

# Neuroplasticity, Brain Reserve, and Aging:

*How to Improve Your Brain Performance*

*by Growing the Size of Your Hippocampus,*

*At Any Age*

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# Neuroplasticity in the Human Brain: Translational Research



## Modifiable factors that alter the size of the hippocampus with ageing

*Majid Fotuhi, David Do and Clifford Jack*

**Abstract** | The hippocampus is particularly vulnerable to the neurotoxic effects of obesity, diabetes mellitus, hypertension, hypoxic brain injury, obstructive sleep apnoea, bipolar disorder, clinical depression and head trauma. Patients with these conditions often have smaller hippocampi and experience a greater degree of cognitive decline than individuals without these comorbidities. Moreover, hippocampal atrophy is an established indicator for conversion from the normal ageing process to developing mild cognitive impairment and dementia. As such, an important aim is to ascertain which modifiable factors can have a positive effect on the size of the hippocampus throughout life. Observational studies and preliminary clinical trials have raised the possibility that physical exercise, cognitive stimulation and treatment of general medical conditions can reverse age-related atrophy in the hippocampus, or even expand its size. An emerging concept—the dynamic polygon hypothesis—suggests that treatment of modifiable risk factors can increase the volume or prevent atrophy of the hippocampus. According to this hypothesis, a multidisciplinary approach, which involves strategies to both reduce neurotoxicity and increase neurogenesis, is likely to be successful in delaying the onset of cognitive impairment with ageing. Further research on the constellation of interventions that could be most effective is needed before recommendations can be made for implementing preventive and therapeutic strategies.

Fotuhi, M. et al. *Nat. Rev. Neurol.* 8, 189–202 (2012); published online 13 March 2012; doi:10.1038/nrneuro.2012.27

## Changing perspectives regarding late-life dementia

*Majid Fotuhi, Vladimir Hachinski and Peter J. Whitehouse*

**Abstract** | Individuals over 80 years of age represent the most rapidly growing segment of the population, and late-life dementia has become a major public health concern worldwide. Development of effective preventive and treatment strategies for late-life dementia relies on a deep understanding of all the processes involved. In the centuries since the Greek philosopher Pythagoras described the inevitable loss of higher cognitive functions with advanced age, various theories regarding the potential culprits have dominated the field, ranging from demonic possession, through 'hardening of blood vessels', to Alzheimer disease (AD). Recent studies suggest that atrophy in the cortex and hippocampus—now considered to be the best determinant of cognitive decline with aging—results from a combination of AD pathology, inflammation, Lewy bodies, and vascular lesions. A specific constellation of genetic and environmental factors (including apolipoprotein E genotype, obesity, diabetes, hypertension, head trauma, systemic illnesses, and obstructive sleep apnea) contributes to late-life brain atrophy and dementia in each individual. Only a small percentage of people beyond the age of 80 years have 'pure AD' or 'pure vascular dementia'. These concepts, formulated as the dynamic polygon hypothesis, have major implications for clinical trials, as any given drug might not be ideal for all elderly people with dementia.

Fotuhi, M. et al. *Nat. Rev. Neurol.* 5, 649–658 (2009); published online 17 November 2009; doi:10.1038/nrneuro.2009.175



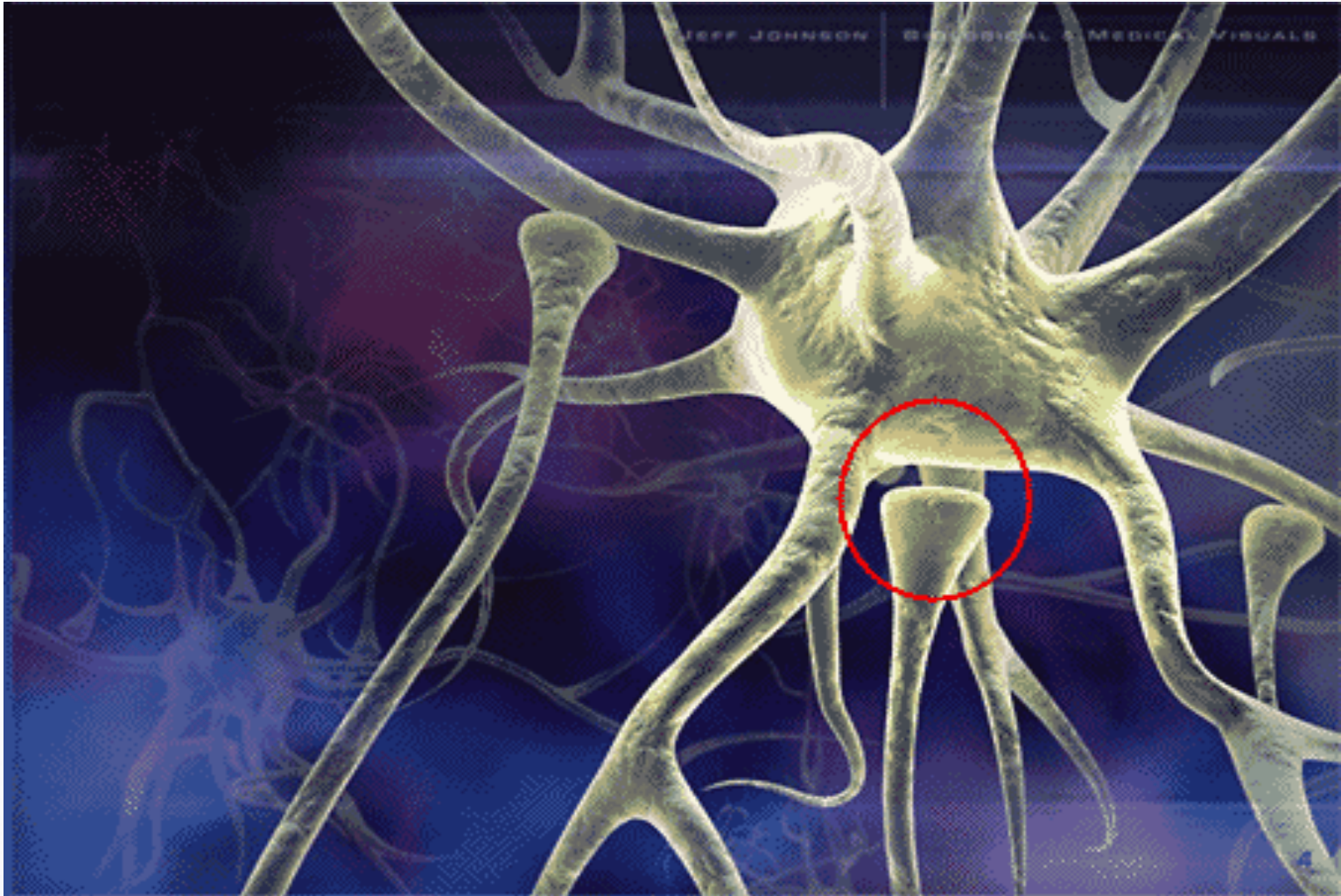
# Objectives

1. Basic Anatomy of Cognition
2. Memory Loss and Brain Atrophy with Aging
3. Brain Reserve & Neurogenesis
4. Six Ways to Grow Your Brain
5. Reversing Brain Atrophy in Elderly, in 3 Months

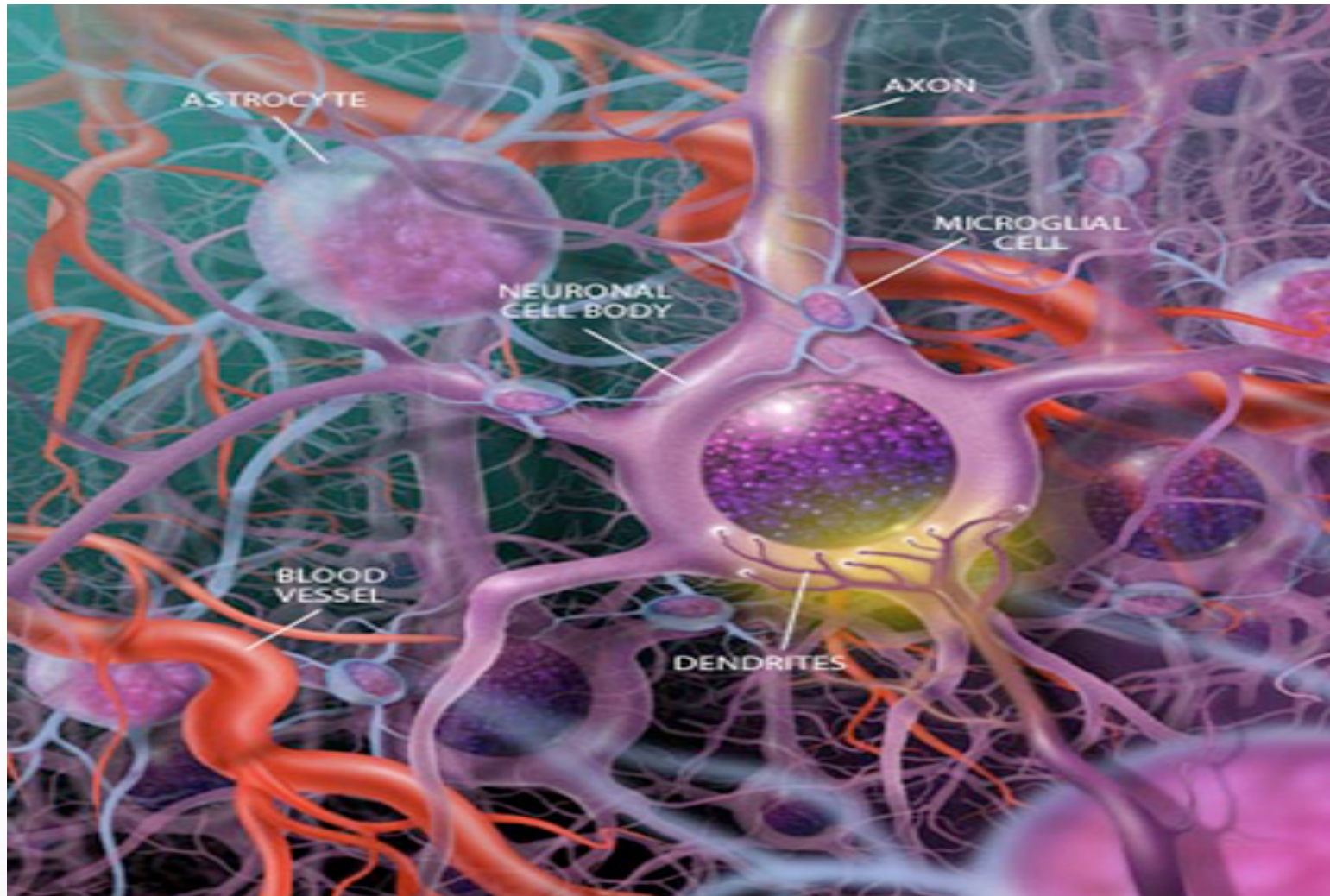




# Synapses

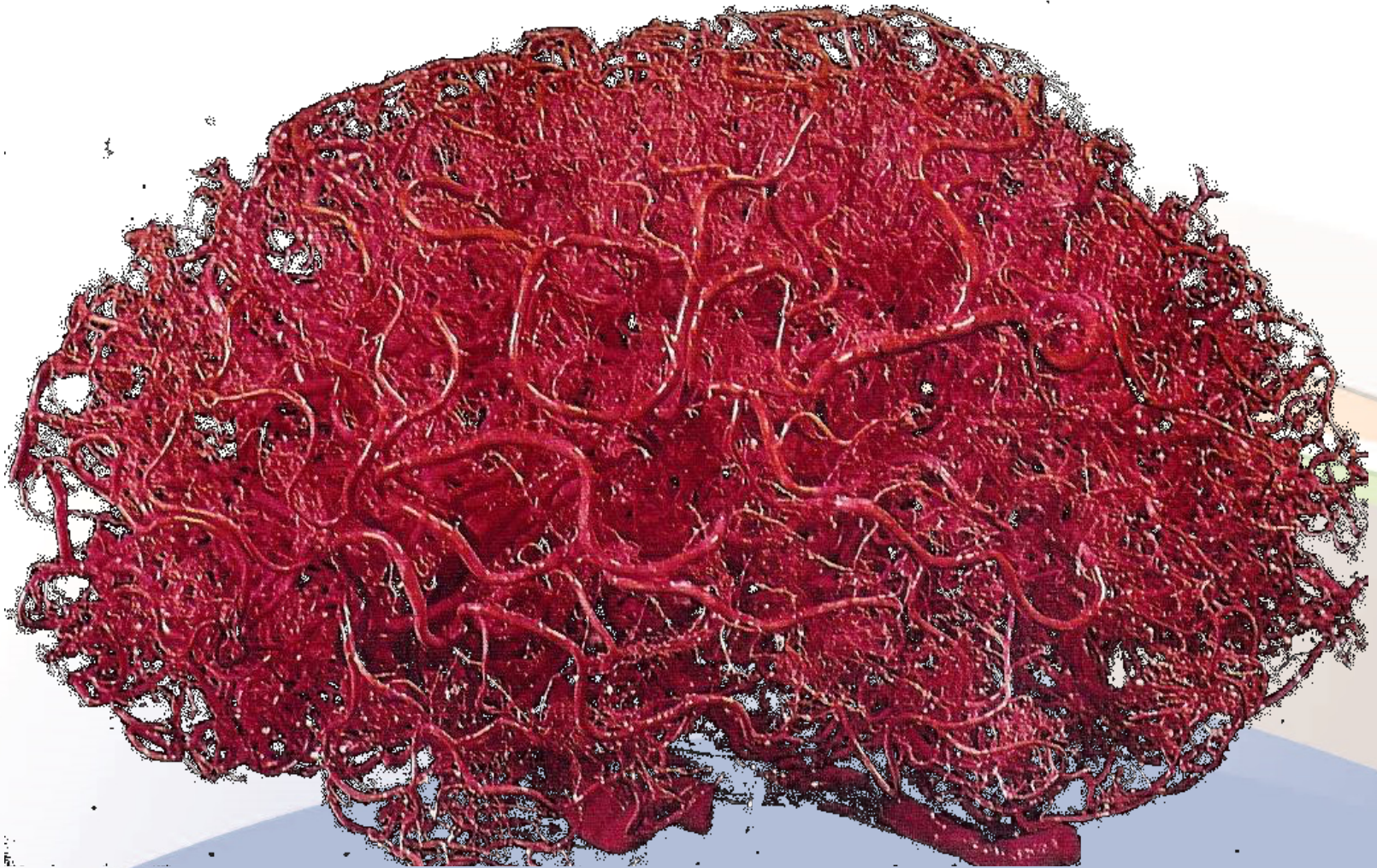


# Blood Vessels





# Brain is a Highly Vascular Organ



# Anatomy of Cognition: Cortex



**Executive function**

**Language**

**Navigation**

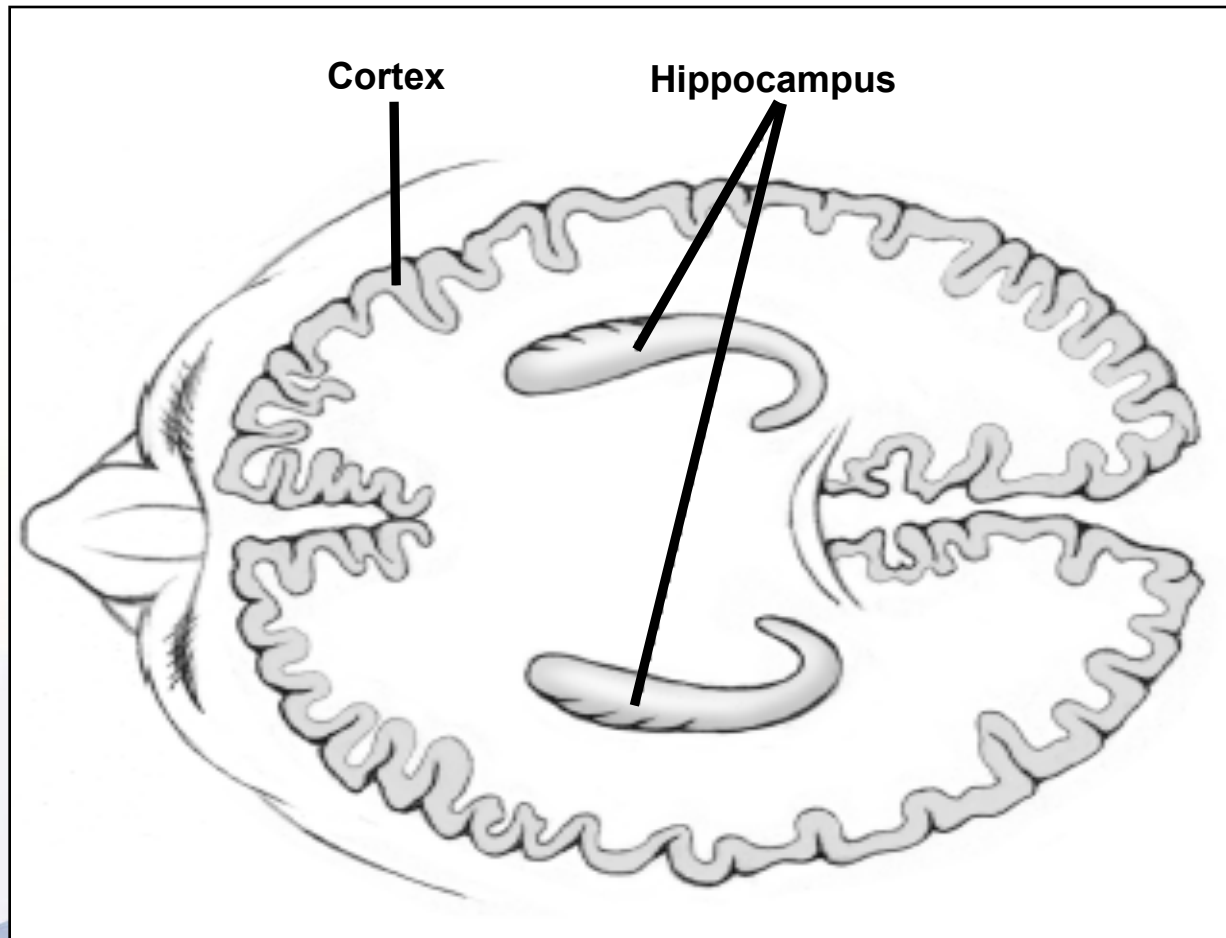
**Calculation**

**Abstract thinking**

**Long term memory**



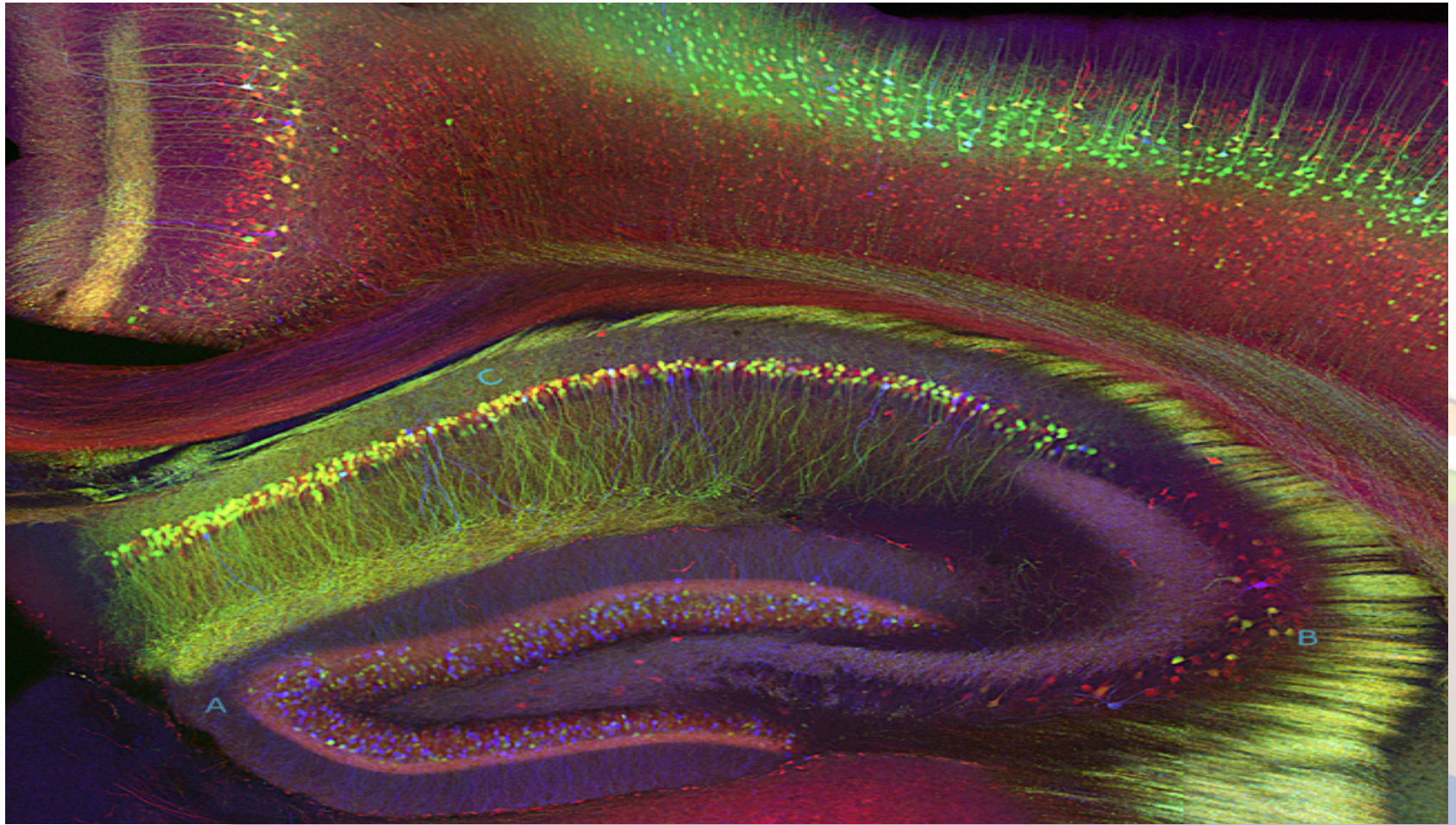
## Short-term Memory & Consolidation: Hippocampus



# Hippocampus



# Neurons in Hippocampus





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# With Aging, Hippocampus Atrophies Faster than the Rest of the Brain

- Hippocampus shrinks by about 0.5% per year after age 40.
- That is the main reason memory lapses become more frequent after age 40.

# What Causes Atrophy in Hippocampus?

**Insomnia**

**Sleep Apnea**

**Obesity**

**Smoking**

**Diabetes**

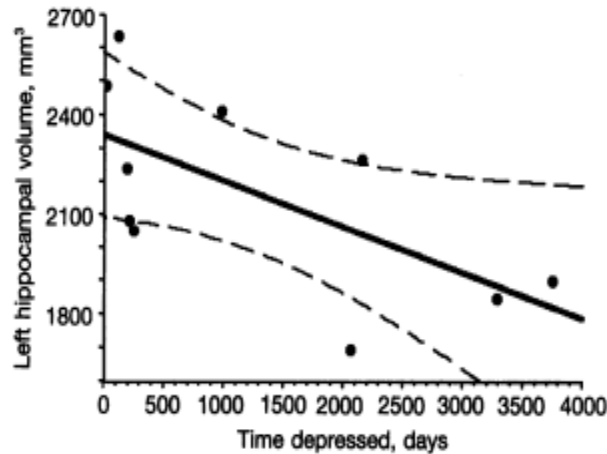
**Concussion**

**Stress**  
**Depression**

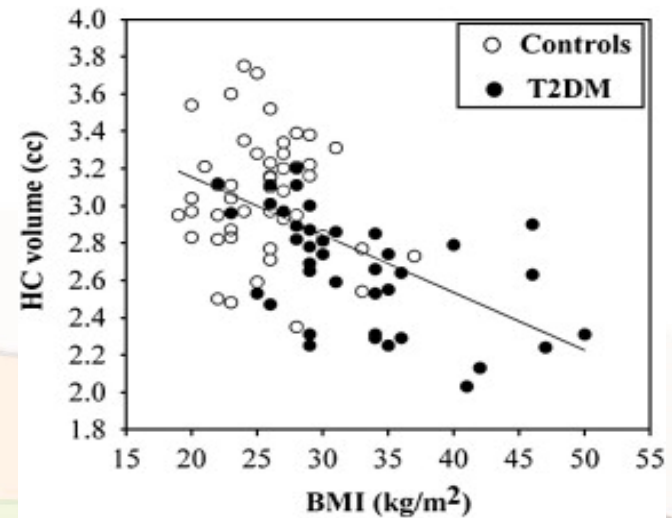
**Alzheimer's**

# More Depression, Obesity, Stress, and Insomnia, Smaller Hippocampus

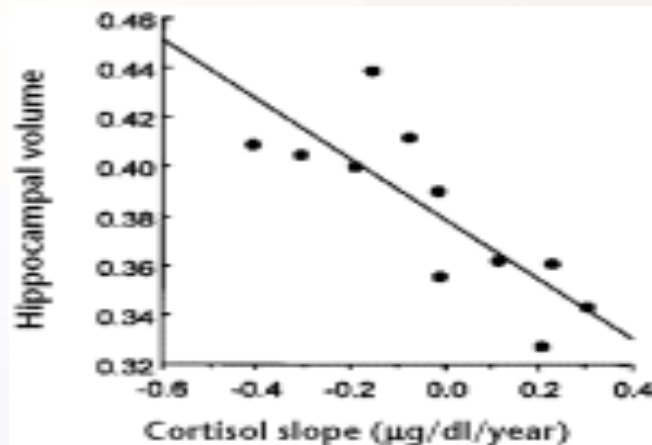
Depression<sup>1</sup>



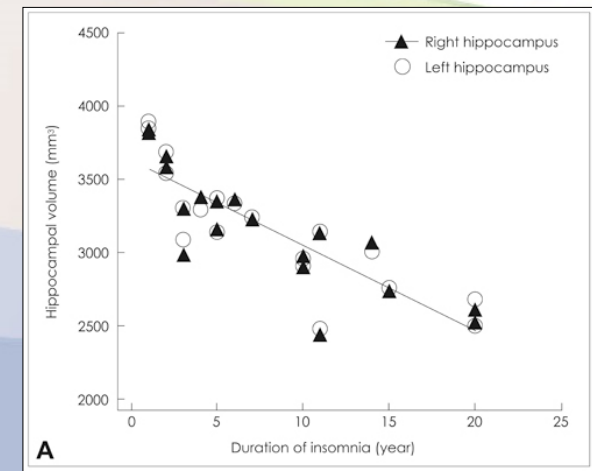
Obesity<sup>2</sup>



Cortisol / Stress<sup>3</sup>



Insomnia<sup>4</sup>



1- Sheline, (1996). *PNAS*, 93(9):3908-13.

2- Brain Research, 2009, Pages 186-194

3- Lupien et al (1998), *Nature NeuroSci* 1 (1), 69-73

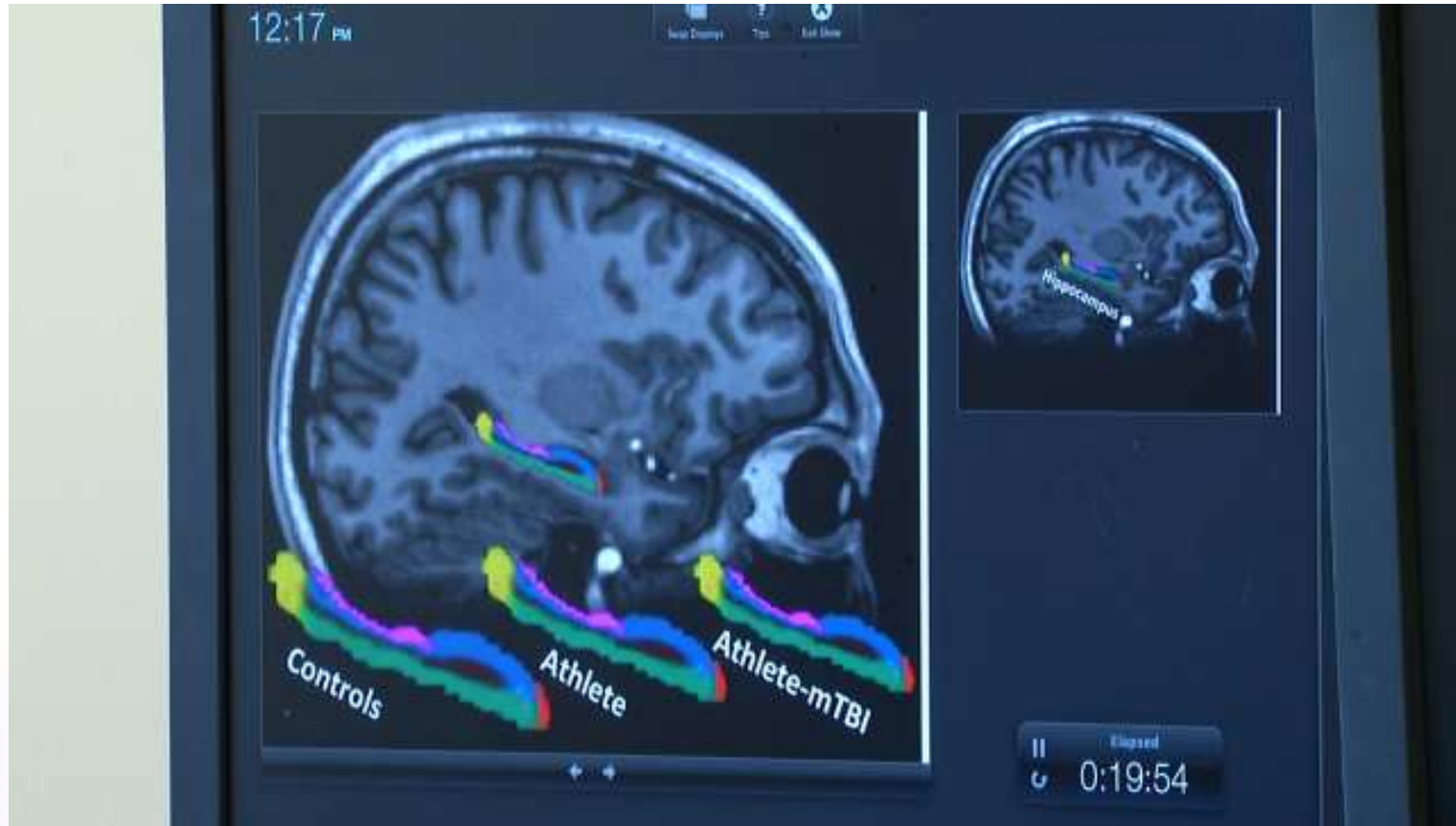
4- Ho et al; *J Clin Neurol*. 2012 Jun;8(2):130-8

# More Traumatic Brain Injury (TBI), Smaller Hippocampus



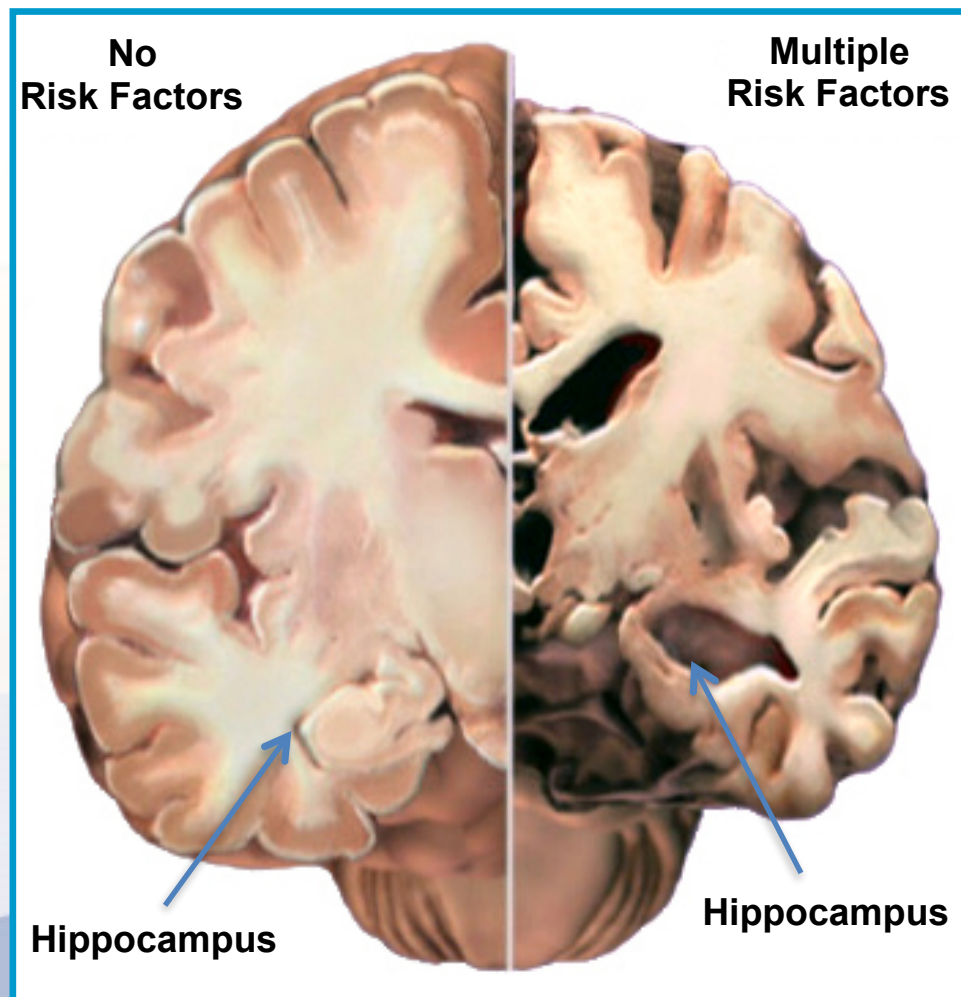


# More Football, Smaller Hippocampus



Singh, JAMA neurology, 2014

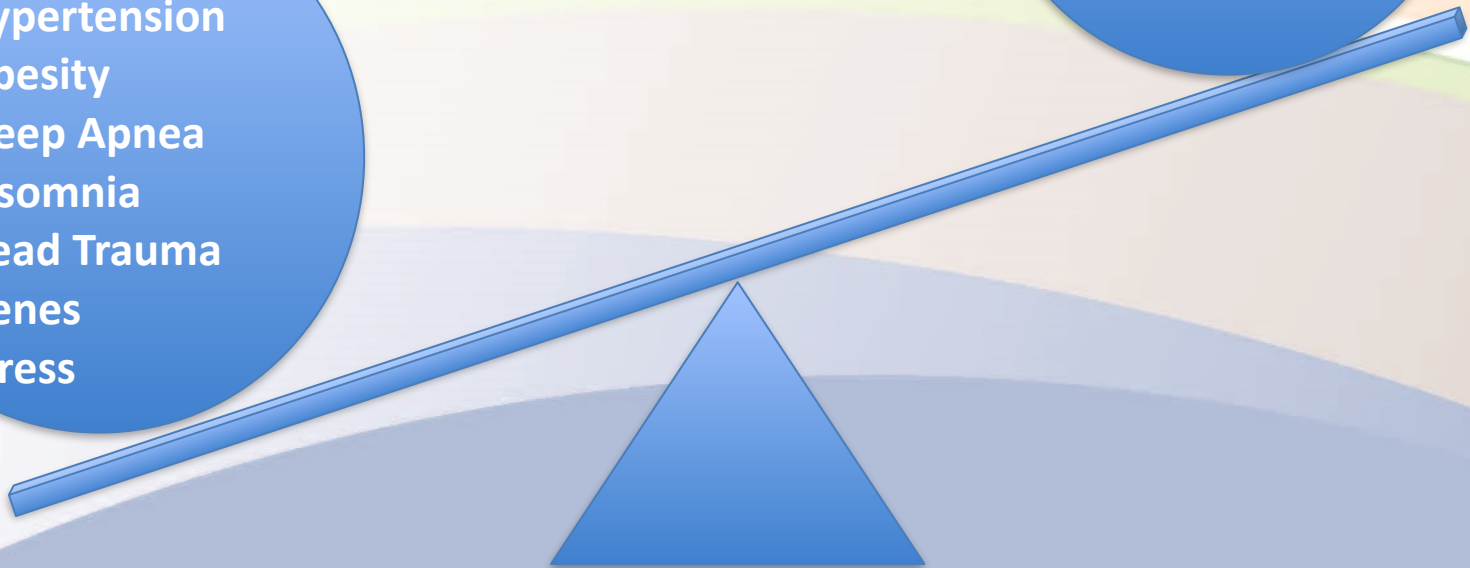
## Multiple Risk Factors, Much Smaller Hippocampus





Diabetes  
Hypertension  
Obesity  
Sleep Apnea  
Insomnia  
Head Trauma  
Genes  
Stress

Brain-healthy Diet  
Physical Fitness  
Cognitive Stimulation  
Quality sleep  
Peace of Mind

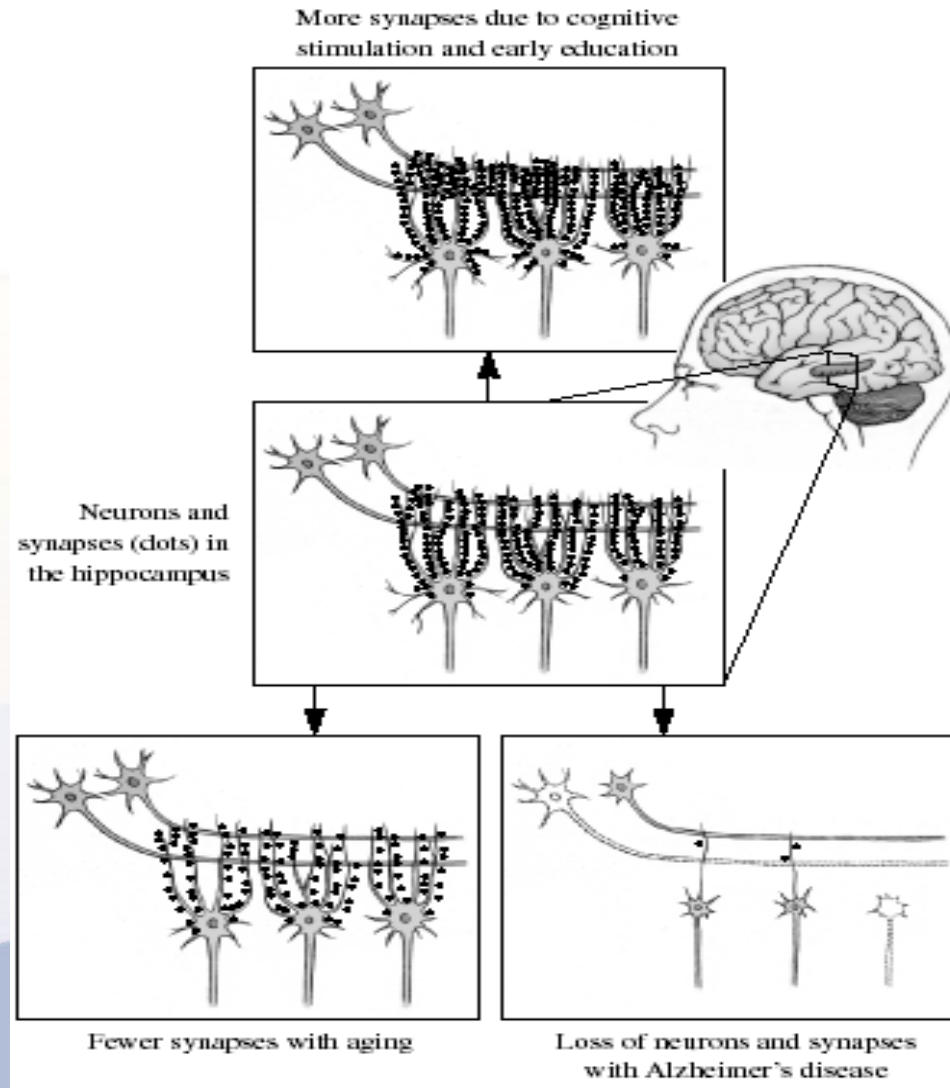


# Objectives

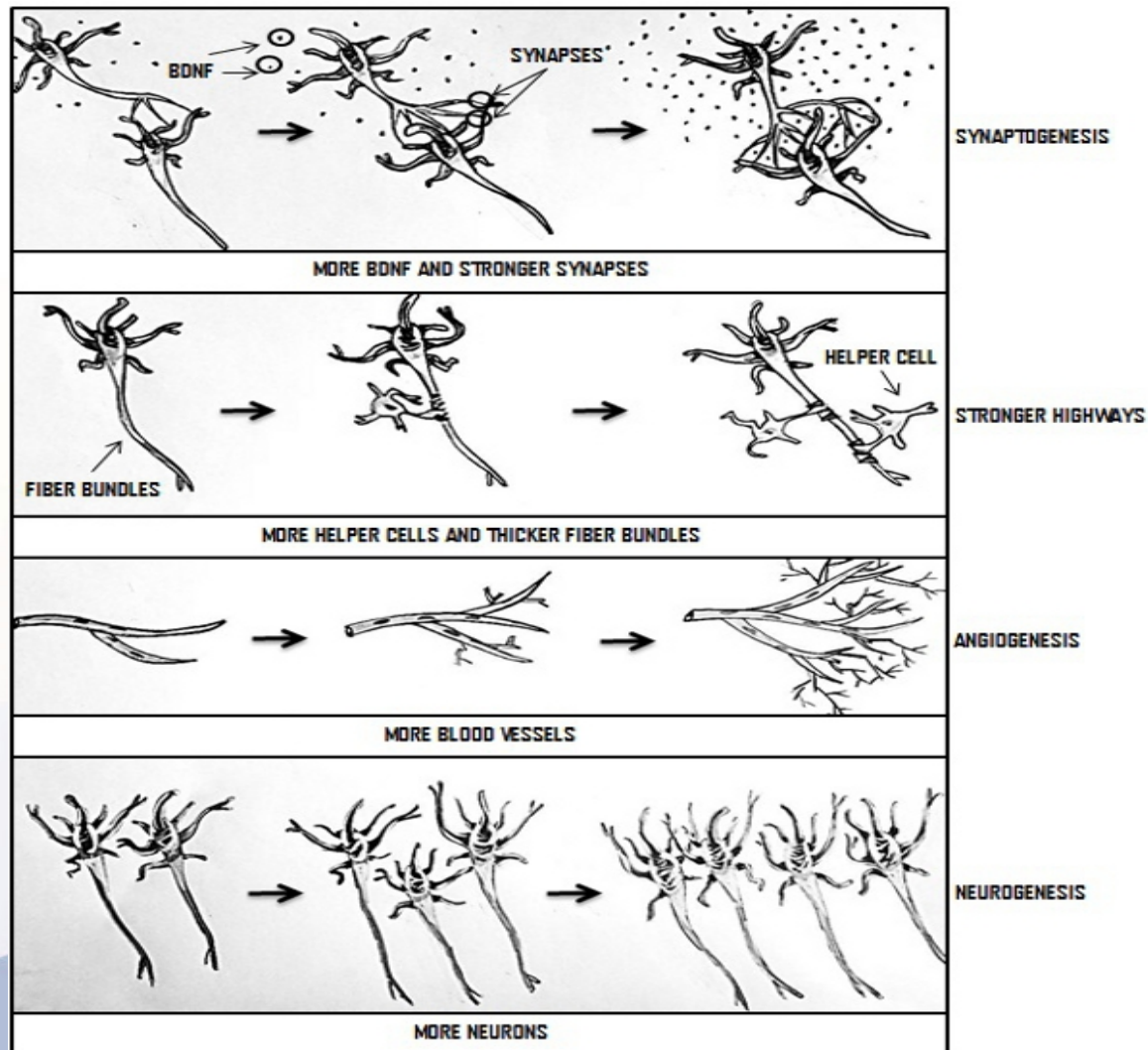
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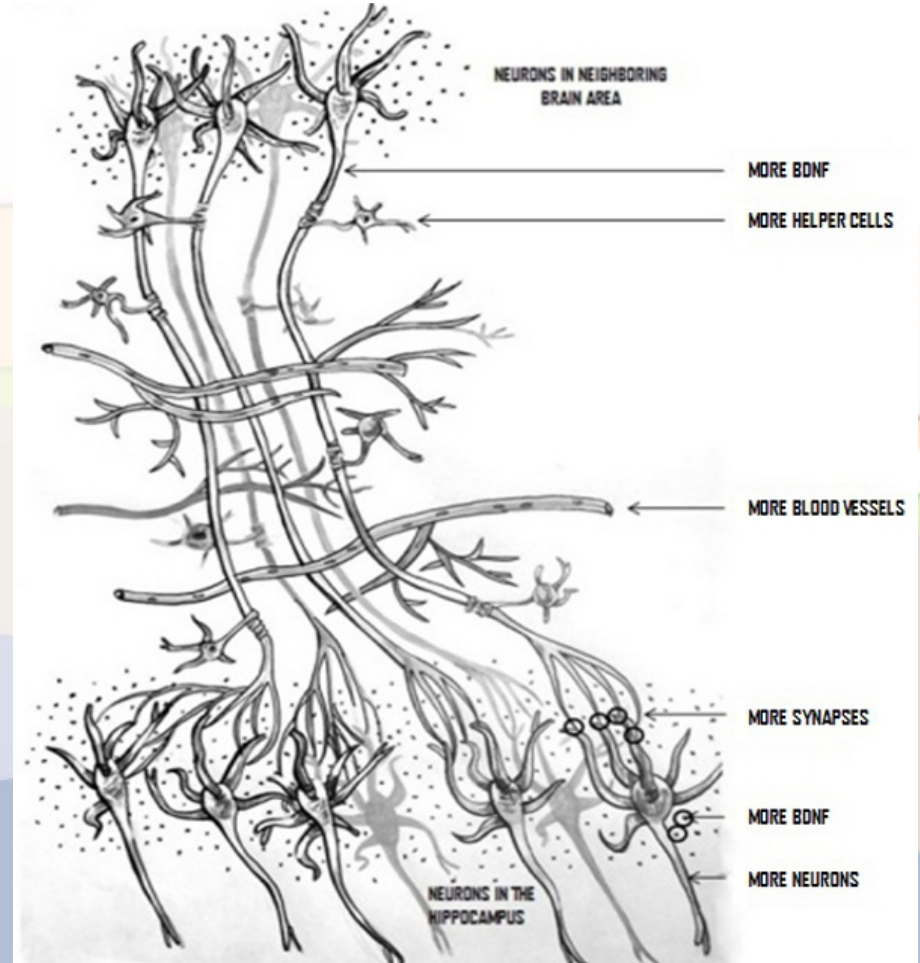
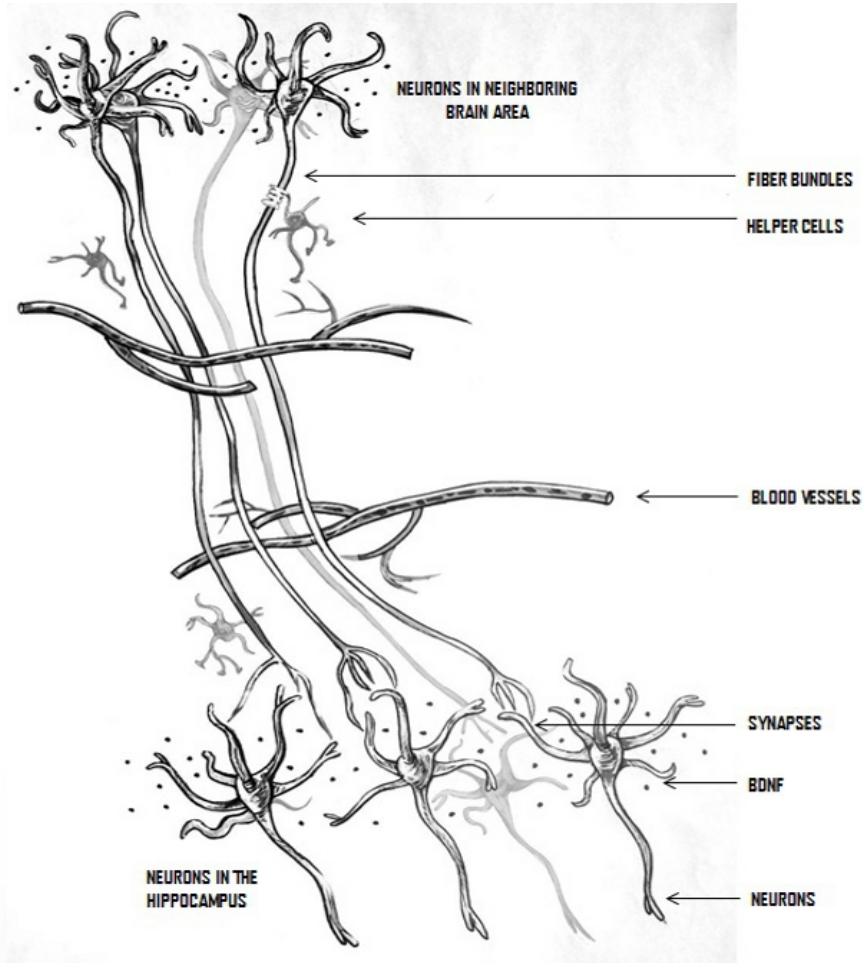
# “Brain Reserve”



# Four Ways to Build a Bigger Brain Reserve

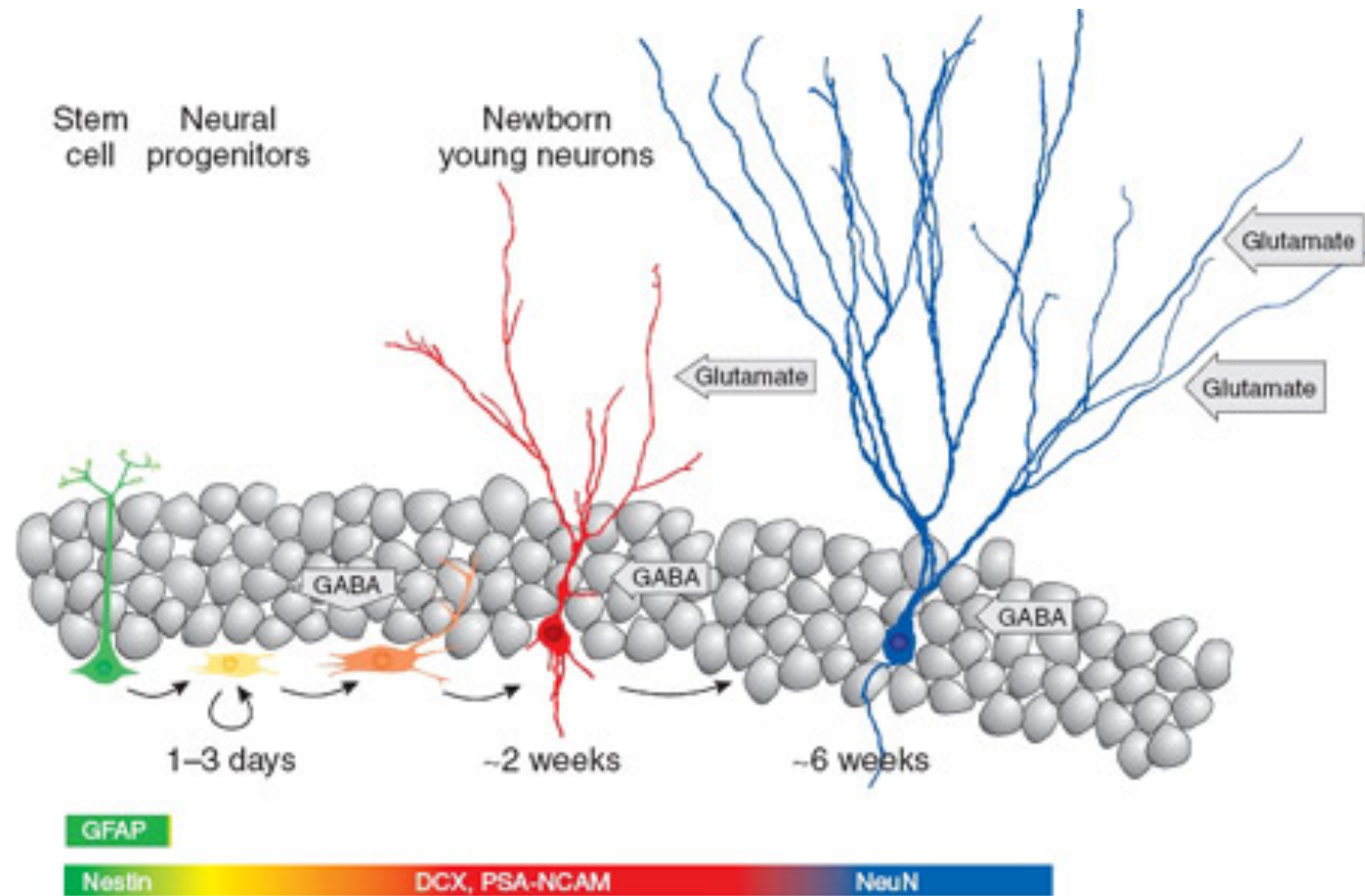


# Before and After



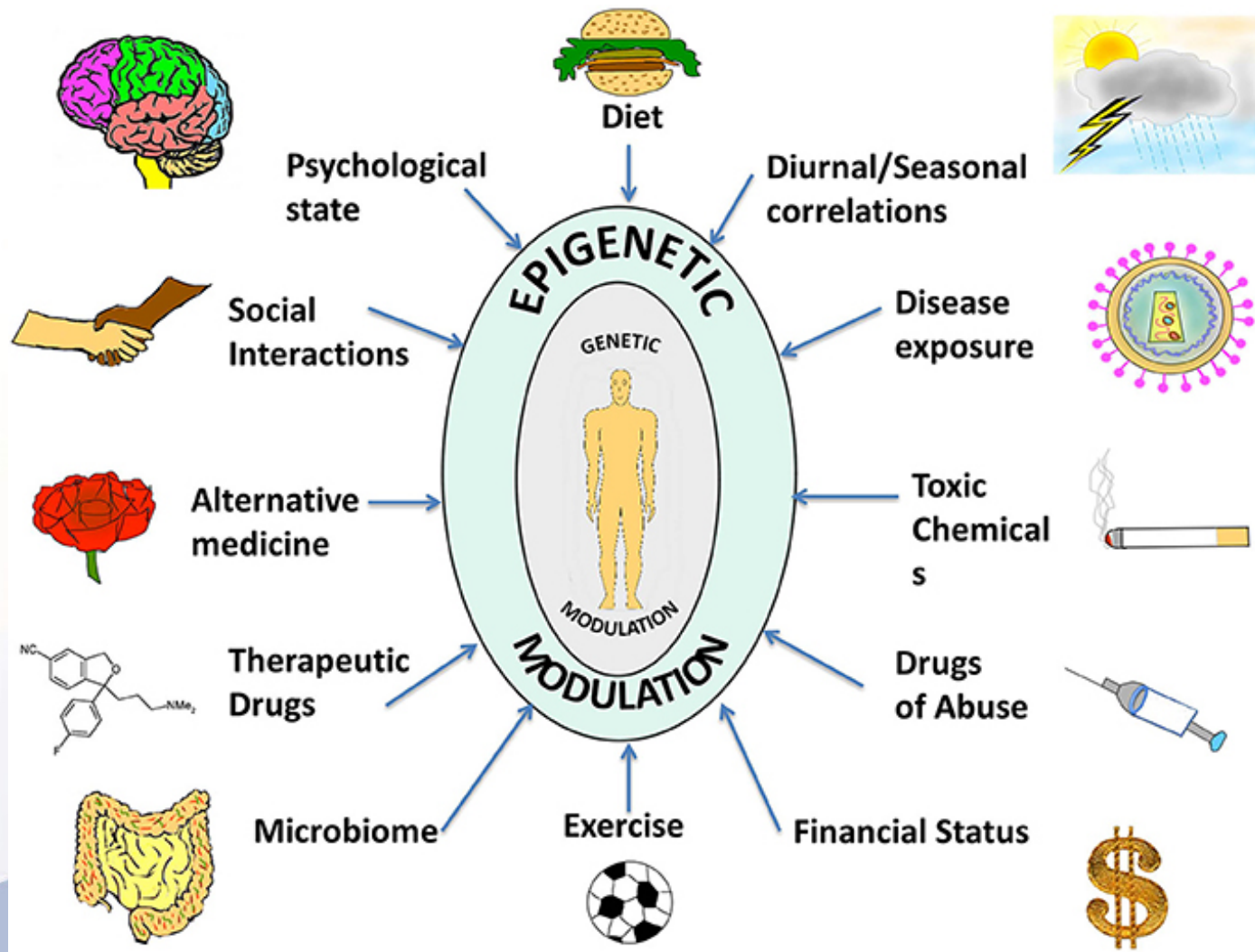


# Neurogenesis in Hippocampus



**Figure 1** Development of newly generated granule cells in the adult hippocampus. Neural progenitors

# Epigenetics: Factors That Modulate Your Genes



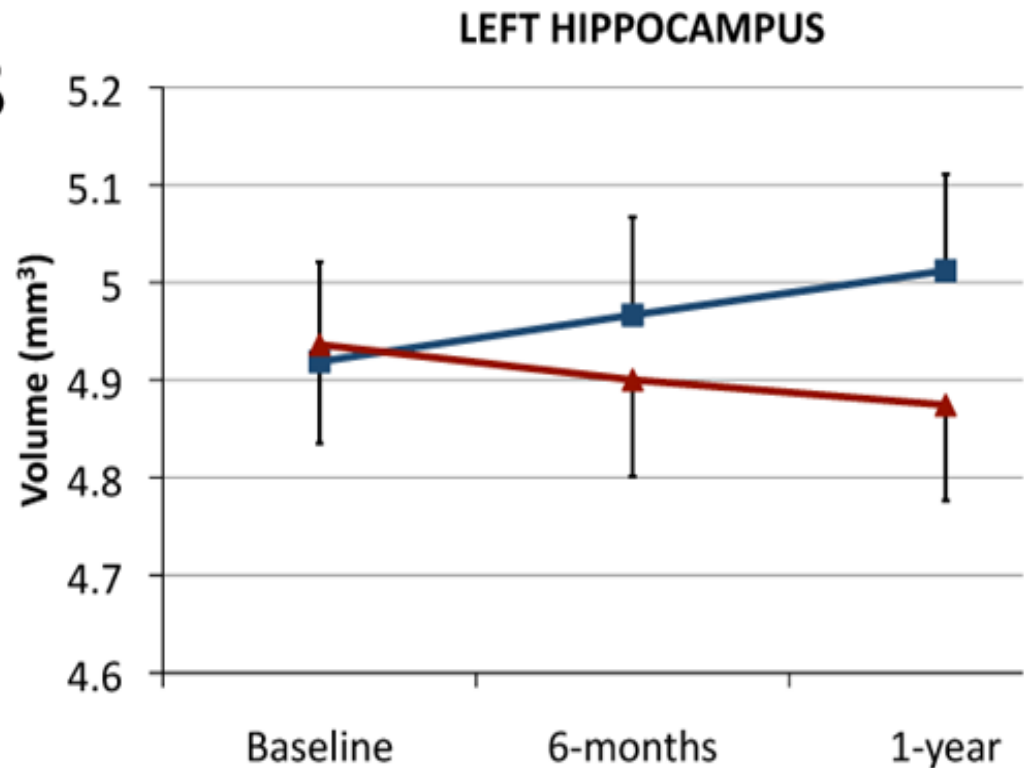
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# 1. More Exercise, Bigger Hippocampus

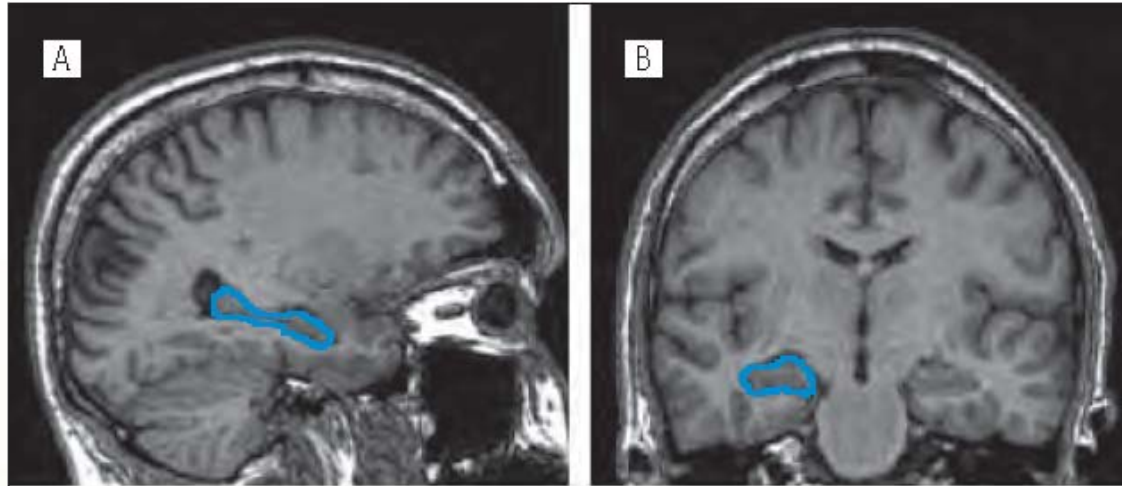
Hippocampus



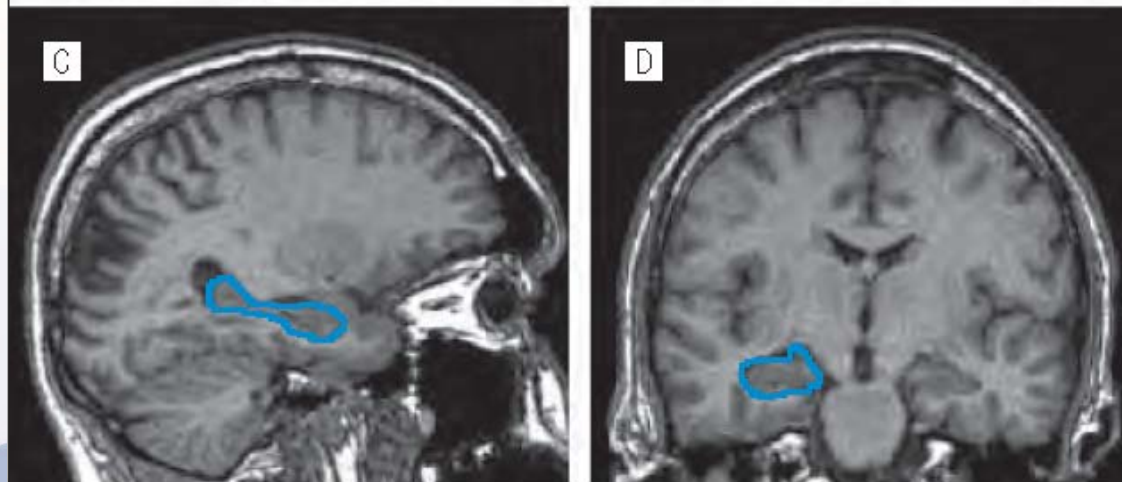
*Erikson, PNAS 2011*

# More Exercise, Bigger Hippocampus, Even After 3 Months

Before

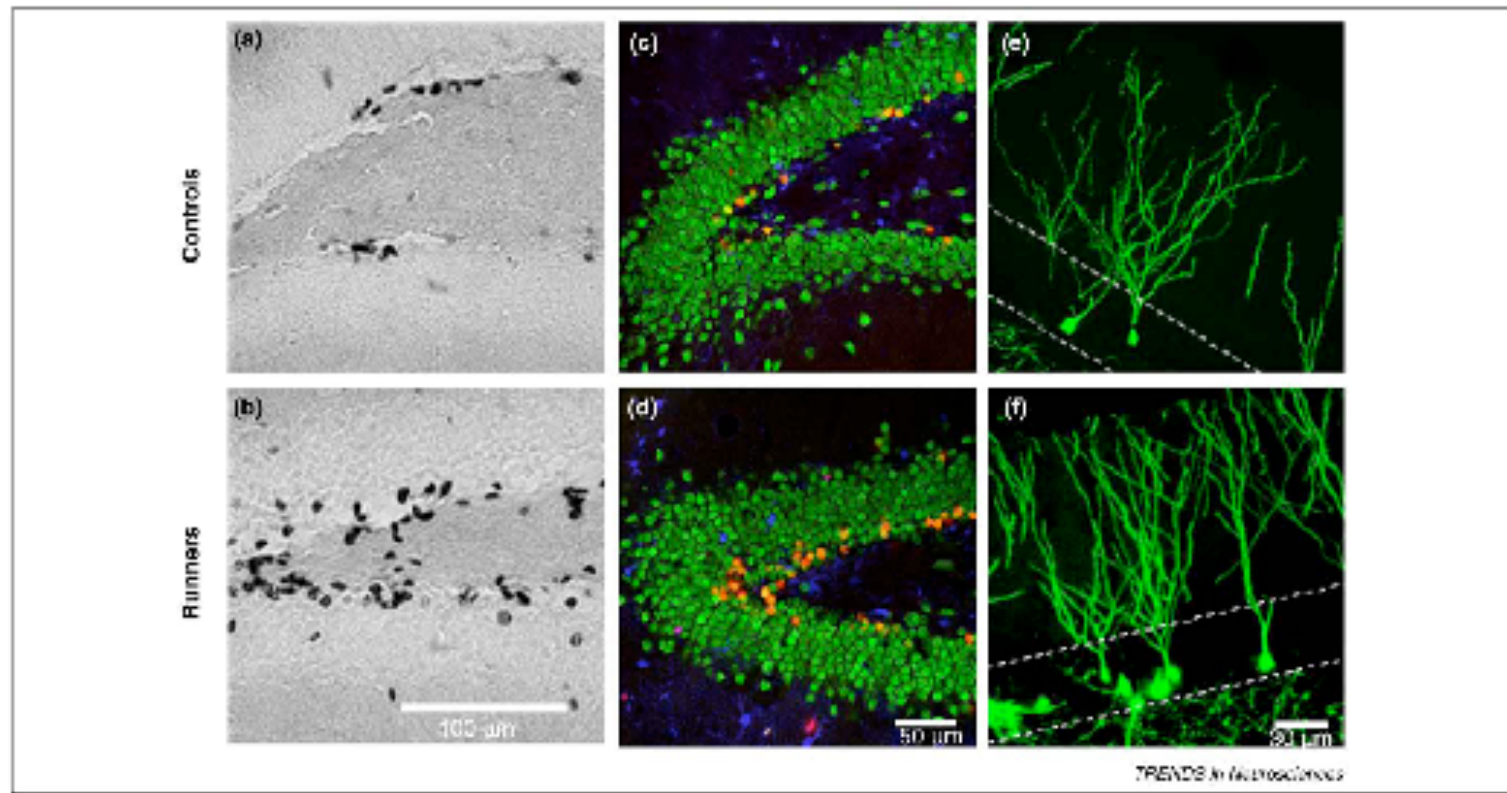


After



*Arch Gen Psychiatry, 2010*

# More Exercise: More Neurogenesis



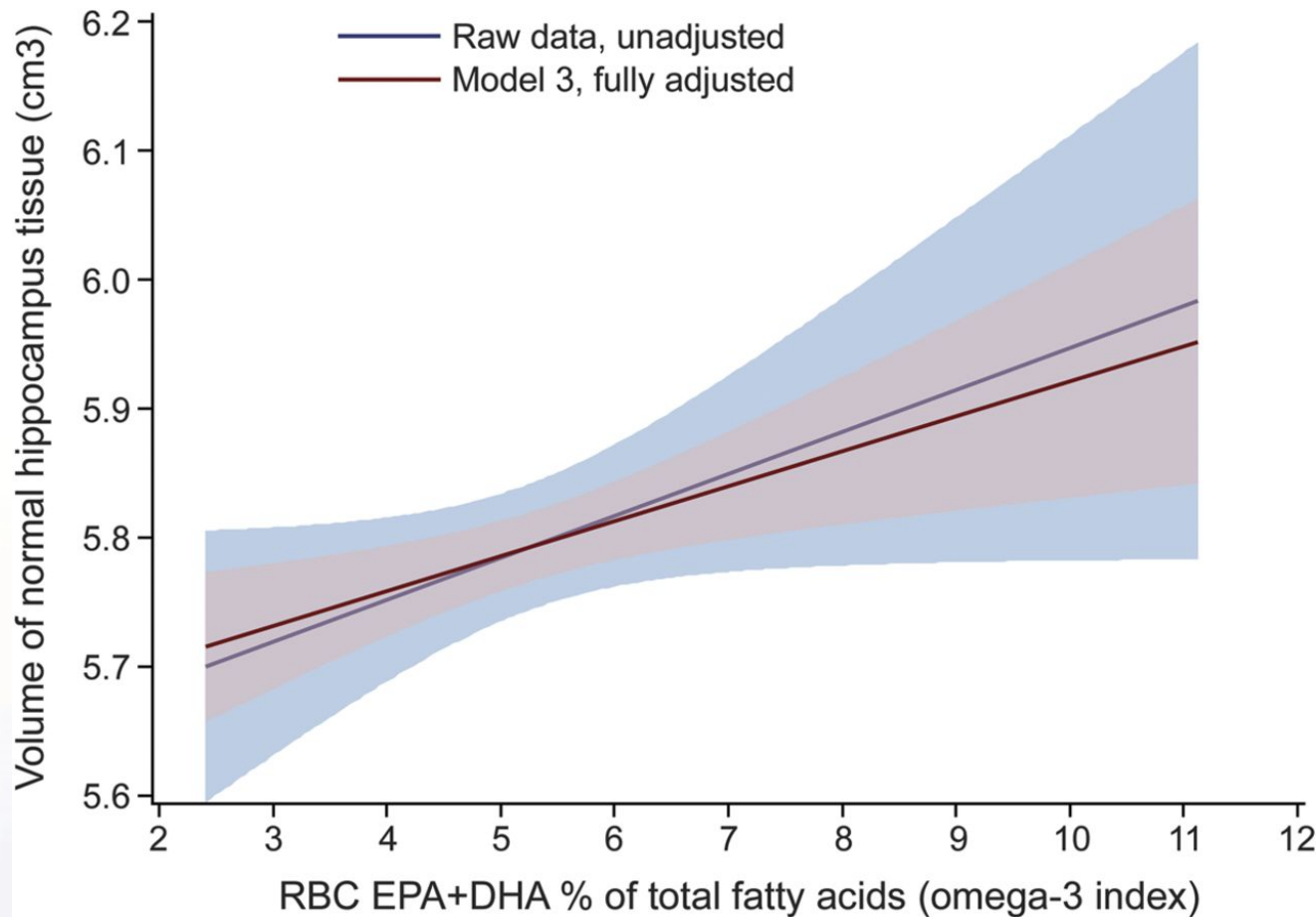
Van Praag, *Trends in Neurosciences*, 2009

## 2. More Omega-3 Fatty Acids, Bigger Hippocampus



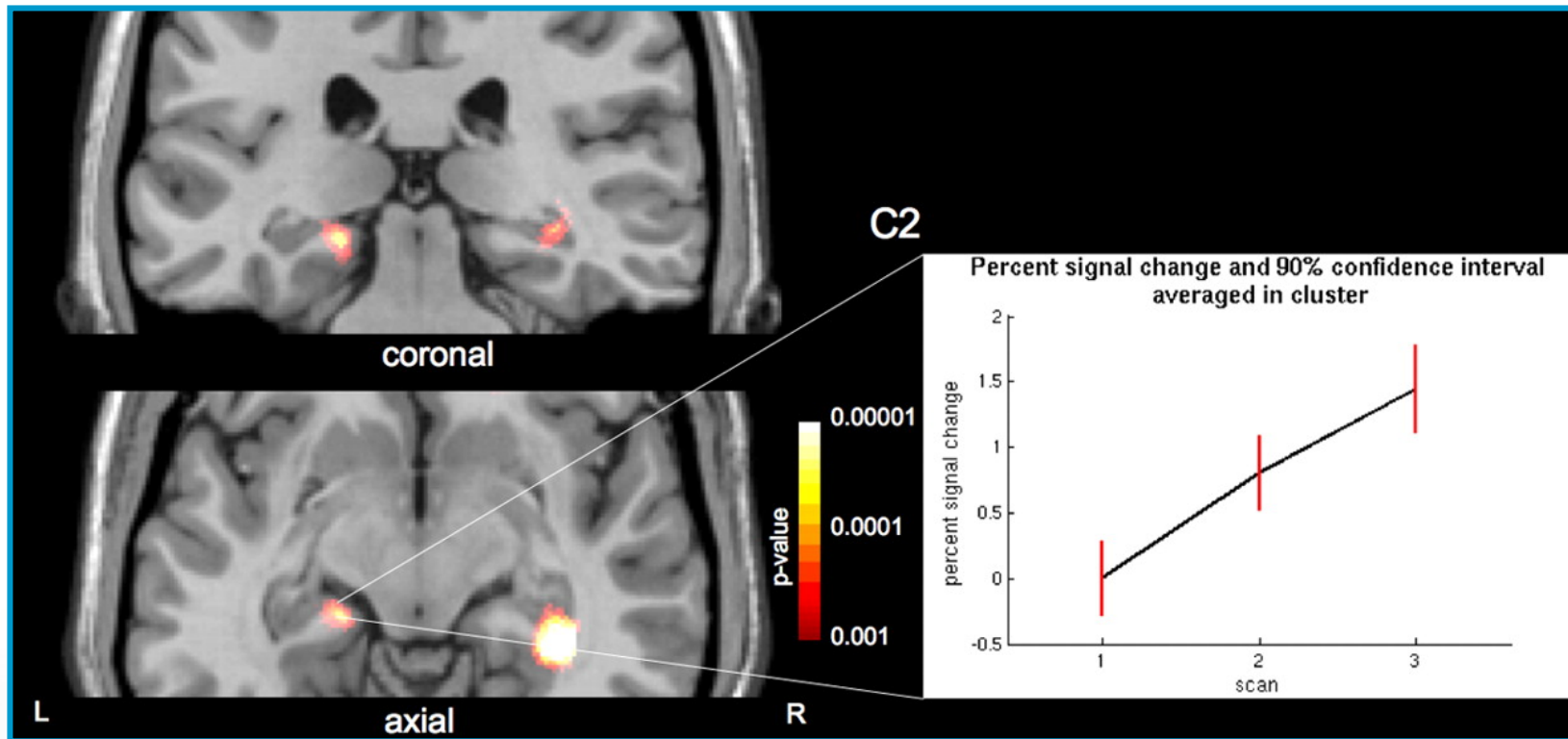


# More Omega-3 Fatty Acids, Bigger Hippocampus



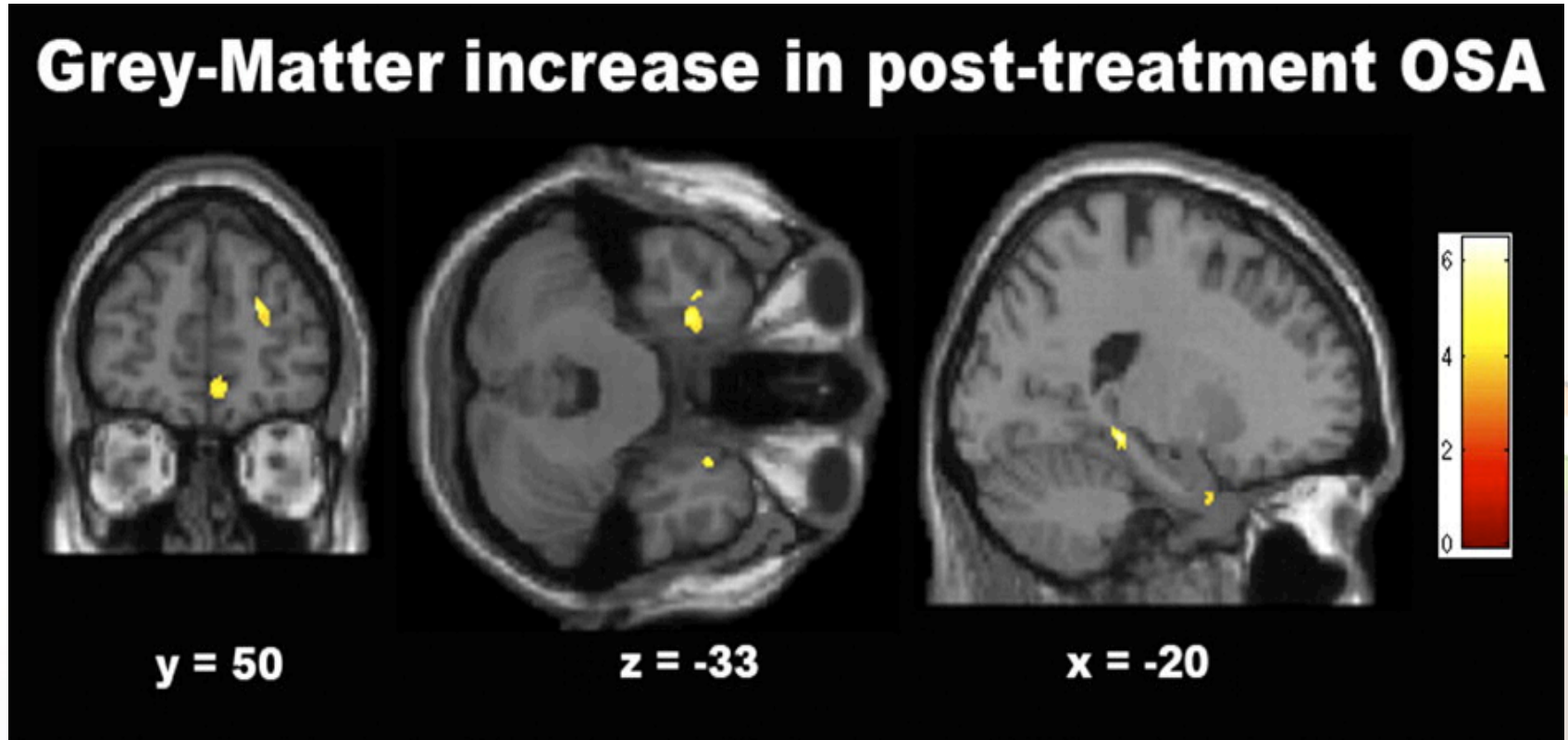
*Pottala J V et al. Neurology 2014;82:435-442*

### 3. More Learning, Bigger Hippocampus



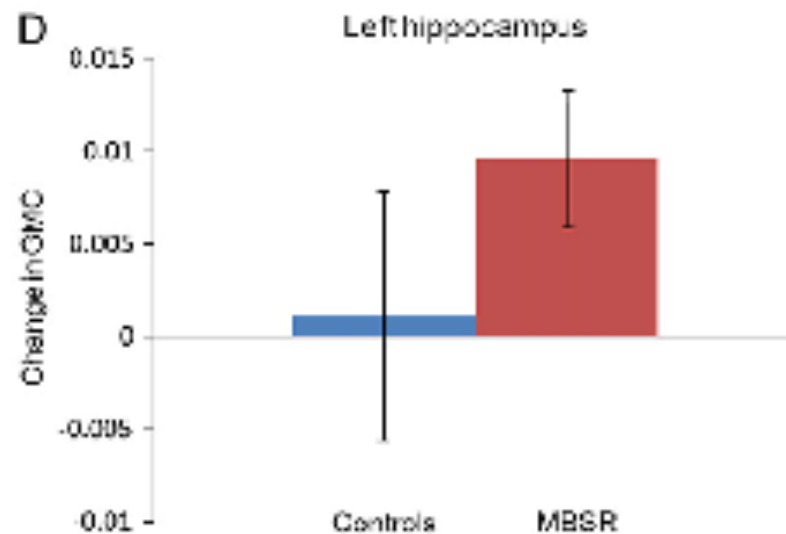
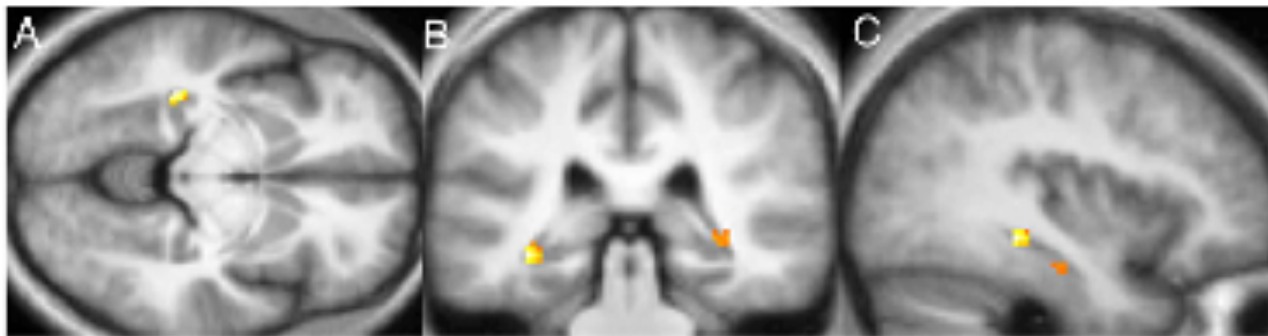
*Draganski et al. J Neurosci 2006; 26:6314–7*

## 4. Better Sleep, Bigger Hippocampus



*Canesa, American Journal of Respiratory Medicine, 2011*

## 5. More Meditation, Bigger Hippocampus

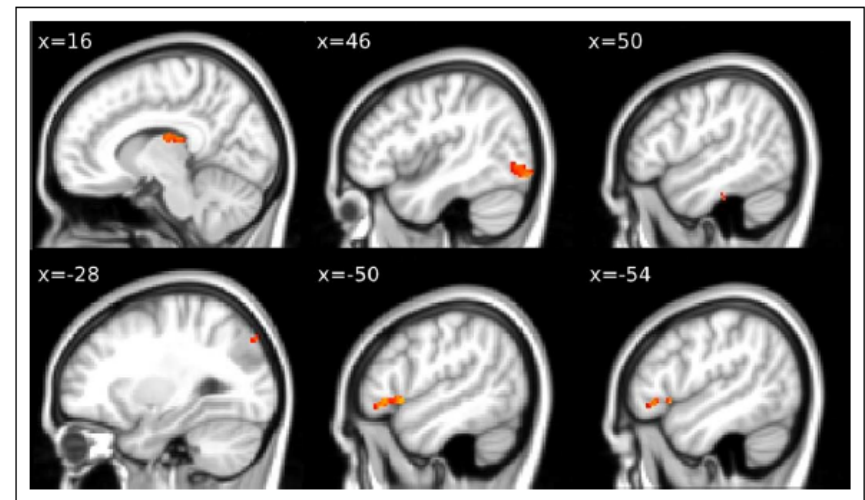
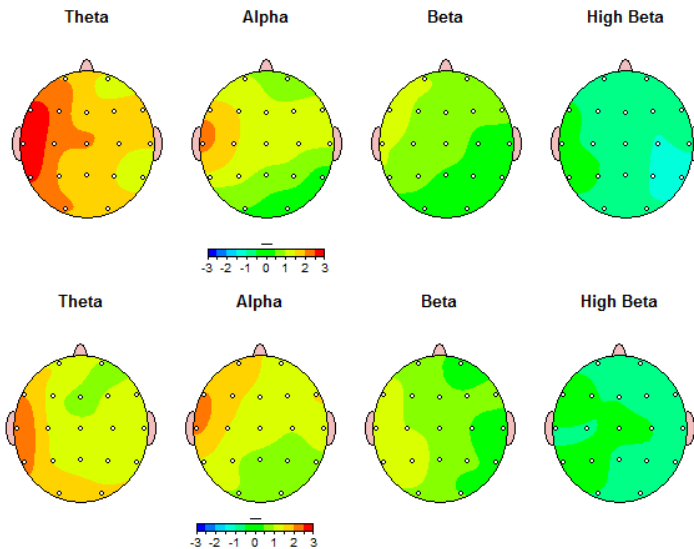


*Holzel , Psychiatric Research, 2011*



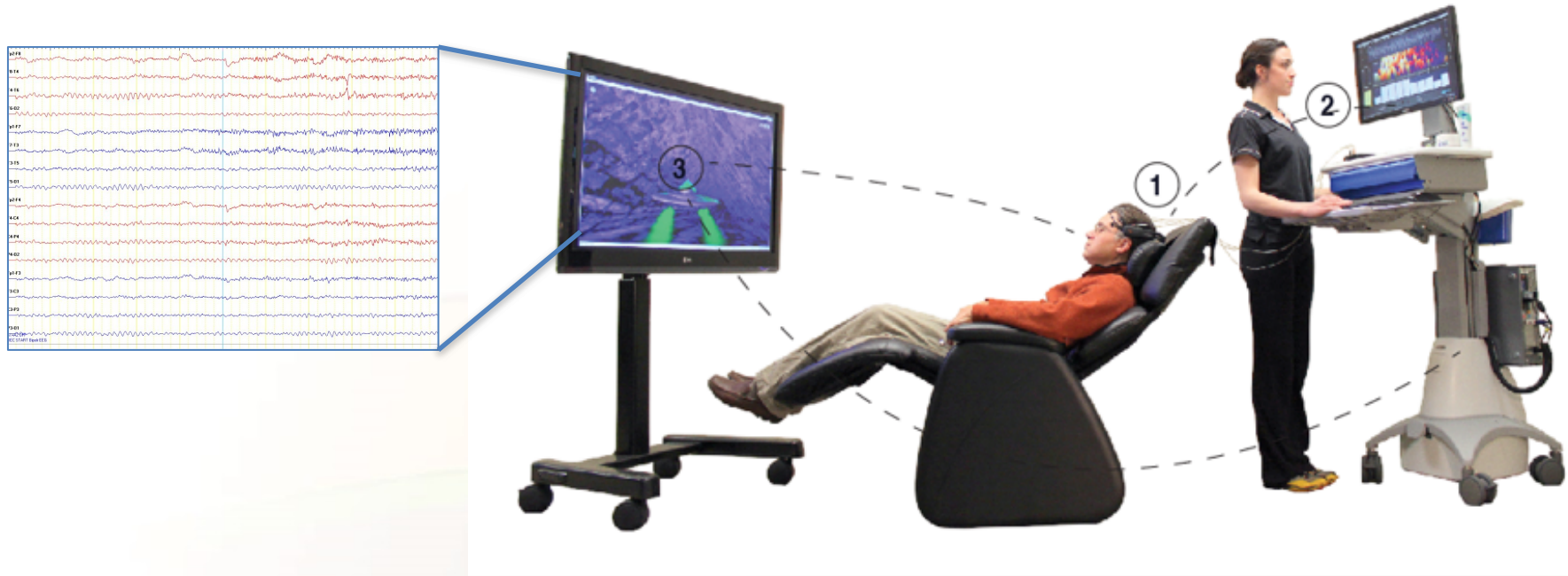
# 6. More Neurofeedback, Bigger Cortex

Z Scored FFT Summary Information



Ghaziri et al. Clin EEG Neurosci 2013; 44 (4) 265-72

# Neurofeedback



- Performed by a certified EEG neurofeedback specialist
- Live EEG feedback is provided through auditory and visual responses to help the patient move brain activity towards an optimal state
- Benefits are long-lasting

# Hippocampus Grows Bigger When You Start Early

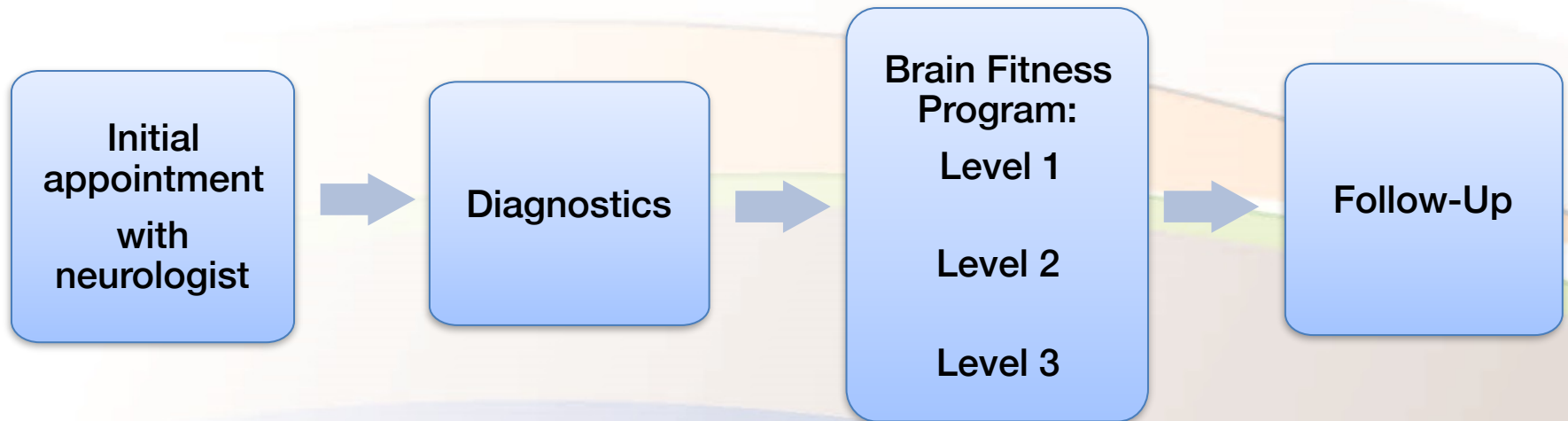


# Objectives

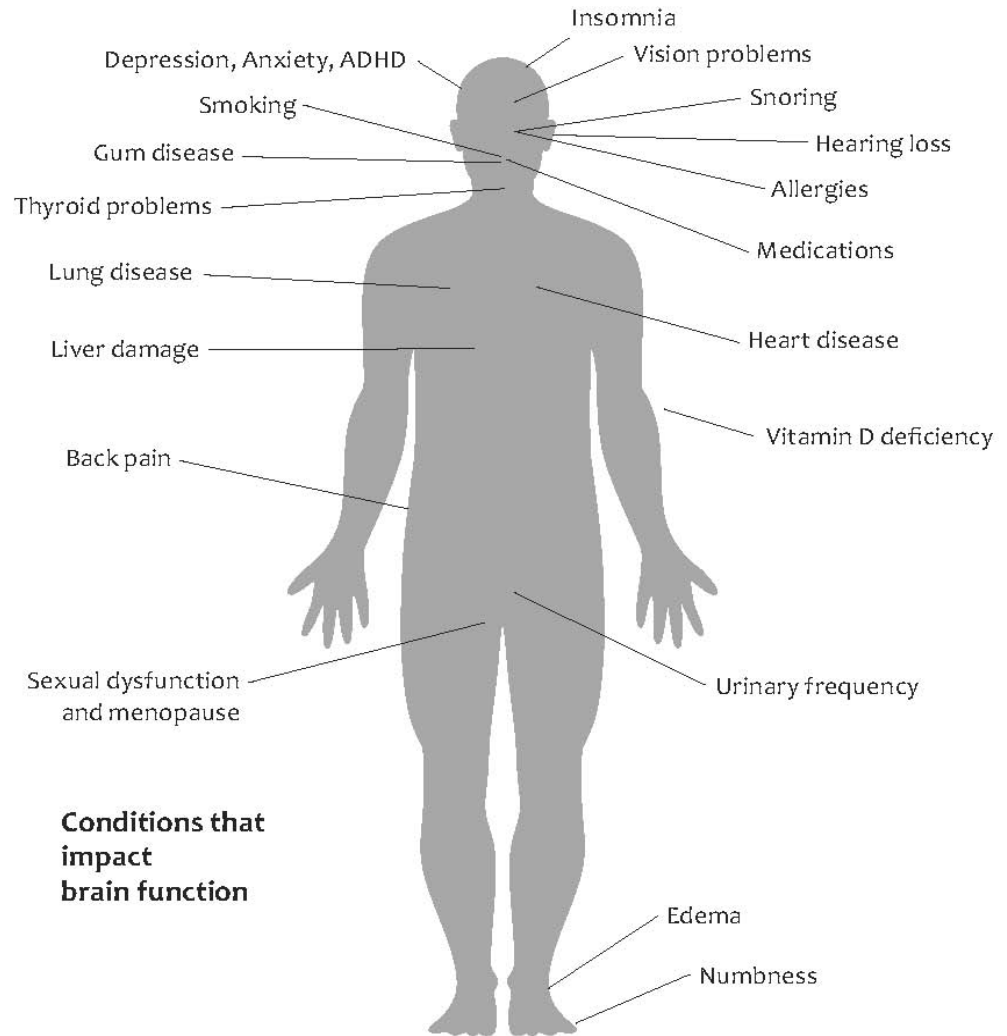
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# Brain Fitness Program: Overview



# Brain Fitness Program: Initial Exam



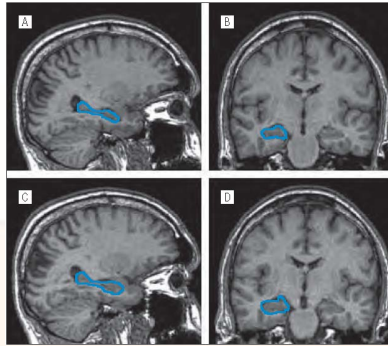
**Conditions that  
impact  
brain function**

# Brain Fitness Program: Comprehensive Diagnostic Tests

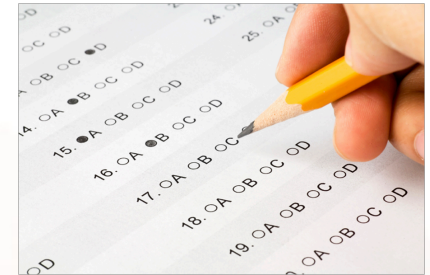
**Cardiopulmonary Testing**



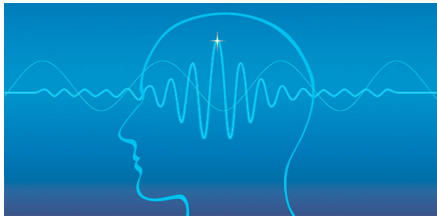
**Brain MRI**



**Neurocognitive Evaluation**



**Brain Mapping qEEG**



**Carotid Ultrasound**



**Blood Test**



**Sleep Health  
Assessment**



# Brain Fitness Program: A Personalized Set of Interventions

**Brain Coaching and Counseling**



**Cognitive Skills Training**



**Neurofeedback Training**



**Weekly Monitoring**

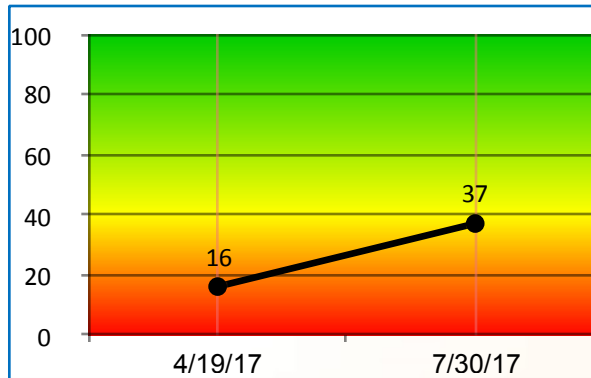




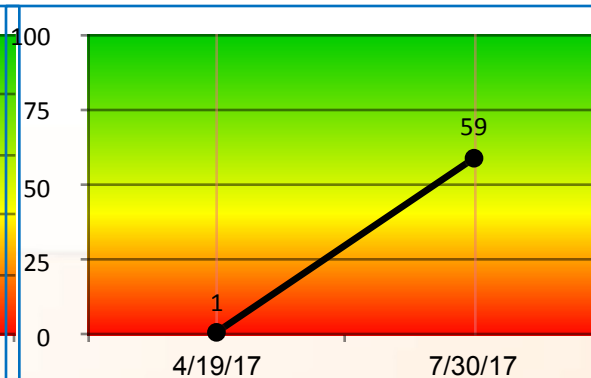
# Brain Fitness Program:

## One example: 69 year old with ?Alzheimer's

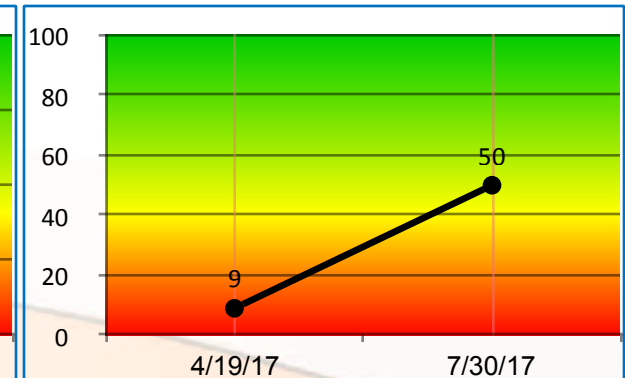
### Memory



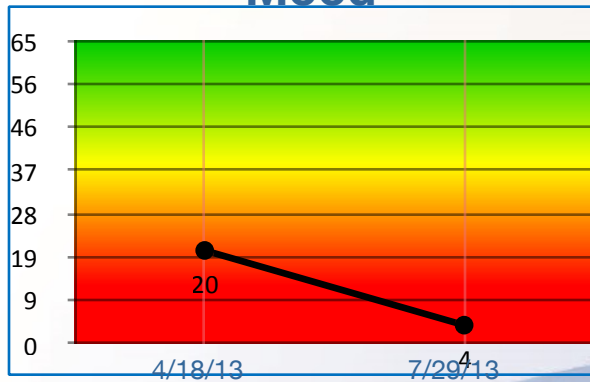
### Attention



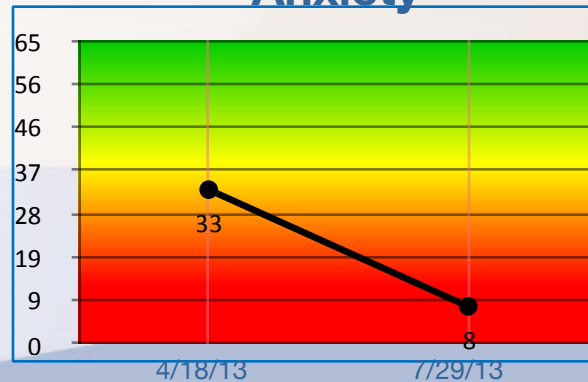
### Concentration



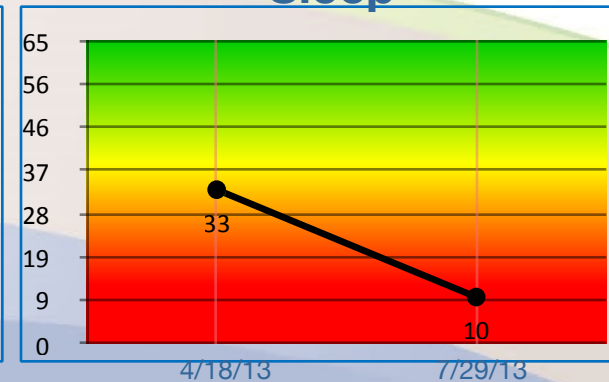
### Mood



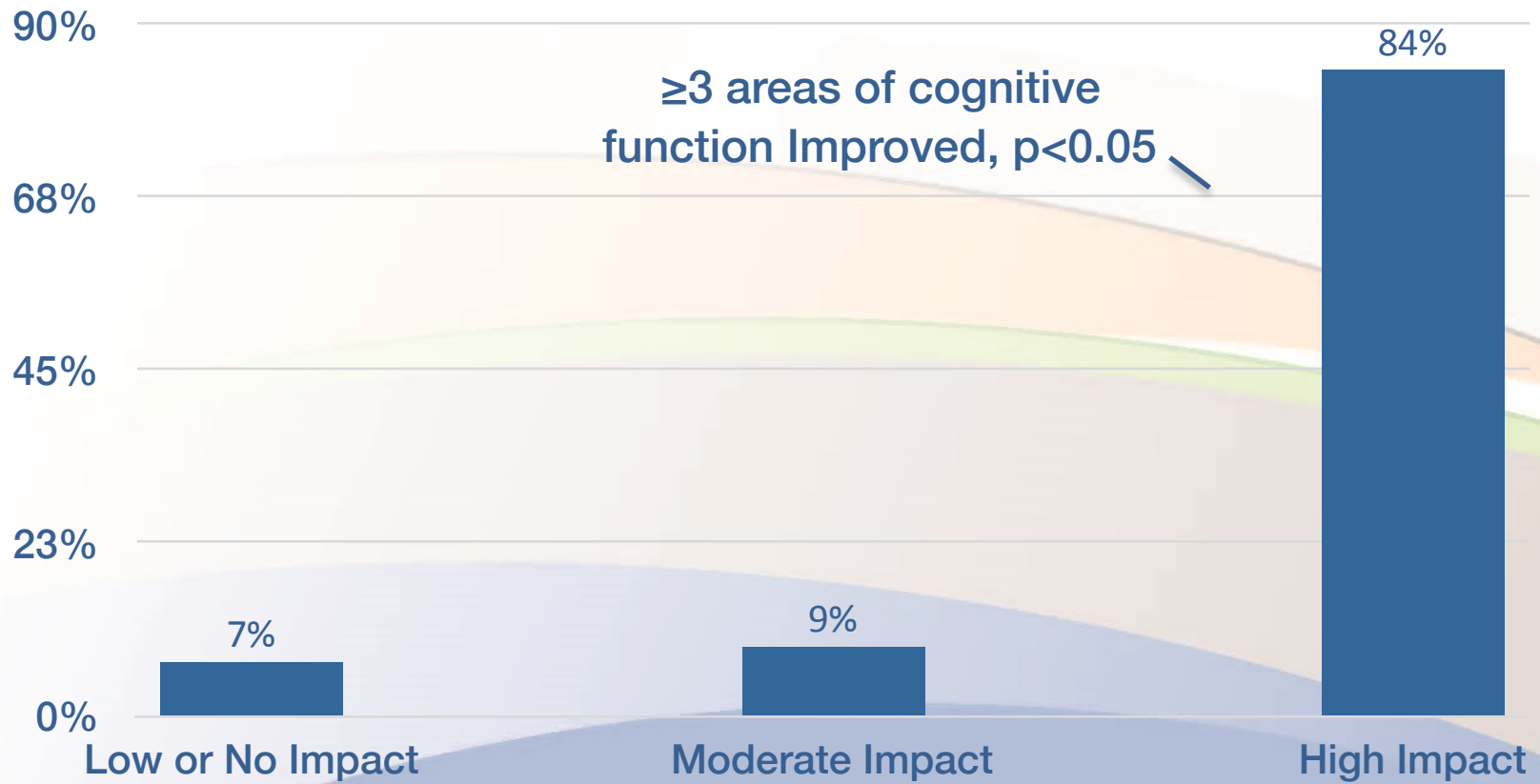
### Anxiety



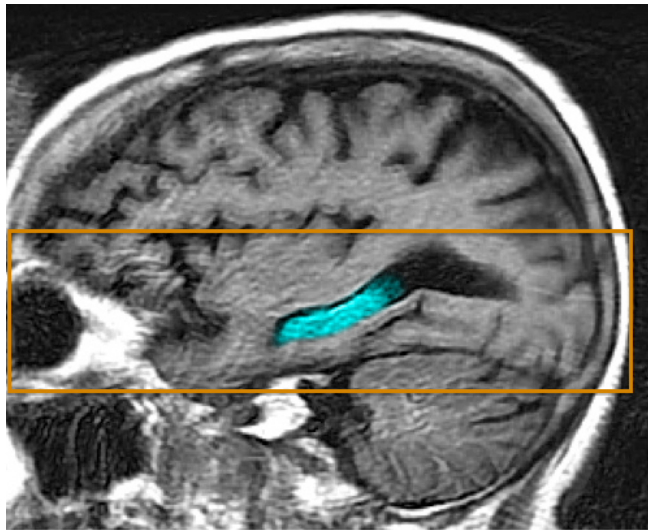
### Sleep



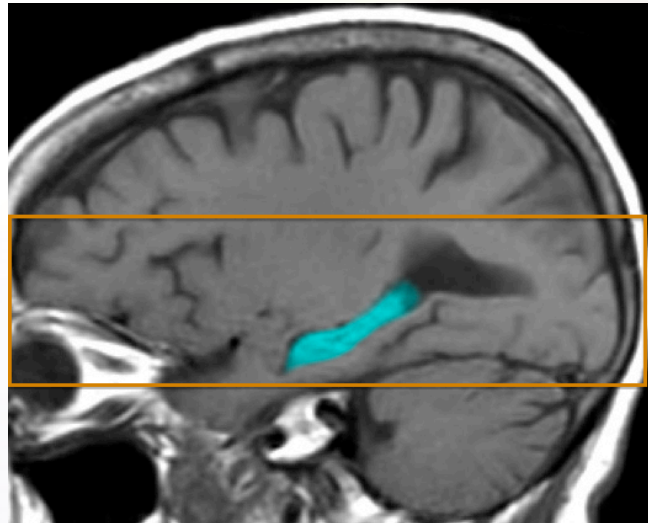
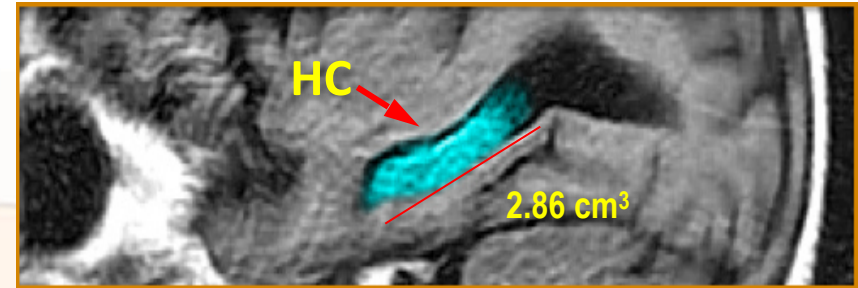
# Brain Fitness Program: Statistical Analysis of 127 Patients with MCI



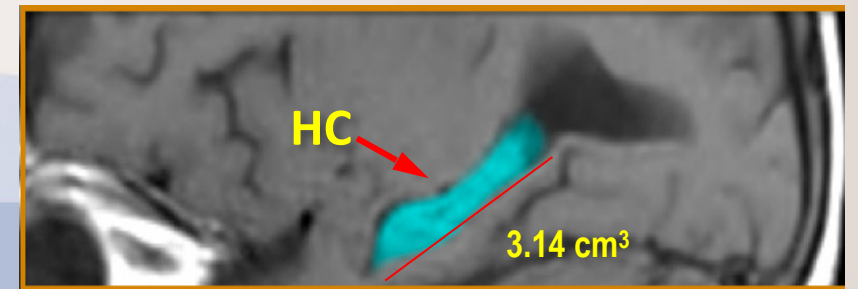
# MRI Results



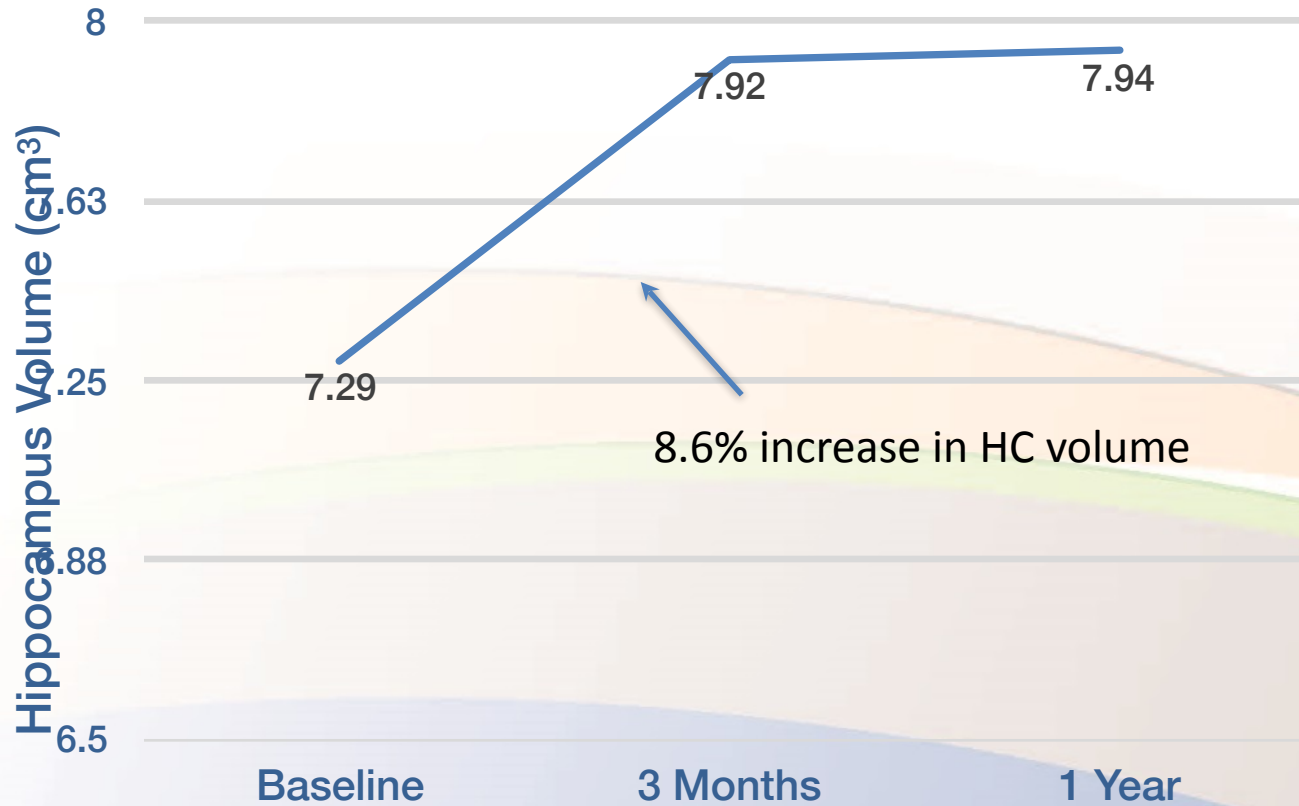
Before



After



# Sustained Benefits



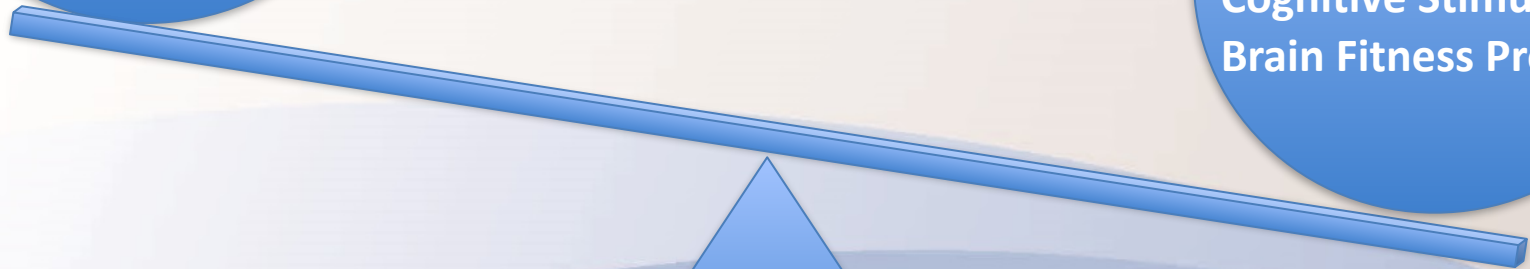
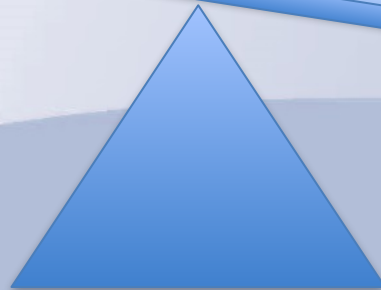




Diabetes  
Hypertension  
Obesity  
Sleep Apnea  
Head Trauma  
Genes  
Stress



Brain-healthy Diet  
Physical Fitness  
Cognitive Stimulation  
Brain Fitness Program



# Summary

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THANK YOU

The background of the slide features a series of overlapping, curved bands of color. From top to bottom, the colors are a very light blue, a warm orange, a pale green, and a darker, muted blue. These bands create a sense of depth and movement, resembling a stylized landscape or a layered effect.