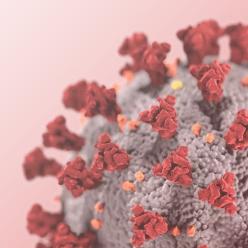
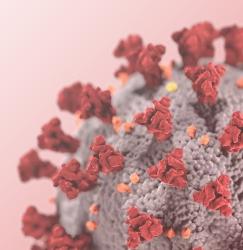
# Biomarkers in Spikopathy: Guiding Treatment

Presented by: Scott Marsland, FNP-C



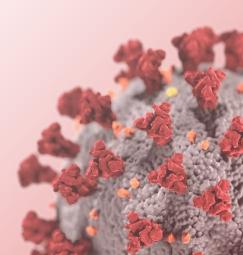
### **CONFLICTS OF INTEREST**

### None



### INTRODUCTION

- No commercially available measure of spike
- Clinical correlation of indirect measure
- Hypotheses re: pathophysiology
- Interventions with clinical benefit
- Implications for practice

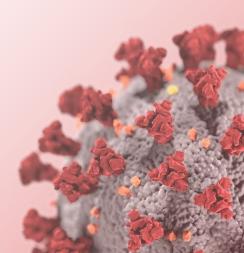


- Mid 30s, female
- Hx of depression, chronic back pain s/p MVC. S/p ACL repair, hand surgery.
- Rx Lexapro, Nuvaring, OTC Vit D, B-12 (Methyl)
- Physically active, passion for snowboarding, living at 9,000 ft altitude, working at 10,000 ft altitude.
- Single, gainfully employed, supportive extended family

- Initial visit was for acute COVID late June 2022
- Single shot J&J
- Treatment in our telemedicine practice, with foundation of FLCCC protocol at that time, including IVM, HCQ, as well as NTZ, Vit D, Quercetin, Zinc, Vit C, Nasopharyngeal/oral spray/gargle
- Initial sx: fever (T Max 101.4F), sore throat, HA, sinus congestion, fatigue, body/muscle aches, brain fog, low oxygen Sa02
- Progression into organizing pneumonia, and high dose Methylpredinsolone was initiated
- Patient was recovering, and abruptly stopped Methylprednisolone
- Uh-oh...

Gestalt of treating acute COVID and post-acute sequelae of COVID (PASC) and vaccine injury

- Complexity
- Brain fog is real: easily overwhelmed
- Close follow up, clear communication
- Citizen scientists: decisive role of patient engagement
- Active learning, flexible application by providers



#### **Clinical Course:**

- Moved from acute COVID directly into PASC, despite tx
- Recurrent acute viral infections, secondary bacterial infections, reactivated EBV, negative by 2/23/23
- Prolonged recovery after each acute episode
- Interval functionality, followed by worsening setbacks
- Gradual acknowledgement and growth towards resolve
- From 6/22 through 2/23, pertinent abnormal labs were elevated RBCs (high altitude), platelets, and low normal B-12 and D3, now at 412 pg/ml and 75.8 ng/mL, respectively
- Many trials of therapy, including: LDN, TA-1 (peptide), Aprepitant, ATP-360.

### **Critical Junctures, 1 of 3**

- Lingering impact on adrenal function from abrupt cessation of high dose steroids. Not Addisonian crisis, but not good.
- Emotional work of processing what is happening in her body, and coming to accept that it wont' resolve without an organized, strategic, layered approach
- Follow up, follow up, follow up: value of a highly professional, knowledgeable, communicative nursing team manifesting itself
- Dietary changes, including elimination of soda
- Breaking isolation with support online from other PASC/ vaccine injured patients.

### **Critical Junctures, 2 of 3**

- Early initiation of EPA: 30% of refractive mental health patients have underlying endothelial inflammation
- POTS symptoms: HR management with Midodrine.
- Measurement of reactivated EBV: initiation of Humacel and Lauricidin, transition to TA-1, incidental NTZ for acute sx.
- Emergence of microclotting as a dominant dynamic, with exertion and cold provoking escalating events of decompensation.
- Progression from anticoagulation with Turmeric, EPA, Aspirin and Nattokinase, to triple anti-coagulation with Plavix and Eliquis mid-January 2023

### **Critical Junctures, 3 of 3**

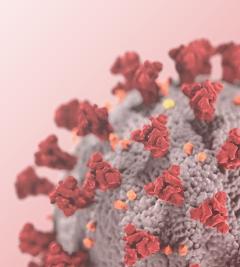
- Implementation of HBOT after triple anticoagulation, with relief of dyspnea, CP, decrease in joint/muscle pain in two weeks
- Identification of a clinically correlated measure: SARS-CoV-2-Semi quantitative antibody 23,192 U/mL
- Improvement of anxiety and cognition with introduction glutamate antagonist (Memantine) early February 2023
- Dynamic of living at high altitude, and coming down to sea level, TWICE!
- GAME CHANGER: Quantum NAC. Back to new normal and rapid, quantifiable drop in spike ab dilution after three weeks.



### EMERGING APPROACHES TO TREATING SPIKE PROTEIN-INDUCED DISEASES

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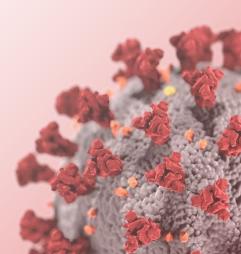




### EMERGING APPROACHES TO TREATING SPIKE PROTEIN-INDUCED DISEASES

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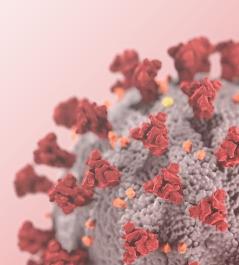




### Dr Gi-Gi and me:

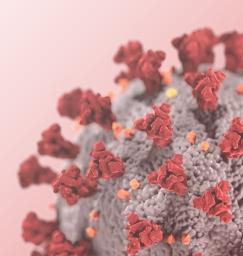
- Medicine and Psychiatry
- Data on >200 patients
- Emerging patterns since December 2022





### **PATHOPHYSIOLOGY**

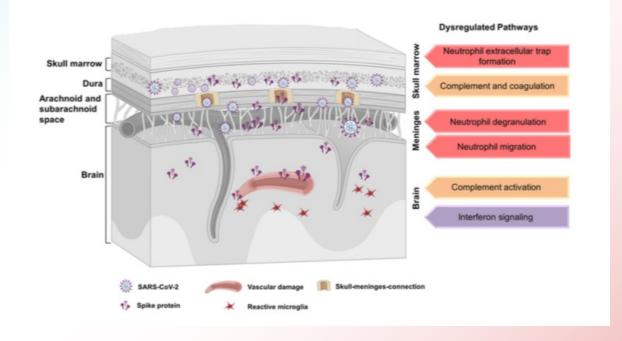
- Brain vs body immunity
- Spike and BBB
- IVM and BBB
- Glutamate and GABA
- Spike and Tau
- EBV

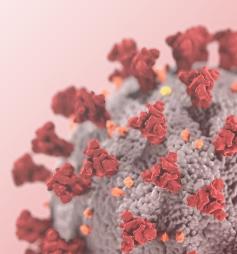


### EMERGING APPROACHES TO TREATING SPIKE PROTEIN-INDUCED DISEASES

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### **Graphical Summary**

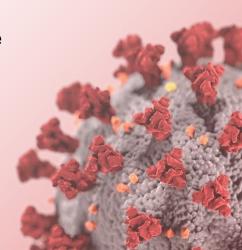




### **GLUTAMATE**

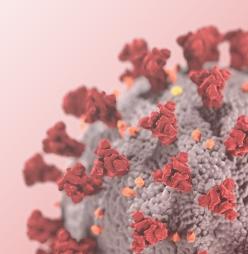
Glutamate is an abundant, powerful excitatory neurotransmitter present almost in all <u>neural</u> synapses in the <u>central nervous</u> <u>system</u> and plays an <u>essential</u> role in the functioning of the <u>brain</u>.

Glutamate levels need to be well regulated, as excessive exposure to glutamate may be toxic to neurons and may damage them leading to ALS.

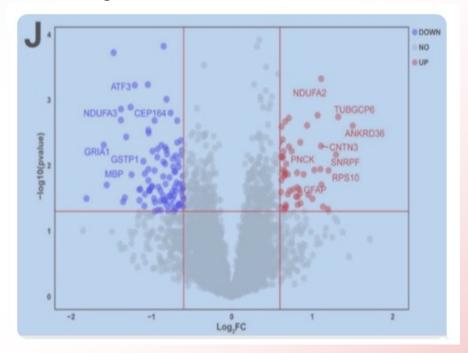


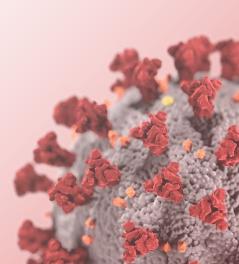
### PATHOPHYSIOLOGY II

- Glutamate channels
- GABA & myelin inhibition
- Neuronal instability, apoptosis



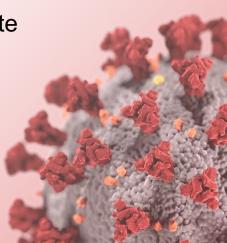
### GRIA1 Downregulated Post-mortem Brains: Mediates Glutamate





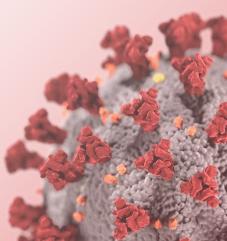
## PATHOPHYSIOLOGY MEETS PHARMACOLOGY: MEMANTINE / NAMENDA

- Low-affinity, voltage dependent, uncompetitive
- NMDA Receptor antagonist
- Interacts under pathologic conditions of >Glutamate
- Commonly used in Psychiatry and Medicine



## PATHOPHYSIOLOGY MEETS PHARMACOLOGY: MEMANTINE / NAMENDA

- Primary side effect is sleepiness
- Primary adverse effects: dizziness, bad dreams and sleep disruption
- Few drug interactions
- May block: nicotinic, acetylcholine, serotonin and sigma-1 receptors
- Inexpensive: ~\$3 for one month supply
- Low dose: up to 5mg daily



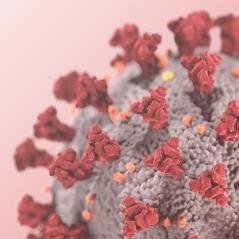
### PATHOPHYSIOLOGY MEETS NUTRACEUTICAL: SHILAJIT / MOOMIYO / FULVIC ACID

- Ayurvedic, in use 3,000 years
- Dark, sticky mineral substance that exudes from the rocks of high mountainous regions during the heat of summer
- Glycine and GABA mimetic actions
- Prevents Tau formation, neural anti-inflammatory, mitochondrial support, increases iron levels, mild diuretic, adrenal support,> testosterone production
- 150mg → 1500mg daily



"If you care, you'll learn one thing from another." Barbara Kingsolver





### **SPIKE VS SPIKE**

### Spike from community infection

- Nucleocapsid
- Lower concentration
- Must pass mucosa

### Spike from injection

No nucleocapsid
Up to 1000x concentration
Bypasses mucosa, becoming
systemic
Cleavage increases affinity

for ACE2

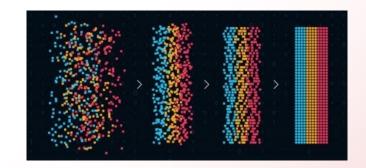
### SPIKE VS SPIKE ANTIBODY

### **Spike**

- Cannot yet be measured with commercially available testing.
- Point-of-care testing in Italy, not brought to market

### **Spike Antibody**

\$100 test from Lab Corp
AND...our data shows clinical
correlation to symptom
burden



### PATTERN RECOGNITION

- In the absence of spike neutralizing efforts, provides some measure of global background spike
- In the presence of spike neutralizing efforts, points towards a reservoir of spike in the brain
- Consistently decreases with intervention
- In symptomatic patients: symptom burden decreases in direct correlation
- Rate of decrease flattens



#### **EMERGING APPROACHES TO TREATING** SPIKE PROTEIN-INDUCED DISEASES

April 28-29, 2023 • Fort Worth, Texas

• SARS CoV-2 Antibody Profile, Nucleocapsid and Spike CPT: Test

code: 160236 and 86769x2

Labcorp Order Code: 160236 and 86769x2

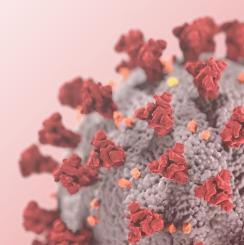
Test code: 160236; 86769x2 Non-fasting, LAB CORP ONLY!

### WHAT IS THE TEST?

#### Ordered Items: SARS-CoV-2 Antibody, IgG; SARS-CoV-2 Semi-Quant Total Ab; Venipuncture

ate Collected: 01/25/2023	Date Received: 01/25/2023	Date Reported: 01/26/2023		Fasting: Yes		
ARS-CoV-2 Antibody, IgG						
Test	Current Result and Flag	Previous Result and Date	Units	Reference Interval		
SARS-CoV-2 Semi-Quant IgG Ab	>800.0		AU/mL	Neg <13.0		
SARS-CoV-2 Spike Ab Interp <sup>A ex</sup>	Positive An Indicate against the SARS-CoV-2 spike protein, including the receptor binding domain (RBD) were detected. It is not yet known what level of antibody to SARS-CoV-2 spike protein correlates to immunity against developing symptomatic SARS-CoV-2 disease.  This assay was performed using DiaSorin Liaison(R) SARS-CoV-2 Trimerie'S IGG assay.					

Test	Current Result and Flag	Previous Result and Date	Units	Reference Interval
SARS-CoV-2 Semi-Quant Total Ab <sup>A, 01</sup>	See Dilution		U/mL	Negative<0.8
SARS-CoV-2 Spike Ab Dilution	8680		U/mL	Negative<0.8
SARS-CoV-2 Spike Ab Interp <sup>A+1</sup>	domain (RBD) were detected. antibody to SARS-CoV-2 spik- developing symptomatic SARS- measure the quantitative le- following vaccination. Such	CoV-2 spike protein receptor It is yet undetermined what protein correlates to immun CoV-2 disease. Studies are u rels of specific SARS-CoV-2 a studies will provide valuable protection from vaccination	level of ity against nderway to ntibodies e insights	



### TIPS FOR SUCCESS

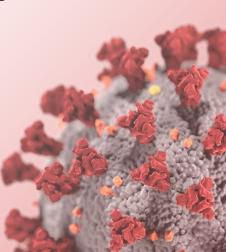
- <u>Lab Corp only</u> (Boston Heart option)
- NOT Quest, NOT local hospital
- Lab Corp will check semi-quant, and if >250 U/mL, will perform a reflex dilution level
- Levels can vary between <0.8 U/mL and >25,000 U/mL
- Patient doesn't need to be fasting

### **INTERPRETATION: PART 1**

- Levels > 400 U/mL often correlate to symptoms, including HA, fatigue, and cognitive impairment
- Levels >5,000 U/mL can correlate to severe changes in mental health and neurological function
- Levels >25,000 U/mL are out of bounds, and concerning
- Unvaccinated patients with levels up to 12,000 U/mL
- Very low or absent levels in decompensated patients may represent a severely compromised immune system

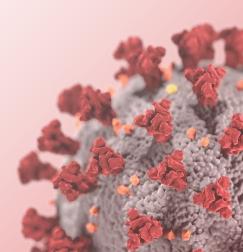
### **INTERPRETATION: PART 2**

- Clinical context is paramount: point 0.0 is testing symptomatic patients
- Early consideration needs to be given to close contacts
- Asymptomatic close contacts with elevated spike ab dilution levels are consistent with "bad batches" and individual autoimmune response
- Spike ab levels in close contacts decrease with spike neutralzing treatment
- This benefits the patient
- And points to...shedding as an real phenomenon



### **TREATMENT**

- Baseline spike neutralization with intermittent fasting (IF), IVM, Nattokinase, Resveratrol
- Add Spermidine to promote autophagy
- Spike ab <1,000 U/mL, consider Shilajit</li>
- Spike ab > 2,000 U/mL, trial Memantine
- Quantum NAC...a game changer?



### **CLINICAL PEARLS**

START MEMATINE

1.25MG

ADVANCE MEMANTINE

5MG

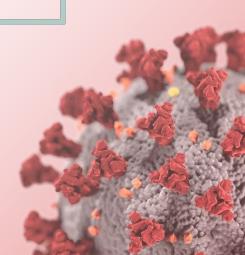
ANTICIPATE
Spike ab >25k?
2-3 months tx

### **STOP MEMANTINE**

Dizziness, bad dreams, sleep interruption, lethargy, abrupt downturn

PLAN TO TRANSITION

Shilajit on deck



### WHAT DO WE SEE?

### DECREASING SPIKE AB

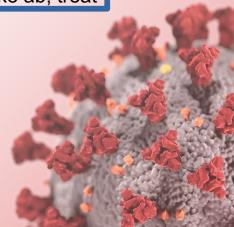
- 2,000 U/mL per month max
- Associated with full interventions &
  - Aggressive IF

## CLOSE CONTACTS (CC)

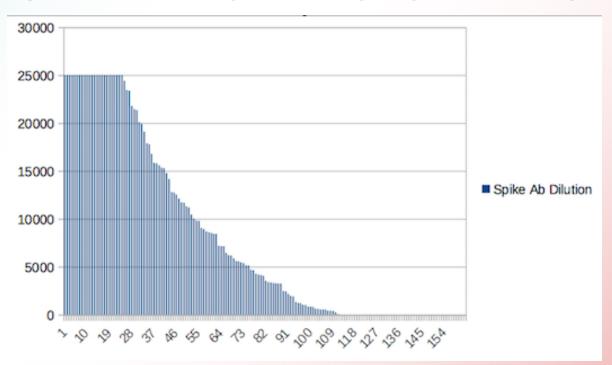
- Signals of shedding
- Patient plateau, check cc
- With > spike ab, treat

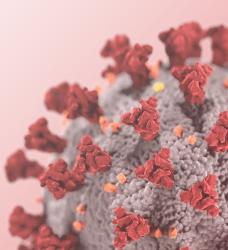
#### RAPID IMPROVEMENT

- Acute sx burden with elevated spike ab
- Reversal of suicidality, homicidality, severe anxiety



### SPIKE ANTIBODY DILUTION LEVELS

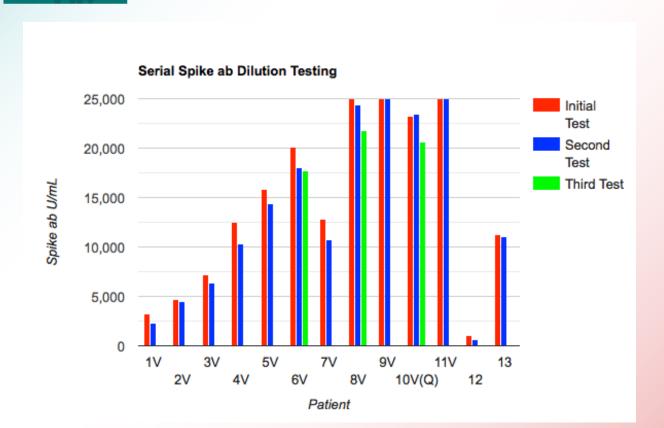


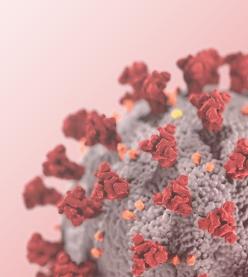




### EMERGING APPROACHES TO TREATING SPIKE PROTEIN-INDUCED DISEASES

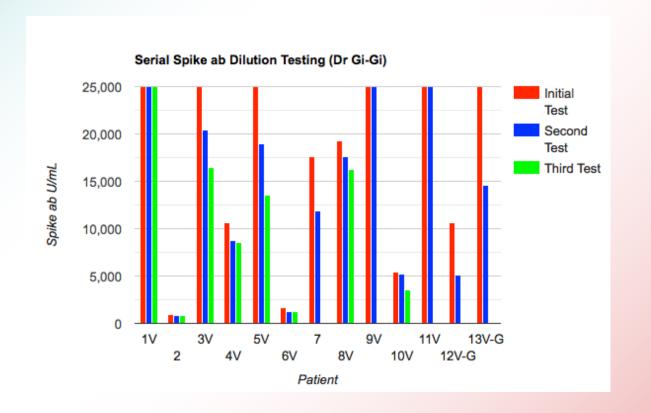
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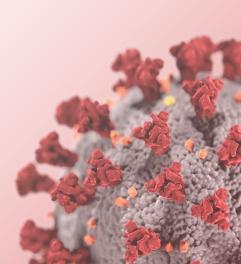




### EMERGING APPROACHES TO TREATING SPIKE PROTEIN-INDUCED DISEASES

April 28-29, 2023 • Fort Worth, Texas





### **EXAMPLES OF SHEDDING**

4,000 U/mL

11,000 U/mL

?

12,000 U/mL

Unvax patient has vax/Boosted spouse has Spike ab 23,500 U/mL

Unvax daughter lives with her works in criminal court

Unvaccinated spouse performed felatio upon vax/boosted spouse

Unvaccinated patient teaches vax/boosted band students

?

5,000 U/mL

500 U/mL

>25,000 U/mL

Snuggled skin-to-skin with friend just boosted

Unvax visited mother in nursing home every day for 1 hour for months

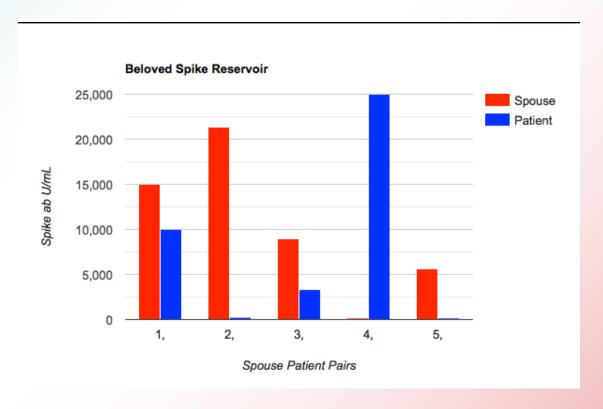
Unvax cared for vax mother at home providing intimate care

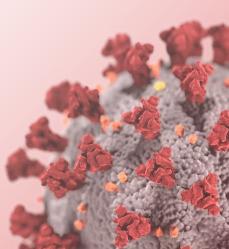
Vax patient living with vax partner



### EMERGING APPROACHES TO TREATING SPIKE PROTEIN-INDUCED DISEASES

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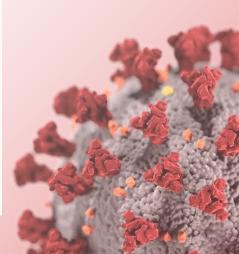


## VAX, SPIKE AB ~4K

#### **BEFORE MEMANTINE**

- Low testosterone
- Low libido
- Erectile dysfunction
- Brain fog
- Poor memory

- Improved libido
- Improved erectile function
- Better energy
- Decreased brain fog
- Improved memory

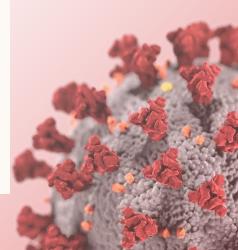


## VAX, SPIKE AB ~5K

#### **BEFORE MEMANTINE**

- Severe back pain
- Nerve pain and weakness in arm and legs
- Difficulty walking

- Improved sleep
- Resolved neuropathic pain
- Resolved brain fog

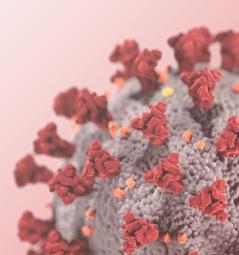


## VAX, SPIKE AB ~7K

#### **BEFORE MEMANTINE**

- Fatigue
- Brain fog
- Memory problems
- Tachycardia

- Improved brain fog
- Improved joint pain
- Reduced paresthesia
- Improved cognition

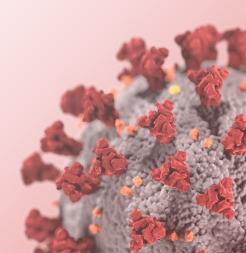


## VAX, SPIKE AB ~18K

#### **BEFORE MEMANTINE**

- Brain fog
- Memory problems
- Dizziness when standing
- Tachycardia
- Depression
- Anxiety
- Headache
- Insomnia

- Resolved HA
- Improved sleep

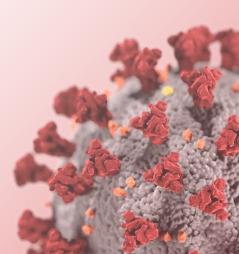


## VAX, SPIKE AB >25K

#### **BEFORE MEMANTINE**

- Brain fog
- Poor sleep
- Lower back pain
- Tinnitus

- Overall feels better
- Less fatigue
- Improved sleep

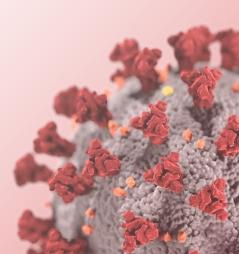


## VAX, SPIKE AB >25K

#### **BEFORE MEMANTINE**

- Numbness and temperature dysregulation in legs
- Poor circulation
- Severe leg pain and cramping
- Stinging in fee

- Less numbness
- Less stinging in feet
- Less hip pain
- Less calf pain
- Overall pain improvement

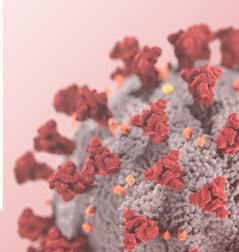


# VAVX, Spike ab > 25 K

#### **BEFORE MEMANTINE**

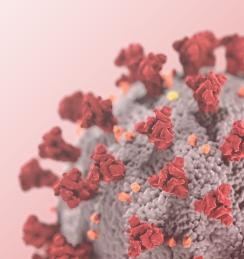
- Neuropathy
- Poor sleep
- Lower energy

- Improved sleep
- Improved energy
- Overall improvement



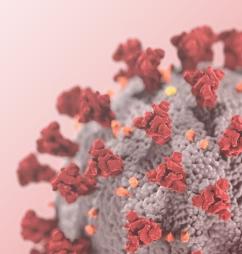
### **IMPLICATIONS FOR PRACTICE**

- Mental health acuity
- Autophagy
- Intermittent fasting
- Layered therapeutics
- Persistence
- Shedding
- Close contacts



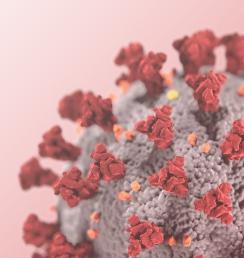
### **EMERGING THERAPEUTIC OPTIONS**

- Quantum NAC www.zerospike.org/en/
- Glutathione IV
- Ashawaganda
- GABA supplementation
- Ozempic (GLP-1 Peptide)
- Mounjari (GLP-1 Peptide)



## **LINGERING QUESTIONS**

- Mechanism of action in brain to lower spike?
- Does auto catalytic amplification explain shedding?
- What are the reasons for variability among individuals?
- What is the impact of depressed immune function on lab values?
- Does adipose tissue represent a spike sink?
- Why do some patients with elevated spike ab not respond well to Memantine or Shilajit?



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## **THANK YOU**

