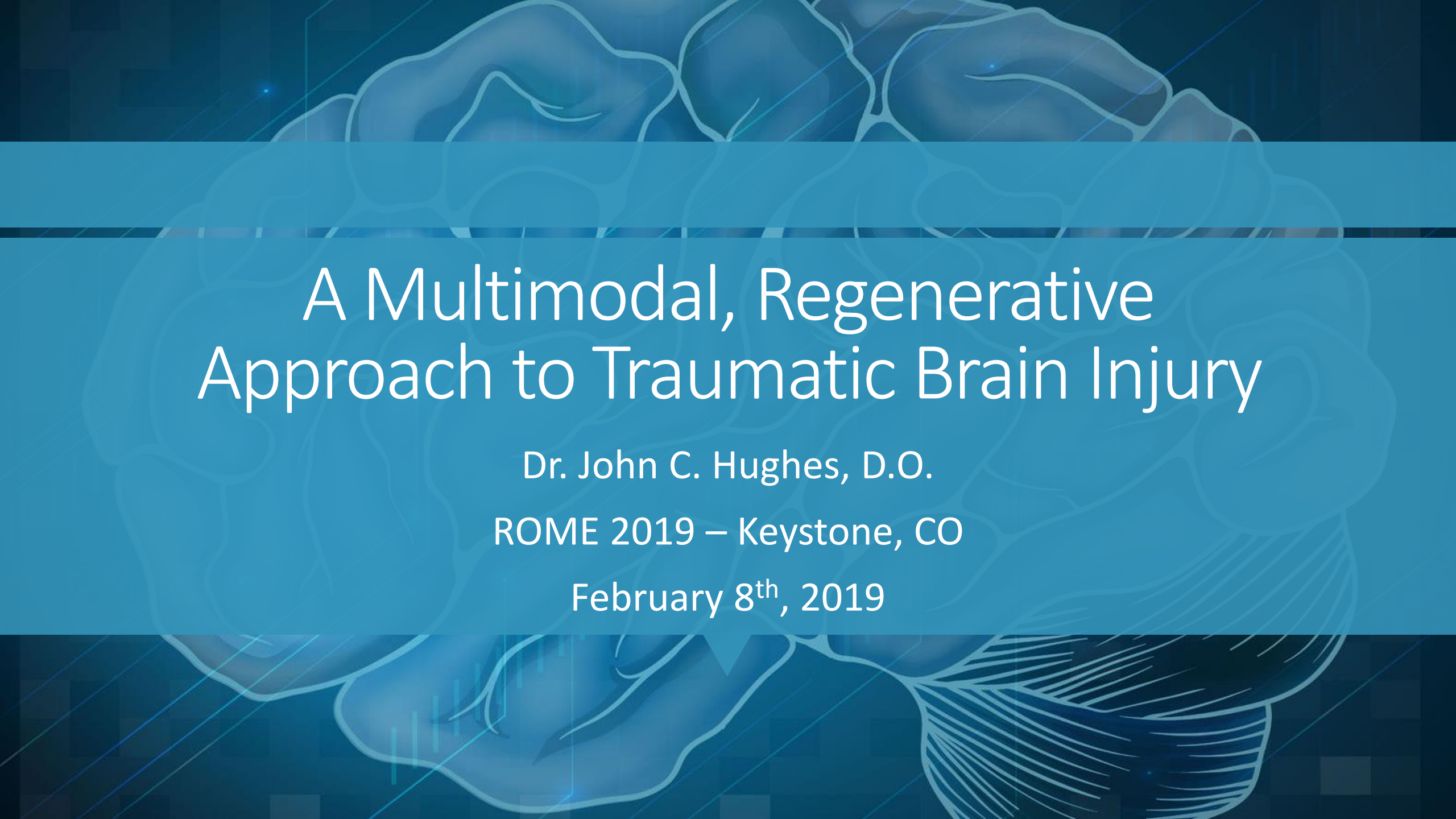

A Multimodal, Regenerative Approach to Traumatic Brain Injury

Dr. John C. Hughes, D.O.

ROME 2019 – Keystone, CO

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Disclosure

The content of this presentation has been peer reviewed for fair balance and evidence based medicine.

Learning Objectives

Define

Define the clinical, biochemical and metabolic effects from TBI

Identify

Identify mainstream and alternative treatments for TBI

Understand

Understand the regenerative model of TBI treatment

Clinical Symptoms from TBI

Physical

Headache
Fatigue
Sleep disorders
Vertigo or dizziness
Tinnitus or hyperacusis
Photosensitivity
Anomia
Reduced tolerance to psychotropic medications
Disorientation
Loss of mobility
Seizures
Loss of smell

Cognitive

Memory decline / loss
Slow reaction time
Inability to pay attention
Executive dysfunction
Slow learning
Interrupted speech
Difficulty understanding
Unable to concentrate
Confusion
Difficulty communicating thoughts
Unable to plan, reason, problem-solve

Psychological

Irritability
Easy frustration
Tension
Anxiety
Affective lability
Personality changes
Disinhibition
Apathy
Suspiciousness
Suicidality
Depression
PTSD

Biochemical and Physiological Responses from TBI

- Disproportional proinflammatory cytokine production and release
- Increased counterregulatory hormones work against the action of insulin

Biochemical and Physiological Responses from TBI

- Hypermetabolic and catabolic states
- Severely impaired nitrogen homeostasis
- Oxidative Stress

Oxidative Stress From TBI

- Impairs cerebral vascular function
- Impairs circulation
- Impairs the energy metabolism
- Damages mitochondria and DNA

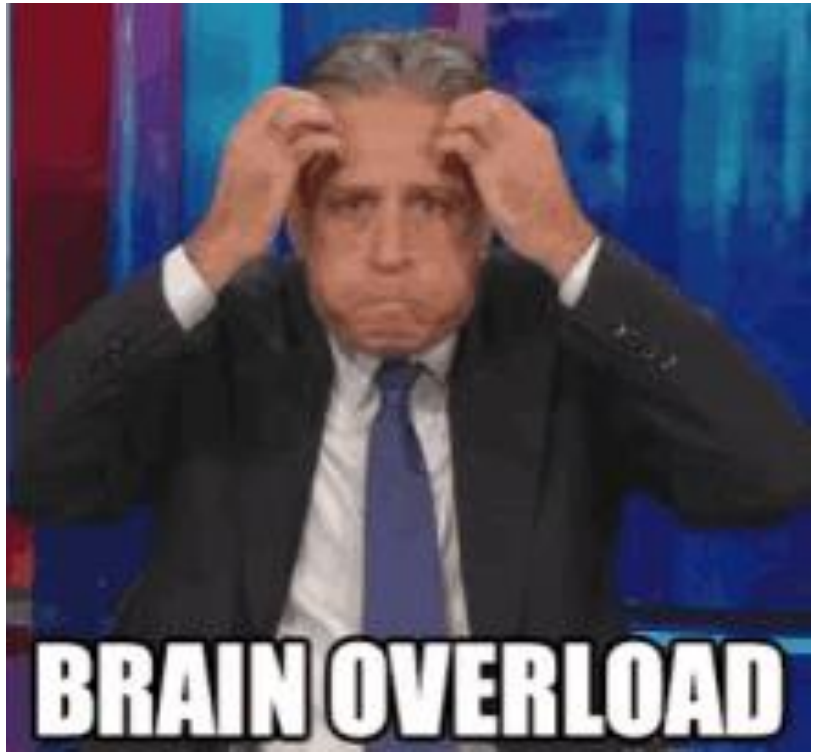
What Happens Metabolically with a TBI?

"The brain is in a metabolic crisis with concussion… potassium ion from inside the cell going extracellularly, calcium ions going intracellularly, neurotransmitters widely released in a chaotic manner.

It takes energy to pump that potassium back, put the neurotransmitters back on so the cell can function."

Dr Robert Cantu, MD, 2013

What Happens Metabolically with a TBI?

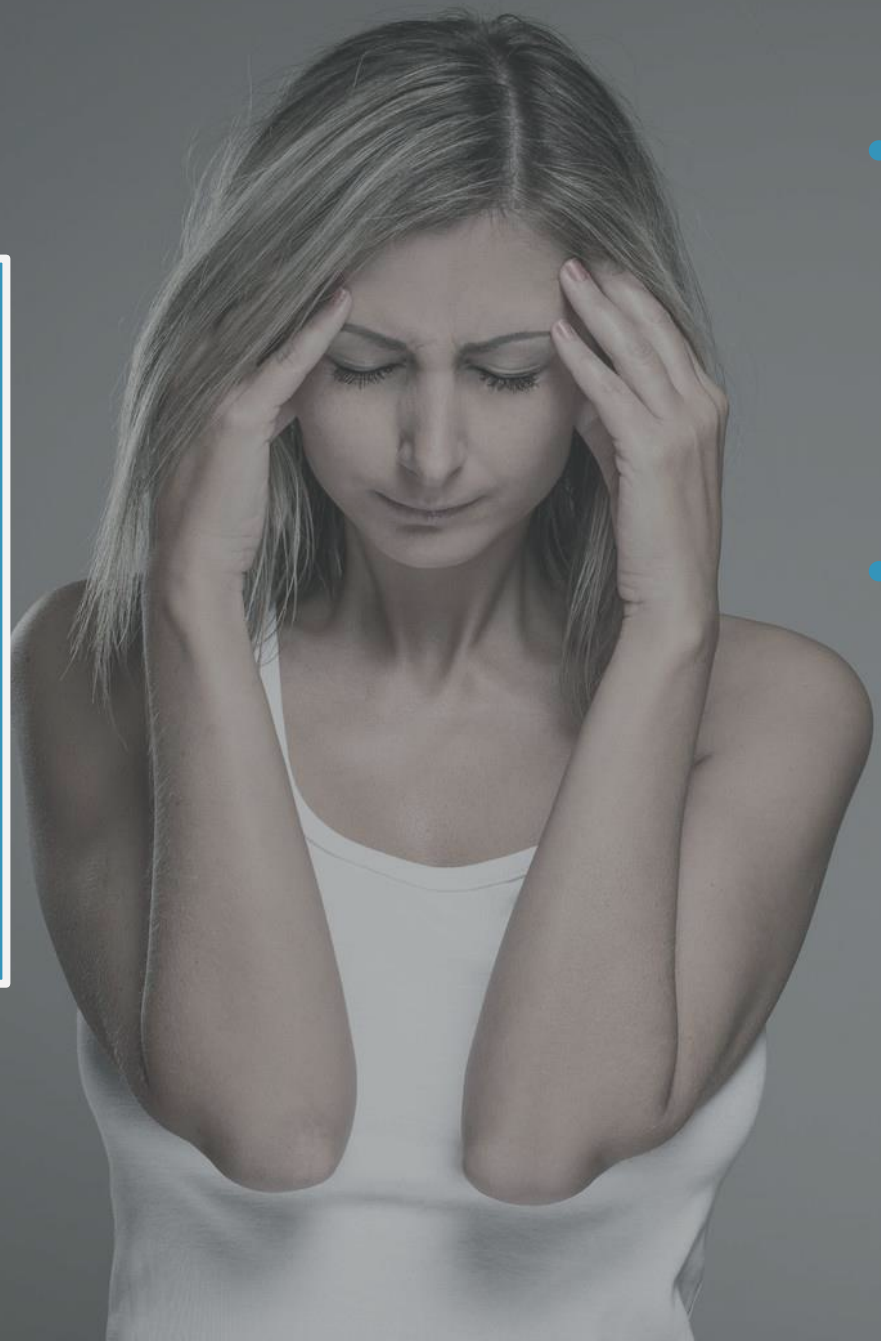


An Energy Crisis

Mainstream Treatments

- Occupational and physical rehabilitation
- Speech therapy
- Pharmaceutical drugs
- Cognitive maintenance exercises
- Patients simply cope with their condition

Alternative Treatments



- Do not seek to regenerate but rather simply treat symptoms
- Do not combine regenerative treatments in a multimodal manner in order to maximize patient benefit

A Multimodal, Regenerative Approach to TBI

It is hypothesized that the practical, effective combination of multiple regenerative TBI therapies can produce synergistic benefits to the patient that exceed the use of one particular TBI treatment.

A Multimodal, Regenerative Approach to TBI

I. Hyperbaric Oxygen Therapy

II. Intranasal Therapies

III. IV Nutrition

IV. Cranial Osteopathy

V. Ketogenic Diet and MCT Oil

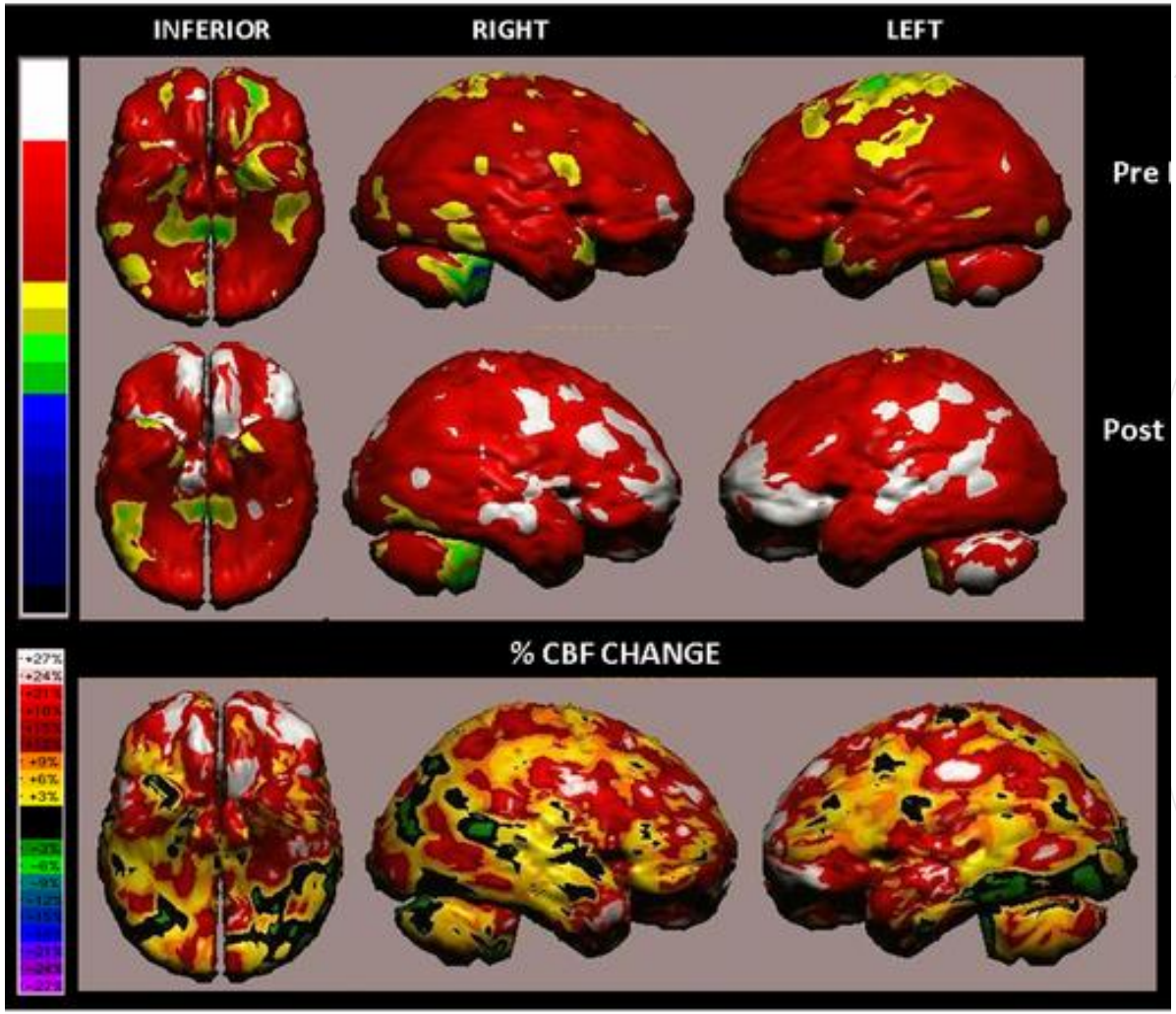
Part I

Hyperbaric Oxygen Therapy (HBOT) for TBI

Hyperbaric Oxygen Therapy (HBOT)

- Allows the body to absorb about 10-15 times its normal supply of oxygen
- Stimulates the growth of tissue, bone and blood vessels, and reduces inflammation

Thom, et al., 2006

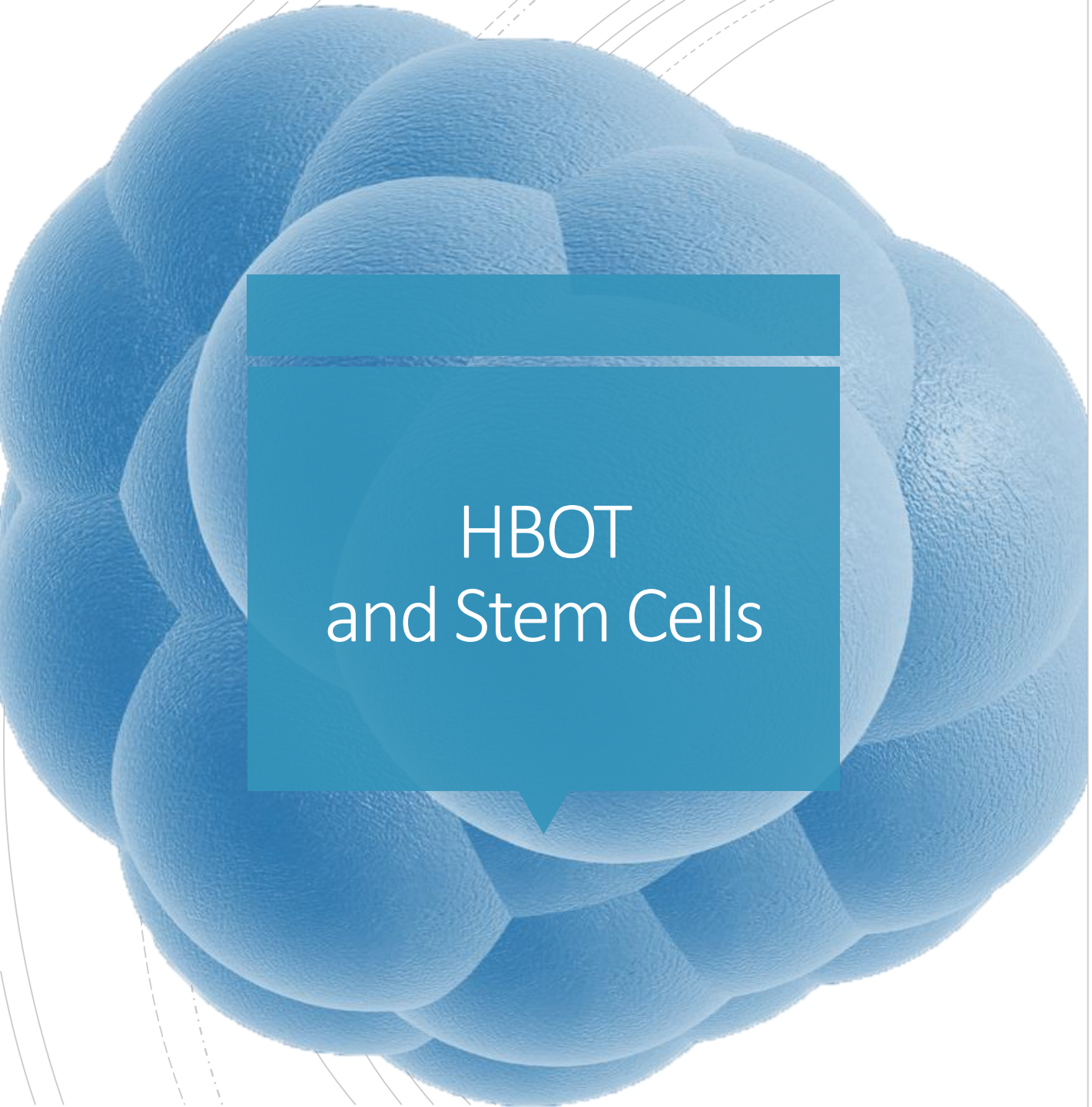


Volume rendered Brain SPECT perfusion maps of a 51-year-old woman suffering from mTBI that had occurred 2 years prior to inclusion in the study

Boussi-Gross et al., 2013

HBOT for TBI

- Induces neuroplasticity
- Increases tissue oxygenation
- Generates new capillary networks
- Restores blood supply
- Increases stem cells in the blood

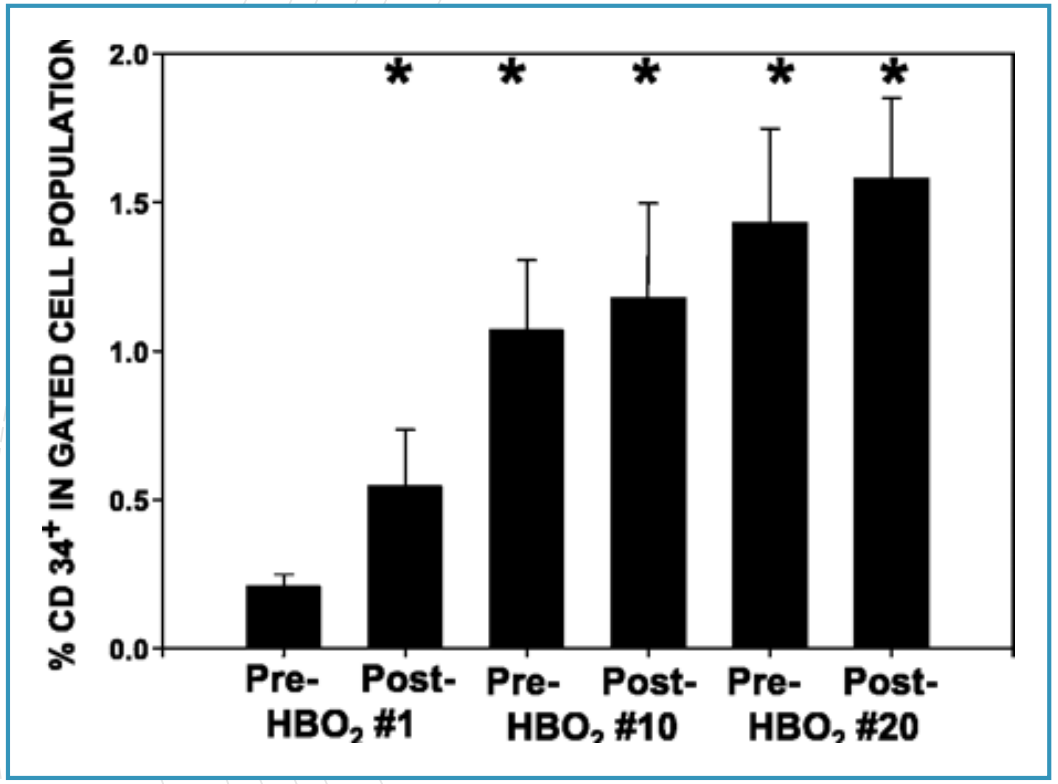


HBOT and Stem Cells

- 2 hours of HBOT triples the patients own circulating stem cells
- 20 sessions of HBOT increases circulating stem cells to 8 fold (800%)

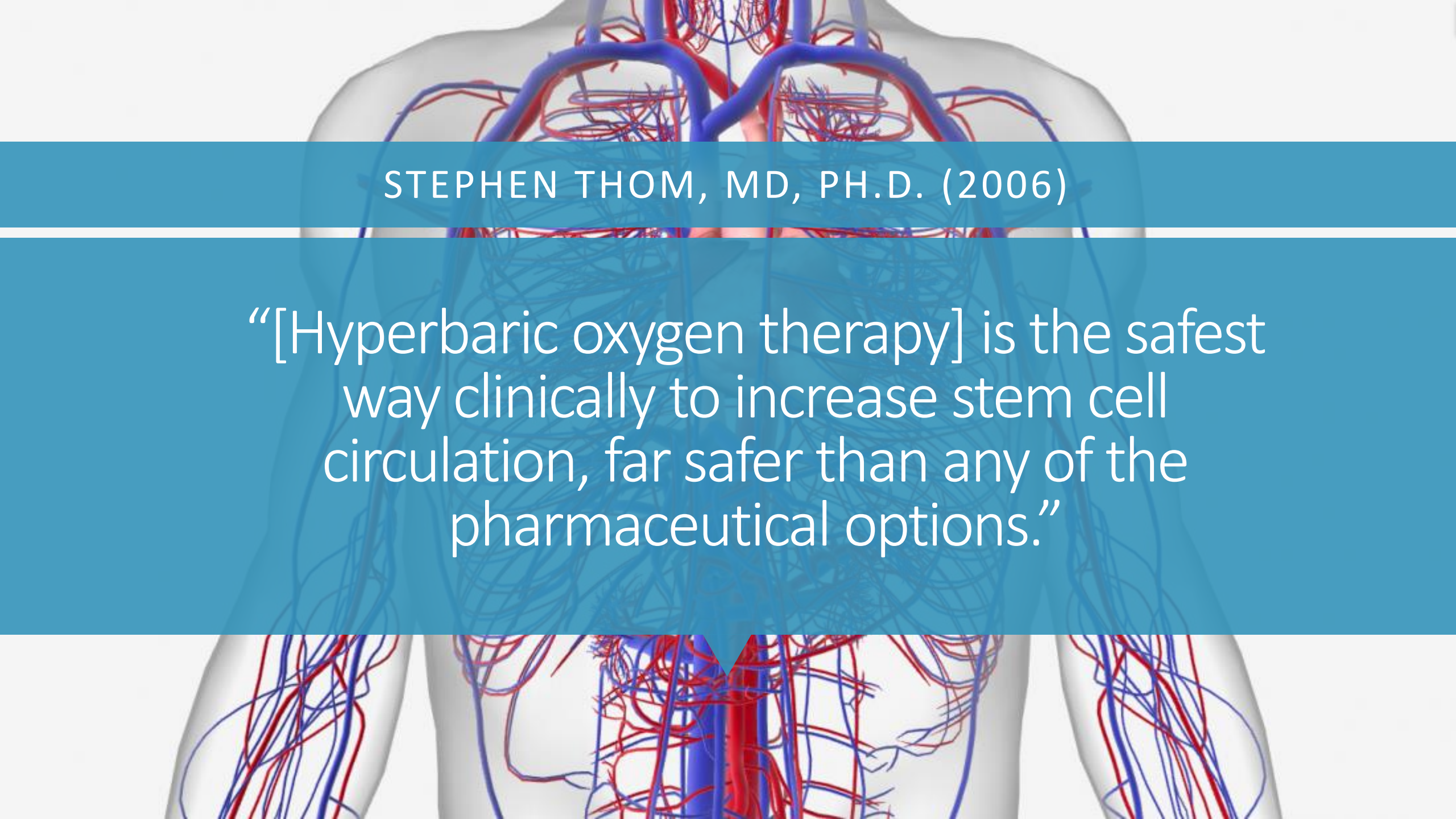
Thom, et al., 2006

Mean CD34+ population in blood of humans before and after HBO2 treatments.



Data are the fraction of CD34+ cells within the gated population using leukocytes obtained from 26 patients before and after their 1st, 10th, and 20th HBO2 treatment.

Thom, et al., 2006



STEPHEN THOM, MD, PH.D. (2006)

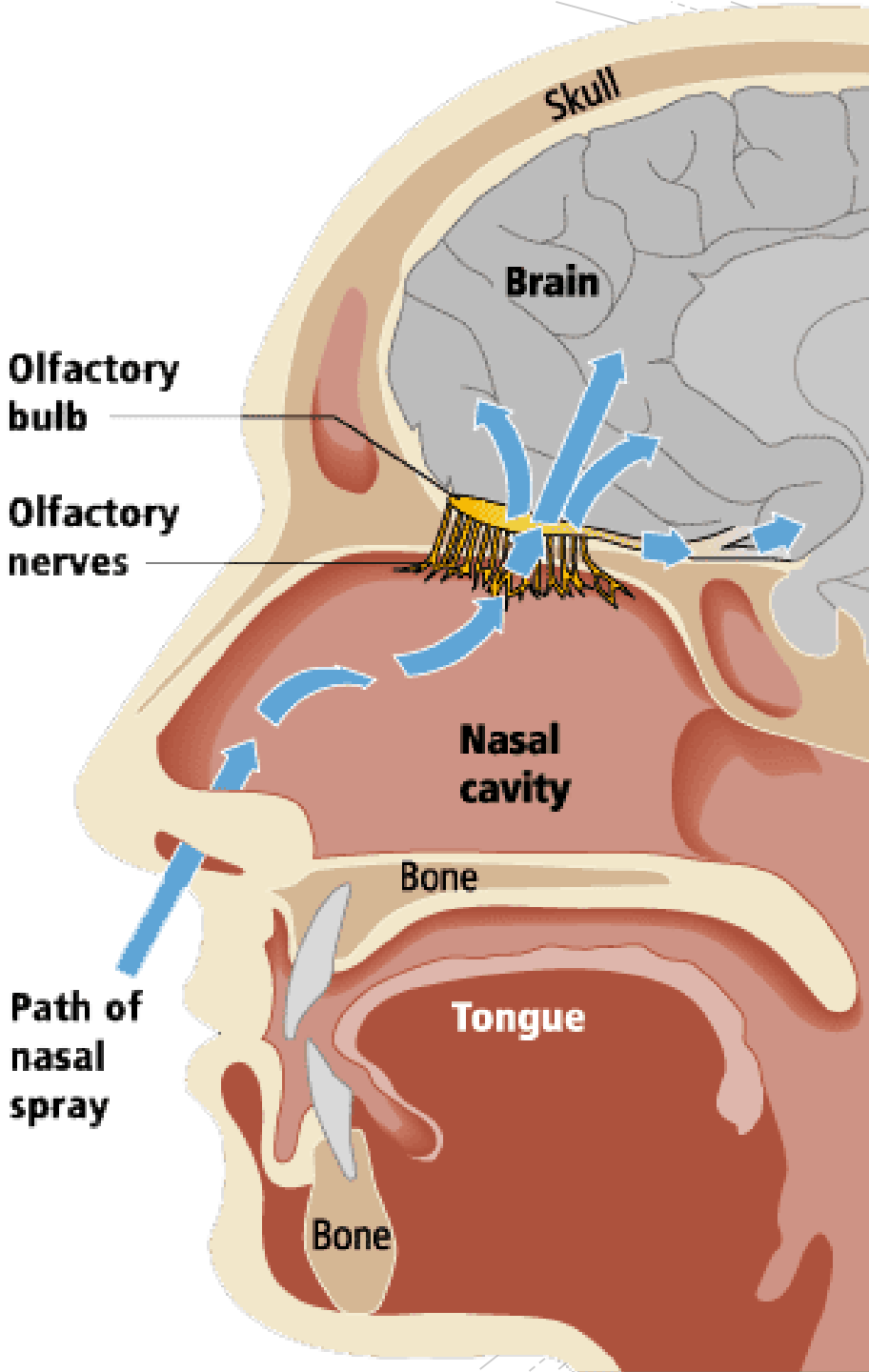
"[Hyperbaric oxygen therapy] is the safest way clinically to increase stem cell circulation, far safer than any of the pharmaceutical options."

Part II

Intranasal Therapies (Insulin, PRP, and Stem Cells) for TBI

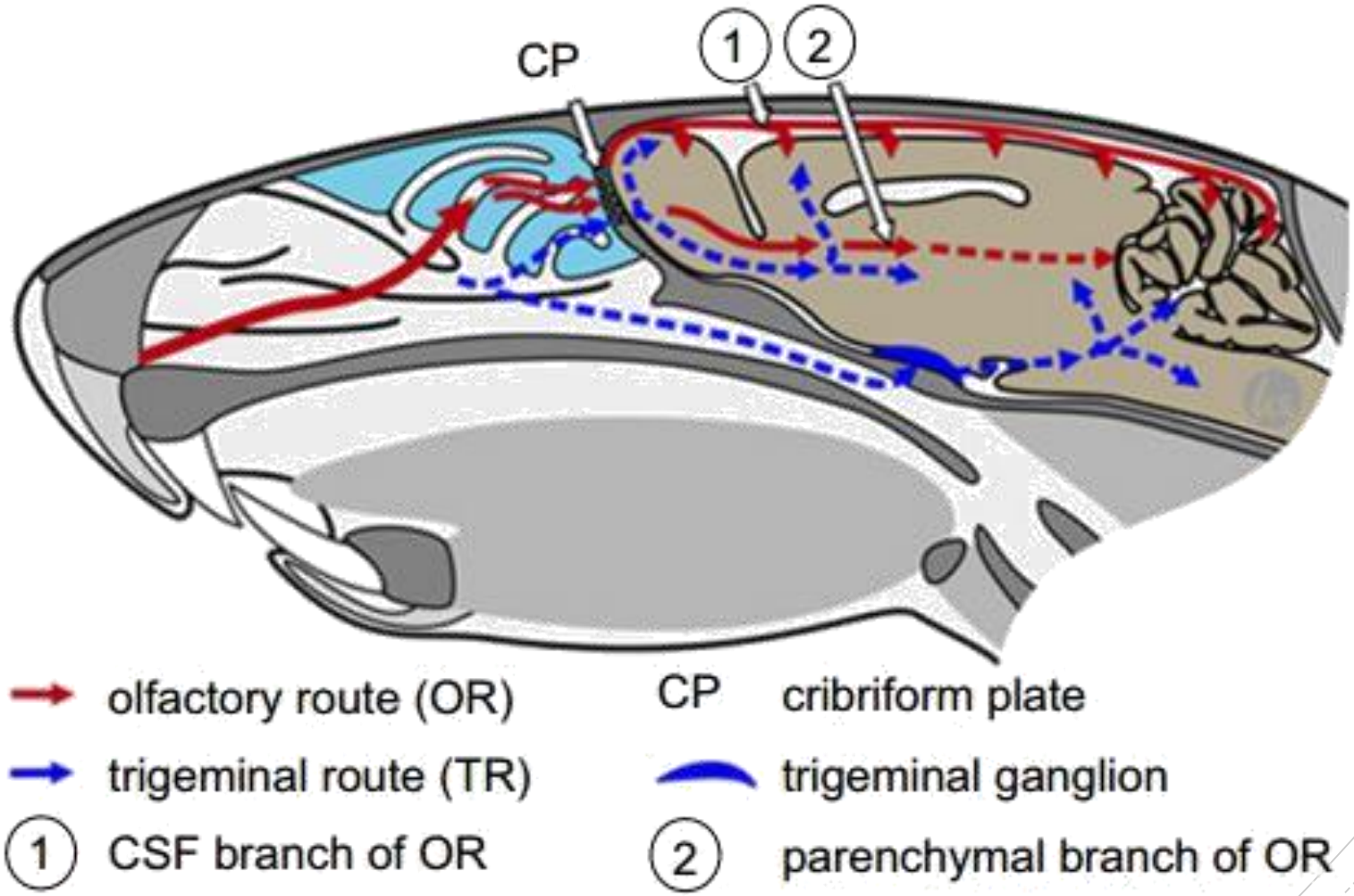
Journey Through the Nose

- Through the olfactory nerves
- Bypasses the blood-brain barrier
- Into the CSF within 10 minutes



Mouse Brain

Solid arrows represent the paths of migration of cells into the brain, dashed arrows reflect possible hypothetical routes of cell delivery

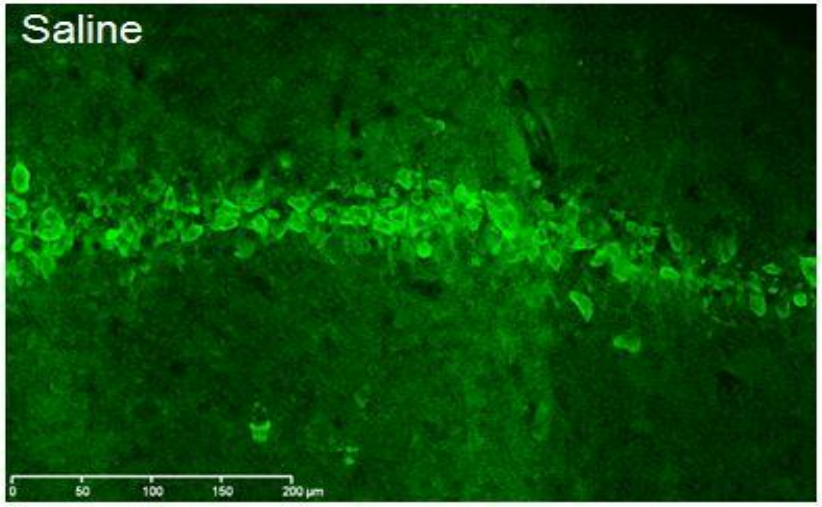


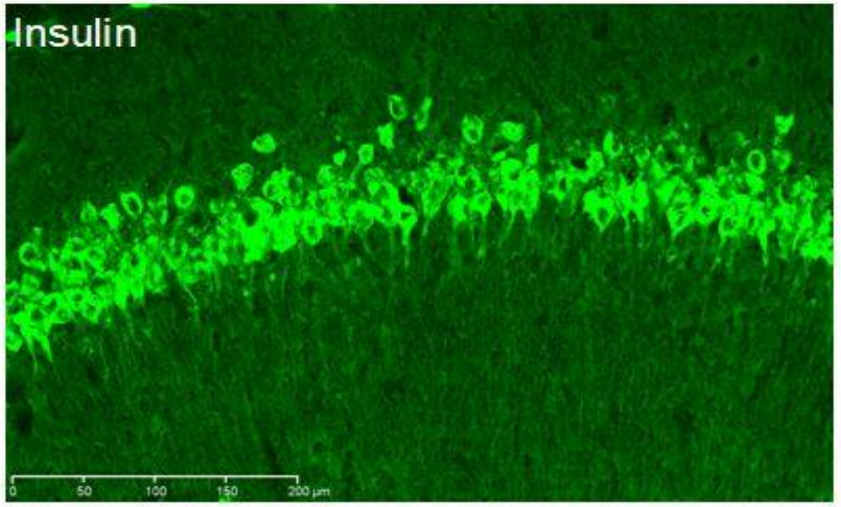
Danielyan, et al., 2014

Intranasal Insulin for TBI

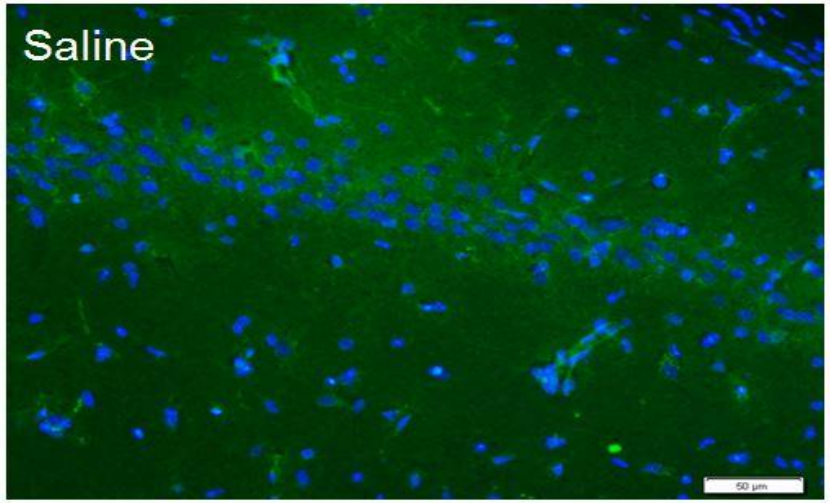
- Improves brain ATP production
- Decreases CSF cortisol
- Improves neuronal viability in the hippocampus
- Increases the expression of anti-inflammatory microglia
- Reduces beta-amyloid and tau protein deposition

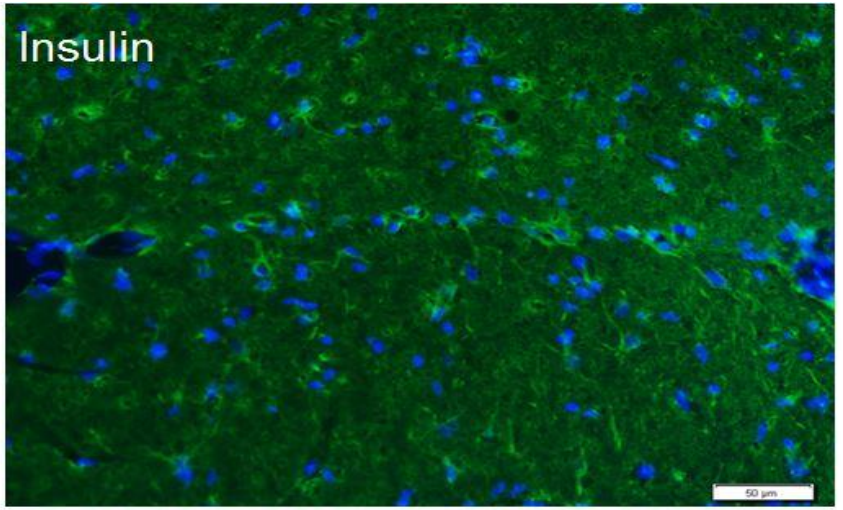
Improved neuronal viability in the hippocampus of the insulin treated rats.





Intranasal insulin increases the expression of anti-inflammatory microglia in the hippocampus





Brabazon, Khayrullina, Frey, & Byrnes, 2014

Platelet Rich Plasma (PRP)

- Autologous plasma contains growth factors and cytokines to aid the injured brain:
  - VEGF, EGF increases angiogenesis
  - PDGF, TGF-p enhance collagen growth
  - IGF-1 stimulates protein synthesis

Platelet Rich Plasma (PRP)

The infusion of concentrated platelets results in an exponential increase in numerous growth factors at the sight of infusion

Plasma cytokines control inflammatory mediators cox1, cox2 and guide stem cells to areas of injury

Intranasal Platelet Rich Plasma (PRP) for TBI

- "Basic fibroblast growth factor infusion enhances injury-induced cell proliferation in the dentate gyrus and improves cognitive function in rats following fluid percussive injury."

- "Other studies have found that infusion of S100β or VEGF can also enhance neurogenesis in the hippocampus and improve the functional recovery of animals following TBI."

Sun, 2014

Peripheral Blood Based Adult Stem Cells

Recently discovered in peripheral blood

PLURIPOTENT adult stem cells

Behave like embryonic stem cells

Give rise to all the cell types

Long lifespan

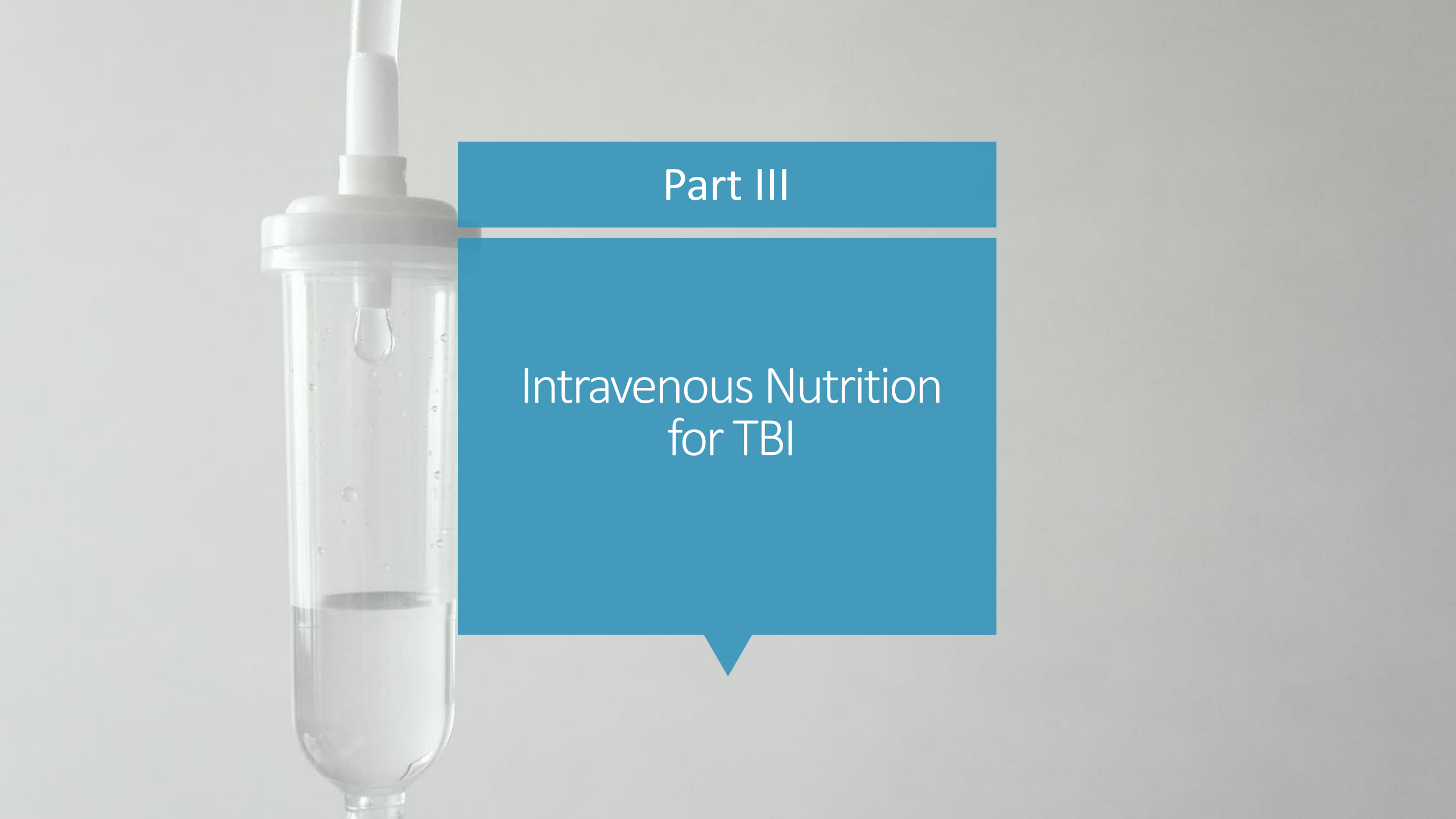
Work in combination with PRP

Intranasal Peripheral Blood Stem Cells for TBI

- Have regenerative and reparative properties
- Have been used to treat ischemic brain damage by reducing gray and white matter loss
- Downregulate neuroinflammatory cytokines

Intranasal Nutrients for TBI

- IN glutathione has been used to reduce oxidative stress and enhance cellular detoxification in Parkinson's disease patients.
- IN methylcobalamin has been shown to improve QEEG Theta activity in ADHD and autism patients.



Part III

Intravenous Nutrition for TBI

IV Nutrition for TBI

- PRP
- Adult peripheral blood stem cells
- NAD+
- Myer's cocktail with potassium, magnesium, calcium, B-complex, B5, B6, and B12, ascorbate, and glutathione

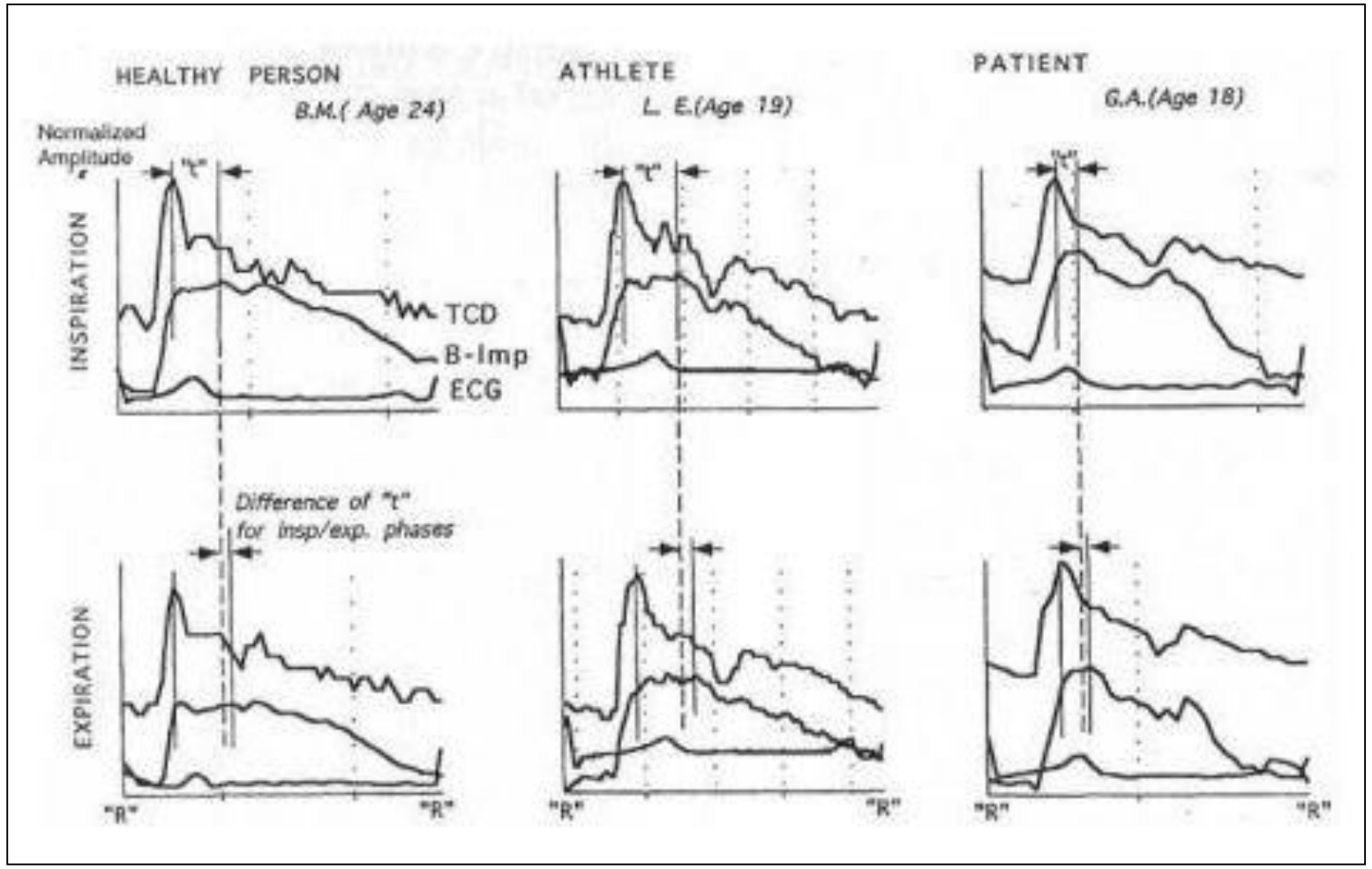
Part IV
Cranial Osteopathy for TBI

Cranial Osteopathy for TBI

- Manual manipulation of the cranial bones and membranes to allow the cerebral spinal fluid to flow properly
- The central nervous system, including the brain and spinal cord, has a subtle, rhythmic pulsation

Cranial Osteopathy for TBI

- This rhythmic pulsation can be blocked in brain injuries - impedes CSF and blood flow
- Effective at treating vertigo and headaches associated with TBIs



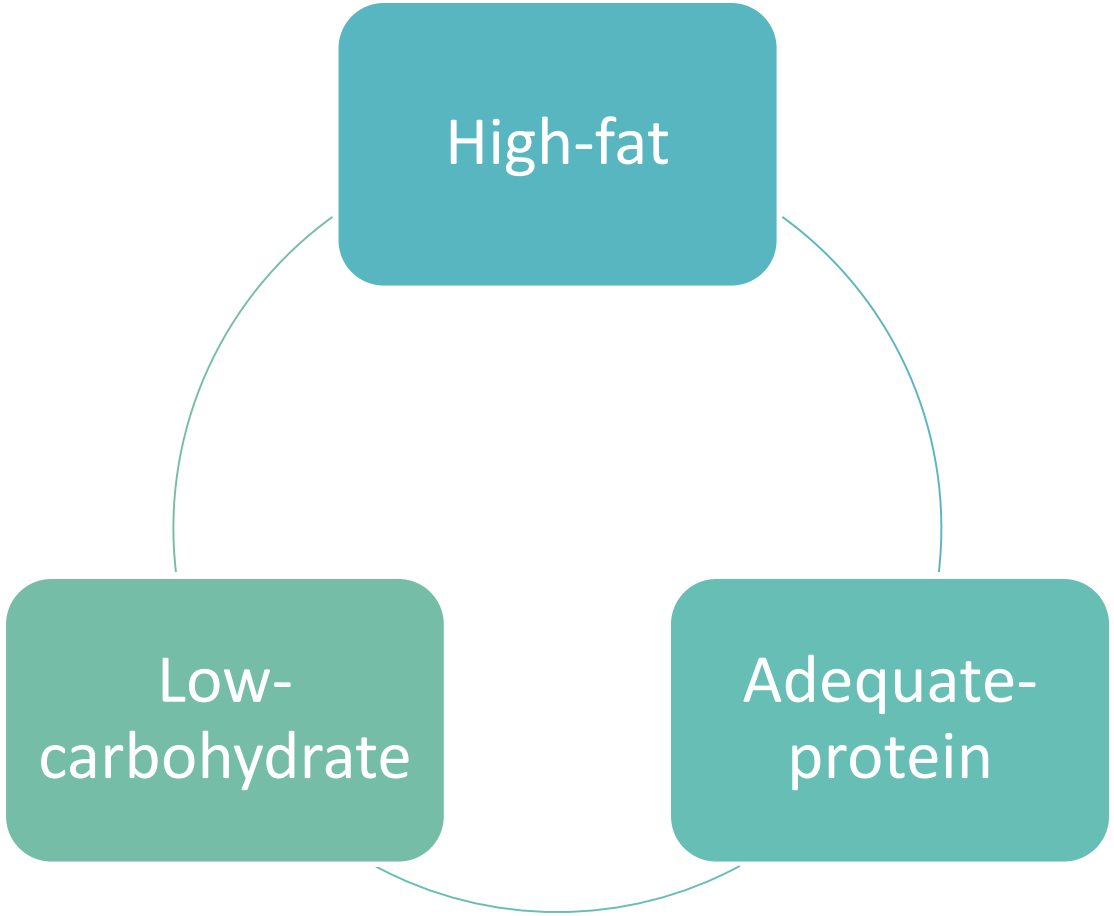
- Time shift between peaks of TCD and B-Imp is determined by the replacement of some portion of CSF out from (or into) zone of B-Imp electrodes.
- This time interval represents the mobility of CSF inside the cranium during the pulse cycle.

Moskalenko, Frymann, Kravchenko, & Weinstein, 2003

Part V

MCT Oils and the Ketogenic Diet for TBI

Ketogenic Diet for TBI



Ketogenic Diet for TBI

DO NOT EAT

- Grains – wheat, corn, rice, cereal, etc.
- Sugar – honey, agave, maple syrup, etc.
- Fruit – apples, bananas, oranges, etc.
- Tubers – potato, yams, etc.

DO EAT

- Meats (organic, pasture-raised, sustainable)
- Above ground vegetables and leafy greens
- High fat dairy
- Nuts and seeds
- Avocado and berries
- Other fats – avocado oil, coconut oil, grass-fed ghee, high-fat salad dressing, saturated fats, etc.

Which burns more even?

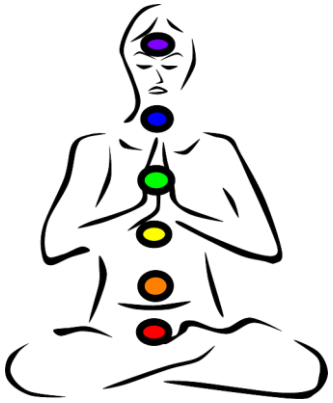
Glucose/Carbohydrates = Kindling

Ketones/Fats = Logs

Ketones are like diesel fuel (Glucose is like gasoline)

- Diesel fuel has a high flash point than gasoline
- Harder to oxidize – Less flammable (excitable)
  - The brain works like a diesel engine
  - Burns more efficiently – lasts longer

Increases Neuroprotection



Increases GABA

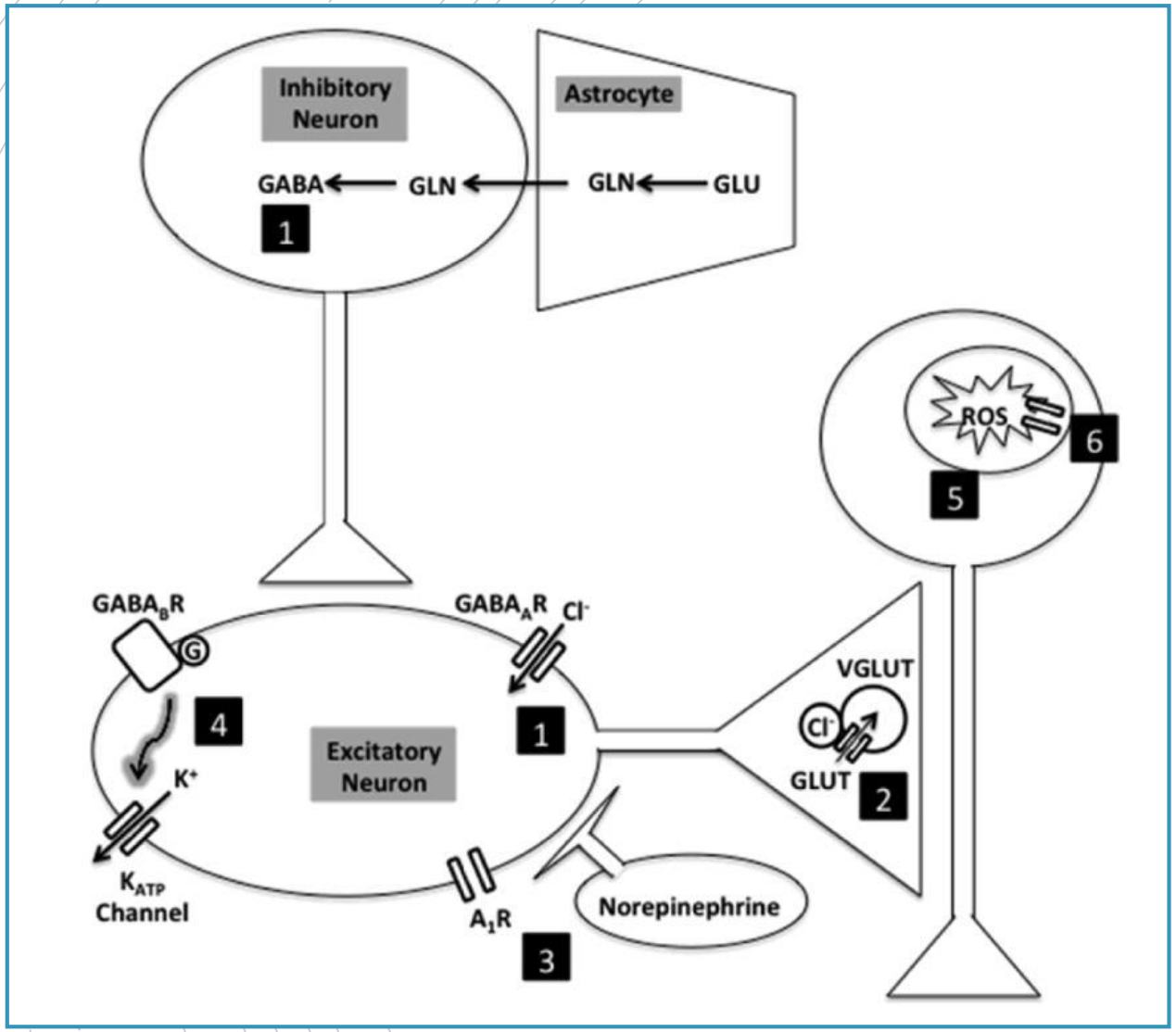
Increases Calming

What else do ketones do?

Decreases Glutamate

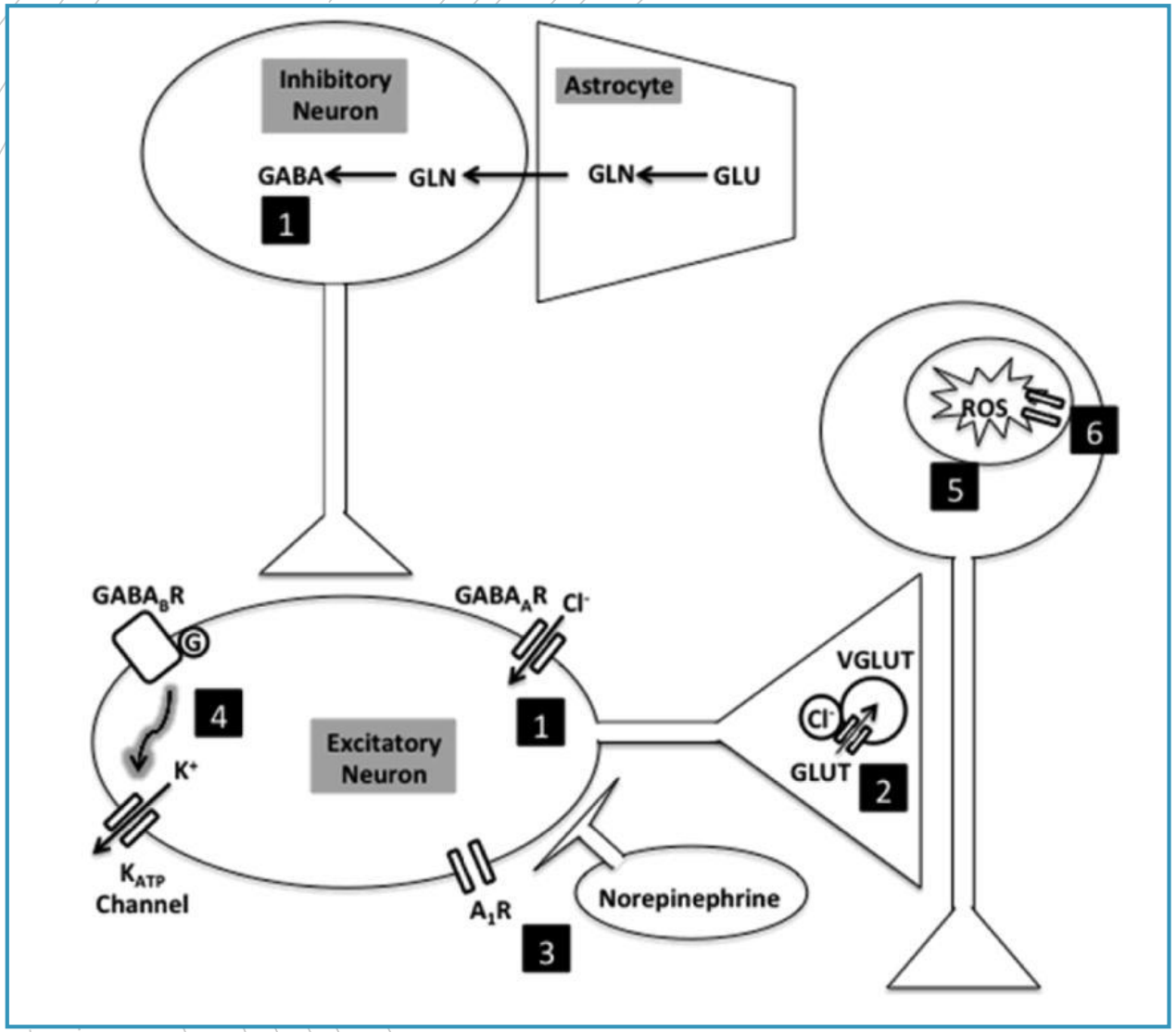
Decreases Depression,
Fear, Anxiety

Decreases Oxidative Stress



- Possible anticonvulsant effects of ketone bodies on the brain
- Increased GABA synthesis
- Decreased glutamate release by competitive inhibition of vesicular glutamate transporters.

McNally & Hartman, 2012



- Increased membrane potential hyperpolarization via KATP channels
- Decreased reactive oxygen species production from glutamate exposure
- Electron transport chain subunit transcription

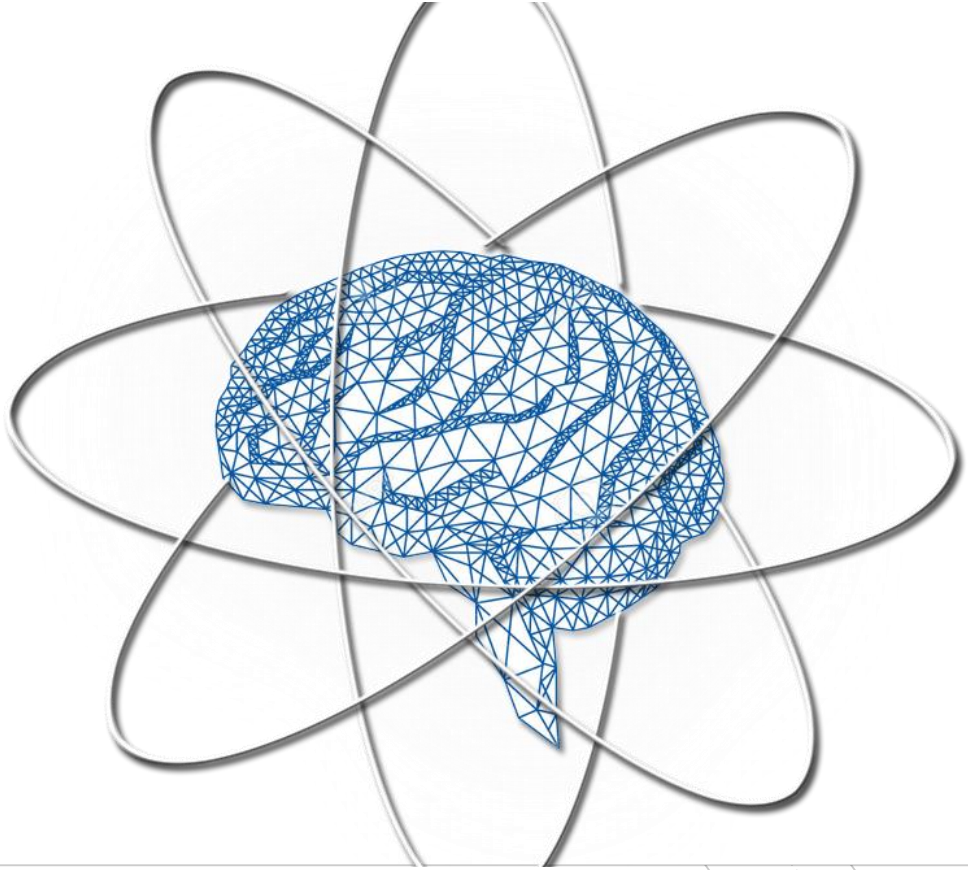
McNally & Hartman, 2012

Neuroprotective Actions of the Ketogenic Diet

- Increases resistance to metabolic stress
- Increases resilience to neuronal loss
- Upregulates energy metabolism genes
- Stimulates of mitochondrial biogenesis

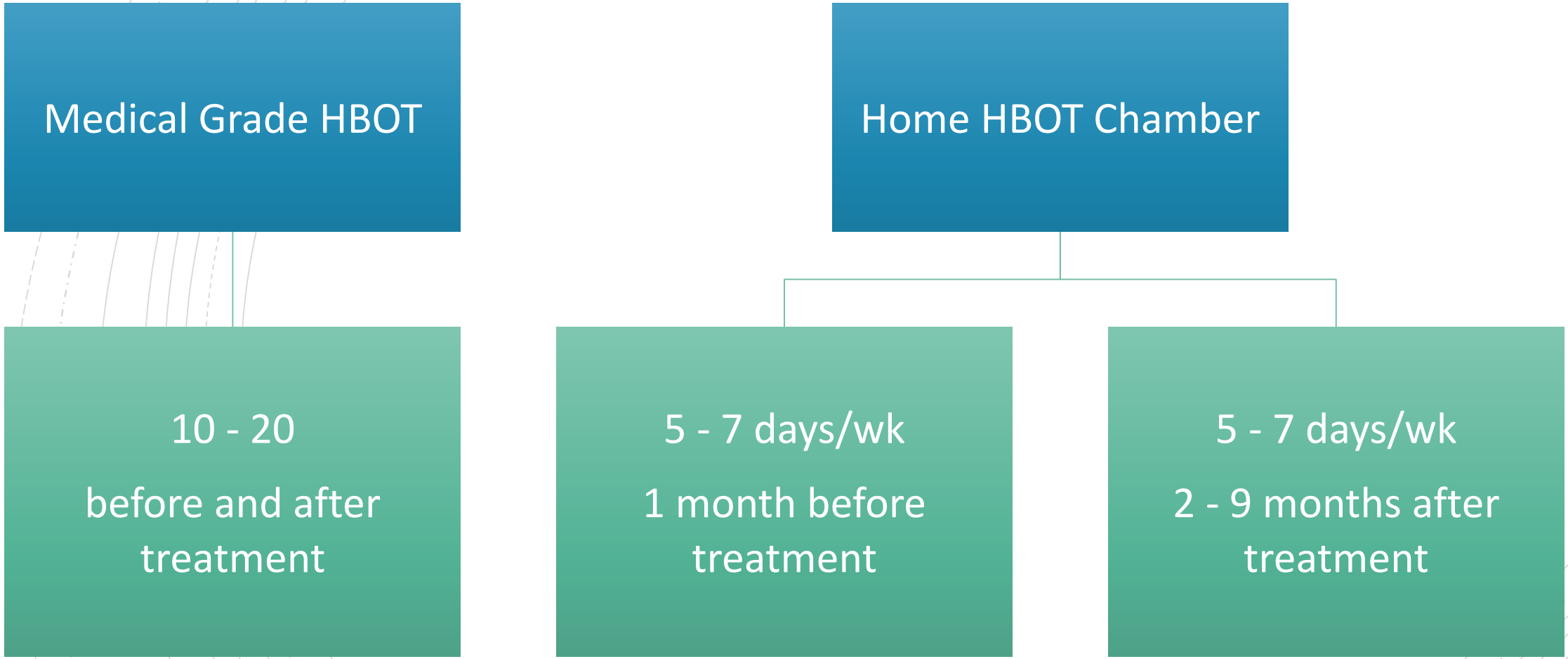
Neuroprotective Actions of the Ketogenic Diet

- Enhances alternative energy substrates
- Promotes synthesis of ATP
- Interferes with glutamate toxicity
- Bypasses the inhibition of complex I in the mitochondrial respiratory chain



The TBI Therapy Protocol

TBI Therapy HBOT Protocol



TBI Therapy 2-Day Program

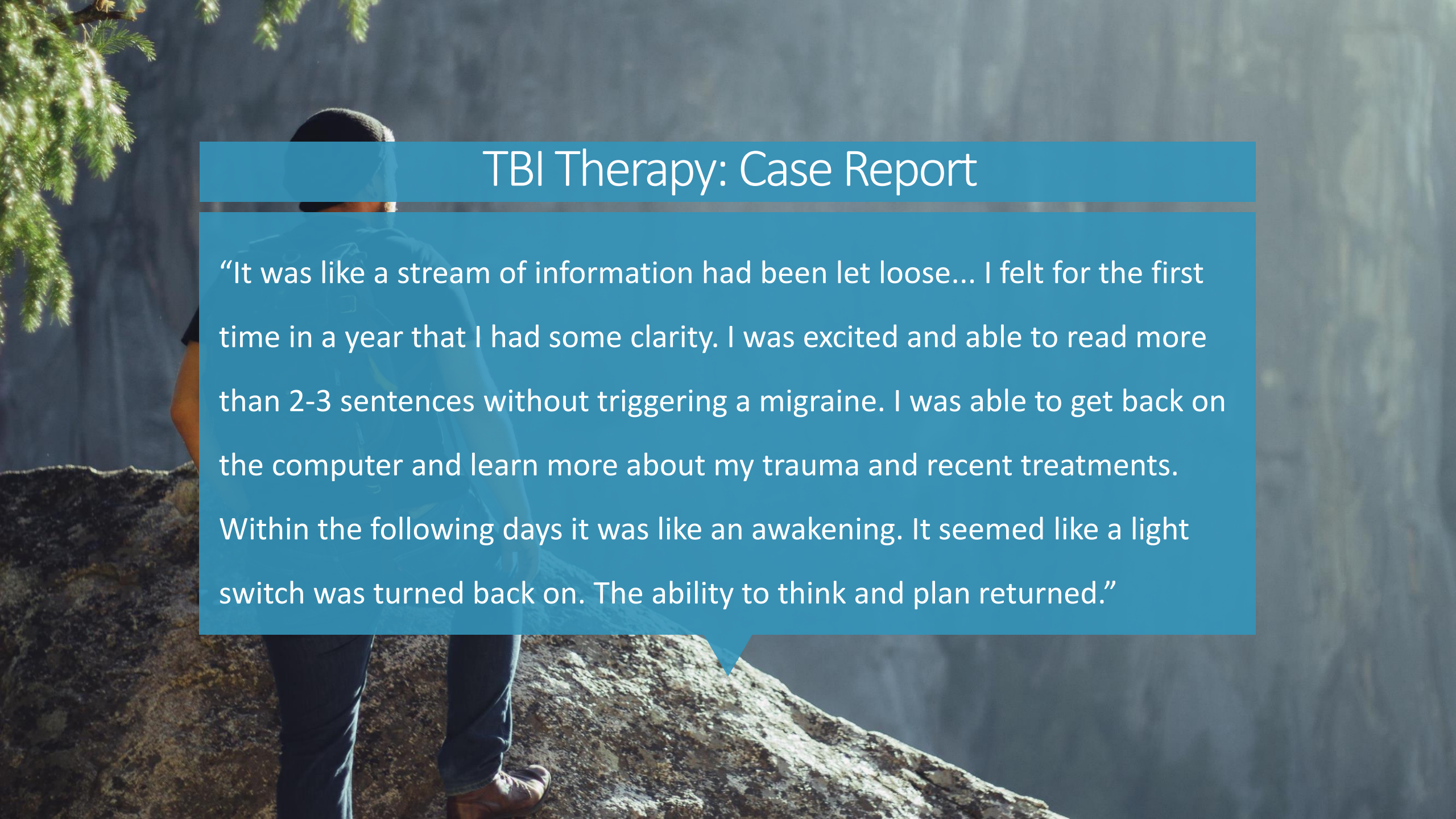
Day 1:	Consultation HBOT Cranial therapy IV therapy Intranasal (IN) PRP and insulin
Day 2:	IV and IN NAD+ IV and IN pluripotent stem cells (VESLs) from the blood HBOT

TBI Therapy 3-Day Program

Day 1:	Consultation HBOT Cranial therapy IV therapy Intranasal (IN) PRP and insulin
Day 2:	IV and IN NAD+ IV and IN pluripotent stem cells (VESLs) from the blood HBOT
Day 3:	IV and IN pluripotent stem cells (VESLs) from the blood HBOT

Case Report: 46 year-old male

Before Treatment:	After Treatment:
• Memory loss • Depression and anxiety • Emotionally unstable • Headaches daily • Inability to carry on conversation • Inability to do math or read • Light and sound sensitivity • Could not drive • Insomnia	• “Memory download” • “An awakening” • Mood and personality improvements • Improvements intellectually, physiologically, and psychologically • Improved ability to read • Able to turn on lights /electronics • Able to drive • Sleep normalized



TBI Therapy: Case Report

"It was like a stream of information had been let loose... I felt for the first time in a year that I had some clarity. I was excited and able to read more than 2-3 sentences without triggering a migraine. I was able to get back on the computer and learn more about my trauma and recent treatments. Within the following days it was like an awakening. It seemed like a light switch was turned back on. The ability to think and plan returned."

TBI Therapy: Case Report

“I felt well enough that I started saying yes again. I also experienced relief from anxiety. TBI Therapy has turned me into a TBI THRIVER, not just a survivor. I’m happy. I enjoy life again, can travel and am doing work in the world that’s more aligned with myself than ever.”

TBI Therapy: Case Report

"The results for me have been are nothing short of MIRACULOUS! Popeye may have his spinach but I have stem cells and PRP! Yes, my brain is strong!"

Out of 100 patients treated, nearly every patient reports:

More mental clarity

Improved memory

Improved executive function/decision making

More stable emotions and less stress

Better ability to cope with pain

More physical and mental energy

Out of 100 patients treated, some patients report:

Less sound and light sensitivity

Improved eyesight

Improved sleep and libido

Improved motor function

(ability to open a clenched fist, ability to walk)

Less muscle spasticity

Conclusion: The Multimodal, Regenerative Approach is a Superior Way to Treat TBI

- The practical, effective combination of multiple regenerative TBI therapies can produce synergistic benefits to the patient superior to mainstream TBI or single modality TBI treatments

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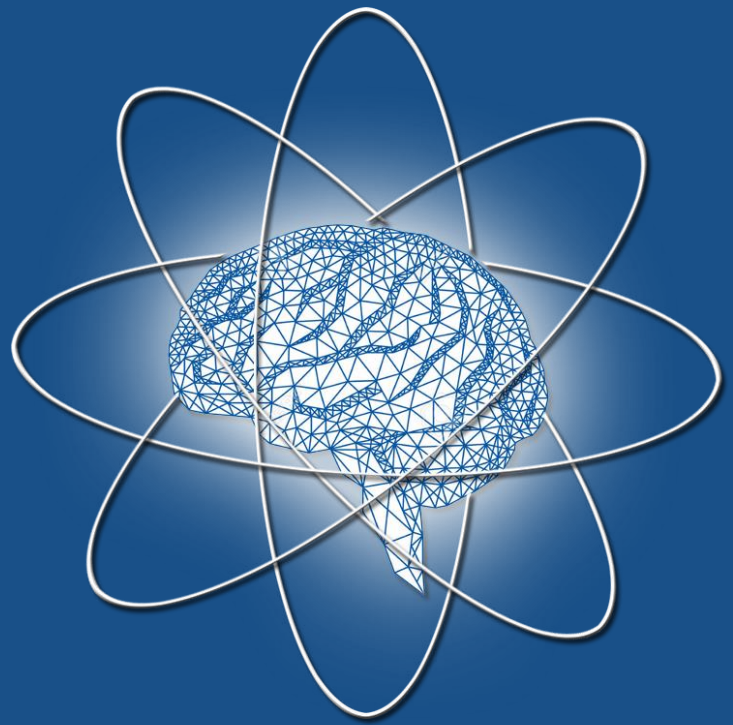

TBI Therapy

Treats TBI patients by combining regenerative therapies: HBOT, stem cells, PRP, and nutritional therapies.

tbitherapy.com

ASPEN INTEGRATIVE MEDICINE

Treats chronic pain and major medical problems using modern and natural medicine.

aspenintegrativemedicine.com