# THE FUNCTIONAL **MEDICINE APPROACH TO LONG COVID**





## OUTLINE

- Introduction
- Long COVID current landscape
- Risk factors, mechanisms, pathophysiology
- Post Viral Recovery Program implementation
- Nutritional supplement selections
- Patient registry outcomes: early data
- Case studies
- Q&A





## LONG COVID PREVALENCE 1

Most Long COVID cases are cases with mild acute illness
1/3 of people with Long COVID have no identified pre -existing conditions

- 104,538,730 reported
   >200,000,000 reported + unreported estimate
- 15-30% people have persistent symptoms @1 month
- 6-10% people have persistent symptoms @3 months
- 1-2% people have persistent symptoms @12 months

## Long COVID Incidence

- ~ 30 million people with Post-COVID @ 1 month
- ~ 10 million people with Post-COVID @ 3 months
- ~ 2 million people with Post-COVID @ 12 months

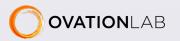




## Current Landscape: 3 Years Into PASC

- Confusing terminology: Chronic COVID, Post Viral Syndrome, Long -Haul COVID, Long
   COVID, and more.
- 200+ symptoms
- No agreed upon definitions: CDC 30 -days; WHO 90-days post-acute infection
- No proven treatments, no standards of care = no guidelines and no trained workforce
- Providers and patients frustrated where to turn for guidance and help?
- NIH RECOVER: research and clinical trials- \$1.2 billion
  - Largely observational studies: no findings published
  - 5 clinical trials planned (2600 total patients, 25-100 sites/study)
  - Drugs (Paxlovid), CBT, exercise, rehab
  - Not a single patient enrolled to-date





## **MORE IS NEEDED**

## A FUNCTIONAL MEDICINE APPROACH





## **Project Timeline**

November 2021 Expert working group assembled February 2022 Social listening study completed mining > 1 million conversations

July 2022 Selected outcomes measures and developed Timeline & Progression of symptoms

September 2022 Finalized specialty supplement recommendations to address 5 drivers of Long COVID.

December 2022 First patients enrolled in virtual group visit program patients. at Hennepin

February 2023 Enrolled first And Health "VIP"

May 2023 registered for the **PVRP** education &

#### January 2022

Prototype rootcause framework &standards for evaluating and treating long COVID

#### April 2022

Presented research to support 5 primary drivers of long COVID

August 2022 Presented 90-day program and protocol (beta)

October 2022 Contracted with NUNM to develop **REDCap** patient registry

#### January 2023

First patients enrolled from Mayo Clinic

#### April 2023

- 100+ patients enrolled in registry with first patients completing 90-day protocol.
- Outcomes data analyzed.
- Completed development of 3part provider education series





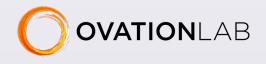
## **Expert Clinical Working Group**



# POST VIRAL RECOVERY IN THE COVID ERA

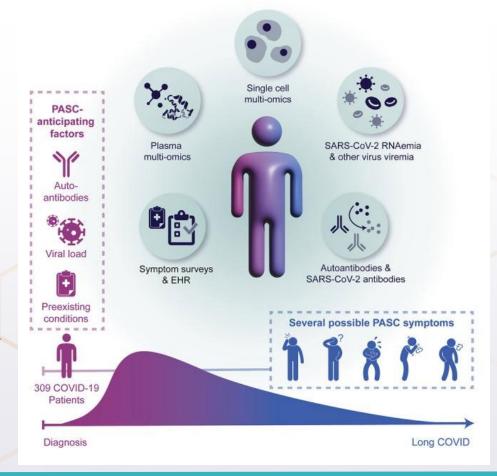
ASSESSING PROGRESSIONAND PERSONALIZING TREATMENT USING A ROOT-CAUSE APPROACH





## Multiple early factors anticipate post-acute COVID-19 sequelae

Su, Y, Yuan, D, et. al. (2022). Multiple early factors anticipate post-acute COVID-19 sequelae. Cell, 185(5). <u>https://doi.org/10.1016/j.cell.2022.01.01</u> <u>4</u>

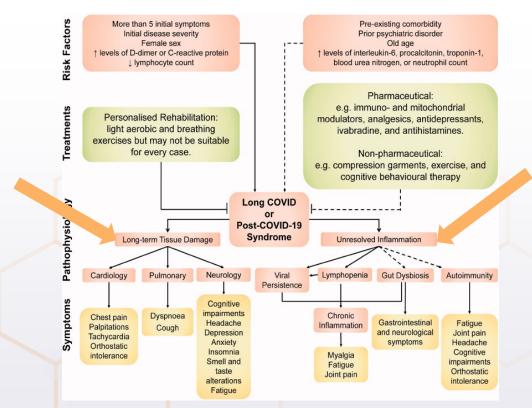






Long COVID or post-COVID-19 syndrome: putative pathophysiology, risk factors, and treatments

Yong, SS (2021). Long COVID or post-COVID-19 syndrome: putative pathophysiology, risk factors, and treatments. Infectious Diseases, 53(10), 737–754. https://doi.org/10.1080/23744235.2021.1924397



An overview of the symptoms, putative pathophysiology, associated risk factors, and potential treatments involved in long COVID. Note: Dashed lines represent areas where evidence is relatively lacking compared to non-dashed lines. (Color online only).

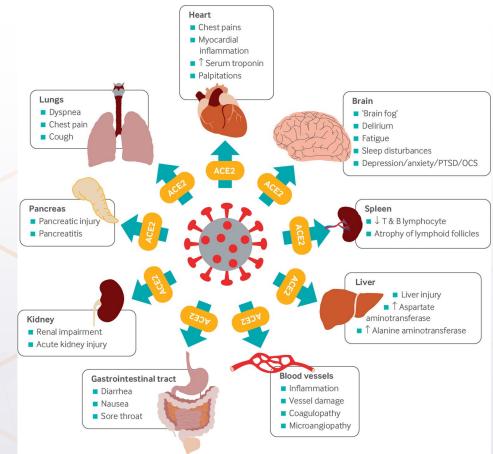




Long covid mechanisms, risk factors, and management

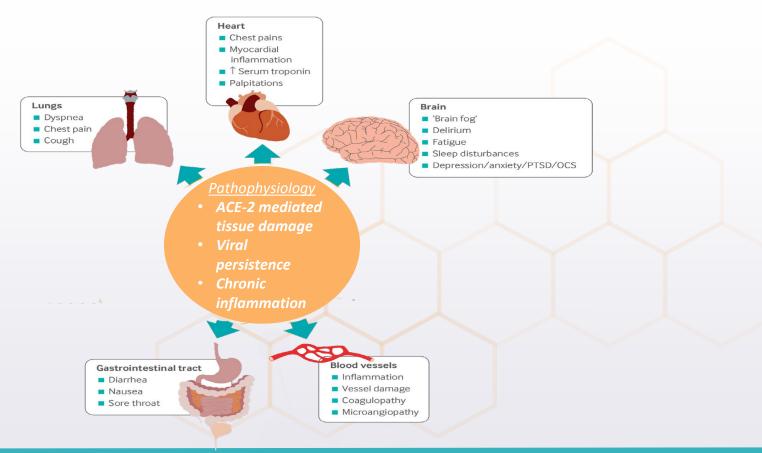
Crook, H, Raza, S, Nowell, J, Young, MK, Edison, P (2021). Long covid—mechanisms, risk factors, and management. BMJ, n1648. <u>https://doi.org/10.1136/bmj.n1648</u>

Multi-organ complications of covid-19 and long covid. The SARS-CoV-2 virus gains entry into the cells of multiple organs via the ACE2 receptor. Once these cells have been invaded, the virus can cause a multitude of damage ultimately leading to numerous persistent symptoms.



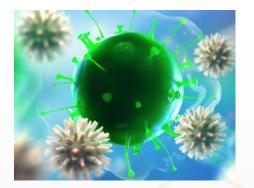




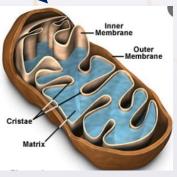








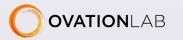
ACE-2 mediated damage Viral persistence Chronic inflammation Gut/Microbiome dysbiosis Mitochondrial dysfunction





Bacteria

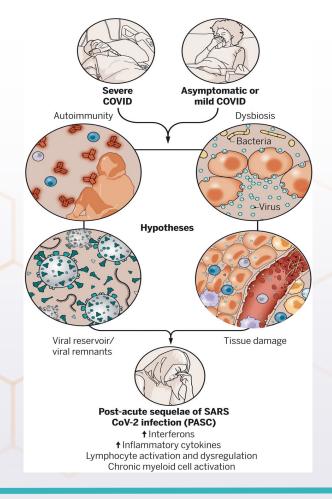
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## Distinguishing features of Long COVID identified through immune profiling

Klein, JB, Wood, JR, Iwasaki, A, et al. (2022). Distinguishing features of Long COVID identified through immune profiling. medRxiv (Cold Spring Harbor Laboratory). https://doi.org/10.1101/2022.08.09.22278592

A fraction of COVID-19 patients with either severe or mild COVID-19 develop a variety of new, recurring, or ongoing symptoms and clinical findings 4 or more weeks after infection. Analyses of immune responses in people with PASC reveal key inflammatory cytokines and cellular activation phenotypes that are significantly elevated over nonPASC convalescent controls. Further studies are needed to identify the drivers of PASC pathophysiology. Illustration: V. Altounian/Science







## Long COVID or Post-acute Sequelae of COVID-19 (PASC): An Overview of Biological Factors That May Contribute to Persistent Symptoms

Proal, AD, VanElzakker, MB. (2021b). Long COVID or Post-acute Sequelae of COVID-19 (PASC): An Overview of Biological Factors That May Contribute to Persistent Symptoms. Frontiers in Microbiology, 12. <u>https://doi.org/10.3389/fmicb.2021.698169</u>

## Long COVID EndoTypes\*

- 1. Multi-Organ Tissue Damage, 2° to acute infection
- 2. Persistent SARS-CoV-2 Infection
- 3. Reactivation of Neurotrophic Pathogens (e.g. HSV, HHV-6, EBV, etc.)
- 4. Microbiome/ Virome Dysregulation

- 5. Autoantibody Production with Molecular Mimicry
- 6. Dysregulated Brainstem & Vagal Nerve Signaling
- 7. Activation of Primed Immune Cells [Hyperinflammation]
- 8. Clotting/ Coagulation Vascular Issues [ACE2]





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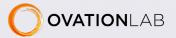
# ACE-2 mediated tissue damage

Chronic Inflammation

Gut/Microbiome dysbiosis Mitochondrial dysfunction

**Viral Persistence** 





## Long COVID: Symptoms



Long COVID / NIH COVID-19 Research. (n.d.). NIH COVID-19 Research. https://covid19.nih.gov/covid-19-topics/long-covid





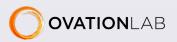
## ACE2-Driven Tissue Damage AND/OR Inflammation/ Immune Dysregulation

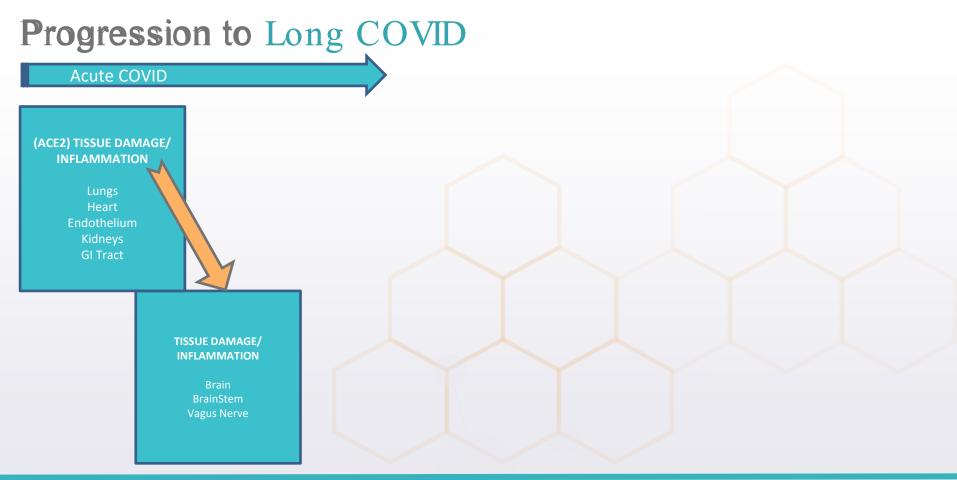
# ACE2 TISSUE DAMAGE

#### **IMMUNE DYSREGULATION**

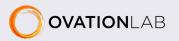
#### \*Chronic Inflammation



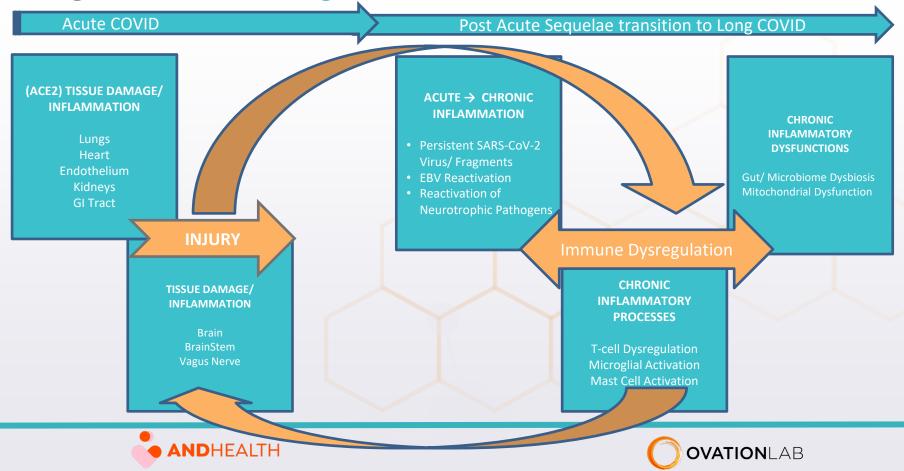


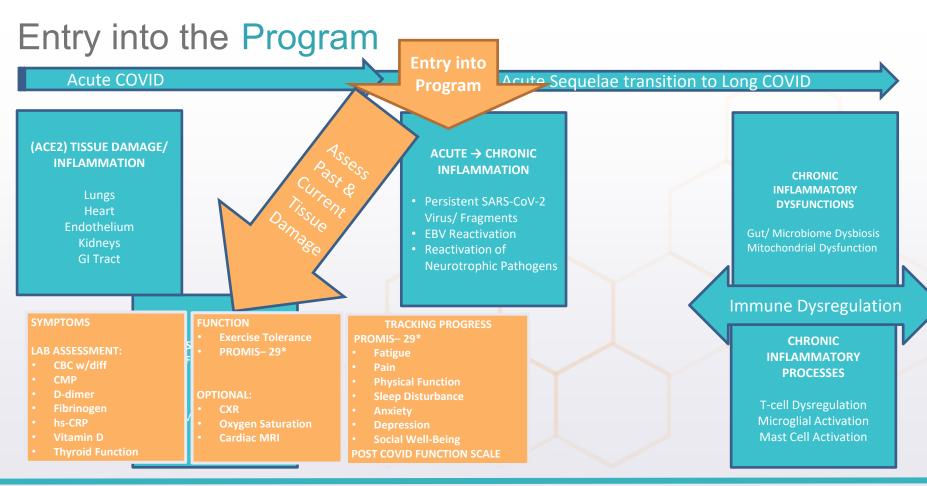






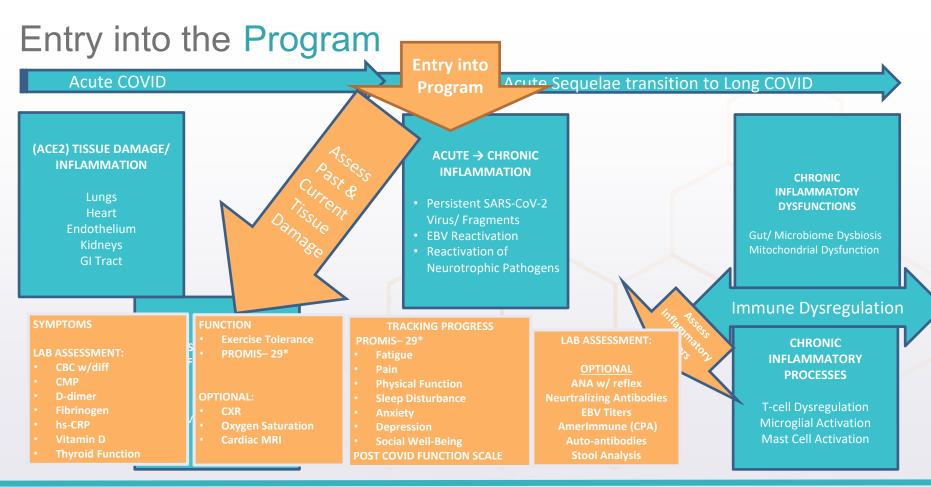
## Progression to Long COVID















## Long COVID Protocol

## ENROLLMENT

- Health History
- Timeline & Progression of Symptoms
- Previous Testing (if available)
- Evaluation of Function (PROMIS-29\*)
- Recovery Goals
- PHASE 1: 90-Day Program
  - Lifestyle Support
  - Baseline Supplements

## LIFESTYLE:

Food & Nutrition
Movement & Exercise
Stress Modification
Social Connection
Sleep
Nutritional Supplementation





## Long COVID Protocol PHASE 1

#### <u>Specialty Nutritional</u> <u>Supplements</u>

Selected to address the primary drivers of long COVID

#### Foundational Supplements

Selected to support optimal health and wellbeing



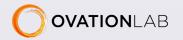


## Long COVID Patient Journey

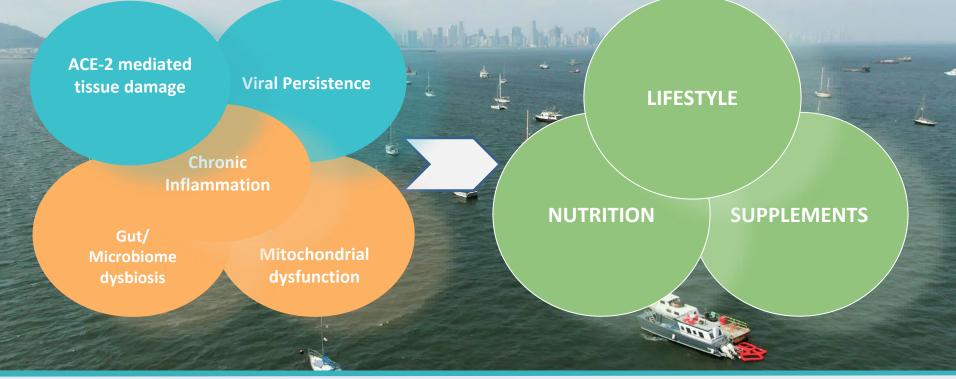
Follow - Up: 30, 60, 90-day patient registry surveys

- Updated timeline and progression of symptoms at 30 -day intervals
- Adherence to food plan, nutritional supplements, and lifestyle recommendations
- PROMIS-29\*
- Post COVID Function Scale (PCFS)





## A Rising Tide Lifts all Boats







# KEY LIFES TYLE INTERVENTIONS OVERVIEW







## LIFESTYLELEARNING OBJECTIVES



Understand the benefits of specific lifestyle factors for recovery from post-viral conditions



Review resources for plant forward nti-inflammatory diet and how to advise using them with participants



Become familiar with the lifestyle handouts provided for sleep, stress, movement, and community



Understand how to implement lifestyle tools clinically as part of the Post - Viral Recovery Program



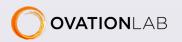


## KEY LIFES TYLE INTERVENTIONS OVERVIEW

- Food & Nutrition
- Sleep
- Movement & Exercise
- Stress Modification
- Social Connection
- Nutritional Supplementation







## WHY LIFESTYLE?

- In a prospective cohort study of 2000 women. Adherence to a healthy lifestyle prior to COVID- 19 infection was inversely associated with risk of Post- COVID Conditions (PCC) in a dose-dependent manner.
- Compared with those who did not have any healthy lifestyle factors, those with 5 or 6 had half the risk of PCC.



Wang S, Li Y, Wang S, Li Y, Yue Y, et al. Adherence to Healthy Lifestyle Prior to Infection and S, Li Y, Yue Y, et al. Adherence to Healthy Lifestyle Prior to Infection and Risk of Post– COVID-19 Condition. JAMA Intern Med. 2023;183(3):232–241.





- Including plant-based foods, colorful and nutrient dense, with high quality protein and fats.
- Eliminating processed foods and drinks.







- Plant-based nutrition is associated with less hospitalization and death from acute covid.
- And greater recovery from acute and post covid.

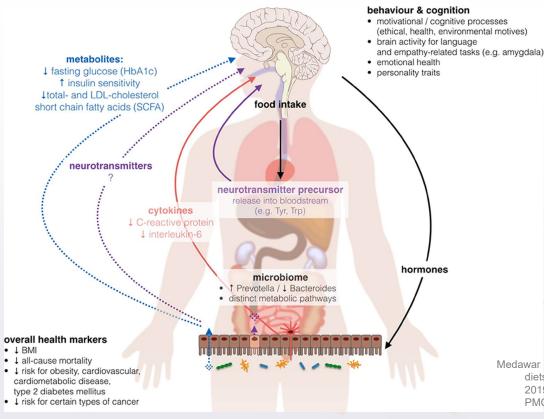
Kim H, Rebholz CM, Hegde S, LaFiura C, Raghavan M, Lloyd JF, et al. Plant-based diets, pescatarian diets and COVID-19 severity: a populationbased case–control study in six countries. BMJ Nutrition, Prevention & Health. 2021 May 18;

Storz MA. Lifestyle Adjustments in Long-COVID Management: Potential Benefits of Plant-Based Diets. Curr Nutr Rep. 2021 Dec;10(4):352-363. doi: 10.1007/s13668-021-00369-x. Epub 2021 Sep 10











Medawar E, Huhn S, Villringer A, Veronica Witte A. The effects of plant-based diets on the body and the brain: a systematic review. Transl Psychiatry. 2019 Sep 12;9(1):226. doi: 10.1038/s41398-019-0552-0. PMID: 31515473; PMCID: PMC6742661.





REVIEW THE PLANT-BASED WHOLE30 RULES

#### The Plant-Based Whole30 Program

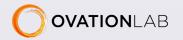
Empowering you to change your health, habits, and relationship with food on your terms. Let's do this.

#### TAKE THE FIRST STEPS



Improve your energy, sleep, cravings, mood,





Resources in the package include, among others

- Program Rules >
- Plant-Based Whole30 Prep Pack >
- Grocery Guide >
- Shopping List >
- Meal Planning >
- Plant-Based Recipes >







## FOOD & NUTRITION

### TIPS for Coaching on Eating for Post Viral Recovery

- Personalize to individual GI symptoms, intolerances and budget
- Use a health coach or nutrition ist
- Advise on time restricted eating
- Rx low glycemic diet
- Optimize fermented foods and prebiotics
- Reduce histamines if symptoms correlate
- Suggest spices for added benefit & taste







## FOOD & NUTRITION: Additional Resources

IFM Lifestyle Course Tools

- Eating a Balanced Diet
- Micronutrients: Phytonutrients
- Mindful and Intuitive Eating
- The Core Food Plan Comprehensive Guide
- Phytonutrient Food Plan Comprehensive Guide







### SLEEP

"COVID-19 and sleep disorders can induce BBB leakage via neuroinflammation, which might contribute to the 'coronasomnia' phenomenon. The new studies suggest that the control of sleep hygiene and quality should be incorporated into the rehabilitation of COVID-19 patients."

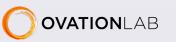
Coiro MJ, Asraf K, Tzischinsky O, Hadar-Shoval D, Tannous-Haddad L, Wolfson AR. Sleep quality and COVID-19-related stress in relation to mental health symptoms among Israeli and US adults. *Sleep Health.* 2021;7(2):127–33

Gupta R, Grover S, Basu A, Krishnan V, Tripathi A, Subramanyam A, et al. Changes in sleep pattern and sleep quality during COVID-19 lockdown. *Indian J Psychiatr.* 2020;62(4):370–378

Jahrami HA, Alhaj OA, Humood AM, et al. Sleep disturbances during the COVID-19 pandemic: A systematic review, meta-analysis, and meta-regression. *Sleep Med Rev*. 2022;62:101591. doi:10.1016/j.smrv.2022.101591

Alzueta E, , et al An international study of post-COVID sleep health. Sleep Health. 2022 Dec;8(6):684-690. doi: 10.1016/j.sleh.2022.06.011. Epub 2022 Sep 23. PMID: 36163137; PMCID: PMC9501615.

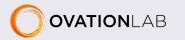




## SLEEP

- Both quality and quantity matter
- Improving sleep hygiene >
- Supplements that support sleep
- Practicing ultradian rhythm
   break or URB >





## SLEEP: Improving Sleep Hygiene

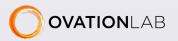
One way to improve sleep quality and quantity is to have daily routines and a bedroom environment that promotes consistent uninterrupted sleep.

Find personalized experiments to achieve sleep goals

Alzueta E,et al. An international study of post-COVID sleep health. Sleep Health. 2022 Dec;8(6):684-690. doi: 10.1016/j.sleh.2022.06.011. Epub 2022 Sep 23

Semyachkina-Glushkovskaya O, Mamedova A, Vinnik V, Klimova M, Saranceva E, Ageev V, Yu T, Zhu D, Penzel T, Kurths J. Brain Mechanisms of COVID-19-Sleep Disorders. Int J Mol Sci. 2021 Jun 28;22(13):6917. doi: 10.3390/ijms22136917. PMID: 34203143;







### SLEEP:

## Additional Resources

IFM Lifestyle Course resources:

- Effects of Poor Sleep
- The Sleep Questionnaire
- Mindfulness for Insomnia and Sleep Disorders
- Suggestions for Better Sleep







## MOVEMENT & EXERCISE >>

- Moderate, regular activity is essential to functioning of immune system, circulation, and tissue oxygenation
- Pacing of movement is essential.





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### MOVEMENTas Post COVID Treatment

POST-COVID-19 SYNDROME

#### Esimated time to resolution

6-12

weeks

8-12

weeks

2?

#### PSYCHOLOGICAL

Depression and anxiety
 Post-traumatic stress

#### NEUROLOGICAL

- Cognitive impairment
- Headache
- · Taste and smell alterations
- Post-traumatic stress
- Sleep disturbances
- · Peripheral neuropathy
- Dizziness
- Delirium

#### CARDIOVASCULAR

- Chest tightness
- Palpitations
- Orthostatic hypotension
- Syncope
- Dysautonomia

#### RESPIRATORY

- Dyspnea
   Chest pain
- Cough

#### MUSCULOSKELETAL

- Fatigue
- Weakness
- Osteoarticular pain
- Muscular pain

#### OTHERS

- Abdominal pain
- Nausea
- Diarrhoea
- Anorexia

#### PSYCHOLOGICAL

- Modulates pain
- t Well-being and mood state

POTENTIAL BENEFITS OF EXERCISE

↓ Stress

#### NEUROLOGICAL

- · Stimulates brain plasticity
- 1 Neurocognitive abilities
- ↓ Cognitive dysfunction
- ↓ Allostatic overload
- t Sleep quality

#### CARDIOVASCULAR

- 1 Mitochondrial biogenesis

- J Blood pressure
- Normalizes dysautonomia

#### RESPIRATORY

- ↓ Dyspnea
- t Oxygen uptake
- t Pulmonary function
- toxydative stress

#### MUSCULOSKELETAL

- ↑ Muscle strength
- † Intermuscular coordination
- tolerance to exercise

#### IMMUNE SYSTEM

- 1 Immune function
- ↓ Pro-inflammatory cytokines
- ↓ Immunosenescence

Jimeno-Almazán A, Pallarés JG, Buendía-Romero Á, Martínez-Cava A, Franco-López F, Sánchez-Alcaraz Martínez BJ, Bernal-Morel E, Courel-Ibáñez J. Post-COVID-19 Syndrome and the Potential Benefits of Exercise. Int J Environ Res Public Health. 2021 May 17;18(10):5329. doi: 10.3390/ijerph18105329. PMID: 34067776; PMCID: PMC8156194.







## TIPS TOGET MOVING

These experiments may help keep your body's stress response in check and help bring more physical activity into your day:

- Personalize it.
- Start small
- Plan it out.
- Step outside.
- Utilize daily activities.
- Take it slow.
- HIIT It.
- Find support.







### MOVEMENT: Additional Resources

IFM Lifestyle Course resources:

- The Exercise History Questionnaire
- Effects of Physical Inactivity
- Tips to Incorporate Mindful Movement Every Day
- SMART Goals for Exercise
- And more







### STRESS MODIFICATION >>

Techniques to modify the impact of stress on augmenting inflammation and decreasing immune function





## **STRESSMODIFICATION**

- Acute COVID is associated with elevated evening cortisol levels.
- Long COVID is associated with low cortisol levels and low activation of the HPA axis.
- Stress modification and adrenal restoration are treatment for post -viral conditions

Lamontagne S.J., Pizzagalli D.A., Olmstead M.C. Does inflammation link stress to poor COVID-19 outcome? Stress Health. 2021;37(3):401-414.

Sandrini L., Ieraci A., Amadio P., Zara M., Barbieri S.S. Impact of acute and chronic stress on thrombosis in healthy individuals and cardiovascular disease patients. Int. J. Mol. Sci. 2020;2

Yavropoulou MP, Tsokos GC, Chrousos GP, Sfikakis PP. Protracted stress-induced hypocortisolemia may account for the clinical and immune manifestations of Long COVID. Clin Immunol. 2022 Dec;245:109133.







## **STRESSMODIFICATION**

Stress triggers are highly individual.

Chronic stress can lead to physical symptoms such as headaches, high blood pressure, sleep issues, and digestive problems. It can also impact emotional and mental health and manifest as depression, anxiety, and panic attacks.







## STRESS: MIND and MENTAL HEALTH

Worsened by stress, mental health and performance impacts post-viral recovery:

- Brain fog is a common symptom this program addresses.
- Mental health conditions are often new or worse post COVID and appropriate referrals and support are imperative to recovery.
- Mindset matters in recovery

Taquet M, et al. Neurological and psychiatric risk trajectories after SARS-CoV-2 infection: an analysis of 2-year retrospective cohort studies including 1 284 437 patients. *Lancet Psychiatry*. 2022;9:815–827. doi: 10.1016/S2215-0366(22)00260-7.

Schou TM, Joca S, Wegener G, Bay-Richter C. Psychiatric and neuropsychiatric sequelae of COVID-19 - A systematic review. Brain Behav Immun. 2021 Oct;97:328-348. doi: 10.1016/j.bbi.2021.07.018. Epub 2021 Jul 30.

R. Zion a, Kengthsagn Louis a, Rina Horii a b, Kari Leibowitz a, Lauren C. Heathcote c d, Alia J. Crum. Making sense of a pandemic: Mindsets influence emotions, behaviors, health, and wellbeing during the COVID-19 pandemic. Social Science & Medicine. Volume 301, May 2022, 114889







## STRESSMASTERY TECHNIQUES >>

Like stress triggers, what helps someone navigate stressful situations varies. Some common experiments include:

- Relaxation techniques
- Meditation
- Finding support from others
- Being physically active (note that
   this can add stress to the body but
   may help address mental and
   emotional stress).

- Spending time in nature
- Diversion tactics
- Gratitude practices
- Laughter, joy, and play
- but Engaging in arts, music, and crafts







**STRESS MODIFICATION:** Additional Resources

IFM Lifestyle Course resources:

- Self-Care questionnaire
- Strategies for Transforming Stress
- Cultivating Mindfulness and Self Awareness
- Mindful Breathing







## STRESS: MIND and MENTAL HEALTH TIPS

- Refer people to mental health providers as appropriate
- Listen deeply, and offer support
- Acknowledge griefover losses
- Set expectations for a variable course of improvement
- Ask specifically what is working/ improving

- Support strengths
- Work with mindset to envision health
- Assist people to find community and reduce isolation

https://www.cdc.gov/coronavirus/2019-ncov/long-term-effects/care-post-covid.html

Burton A, Aughterson H, Fancourt D, Philip KEJ. Factors shaping the mental health and well-being of people experiencing persistent COVID-19 symptoms or 'long COVID': qualitative study. BJPsych Open. 2022 Mar 21;8(2):e72.



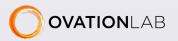


## SOCIAL CONNECTION >>

People impacted by long COVID reflect how isolated they continue to feel as others move on with lifting of the pandemic restrictions.

Connecting participants to community, and especially to support of people experiencing similar losses is essential to recovery.





## **COMMUNITY RESOURCES**

• https://www.weforum.org/agenda/2021/08/lon

g-covid-symptoms-illness-pandemic

- https://www.survivorcorps.com
- <u>https://www.longcovid.org</u>
- <u>https://longcovidalliance.org</u>







## SOCIAL CONNECTION: Additional Resources

IFM Lifestyle Course resources:

- Importance of Social Support and Relationships
- The Self- Care Questionnaire
- Practicing Gratitude with the IFM Gratitude Journal







# **POST VIRAL RECOVERY:** NUTRITIONAL SUPPLEMENTS





# **SUPPLEMENT** TARGETING

◆ ACE-2 mediated damage (tissue ♦ Viral persistence Chronic inflammation Mitochondrial dys function Gut / Microbiome dysbiosis

After simplifying the Post COVID Recovery Framework, we found ourselves needing a curated suite of nutritional product solutions to target the <u>five</u> main drivers of Long COVID.





# LONG COVID FRAMEWORK







#### **Foundational Supplements**

MitoCore [MVit + AntiOx] (2x/day 90 days) Vitamin K2 with D3 (up to 10,000IU) Magnesium Glycinate (200mg) 2x/day 90 days) Quercetin (500mg 2x/day 90 days) Coenzyme Q10 (100mg/day 90) days)

MitoCore key features:

Comprehensive MVI with Antioxidant and Mitochondrial Support

- NAC 500 mg
- Glucoraphanin 40 mg
- Alpha Lipoic Acid, Resveratrol and EGCG
- Acetyl L-carnitine 500 mg, Magnesium
- Manganese, Zinc

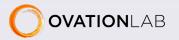
Vitamin K2 with D3 dosing based on levels:

- serum level-<40 ng/mL, use 10000 IU;</p>
- ✤ if 40-60 ng/mL, use 5000 IU daily
- ✤ if >60 ng/mL no additional needed

Quercetin:

- Immunomodulatory and may improve T Reg function
- Demonstrated anti-viral activity against both RNA and DNA viruses

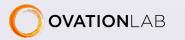




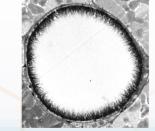
Arterosil: Endothelial, Blood Vessel, Cardiovascular Tissue Damage











Arterosil: Protects and repairs the Endothelial Glycocalyx (EGx)

- Selectively permeable barrier that prevents blood components from sticking to the vessel wall
- Harbors Super Oxide Dismutase (extracellular SOD)
- Regulates vascular permeability, inflammation, coagulation, and fluid balance

Active Ingredient: Rhamnan Sulfate





**return** HEALTHY ITIS" Response to Die Off Re **ITIS**: Natural Anti -inflammatory, Scutulleria

ΓIS Formula:	Supplement Fac	ts	
Reduces pain & inflammation	Serving Size: 4 Vegetarian Capsules Servings Per Container: 30 Amount Per Serving		DV.
Maintains comfortable joint & muscle	Skullcap Extract (Scutellaria baicalensis, root, wildcrafted)		
function	Celery Seed Powder (Apium graveolens)		
	Tart Cherry Extract 4:1 (Prunus cerasus, fruit)		*
oothes an irritated immune system	Cat's Claw Powder		
	(Unicaria tomentosa, wildcrafted bark) Boswellia serrata Extract		-
ey Ingredients:	(Standardized to 65% Boswellic Acid, Non-GMO) Devit's Claw Root Powder		
	(Harpagophytum procumbens, Non-GMO, Organic)		
Scutellaria (Chinese skullcap)	Bromelain (600 GDU/g) Feverfew Powder (Tanacetum parthenium, herb, Organic)		
· · · · · · · · · · · · · · · · · · ·	Ouercetin Dihydrate		
Curcumin	BioPerine Black Pepper Extract (Piper nigrum, fruit)		
$D = 11^{\circ} (C = 1^{\circ})$	Ginger Essential Oil (Zingiber officinale, root)		
Boswellia serrata (frankincense)	Curcumin Extract 95%	202.5 mg	6
Copaiba	(Curcuma longa root extract)	202.5 mg	
1	Roman Chamomile Essential OII	20 mg	
Roman Chamomile	1 - % Daily Value * - Daily Value Not Established		
	Other Ingredients: Cellulose (capsule).	SP2721-12	0







#### Tollovid: Blocks viral replication capacity

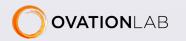
Coronaviruses use their signature "spikes" to fuse themselves to a cell's outer membrane via the ACE2-receptor to gain entry inside.

The spike protein is primed for access using a 3CL protease
By tightly binding to this protease enzyme, its ability to cleave is blocked.
This shuts down the entire virus-making apparatus, leaving it unable to enter the cell and replicate.



Active Ingredient: Gromwell Root







MegaSporeBiotic: Maintain healthy gut barrier function

#### MegaSporeBiotic

- Probiotic blend of 5 Bacillus spores that have been shown to maintain healthy gut barrier and immune function
- Aims to RECONDITION the gut instead of reseeding with probiotic strains that cannot survive digestion or colonize the gut.

Active Ingredient: Bacillus spores







Gut Health Support, Microbial Diversity

#### MegaPreBiotic

- Contains non-digestible oligosaccharides that can increase microbial diversity and selectively feed beneficial bacteria like Akkermansia muciniphila, Faecalibacterium prausnitzii, and Bifidobacteria
- REINFORCES the beneficial microbial changes created by MegaSporeBiotic + to promote a strong and diverse microbiome..

Active Ingredient: non-GMO green and gold kiwi fruit, non-GMO corn cob, and rBST-free cow's milk





# FIRST 30-DAYS SUPPLEMENT PROTOCOL

#### Arterosil

- 2 caps 2x/day
- ITIS Formula
  - 2 caps 3x/day
- Tollovid

Specialty Nutritional Supplements

- 3 caps 4x/day
- MegaSporeBiotic
  - 1 cap/day for 7 days, then 2caps/day to complete one bottle
- MegaPrebiotic
  - N/A

Foundational Supplements

- MitoCore [MVit + AntiOx]
- 2 caps 2x/day
- Vitamin K2 with D3
  - 1 cap 2x/day with food
- OmegaGenics® EPA-DHA 1000
  - 1000 mg 2x/day
- Designs for Health® Magnesium Chelate
  - 200 mg 2x/day
- Vital Nutrients Quercetin 250 mg
  - 2 caps 2x/day
- Vital Nutrients Coenzyme Q10 100 mg
  - 1 cap once/day





# **31-TO-60 DAYS SUPPLEMENT PROTOCOL**

Supplements

Foundational

#### ♦ Arterosil\*

1 cap 2x/day

- ITIS Formula
  - 2 caps 3x/day

#### Tollovid\*

nal

Special

Supplements

2 caps 4x/day for 7 days then 1 cap 2x/day for 30 days

#### MegaSporeBiotic\*

2caps/day to complete one bottle

- MegaPrebiotic\*
  - 3 caps/day

#### MitoCore [MVit + AntiOx]

- 2 caps 2x/day
- Vitamin K2 with D3
  - 1 cap 2x/day with food
- OmegaGenics® EPA-DHA 1000
  - 1000 mg 2x/day
- Designs for Health® Magnesium Chelate
  - 200 mg 2x/day
- Vital Nutrients Quercetin 250 mg
  - 2 caps 2x/day
- Vital Nutrients Coenzyme Q10 100 mg
  - 1 cap once/day





# 61-TO-90 DAYS SUPPLEMENT PROTOCOL

#### Arterosil

1 cap 2x/day

- ♦ ITIS Formula
  - 2 caps 3x/day
- Tollovid\*

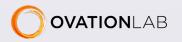
Specialty Nutritional Supplements

- 1 cap 2x/day for 7 days
- MegaSporeBiotic\*
- MegaPrebiotic
  - 3 caps/day

Supplements Foundational

- MitoCore [MVit + AntiOx]
- 2 caps 2x/day
- Vitamin K2 with D3
  - 1 cap 2x/day with food
- OmegaGenics ® EPA-DHA 1000
- 1000 mg 2x/day
- Designs for Health® Magnesium Chelate
- 200 mg 2x/day
- Vital Nutrients Quercetin 250 mg
  - 2 caps 2x/day
- Vital Nutrients Coenzyme Q10 100 mg
  - l cap once/day





# Patient Registry Early Outcomes



#### Benchmarking Our Success

0.5 - 1.0 point decline = clinically significant

\$ >1.0 point decline =
excellent

## Patient Outcomes

#### 4.5 months after first patient enrollment

#### Post COVID Function Scale

- Patients (n=8) achieved an average decline of 1.5 points.
- ◆ 5 of 8 patients reported total resolution over the 90-day program.
- No significant adverse events to date. 2 reports Tollovid GI intolerance (upset stomach).

PROMIS-29: 7 domains of function plus a global pain measure

- Physical Function: increased from 15th to 42nd percentile
- Social Participation: increased from 17th percentile to 63rd percentile
- Anxiety: decreased from 80th to 34th percentile
- Depression: decreased from 73rd to 46th percentile
- Fatigue: decreased from 82nd to 58th percentile
- Sleep Interference: decreased from 58th to 52nd percentile
- Pain Interference: decreased from 63rd to 41st percentile





*"My blood oxygen used to go down to 82% while doing PT at the hospital little effort. Now I stay over 90% O2 sat with a lot more movement in 2 weeks on the supplements."* 

*"The group visits tools give me power – I have agency when I have scary and dangerous symptoms, they no longer overwhelm me."* 

*"My body shape is changing, and I'm convinced it (the protocol) is helping everything."* 

#### **Early Patient Quotes**

- \* "My migraines are gone now, just small daily headaches remain, they are less."
- \* "My chest pain was really bothering me, I almost went to the ER again but they never find anything. I took an Arterosil and my pain reduced enough to stay home. Now that I'm on it 2 weeks I don't have those pains anymore." (Patient cried with gratitude when sharing this).
- \* "My pain with exertion is less, and I can move more without being set back."
- ✤ "I have much more mental clarity."
- "I have much better energy and much better mood."
- \* "My muscle fatigue is much less since starting the program."
- I am done with the Tollovid. It helped my energy. On the Arterosil now for a month I continue to have a lot less pooling and swelling in my legs, and my POTS symptoms are less."
- I know more than my regular doctors about my condition. I wish more knew about mitochondria, vagal tone, cortisol and adrenal health, Ace-2 receptors and how it all affects long COVID symptoms."
- "I no longer feel alone in what I am dealing with. Hearing everyone else's stories heals me. Hearing everyone's tools and ways of coping gives me hope and options."
   (6 people emphasize this).







# CASE STUDY: CS

Dr. Patrick Hanaway, MD

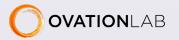
# CS Demographics

\$51yo male, business executive
Height = 6' 3<sup>1</sup>/<sub>2</sub>"
Weight = 297#
BMI = 36.6



Vaccine x1 (Pfizer) April, 2021
COVID Jan, 2022







## CS History / Timeline

COVID #1 = Jan, 2022

- Very Sick, Hospitalized x 8d, O2 support (no ventilator)
- \* Fluid on Lungs, recurrence of Lymphoma
- ♦ ChemoTx May,  $2022 \rightarrow August$ , 2022
- Vaccine (Pfizer) #2 Sept, 2022

#### COVID #2 = Jan, 2023

- \* Milder, received Paxlovid
- ◆ Hospitalized x4d for ↓ WBC, received Neupogen
- ◆ Development of 'Brain Fog' and ↓ Memory (without fatigue)
- Primary Concern is "loss of words . . . mid-sentence"







# CS Diagnoses

Meds:

Valacyclovir (s/p Shingles)

Supplements:

MultiVitamin (generic)







### **CS** Treatments

Week 1 – Fasting Mimicking Diet

- Week 2 Begin Lifestyle Modification & CGM
- Week 3 Begin Protocol
  - \* Tollovid x 30 days (3 caps QID) with one month taper
  - ITIS' 2 caps TID x 90d
  - \* Arterosil 2 caps BID x 30d, then 1 cap BID x 60d
  - VascaNOX1 cap BID x 90d
  - Mitocore (B-Vitamin + AntiOx) 2 caps BID x 90d
  - Vitamin D 5000IU qD x 90d
  - EPA/ DHA 1000mg 2 caps qD x 90d
  - Vitamin C 500mg TID x 90d



Whole30 Food Plan Gradual Exercise Plan





### CS Follow-up/Outcomes

Long COVID x 4 months

#### 4/12

♦ Dietary modification x 2 weeks  $\rightarrow$  Weight  $\downarrow 10#$ 

#### 5/2

"The results of my scans indicate that I am still in remission and that previously impacted lymph nodes are all within normal size. The even more impressive news is that my blood counts are better than they have been for 17 years."

#### 5/15

• Weight Loss =  $\downarrow 35\#$ 

"I feel better than I've felt in years. I don't experience 'word loss' like I did. The changes have been profound."









Group Medical Visits

**Dr. Kara Parker, MD** Faculty, Department of Family Medicine Director of Group Medical Visits Hennepin Healthcare Whittier Clinic

# **PVRP Group Medical Visit Components**



Scaled education on the benefits of and resources for specific Lifestyle factors for recovery after COVID

Group instruction and exploration on Whole 30, plant based anti inflammatory diet tailored to each person's needs

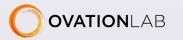


Group teaching on the specialty and foundational supplements for post Viral recovery



Time for support, connection, questions, and group practices for each part of the program





### PVRP Group Visit Our Topics

- Overview of program, Intros, Patient Post COVID story
- Food & Nutrition
- Sleep & Mood
- Movement & Exercise
- Stress Modification
- \*Brain and Microbiome
- Mitochondria







### PVRP Group Medical Visits Our Structure:

\*7 week series, 2 hour virtual visits Start with a Mind Body exercise to center \*Ask a question relevant to each weekly topic Teach on physiology, science, and tips and encourage discussion and sharing End with a long form mind-body exercise and closing take-aways or actions \*Empower participants to practice self-care during the visits







### PVRP Group Medical Visits Stats/ Insights:

7/2020 to current, 10 groups, average 12 pts 2 recent groups using the protocol

- Patients who feel alone, isolated, and misunderstood benefit from connection
- The Lifestyle and supplements (protocol) fit well in a GV format
- I have learned on care of pts w LC by listening to pts w LC share needs & wins
  Benefits multiply for pts and practitioners in the

care and presence of others







- \$59 yo female with PMH depression and chronic constipation and 3 Pfizer n Biotech vaccines including a booster 11/21.
- COVID + Dec 21 by pos home PCR test.
- Had flu like symptoms cared for at home without complication
- Began to develop new symptoms by 3 weeks







By one month reported:

- ✤ Severe fatigue
- Cognitive changes
- Vision changes
- New daily headache
- ✤ Joint pain
- Muscle pain
- Disturbance of sleep
- Mood change
- Loose stools







Worked with her primary care to address symptoms with no change by 6 months, sought referral

\*Was sent to our PM&R Post COVID clinic, and while on a waitlist was also referred to our Long COVID Group Visits Program and started 9/22







Attended all 7 GV sessions:



- \* Visit 1 Intro and Covid Story. TD shared her grief at fxn loss
- Visit 2 Stress: Learned extended exhale breathing & shaking
- Visit 3 Nutrition: Learned whole 30 recipes and reduced grains
- Visit 4 Mitochondria: Started coenzyme Q 10, healthy fats
- Visit 5 Sleep: Practiced Body Scan to fall asleep
- Visit 6 Brain and Biome: Tried humming & prebiotic foods
- Visit 7 Movement: Learned tips on pacing & theraband exercises



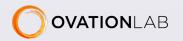


At completion of the GV Series she went from "40%" to "60%
She then started TOLLOVID and Arterosil

≻by day 5 she walked to and around the lake!★Finished a 10 day regimen TOLLOVID

reported HA gone, brain clear, concentration back, reading
constipation returned (a sign she was back to herself)!
ITIS was given and she did not tolerate it, stopped it
Added MegaSpore prebiotics and aloe and stools normalized, making her feel better than pre-covid condition.



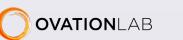


★4 months later had a return of fatigue when she:
>had added back grains
>had a mild URI
>smelled mold in her apartment after heavy rain

\*Symptoms resolved after she:

- ≻reduced grains
- Filtered air, got tested and remediated and
   restarted TOLLOVID for 10 days









# RESOURCES

www.ovationlab.com/PVRPresources



# What's Next?

Patient Registry Goal: 100 new patients who complete the 90 -day program based using updated protocol by 9/30/2023.

 Interested in joining our learning community?
 Complete and submit interest form here

#### **Next Steps**

- Special Topics Workshops
- Master Classes
- Learning Community Case Studies
  - Learn from our working group and your colleagues
  - Present your own patient cases





#### **Question and Answer Session**





#### Recommended Education Programs and Online Courses

The American Academy of Anti-Aging Medicine (A4M)

Use the code VIRAL200 for \$200 off your registration fee Longevity Fest 2023 – December 14-16, 2023

The Academy of Integrative Health & Medicine (AIHM)



Use the code OVATION100 for \$100 off your registration fee to the October Conference Annual Conference - October 5-8, 2023

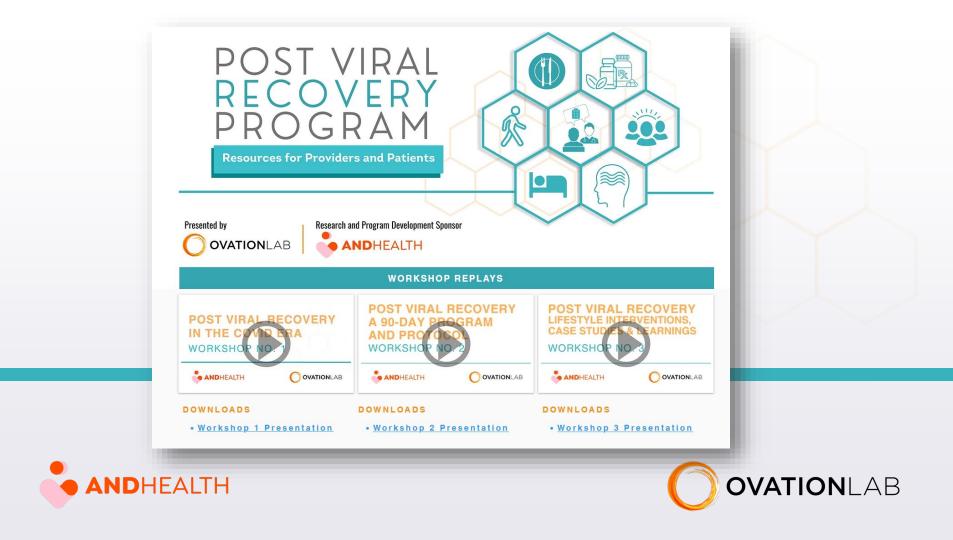
#### The Institute for Functional Medicine (IFM)



Use the code OVATIONLAB10 for 10% off your registration fee <u>Applying Functional Medicine in Clinical Practice</u> <u>May 2023 (AFMCP)</u> - online Lifestyle: The Foundations of Functional Medicine – on-demand, online







#### POST VIRAL RECOVERY PROGRAM Resources for Providers and Patients





#### RECOMMENDED EDUCATION PROGRAMS AND ONLINE COURSES

Check-out the links to upcoming conferences and online programs offered by the following education organizations.

- The American Academy of Anti-Aging Medicine (A4M)
  - Use the code <u>VIRAL200</u> for \$200 off your registration fee<sub>.</sub>
  - Longevity Fest 2023
- The Academy of Integrative Health & Medicine (AIHM) Use the code <u>OVATION100</u> for \$100 off your registration fee to the October Conference
  - <u>Annual Conference, October 5-8, 2023</u>

#### The Institute for Functional Medicine (IFM)

Use the code **OVATIONLAB10** for 10% off your registration fee

- Applying Functional Medicine in Clinical Practice™ May 2023 (AFMCP)
- AIC 2023: Advancements in Clinical Research and Innovative
   Practices in Functional Medicine
- Lifestyle: The Foundations of Functional Medicine

#### Personalized Lifestyle Medicine Institute (PLMI)

• Is COVID Long-Haul a disease of the gut

NIH Researching COVID to Enhance Recovery (RECOVER)

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#### PATIENT CASES - Coming

#### OR PATIENTS

Eullscript Patient Protocol

#### WHOLE30 PROGRAM

Program Rules
 Plant-Based Whole30 Prep Pack
 Grocery Guide
 Shopping List

Meal Planning
 Plant-Based Recipes

#### LIFESTYLE INTERVENTION RESOURCES

How to Improve Sleep Hygiene
 Circadian Rhythms
 Stress Management

Physical Activity and Stress
 Exercise Tolerance Assessment for
 Exercise Fitness & HR Recovery
 Instructions (from Share Care)

WHO Support for Rehabilitation

- The Levine Protocol For Exercising With POTS - Better By The Beat
   Instructions for POTS Exercise
- Program—Children's Hospital of Philadelphia the Structure of the Training Calendars

 <u>Coronavirus Recovery: Breathing</u> <u>Exercises | Johns Hopkins Medicine</u>
 Bouncing Back From COVID-19

#### SAMPLE SURVEY INSTRUMENTS

- LongCOVID Patient Intake Form
   WHO Post COVID-19 CRF
- <u>Timeline and Progression of</u>

TOOLS

Application to Participate in Post

community and patient registry.

Viral/Long COVID learning

Protocol Application with

· If you do not have an active

Fullscript account, please use this

link to create your no-cost account.

• If you would like to prescribe the

specialty products only, please

. If you would like to prescribe the full

· Dispensary & individual patient

protocol or make modifications to it,

Fullscript

use this link.

discounts

please use this link.

- Symptoms Table
- PROMIS29
- Post COVID Function Scale







YOU

THANK



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