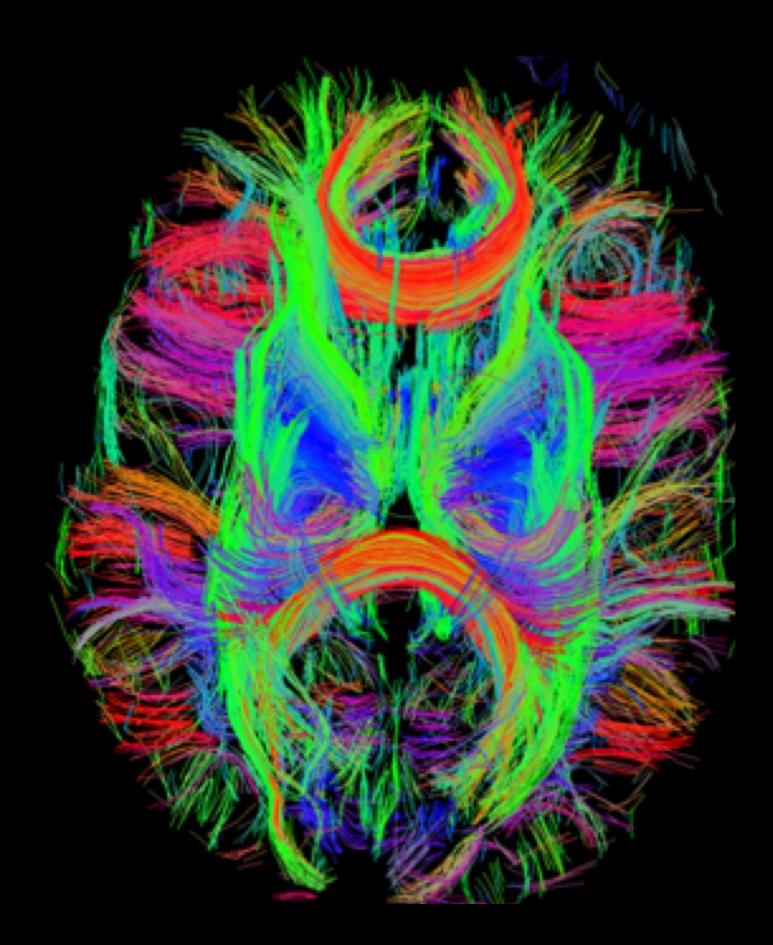
# Functional Connectivity and the Ageing Brain

- 5 november 2020
- European Association of Geriatric Psychiatry (EAGP)

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### Program

- Functional connectivity: what is it?
- Resting-state: why?
- Changes in functional connectivity during ageing
- Amyloid-beta and functional connectivity



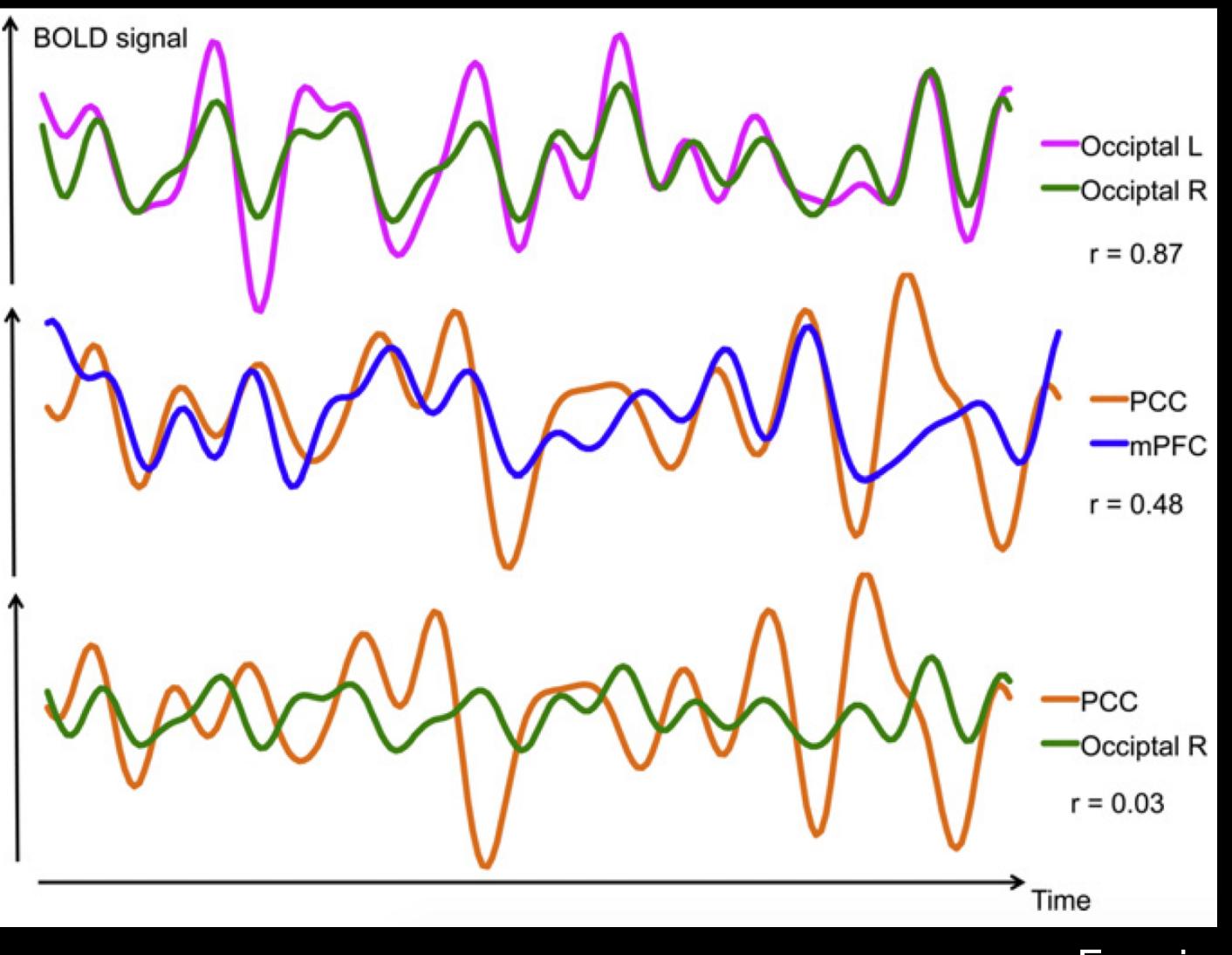
### Structural Connectivity



www.humanconnectome.org www.humanconnectomeproject.org



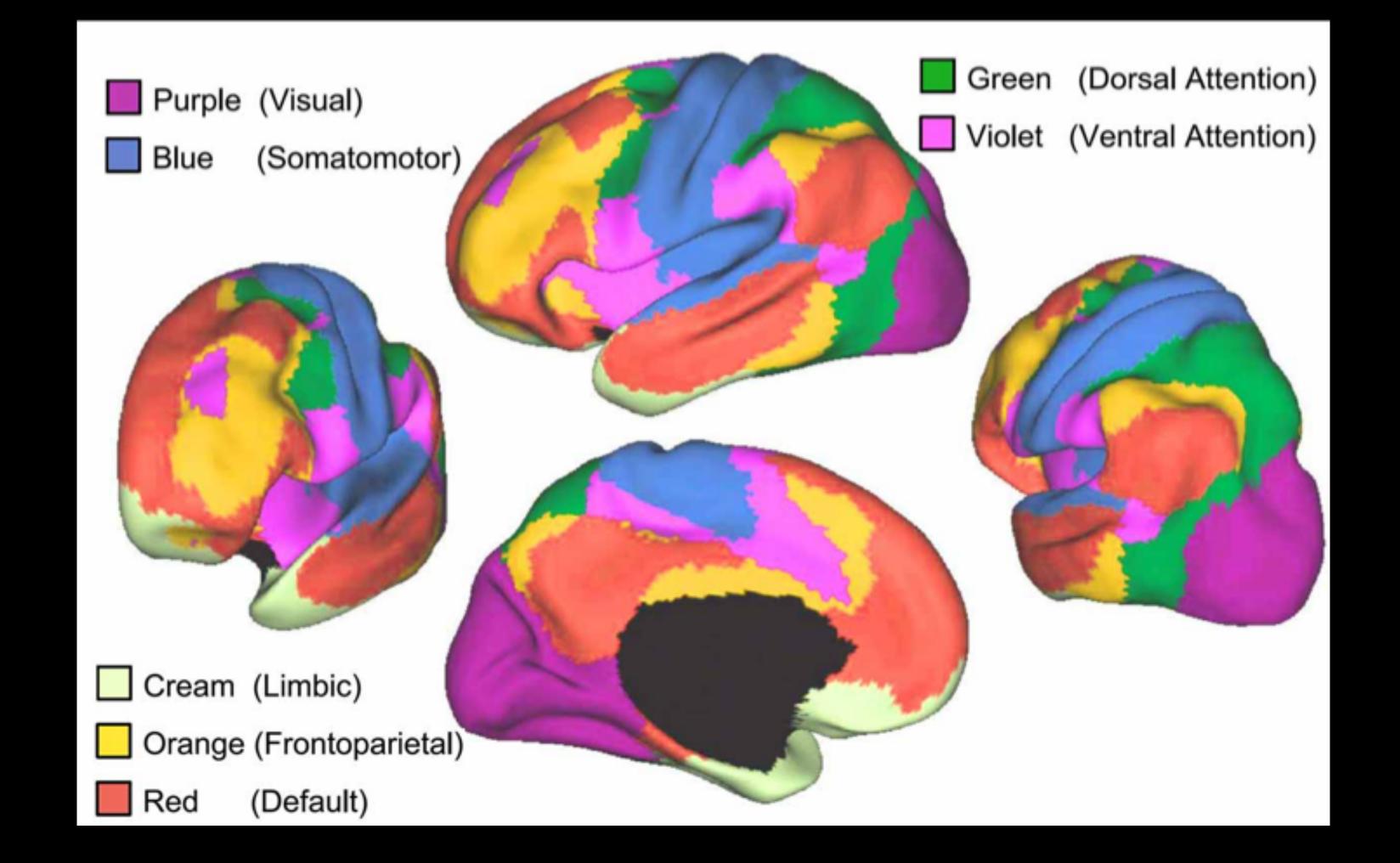
### **Functional Connectivity**



Ferreira & Busatto et al., 2013



### **Networks**





## Resting-state

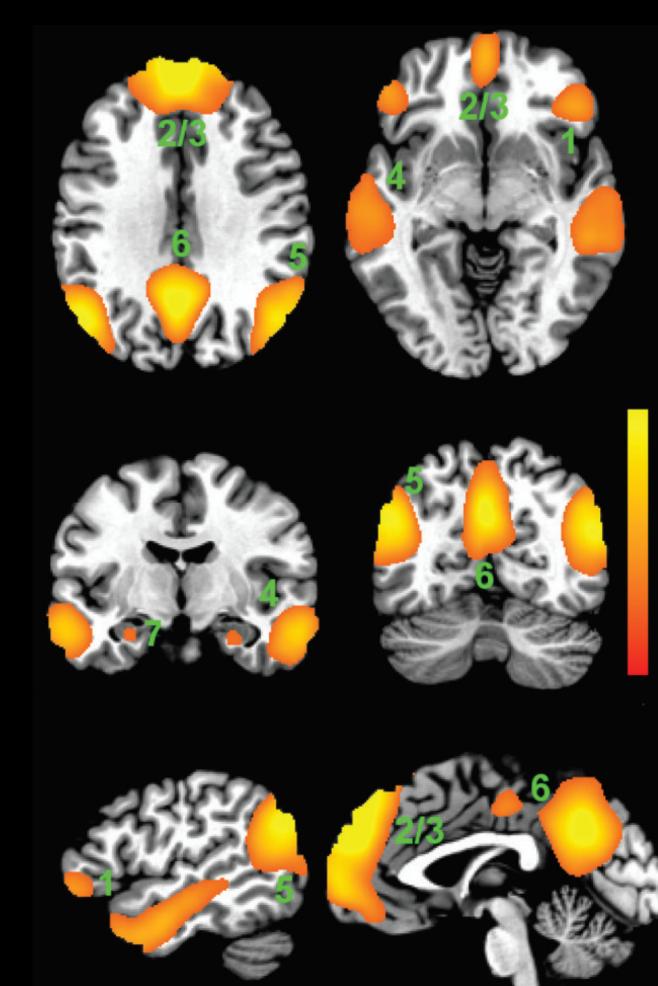


Photo by Zachary Nelson on Unsplash

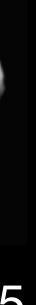


### Default Mode Network

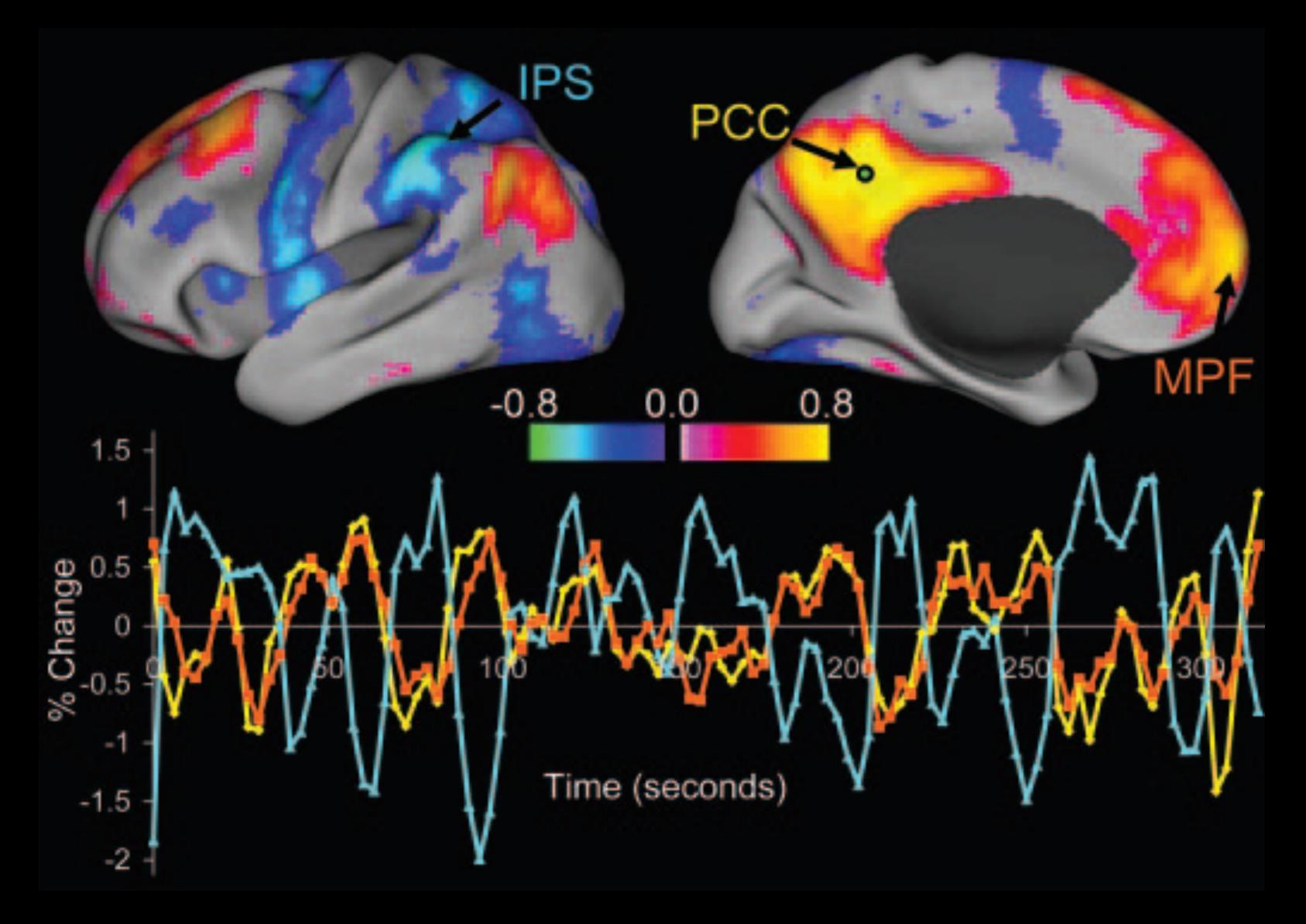
'We posit that when an individual is awake and alert and yet not actively engaged in an attention-demanding task, a **default** state of brain activity exists that involves, among other areas, the MPFC and the posterior cingulate and precuneus. Information broadly arising in the external and internal milieu is gathered and evaluated. When focused attention is required, particularly if this activity is novel, activity within these areas may be attenuated.' (Raichle et al., 2001)



Raichle, 2015



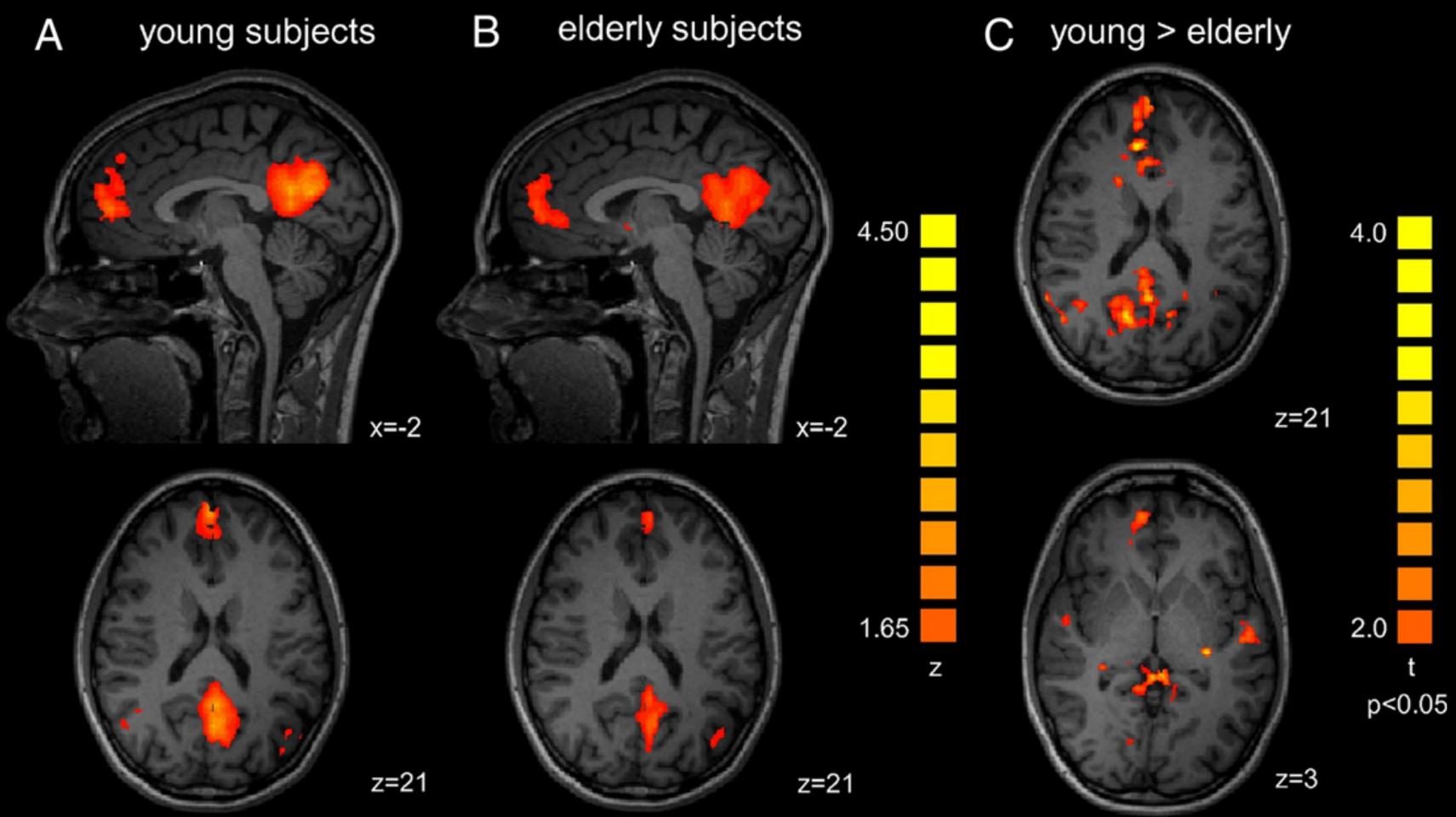
### **Correlations and Anticorrelations**

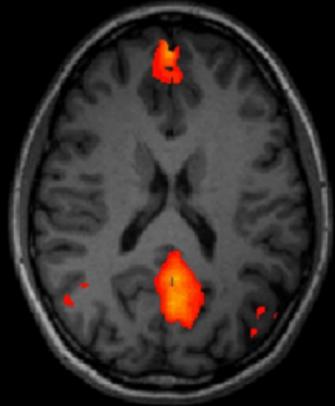


### Fox et al., 2005



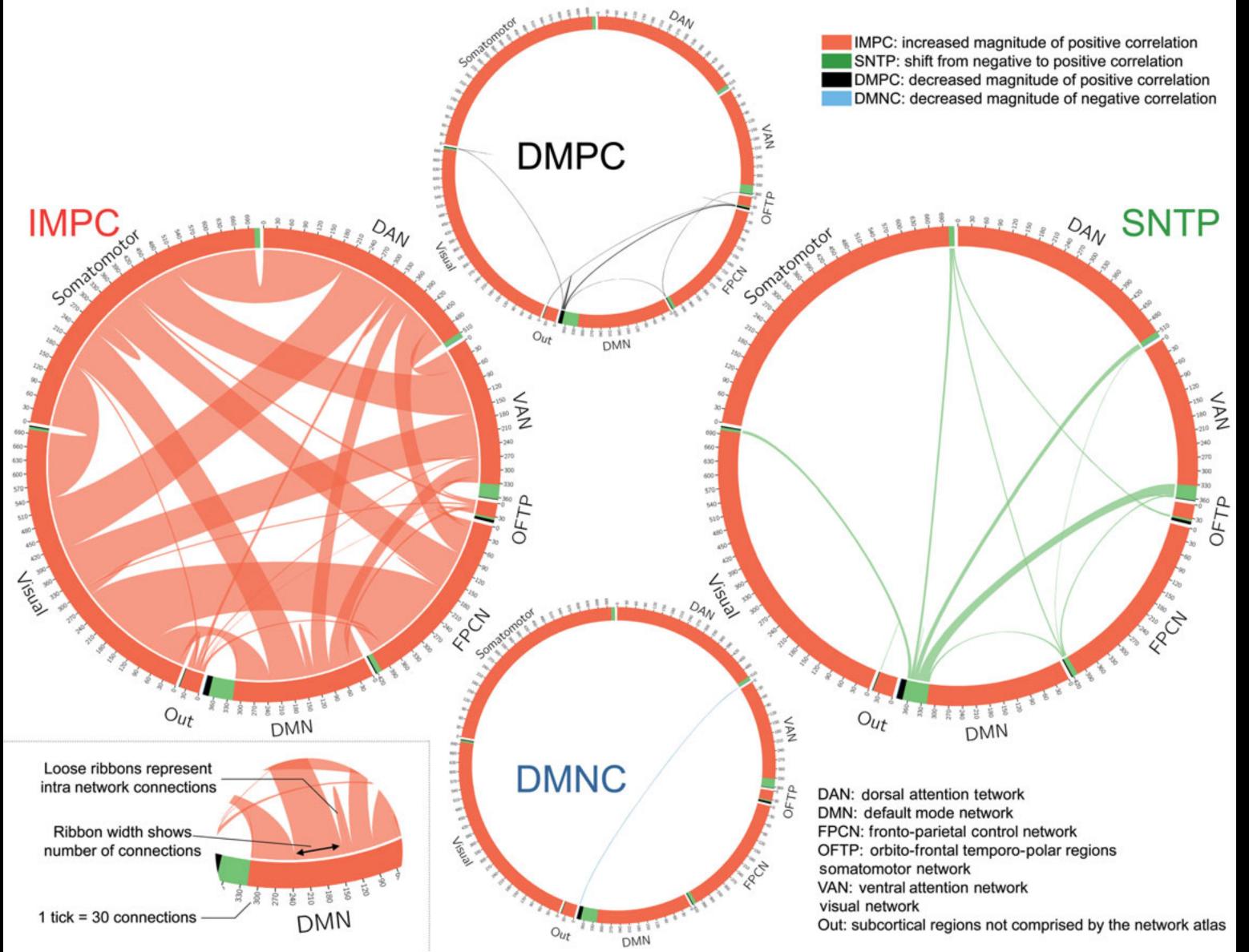
## DMN and Ageing



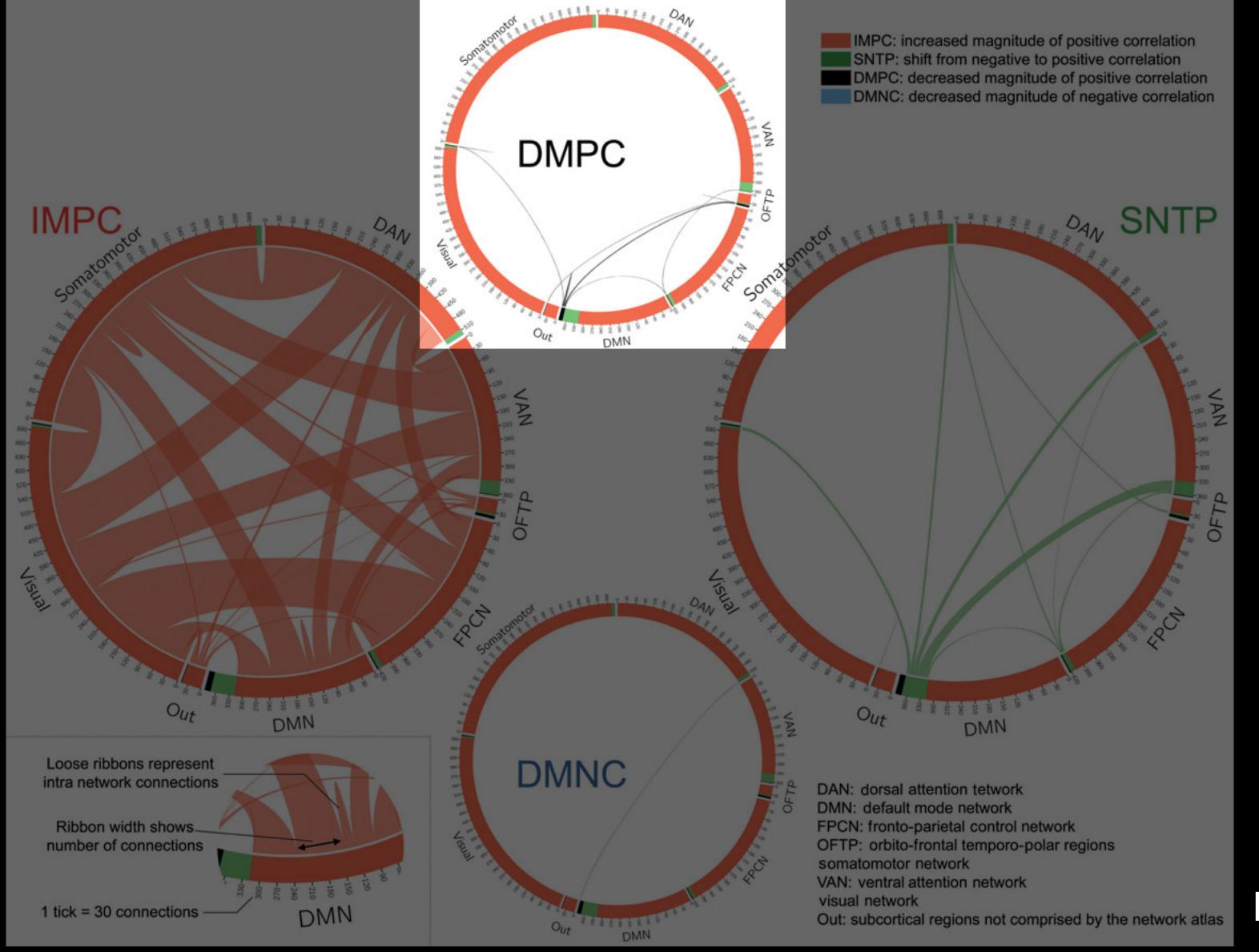


Koch, et al 2010

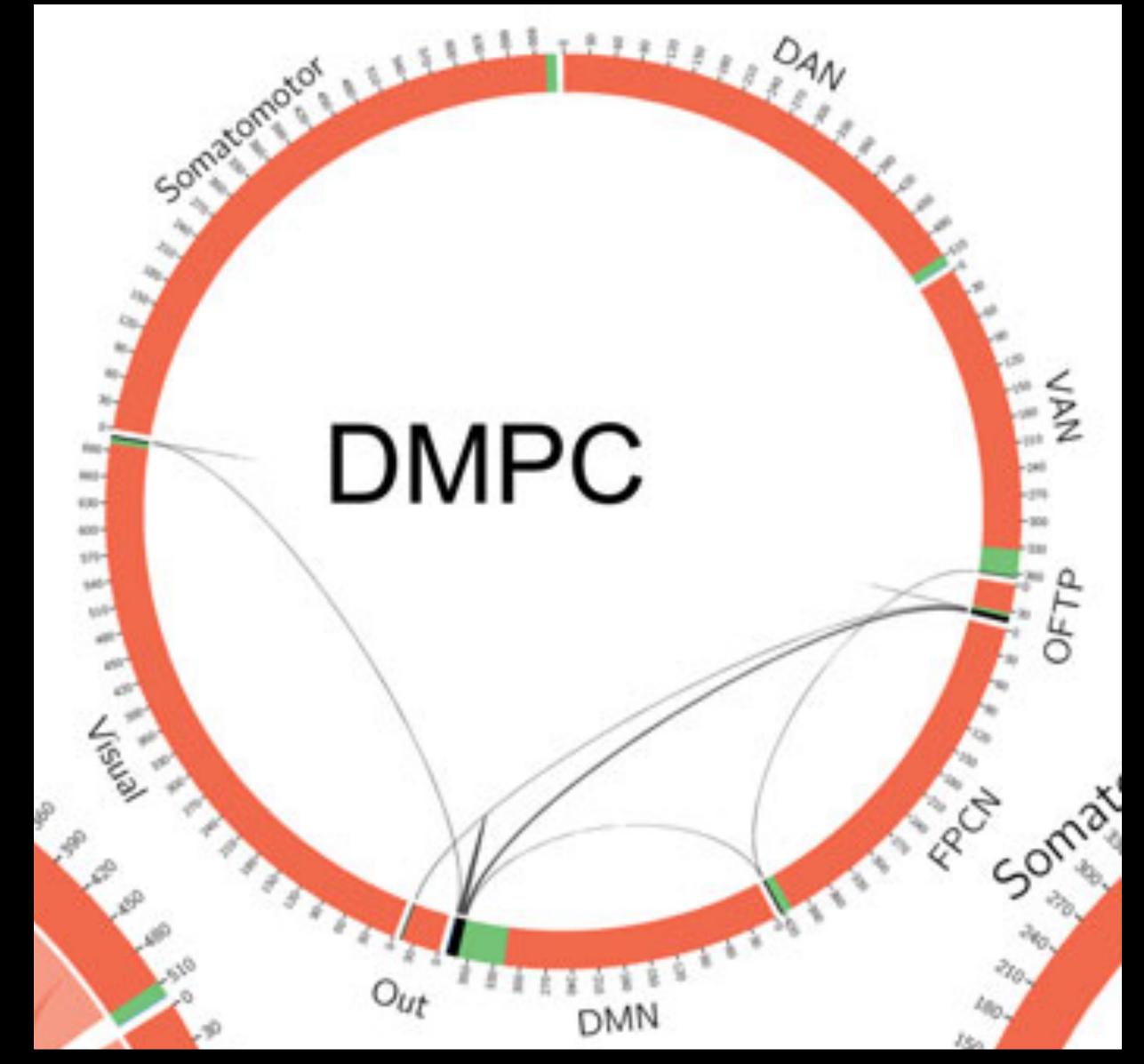




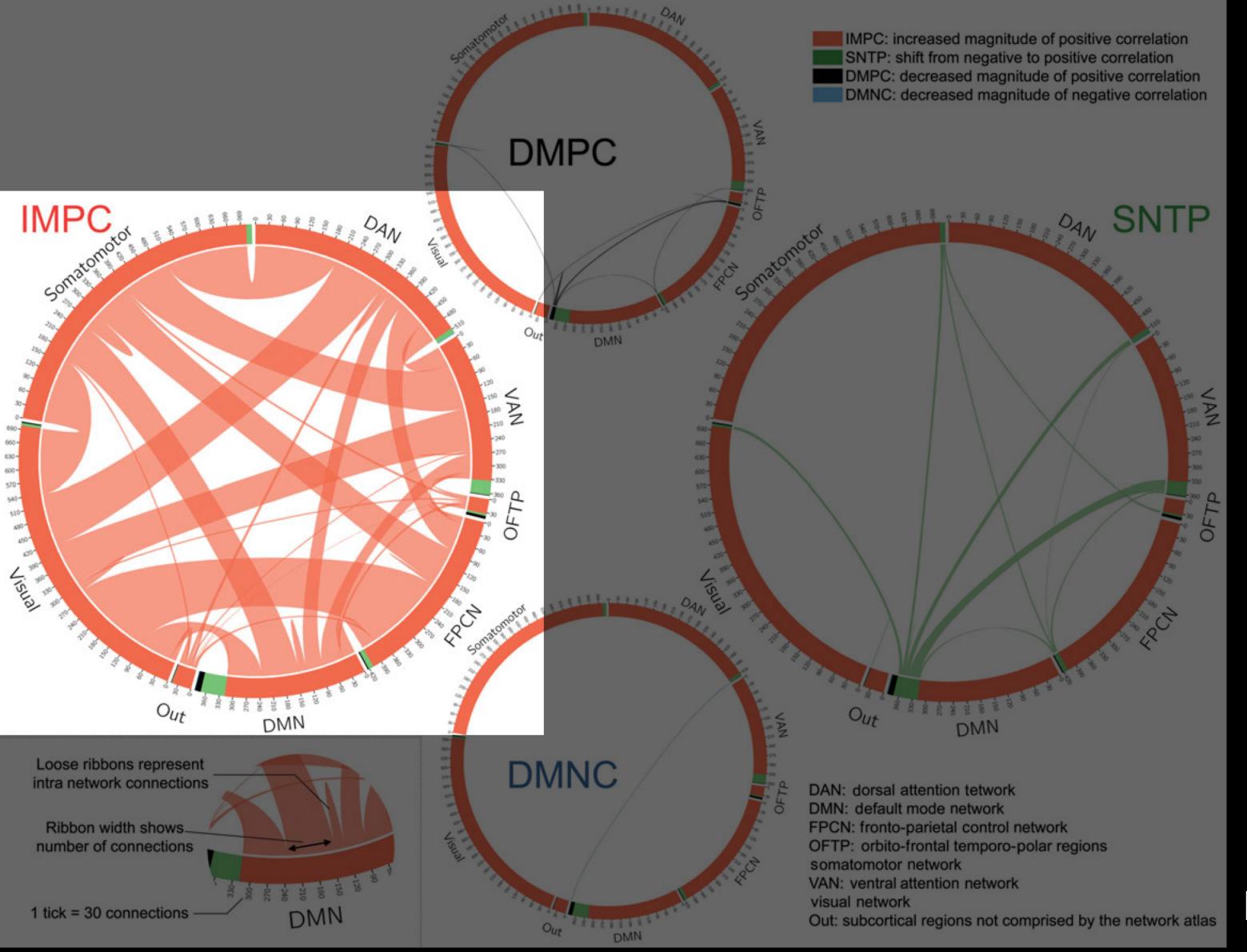




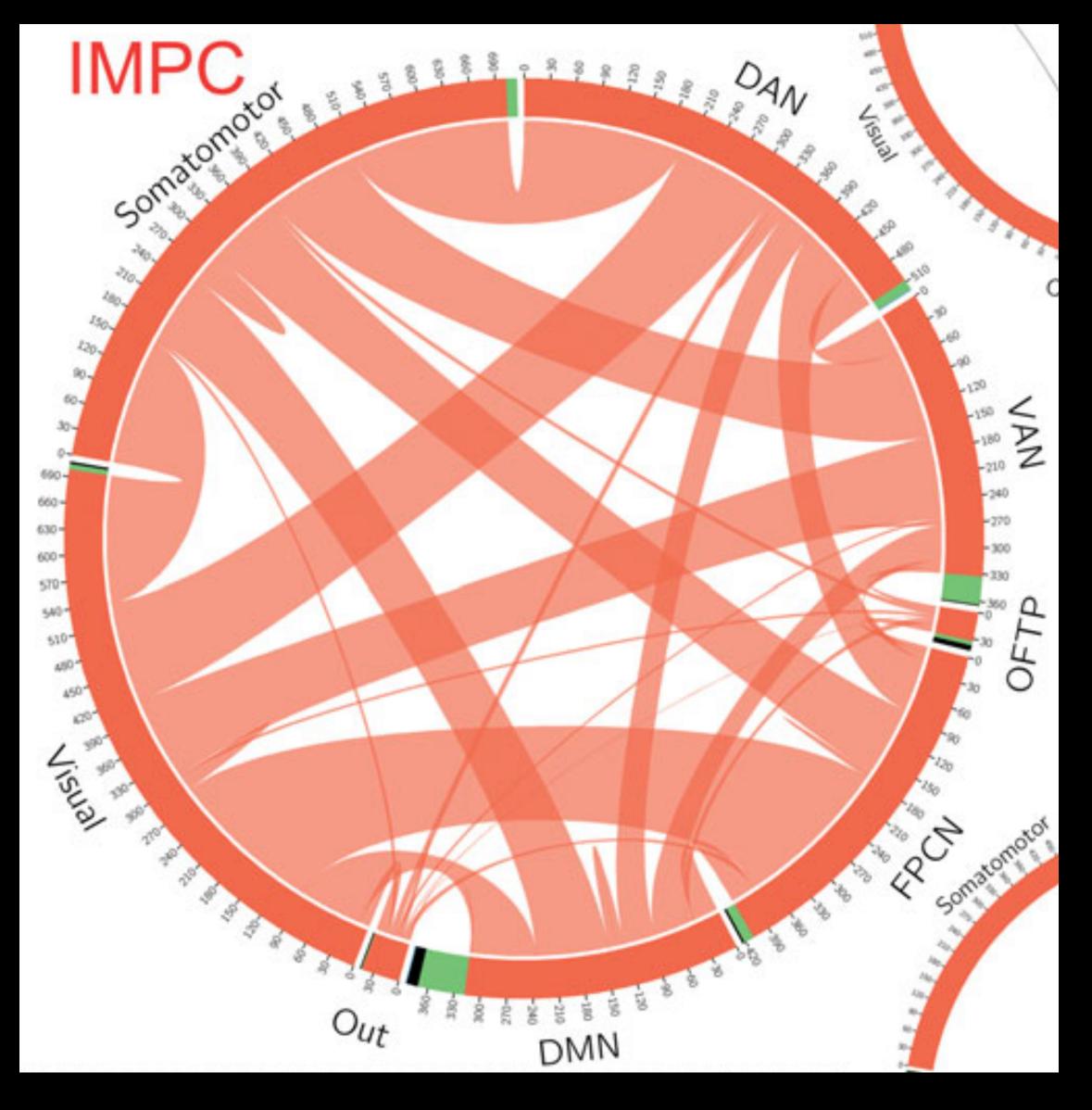




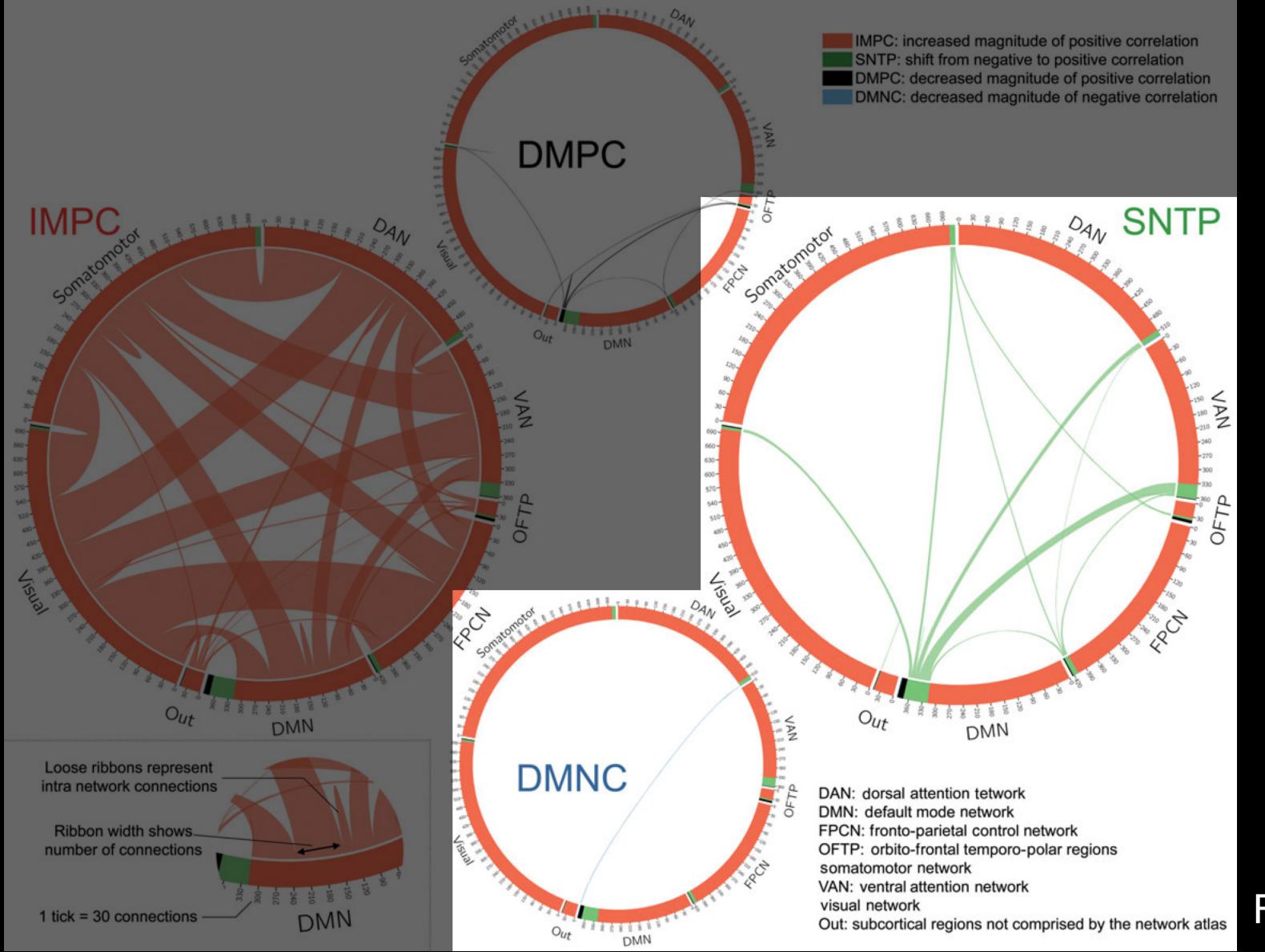




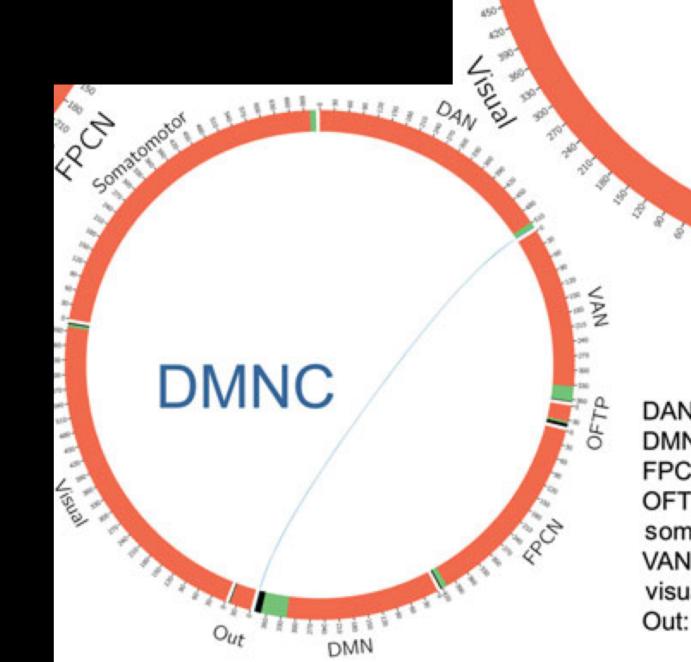




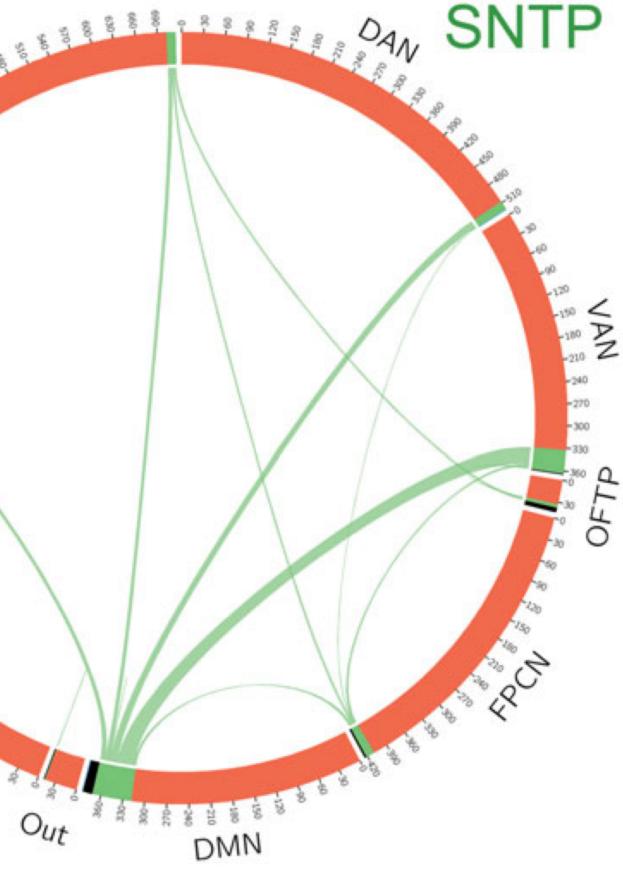








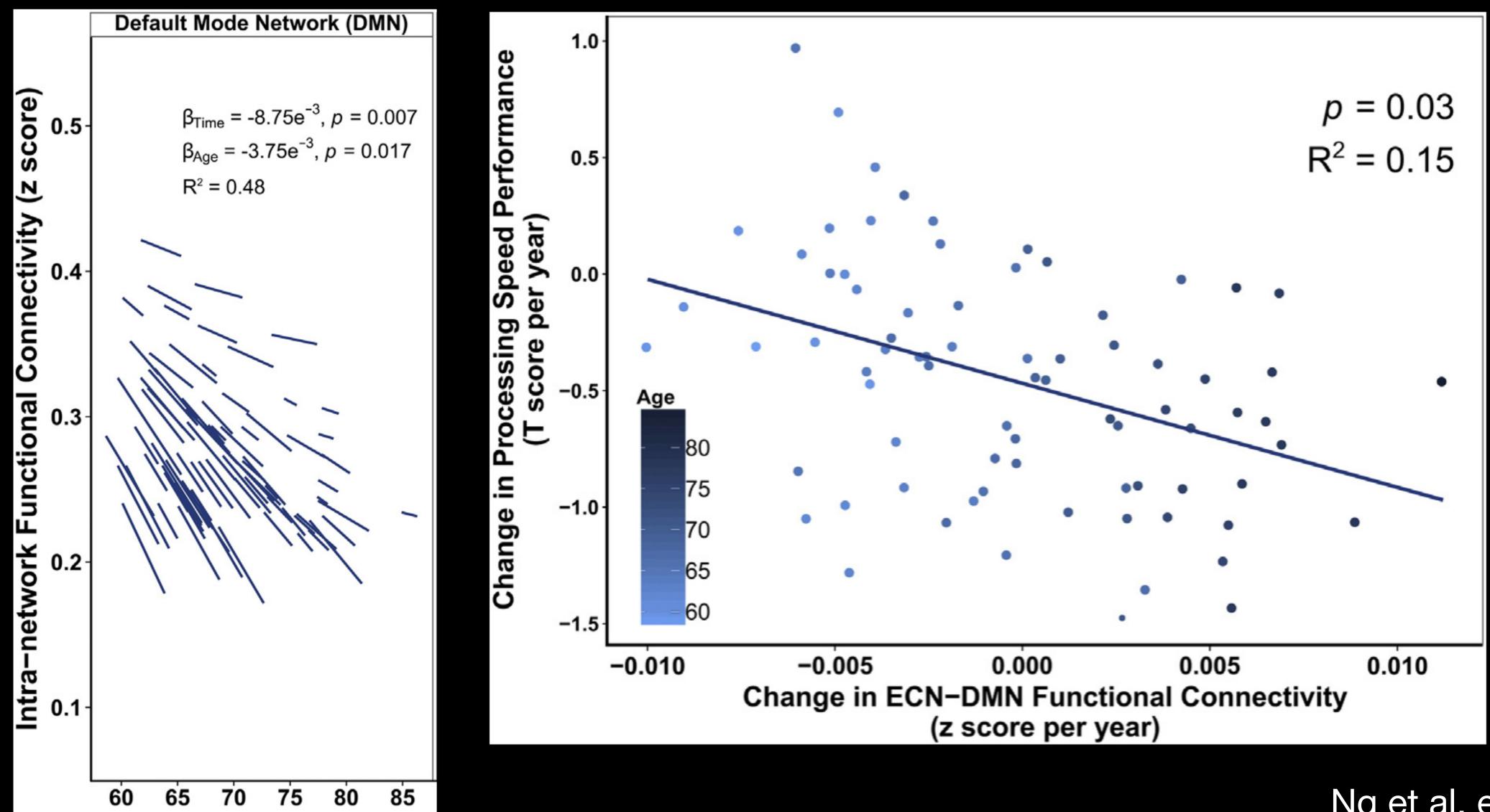
660-



- DAN: dorsal attention tetwork DMN: default mode network FPCN: fronto-parietal control network OFTP: orbito-frontal temporo-polar regions
- somatomotor network
- VAN: ventral attention network
- visual network
- Out: subcortical regions not comprised by the network atlas



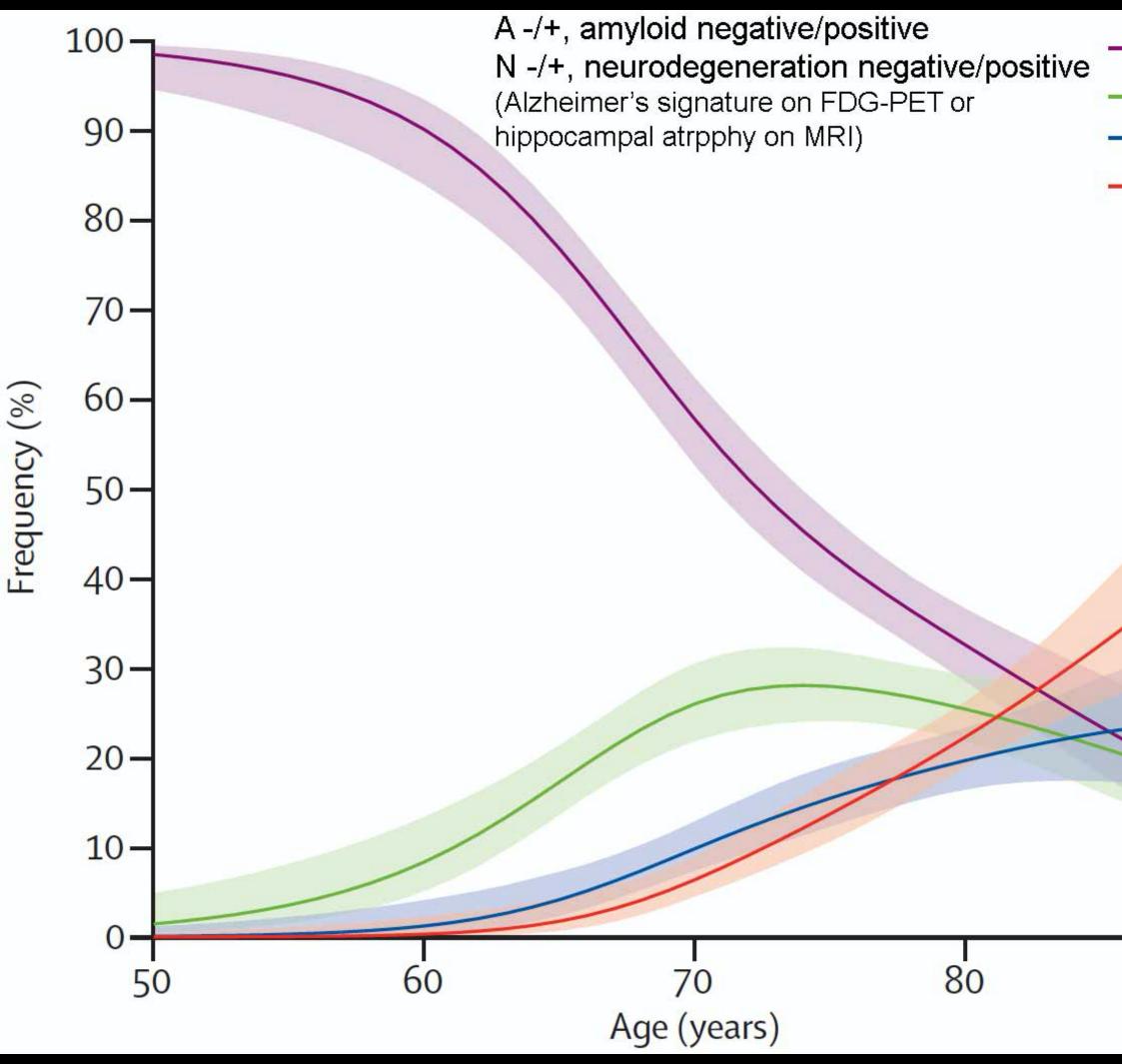
# Longitudinal study



Ng et al, et al 2016



## **Amyloid-beta and Ageing**



- A<sup>-</sup>N<sup>-</sup>  $--- A^+N^-$ — A<sup>-</sup>N<sup>+</sup> — A<sup>+</sup>N<sup>+</sup>

90

"By age 89 years, about 83% of individuals with normal cognitive function had levels of amyloidosis, neurodegeneration, or both that were similar to those seen in mild Alzheimer's disease [...] and people are able to maintain normal cognitive function despite these abnormalities. Typical cognitive ageing, defined as remaining free of dementia, is therefore most often characterized by the presence rather than the absence of these imaging abnormalities."

Jack et al, et al 2014

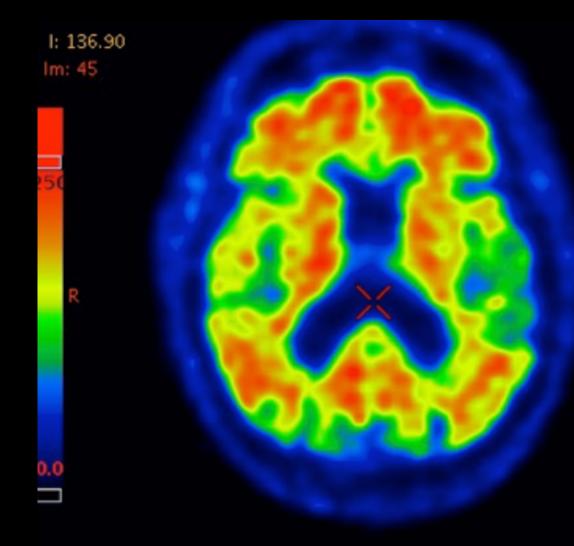


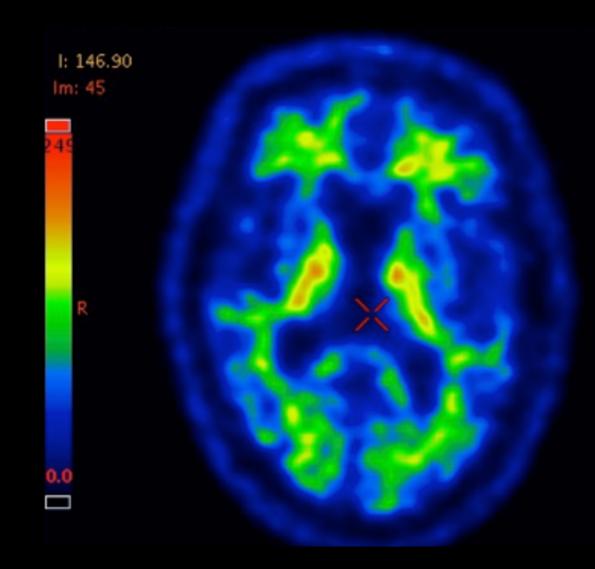




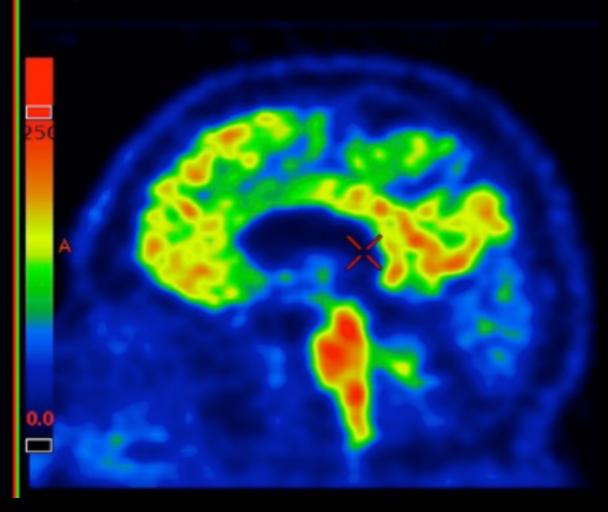


### Amyloid-beta imaging





R: 0.59

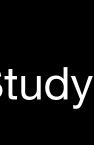




#### **18F-Flutemetamol**

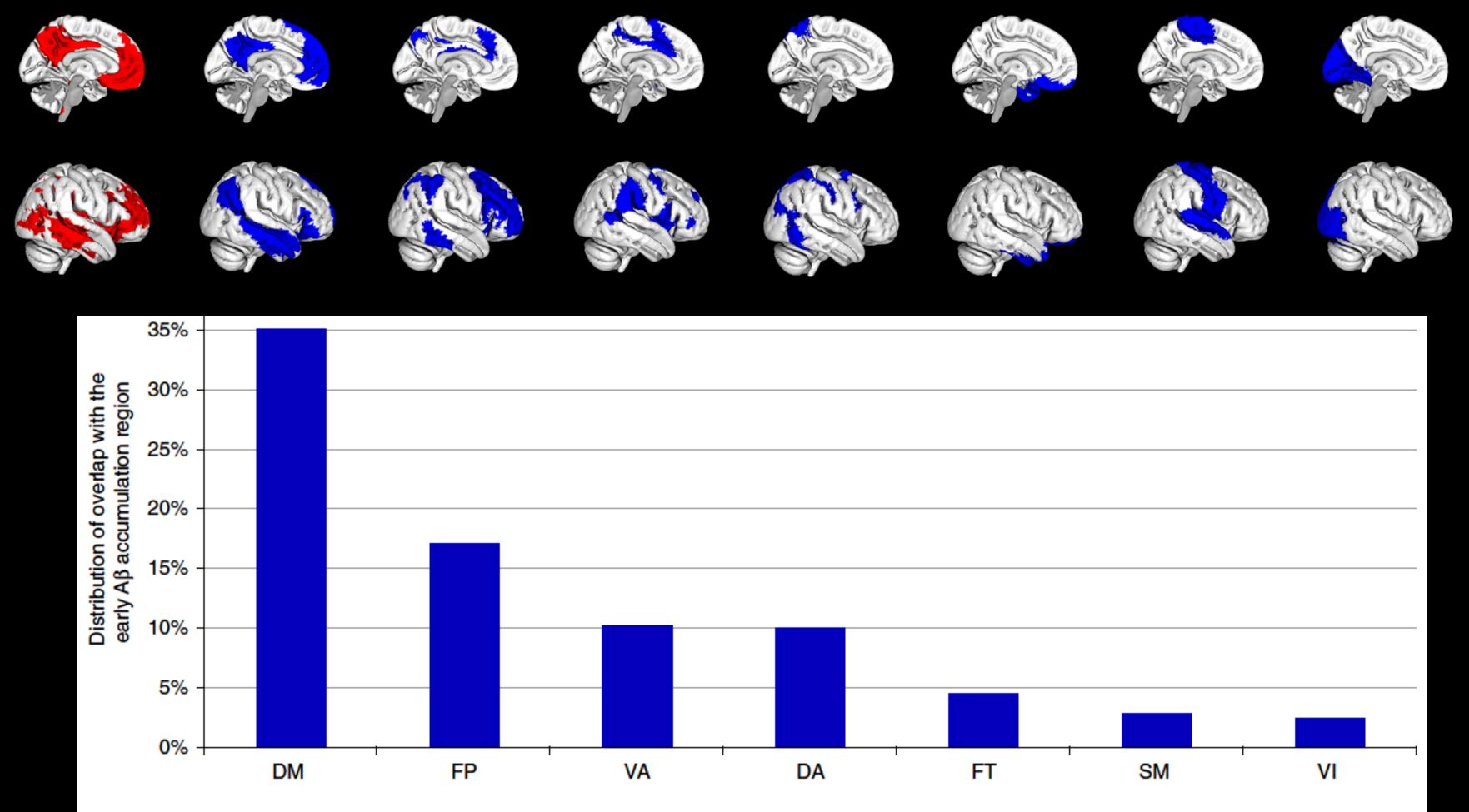
#### The Swedish Biofinder Study

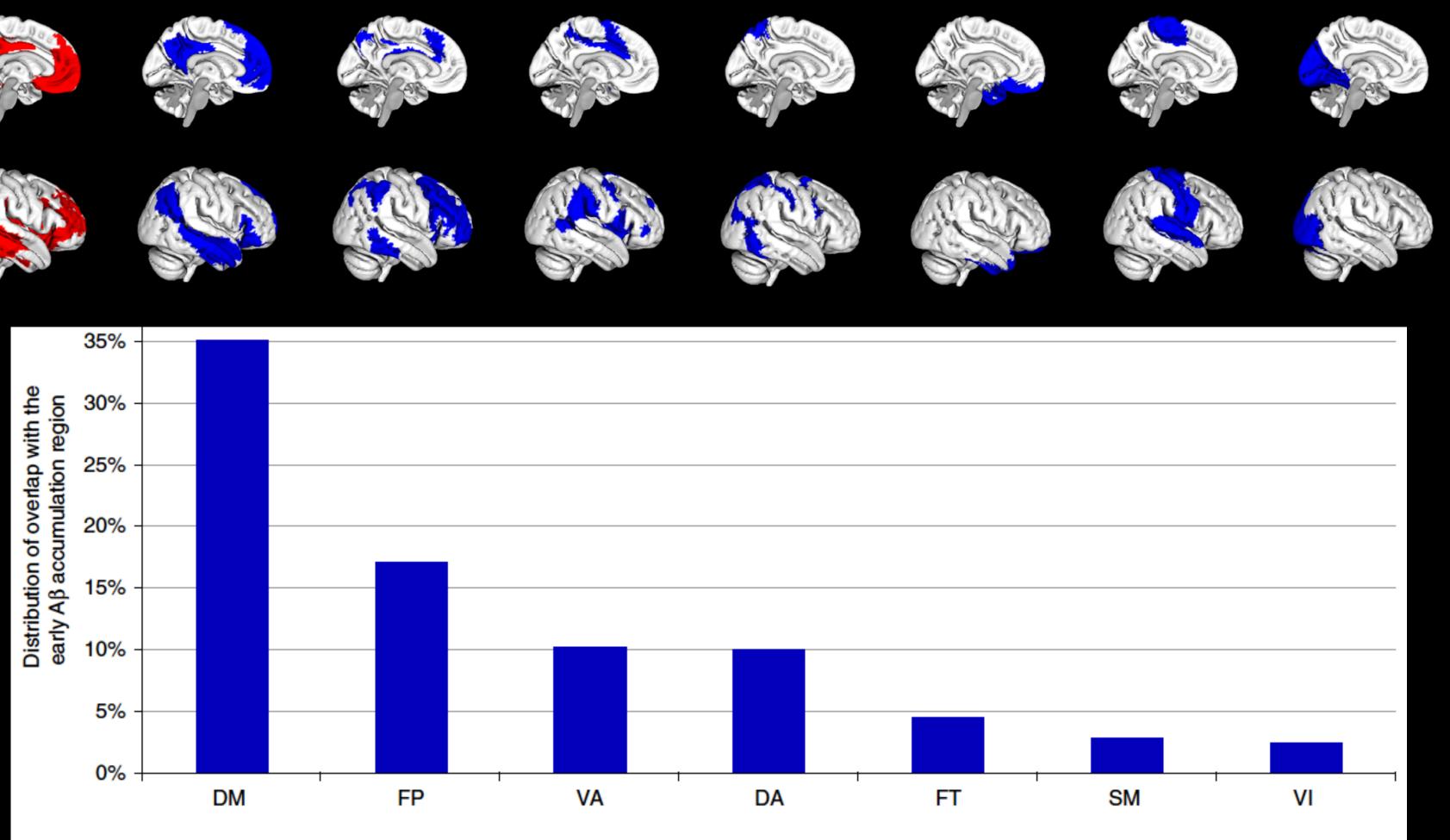
http://biofinder.se/data-biomarkers/amyloid-pet-imaging/





### Amyloid-beta and Default mode network

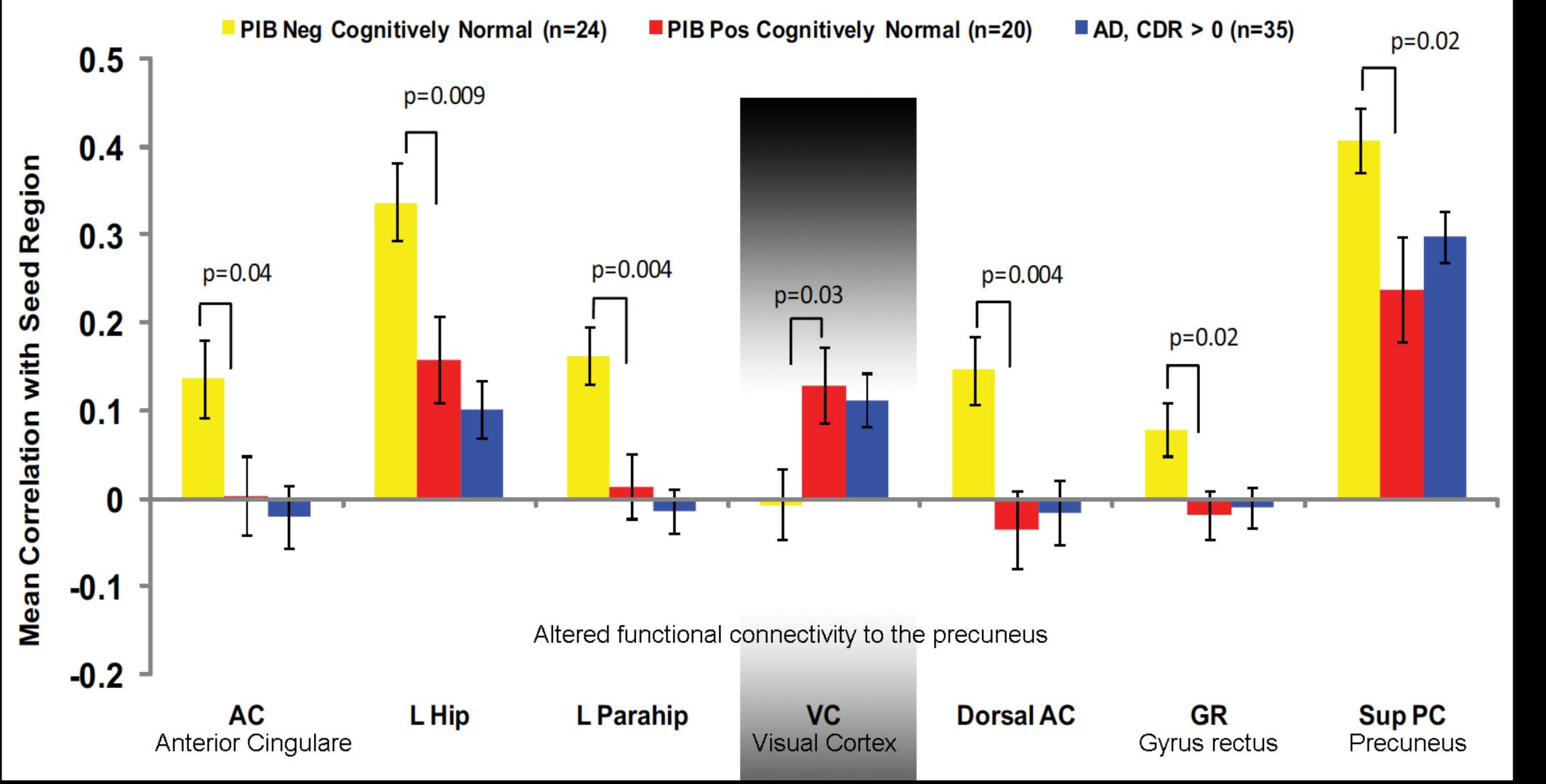




Palmqvist et al., 2017

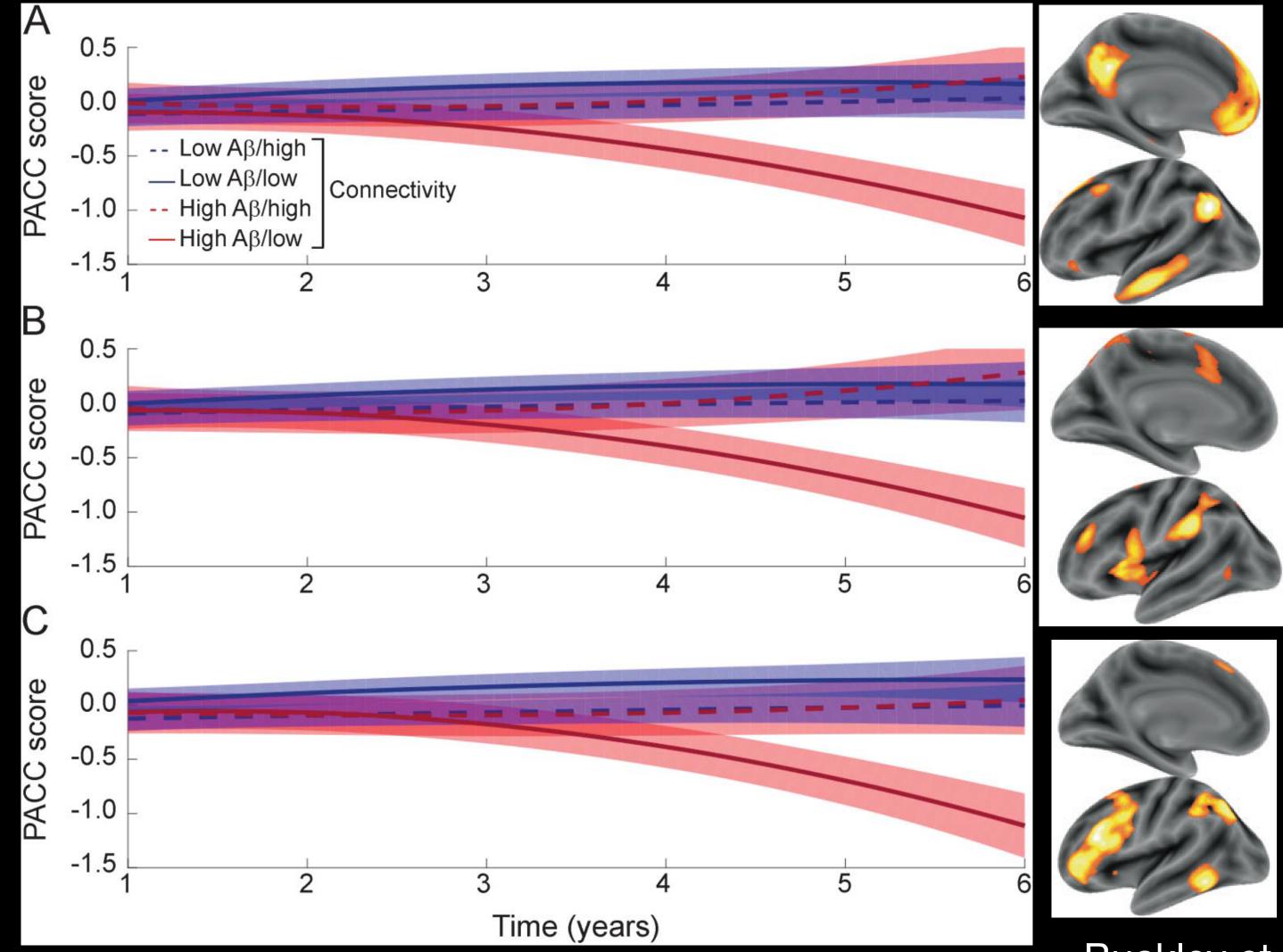


### Amyloid-beta and Default mode network





### Amyloid-beta, Connectivity and Prognosis



DMN

Salience network

**Control network** 

Buckley et al 2017. Shaw et al 2015.







# Wrapping up

- Aging is associated with :
  - decreases in functional connectivity within the DMN
  - decreased segregation between networks
  - loss of anticorrelations

- Amyloid-beta deposition  $\bullet$ 
  - Earliest accumulation: DMN
  - Impact on functional connectivity ightarrow

Mechanism: Amyloid-beta excitotoxicity? Connectivity as an cause? Compensation?

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Thanks!