

Propojení výuky oborů Molekulární a
buněčné biologie a Ochrany a tvorby
životního prostředí OPVK
(CZ.1.07/2.2.00/28.0032)

Can We Make Our Brains More Plastic?

SCIENCE

5 OCTOBER 2012

VOL 388

Kateřina Kubáňová



INVESTICE DO ROZVOJE VZDĚLÁVÁNÍ

„ A fully plastic brain is not very helpful. It learns everything but remembers nothing.“

Gred Kempermann, neuroscientist

Center for Regenerative Therapies Dresden

German Center for Neurodegenerative Diseases

- too much plasticity may also play a role in some neurological disorders, including epilepsy and schizophrenia

□ understanding the flexibility of certain parts of the brain:

□ studying the development of sensory system

□ ↘ uncovered a network of genes and proteins that influence critical periods

□ **critical periods** = windows of time in which the brain is primed for certain types of input

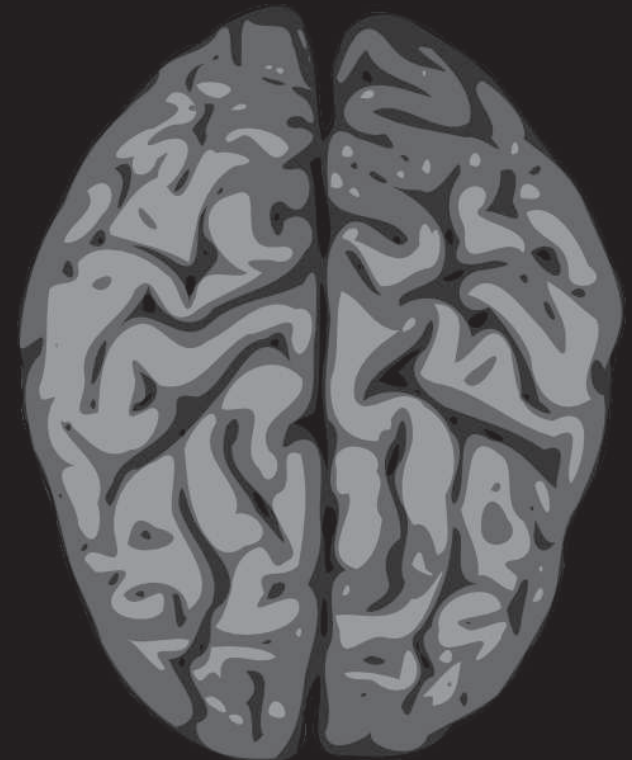
Critical Periods

□ brain becomes wired for certain tasks

↳ turning the signals received from eyes into recognizable images

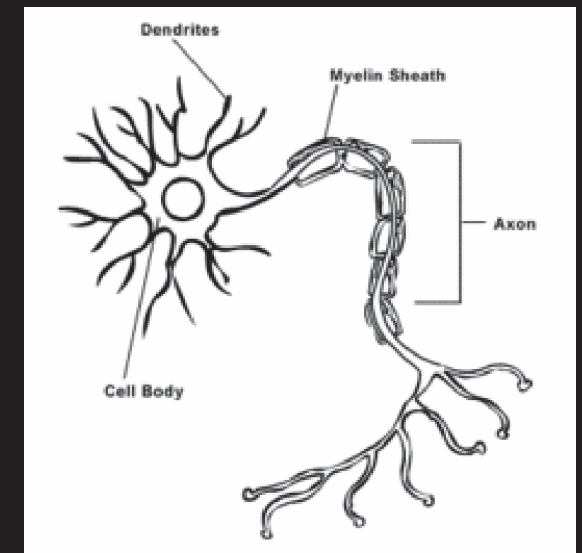
↳ distinguishing sounds present in spoken language

□ lack of inputs during a critical period → hard recovering



Critical Periods

- **earliest** → development govern senses such as sight, hearing and balance
- **later** → higher-order skills such as language acquisition and social interactions
- **most important** → connection the neurons make with each other



Critical Periods close

- ❑ decrease of the plasticity-driving signals
- ❑ produce of signals that limit new connections between cells

- ❑ scientists used genetic tricks to remove the brakes on brain plasticity in mice
 - the critical periods last well into adulthood

„Just take away the brakes and the brain can perhaps recover its lost capabilities.“

Carla Shatz, neuroscientist

Stanford University in Palo Alto, California

□ in lab animals it is possible

↳ researchers bred mice that lack some of the genes that act as plasticity brakes

Knock-out mice

- ❑ mutant mice recovered from stroke better
- ❑ several tests for neural activity
- ❑ good performance on the rotarod
- ↘ motor skills test for lab mice
- ❑ range of behavioral tests
- supermice

„That’s certainly not the whole story. There has to be some downside.“

Carla Shatz, neuroscientist

Stanford University in Palo Alto, California

- ❑ much rewiring can lead to short circuits in the brain → seizures
- ❑ knock-out mice responded to smaller dose of seizure-inducing drugs

- in humans → result of the unleashing brain plasticity might be epilepsy
- **epilepsy** → much more common in childhood
- closing critical periods may also provide a firm foundation for further brain development
- missing plasticity brakes are suspected not only in epilepsy but also in schizophrenia and Alzheimer's disease

□ brain plasticity can be augmented without completely removing the brakes

□ certain kinds of sensory signals can rewire adult brains

↳ mainly **sound** and **touch**

Michale Merzenich neuroscientist, University of California

□ specially designed computer games can improve performance on memory and other cognitive tasks in both children and older adults

↳ even months after the training stops

Daphne Bavelier neuroscientist, University of Geneva, Switzerland

- playing action video games can improve vision and several kinds of cognitive skills

The succes of games

- linked to the brain's reward and attention systems
- several of the molecules identified as plasticity brakes involve these pathways

□ two drugs enhance attention:

□ fluoxetine (known as Prozac)

□ Aricept

↳ can lengthen or even reopen critical periods in experimental mice

□ both drugs in clinical trials for reversing the effects of lazy eye in childhood

□ fluoxetine → helped stroke patients recover lost motor skills

Fluoxetine

- influences the growth of new neurons
- most neurogenesis stops in childhood
- **two areas of the brain** keep producing new neurons:

↳ **subventricular zone** (connects to olfactory bulb)

↳ **subgranular zone** of the dentate gyrus (part of the hippocampus)

- several way how to boost the production of the new neurons in these regions
 - ↳ increase physical exercise
 - ↳ exposure to unfamiliar or complex environments

- fluoxetine and other antidepressants that act through the dopamine pathway also increase the neuronal birthrate and may keep the newborn neurons flexible longer

- what this ongoing production of neurons means for the brain is unclear
- **idea** → new neurons may aid the brain in adjusting to new environments, perhaps by helping the brain detect unfamiliar aspects of an otherwise familiar setting
- **new neurons** → have their own critical period, lasting roughly 4 weeks, during which they are particularly excitable (fluoxetine might lengthen this period)

Propojení výuky oborů Molekulární a buněčné biologie a Ochrany a tvorby
životního prostředí OPVK (CZ.1.07/2.2.00/28.0032)

BLOOD TO BLOOD

KAROLÍNA VAVROUŠKOVÁ

ALŽBĚTA ZLOCHOVÁ

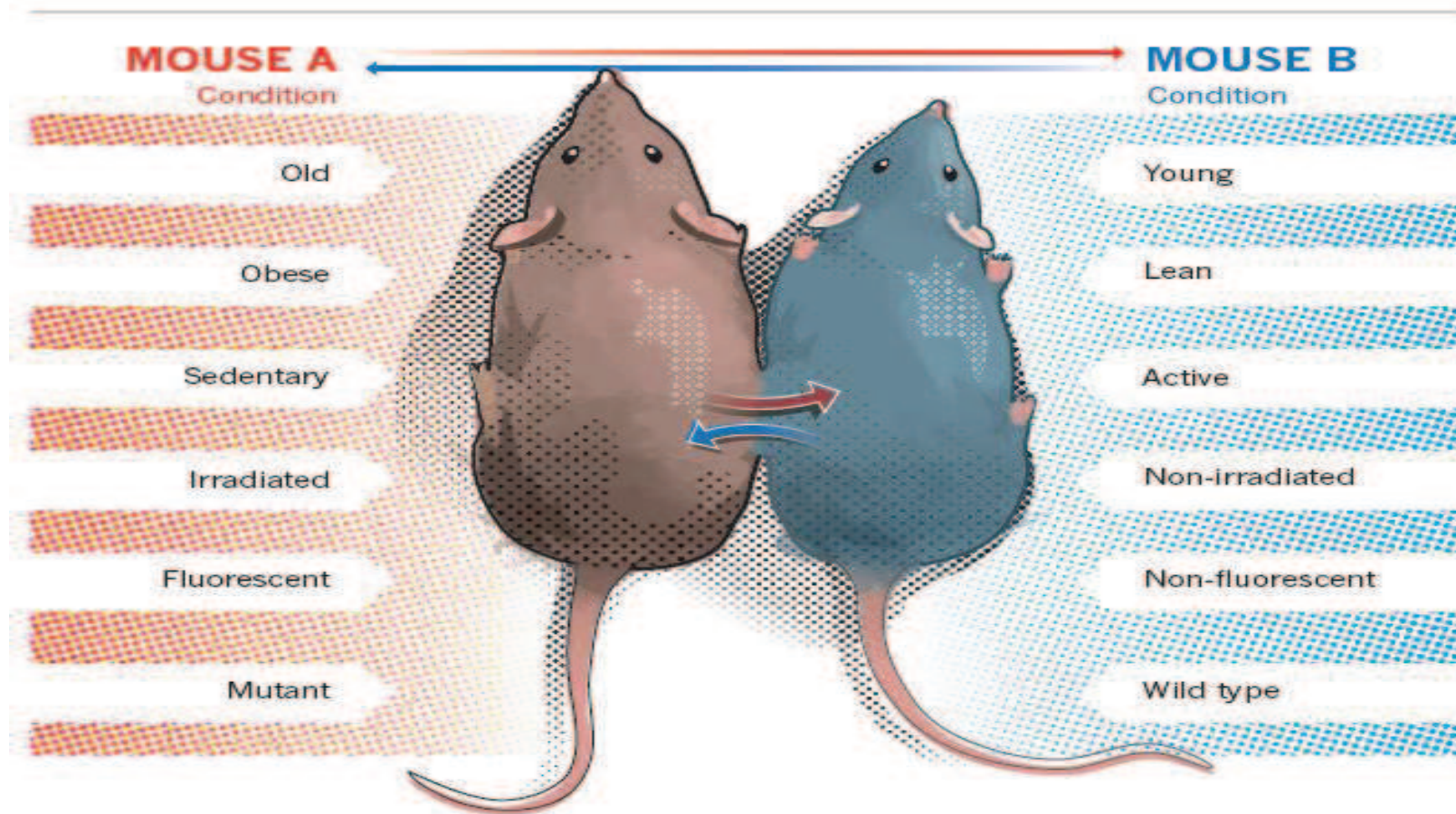
MBB

-
- Nature
 - 22 January 2015
 - Megan Scudellari

Parabiosis

- 150-year-old surgical technique that unites the vasculature of two living animals
- From Greek *PARA* → alongside
- *BIOS* → life

- Test what circulating factors in the blood one of the animal do when they enter another animal



Experiments

- Share the circulatory system of an old mouse and young mouse
 - Remarkable results → heart, brain, muscles
 - Old mice → stronger, smarter and healthier
 - Bring new life to old bodies
-
- Now → scientists have begun to identify the components of young blood
 - September → a clinical trial in California-the first who have started testing young blood and Alzheimer's disease

1864

- ❑ Physiologist Paul Bert → removed the strip of skin of two rats and stitched the animals together
- ❑ Hoped they could create a shared circulatory system
- ❑ He found that fluid injected into a vein of one rat passed easily into the other
- ❑ Won an award in 1866
- ❑ After him → one team ruled out the idea that dental cavities are results of sugar in the blood by using a pair of parabiosed rats, only one was fed by glucose
- ❑ The rats → same blood glucose levels, but only one rat had dental cavities

1956

- Clive McCoy from Cornell University in New York → the first one who apply parabiosis to the study of ageing
- 69 pairs of rats, all of different ages
- For example 1,5-month-old was paired with 16-month old rat → the equivalent of 5-year old human with a 47-year old
- Not successful → 11 pairs died, rats were not adjusted and they started eating each other
- Must be socialized with each other
- Results → the older animals' bones became similar in weight and density of the bones of younger rats

1972

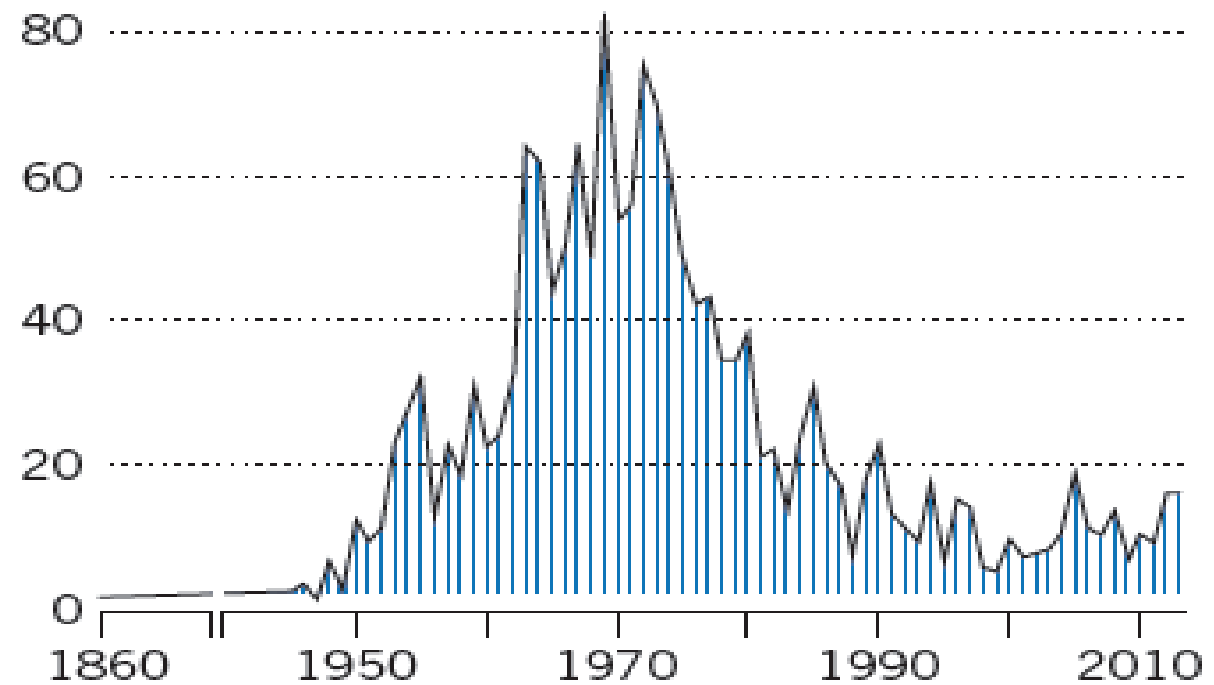
- Two researches at Univeristy of California studied lifespans
- Older partners lived for four to five months longer → the young blood might affect longevity

Despite these findings, parabiosis fell out of use

Publications on parabiosis

Parabiosis gained popularity during the 1960s and 1970s, but eventually fell out of wide practice.

Fell out of
favour after
1970s



Blood components

- ❑ **What exactly in the blood is responsible for the rejuvenating effects?**
- ❑ OXYTOCIN → a hormone of love
- ❑ Young blood can also help to form new neurons and reverse age-related thickening of walls of the heart

-
- ❑ start screening the proteins of young blood → GDF11
 - ❑ → increase the strenght and stamina of muscles

 - ❑ Start screening plasma → plasma activates brain plasticity in older mice and increase memory

Future...

- ❑ Alzheimer disease → neuron loss
- ❑ Experiment → condition of one patient got better after he received a plasma transfusion

- ❑ Six of out a planned 18 people with Alzheimer's, all aged 50 or above, have already begun to receive plasma harvested from men aged 30 or younger

Negatives

❑ Scientist are afraid of failure

❑ For now → any claims that young blood or plasma will extend lifespan are false → the data are just not there

Thank you for your attention 😊