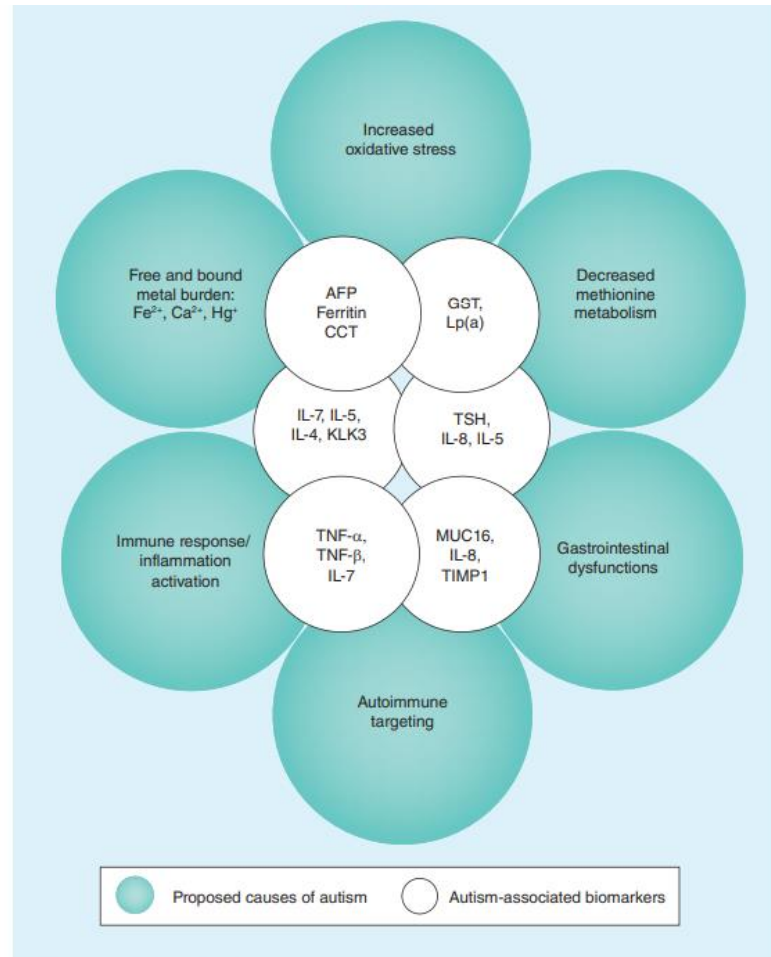




## Newborn screening for autism: in search of candidate biomarkers

**Background:** Autism spectrum disorder (ASD) represents a wide range of neurodevelopmental disorders characterized by impairments in social interaction, language, communication and range of interests. Autism is usually diagnosed in children 3–5 years of age using behavioral assessments. Newborn screening for ASD at birth would be beneficial for early initiation of treatment. **Aim:** This retrospective study sought to identify newborns at risk for ASD. **Methods:** This study utilized stored frozen specimens from ASD children and controls. The 15 candidate biomarkers were then discussed regarding their association with ASD. **Results:** Three sets of five biomarkers associated with ASD were found that differed from control groups. **Conclusion:** This study determined that a statistically selected panel of 15 biomarkers successfully discriminated presumptive newborns at risk for ASD from those of nonaffected controls.

Mizejewski, G. J., Lindau-Shepard, B., & Pass, K. A. (2013). *Newborn screening for autism: in search of candidate biomarkers*. *Biomarkers in Medicine*, 7(2), 247–260. doi:10.2217/bmm.12.108



**Figure 3. A circle model of proposed causes of autism versus screen-selected biomarkers.** A circle model of the proposed six causes of autism is shown on the outer circles, while the inner overlapping circles indicate the biomarkers associated with the causative metabolic or immune response/inflammation events. The overlap of the outer and inner circles indicates the degree of association of the two agents.

The proper use of epigenetics here is to raise your “Index of Suspicion” of pathways that would be compromised under an oxidative stress load.

The benefit of this type of research is that the epigenetics point to possible specific dysfunctions under an oxidative stress load.

These dysfunctions are identified as root causes of autism. All very true.

The knowledge of these will allow you to intervene in the presence of the pathology.

Of great benefit is to know the probabilities so that you can prevent the occurrence of pathology.

Mizejewski, G. J., Lindau-Shepard, B., & Pass, K. A. (2013). *Newborn screening for autism: in search of candidate biomarkers. Biomarkers in Medicine*, 7(2), 247–260. doi:10.2217/bmm.12.108





Is this premise correct??.....NO

## Newborn screening for autism: in search of candidate biomarkers

**Background:** Autism spectrum disorder (ASD) represents a wide range of neurodevelopmental disorders characterized by impairments in social interaction, language, communication and range of interests. Autism is usually diagnosed in children 3–5 years of age using behavioral characteristics; thus, diagnosis shortly after birth would be beneficial for early initiation of treatment. **Aim:** This retrospective study sought to identify newborns at risk for ASD utilizing bloodspot specimens in an immunoassay. **Materials & methods:** The present study utilized stored frozen specimens from ASD children already diagnosed at 15–36 months of age. The newborn specimens and controls were analyzed by immunoassay in a multiplex system that included 90 serum biomarkers and subjected to statistical analysis. **Results:** The results of the study showed that 10% of the newborn specimens were found that differed from their association with ASD. **Conclusion:** The study successfully discriminated presu

**“diagnosis** shortly after birth would be beneficial for early initiation of treatment”

**Presumption is that *you are born* with Autism. Although this entire paper refutes that premise, it is a commonly held belief.**

Mizejewski, G. J., Lindau-Shepard, B., & Pass, K. A. (2013). *Newborn screening for autism: in search of candidate biomarkers*. *Biomarkers in Medicine*, 7(2), 247–260. doi:10.2217/bmm.12.108



# GLITCH: The Definition of “DIAGNOSIS”

“investigation or analysis of the cause or nature of a condition, situation, or problem”

\*

## SYMPTOM(S)

- ▶ Sore throat
- ▶ IBS
- ▶ “Hyperacidity”
- ▶ Effects of root causes

## DIAGNOSIS

- ▶ Strep Throat
- ▶ Celiac Disease
- ▶ H.Pylori induced gastritis
- ▶ Root Causes

A descriptive "diagnosis" of the syndrome often leads to suboptimal clinical results as the investigation into root causes ceases.

"Diagnosing" an infant at birth with autism based on genetic polymorphisms may have the *counterproductive effect of labeling, the feeling that nothing can be done, and the parents may be dissuaded from making healthy choices for the child that would prevent Autism.*

*“Newborn screening for autism: in search of candidate biomarkers”\**  
The Learning Points.

The authors prove, on an acceptable scientific basis:

- That autism has root causes.
- There may be predictive power in determining the genetic SNPs.
- In so doing, within their pathways, the SNPs provide a map for the clinician in either treatment or prevention.



# What have we learned?



Your genes are not your destiny



Proper knowledge of SSNPs will raise your index of suspicion for certain types of pathology. Knowledge of the necessities of the biochemical pathways give you actionable treatment possibilities.



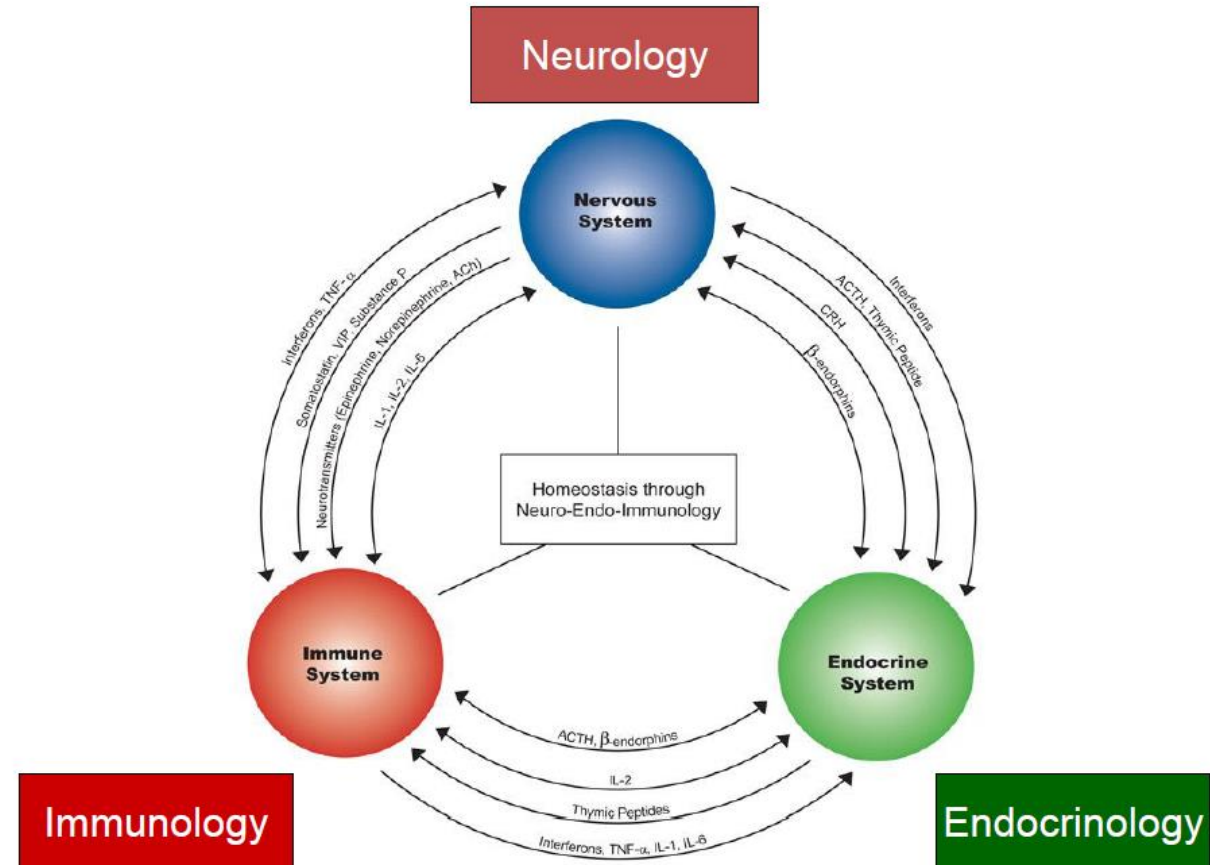
It's simply one set of data.

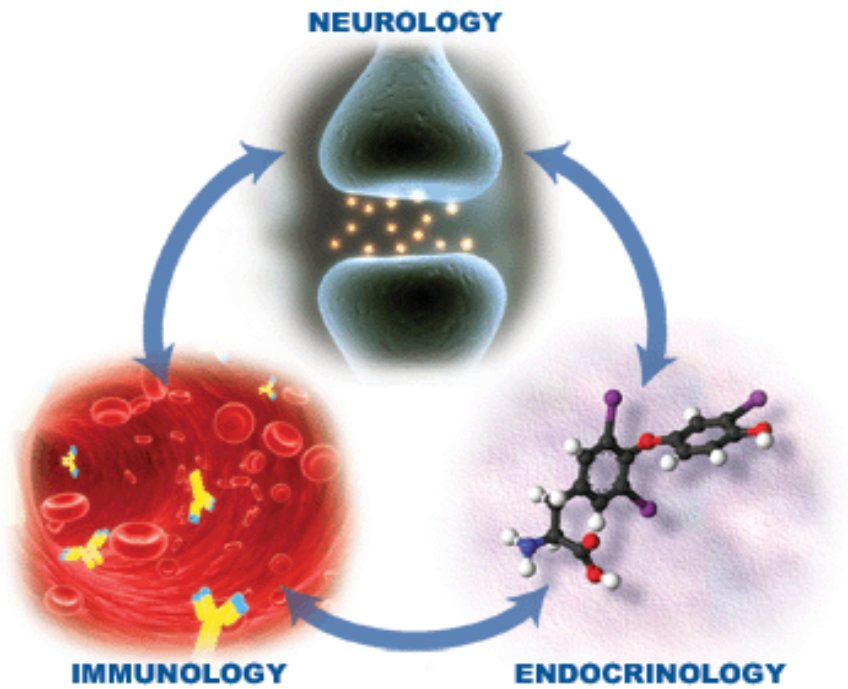
# Neuro-Endo-Immunology

## The “NEI Supersystem”

### Shift from Linear to Integrated Medicine

Not separate systems





## Each System has Unique Biomarkers

**The Neurological System's Biomarkers:**  
Neurotransmitters (Serotonin, GABA, Glutamate, Dopamine, Epinephrine, Norepinephrine, PEA)

**Endocrinology (glands, etc.) biomarkers:**

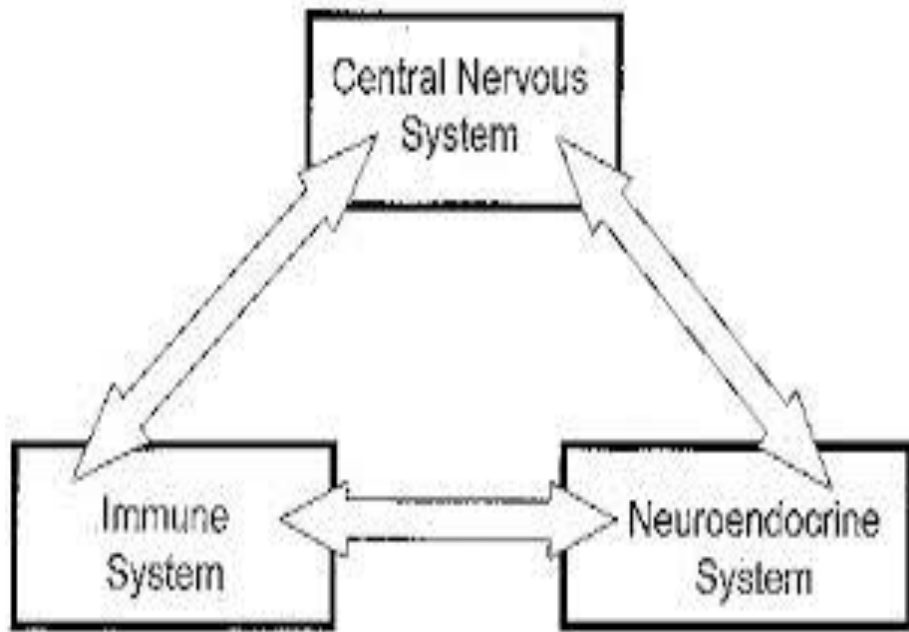
Hormones (estrogen, progesterone, insulin, thyroxine, etc.)

**The Immune system's biomarkers:**

Cytokines (IL-12, TNF- $\alpha$ , TGF- $\beta$ , IL-4, IFN- $\gamma$ )

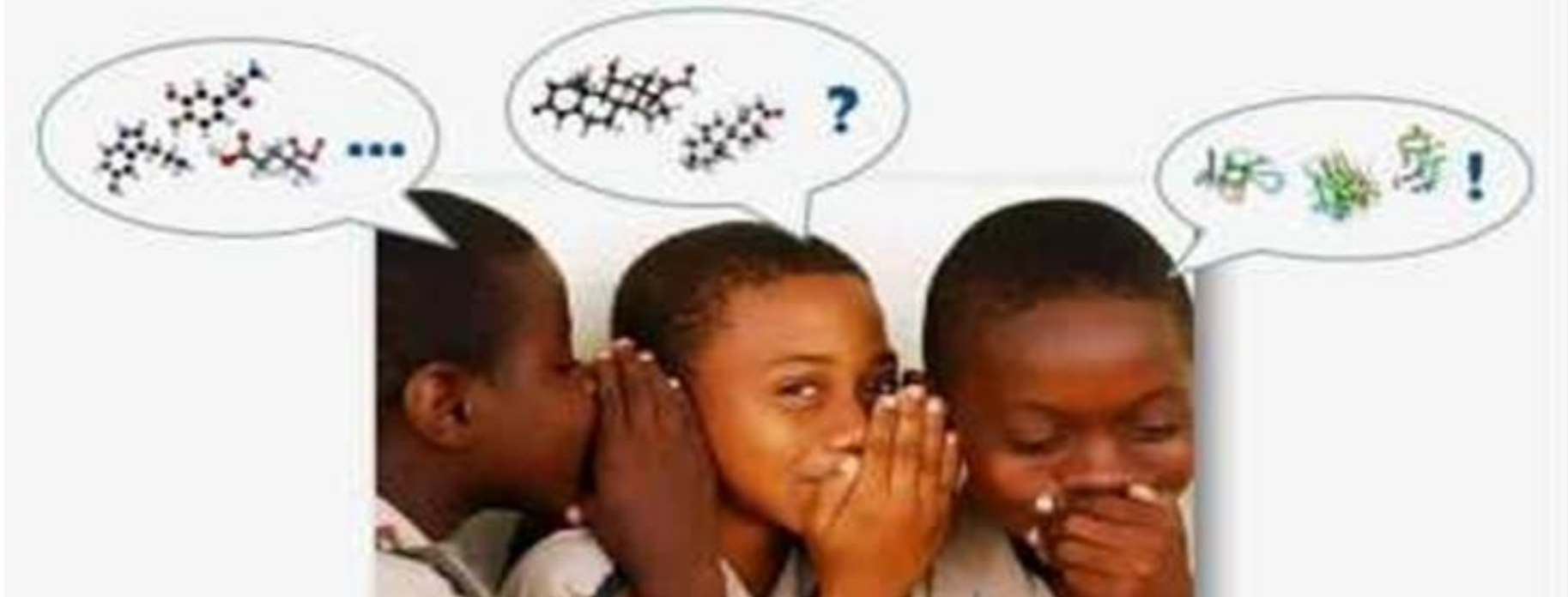


# All systems have receptors for each others' systems



- Neuro system has receptors for cytokines and hormones
- Endo system has receptors for neurotransmitters and cytokine
- Immune system has receptors for neurotransmitters and hormones

The Neuro, Endo, and Immune systems are in constant conversation



# Neuroendocrine Interactions in the Immune System

[Dennis D. Taub](#)<sup>†</sup>

► [Author information](#) ► [Copyright and License information](#) [Disclaimer](#)

The publisher's final edited version of this article is available at [Cell Immunol](#)

See other articles in PMC that [cite](#) the published article.

## Abstract

Substantial evidence now exists supporting the bidirectional communication between the neuroendocrine and immune systems. A number of hormonal and neuropeptide mediators have been shown to influence immune development and function in healthy, aged and diseased individuals. Immune receptors for many of these ligands and similarly, receptors for cytokines and growth factors have been identified on cells within the central nervous and endocrine systems. During times of stress or injury, each of these systems come into play and transmits messages to one another. The lines of communication between the immune system and these various neuronal and endocrine organ systems constitute specific axes of interactions, which have been shown to have a profound impact on immune function, disease development and susceptibility to infections and disease. In this Special Issue, experts in neuroendocrine immunology have provided comprehensive reviews on the current advances in this area of research as well as commentary on relevance of the various axes in controlling immunity and disease development.

**Keywords:** Neuroendocrine, Immunity, Hormones, Neuropeptides, Hypothalamic-Pituitary-Adrenal (HPA), Stress, Thymus, Sympathetic Nervous System

*Substantial evidence now exists supporting the bidirectional communication between the neuroendocrine and immune systems.*

*Taub DD. Neuroendocrine interactions in the immune system. Cell Immunol. 2008;252(1-2):1–6. doi:10.1016/j.cellimm.2008.05.006*

Format: Abstract ▾

*Adv Exp Med Biol.* 2017;996:123-134. doi: 10.1007/978-3-319-56017-5\_11.

## **Psycho-Neuro-Endocrine-Immunology: A Psychobiological Concept.**

França K<sup>1,2</sup>, Lotti TM<sup>3</sup>.

### **Author information**

- 1 Centro Studi per la Ricerca Multidisciplinare e Rigenerativa, Università Degli Studi "G. Marconi", Rome, Italy.
- 2 Institute for Bioethics & Health Policy; Department of Dermatology & Cutaneous Surgery; Department of Psychiatry & Behavioral Sciences, University of Miami Miller School of Medicine, Miami, FL, USA.
- 3 Centro Studi per la Ricerca Multidisciplinare e Rigenerativa, Università Degli Studi "G. Marconi", Rome, Italy. professor@torellolotti.it.

### **Abstract**

Psycho-Neuro-Endocrine-Immunology (P.N.E.I.) is a scientific field of study that investigates the link between bidirectional communications among the nervous system, the endocrine system, and the immune system and the correlations of this cross-talk with physical health. The P.N.E.I. innovative medical approach represents a paradigm shift from a strictly biomedical view of health and disease taken as hermetically sealed compartments to a more interdisciplinary one. The key element of P.N.E.I. approach is represented by the concept of bidirectional cross-talk between the psychoneuroendocrine and immune systems. The Low Dose Medicine is one of the most promising approaches able to allow the researchers to design innovative therapeutic strategies for the treatment of skin diseases based on the rebalance of the immune response.

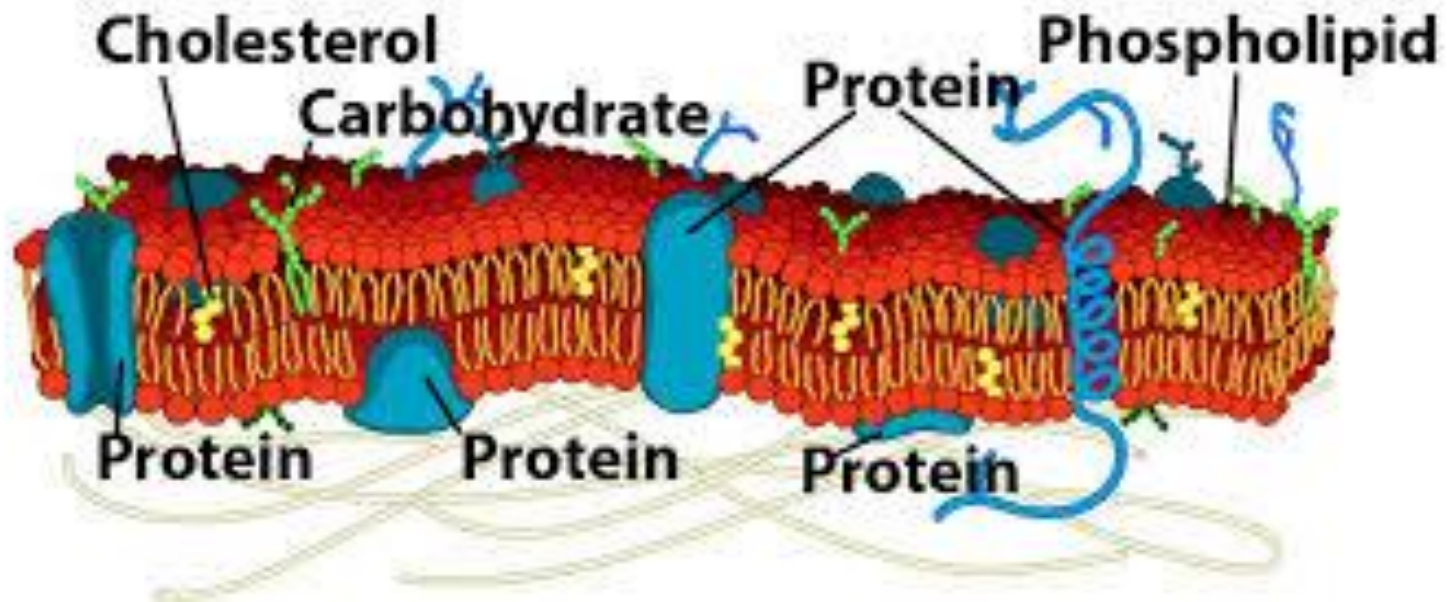
Psycho-Neuro-Endocrine-Immunology (P.N.E.I.) is a scientific field of study that investigates the link between bidirectional communications among the nervous system, the endocrine system, and the immune system and the correlations of this cross-talk with physical health.

<https://www.ncbi.nlm.nih.gov/pubmed/29124696>

# What have we learned about the NEI supersystem

- ▶ We have learned that the neurological, endocrine, and immune systems constantly communicate with one another.
- ▶ The mechanism of this is the fact that each system has the receptors for the other systems' biomarkers.
- ▶ When one system is compromised it will affect the other systems.
- ▶ The NEI super system removes diagnosis and treatment from a single discipline to a more multidisciplinary approach. Dare I say, a more holistic approach?





## Cell Membrane Integrity

Important and often overlooked!

*Source: sciencemusicvideos.com*

# The “Master” of the Cell Physiology The Cell Membrane

The cell membrane is selectively permeable protecting homeostasis.

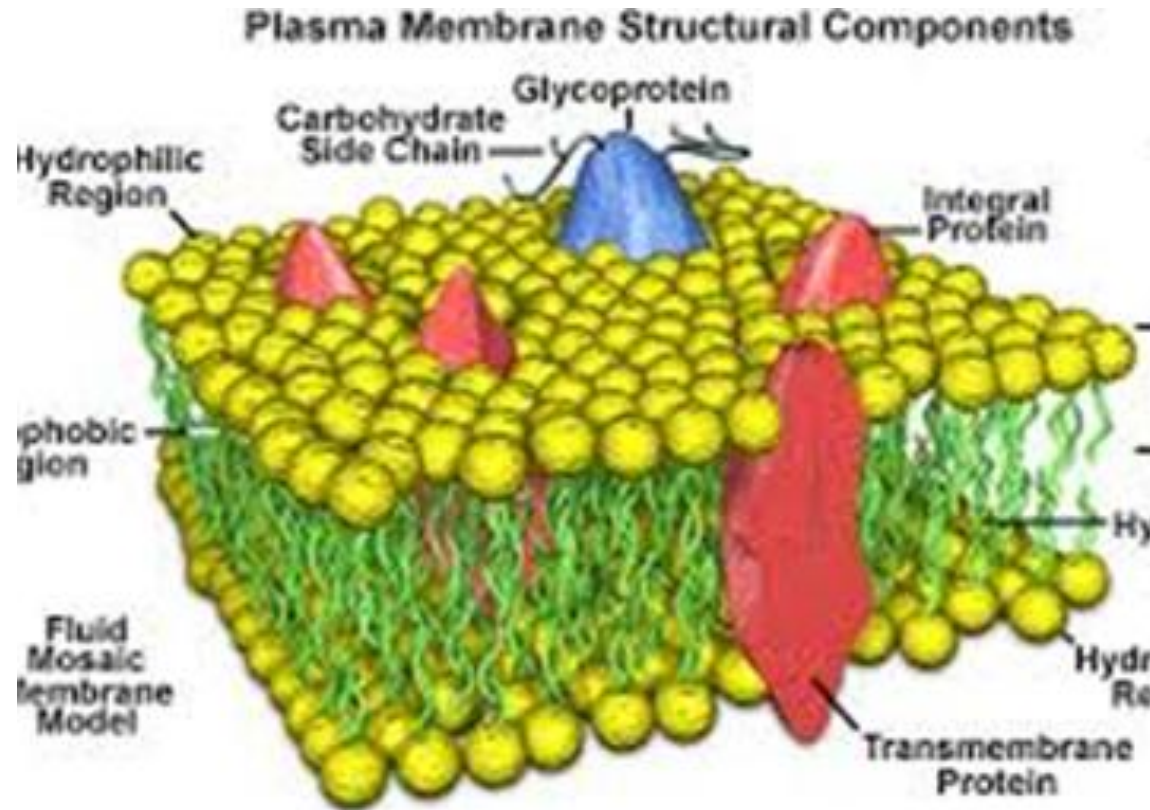
Allows for the sending of messages throughout the body (depolarization/repolarization of neural impulses)

Contains integral proteins, transmembrane proteins (acting as pumps or channels)

Contains receptors that provide for the signaling of various biological functions.

Plays a key role in the immune process.

# “Leaky” Cell Membranes

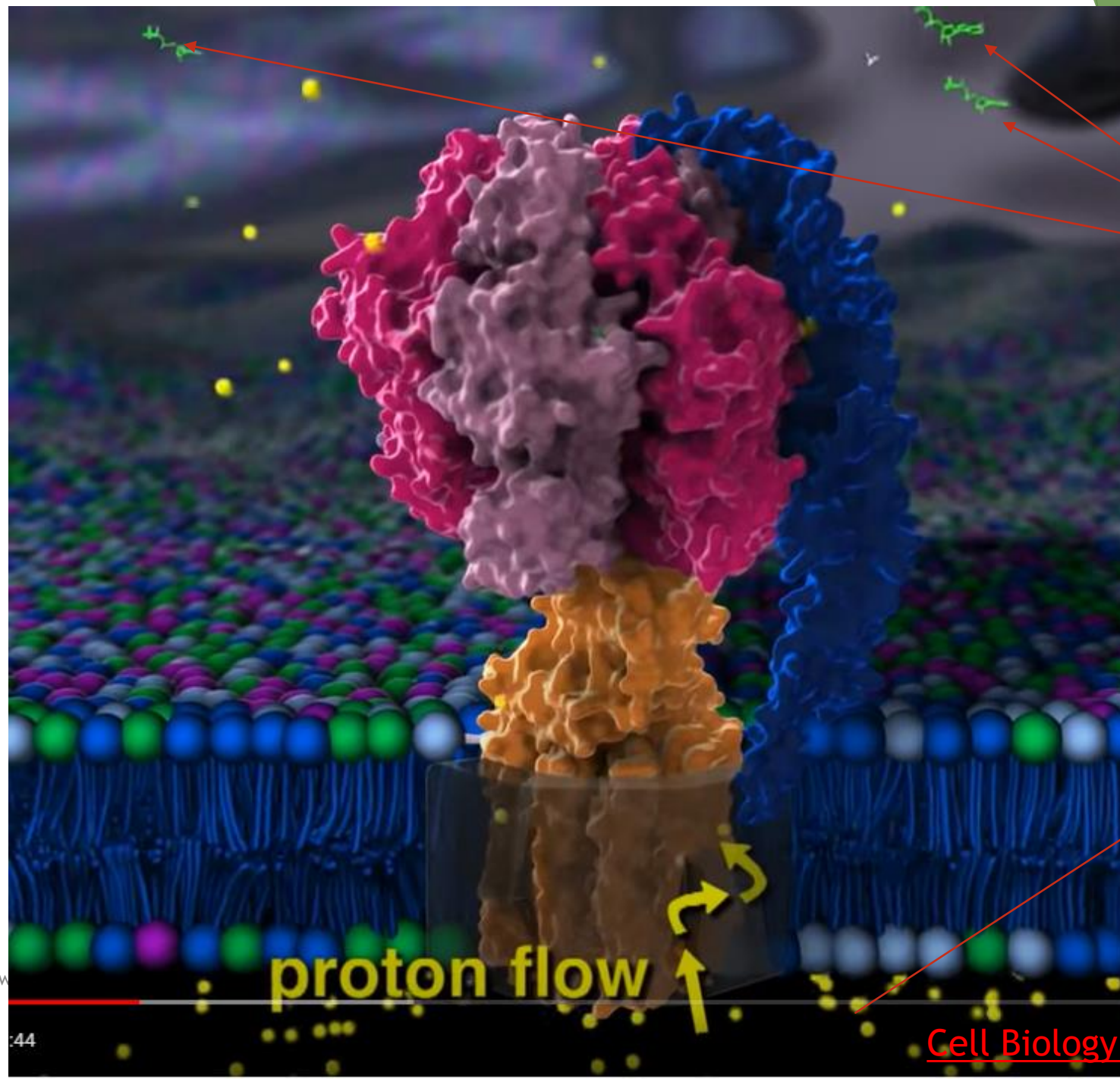


- ▶ Interrupts all physiological processes including:
- ▶ Neural transmission, integral and transmembrane protein function.
- ▶ Most importantly, interferes with the immune system's ability to differentiate between self and non-self.



# ATP Synthase Complex V

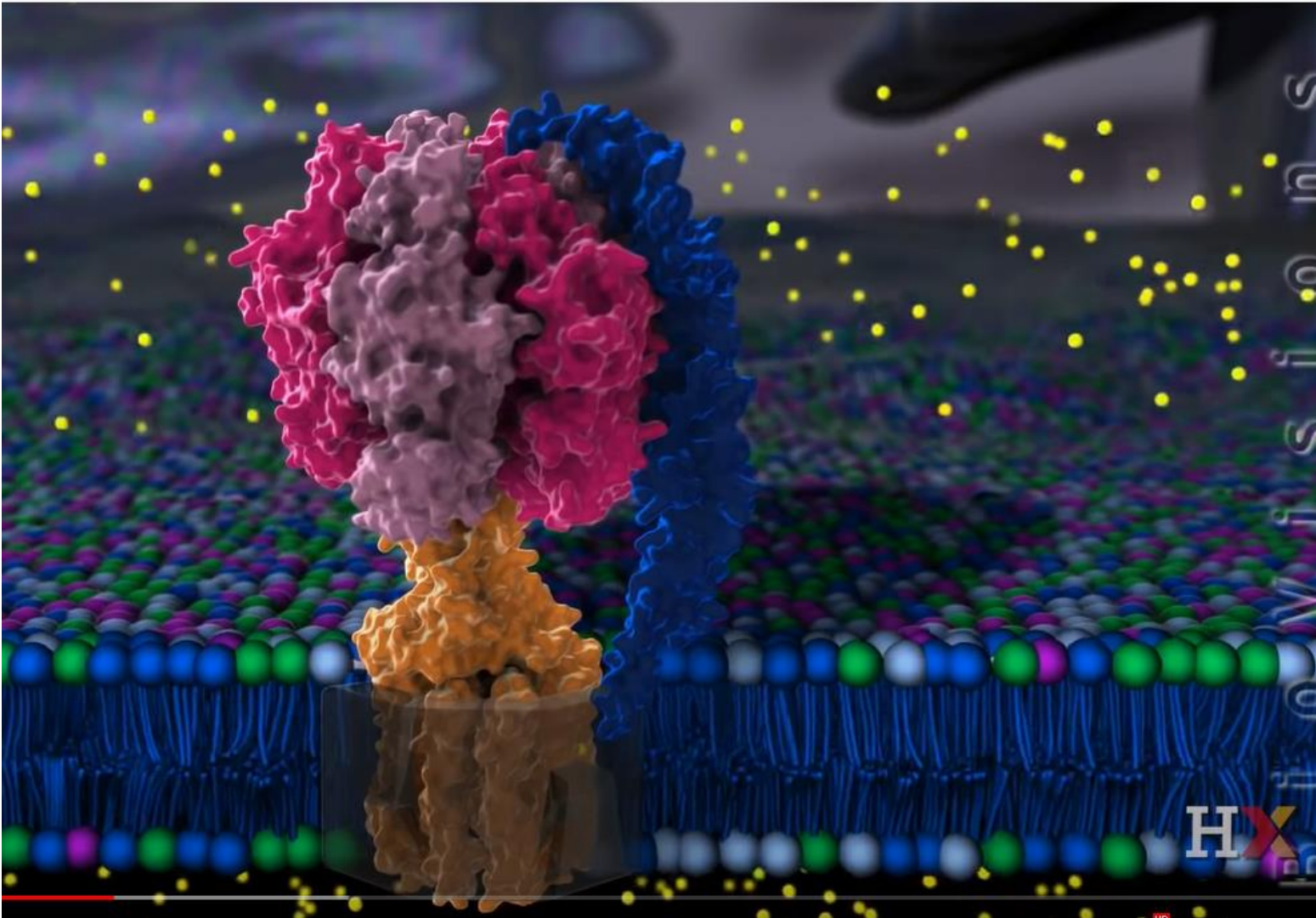
Patent Cell  
Membrane



ATP  
(energy)

High Energy  
Protons

proton flow



Leaky Membrane

Protons Leak  
Through

Energy  
production  
stops and cell  
death occurs

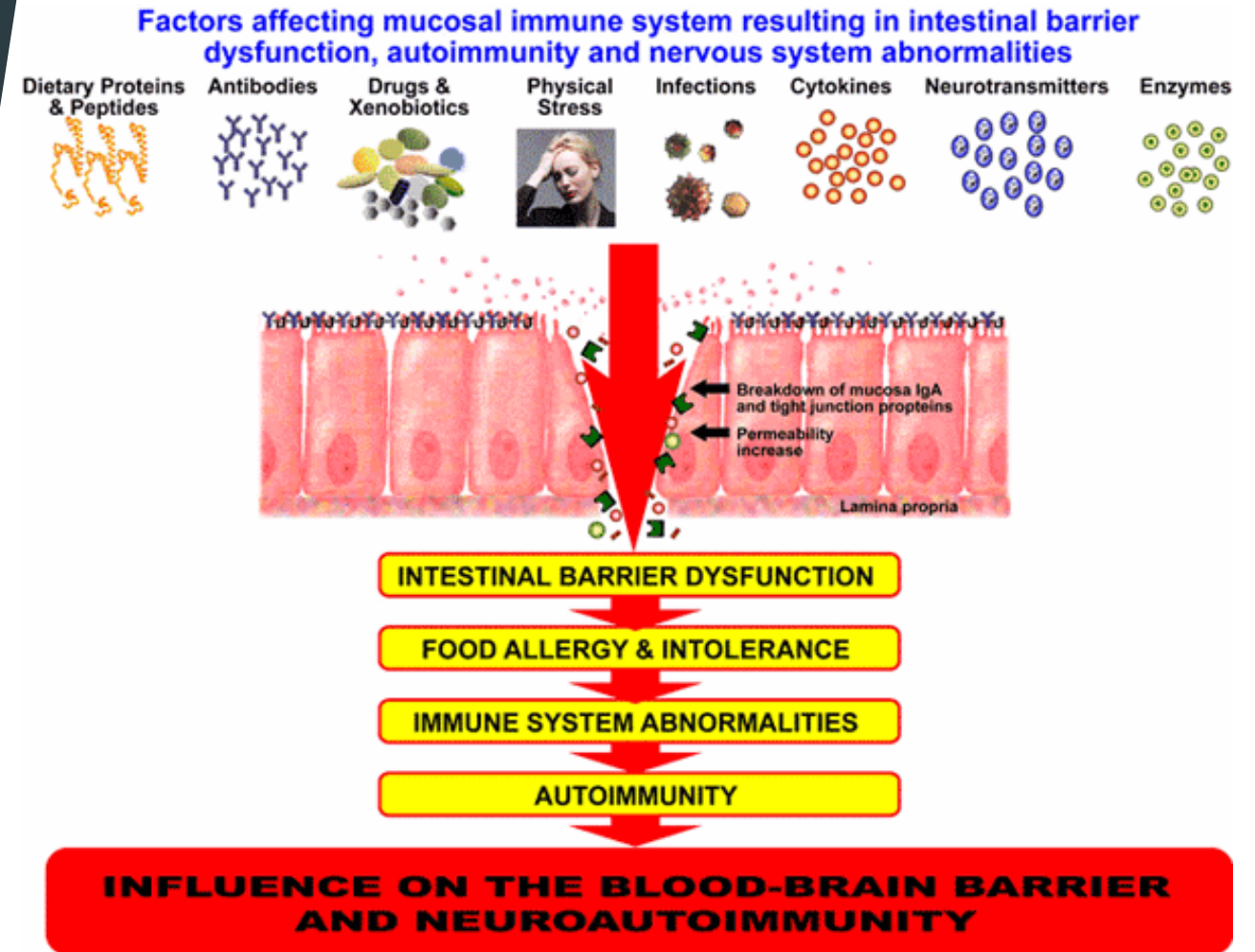
Cell Biology: Mitochondria | edX



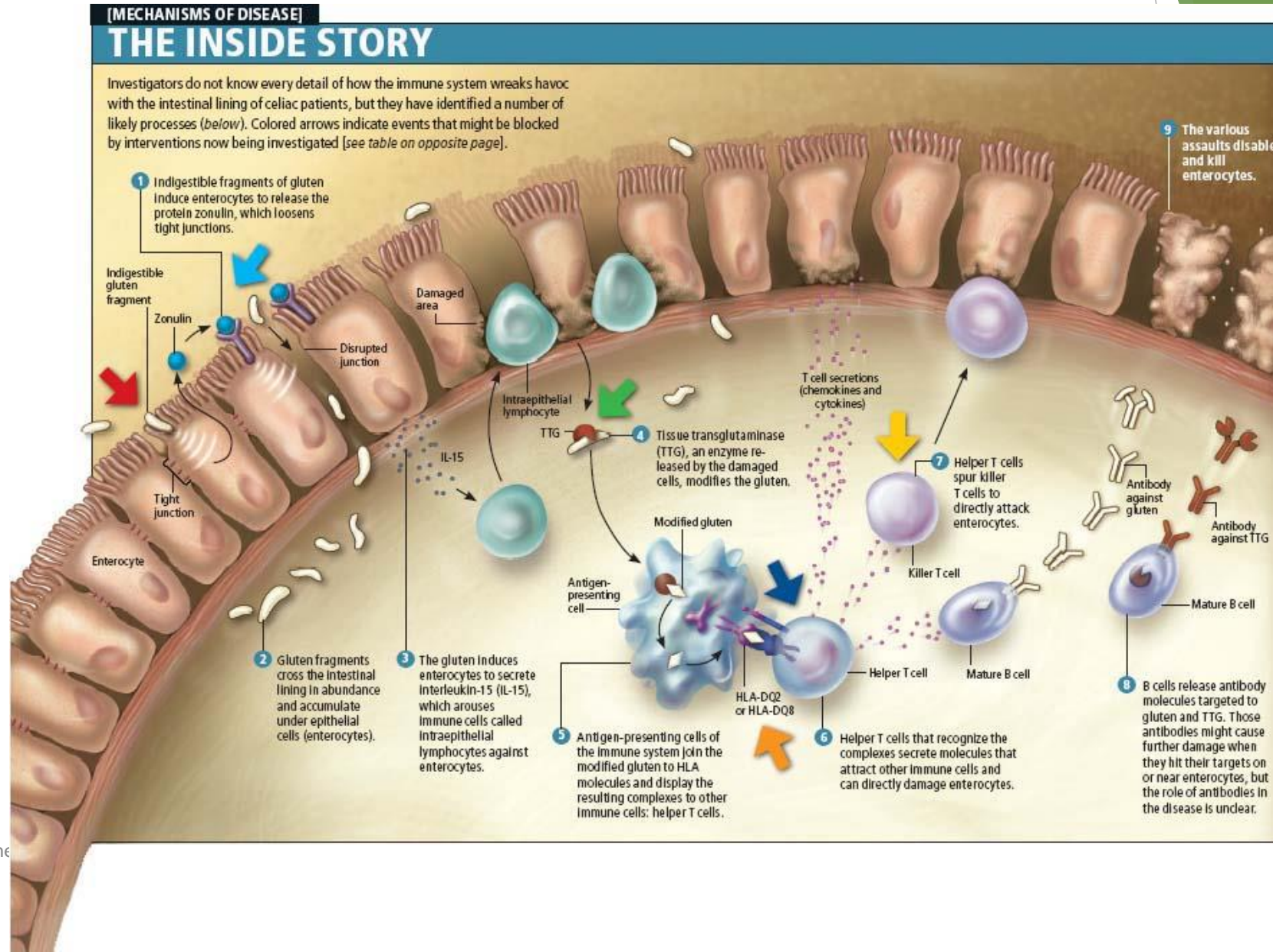
## ► Leaky Gut Syndrome

- The most common causation of chronic inflammation
- AND
- The easiest to fix.

► <http://www.glutenfreesociety.org/gluten-free-society-blog/leaky-gut-syndrome-is-gluten-at-the-root>



# For Those who like things complex 😊





## Gut–brain axis: how the microbiome influences anxiety and depression

Jane A. Foster , Karen-Anne McVey Neufeld

 Show more

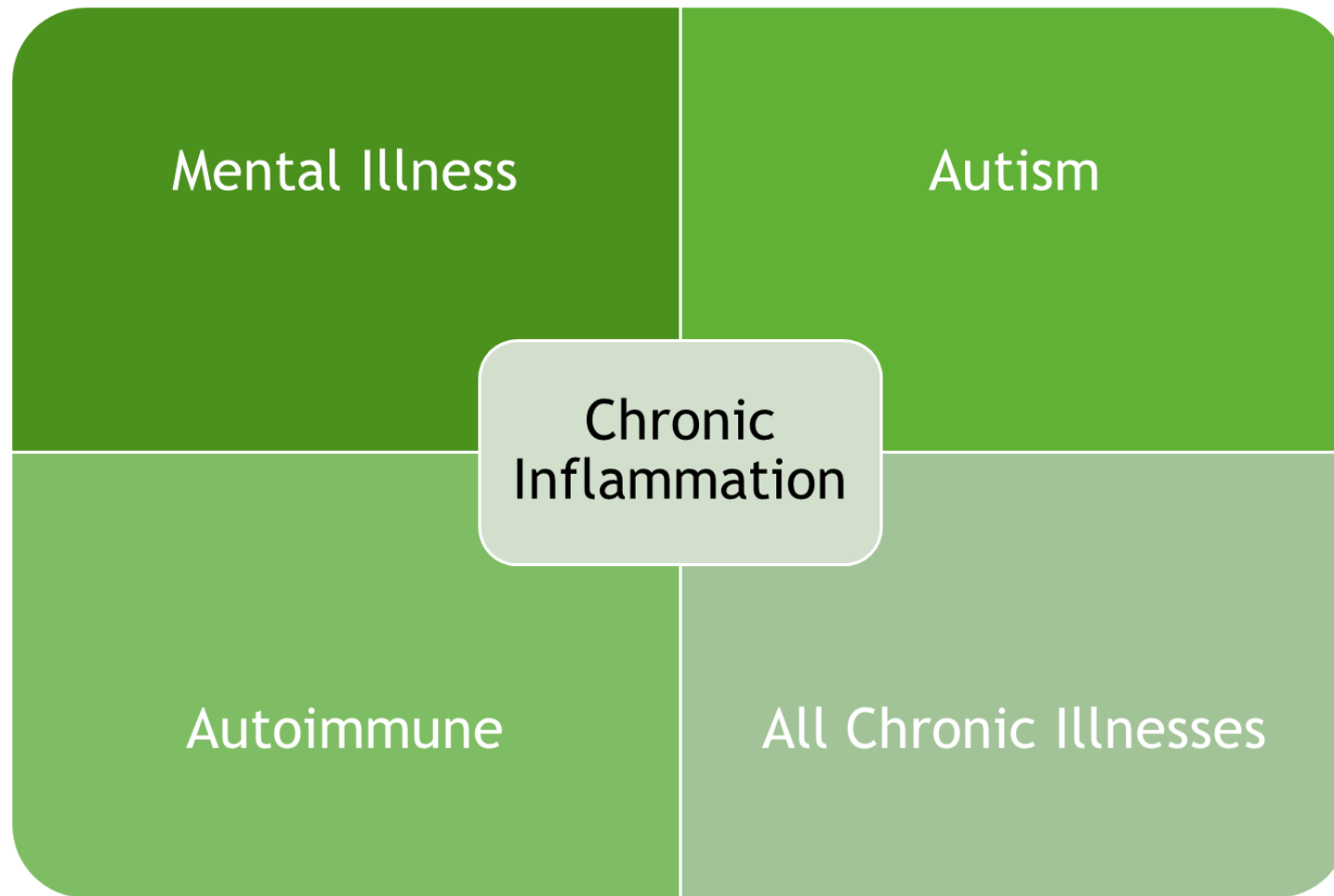
<https://doi.org/10.1016/j.tins.2013.01.005>

[Get rights and content](#)

Within the first few days of life, humans are colonized by commensal intestinal microbiota. Here, we review recent findings showing that microbiota are important in normal healthy brain function. We also discuss the relation between stress and microbiota, and how alterations in microbiota influence stress-related behaviors. New studies show that

New studies show that bacteria, including commensal, probiotic, and pathogenic bacteria, in the gastrointestinal (GI) tract can activate neural pathways and central nervous system (CNS) signaling systems. Pathogenic bacteria, in particular, can activate neural pathways and central nervous system (CNS) signaling systems, leading to anxiety and depression. Future animal and human studies on the microbiota–gut–brain axis may lead to the development of new treatments for mental health disorders.

Jane A. Foster, Karen-Anne McVey Neufeld, Gut-brain axis: how the microbiome influences anxiety and depression, *Trends in Neurosciences*, Volume 36, Issue 5, 2013, Pages 305–312, ISSN 0166-2236,

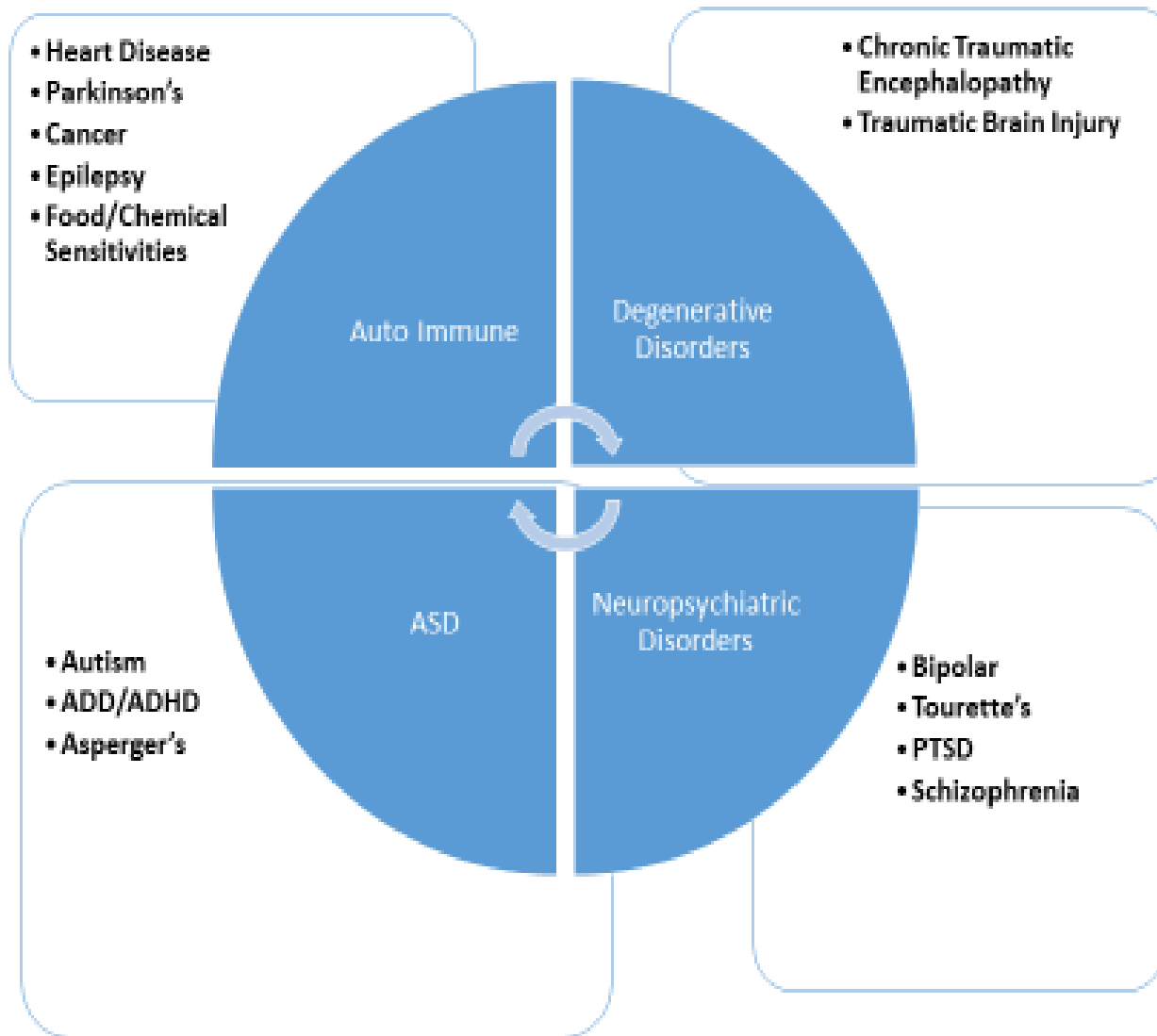


Chronic Inflammation and CIRS (Chronic Inflammatory Response Syndrome) are equivalents

*Source: "Leaky Gut, Leaky Cells, Leaky Brain" Used with permission of the authors*



## Reference for the last slide



Source:  
Robert K. Naviaux,  
*Metabolic features of the cell danger response*,  
*Mitochondrion*,  
Volume 16,  
2014,  
Pages 7-17,  
ISSN 1567-7249,  
<https://doi.org/10.1016/j.mito.2013.08.006>.



# How to fix a Leaky Gut

## *Principles:*

We need to fully digest foods. Undigested foods make up most of the antigens that enter our bodies. (*Digestive Enzymes*)

We need to re-create the mucus layer in the gut. The mucus layer is where the microbiome lives, what they eat, and where they do their work. The mucus layer traps antigens, toxins, xenobiotics and forms the initial layer of protection. (*FOS, GOS, XOS, HOS*)

We need to repair the cells and the tight junctions. This area is our second layer of defense preventing the entry of the above mentioned into our bodies. (*Butyrate, Zn Carnosine, SBI*)

We need to re-populate the GI tract with an adequate diversity of biota. (*Probiotics*)

# Miraculous (Almost) Leaky Gut Mucosal Butter

- ▶ **Leaky Gut Mucosal Butter** is a simple self-blended product that is showing rapid improvement in cellular repair. It is safe, easy to source, and easy to make.
- ▶ Who should use this:
  - ▶ Those who want to resolve their cell wall integrity issues that have led to chronic inflammatory symptoms like:
    - ▶ Autoimmune diseases
    - ▶ ASD
    - ▶ Multiple and Extensive Food Intolerances
    - ▶ ME/CFS
    - ▶ Fibromyalgia
    - ▶ Almost any inflammatory condition

# Leaky Gut Mucosal Butter Ingredients

- ▶ Organic Extra Virgin Olive Oil ½ cup
- ▶ Organic Salted Butter ¼ cup (if dairy sensitive use ¼ cup coconut, almond, or hempseed oil instead)
- ▶ Unpasteurized Natural Honey: 2 Tablespoons
- ▶ Probiotics 5 capsules
- ▶ Zinc L Carnosine 4 capsules (Seeking Health, Pure Encapsulations)
- ▶ Sialex (Ecological Formulas): 4 capsules
- ▶ SunButyrate TG (Pure Encapsulations): 1 oz.
- ▶ OPTIONAL: L Glutamine powder: 4 scoops.
- ▶ **FOR MANY OF MY PATIENTS, INCREASED GLUTAMINE = INCREASED GLUTAMATE AND NEURAL EXCITATION. USE WITH CAUTION IF YOU SUFFER FROM ANXIETY, OCD, OR ANY OF THE OTHER EXCITATORY DISORDERS. IT'S OK TO OMIT THIS IF YOU ARE UNSURE.**

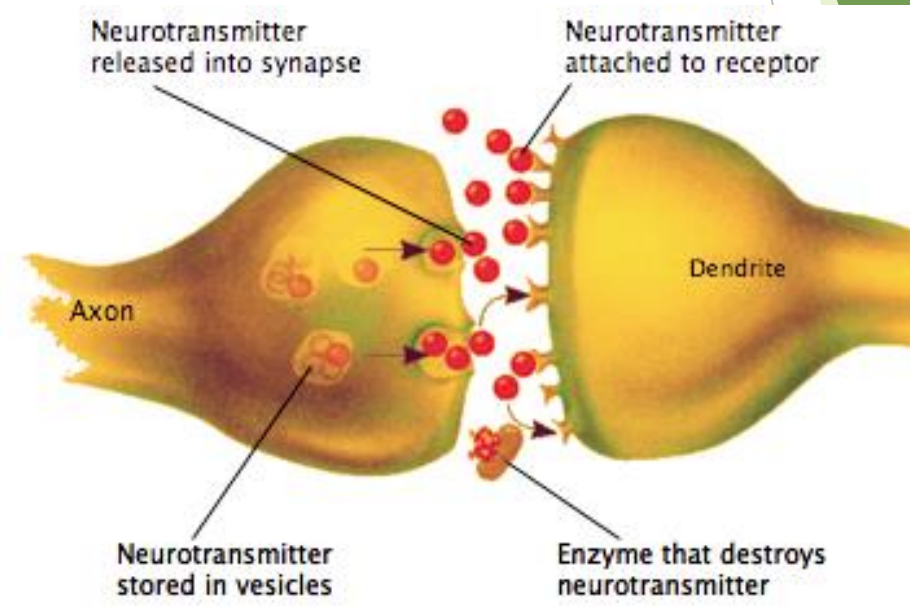
- ▶ INSTRUCTIONS:
- ▶ Open the capsules and pour out the powder into the blender. Add the oil, butter, and honey.
- ▶ Blend on high for approximately two minutes or until the substance is smooth. Then refrigerate. By morning will be able to use the “butter”.
- ▶ NOTE: You can take the butter “straight” or put on any food that you desire (toast, etc.)

Add Serum Derived Bovine Immunoglobulin Isolate\* (6 caps) when there is low SIgA

\*Hammad Liaquat, Munish Ashat, Abigail Stocker, Lindsay McElmurray, Karen Beatty, Thomas L. Abell, Gerald Dryden, Clinical Efficacy of Serum-Derived Bovine Immunoglobulin in Patients With Refractory Inflammatory Bowel Disease, The American Journal of the Medical Sciences, Volume 356, Issue 6, 2018, <https://doi.org/10.1016/j.amjms.2018.08.019>.

# Neurotransmitters

- ▶ *Neurotransmitters*, also known as chemical messengers, are endogenous chemicals that enable neurotransmission.
- ▶ They transmit signals across a chemical synapse, such as a neuromuscular junction, from one neuron (nerve cell) to another "target" neuron, muscle cell, or gland cell.



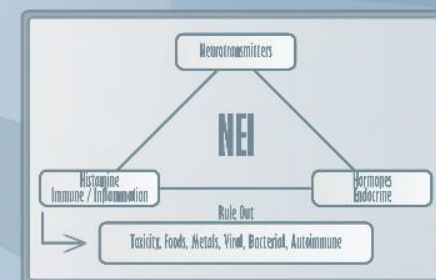
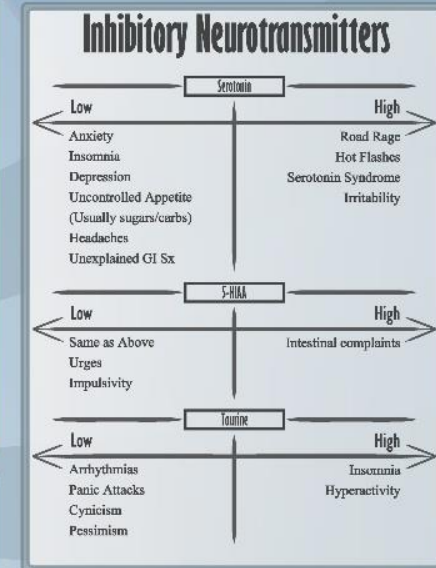
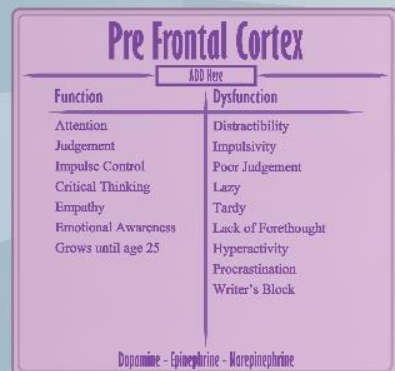
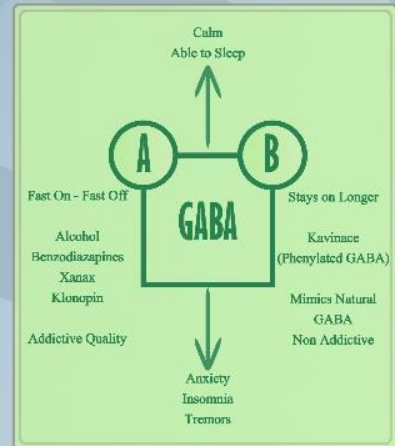
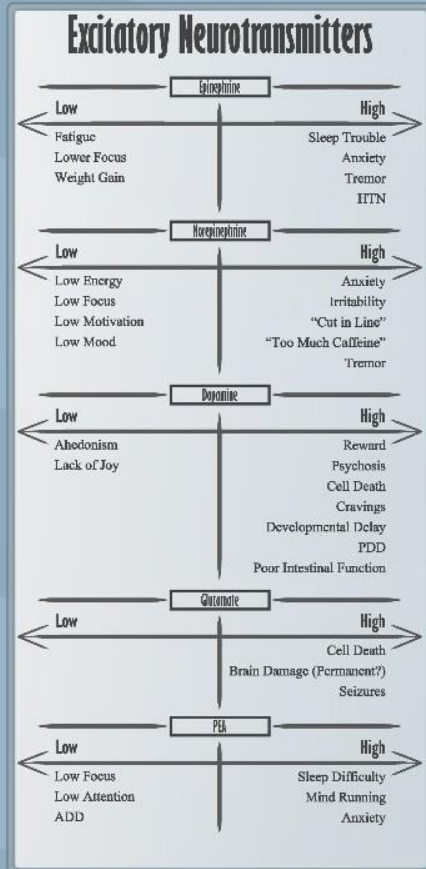


# Excitatory

vs.

# Inhibitory

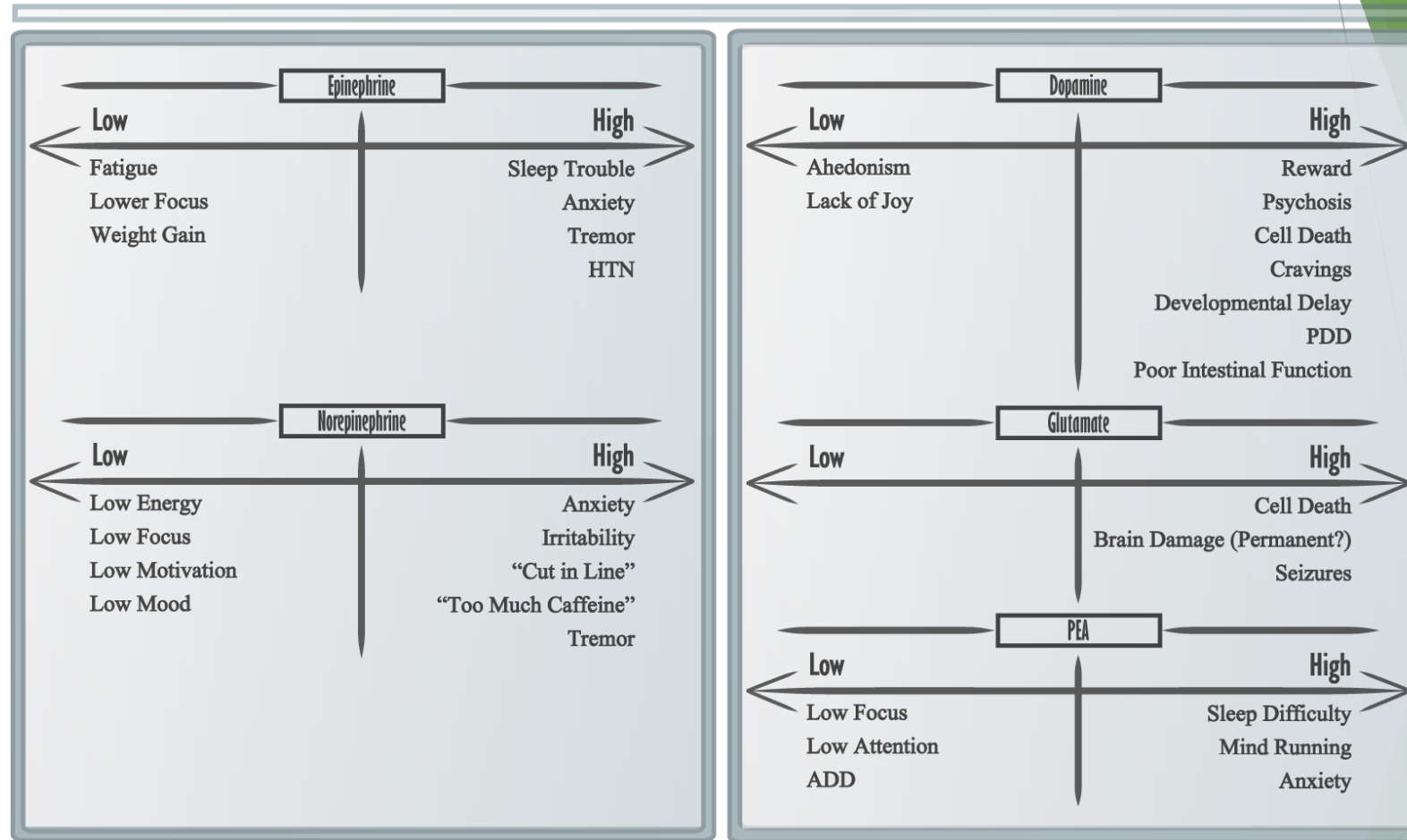




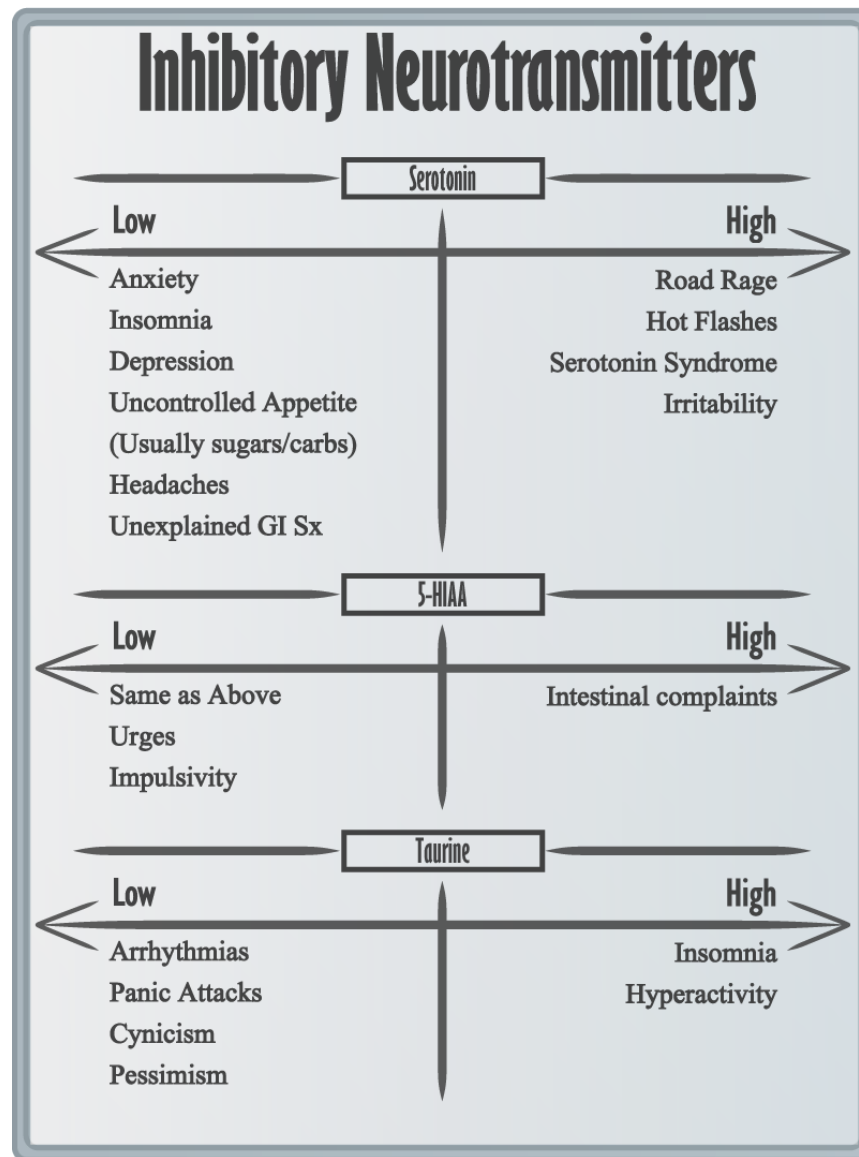
# BRAIN WALL



# Excitatory Neurotransmitters



*Think of these as the substances that keep you awake and give you energy*



***Think of these as the substances that “calm” the nervous system***

# Case Studies

# Case Study....Alyssa

*8 Year Old Female with visual distortions. Mom initially contacted presenter with the possible need for Irlen Glasses due to visual distortions.*

*Also c/o “bad gut”. Pain upon eating gluten, soy or almost anything else.*

*After questioning, Hallucinations (Auditory, Olfactory & Visual) were identified.*

*Advised mom to obtain a standard work up for 2 basic reasons:*

- Sometimes there are conditions that are easily corrected or are better treated by a different specialist. And...*
- Olfactory hallucinations are secondary to a brain tumor, unless proven otherwise*



# Standard Medical Work Up

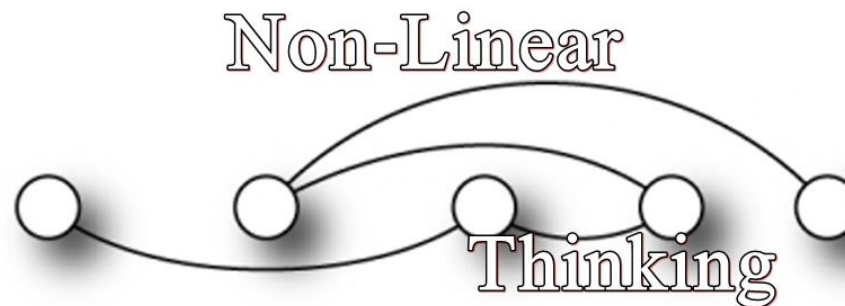
## *Standard work-up*

- ▶ *Brain CT*
- ▶ *MRI*
- ▶ *labs for thyroid, CBC, Complete Medical Profile, etc.*
- ▶ *Mom was instructed to return to me if the tests were negative for pathology or signs of obvious illness.*
- ▶ *In other words, if she was to be placed on anti-psychotics, let me help.*

## *Results:*

- ▶ *CT of the Brain-negative for pathology.*
- ▶ *MRI of the brain-negative for pathology*
- ▶ *Entire laboratory analysis within reference ranges(A.K.A. -Normal)*
- ▶ *The only treatment options offered were progressive use of psychotropic agents leading to atypical antipsychotic medications.*
- ▶ *Outlook: GUARDED No expectation of a normal life.*

# What Now? There are So Many Possibilities... We Need Direction



# Epigenetics Can Help Point The Way

Notice I said, *Help Point the Way*

Most effective use of genetic information

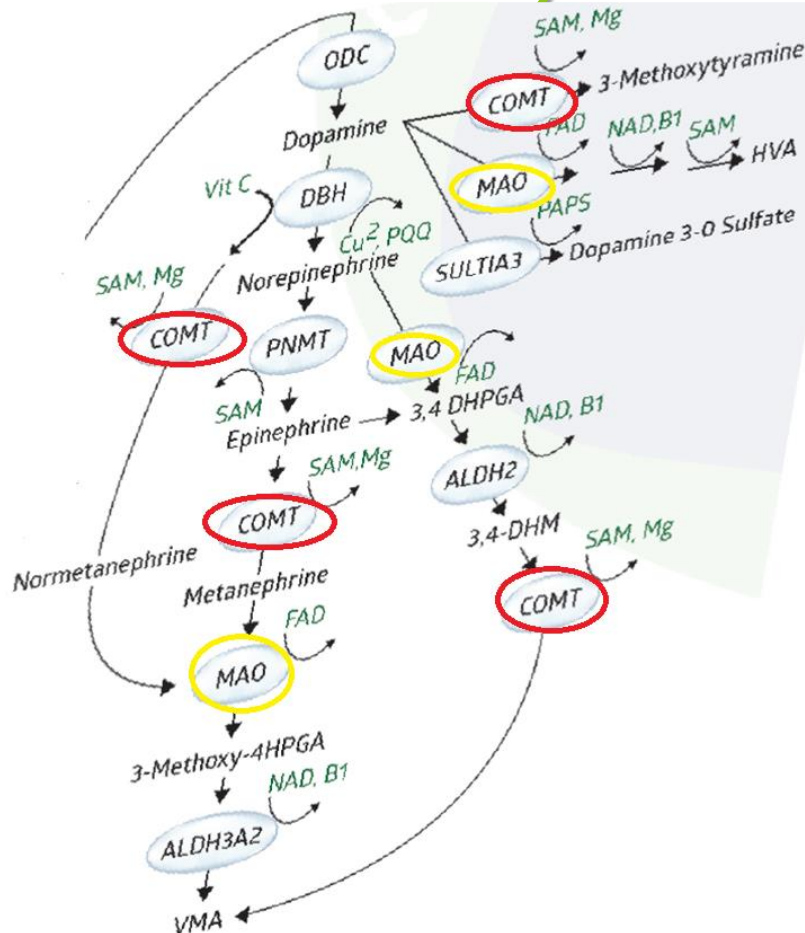
- ▶ To raise your index of suspicion of pathophysiology in certain areas of your patient's physiology
- ▶ If you know the pathways and there are a significant level of polymorphisms, then...
- ▶ That pathway(s) may not function well under oxidative stress

Say Ye..."But I have to learn all those genes and snps....there are a MILLION of them!"

- ▶ Yea, yea so you say...but maybe we can focus our analysis (so we don't end up in paralysis, or worse)?

# EXCITATION CAN CAUSE THESE SYMPTOMS, WHICH SNPS ARE IMPORTANT TO CONSIDER?

## COMT, MAO



COMT	<a href="#">rs6269</a>	G	AA	-/-
COMT -61 P199P	<a href="#">rs769224</a>	A	AG	+/-
COMT H62H	<a href="#">rs4633</a>	T	TT	+/+
MAO A R297R	<a href="#">rs6323</a>	T	GT	+/-

SNPs slow down the metabolism (drainage) of catecholamines and eventually, they will “overflow”



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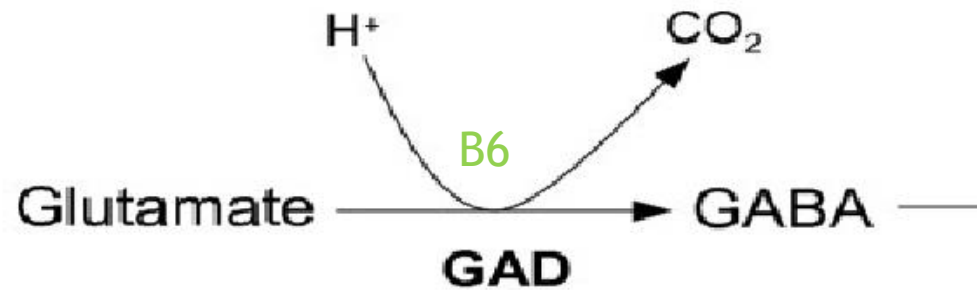
Papaleo, Francesco et al. “Genetic Dissection of the Role of Catechol-O-Methyltransferase in Cognition and Stress Reactivity in Mice.” The Journal of neuroscience : the official journal of the Society for Neuroscience 28.35 (2008): 8709-8723. PMC. Web. 30 July 2015.

Simpson, Eleanor H. et al. “Genetic Variation in COMT Activity Impacts Learning and Dopamine Release Capacity in the Striatum.” Learning & Memory 21.4 (2014): 205-214. PMC. Web. 30 July 2015.

# INCREASED GLUTAMATE CAN CAUSE EXCITATION

## What SNPs can cause that?

### GAD



*GAD1 (At5g17330)*  
*GAD2 (At1g65960)*  
*GAD3 (At2g02000)*  
*GAD4 (At2g02010)*  
*GAD5 (At3g17760)*

GAD1	<a href="#">rs2058725</a>	C	CC	+/+
GAD1	<a href="#">rs3791851</a>	C	TT	-/-
GAD1	<a href="#">rs3791850</a>	A	AA	+/+
GAD1	<a href="#">rs12185692</a>	A	CC	-/-
GAD1	<a href="#">rs3791878</a>	T	GG	-/-
GAD1	rs10432420	A	AA	+/+
GAD1	<a href="#">rs3828275</a>	T	CC	-/-

Association between glutamic acid decarboxylase genes and anxiety disorders, major depression, and neuroticism.  
Mol Psychiatry. 2006 Aug;11(8):752-62. Epub 2006 May 23.  
Hetteima JM1, An SS, Neale MC, Bukszar J, van den Oord EJ, Kendler KS, Chen X.



## Association between glutamic acid decarboxylase genes and anxiety disorders, major depression, and neuroticism.

Mol Psychiatry. 2006 Aug;11(8):752-62. Epub 2006 May 23.

Hettema JM<sup>1</sup>, An SS, Neale MC, Bukszar J, van den Oord EJ, Kendler KS, Chen X.

### Abstract

Abnormalities in the gamma-aminobutyric acid (GABA) neurotransmitter system have been noted in subjects with mood and anxiety disorders. Glutamic acid decarboxylase (GAD) enzymes synthesize GABA from glutamate, and, thus, are reasonable candidate susceptibility genes for these conditions. In this study, we examined the GAD1 and GAD2 genes for their association with genetic risk across a range of internalizing disorders. We used multivariate structural equation modeling to identify common genetic risk factors for major depression, generalized anxiety disorder, panic disorder, agoraphobia, social phobia and neuroticism (N) in a sample of 9270 adult subjects from the population-based Virginia Adult Twin Study of Psychiatric and Substance Use Disorders. One member of each pair of monozygotic twins was randomly selected as a case or control based on the results of the analysis. The resulting sample was used in a two-stage association study in which candidate genes were tested for replication in a second sample. In the GAD1 region demonstrated in the first sample, a common high-risk haplotype was identified in all 1128 subjects indicated that they formed a common high-risk haplotype that was significantly over-represented in cases (P=0.003) with effect size OR=1.23. Out of 14 GAD2 markers screened in stage 1, only one met the threshold criteria for follow-up in stage 2. This marker, plus three others that formed significant haplotype combinations in stage 1, did not replicate their association with the phenotype in stage 2. Subject to confirmation in an independent sample, our study suggests that variations in the GAD1 gene may contribute to individual differences in N and impact susceptibility across a range of anxiety disorders and major depression.

Abnormalities in the gamma-aminobutyric acid (**GABA**) neurotransmitter system have been noted in subjects with mood and anxiety disorders.

# ROS, Aldehydes (Yeast)

SOD2	<a href="#">rs2758331</a>	A	AC	+/-	
SOD2	rs2855262	T	CT	+/-	
SOD2 A16V	<a href="#">rs4880</a>	G	AG	+/-	
PON1 Q192R	<a href="#">rs662</a>	C	CT	+/-	

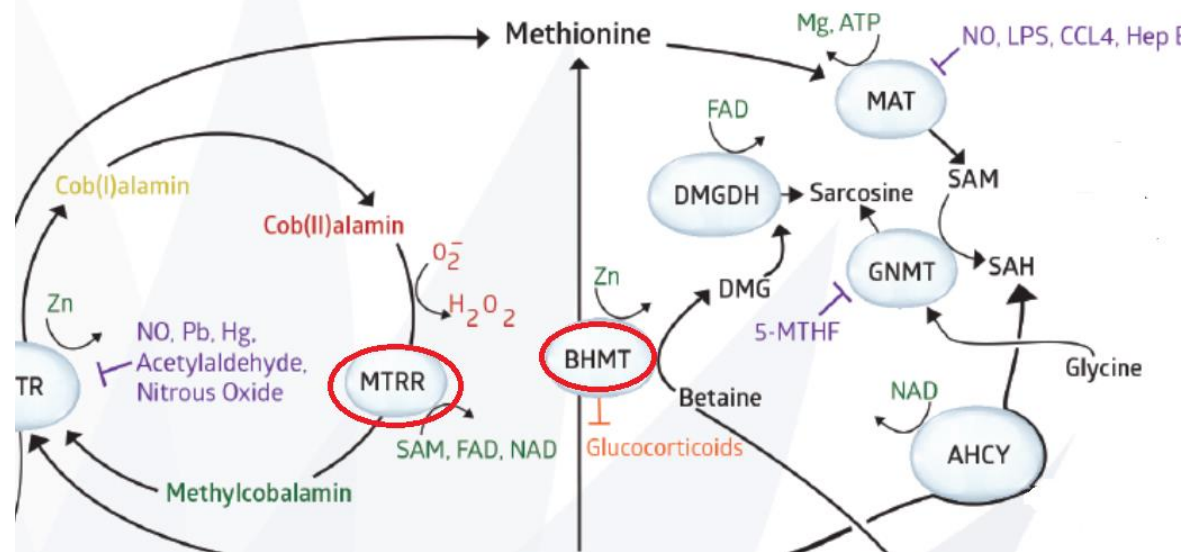
SOD suspect mitochondrial involvement. Involved in MCS

PON1 Organophosphates (Patient lives in a farming community)

Suspect difficulty in metabolizing aldehydes. Also involved in MCS

NAT2 A803G (K268R)	<a href="#">rs1208</a>	G	AG	+/-	
NAT2 C190T (R64W)	<a href="#">rs1805158</a>	T	CC	-/-	
NAT2 G590A (R197Q)	<a href="#">rs1799930</a>	A	AG	+/-	
NAT2 G857A (G286E)	<a href="#">rs1799931</a>	A	GG	-/-	
NAT2 T341C (I114T)	<a href="#">rs1801280</a>	C	CT	+/-	

# BHMT



MTRR R415T	-/-
MTRR-11 A664A	+/-
MTRR	-/-
MTRR	+/-

BHMT-02	+/-
BHMT-04	+/-
BHMT-08	+/-
BHMT R239Q	-/-

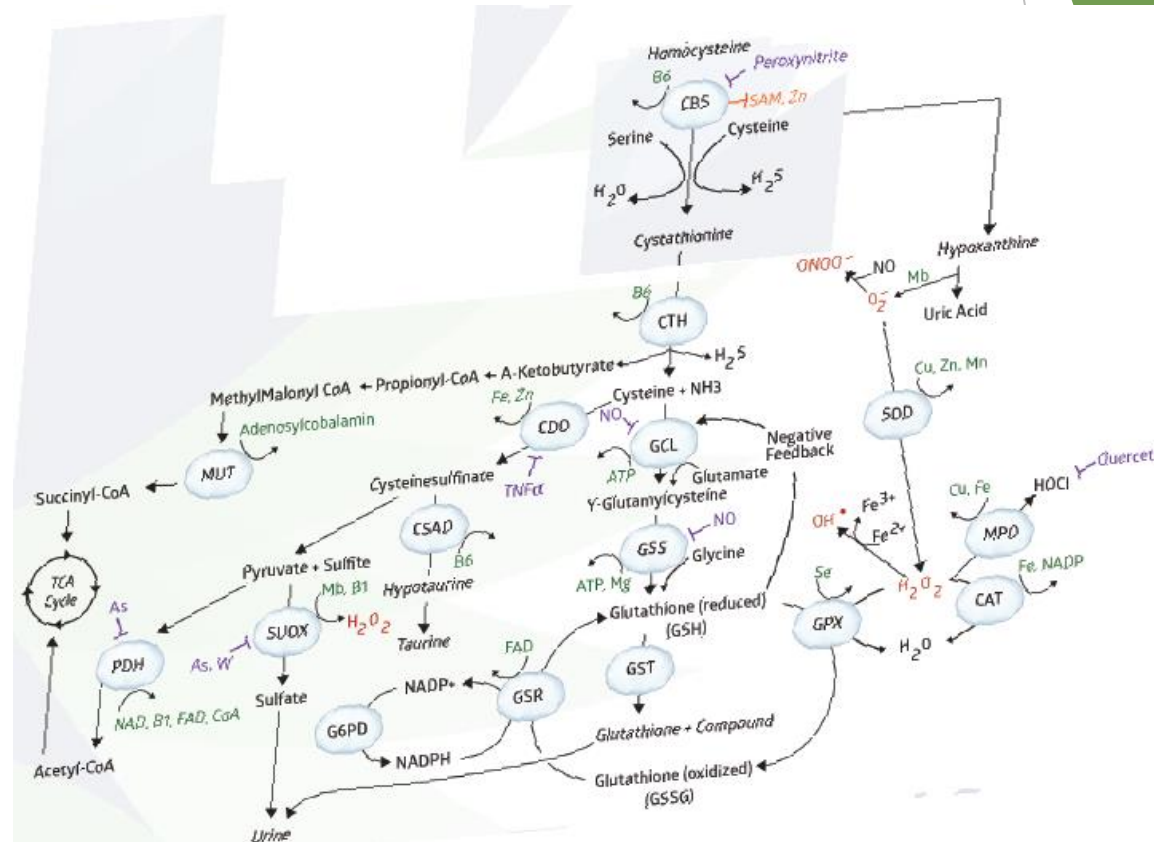
Pearl: Patients like this will internalize stress and/or have chronic dysthymia. People with this pattern who have PTSD will respond better to EMDR than psychotherapy (talk therapy)



Obeid, Rima. "The Metabolic Burden of Methyl Donor Deficiency with Focus on the Betaine Homocysteine Methyltransferase Pathway." *Nutrients* 5.9 (2013): 3481-3495. *PMC*. Web. 30 July 2015.

# TRANSULFURATION

CBS A13637G	+/-
CBS A360A	+/-
CBS C19150T	-/-
CBS C699T	-/-
CBS N212N	-/-



After questioning and review of labs, the transsulfuration pathway did not seem to express in this patient. When it does express you may see brain fog, high ammonia on lab tests and/or high taurine on NT testing.

# FUT2 & IGA

FUT2	<a href="#">rs492602</a>	G	AG	+/-	
FUT2	<a href="#">rs601338</a>	A	AG	+/-	
FUT2	<a href="#">rs602662</a>	A	AG	+/-	

**FUT2 has possible contribution to imbalances in the gut microbiome and B12**

**Tendency toward food allergies especially with leaky gut syndrome**

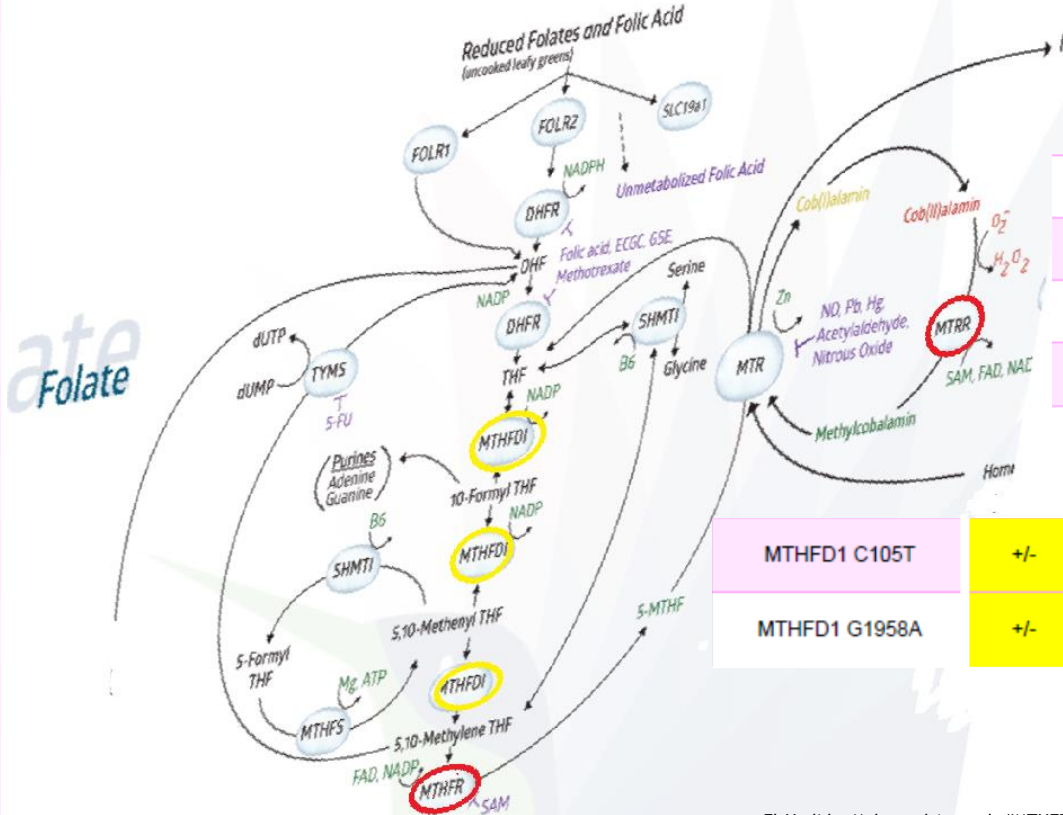
## IgA Snps

TRAF1	<a href="#">rs3761847</a>	G	AG	+/-	
IRF5	<a href="#">rs4728142</a>	A	AG	+/-	
IGF1R	<a href="#">rs2229765</a>	A	GG	-/-	
IFIH1 (HLA)	<a href="#">rs1990760</a>	C	CT	+/-	
HLA	<a href="#">rs9271366</a>	G	AG	+/-	
CFH	<a href="#">rs6677604</a>	A	GG	-/-	
HLA-DQA2	<a href="#">rs9275224</a>	A	AG	+/-	
MTC03P1	<a href="#">rs9275596</a>	C	CT	+/-	
PSMB8 / TAP1 / TAP2	<a href="#">rs9357155</a>	A	GG	-/-	
HLA-DPB2 / COL11A2P	<a href="#">rs1883414</a>	A	GG	-/-	



# METHYLATION

MTHFR C677T	+/-
MTHFR A1298C	+/-
MTHFR G1793A (R594Q)	+/-
MTHFR	+/-
MTHFR	-/-
MTHFR	+/-
MTHFR	+/-
MTHFR	+/-
MTHFR	+/-
MTHFR	-/-
MTHFR	-/-



MTRR R415T	-/-
MTRR-11 A664A	+/+
MTRR	-/-
MTRR	+/+

MTHFD1 C105T	+/-
MTHFD1 G1958A	+/-

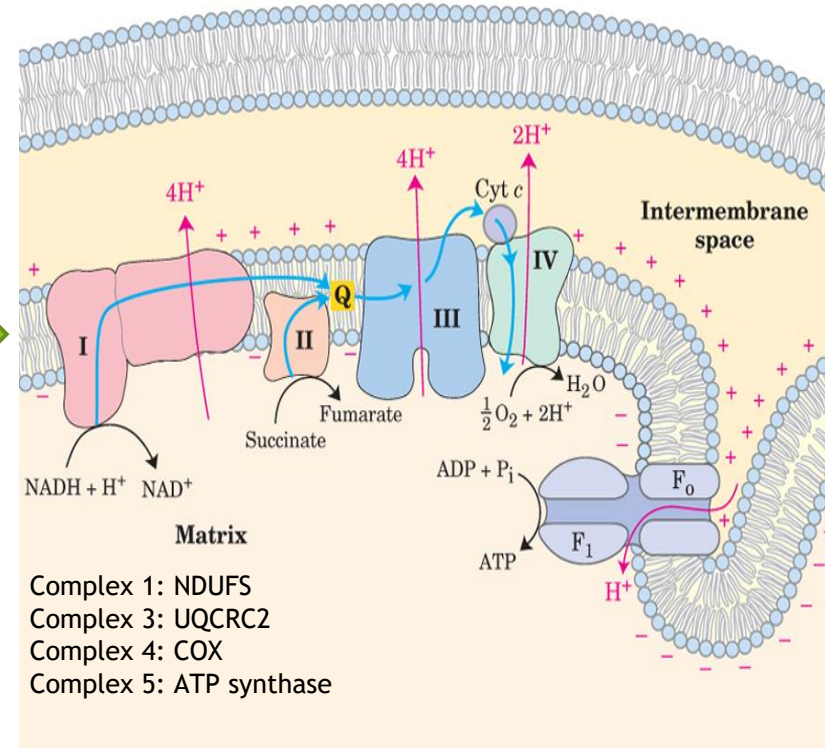
El-Hadidy, Mohamed A. et al. "MTHFR Gene Polymorphism and Age of Onset of Schizophrenia and Bipolar Disorder." *BioMed Research International* 2014 (2014): 318483. PMC. Web. 30 July 2015.

Neurosci Biobehav Rev. 2013 Sep;37(8):1597-610. doi: 10.1016/j.neubiorev.2013.06.006. Epub 2013 Jun 18. The role of COMT gene variants in depression: Bridging neuropsychological, behavioral and clinical phenotypes. Antypa N1, Drago A, Serretti A.

(c) Dr Jess P Armine (unless otherwise noted) 2014

# MITOCHONDRIAL COMPLEX 1-THE MOST IMPORTANT

NDUFS7	rs2332496	A	AG	+/-
NDUFS7	rs7254913	G	AA	-/-
NDUFS7	<a href="#">rs1142530</a>	T	TT	+/+
NDUFS7	<a href="#">rs7258846</a>	T	TT	+/+
NDUFS7	<a href="#">rs11666067</a>	A	AA	+/+
NDUFS7	rs2074895	A	AA	+/+
NDUFS7	rs809359	G	AA	-/-
NDUFS8	rs4147776	C	AA	-/-
NDUFS8	rs1122731	A	AG	+/-
NDUFS8	rs999571	A	AG	+/-
NDUFS8	rs2075626	C	CT	+/-
NDUFS8	rs3115546	G	TT	-/-
NDUFS8	rs1104739	C	AC	+/-
NDUFS8	rs1051806	T	CT	+/-



NADH-ubiquinone oxidoreductase (NDUFS) - GSSG will block the entry of the electron donors into the electron transport chain

# What is Expressing?

## Pointers to the Diagnoses

Complaint/ Symptoms	Snps	Index of Suspicion high for these root causes	Downstream effects	Questions to ask	Testing
Hallucinations (excitation)	COMT, MAO, MTHFD1, GAD, MTHFR, MTRR, MTR	Immune issues, microbial involvement	Neurotransmitter Imbalance	Voices “chattering” or screaming (intrusive thoughts)	NT test, Tests for: Lyme, Co-infections, Viruses, parasite, Candida, etc.
“Bad Gut”	IgE, IgA, IgG DAO, HNMT, HDC. HRH, FUT2	Leaky Gut Syndrome	Immune Upregulation, Immune dysregulation, Dysautonomia, Histamine Intolerance	Relationship of symptoms to to food intake, color/frequency of BM,	Food Allergy Tests, Organic Acid Test, Cross Reactivity Testing
Bad Gut	NAT/ALDH (aldehyde metabolism)	Yeast (acetyl-aldehydes)	Neural upregulation, adrenal fatigue,	How does patient react to ETOH intake? Coated tongue?	Stool, Antibody Testing, B5 level
Mitochondrial Dysfunction	NDUFS, COX, UQCRC2, ATP	GSSG, Oxidative stress	Fatigue, lack of healing ability	Ask about fatigue, lack of ability to heal,	ATP, ADP conversation, GSSG, reduced GSH, anti oxidant testing (SOD), Thyroid panel
Methylation	MTHFD1, MTHFR, MTRR, MTR	All of the Above	General lack of ability to heal	(too broad, many symptoms)	Organic Acid Testing, cellular micronutrient analysis

# Alyssa's SNPs Indicated Probable Issues in the Following Areas...

## Areas/pathways

- ▶ Neurotransmitters
- ▶ Leaky Gut Syndrome
- ▶ Aldehyde Metabolism
- ▶ Methylation
- ▶ Mitochondrial function

## How do we use this information?

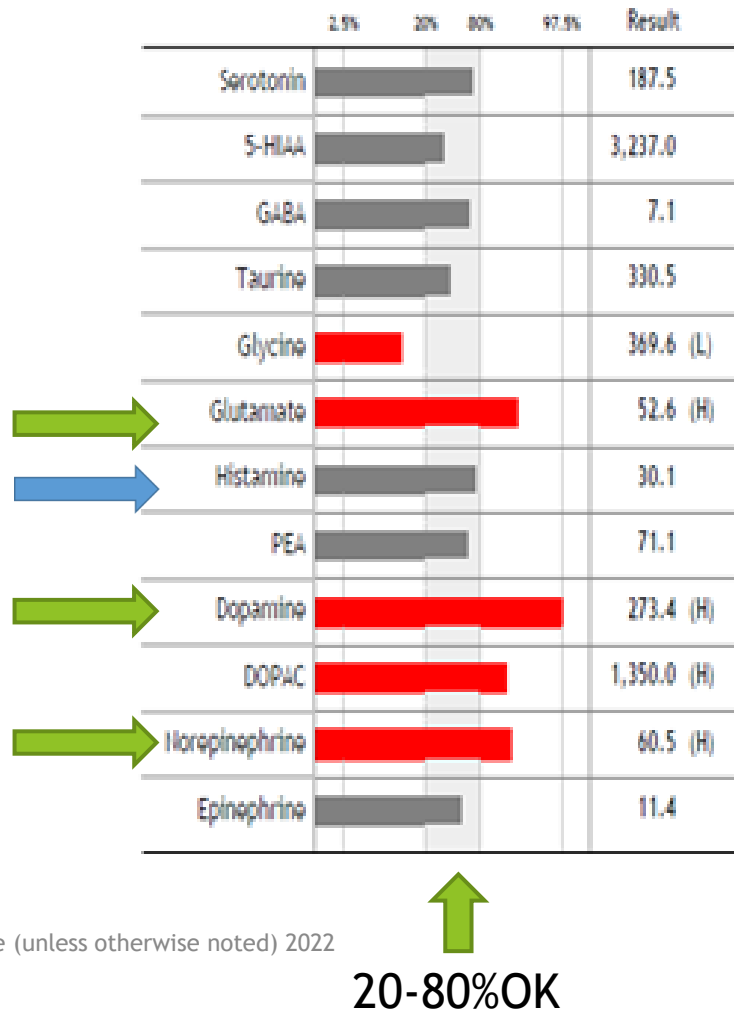
- ▶ **Correlate, correlate, correlate!** Or, if all the dogs are barking up a tree, don't yell at the dogs...look up the tree!
- ▶ Use the estimated function of the enzymes (snps) and compare them to:
  - ▶ Symptoms
  - ▶ Personal/family Hx
  - ▶ Clinical observations
- ▶ Use the estimated function of the enzymes (snps) to:
  - ▶ Raise index of suspicion of root cause(s)
  - ▶ Help identify downstream effects
  - ▶ Determine which tests will solidify diagnoses
  - ▶ Ultimately, assist you in creating an individualized, successful treatment plan



- (c) Dr Jess P Armine (unless otherwise noted) 2022



# Alyssa's NTs...Where is she?



(c) Dr Jess P Armine (unless otherwise noted) 2022

## ► Inhibitory/Excitatory Balance

- Visually compare the rough levels of inhibitory NT's (Serotonin, GABA, Taurine, Glycine) with the excitatory NT's (Glutamate, Histamine, PEA, Dopamine, Norepi, Epi)
- The excitatory NT's "outweigh" the inhibitory NT's
- The "Net Result" is an excited nervous system.

## ► Hallucinations

- Always from over excitation
- Classically, high dopamine causes hallucinations (*but not always*)
- Glutamate, Histamine, PEA, Dopamine, Norepi, Epi can all cause "excitotoxicity" or perhaps it's the combination

## ► Adrenal Fatigue

- Give indicator of how long her root causes have been present

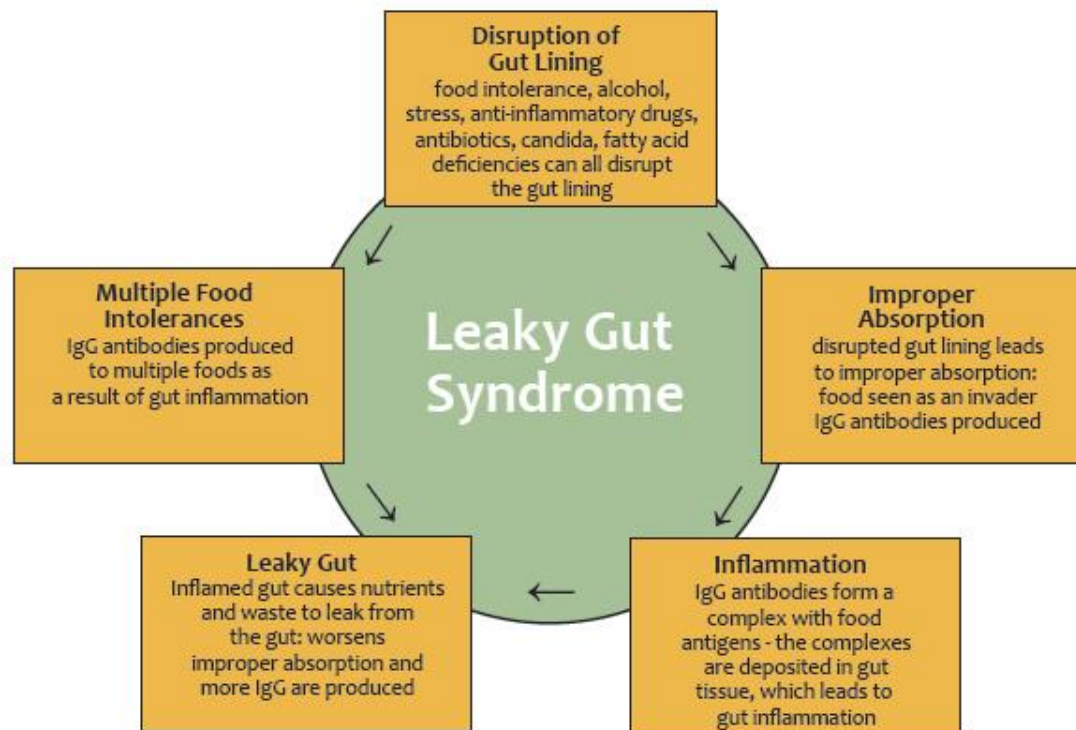


# Who is Upregulating The Nervous System

We Need To Elucidate

# Leaky Gut Syndrome

Hints: “bad gut” on history; IgA/IgG/IgE, SHMT, FUT2 SNPS; Food Allergy Testing



Source: <http://allergytreatmentservices.com/digestion.html>

## Net Result...INFLAMMATION



## Candida OverGrowth Symptoms

### ANXIETY

HeadAches-Migraines

### VAGINITIS

### EXCESSIVE FATIGUE

### ACNE

### DIZZINESS

Athlete's Foot

low sex drive

### ALCOHOL CRAVINGS

Inability to Concentrate

HyperActivity

### MOOD SWINGS

Sinus Inflammation

Poor Memory

Cognitive Impairment

learning difficulties

### ITCHING

### ECZEMA

### DEPRESSION

### PMS

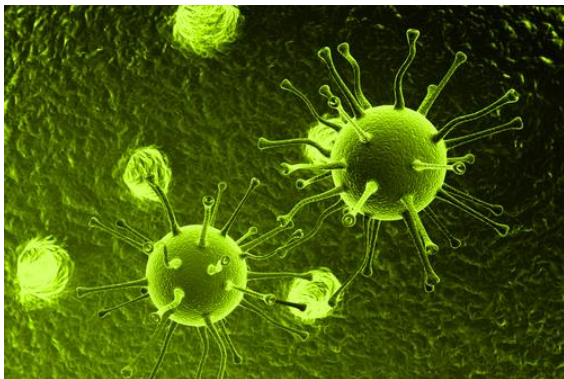
### PERSISTENT COUGH

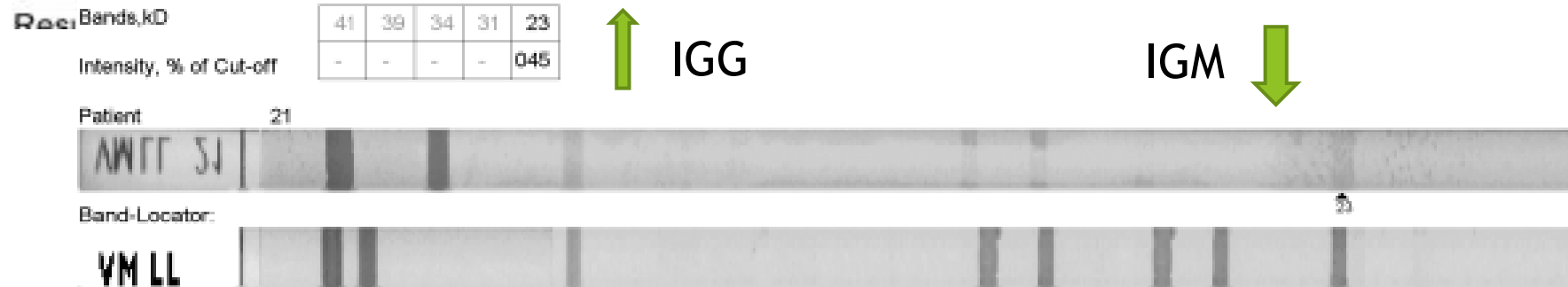
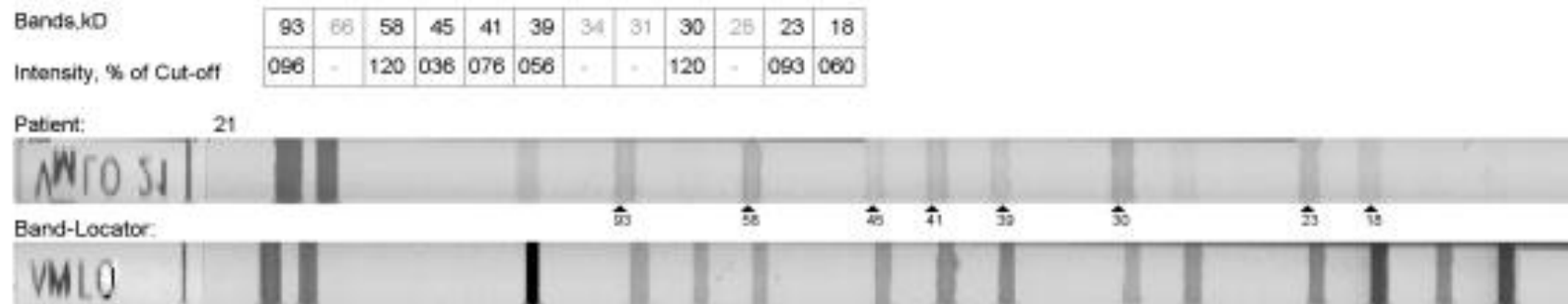
chronic pain

Irritability

muscle weakness

# Microbial Involvement

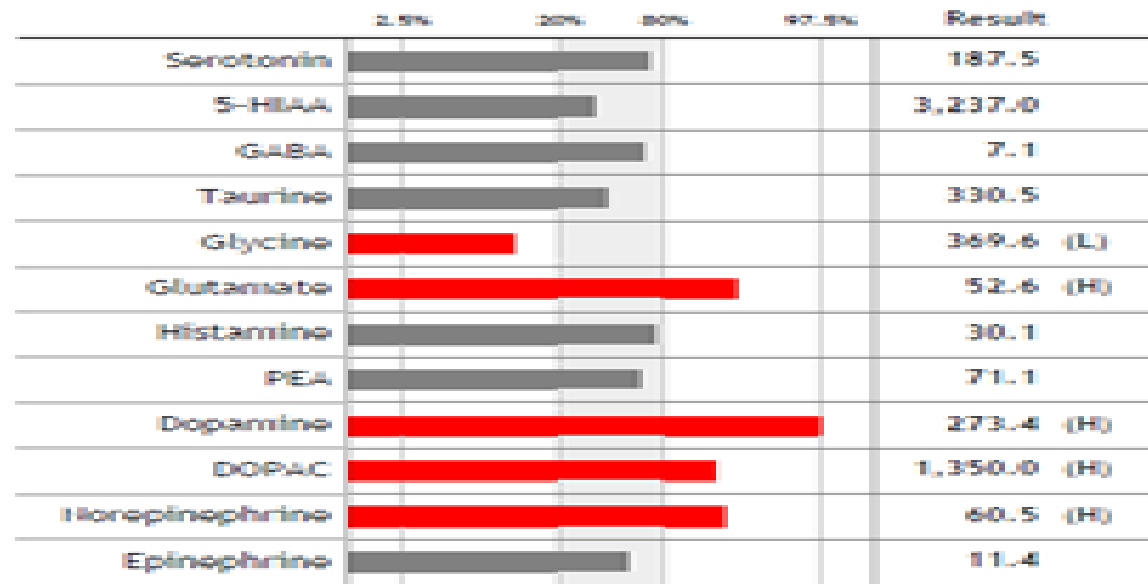




↑ IGG

IGM ↓

Results for this specimen:





# Lyme Disease: Adult Symptoms

Close Window

## Fast Facts

- Lyme is fastest growing vector-borne disease
- 85% do not recall tick bite
- Less than 70% of people develop a rash
- Treatment should begin without testing if rash is present
- Lab tests may be negative in the first 4-6 weeks

## Early symptoms

- Flu-like illness (fever, chills, sweats, muscles aches, fatigue, nausea and joint pain)
- Rash (10% have EM rash)
- Bell's palsy

CHILDREN'S SYMPTOMS

## Later Symptoms

- Headache
- Stiff neck
- Light or sound sensitivity
- Cognitive impairment
- Sleep disturbance
- Depression, anxiety, or mood swings
- Arthritis
- Fatigue
- Abdominal pain, nausea, diarrhea
- Chest pain, palpitations
- Shortness of breath
- Tingling, burning or shooting pains



# Alyssa's Labs and Dx

## Lab

- ▶ Alyssa was extensively tested and found to have antibodies to Yeast and HHV6.
- ▶ A Western Blot for Lyme was positive in my opinion.
- ▶ Numerous food allergies by IgG testing.
  - Concentrations were in Gluten, Dairy and Yeast areas.

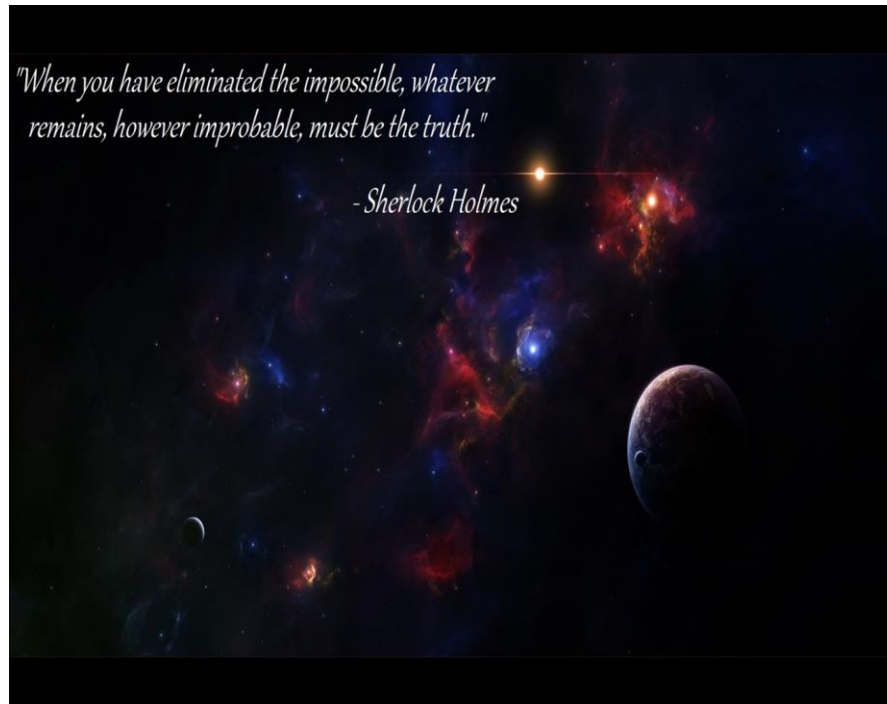
## Working Diagnoses

- ▶ Lyme Disease (neural) leading to neural upregulation
- ▶ Yeast overgrowth (gut) releasing acetaldehydes (neurological irritant)
- ▶ Leaky Gut Syndrome (food allergies, immune upregulation)
- ▶ Viral Syndrome (neural dysregulation)

# CHECKPOINT:

## Root Cause vs. Downstream Effect...

### Ask Yourself...

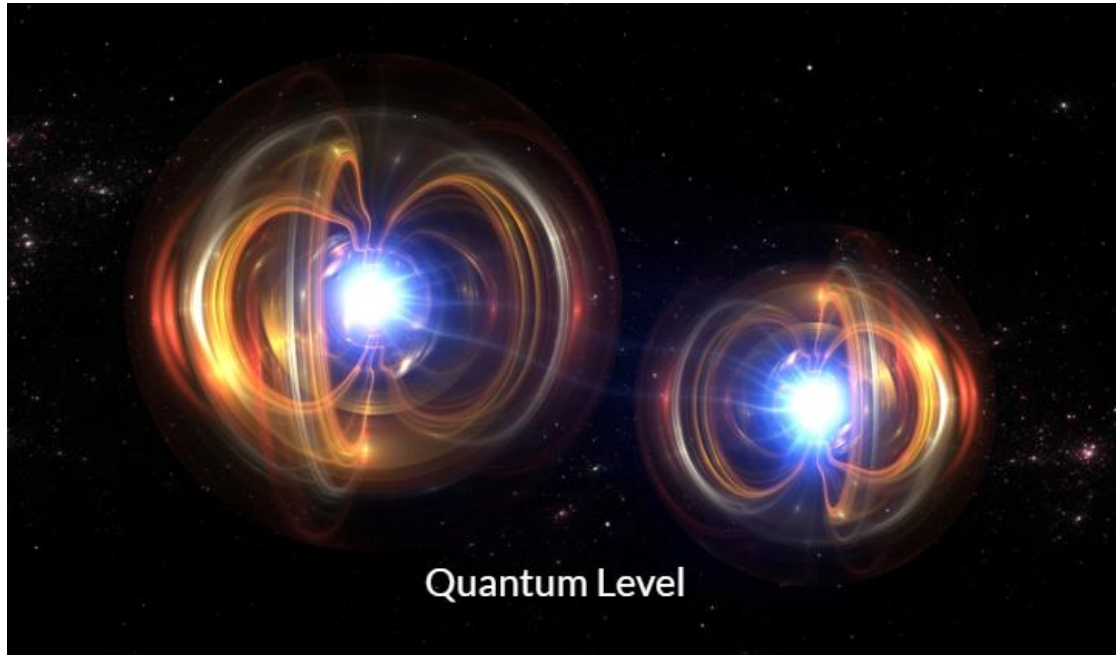


- ▶ Leaky Gut Syndrome can and does lead to immune upregulation/dysregulation. This is evidenced by the numerous food allergies and the patient's GI symptoms.
  - ▶ *Can this cause the increase in catecholamines and hallucinations?*
- ▶ Lyme and HHV6 attack the neural cells.
  - ▶ *Can this cause the increase in catecholamines and hallucinations?*
- ▶ Yeast overgrowth causes increased levels of acetaldehyde. Combined with the NAT2 snps.
  - ▶ *Can this cause neural inflammation resulting in hallucinations?*
- ▶ Answer: **Yes to all of the above.**

# What To Do? When to Do it? Keep It “Foundational”

***“Reduce Stress,  
Heal the Cells,  
Heal the Gut,  
Kill the Bugs!!”***

# You May Understand Physiology Down to the Quantum Level But You Can Only Intervene at the Global Level







Turn This



Into This

TREAT ROOT CAUSES AND DOWNSTREAM EFFECTS

TREATMENT  
(LESSEN THE TRAFFIC)

# Treatment

## Step 1: Cellular & GI Repair

### ► Principles:

- We need to fully digest foods. Undigested foods make up the majority of the antigens that enter our bodies.
- We need to re-create the mucus layer in the gut. The mucus layer is where the microbiome live, what they eat, and where they do their work. The mucus layer traps antigens, toxins, xenobiotics and forms the initial layer of protection.
- We need to repair the cells and the tight junctions. This area is our second layer of defense preventing the entry of the above mentioned into our bodies.
- We need to re-populate the GI tract with an adequate diversity of biota.
- Demulcent herbs to recreate the mucus layer\*
- Phospholipids to support cellular repair
- Digestive enzymes to assure full breakdown of foods and prevent creation of antigens.
- DAO enzyme to help break down histamine
- Probiotics (soil based with *S. Boulardii*)
- Absorbable Vitamins and Minerals

# Treatment

## Step 2: Kill the Bugs

*The Quintessential Devils in this Matter*

- ▶ Child was co treated by myself and an integrative pediatrician
- ▶ GI Repair program was conducted for a period of 3 months
- ▶ Thereafter, we went after the bugs. There was some disagreement as to the form of treatment (whether to insert a PICC line and use rotating antibiotics or use other available non-pharmaceutical options)
- ▶ The parents were given full information, pros, cons, etc. by the pediatrician and myself
- ▶ The parents chose the latter.

# Treatment

## Step 3: Retest

- ▶ Three months after biocidal treatment was initiated, testing was done again at the same lab.
- ▶ All results were negative
- ▶ All symptoms were gone



*The Sleuthing Was Worth It!*

# Alyssa is now in College

She's an Honor Student, Star Athlete, and a beautiful young woman. 18 years old



8 y/o

## **Reality:**

*Hallucinations were a expression of genetic predisposition caused by neural excitation and immune upregulation secondary to infections and leaky gut syndrome. All "kept going" by chronic CDR.*

12y/o



16 y/o



## **Prospect:**

*A life on antipsychotic meds*



## **Result:**

**A life saved**



The background features a dark, reflective surface with several bright green neon lines. These lines are arranged in a series of vertical, slightly curved paths that converge towards a central point. A single, very bright and straight green line runs vertically down the center, creating a strong focal point. The light from these lines reflects on the surface below, creating a shimmering effect.

# Chelsi

Story of Hope

# Started with Febrile Seizures at 10 months old

## ► 1994

- **4 seizures in 3 years.** Normal birth. First seizure at 10 months old during high fever. Maybe 4 more seizures throughout the next 3 years, but only during high fever.

## ► 1997

- **A Few absence.** In August, started with freezing type seizures where she would bend her arms up, turn her head and eyes to the right and gently, rhythmically quiver for 3 to 10 seconds with head and eyes to the right. Only happens when she's playing, and if she playing hard enough to get breathless, they would last longer (up to a minute) and she would collapse after.
  - Started on **Zarontin** – didn't do much (started absence seizures)
  - **Depakote** added – seizure free for 2 months, weaned off **Zarontin**. As soon as she was on a lower dose, myoclonic jerks started

## ► 1998

- **2 febrile/lots of myoclonic.** MRI at Childrens 5/98 diagnosed myoclonic with other generalized activity.
  - **Depakote** level 138, weaned to introduce **Lamictal**
  - Started **Lamictal** 25mg, ½ tab bid
  - 2-3 weeks later, 2 tonic clonic seizures 15 minutes apart (July)
    - Became severely ataxic and ended up in Children's for 4 days.
      - **Thrombocytopenia** - Platelet count in the 60's, got a rash, **Depakote** decreased, **Lamictal** discontinued
- **Video EEG** 5/28 & 5/29 Generalized slowing and myoclonic seizures. Tendency for generalized atypical SW discharges to the rt hemisphere.

## ► 1999

- **No Seizures.**
  - **Depakote** 250mg AM, 125mg PM, **Zarontin** 250mg – Had been seizure free for several months.
  - Was under care of Marcio Sotero, started seeing Dr. Graf locally through Childrens 10/99
  - Seizure free for over 1 year. Drugs decreased because of sleepiness/cognitive difficulties.

## ► 2006

- **9 tonic clonic seizures**
- Seizures 12/26, 10/9, 9/22, 8/1, 8/10, 7/20, 4/9, 3/14, 2/28, (nothing 10 months prior)
  - **First seizure EVER not related to HEAT**...and continued to have them for no reason.
  - Seizures more convulsive
- 6/20 Increased **Zonegran** to 175 in the evening, **Keppra** 750 BID.
  - March is when the frequency really picked up and seizures were unpredictable and harder.
  - Started mensus for the first time October 6, 2006
  - **Zonegran** 125 in eve and **Keppra** 1000 bid. Drugs switched in hopes of less absence, but result was more grand mal.
- MRI at Childrens 8/25 - normal

# Number and type of seizures increased through the years

## ▶ 2010

### ▶ **47 tonic clonic seizures**

- **Zonegran** 300 mg /day (tried to stop...it didn't go well- seizures increased and came at odd times)
- **Keppra** 1750 bid
- **Clobezam** down to 7.5 mg/day

## ▶ 2011

### ▶ **74 tonic clonic seizures**

- ▶ **Video EEG** done at Harborview Epilepsy Center.

Confirmed generalized seizures.

- ▶ **NutrEval** 10/11 – NOTE still low on [alanine](#), [lysine](#), [glycine](#), [senine](#) (see 2005 Swedish tests)

- ▶ **Hair** sent to Doctor's Data for Toxic testing: High in Uranium, Copper, Cobalt, Lead

- ▶ **Blood** tests showed Chloride High, Alkaline Phosphatase low

- ▶ **DNA** tested for C677T and A1298C mutations (negative)

- ▶ **FOOD** allergy tested 10/11 – High antibodies for Cheese, Milk, Whey, beef, lamb, pork

- **Zonegran** 200 mg bid

- **Keppra** 1000 bid

- **Clobezam** 12.50.

- **Banzel** ? mg – Started falling forward. Many daytime seizures and injuries and stitches. Weaned off between February & July

## ▶ 2012

### ▶ **101 tonic clonic seizures (probably 10% just tonic)**

- ▶ **NutrEval** 10/12 – NOTE still low on [alanine](#), [lysine](#), [glycine](#), [senine](#) (see 2005 Swedish tests)

- ▶ Jan to June, 90% of seizures between 2pm&6pm. Many injuries and stitches.

- ▶ Medical Marijuana tried 6/26/12 to 12/12. Switched seizures to her sleep (perhaps because she was getting sugar in the brownies?), but frequency increased. Seizures would start at 9:38pm and sometimes had 2 or 3 per night.

- **Zonegran** 200 mg bid

- **Keppra** 1000 bid

- **Clobezam** 10 mg bid

- **Vimpat (Lacosamide)** 125mg /day (I believe this is why they switched to daytime)

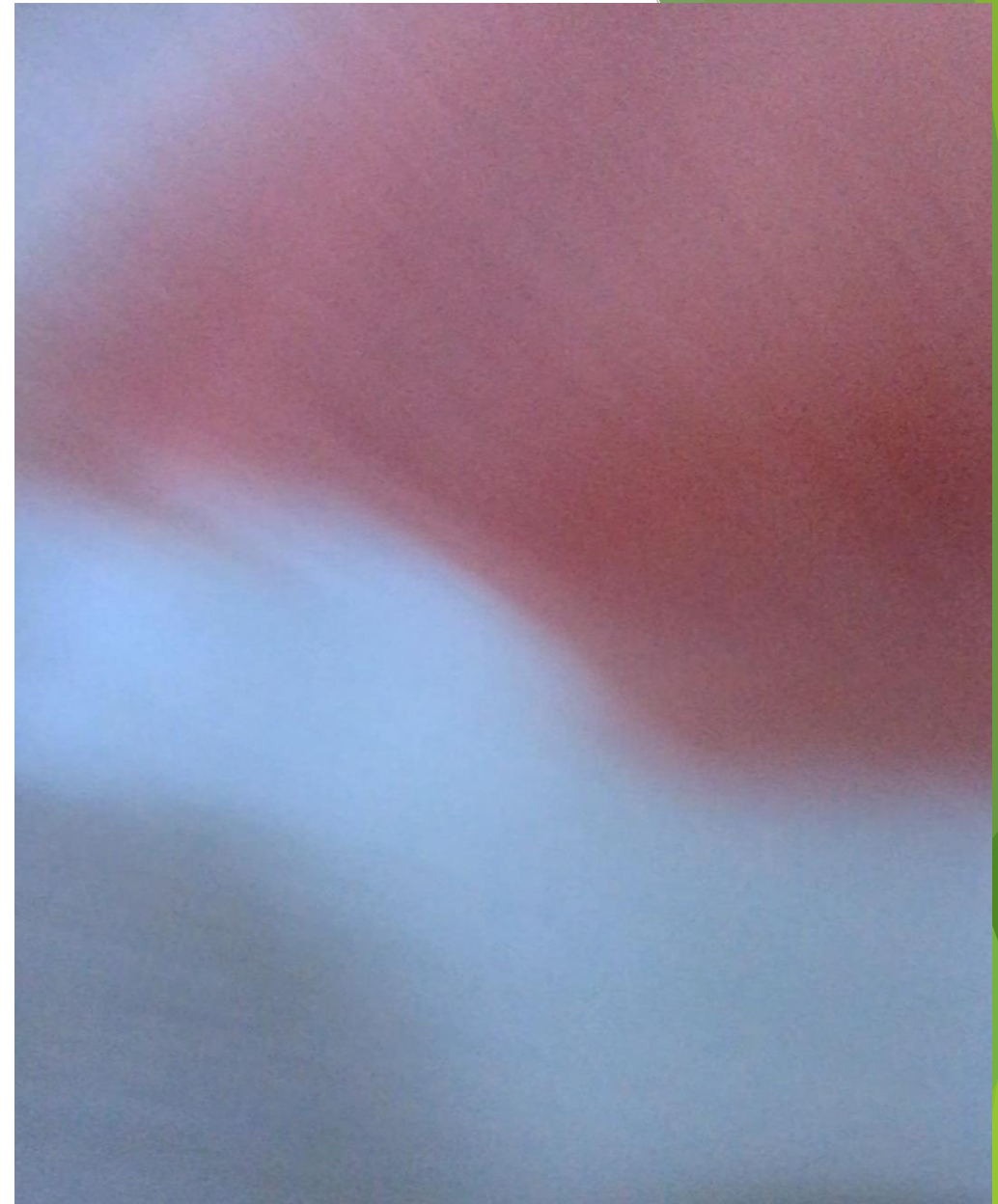
## ▶ 2013

### ▶ **146 tonic clonic seizures**

- **I BELIEVE INCREASES FOR THE LAST 3 YEARS WERE FROM BANZEL, VIMPAT & RUFINIMIDE.**

Depakote added back in because Dr. Miller saw this history and thought it was worth going back. I disagree.

What she  
fought.  
Tonic Clonic  
seizures



### One year report for Chelsi Ownby

Date created: March 17, 2020

Start Month: April 2019

End Month: March 2020

Weight: 106 lbs. (as of June 12, 2015)

Birth date: March 11, 1994

Current Medications: Depakote 1000.00 mg Daily

Keppra 1250.00 mg Daily

Zonegran 50.00 mg Daily

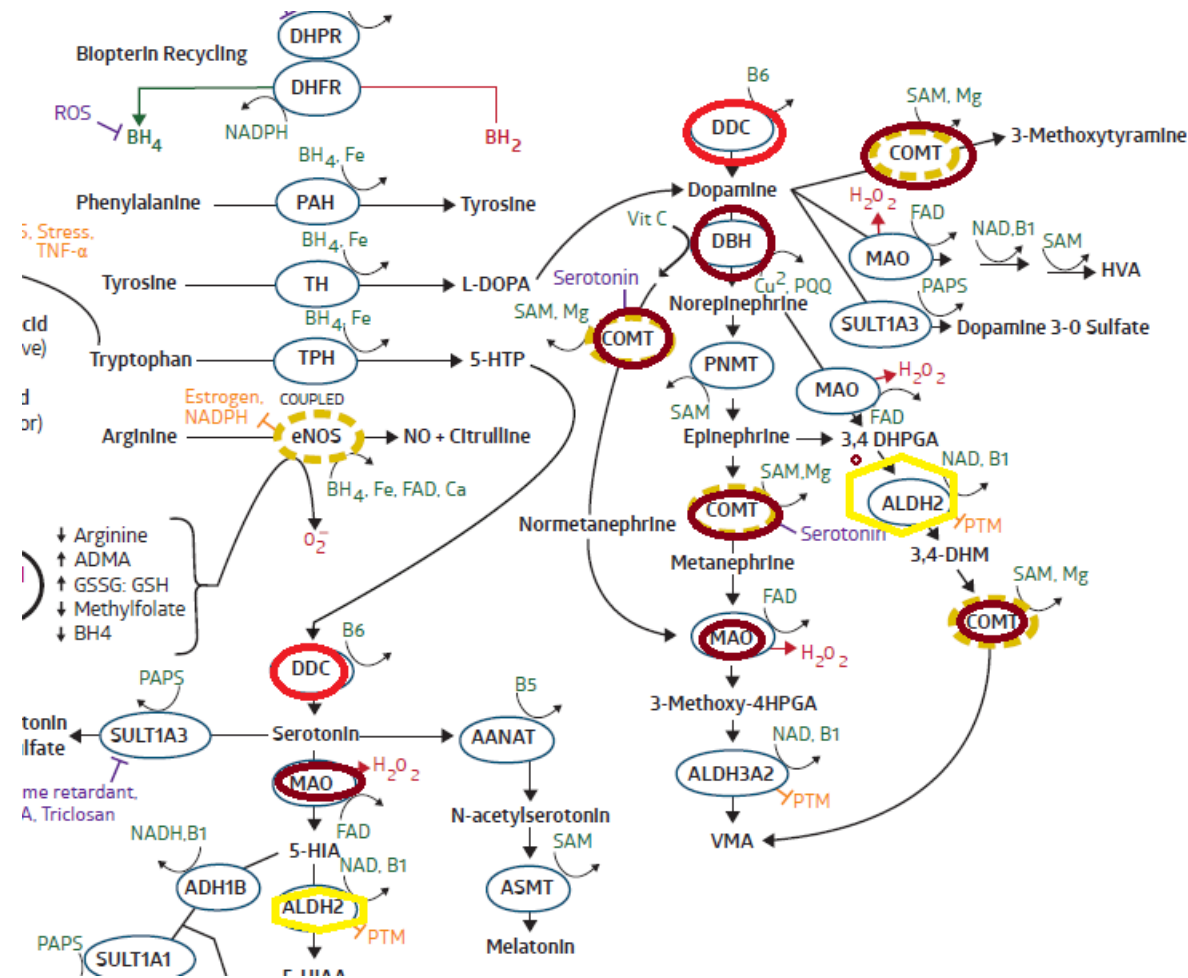
April, 2019		May, 2019		June, 2019	
Week 1	4	Week 1	5	Week 1	
Week 2	5	Week 2	4	Week 2	
Week 3	3	Week 3	4	Week 3	
Week 4	5	Week 4	3	Week 4	
Total: 17		Total: 16		Total:	
July, 2019		August, 2019		September, 2019	
Week 1	3	Week 1	5	Week 1	
Week 2	3	Week 2	4	Week 2	
Week 3	2	Week 3	4	Week 3	
Week 4	4	Week 4	5	Week 4	
Total: 12		Total: 18		Total:	
October, 2019		November, 2019		December, 2019	
Week 1	4	Week 1	6	Week 1	
Week 2	4	Week 2	6	Week 2	
Week 3	3	Week 3	3	Week 3	
Week 4	9	Week 4	7	Week 4	
Total: 20		Total: 22		Total:	
January, 2020		February, 2020		March, 2020	
Week 1	3	Week 1	15	Week 1	
Week 2	4	Week 2	4	Week 2	
Week 3	7	Week 3	4	Week 3	
Week 4	9	Week 4	5	Week 4	
Total: 23		Total: 28			

25

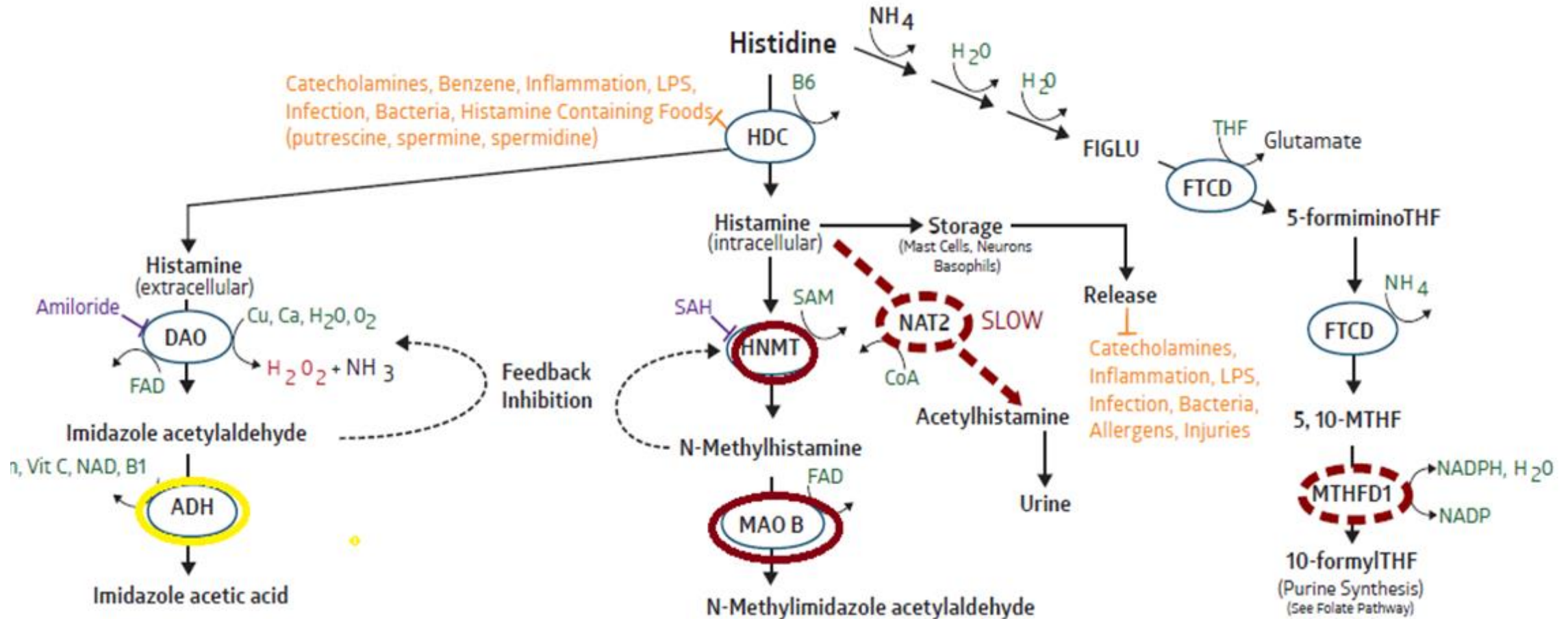
Number of monthly seizures steadily increased from 4/2019 through 3/2020



## Genetic Predisposition toward Excitation

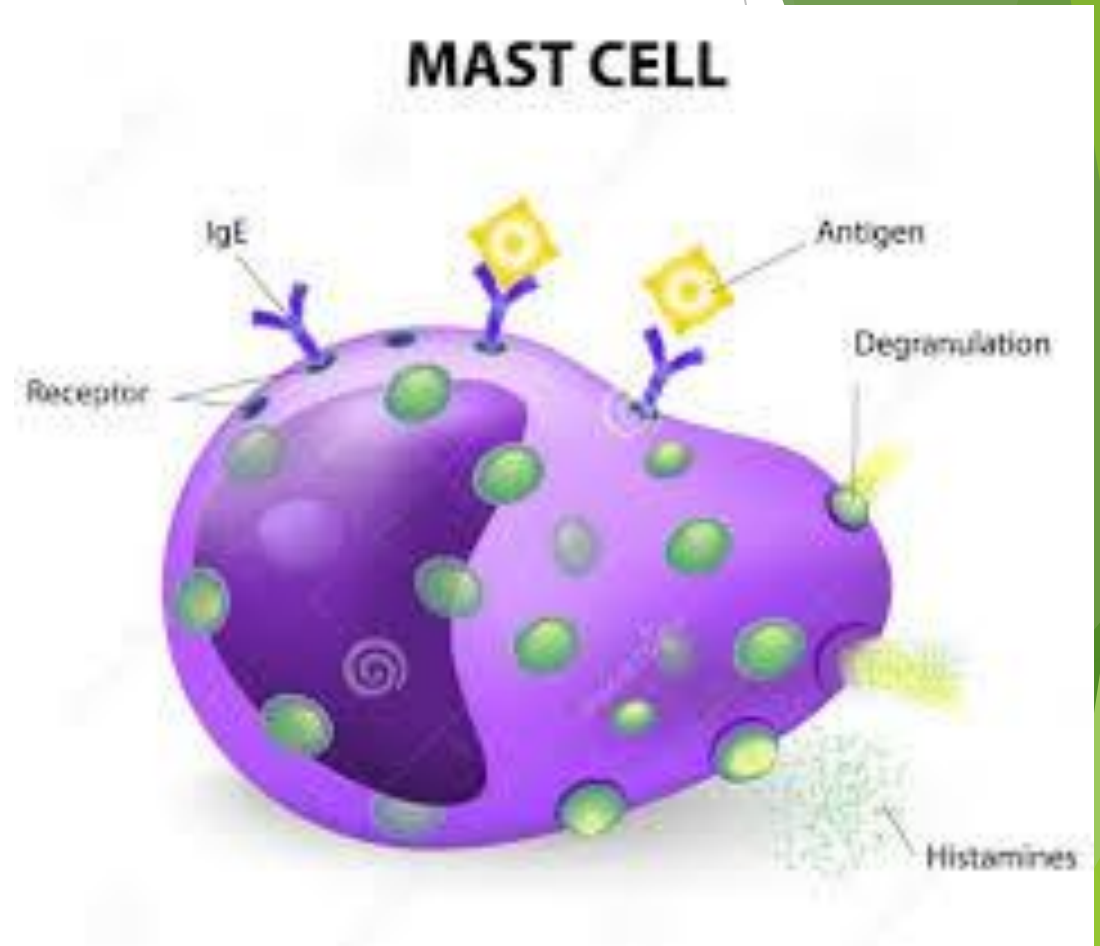


# Predisposition to High Histamine



# Histamine: The Major Offender

- ▶ Histamine is most commonly known for its role in allergic reactions
- ▶ Also involved in neurotransmission and can affect your emotions and behavior as well.
- ▶ Histamine helps control the sleep-wake cycle and promotes the release of epinephrine and norepinephrine.
- ▶ High histamine levels have been linked to obsessive compulsive tendencies, depression, and headaches.
- ▶ Low histamine levels can contribute to paranoia, low libido, fatigue, and medication sensitivities.





## Toxic & Essential Elements; Hair

TOXIC METALS				PERCENTILE	
		RESULT µg/g	REFERENCE INTERVAL	68 <sup>th</sup>	95 <sup>th</sup>
Aluminum	(Al)	3.4	< 8.0		
Antimony	(Sb)	< 0.01	< 0.066		
Arsenic	(As)	0.036	< 0.060		
Barium	(Ba)	0.93	< 1.5		
Beryllium	(Be)	< 0.01	< 0.020		
Bismuth	(Bi)	0.17	< 2.0		
Cadmium	(Cd)	0.019	< 0.060		
Lead	(Pb)	0.80	< 0.60		
Mercury	(Hg)	0.11	< 0.40		
Platinum	(Pt)	< 0.003	< 0.005		
Thallium	(Tl)	< 0.001	< 0.002		
Thorium	(Th)	< 0.001	< 0.002		
Uranium	(U)	1.0	< 0.060		
Nickel	(Ni)	0.38	< 0.30		
Silver	(Ag)	0.13	< 0.18		
Tin	(Sn)	0.32	< 0.30		
Titanium	(Ti)	0.53	< 0.60		
Total Toxic Representation					

Hair Analysis: 6-week overview of what the body is EXCRETING.

Serum: Recent

RBC: approximately 3 months

WBC: approximately 6 months

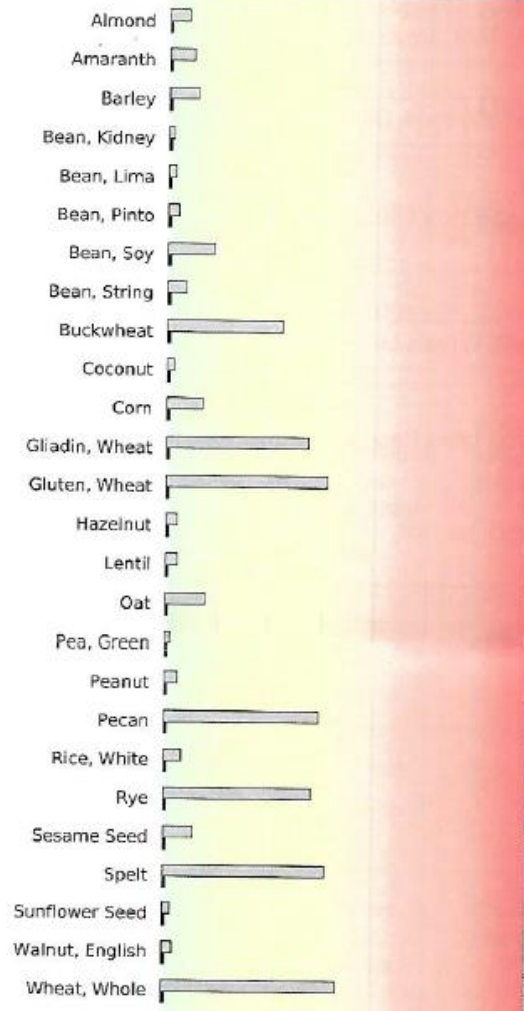
ESSENTIAL AND OTHER ELEMENTS								
		RESULT µg/g	REFERENCE INTERVAL	PERCENTILE				
				2.5 <sup>th</sup>	16 <sup>th</sup>	50 <sup>th</sup>	84 <sup>th</sup>	97.5 <sup>th</sup>
Calcium	(Ca)	1050	350- 1000					
Magnesium	(Mg)	240	35- 120					
Sodium	(Na)	390	18- 180					
Potassium	(K)	16	8- 75					
Copper	(Cu)	170	11- 37					
Zinc	(Zn)	190	150- 230					
Manganese	(Mn)	0.47	0.08- 0.60					
Chromium	(Cr)	0.44	0.40- 0.65					
Vanadium	(V)	0.13	0.020- 0.075					
Molybdenum	(Mo)	0.032	0.025- 0.060					
Boron	(B)	0.61	0.20- 1.2					
Iodine	(I)	0.66	0.25- 1.3					
Lithium	(Li)	0.013	0.007- 0.020					
Phosphorus	(P)	165	150- 220					
Selenium	(Se)	0.61	0.70- 1.1					
Strontium	(Sr)	2.4	0.86- 6.2					
Sulfur	(S)	44900	44000- 50000					
Cobalt	(Co)	0.23	0.005- 0.040					
Iron	(Fe)	8.1	7.0- 16					
Germanium	(Ge)	0.035	0.031- 0.040					
Rubidium	(Rb)	0.018	0.006- 0.060					
Zirconium	(Zr)	0.57	0.025- 0.50					

Patient: Chelsi R Ownby  
 Accession #: 201155514  
 Sex: F Age: 18  
 Date of Birth: 1994-03-11  
 Collected: 2011-10-03 Received: 2011-10-04 Completed: 2011-10-12 CLIA #: 50D0965661 © US BioTek Laboratories

IgG   
 IgE 

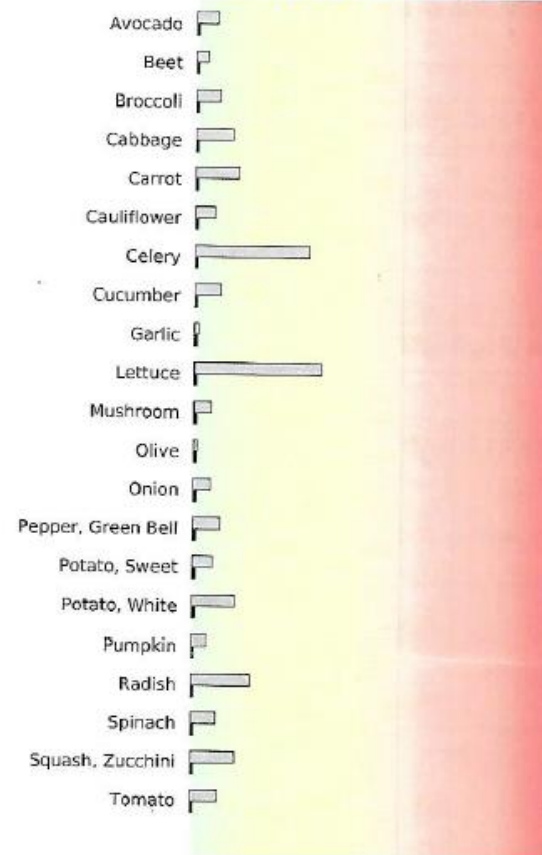
### Grains/Legumes/Nuts

0 I II III IV V VI



### Vegetables

0 I II III IV V VI



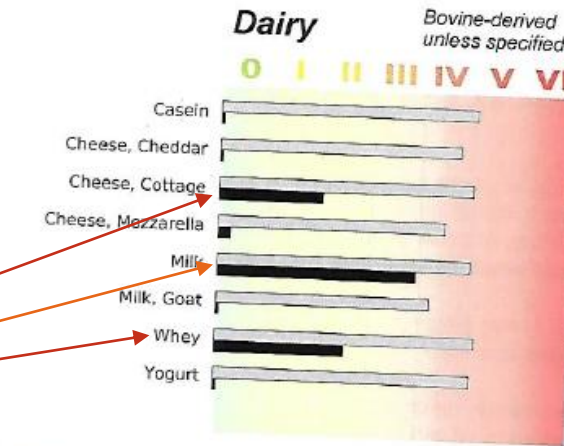
Leaky Gut!



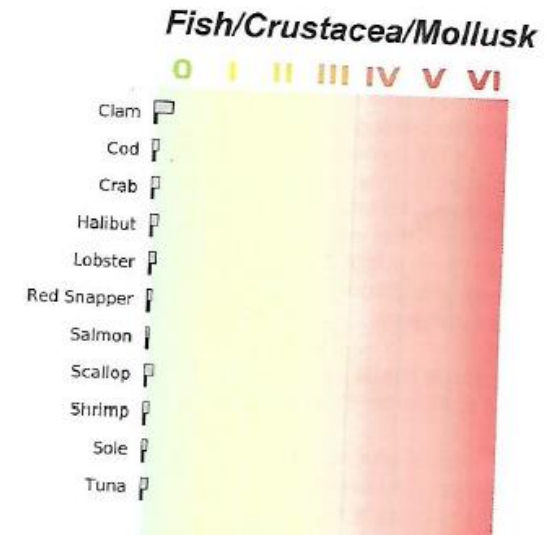
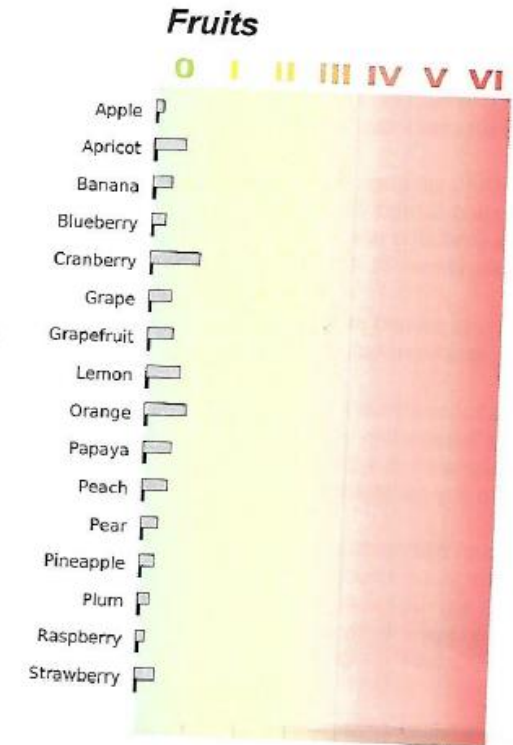
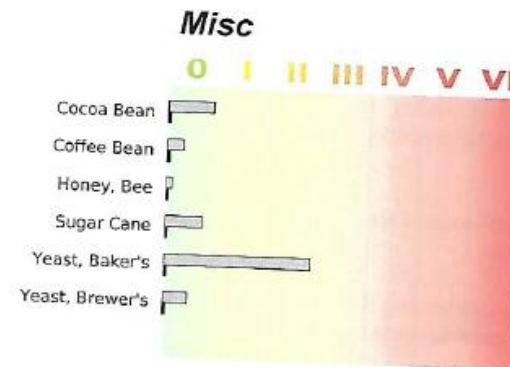
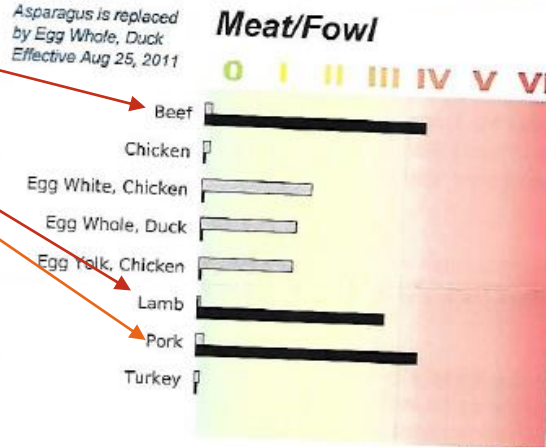


Histamine


Leaky Gut

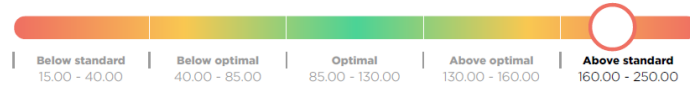



Asparagus is replaced by Egg White, Duck Effective Aug 25, 2011

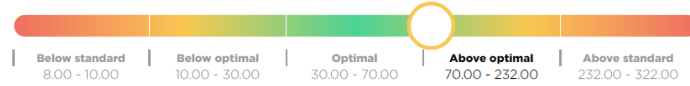


## IRON MARKERS

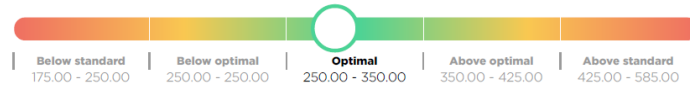
Iron - Serum   
213.00 µg/dL




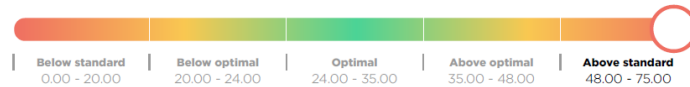
Ferritin   
79.00 ng/mL



TIBC  
285.00 µg/dL



% Transferrin saturation   
75.00 %



Review > [Nat Rev Microbiol. 2008 Jul;6\(7\):541-52. doi: 10.1038/nrmicro1930.](#)

# Viral infection and iron metabolism

Hal Drakesmith <sup>1</sup>, Andrew Prentice

Affiliations + expand

PMID: 18552864 DOI: [10.1038/nrmicro1930](#)

## Abstract

Fundamental cellular operations, including DNA synthesis and the generation of ATP, require iron. Viruses hijack cells in order to replicate, and efficient replication needs an iron-replete host. Some viruses selectively infect iron-acquiring cells by binding to transferrin receptor 1 during cell entry. Other viruses alter the expression of proteins involved in iron homeostasis, such as HFE and hepcidin. In HIV-1 and hepatitis C virus infections, iron overload could be partly caused by the viruses themselves. These interactions might suggest new mechanisms for iron overload.

iron overload is associated with poor prognosis and could be partly caused by the viruses themselves.

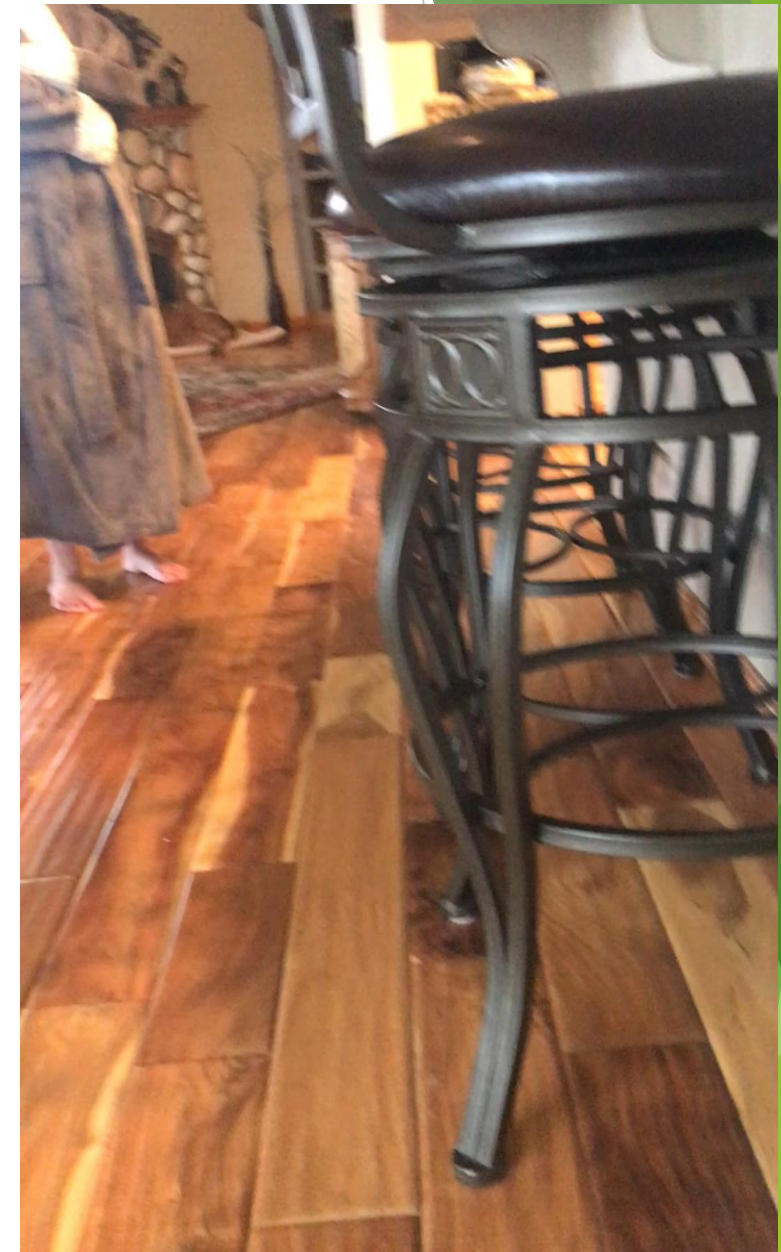
Similar articles

Drakesmith H, Prentice A. Viral infection and iron metabolism. *Nat Rev Microbiol.* 2008 Jul;6(7):541-52. doi: 10.1038/nrmicro1930. PMID: 18552864.

# 2019

- ▶ 2019
- ▶ **201 tonic clonic seizures**
- ▶ Wandering around after seizure got much worse. Seizures coming later & later.
- ▶ Frankincense & Copaiba essential oils tried in February. Had a situation where extra Frankincense was used and had involuntary "dancing" movements-she couldn't stop and was in distress (have video). Discontinued. Seizures started getting later (1am & 2am) and sometimes having 2 or 4 per night. 1<sup>st</sup> fall in over 2 years. Awake – standing on wood floor and went backward. Bizarre episode where she saw her belly in the mirror and had her 1<sup>st</sup> ever total freak out. Seizures started at ANY time of the night/morning 2:30am to 8:30am. Unprecedented! Things spun out of control this year, especially Oct to Dec. We have not been able to regain any normalcy (9:30 to 11pm) in seizures, even though we are back on doses of all 3 drugs same as 2015 . Looking into EMF's and Wifi (#Steelesarmy) and Dr. Jess Armine.

(c) Dr Jess P Armine (unless otherwise noted) 2022



February, 2020	
Week 1	9
Week 2	4
Week 3	4
Week 4	5
Total: 22	



March, 2020	
Week 1	6
Week 2	8
Week 3	7
Week 4	4
Total: 25	

April, 2020	
Week 1	7
Week 2	3
Week 3	6
Week 4	5
Total: 21	

May, 2020	
Week 1	6
Week 2	5
Week 3	6
Week 4	10
Total: 27	

June, 2020	
Week 1	6
Week 2	4
Week 3	6
Week 4	10
Total: 26	

July, 2020	
Week 1	0
Week 2	1
Week 3	5
Week 4	7
Total: 13	

August, 2020	
Week 1	6
Week 2	19
Week 3	5
Week 4	8
Total: 38	

September, 2020	
Week 1	4
Week 2	5
Week 3	8
Week 4	8
Total: 25	

October, 2020	
Week 1	3
Week 2	4
Week 3	3
Week 4	3
Total: 13	

November, 2020	
Week 1	1
Week 2	1
Week 3	1
Week 4	2
Total: 5	

December, 2020	
Week 1	1
Week 2	1
Week 3	1
Week 4	2
Total: 5	

January, 2021	
Week 1	1
Week 2	1
Week 3	1
Week 4	1
Total: 4	



Feb 2020 to Feb 2021

# FMP treatment started January 2020

- ▶ Foundational treatment consisting of liposomal vitamins and minerals.
  - ▶ Treatment for leaky gut syndrome.
  - ▶ Mitochondrial support.
  - ▶ Cell membrane support
- ▶ This patient progressed very slowly. The hundreds (maybe thousands) of seizures did significant damage to her central nervous system. There was no single root cause that we could identify so we concentrated on optimizing her cell function, biochemical pathways, and balancing her neurotransmitters.
  - ▶ It worked!



## Playing Cards 2021



Late 2021

- ▶ Chelsi is 27 years old and continues to progress albeit slowly.
- ▶ I want to give a shout out to her parents (and dog). They are the finest, most dedicated people I've had the pleasure of working with in my career.



Just simply-Thank You, for  
saving Chelsie's life!

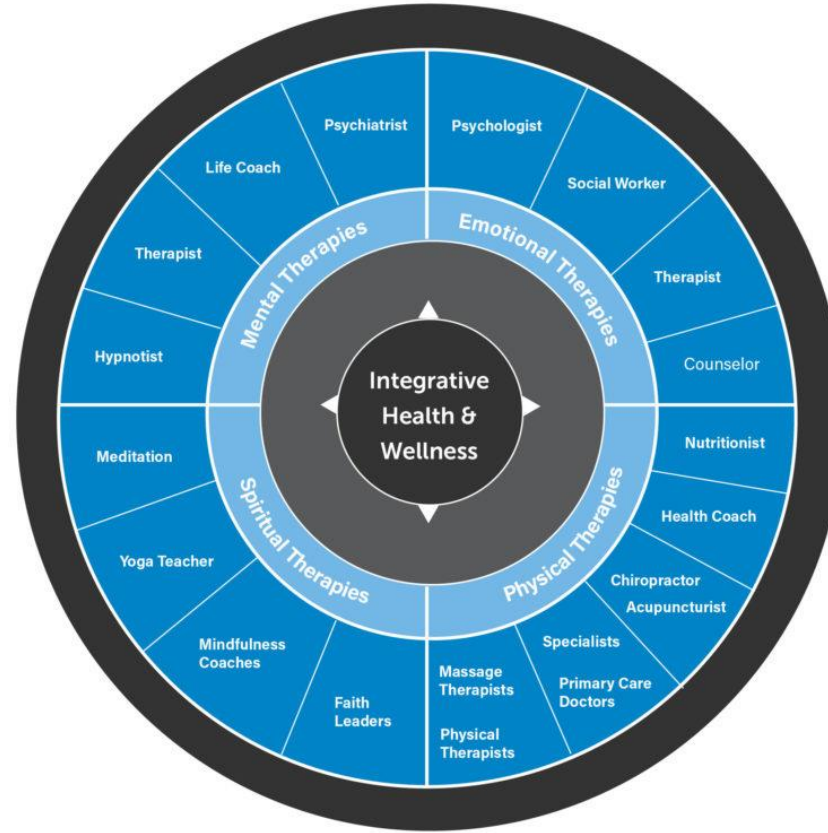
Love,

Sue & Chelsi





# *If we think differently and Work **TOGETHER***



## ***...we can change lives!!***

# Tips on Picking out a Practitioner





***“Listen to your patient, he is telling you the diagnosis”***

Sir William Osler, Bt  
Founder Father of Johns Hopkins Medical Center\*

\*Tuteur, Amy (November 19, 2008). [“Listen to your patient”](#). The Skeptical OB. Retrieved April 9, 2012.

***REMEMBER, In Real Estate, It’s “Location, Location, Location.”  
In Health Care it’s, “History, History, History!”***

# Words <sup>OF</sup> WISDOM



Treats the Patient... NOT the Test

# You can't do this alone....TIPS on How to pick a practitioner

- Certifications beyond their original degree / license
- An eclectic knowledge base so you are afforded true holistic care
- They're focused on finding root causes, the downstream effects, and creating an individualized manner of treating **YOU!** Not "your condition".
- They use lab tests appropriately and know how to **Interpret** (not just read) them. But most of all...
- **They will take time to listen and explain**



Interview the practitioner before committing yourself to their care.

**Assure yourself that he/she is as committed to your healing as you are and will take the time and effort to return you to life.**

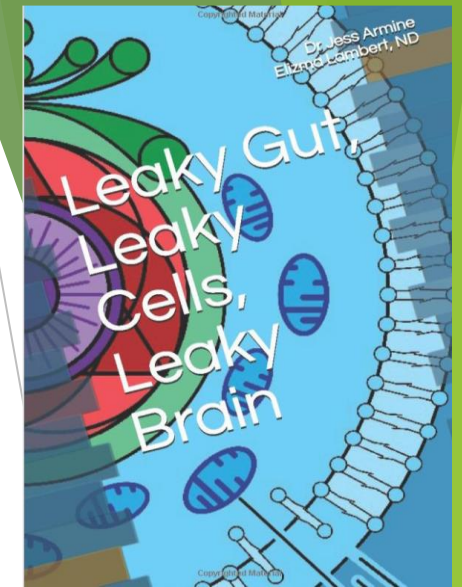


# Chronic Illnesses Take a Long Time to Heal.

- ▶ The magic pill does not exist.
- ▶ Working with a practitioner, embracing the healing process, and participating in your recovery and you **WILL HEAL!!!**

# Resources

- ▶ [Nutritionist \(ldnresearchtrust.org\)](http://Nutritionist(ldnresearchtrust.org))
- ▶ [Leaky Gut, Leaky Cells, Leaky Brain: Where to go when all hope is lost!: Armine, Jess, Lambert, Elizma: 9798747272231: Amazon.com: Books](#)
- ▶ USA: [Rupa Health | A simpler way to order specialty labwork.](#)
- ▶ UK/EU: <https://functionaldx.com>, [Welcome to Biolab Medical Unit London UK](#), [Regenerus Labs](#)





# Evidenced Based References

**Butyrate:** are important as food for cells lining the mammalian colon (colonocytes). Without butyrates for energy, colon cells undergo autophagy (self digestion) and die.<sup>1</sup>

<https://www.ncbi.nlm.nih.gov/pubmed/27346602>

<https://www.ncbi.nlm.nih.gov/pubmed/25875123>

**Honey:** Today, honey has been scientifically proven for its antioxidant, regulation of glycemic response, antitumor, antimicrobial, anti-inflammatory, and cardiovascular potentiating agent. It can be used as a wound dressing and healing substance.

<https://www.ncbi.nlm.nih.gov/pubmed/29101693>

**Zinc Carnosine:** *Involved in the reversal of neurodegenerative diseases, gastrointestinal conditions, antioxidant, metal chelating, anti-crosslinking, and anti-glycation activities*

<https://www.ncbi.nlm.nih.gov/pubmed/29382141>

<https://www.ncbi.nlm.nih.gov/pubmed/24247360>

<https://www.ncbi.nlm.nih.gov/pubmed/25846004>

**Sialic acid** are cytoprotectors, mainly localized on the surface of cell membranes with multiple and outstanding cell biological functions. Provides mucin for the GI tract.

<https://www.ncbi.nlm.nih.gov/pubmed/30509400>

- NEI:  
<https://www.neurorelief.com/index.php?p=cms&cid=108&pid=85&type=1>
- Brain Basics:  
<http://www.nimh.nih.gov/health/educational-resources/brain-basics/brain-basics.shtml>
- The Brain from Top to Bottom:  
[http://thebrain.mcgill.ca/flash/i/i\\_01/i\\_01\\_m/i\\_01\\_m\\_ana/i\\_01\\_m\\_ana.html](http://thebrain.mcgill.ca/flash/i/i_01/i_01_m/i_01_m_ana/i_01_m_ana.html)
- Neurotransmitters, An Introduction:  
<http://mybrainnotes.com/serotonin-dopamine-epinephrine.html>
- **Epigenetics of depression.** Lolak S, Suwannarat P, Lipsky RH. Prog Mol Biol Transl Sci. 2014;128:103-37. doi: 10.1016/B978-0-12-800977-2.00005-X. PMID: 25410543

- ▶ Cell Membrane Rap: [Cell Membranes Rap - YouTube](#)
- ▶ The Cell Song: [The Cell Song - YouTube](#)
- ▶ Mitochondria Song: [Electron Transport Chain \(Music Video\) - YouTube](#)
- ▶ Enzymes [Enzymes: Mr. W's Enzyme Song - YouTube](#)
- ▶ [Sciencemusicvideos.com](#)

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Dr Jess treats patients and mentors practitioners worldwide.

Dr Jess offers prospective patients a complimentary 30 min “get acquainted” session.



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“HEAL THE SICK,  
FEED THE HUNGRY,  
SHELTER THE WEARY,  
DEFEND THE WEAK”

