

# 3rd European Initiative for Exercise in Medicine

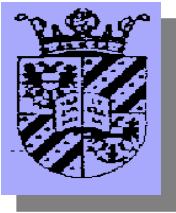
## Physical activity and Brain function

Gertjan van Dijk

Center for Behaviour & Neurosciences and

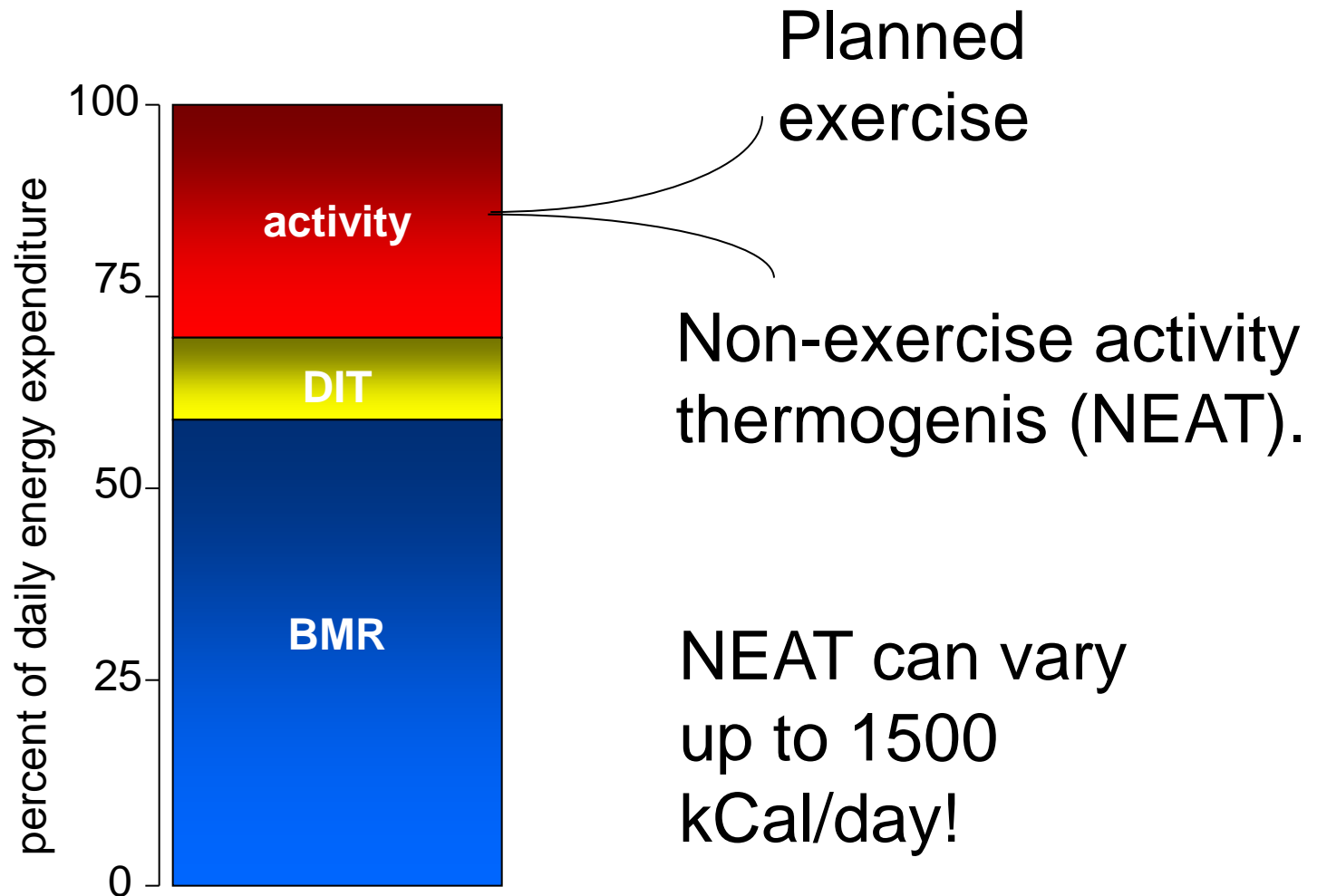
Center for Isotope Research

University of Groningen, the Netherlands

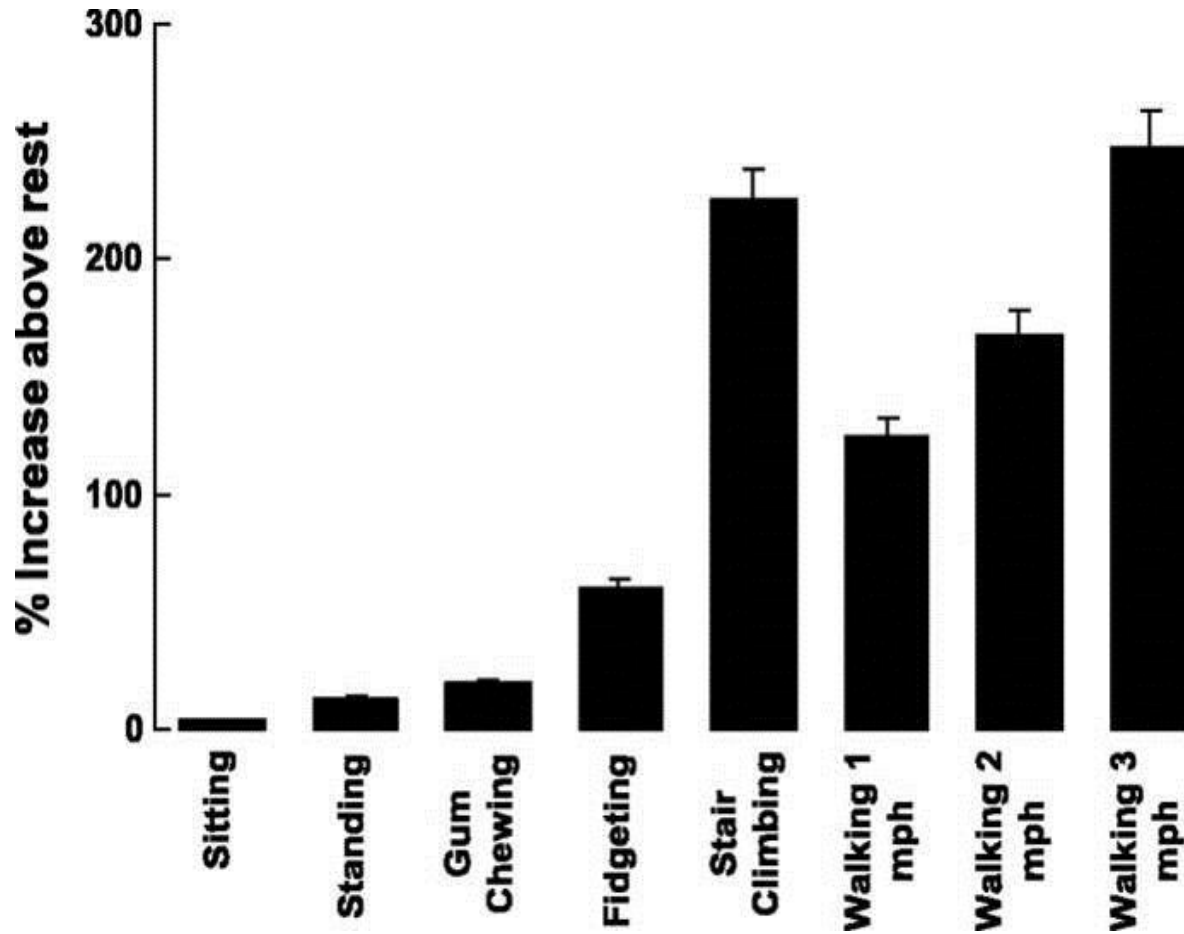




# Daily energy expenditure

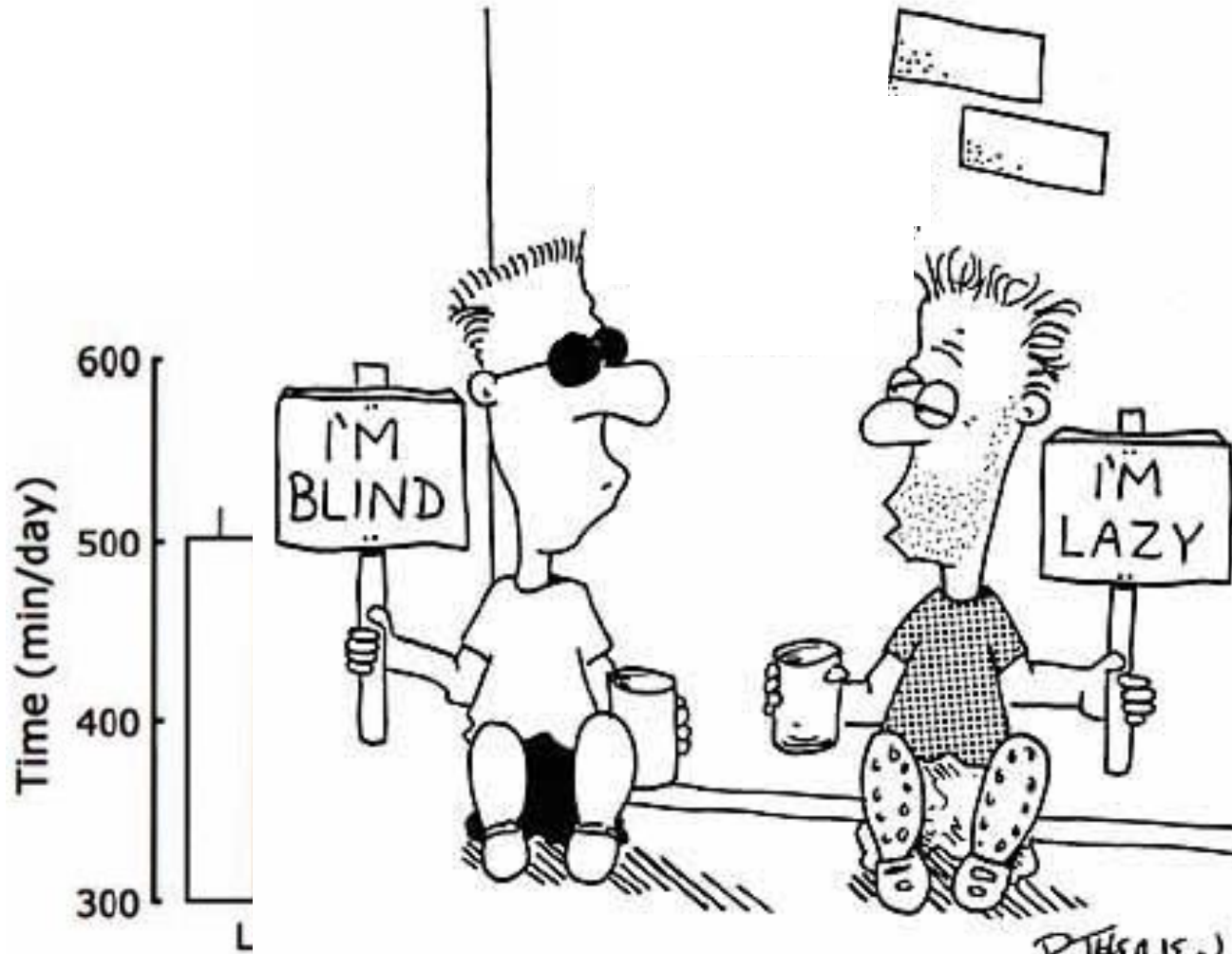


# Energy expenditure of varied low-level activities (James Levine)



Levine, J. A. Am J Physiol Endocrinol Metab 286: E675-E685 2004;  
doi:10.1152/ajpendo.00562.2003

# Changes in NFAT and weight gain



**“No, really... I was born this way.”**

cts were  
nin  
ay  
s were  
nin  
ay  
(lying)  
identical  
ovement  
ely  
ith fat

# Weight gain = energy intake (EI)- expenditure (EE)

1920' vs 2000:

Gradual decline of EE, but  
almost no difference in EI

the Netherlands in 2011:

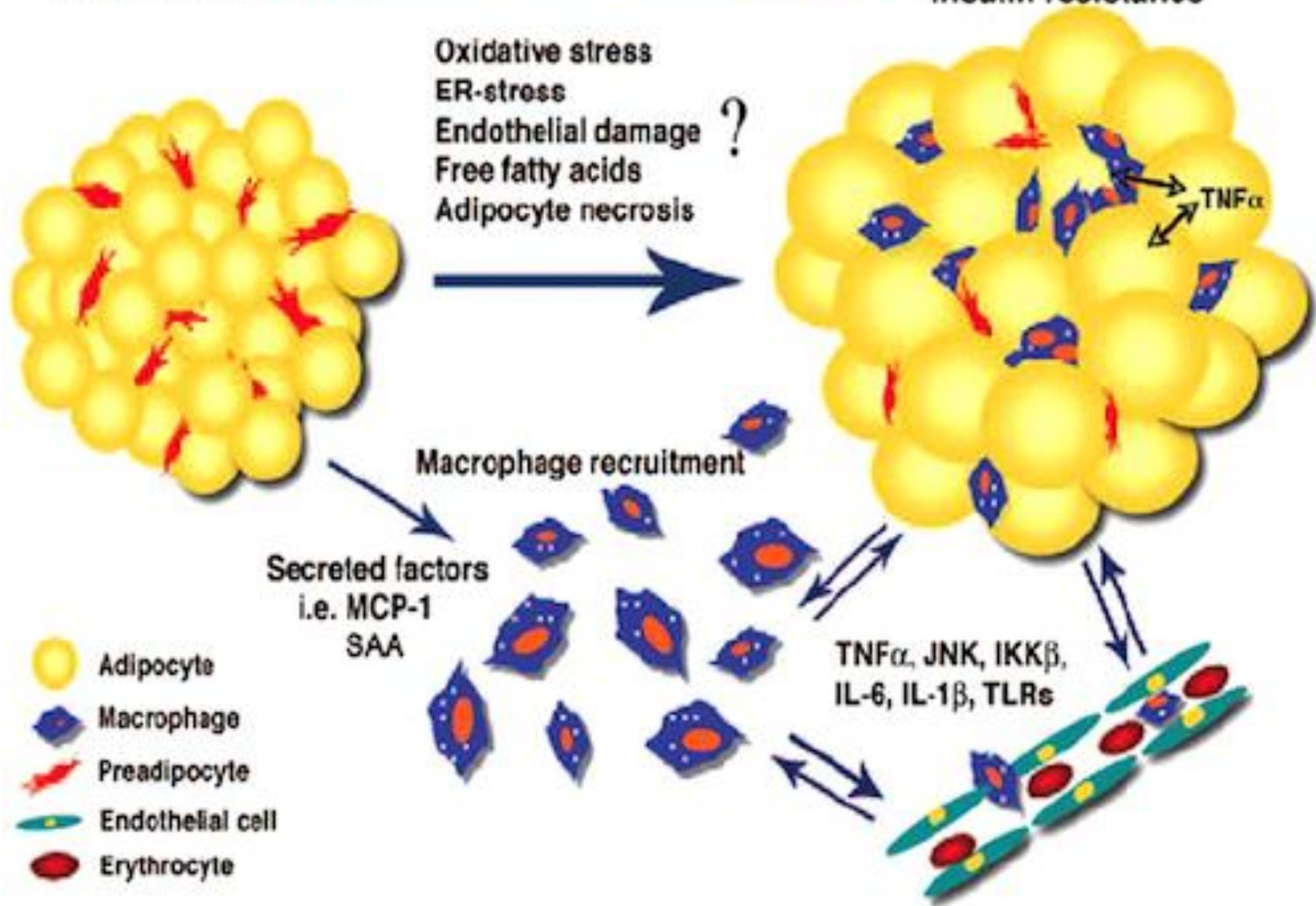
less than 40% meets the  
minimal requirement:  
5x per week 30 min/day  
moderate exercise



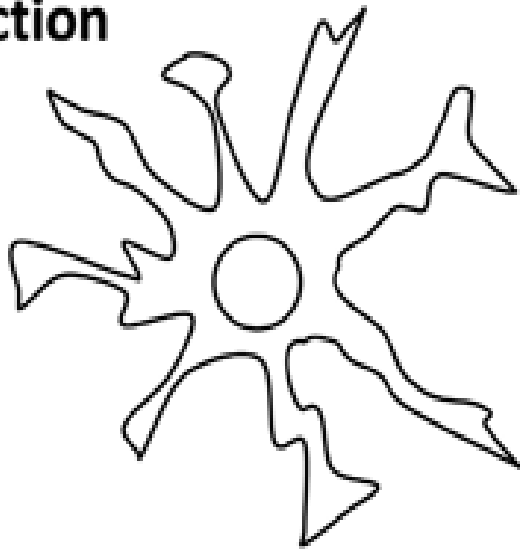
Weight gain

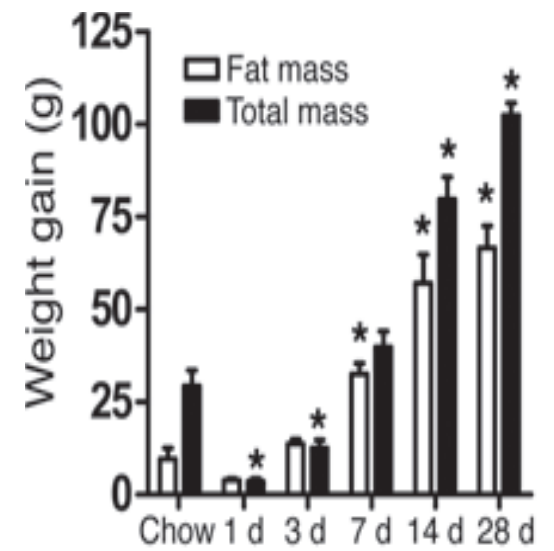
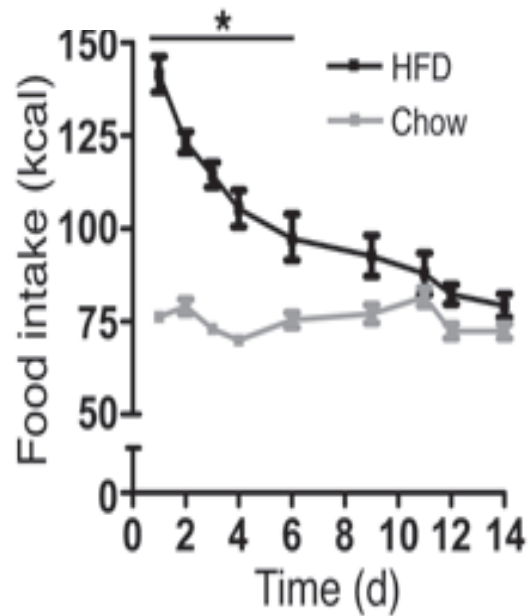
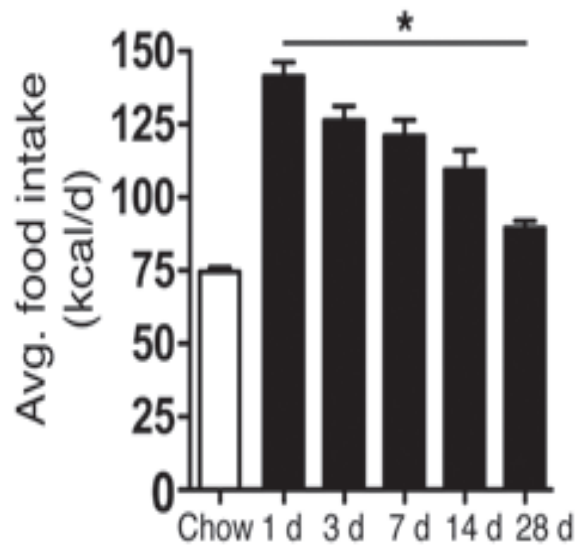


Adipose tissue inflammation  
Insulin resistance

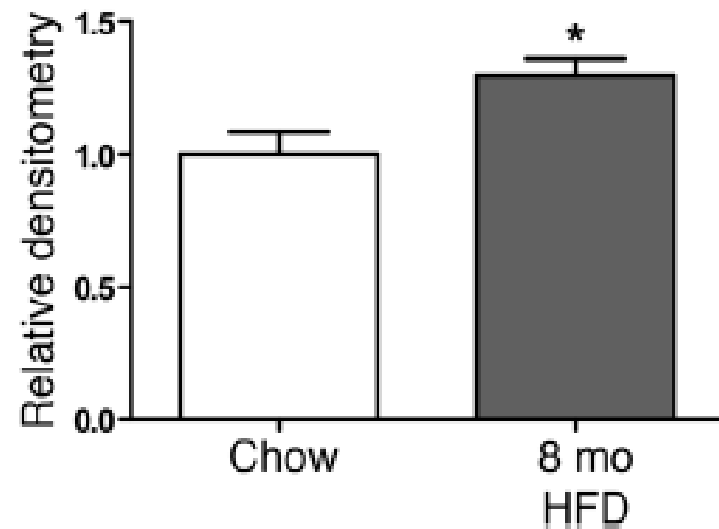
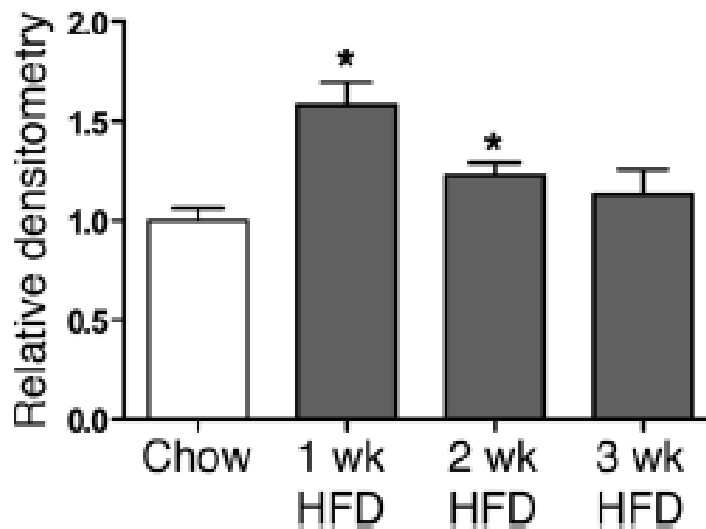
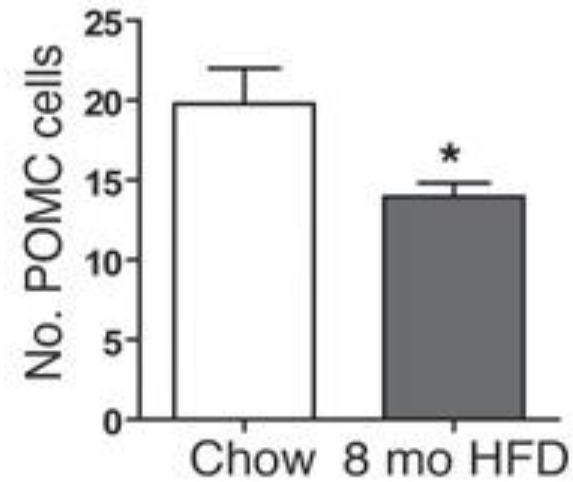
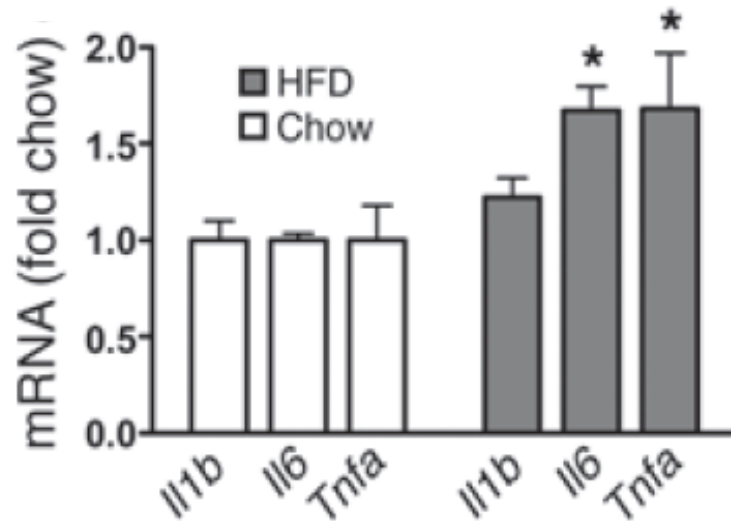


**Astrocytes preserve  
and promote CNS  
function**

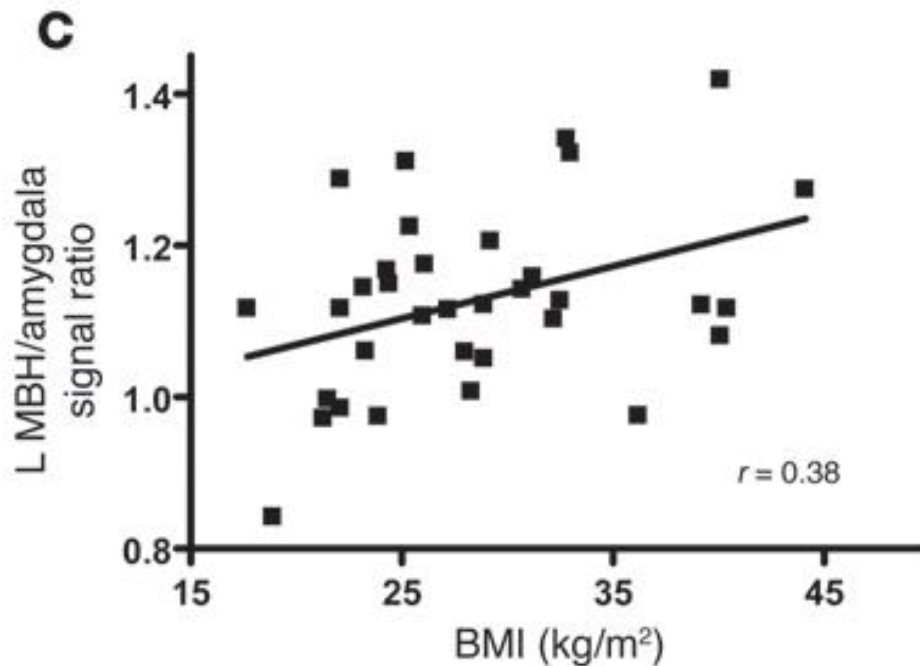
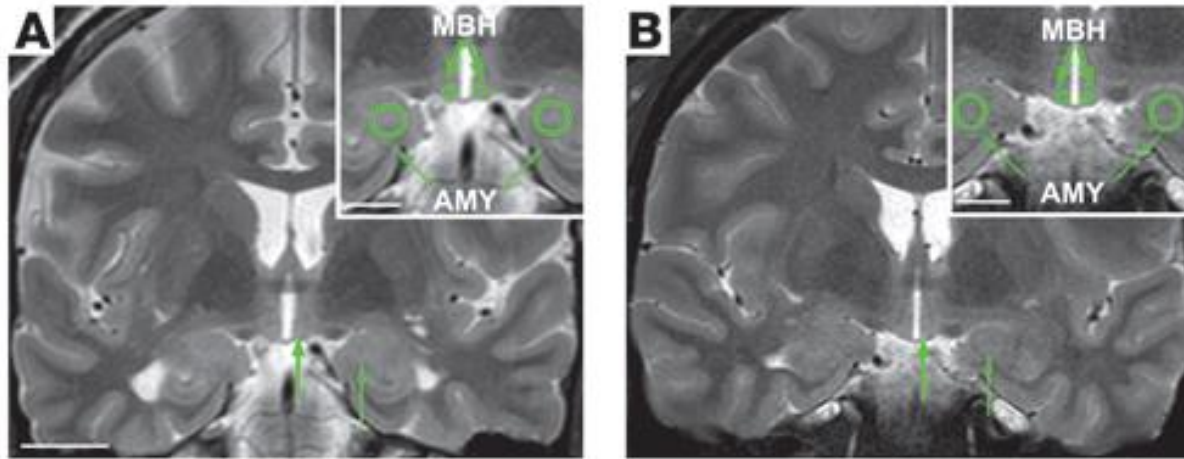




# Brain inflammation by HF diet



# Gliososis in obese subjects



# Association of diabetes mellitus and dementia: The Rotterdam Study

Diabetologia (1996) 39: 1392–1397

A. Ott<sup>1</sup>, R.P. Stolck<sup>1</sup>, A. Hofman<sup>1</sup>, F. van Harskamp<sup>2</sup>, D.E. Grobbee<sup>1</sup>, M.M.B. Breteler<sup>1</sup>

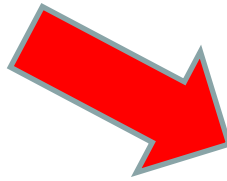
<sup>1</sup> Department of Epidemiology & Biostatistics, Erasmus University Medical School, Rotterdam, The Netherlands

<sup>2</sup> Department of Neurology, Erasmus University Medical School, Rotterdam, The Netherlands

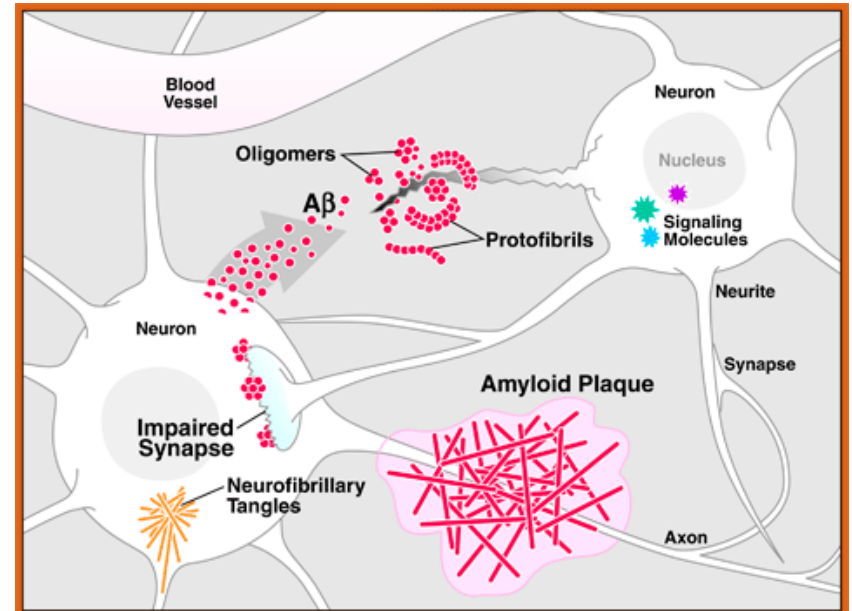


Type  
**Diabetes**  
**1**

Type  
**Diabetes**  
**2**



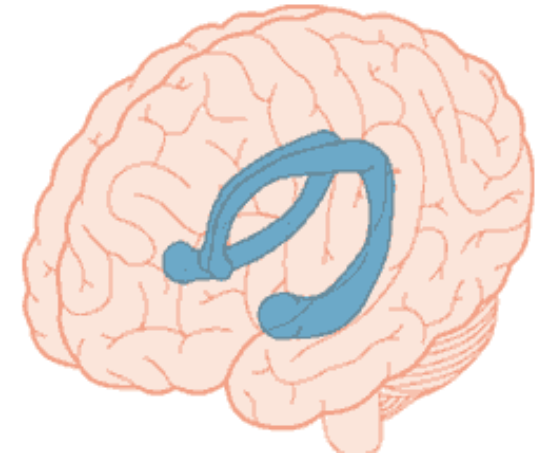
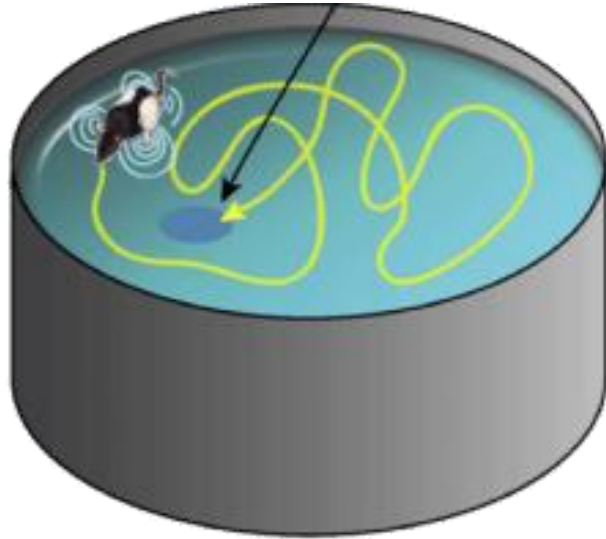
2-fold higher risk!!



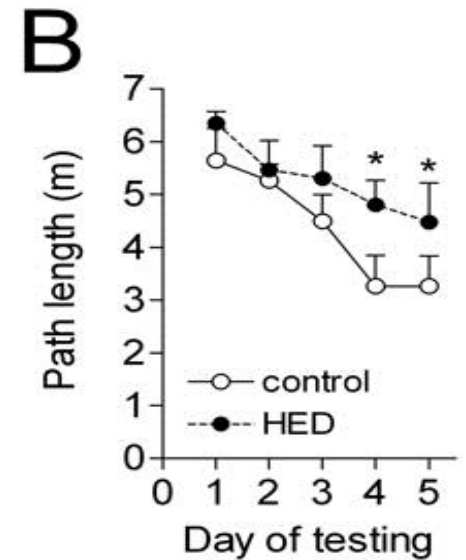
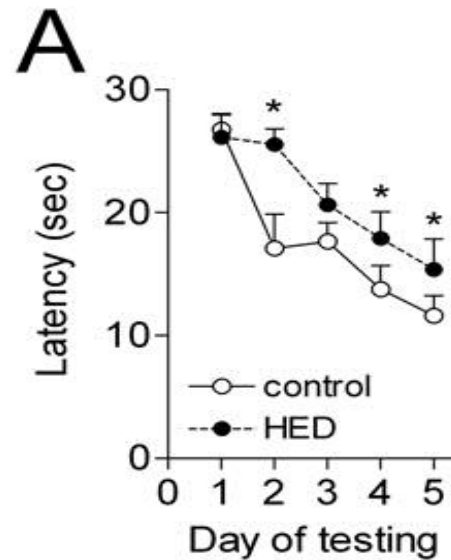
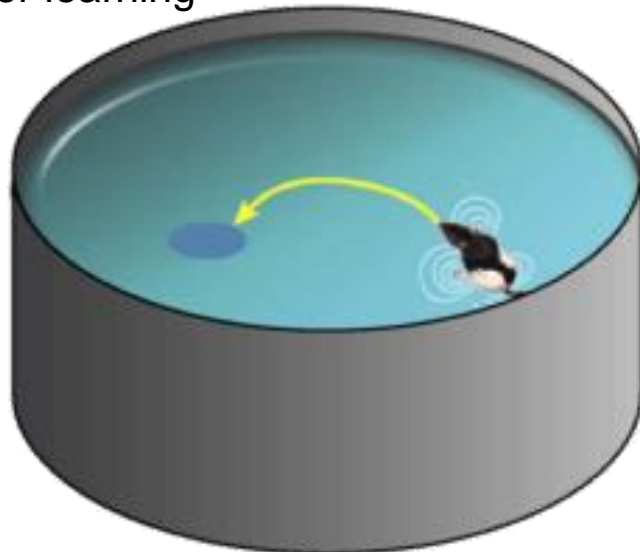
# Spatial memory

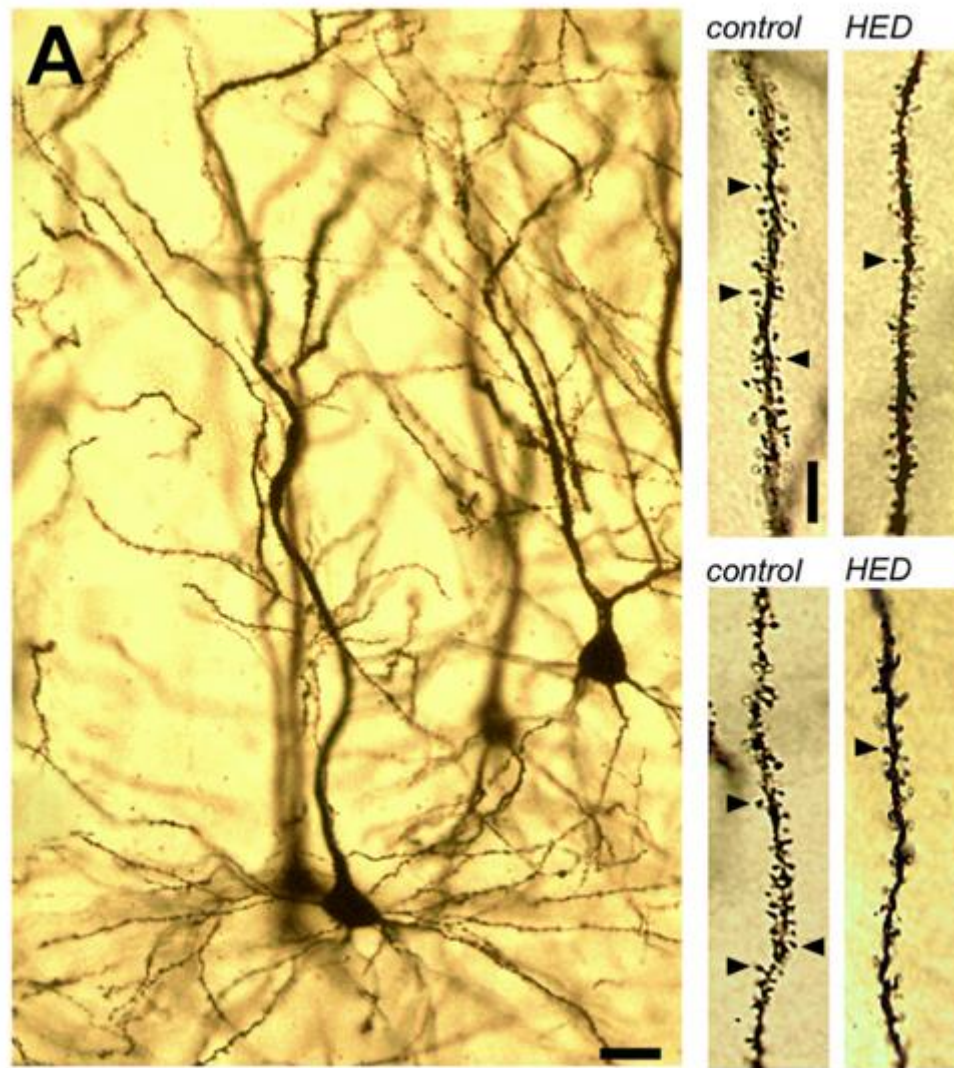
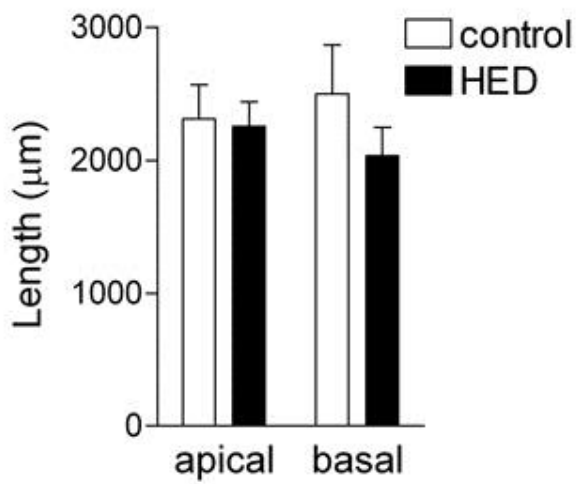
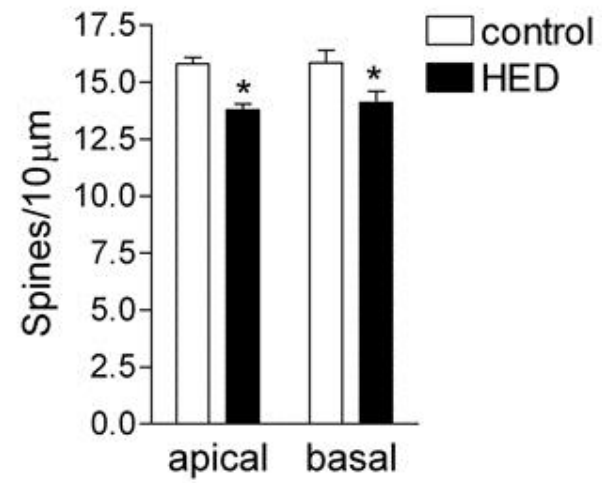
During learning

Hidden platform



After learning

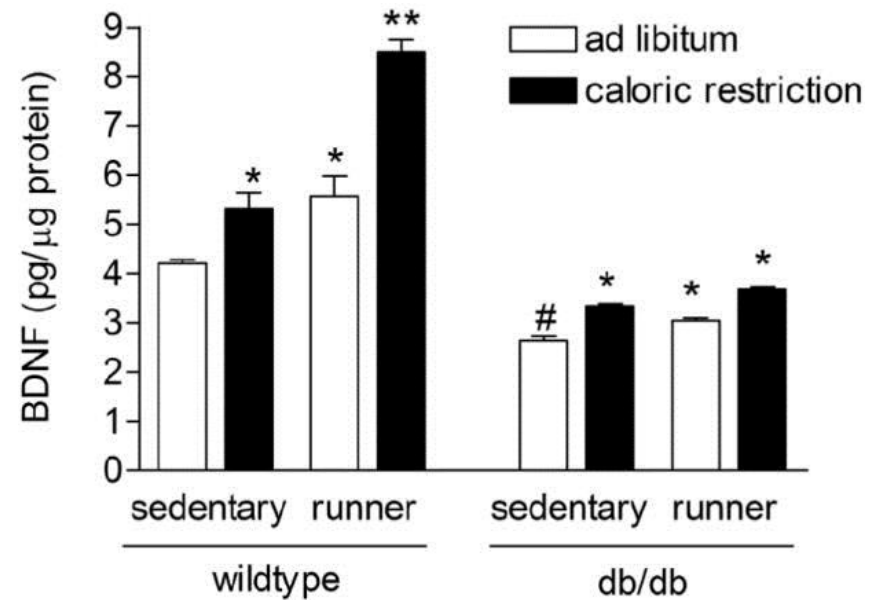
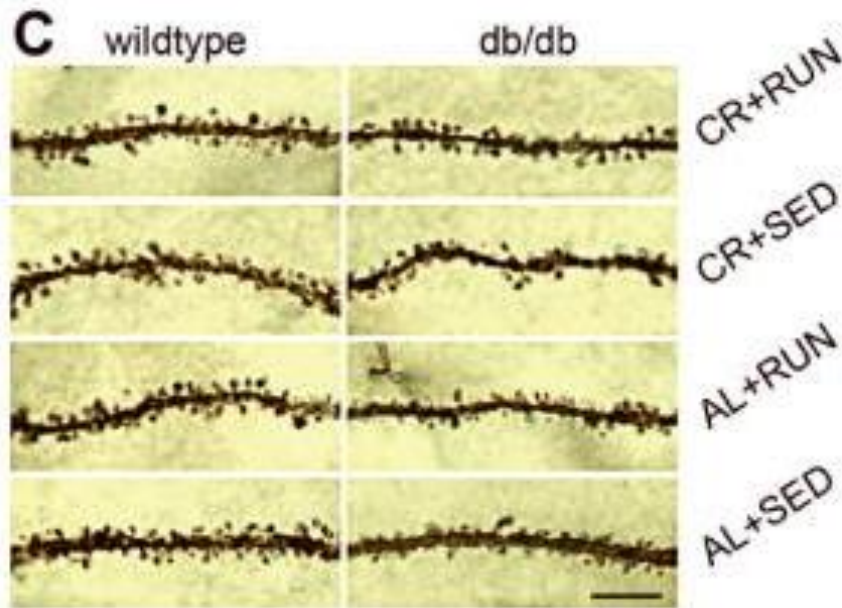
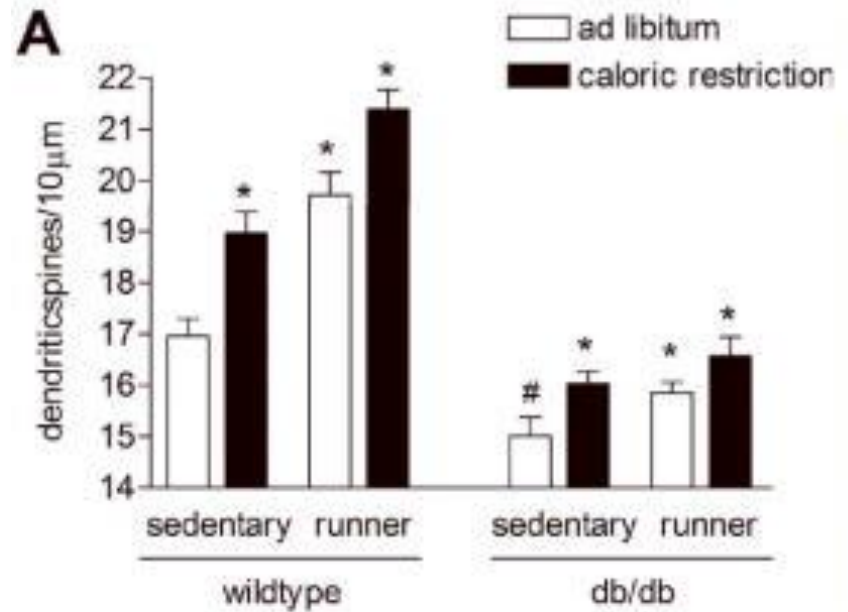




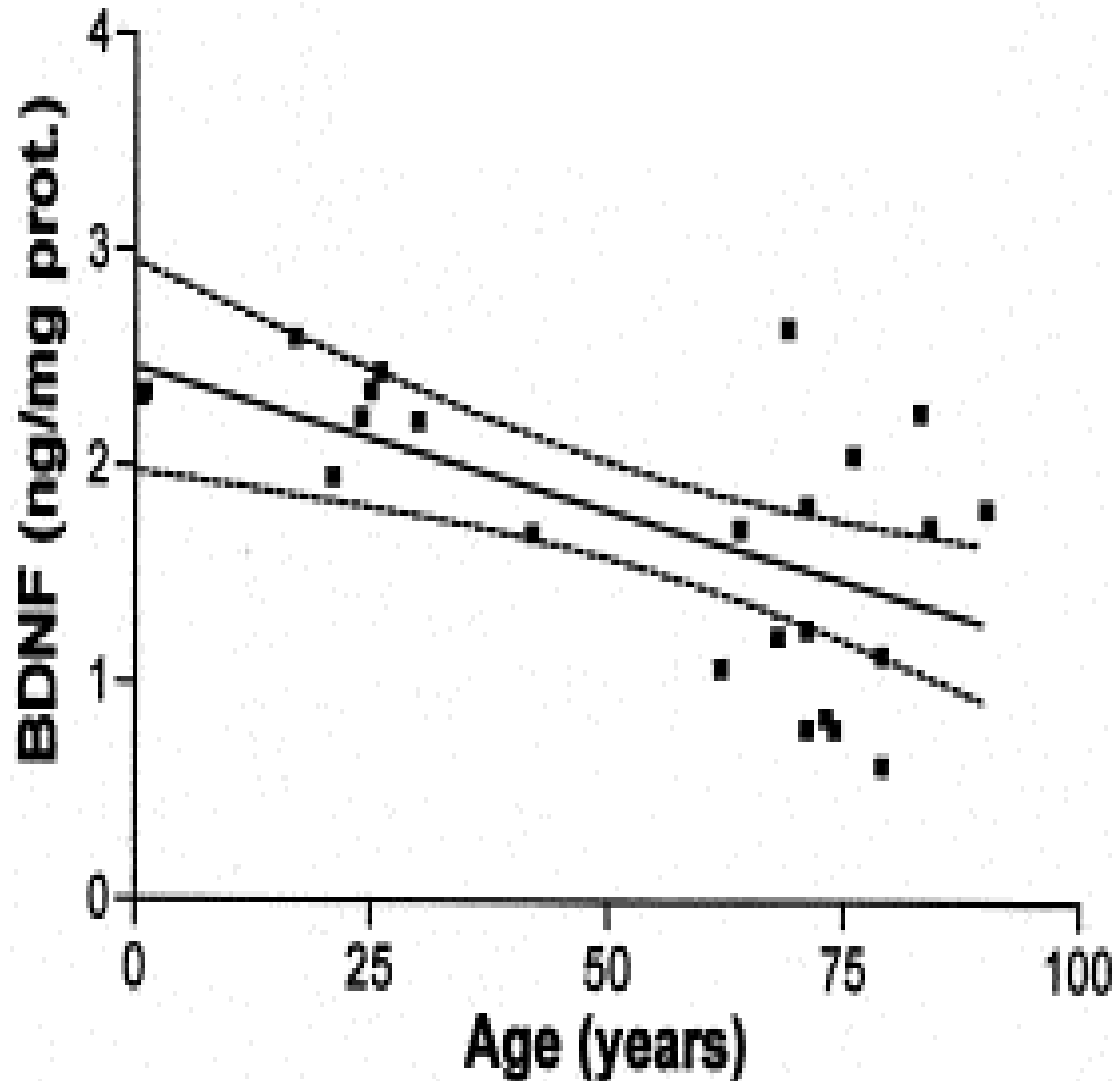
# Is there hope???



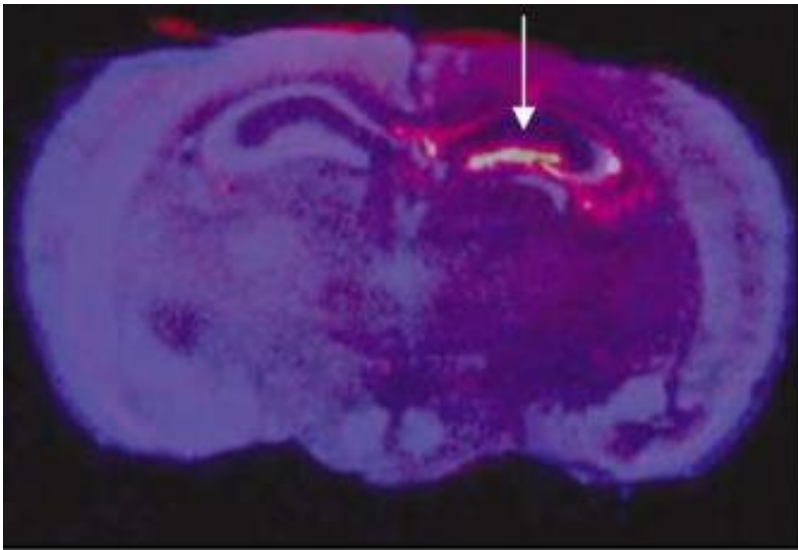




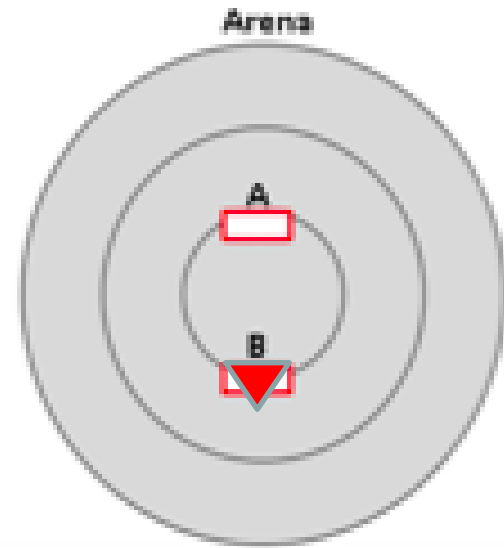
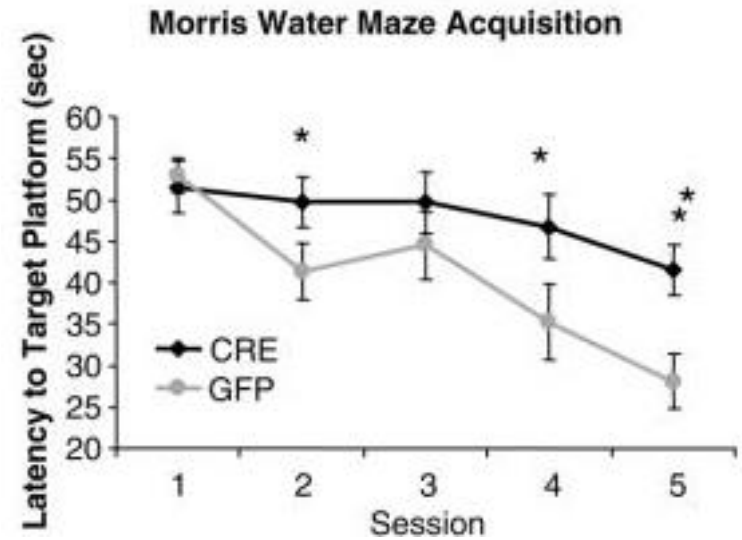
# Ageing decrease BDNF



# Site-specific deletion of BDNF-gene in hippocampus by Cre-Lox



Heldt et al Mol Psychiatry 2007



# What if one is not banning junk food, but tries to compensate with physical activity?

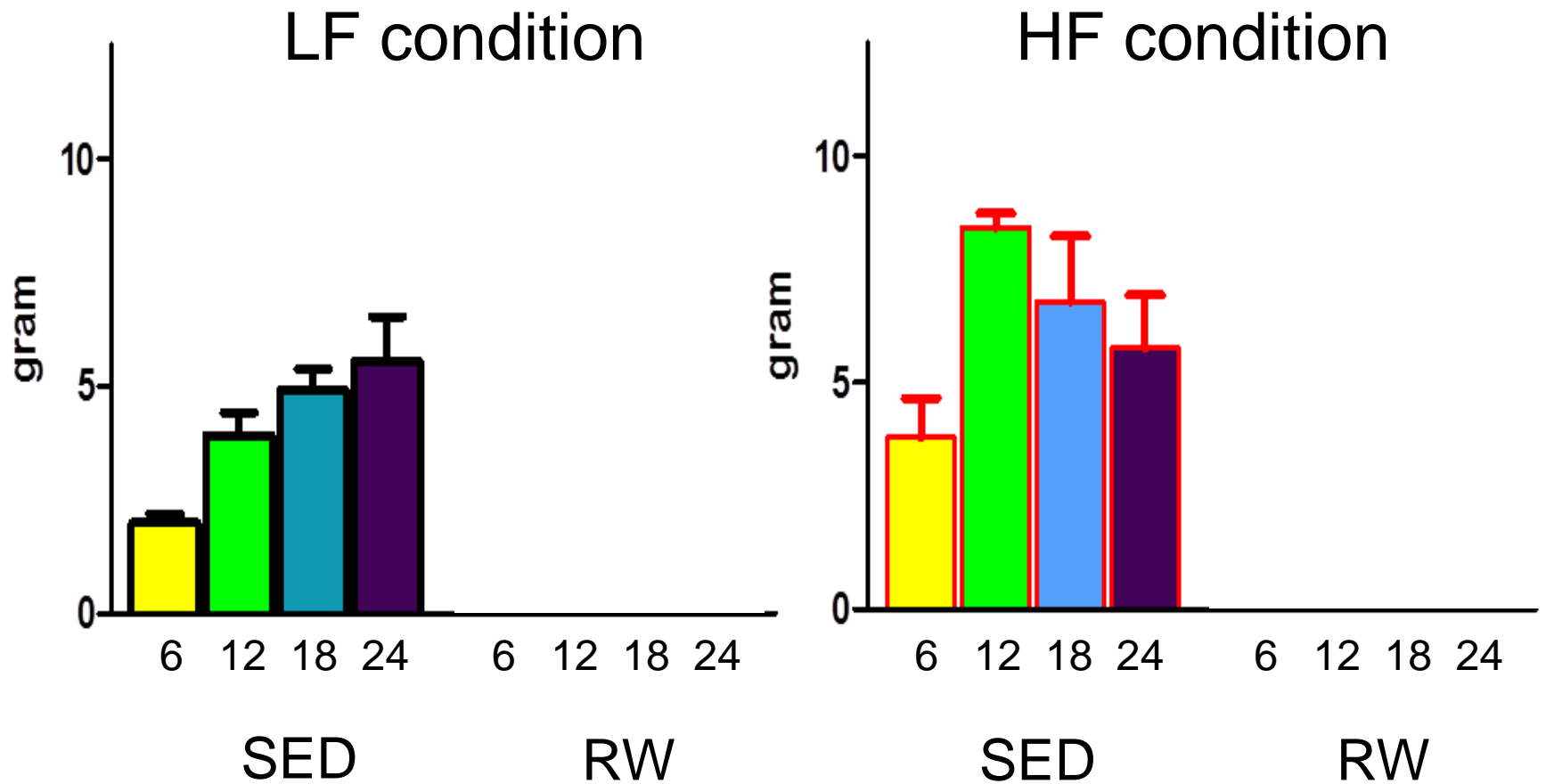


*“Effect of diet and physical activity on the systems biology of ageing”*

Male C57 Bl6 mice housed under:  
LF - HF SED  
LF - HF RW  
LF - HF CR

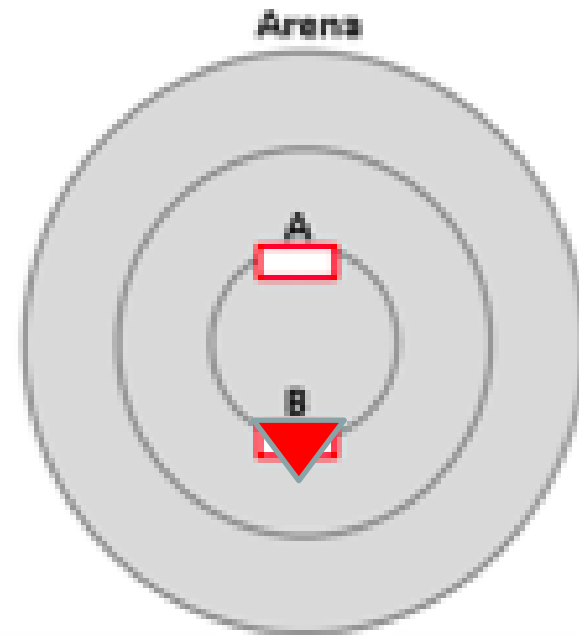
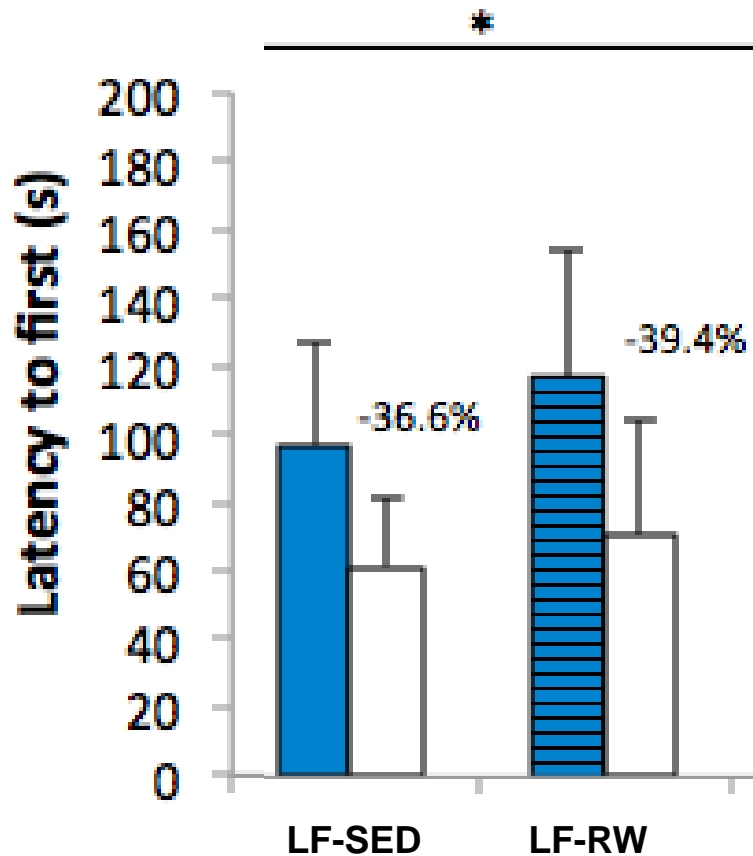
-sacrifice at 6, 12, 18 and 24 mnth  
-lifespan groups

# Visceral fat



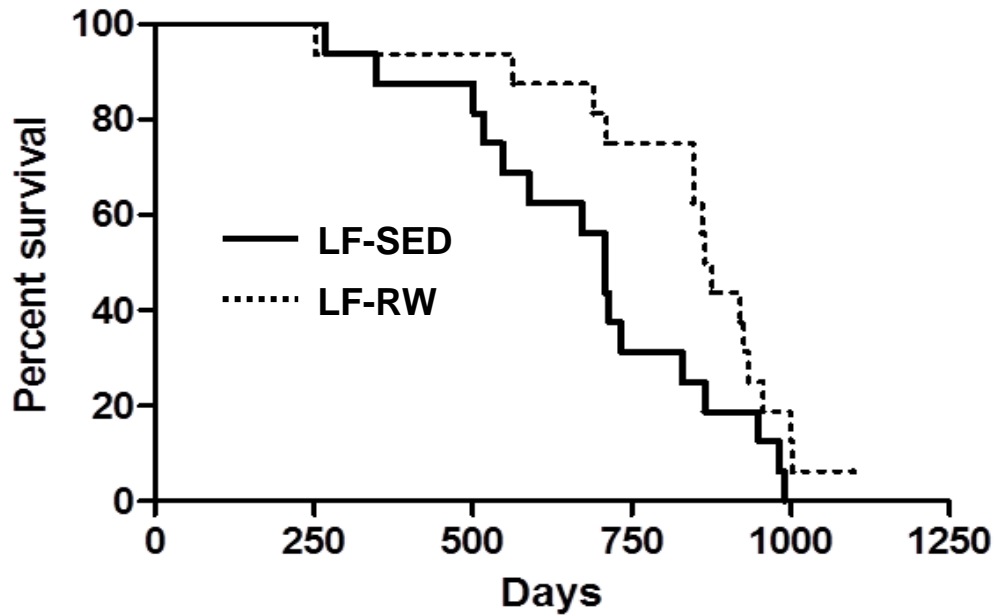
# Behavioral tests: NOR

(age 21 months)



FO NO

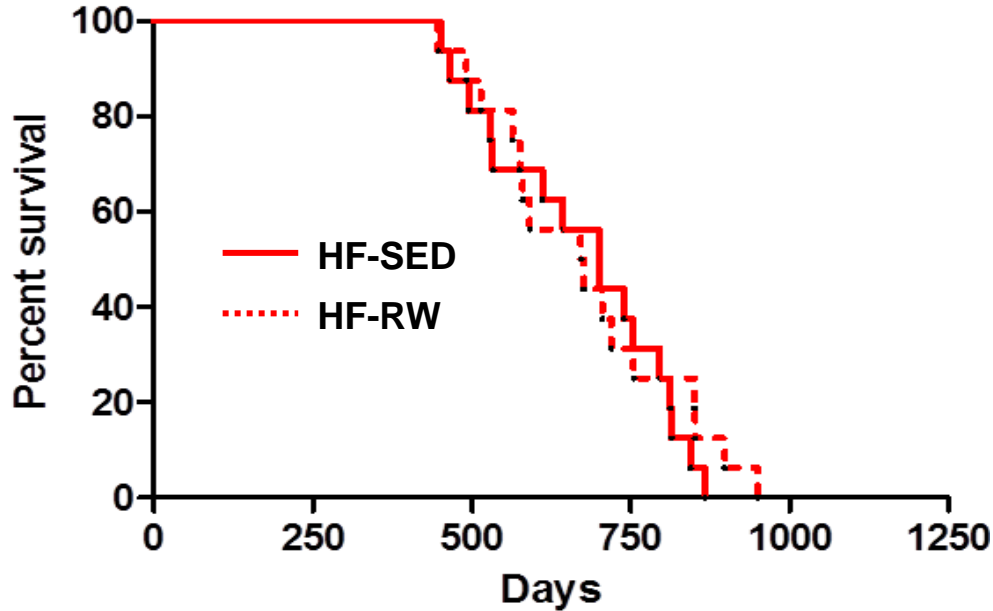
LF diet



Median life span  
LF-SED 707 days  
LF-RW 870 days

Maximum lifespan  
LF-SED 990 days  
LF-RW > 1120 days

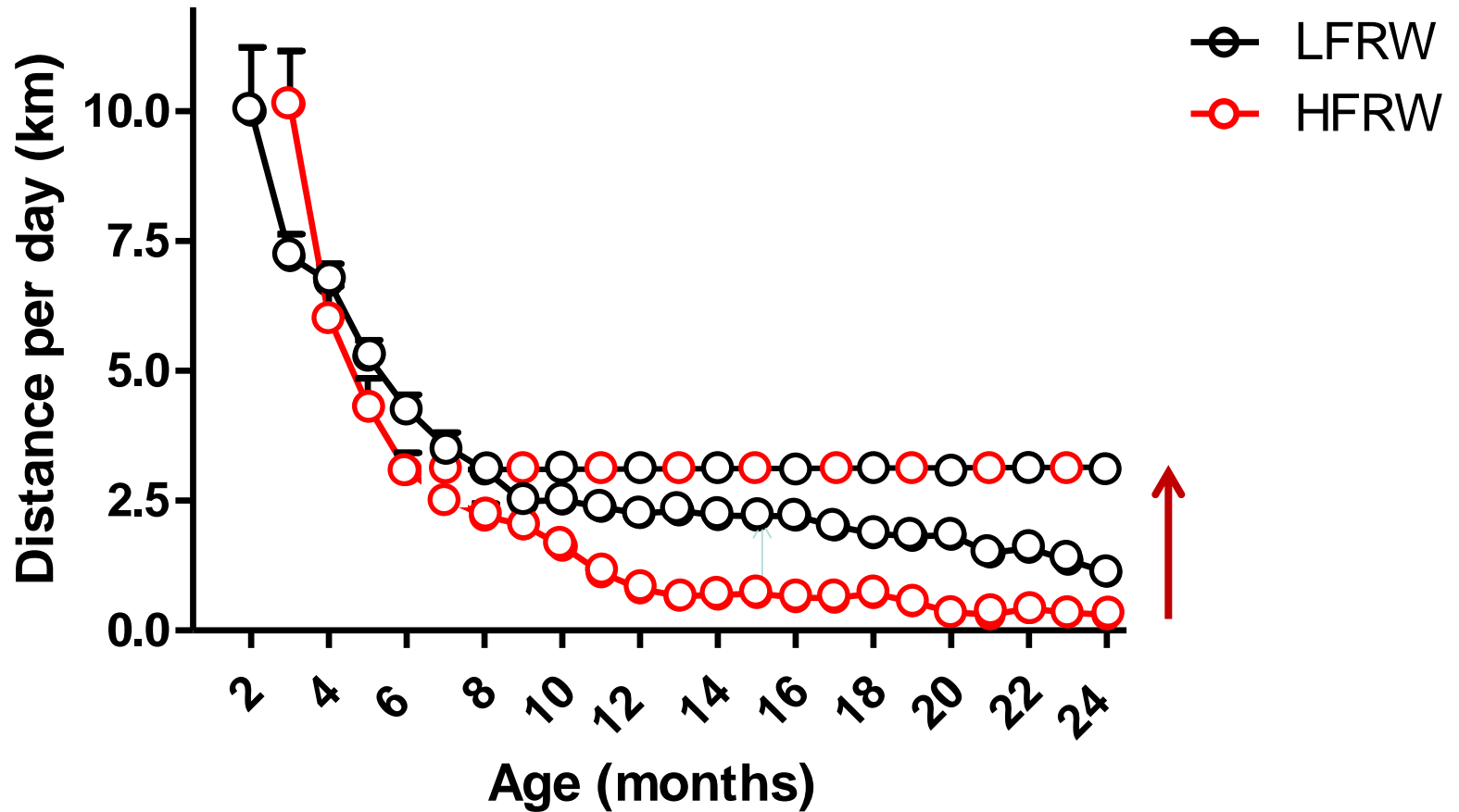
HF diet



Median life span  
HF-SED 700 days  
HF-RW 674 days

Maximum life span  
HF-SED 887  
HF-RW 950

# Running wheel activity





# Semmelweis University-Budapest, Hungary

## Faculty of Physical Education and Sport Sciences

### *Research Institute of Sport Sciences*

- Muscular mechanisms and adaptation in physical exercise
- Physical activity and health, health promotion, sport performance
- Neurodegenerative illnesses
- Ageing and well being through exercise



Laboratory of Endurance Physiology



Biochemistry Laboratory



Laboratory animal house



# Effects of life-long exercise on brain ageing

## **Life-long exercise protocol:**

Male Wistar rats

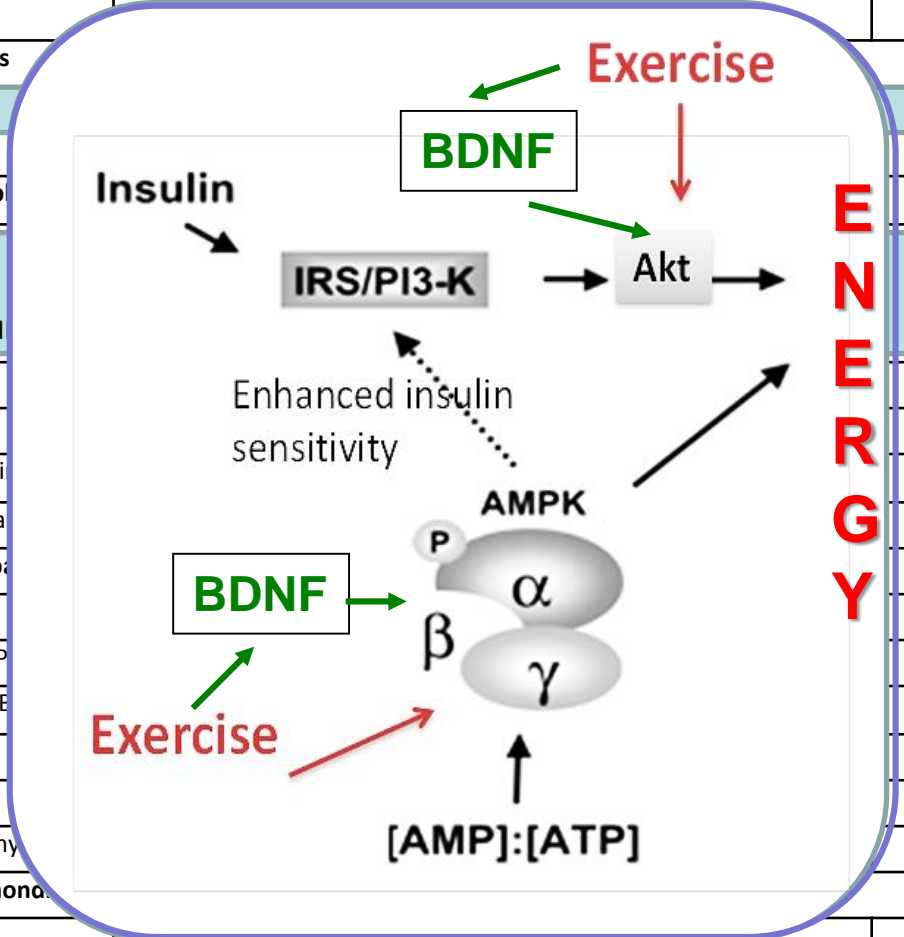
Moderate intensity treadmill running (15 m/min)

3 training sessions per week (3\*60 min)

From the age of 3 month to 24 months

proteins	sedentary	exercise	% change to sedentary controls	p=
<b>Cholinergic brain</b>				
CHAT				0.024
<b>Chronic factors</b>				
BDNF				0.019
DCX				0.012
<b>Energy supp</b>				
Glut-1				0.049
p-Akt/Akt				0.047
p-AMPK/AM				0.012
<b>Synaptic support</b>				
myloidPP				0.055
synaptophysin				0.079
p-synapsin/synap				0.045
<b>MAPK-CREB signal p</b>				
ERalpha				ns
p-MAPK/MAP				ns
p-CREB/CREB				ns
<b>Oxidative stress</b>				
ROS				0.045
Protein Carbonyl				0.019
<b>Immune and mitochond</b>				
NFkappaB				ns
PGC-1	3.17 ± 0.38	3.79 ± 0.24	20	ns
NRF-1	1.14 ± 0.14	1.59 ± 0.13	39	ns
mTTFA	2.68 ± 0.30	2.98 ± 0.22	11	ns

**Short Summary Molecular aspects**



# Conclusion

- Feeding a HF diet causes brain inflammation, and increases the risk for neurodegenerative disorders.
- Physical activity improves physical and mental fitness, and extends life span when feeding a healthy fibered LF diet, but not when feeding a HF diet.
- HF feeding reduces voluntary physical activity.

# Thanks to

## University of Groningen

- Drs F Reijne
- Dr B Talarovicova
- Prof B. Groen
- Prof B. Bakker
- Prof A Scheurink
- Prof H Meijer
- Prof E van der Zee
- Prof U Eisel
  
- students

## Semmelweis University

- Prof C. Nyakas



Me? In 50  
years from  
now??

