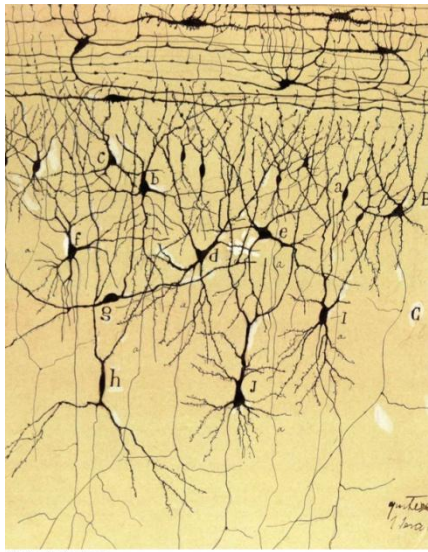




НЕРВНЫЕ КЛЕТКИ ВОССТАНАВЛИВАЮТСЯ ИЛИ КАК УЛУЧШИТЬ ПАМЯТЬ

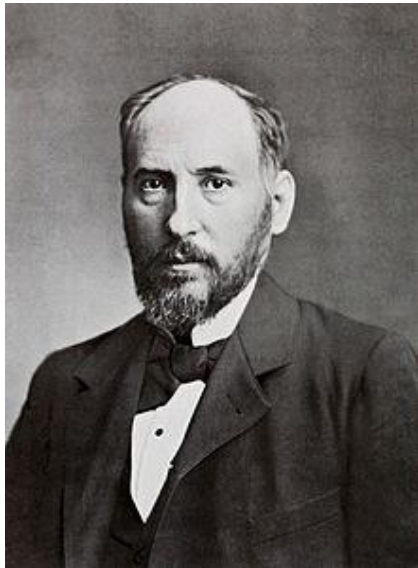
Выполнила: студентка 6 курса лечебного факультета Буган Николетта.



Santiago Ramón y Cajal
Capas 1ª y 2ª de la corteza optativa de la circunvolución del hipocampo del bife, n. 1901
© Herederos de Ramón y Cajal

ИСТОРИЯ ВОПРОСА

"Как только развитие закончено, рост и регенерация аксонов и дендритов прекращаются. Центры взрослого мозга представляют собой нечто установленное, законченное и неизменное. Всё может умереть, ничто не может быть восстановлено», – Рамон-и-Кахаль, 1913



Нобелевская премия по физиологии или медицине 1906 «в знак признания трудов о строении нервной системы».



ПЕРВЫЕ ДОГАДКИ



- **J. Altman.** *Are new neurons formed in the brains of adult mammals?* *Science* 1962; 135, 1127–1128 (первое предположение о нейрогенезе в гиппокампе у мышей, получивших травму).
- **F. Nottebohm.** *From bird song to neurogenesis.* *Scientific American* 1989. 260: 74-9 (с наступлением сезона спаривания у самок канареек в ядрах мозга, связанных с вокализацией и обучением, резко возрастает число нейронов).
- **Eriksson P. S., Perfilieva E., Bjork-Eriksson T., et al.** *Neurogenesis in the adult human hippocampus.* *Nat. Med.* 1998;4: 1313–1317 (посмертный анализ активности гиппокампальных нейронов у людей, при жизни принимавших 5-бромодезоксиуридин).

«Это было реальным шоком, потому что нам всегда преподавали, что взрослый мозг сохраняет тот же самый размер, с теми же самыми клетками, навсегда. Это было неоспоримым фактом знания мозга. Как их могло стать больше? Это противоречило всему, что я когда-либо изучал...» - писал Ноттебом.

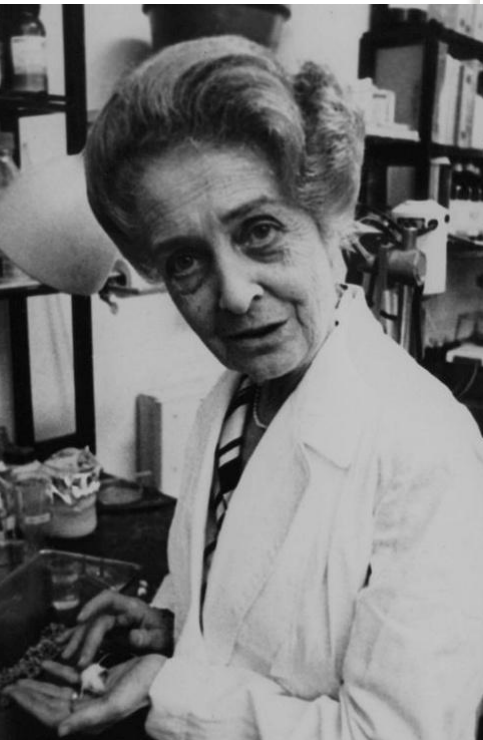




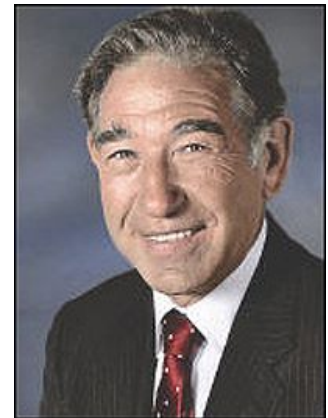
"The effects of wing bud extirpation on the development of the central nervous system in chick embryos", *Journal of Experimental Zoology*, 1934; 68:449-494.

“Для нормального развития нервной системы необходима гибель нервных клеток, только специальным образом запрограммированная”.

Hamburger Viktor and Rita Levi-Montalcini. "Proliferation, Differentiation and Degeneration in the Spinal Ganglia of the Chick Embryo under Normal and Experimental Conditions." *Journal of Experimental Zoology*, 1949: 457–501

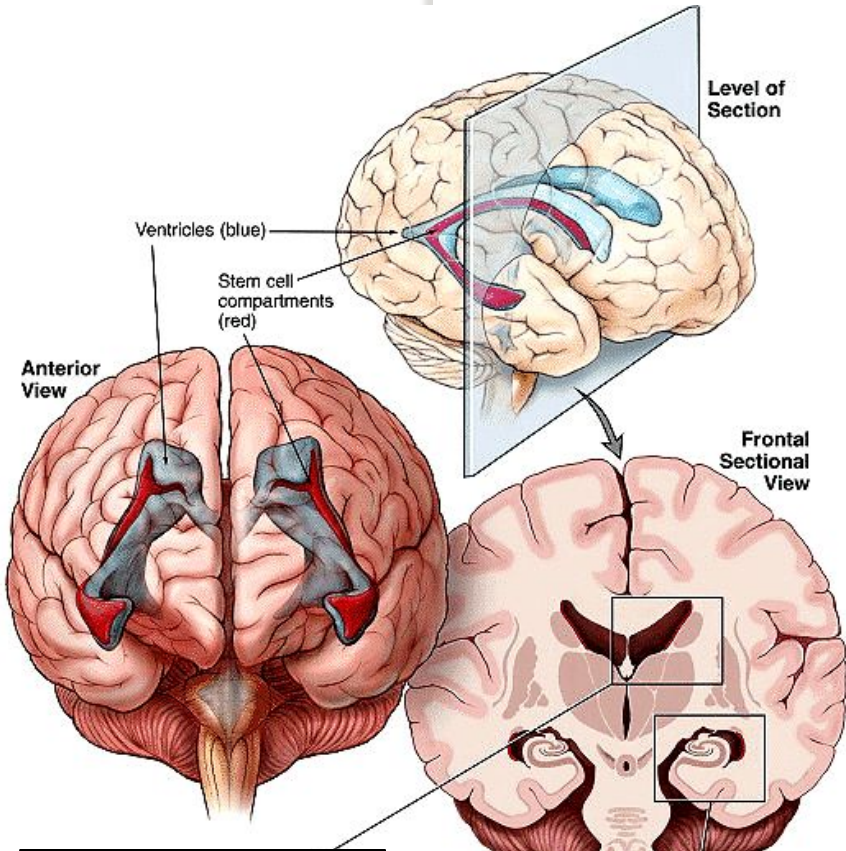


*Опухолевые мышечные ткани, куриная кровь, экстракт эмбрионов и... чувствительные ганглии эмбриона = необходимые ингредиенты для открытия **фактора роста нервов...** и фактора роста эпителия.*



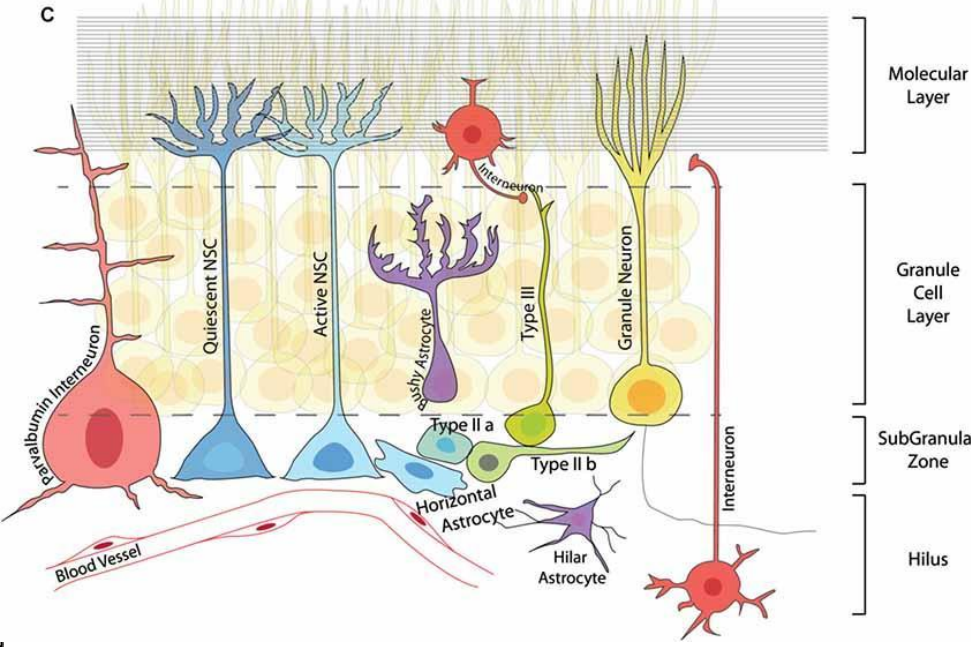
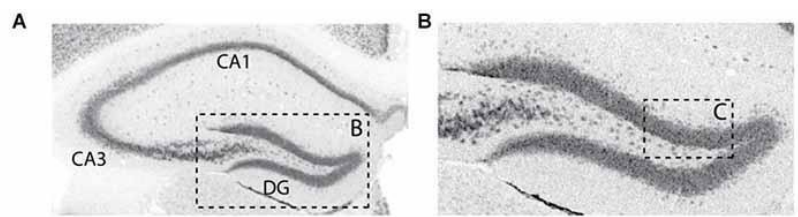
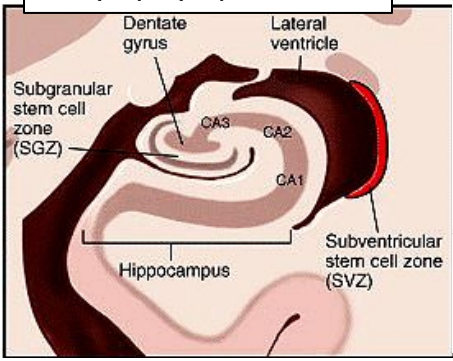
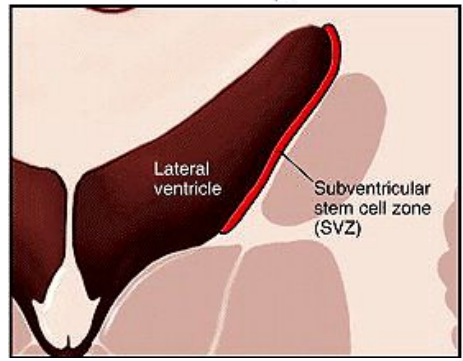
Нобелевская премия по физиологии или медицине 1986 «в знак признания открытий, имеющих важнейшее значение для раскрытия механизмов регуляции роста клеток и органов».

ОСНОВЫ НЕЙРОГЕНЕЗА



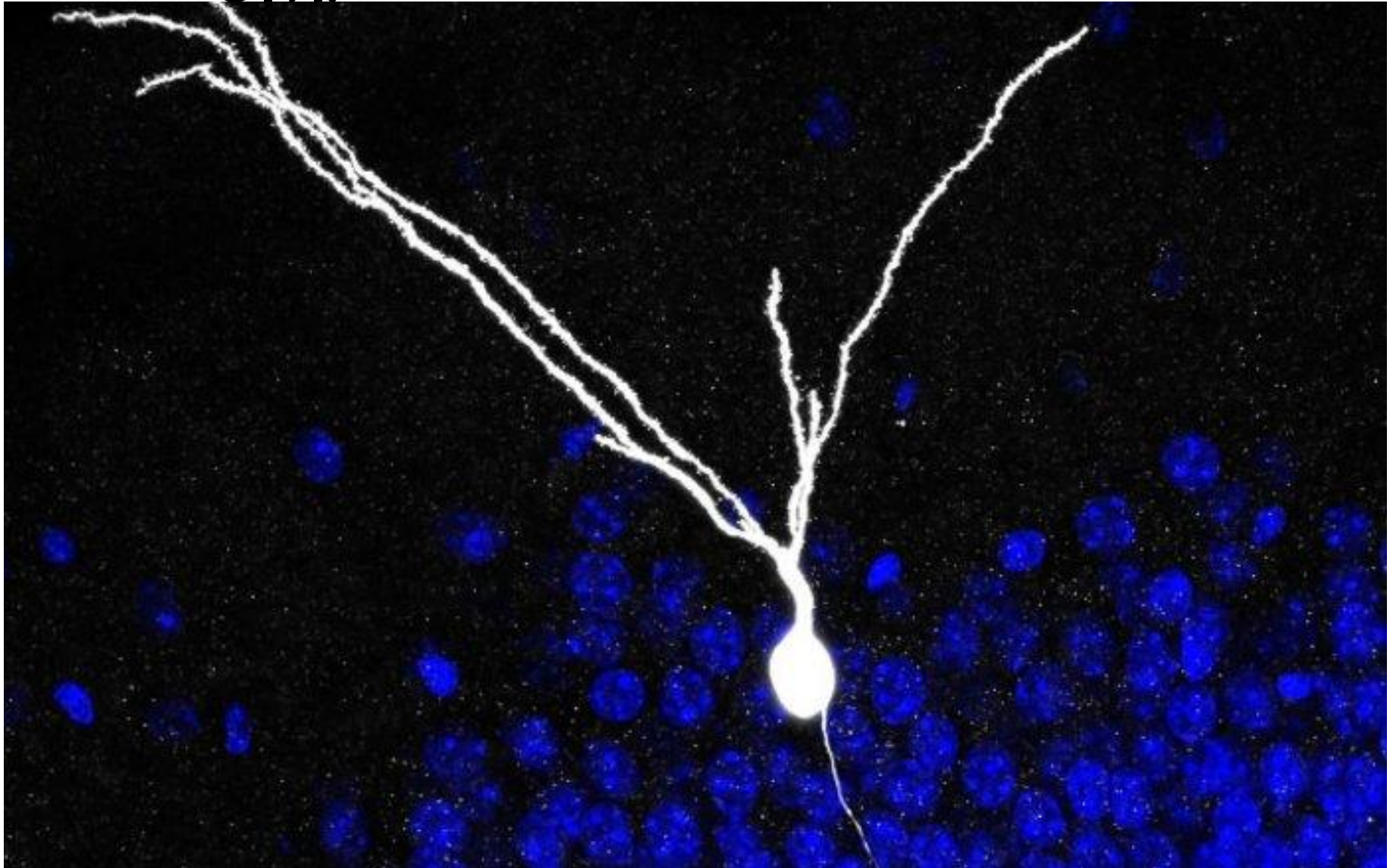
Субвентрикулярная зона

Субгранулярная зона



МЕХАНИЗМЫ НЕЙРОПЛАСТИЧНО СТИ:

- спраутинг (рост отростков нейронов)
- арборизация (ветвление дендритов)
- изменение синаптической проводимости (резервы)
- нейрогенез (стволовые клетки)

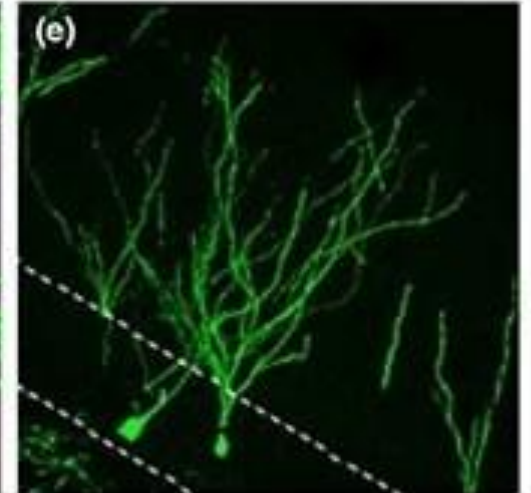
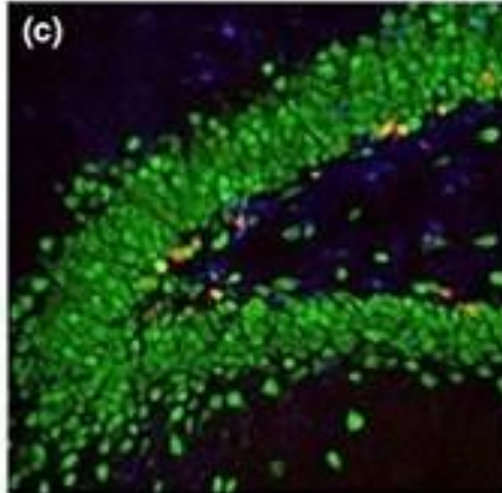
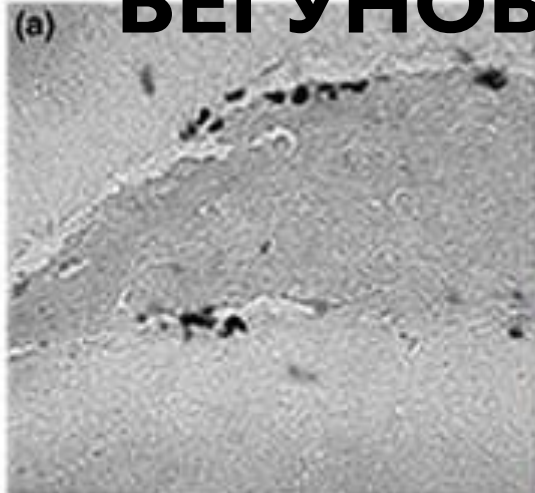




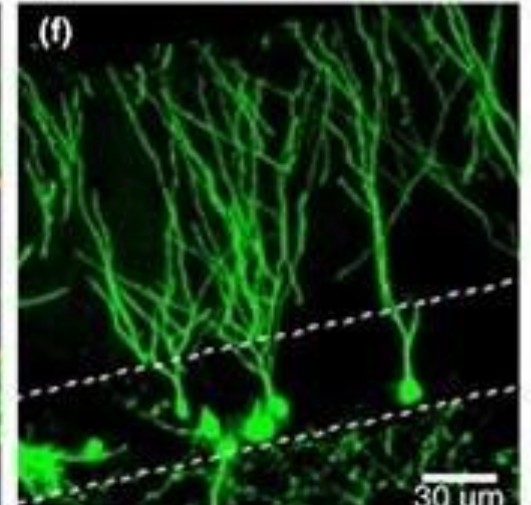
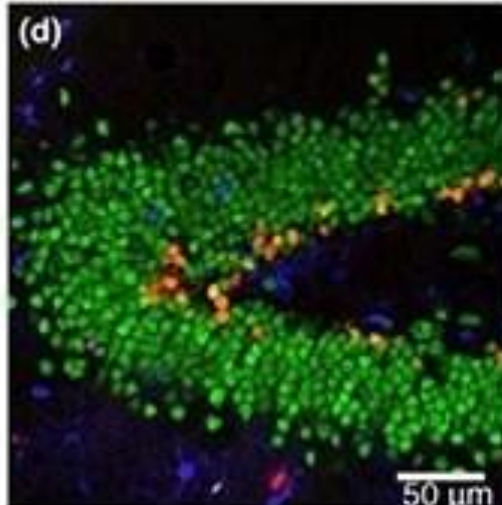
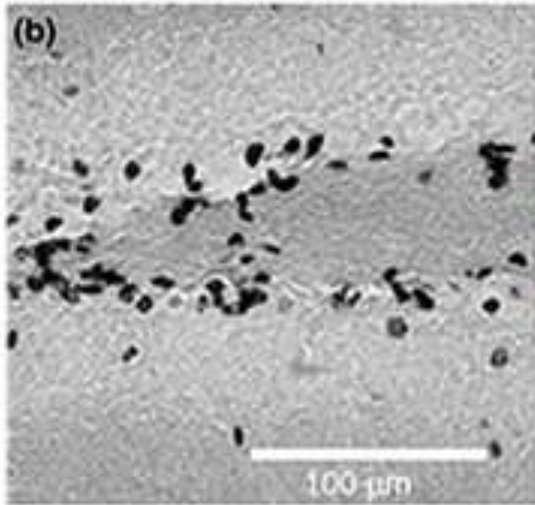
Saive A-L, Royet J-P, & Plailly J. A review on the neural bases of episodic odor memory: from laboratory-based to autobiographical approaches. *Frontiers in Behavioral Neuroscience* 8 (2014). doi: 10.3389/fnbeh.2014.00240

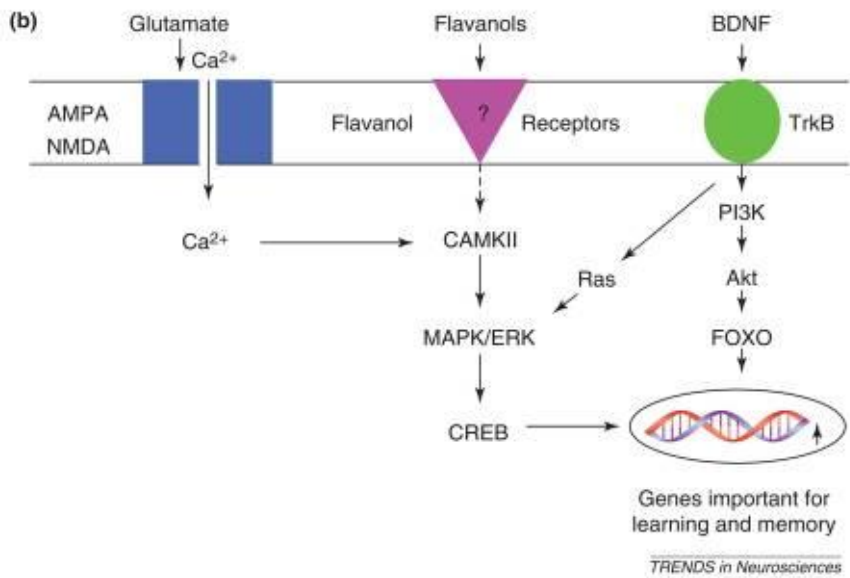
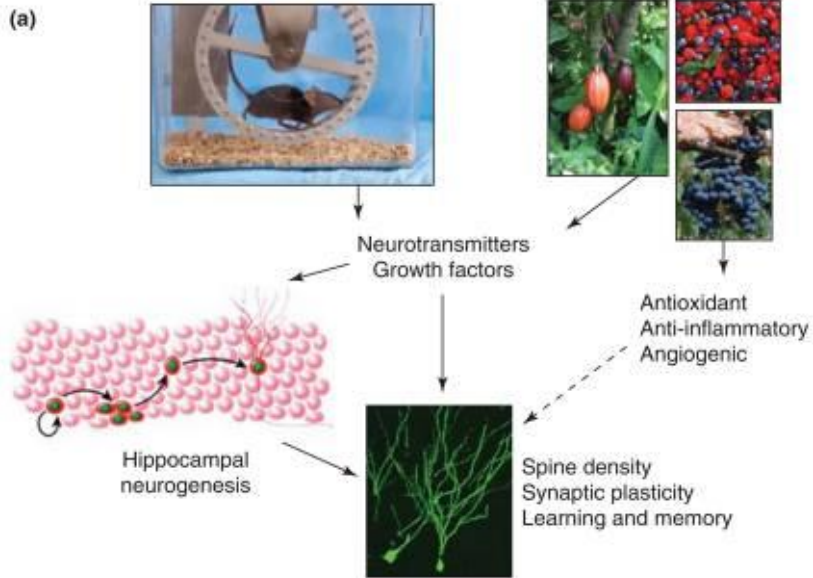
НЕЙРОГЕНЕЗ У КРЫС-БЕГУНОВ

Controls



Runners

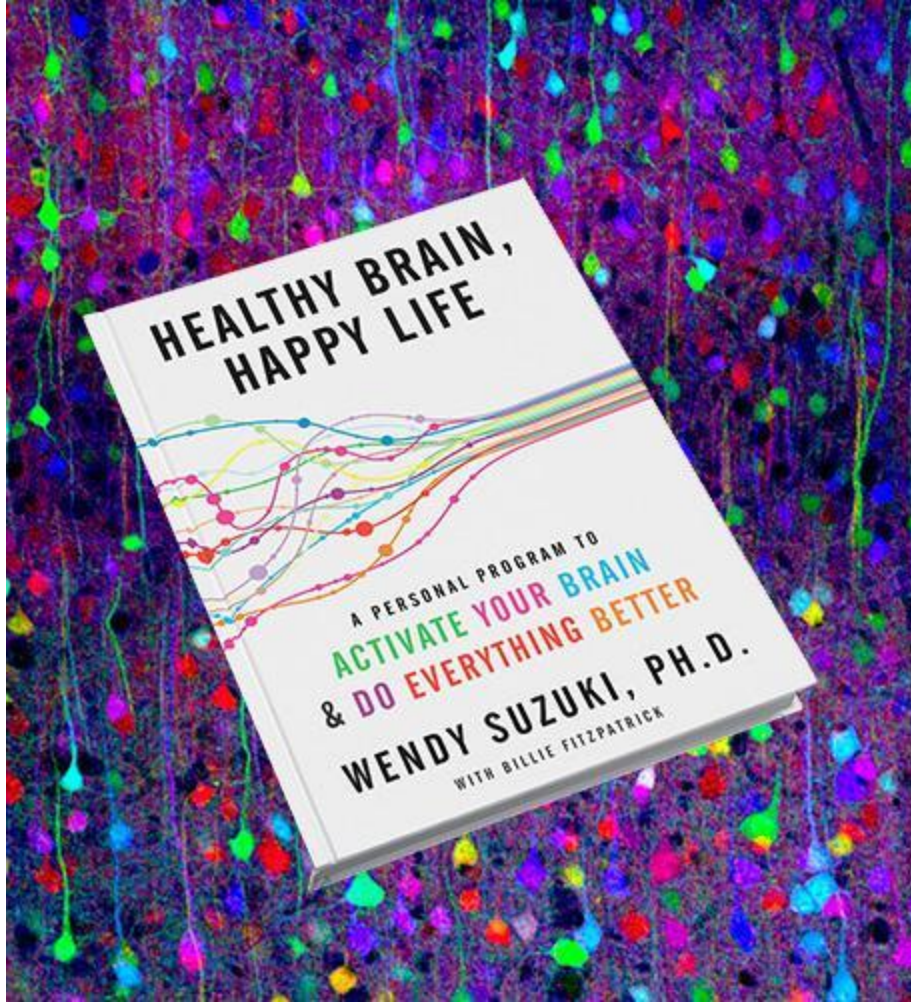




Henriette van Praag. Exercise and the brain: something to chew on. *Trends Neurosci.* 2009 May; 32(5): 283–290.



Molteni R, Barnard RJ, Ying Z, Roberts CK, Gómez-Pinilla F. A high-fat, refined sugar diet reduces hippocampal brain-derived neurotrophic factor, neuronal plasticity, and learning. *Neuroscience*. 2002;112(4):803-14.



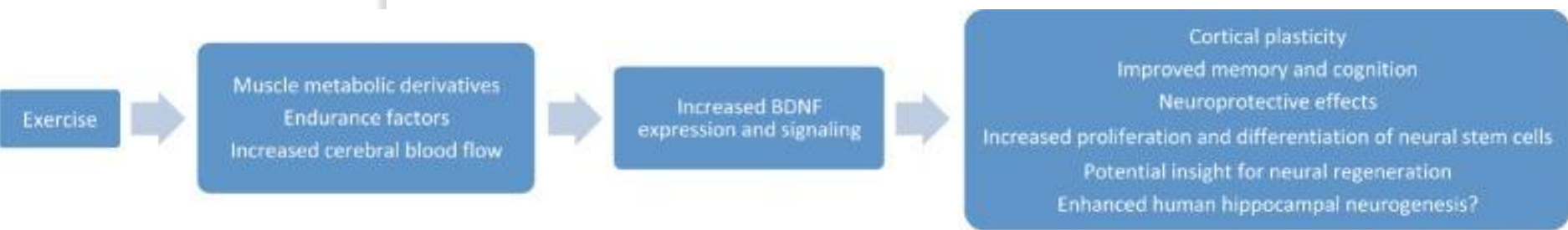
Brain and Behavioral Changes	Exercise Protocol 	Time Course Post-Exercise Cessation								Anatomical Sites
		20 min	40 min	60 min	80 min	100 min	120 min	...	24 hrs +	
A. Behavioral Changes										
<i>Prefrontal-Dependent Cognitive Function</i> ²⁷	50 min cycling at 85% of age-predicted MHR, with additional 5 min warmup and cooldown	[120 min]								
<i>Mood</i>										
Enhanced positive mood states ²⁸	A range of exercise modes at varying intensities (low to very high) for a duration of 7 to over 75 min	[24 hrs]								
Decreased negative mood states ²⁹	30 min of aerobic exercise class at age-predicted 60-80% of MHR	[24 hrs]								
<i>Stress Reduction</i> ³⁰	60 or 120 min of cycling at 50-55% of VO ₂ max, with additional 3 min warmup	[3 hrs]								
B. Neurophysiological Changes										
<i>Extracellular Recording</i>										
Increased theta ⁷⁰	5 min of treadmill running at speeds ranging from 1 to 22 cm/sec	[6 min]								Hippocampus
<i>EEG</i>										
Increased delta, theta, alpha and beta ⁶⁶	A range of exercise modes at various intensities for a duration of 2 to 60 min	[30 min]								Frontal, Temporal, Parietal and Occipital Cortices
Increased P300 amplitude ⁸⁸	20 min of treadmill walking at 60% of age-predicted MHR	[25 min]								
Decreased P300 latency ⁸⁸	20 min of treadmill walking at 60% of age-predicted MHR	[25 min]								
<i>fNIRS</i>										
Increased activation during task performance ¹⁰⁰	10 min of cycling at 50% of VO ₂ max	[15 min]								Prefrontal Cortex
Increased HAROLD during task performance ¹⁰⁵	10 min of cycling at 50% of VO ₂ max	[15 min]								
<i>TMS</i>										
Excitation ¹¹⁶	20 min of cycling at 65-70% of age-predicted MHR, with additional 5 min warmup	[30 min]								Primary Motor Cortex
Inhibition ¹¹⁶	20 min of cycling at 65-70% of age-predicted MHR, with additional 5 min warmup	[30 min]								
<i>fMRI</i>										
Increased hippocampal pattern similarity ¹¹³	25 min of high-intensity Interval cycling at 80% of age-predicted MHR, with additional 5 min warmup and cooldown	[48 hrs]								Hippocampus, Frontal and Temporal Cortices
Altered activation to an acute stressor ¹¹⁴	30 min of treadmill walking/running at 60-70% of VO ₂ max	[90 min]								
C. Neurochemical Changes										
<i>Lactate</i> ¹²⁹	Graded exercise test on a bicycle ergometer to 85% of age-predicted MHR	[40 min]								Cortex
<i>Cortisol</i> ¹⁴¹		[120 min]								Hippocampus, Hypothalamus, Pituitary gland, Frontal Cortex
<i>Neurotrophins</i>										
BDNF ¹⁷⁹	30 min treadmill running at low- (15 m/min) or moderate-intensity (25 m/min)	[90 min]								Hippocampus, Prefrontal Cortex, Striatum
IGF-1		?								
VEGF		?								
<i>Neurotransmitters</i> (see 197 for review)										
Dopamine ²⁰¹	20 min treadmill running at 12 m/min	[120 min]								Hippocampus, Prefrontal Cortex, Striatum, Midbrain, Brainstem, Pons, medulla, Hypothalamus, Cortex, Preoptic area
Norepinephrine ²²²	120 min treadmill running at 25 m/min at a 3% slope	[70 min]								
Serotonin ²¹⁴	120 min treadmill running at 25 m/min	[120 min]								
Acetylcholine										
GABA ²⁵⁵	Graded exercise test on a bicycle ergometer to ≥80% age-predicted MHR,	[20 min]								
Glutamate ²⁰¹	20 min treadmill running at 12 m/min	[40 min]								
<i>Endogenous Opioids</i> ²⁴⁰	2 hrs of endurance running (21.5±4.7 km)	[30 min]								Frontal, Temporal, and Parietal Cortices, Cerebellum, Basal Ganglia
<i>Endocannabinoids</i>		?								

Basso JC, Suzuki WA. The Effects of Acute Exercise on Mood, Cognition, Neurophysiology, and Neurochemical Pathways: A Review. *Brain Plast.* 2017;2(2):127-152. Published 2017 Mar 28. doi:10.3233/BPL-160040

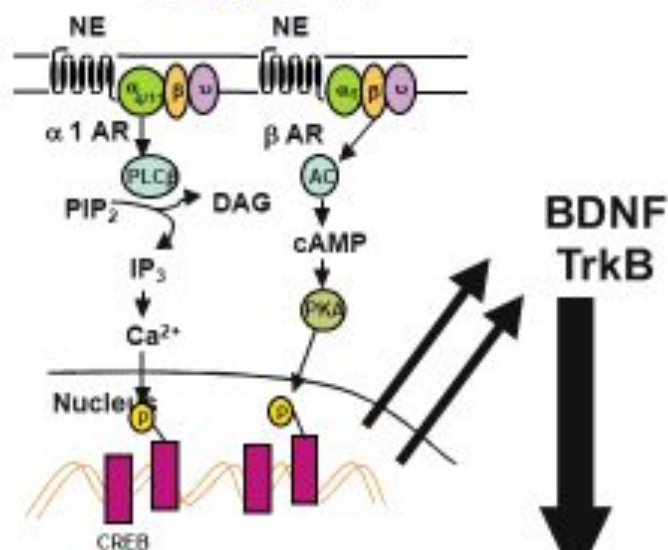
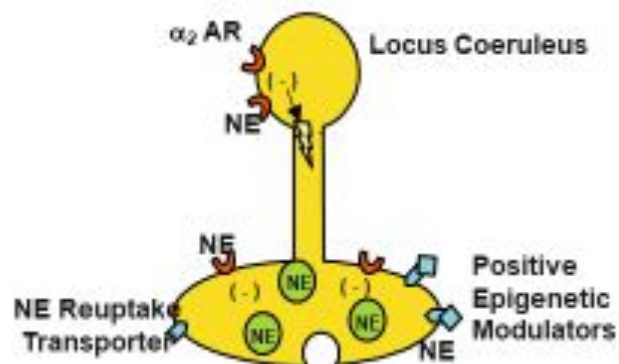


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Sleiman SF, et al. Exercise promotes the expression of brain derived neurotrophic factor (BDNF) through the action of the ketone body beta-hydroxybutyrate. *Elife.* 20165.



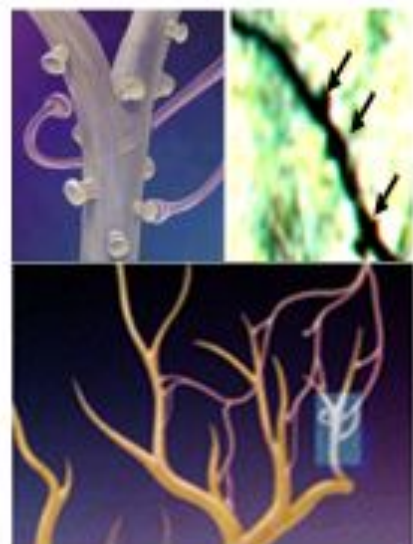
Genomics 101: BDNF and Brain Function



BDNF promotes the growth and survival of brain cells:

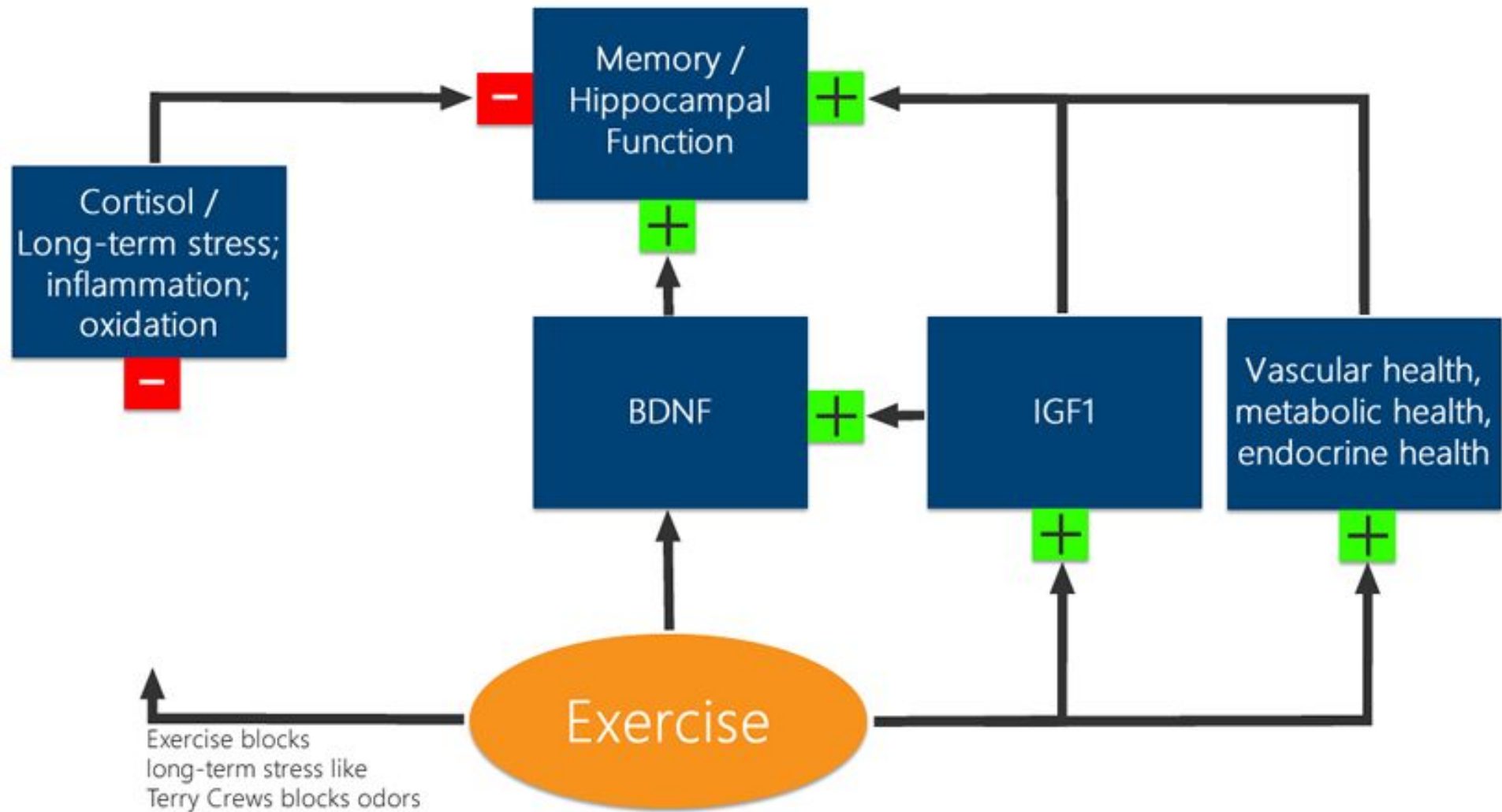
- Restores/enhances neural connectivity
- Promotes healthy cognitive functioning.
- Produces critical neurotransmitters
- Improves executive functioning and memory

Low
BDNF

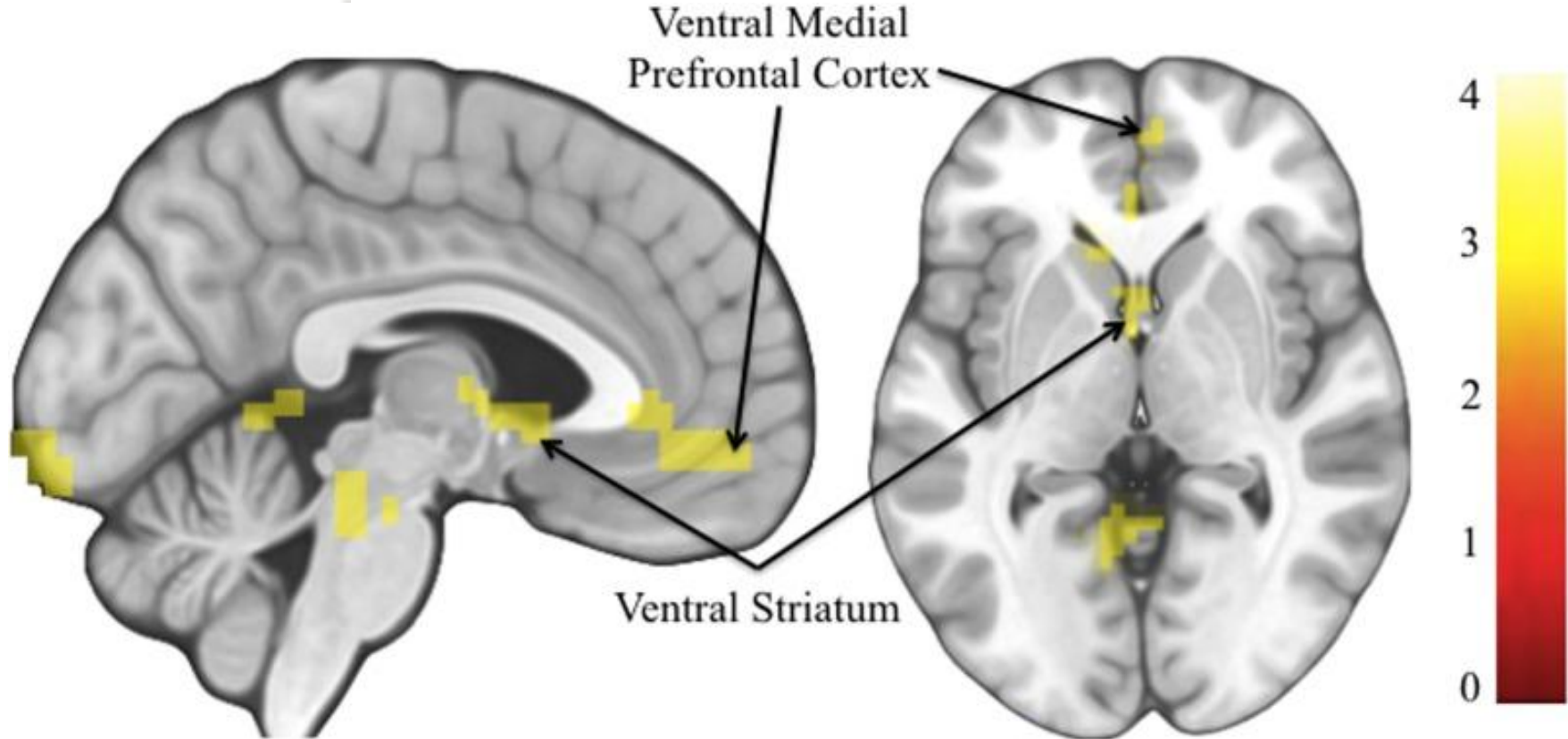


High
BDNF





Basso JC, Suzuki WA. The Effects of Acute Exercise on Mood, Cognition, Neurophysiology, and Neurochemical Pathways: A Review. *Brain Plast.* 2017;2(2):127-152. Published 2017 Mar 28. doi:10.3233/BPL-160040



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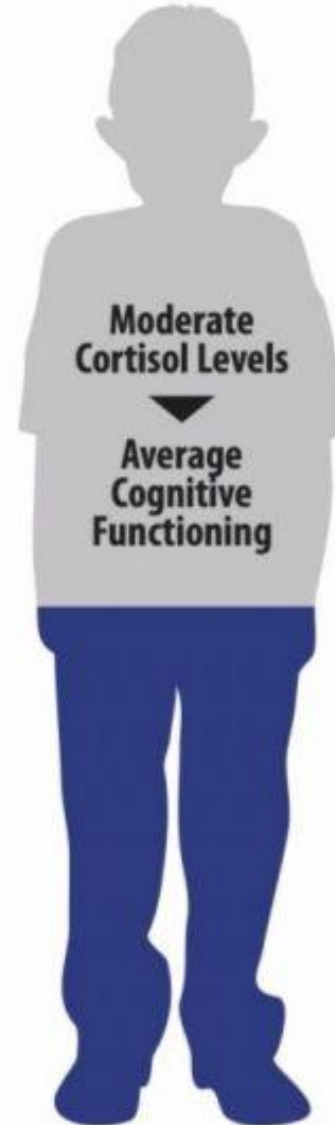
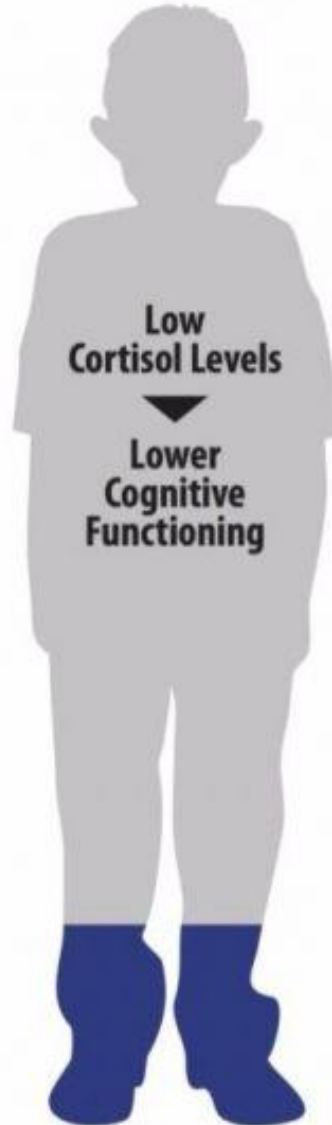
At Risk Environment:

Family Instability and
Parental Emotional
Unavailability



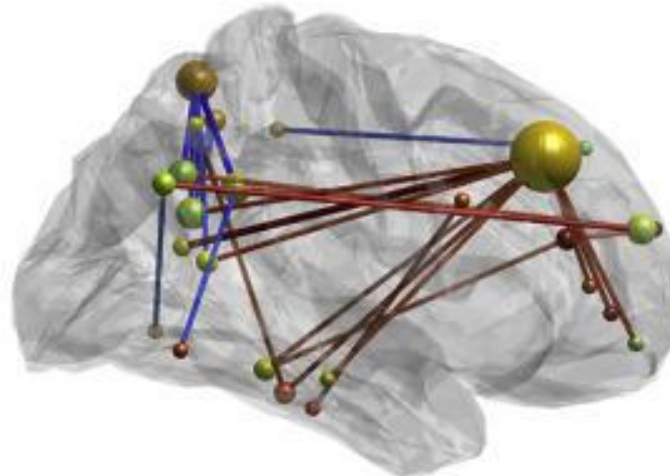
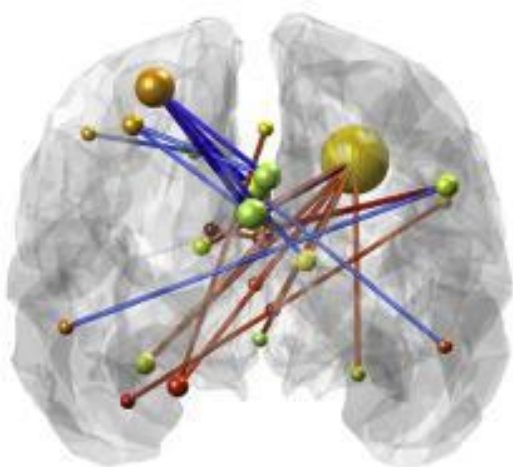
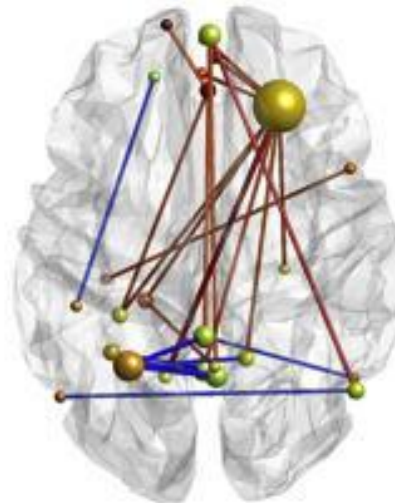
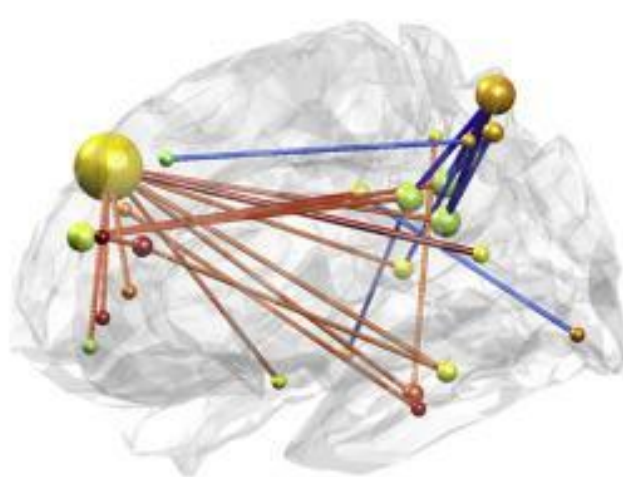
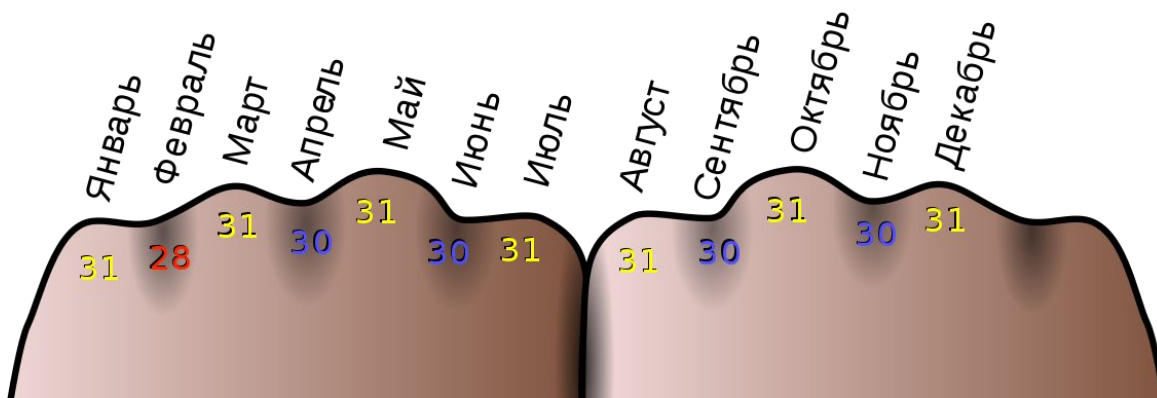
At Risk Environment:

Parental
Emotional
Unavailability

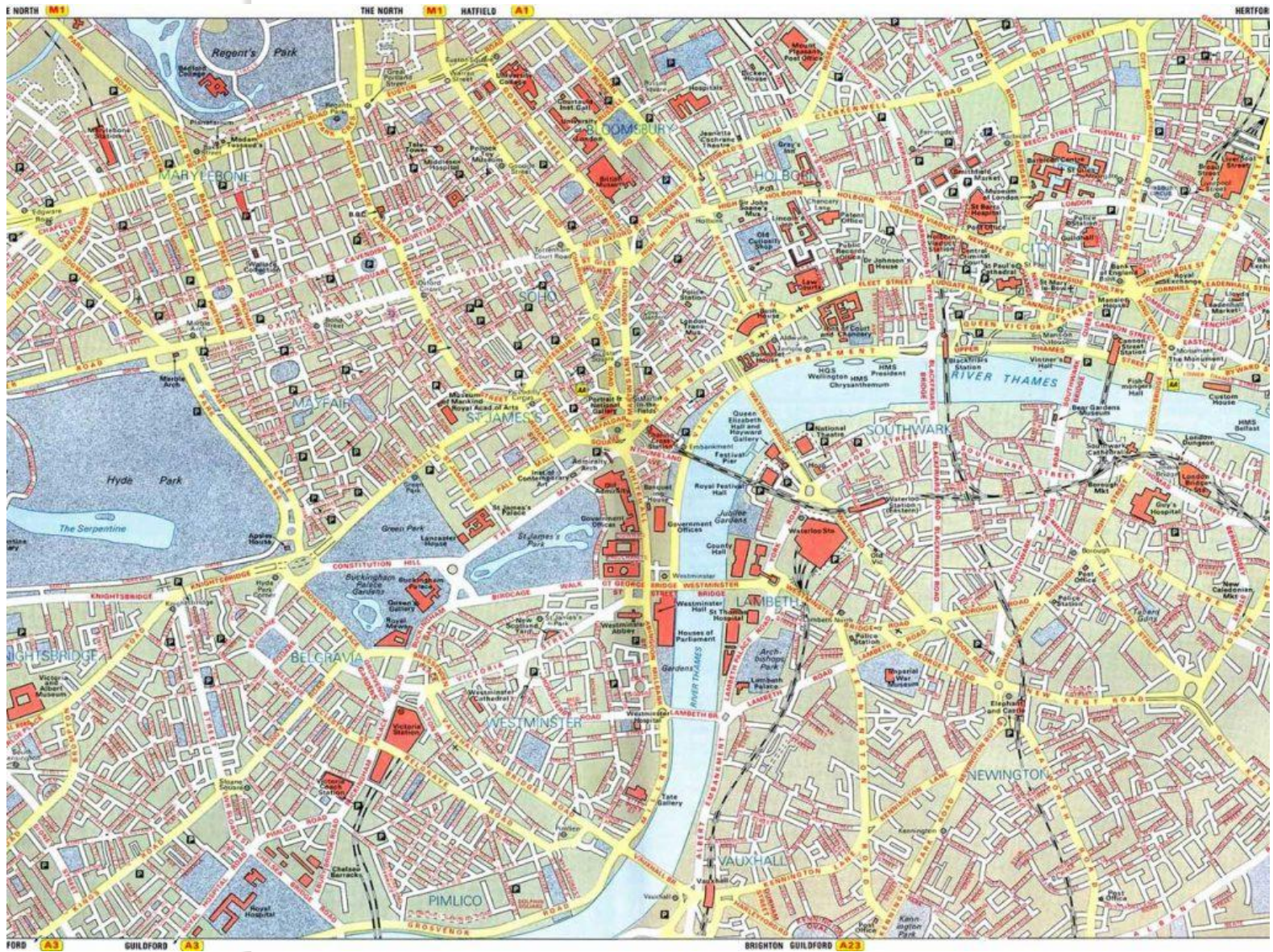


ЛОКУСНАЯ ТРЕНИРОВКА ПАМЯТИ





	mnemonic training	active control	passive control
pre-training 20 min	26.5 ±16.2	31.3 ±14.8	29.5 ±16.1
post-training 20 min	62.5 ±11.1	42.7 ±17.1	36.6 ±19.5
training change 20 min	+35.9 ±17.0	+11.4 ±11.8	+7.1 ±13.3
pre-training 24 h	16.5 ±14.0	19.6 ±12.7	18.7 ±15.7
post-training 24 h	55.7 ±16.9	31.1 ±18.6	21.8 ±19.1
training change 24 h	+39.2 ±17.8	+11.4 ±12.7	+3.1 ±10.7
4 month follow-up 15 min	50.3 ±16.5	30.4 ±9.5	27.4 ±9.8
4 month change 20/15 min	+22.4 ±18.9	+0.5 ±11.8	-2.2 ±11.4



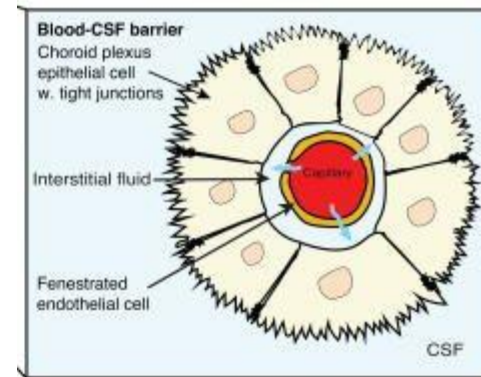
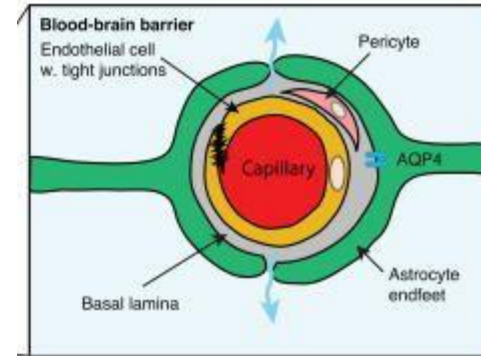
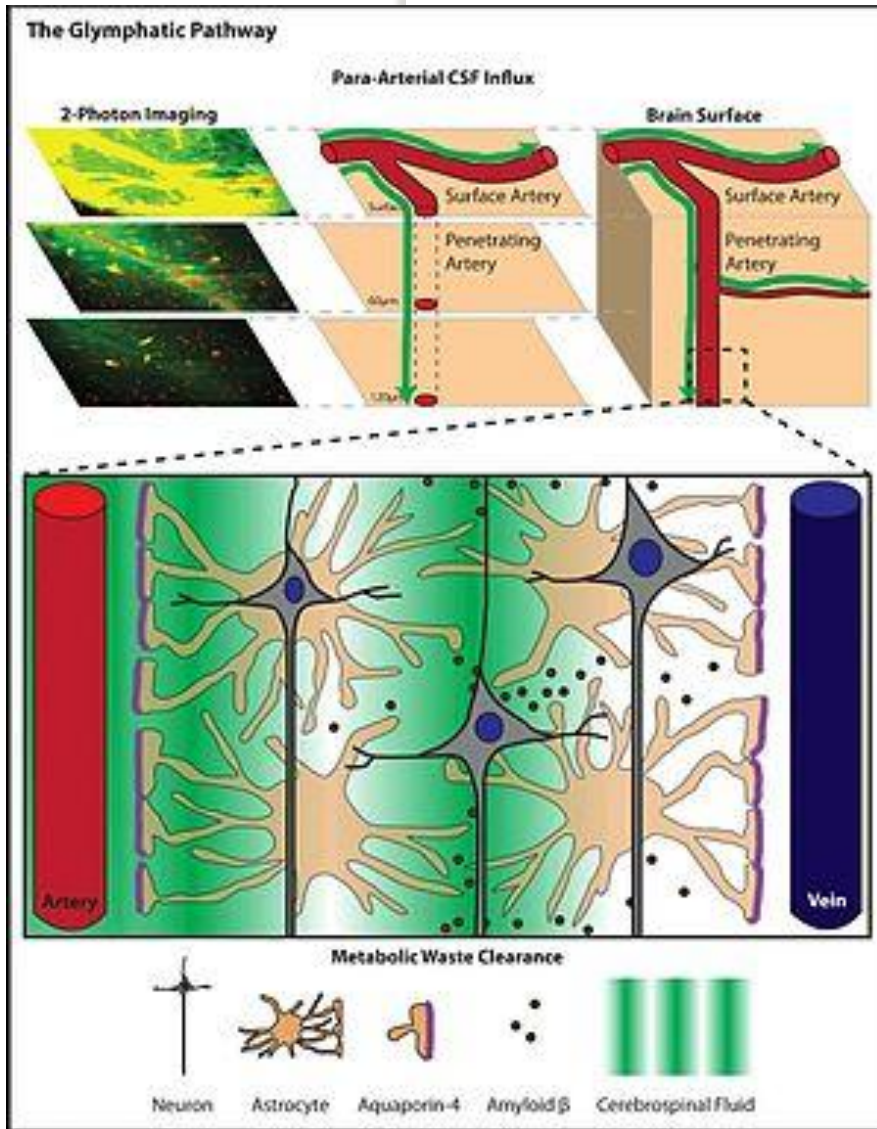
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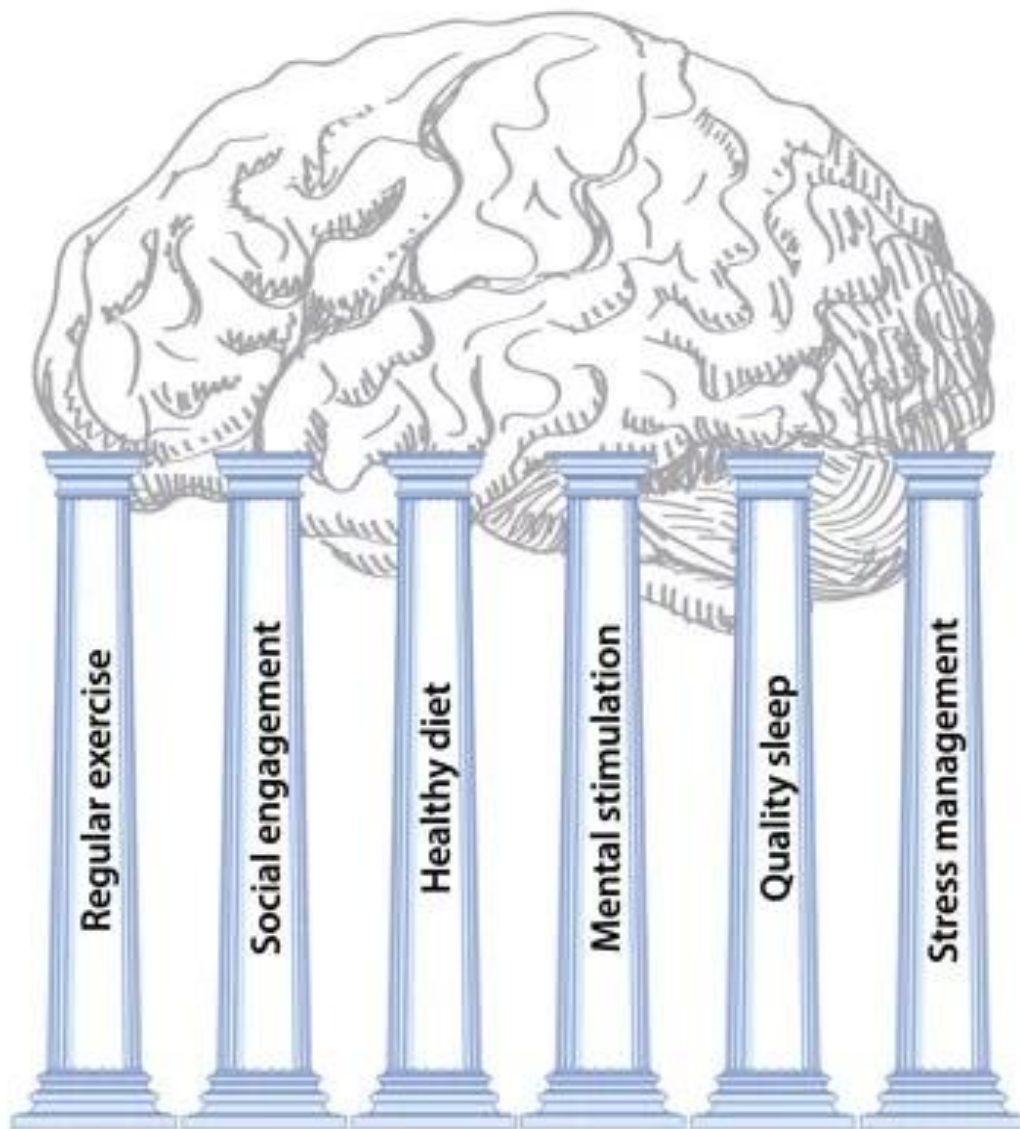


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Lee H, Xie L, Yu M, et al. The Effect of Body Posture on Brain Glymphatic Transport. *J Neurosci.* 2015;35(31):11034-44.

The 6 Pillars of Alzheimer's Prevention



**СПАСИБО ЗА
ВНИМАНИЕ!**