

## Is bewegen goed voor het Brein ?

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## Inhoud

Bewegen – cognitie  
Mechanismen – neurogenese  
Welke soort beweging ?  
Bewegen – cognitie en pollutie

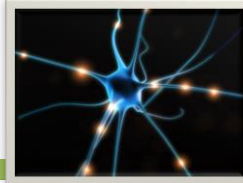
## Health benefits of exercise

- Obesity
- Diabetes
- Cardiovascular diseases
- Hypertension
- COPD
- Osteoarthritis
- Osteoporosis
- ....
- Cognition
- Parkinsons disease
- Alzheimers disease
- Stroke
- Depression
- Spinal cord injury
- ...



## Breïn

Ca 1300 gram  
1.10<sup>11</sup> cellen (honderd miljard)  
Elk 5000 à 10000 verbindingen  
→ 1.10<sup>15</sup> synapsen  
150.000km zenuwvezels

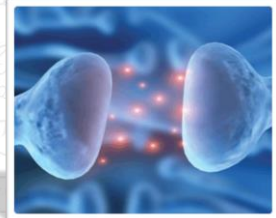


## The Brain needs energy !

Brain uses 130g Glucose/day  
@ rest → high % available Glucose  
Glycogen : stored in Astrocytes  
Lactate primary source for neurons

## Neurotransmitters

- Serotonin (5-HT)
- Dopamine
- Noradrenaline
- Acetylcholine
- GABA
- Glutamate
- ...



## Cognitie

De onderliggende operaties die door het centrale zenuwstelsel worden gebruikt om informatie te verwerken.

- Geheugen
- Aandacht
- Perceptie
- Problem solving
- Reactie tijd
- Snelheid uitvoering van handelingen en bewegingen



## Bewegen & school prestaties

### Aerobic Fitness

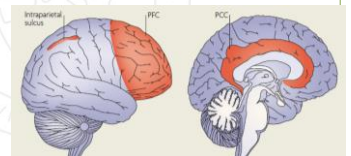
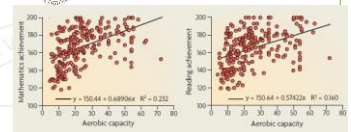
→ Positieve relatie met schoolprestaties

BMI → Negatieve relatie

### Relevante neurale netwerken

→ Prefrontal Cortex

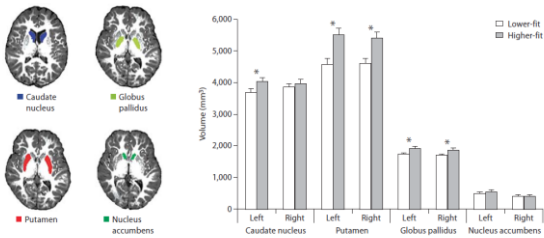
→ Parietal/Posterior Cingulate Cortex



Hillman et al 2008

## Basal ganglia volume pre-adolescent children

Duidelijke associatie met *aerobic fitness*



Chaddock et al 2010

## Physical Activity intervention

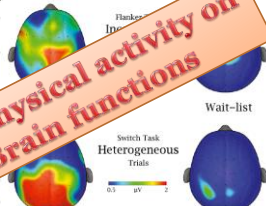
- N= 221
- 7-9yr
- 9 month intervention RCT (waiting list)
- PA : after school recreation & sports
- Measuring Brain Activity - ERP
- Accuracy & reaction time (executive control)



Hillman et al, 2014

## Results

- Fitness ↑
- Cognitive tests ↑
- Attention ↑
- Cognitive flexibility ↑



→ Positive effect of physical activity on cognition and brain functions for conditions that require great executive control and stability of Brain Functions

Hillman et al, 2014





## Cycling desks

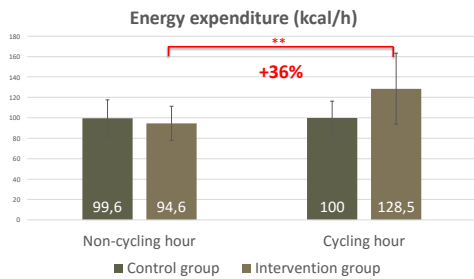
Sports Med (2014) 44:1261-1273  
DOI 10.1007/s40279-014-0202-x

SYSTEMATIC REVIEW

### Active Workstations to Fight Sedentary Behaviour

Tine Torbeyns · Stephen Bailey · Inge Bos · Romain Meeusen

## Results: EE



\*\* p ≤ 0.001



## Cycling desk : typing, cognitive performance, brain activity

- **23 office workers**
- 'normal' typing
- Typing @ cycling desk

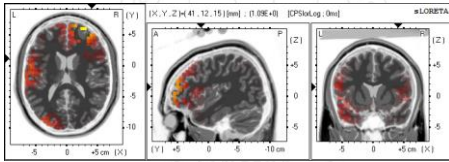
- Typing performance =
- Word recognition =
- Reaction time ↑
- Selective attention ↑
- Sustained attention ↑



Torbeyns, Meeusen et al submitted

## Brain activity

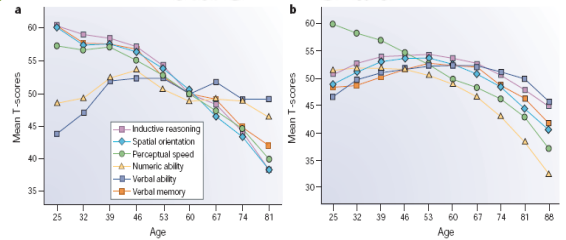
- Difference in brain activity between 'sitting' & 'cycling' desk
- More brain activity – more active Brains



Torbeyns, Meeusen et al submitted

## Cognitieve prestaties dalen met de leeftijd

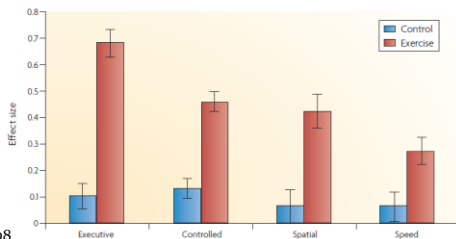
- Cross sectional
- Longitudinal



(Hedden & Gabrieli, 2004)

## Exercise effect : Cognitie ouderen

**Meta-analysis** : positieve invloed van inspanning op cognitie



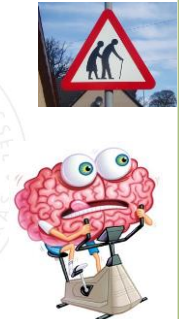
Hillman et al 2008

## Exercise, Brain Volume, Cognition

- 1 yr physical activity or stretching
- 2 x 60 ppn
- Gemidd lftd : 65.6 – 67.5yr
- 40 min wandelen; of stretching
- 2-3/Wk

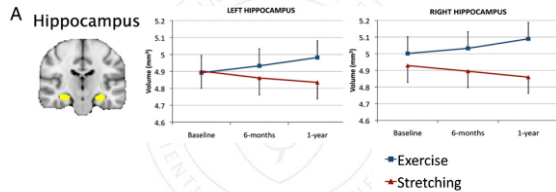
→ 'duur'-inspanning

Erickson et al 2011



## 1 yr aerobic exercise (RCT)

Hippocampus blijft 'plastisch' op latere lftd

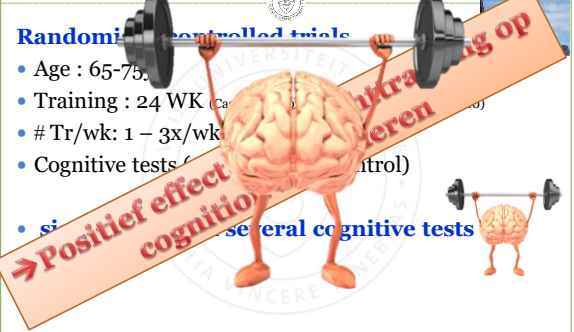


Erickson et al 2011

## Kracht Training & Cognitie

Randomized controlled trials

- Age : 65-75
- Training : 24 WK (Cardio)
- # Tr/wk: 1 - 3x/wk
- Cognitive tests (Control)
- si several cognitive tests



Cassilhas et al 2007; Liu-Ambrose et al 2010, 2012; Davis et al 2010

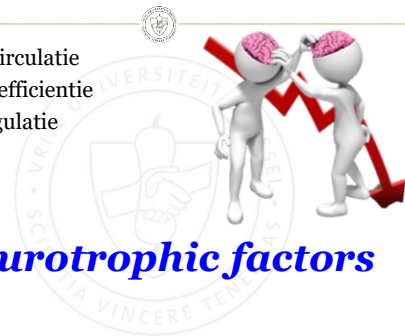
Fysieke inspanning beschermt tegen de cognitive vermindering en verbetert 'leren & onthouden'

Mechanismen ??

## Mechanismen ?

Cerebrale circulatie  
Neuronale efficiëntie  
Glucose regulatie  
...

**Neurotrophic factors**



## Exercise & Neurogenesis

Aanmaak van nieuwe hersencellen

→ is dit een Dogma ?

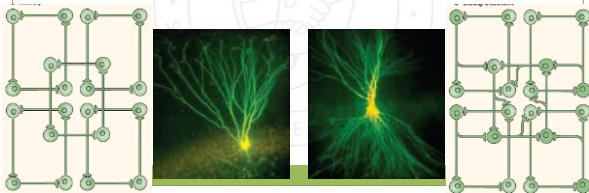
## Hersencellen sterven af ...

en worden  
Neurogene  
BESTAA  
→ Enkel tij  
Ramon y C  
fasen in c  
→ en die e  
Vanaf 1960  
Eind vorige



Elke dag worden er duizenden nieuwe cellen toegevoegd aan het brein van zoogdieren

→ procentueel weinig in vergelijking tot het totaal aantal hersencellen, maar dit geeft aan dat er een duidelijk structurele verandering mogelijk is



## Vermindering neuronen

acute & chronische stress;  
veroudering

- Glucocorticoiden, morfine, heroïne, alcohol, ...
- Pathologische omstandigheden : Alzheimers Disease, Parkinson's Disease, ...

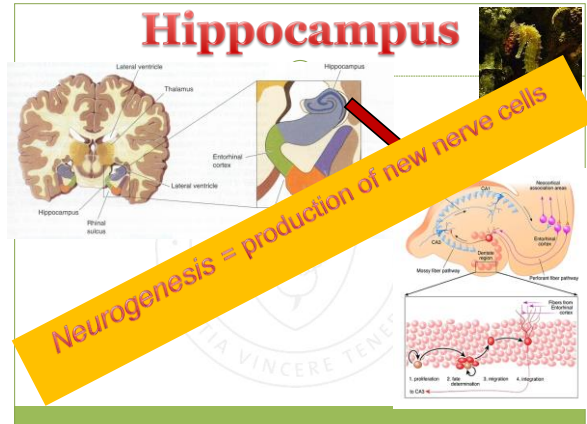


## Nieuwe hersencellen

Kunnen enkele dagen overleven, maar sommige blijven een leven lang bestaan

→ De hersencellen die op **volwassen leeftijd** worden aangemaakt hebben voornamelijk hun belang bij **leerprocessen en geheugen**.

## Hippocampus



## Brain Derived Neurotrophic Factor (BDNF)

Is de meest wijd verspreide **neurotrophine** in de hersenen

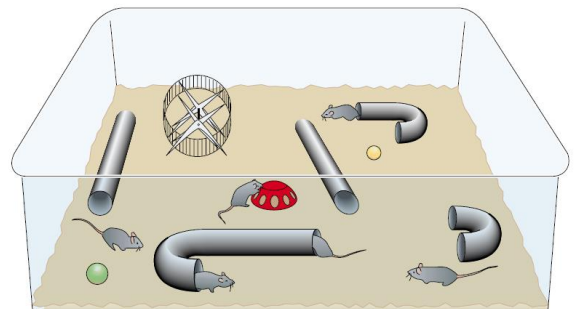
Het bevordert de **groei** en het **onderhoud** van verschillende neuronale systemen

Is betrokken bij **leerprocessen**

BDNF expressie is verminderd bij patiënten met **Alzheimer's disease**

## Enriched environment = verrijkte omgeving

speeltjes, loopwielletjes, grotere kooien, ...





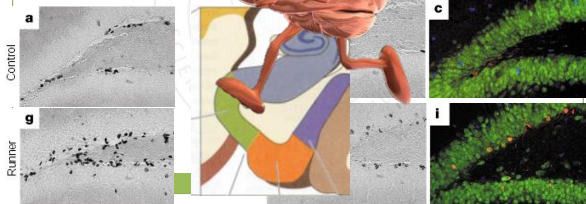
## Voluntary running

(Van Praag et al Nat neurosci 1999)

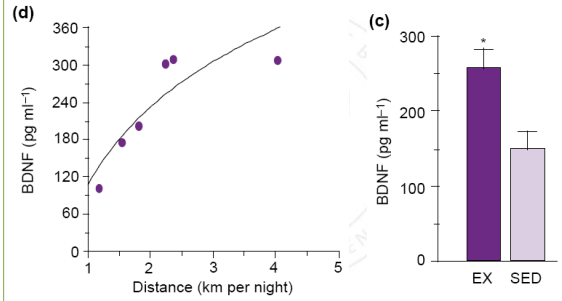


Zal het overleven van nieuwe neuronen bevorderen = cell proliferatie  
Of neurogenese (van nieuwe cellen) stimuleren

One day Runners 4 weeks



## BDNF expression correlates with the distance run per night



(Cotman & Berchtold TINS 2002)

## Wat bij de mens ?

- Verschillende studies hebben in serum verhoogde BDNF concentraties gevonden
- Is dit een indicator van wat er in de hersenen gebeurt ?



Neuroscience Letters 484 (2010) 150–154

Contents lists available at ScienceDirect  
Neuroscience Letters

Influence of citalopram and environmental temperature on exercise-induced changes in BDNF  
Maaike Goekint<sup>a,b</sup>, Bart Roelands<sup>a,b</sup>, Elsa Heyman<sup>a,c</sup>, Rose Njemini<sup>d</sup>, Romain Meeusen<sup>a,\*</sup>

RESEARCH REVIEW ARTICLE

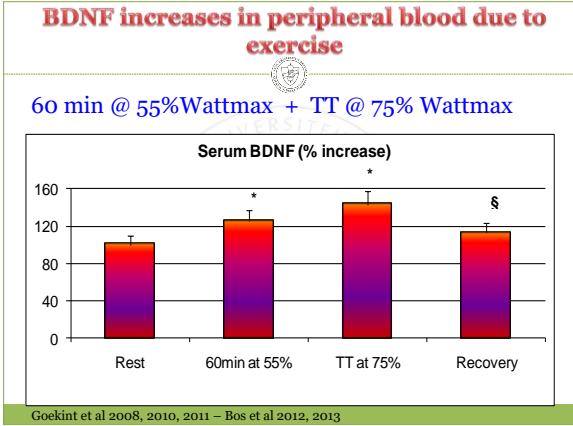
Neuroplasticity – Exercise-Induced Response of Peripheral Brain-Derived Neurotrophic Factor  
A Systematic Review of Experimental Studies in Human Subjects

neurotrophic factor ?  
Kristol Knaepen<sup>1</sup>, Maaike Goekint<sup>1,2</sup>, Elsa Marie Heyman<sup>1,3</sup> and Romain Meeusen<sup>1</sup>  
Maaike Goekint<sup>a,b</sup>, Bart Roelands<sup>a,b</sup>, Kevin De Pauw<sup>a</sup>, Kristel Knaepen<sup>a</sup>, Inge Bos<sup>a,d</sup>, Romain Meeusen<sup>a,\*</sup>

Psychobiology and Behavioral Strategies

No Influence of Noradrenaline Manipulation on Acute Exercise-Induced Increase of Brain-Derived Neurotrophic Factor

Goekint et al 2008, 2010, 2011  
MAAIKE GOEKINT<sup>1,2</sup>, ELSA HEYMAN<sup>1,3</sup>, BART ROELANDS<sup>1</sup>, ROSE NIEMINI<sup>4,5</sup>, IVAN BAUTMANS<sup>1,4,5</sup>, TONY METS<sup>1,5</sup>, ROMAIN MEEUSEN<sup>1</sup>



### BDNF & Strength training

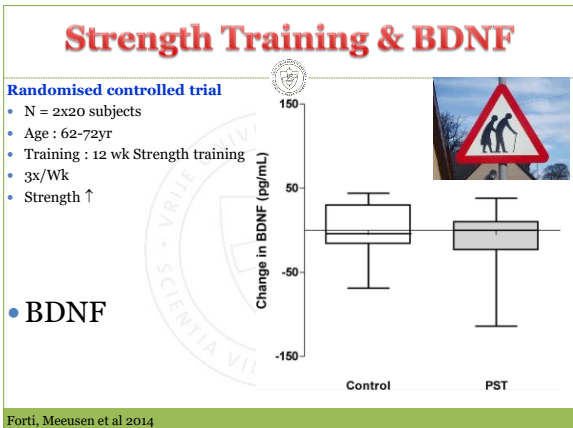
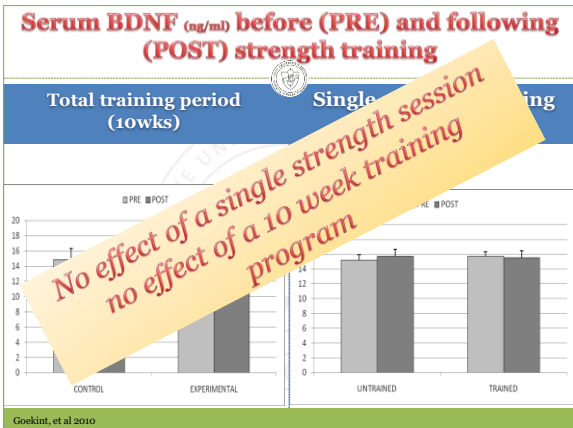
ORIGINAL ARTICLE

#### Strength training does not influence serum brain-derived neurotrophic factor

Maike Goekint · Kevin De Pauw · Bart Roelands · Rose Njemini · Ivan Bautmans · Tony Mets · Romain Meeusen

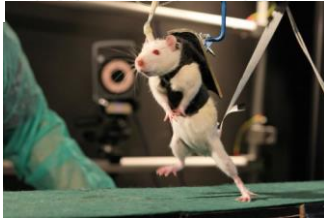
Training program during 10 weeks, with a frequency of 3 training sessions per week

Goekint, et al 2010



## Exercise, cognition & the Brain, ...

### Mechanisms ?



## Cognition & Strength training

Eur J Appl Physiol (2010) 110:285–293  
DOI 10.1007/s00421-010-1461-3

ORIGINAL ARTICLE

Strength training does not influence serum brain-derived neurotrophic factor

Maijke Gokint · Kevin De Pauw · Bart Roelands · Rose Niemi · Ivan Raaijmakers · Tony Meys

**Training study in young & old adults → no increase in peripheral BDNF**  
**But**  
**Improvement in cognition**  
 ?? Mechanisms → animal study

Gokint, et al 2010, Forti et al 2014, Casilhas et al 2007, Liu-Ambrose et al 2010, 2012, Davis et al 2010

## Endurance – Resistance

### 8 wks training



Casilhas, Meeusen, et al. *Neuroscience* 2012

## Learning & Memory : Aerobic & Resistance Training

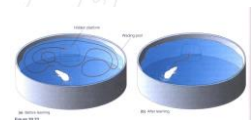
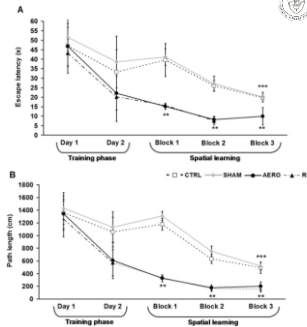
8 wks training

Rats

Water Maze

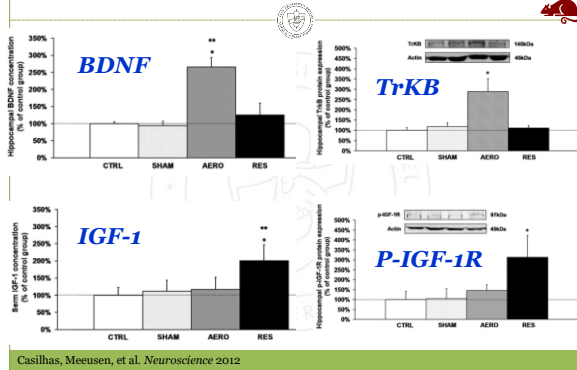
Brain mechanisms

Cognition improved



Casilhas, Meeusen, et al. *Neuroscience* 2012

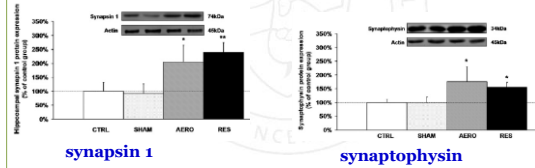
## Mechanisms



## Mechanisms

Aerobic and Resistance training for 8 weeks increases learning and spatial memory in a similar manner.

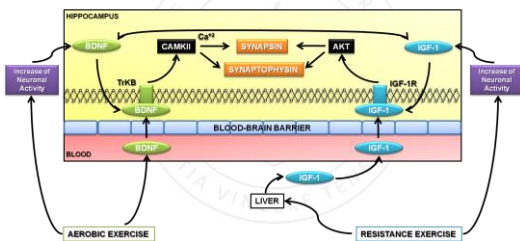
- **Aerobic exercise** modulates BDNF/TrkB and -CaMKII
- **Resistance training** acts on the CNS via IGF-1/IGF-1R and AKT pathway
- Both induced the expression of **synapsin 1** and **synaptophysin**, which might contribute to improved spatial learning and memory.



## Aerobic or Resistance Training ?

Aerobic and Resistance training for 8 weeks

→ **Synaptic Plasticity**



## Dus, ...

- Inspanning is goed voor het Brein !
- Inspanning bevordert de cognitie zowel op jonge als op oudere leeftijd
- Duidelijk bewijs dat men nieuwe neuronen aanmaakt tijdens inspanning
- BDNF speelt hierbij een belangrijke rol
- Maar er is een limiet
- ... is dit altijd het geval ?

## Exercise & Pollution



## Particular Matter (PM)

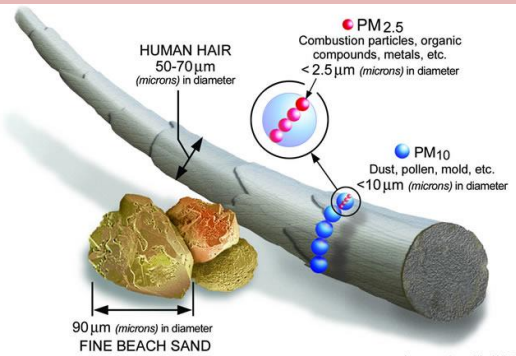


Image courtesy of the U.S. EPA

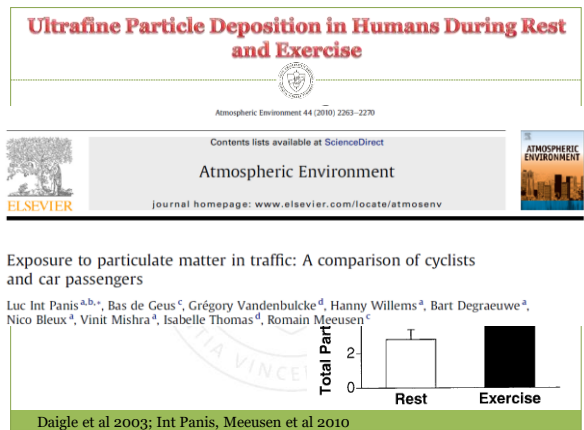
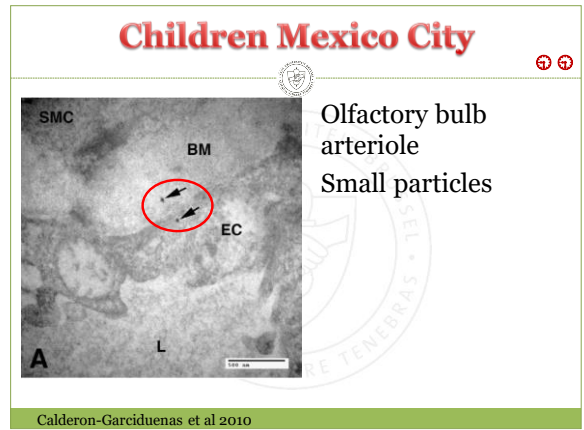
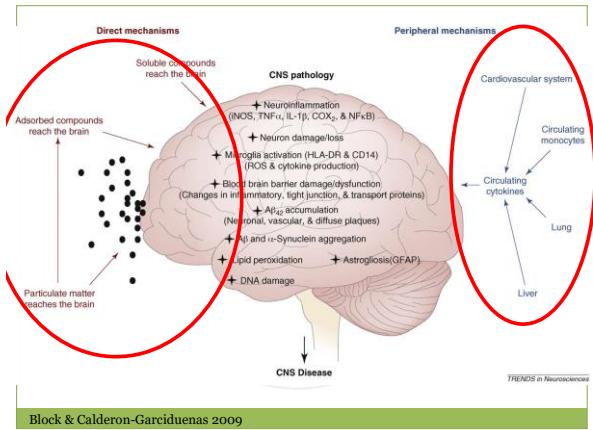
## Pollution

### Long-term exposure to traffic-related air pollution :

- Cardiovascular & pulmonary disease
- **Mild Cognitive Impairment in Elderly** (Ranft et al 2009) → distance < 50m from busy traffic road
- PM10 predicts cognitive function (Chen 2009)

## Pollution & the Brain





# Exercise, Pollution & BDNF

Neuroscience Letters 580 (2015) 110–112  
Contents lists available at ScienceDirect  
Neuroscience Letters  
ELSEVIER  
journal homepage: www.elsevier.com/locate/neulet

No exercise-induced increase in serum BDNF after cycling near a major traffic road

I. Bos<sup>a,b</sup>, I. Jacobs<sup>a</sup>, T.S. Nawrot<sup>a,c</sup>, B. de Geus<sup>a</sup>, R. Toes<sup>a</sup>, L. Ist Paris<sup>a,d</sup>, B. Degrauwe<sup>a</sup>, R. Meerssens<sup>a,e</sup>

<sup>a</sup> Institute of Environmental Health 2010, <sup>b</sup> Institute of Environmental Health 2010, <sup>c</sup> Institute of Environmental Health 2010, <sup>d</sup> Institute of Environmental Health 2010, <sup>e</sup> Institute of Environmental Health 2010

ENVIRONMENTAL HEALTH

**RESEARCH** Open Access

Subclinical responses in healthy cyclists briefly exposed to traffic-related air pollution: an intervention study

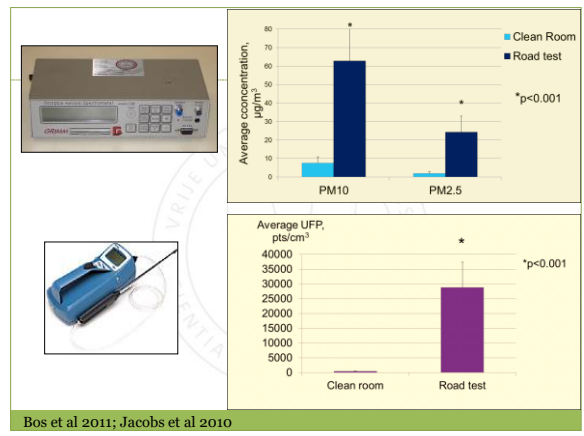
Lutz Jaeger<sup>1</sup>, Sara H. Nauman<sup>2</sup>, Sas de Waard<sup>3</sup>, Boran Meusen<sup>3</sup>, Bart Degrauwe<sup>4</sup>, Alfred Bernard<sup>5</sup>, Muhammad Sighi<sup>6</sup>, Boris Nemery<sup>7</sup>, Luc H. Van<sup>8</sup>

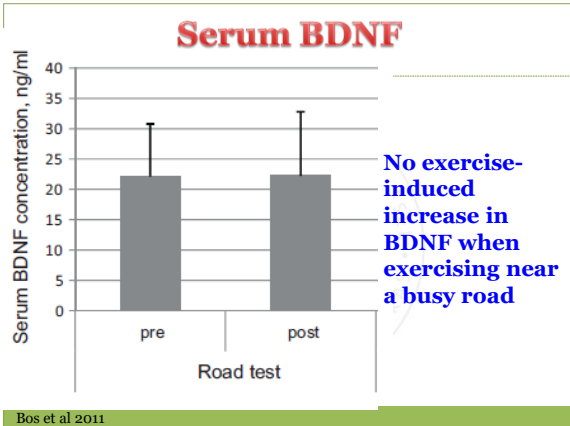
## ANTWERP RING ↔ CLEAN ROOM



**Clean Room**

Bos et al 2011; Jacobs et al 2010





### Start - to - Run campaign

10 weeks start a Run (SoR) cycle

#### Subclinical Effects of Aerobic Training in Urban Environment

INGE BOS<sup>1,2</sup>, PATRICK DE BOEVER<sup>1,3</sup>, JEF VANPARIJS<sup>2</sup>, NATHALIE PATTYN<sup>2,4</sup>, LUC INT PANIS<sup>1,5</sup>, and ROMAIN MEEUSEN<sup>2</sup>

<sup>1</sup>Environmental Risk and Health, Flemish Institute for Technological Research (VITO), Mol, BELGIUM; <sup>2</sup>Human Physiology and Sports Medicine, Vrije Universiteit Brussel, Brussels, BELGIUM; <sup>3</sup>Centre for Environmental Sciences (CMK), Hasselt University, Diepenbeek, BELGIUM; <sup>4</sup>Vital Signs and Performance Monitoring (VSPM) Research Unit, Royal Military Academy, Brussels, BELGIUM; and <sup>5</sup>Transportation Research Institute (IMOB), Hasselt University, Diepenbeek, BELGIUM

PTRAK *Med. Sci. Sports Exerc.*, Vol. 45, No. 3, pp. 439-447, 2013.

Bos et al, 2013

### Biological outcome measures

Fitness level: cooper test

Cognitive performances

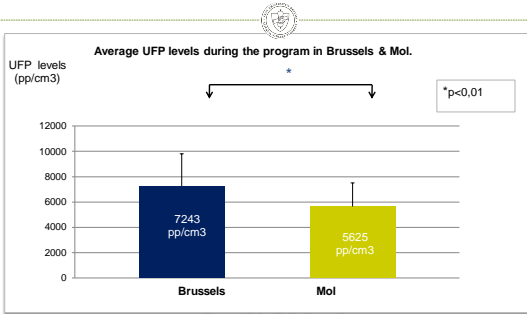
Blood leukocyte counts  
Serum BDNF level  
whole genome gene expression

eNO measurements

Bos et al, 2013



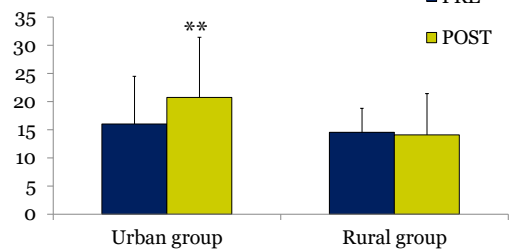
## Results: UFP levels



Bos et al., 2013

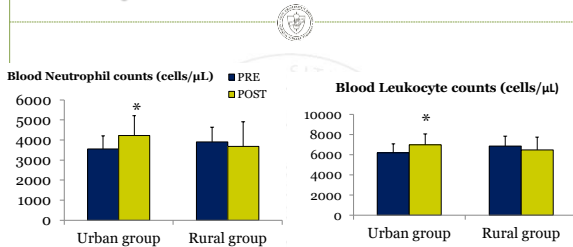
## Respiratory inflammation

Exhaled NO (ppb)



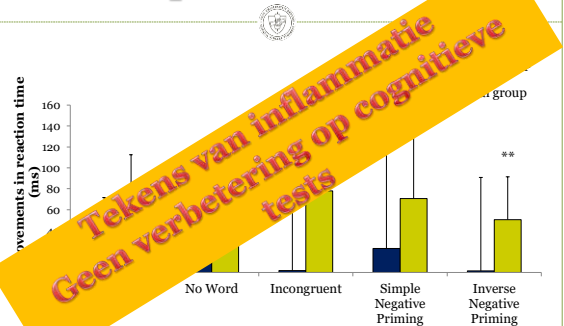
Bos et al., 2013

## Systemic inflammation



Bos et al., 2013

## Stroop Color Word Test



Bos et al., 2013

## Dus,

- Inspanning in een vervuilde omgeving onderdrukt de perifere expressie van BDNF
- Training (10wkn) :
  - Verbeterd de conditie 😊
  - Systemische en Respiratoire inflammatie (pollutie)
  - Geen verbetering op cognitieve tests
- → Mechanismen ??

## Mechanisms

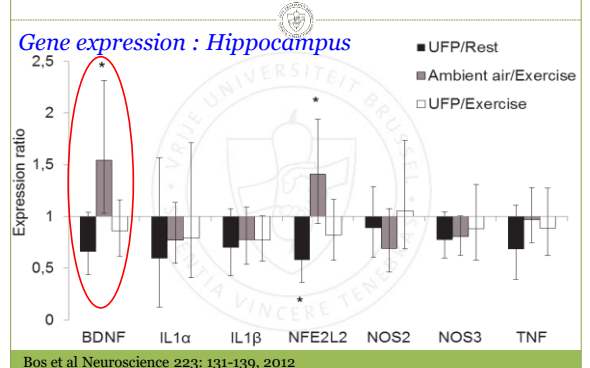


### Gene expression changes in rat brain tissue after acute exposure to ultrafine particles during exercise and rest

■ Total UFP-emission:  $10^6$  -  $10^7$  parts/cm<sup>3</sup>  
 ■ CO<sub>2</sub> conc: 1500ppm (< TWA)

Bos et al 2012

### No up-regulation BDNF in hippocampus rats exposed to UFP during exercise



# Muizen in de Craeybecks Tunnel

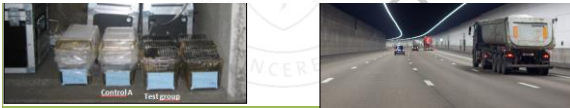
Inhalation Toxicology, 2012; 24(10): 676-686  
 © 2012 Informa Healthcare USA, Inc.  
 ISSN 0895-8378 print/ISSN 1191-7691 online  
 DOI: 10.1109/08958378.2012.714004



RESEARCH ARTICLE

## Changed gene expression in brains of mice exposed to traffic in a highway tunnel

Inge Bos<sup>1,2</sup>, Patrick De Boever<sup>1,3</sup>, Jan Emmerechts<sup>4,5</sup>, Jurgen Buekers<sup>1</sup>, Jeroen Vanoirbeek<sup>5</sup>, Romain Meeusen<sup>2</sup>, Martine Van Poppel<sup>1</sup>, Benoit Nemery<sup>5</sup>, Tim Nawrot<sup>1,5</sup>, and Luc Int Panis<sup>1,6</sup>



Bos et al MSSE, 2012

# Results

- Air quality: Tunnel vs. Traffic control Centre
  - The average concentration of measured pollutants

Pollutant	Tunnel	Traffic Control centre	Factorial difference
PM2.5	55.1 µg/m <sup>3</sup>	29.4 µg/m <sup>3</sup> * <sup>1</sup>	±2
EC	13.9 µg/m <sup>3</sup>	2.04 µg/m <sup>3</sup> * <sup>2</sup>	±7
OC	23.1 µg/m <sup>3</sup>		
UFP	150 000 particles/cm <sup>3</sup>	6 000 particles/cm <sup>3</sup>	±25

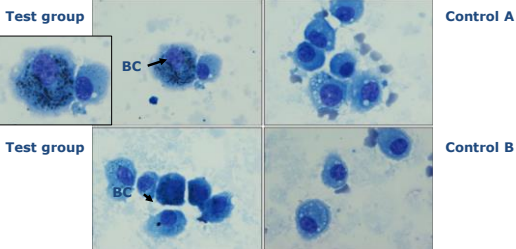
\*<sup>1</sup> Monitoring station in City of Antwerp

\*<sup>2</sup> Reported in Flanders (Vercauteren, 2011)

Bos et al MSSE, 2012

# Results

- Carbon loading of alveolar macrophages

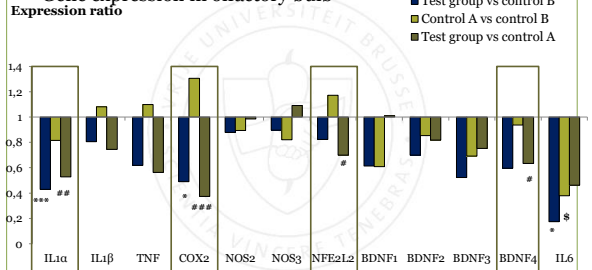


- Physiological signs of inflammation

Bos et al MSSE, 2012

# Results

- Gene expression in olfactory bulb



Bos et al MSSE, 2012

