

# Detection of preclinical Alzheimer's disease for the preemptive therapy



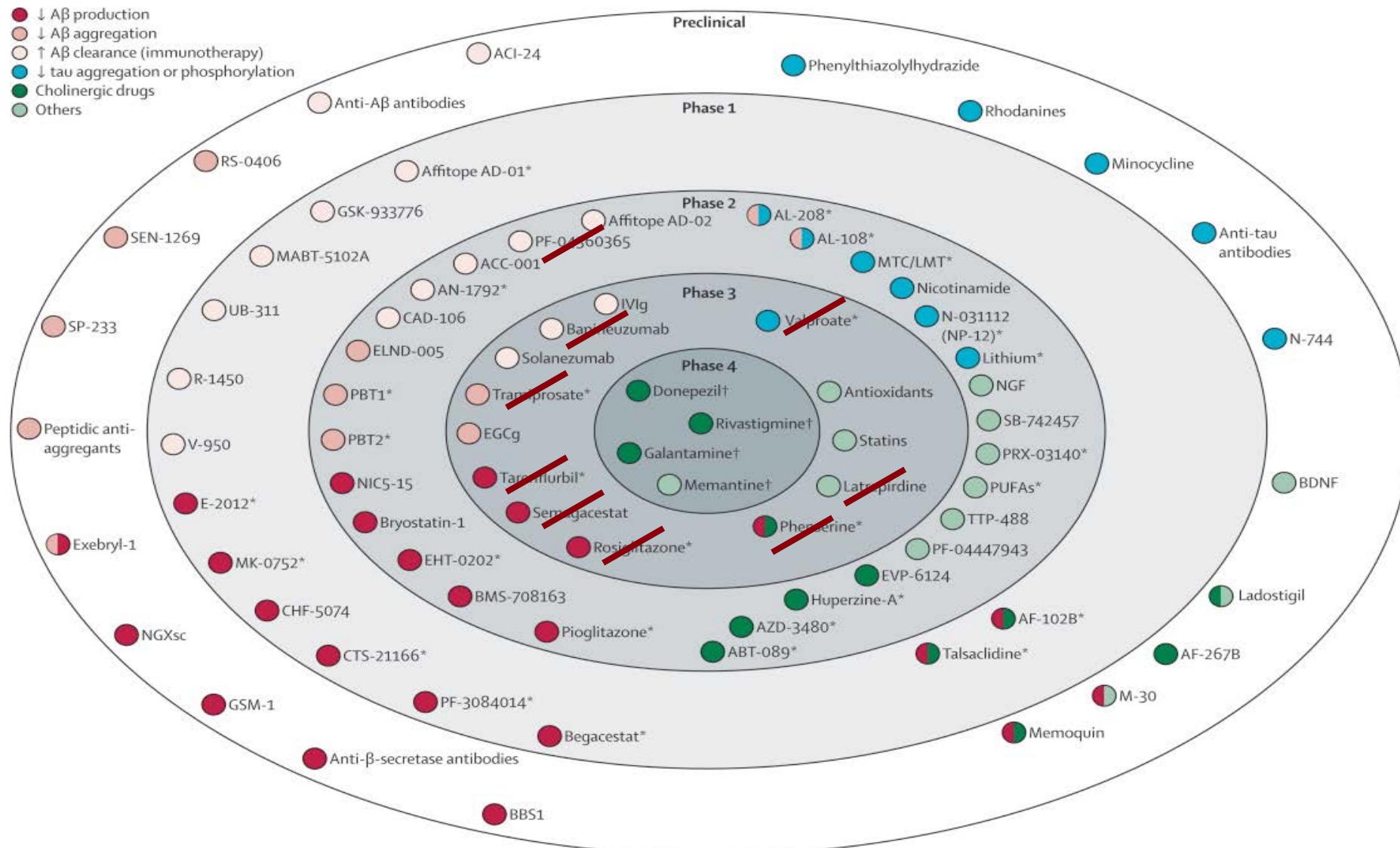
National Center for Geriatrics and Gerontology  
Center for Development of Advanced Medicine for Dementia  
Katsuhiko Yanagisawa

# *Stop Alzheimer Disease Before It Starts !*



Greg Miller, Science 337, 790-792, 2012

- ↓ A $\beta$  production
- ↓ A $\beta$  aggregation
- ↑ A $\beta$  clearance (immunotherapy)
- ↓ tau aggregation or phosphorylation
- Cholinergic drugs
- Others



# A tangled web of targets

Drugs in development for Alzheimer's disease take aim at a variety of neural mechanisms.  
But despite a wealth of possibilities, there have been few successes.

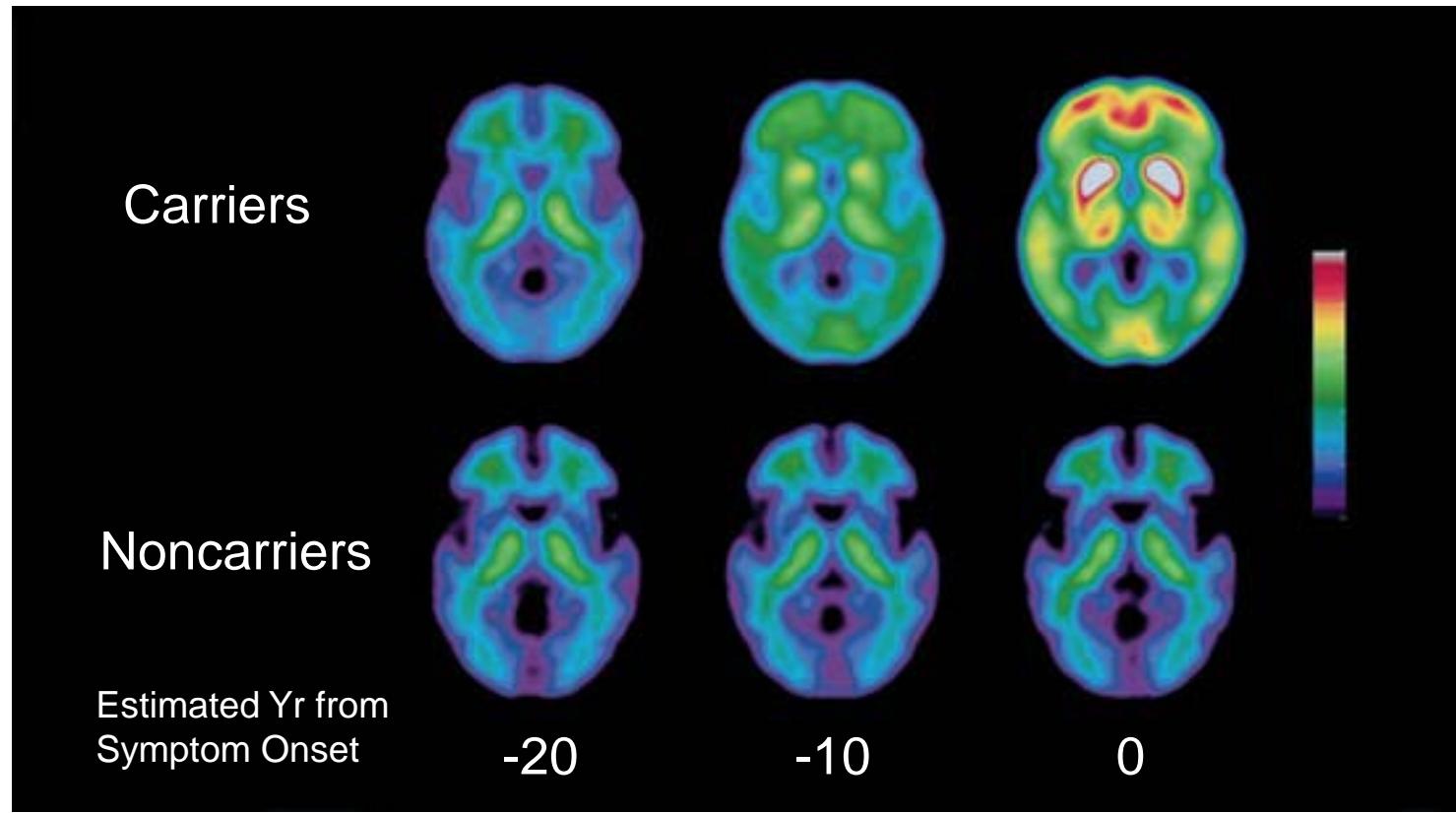
Lauren Gravitz



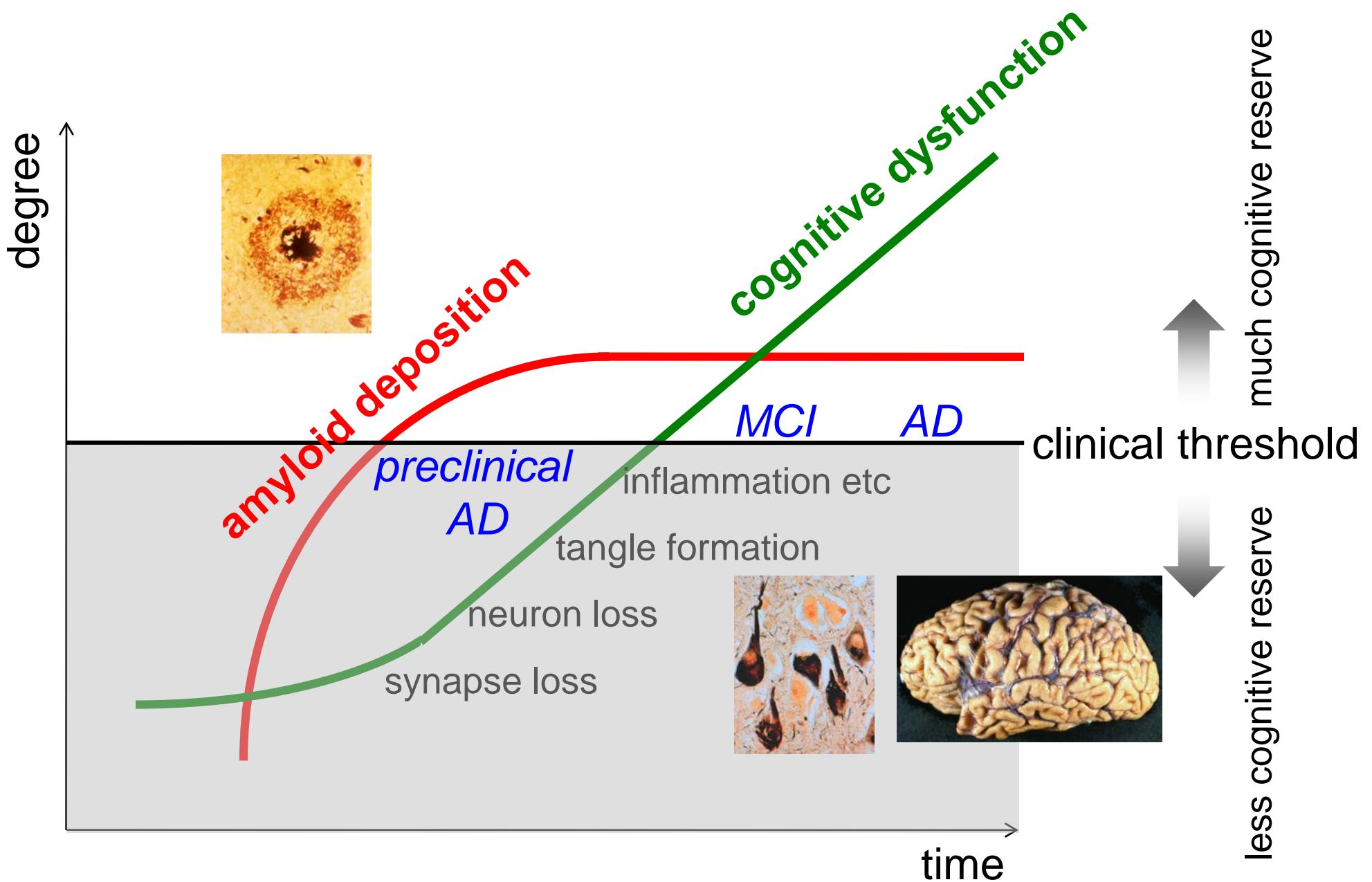
*too late for therapeutic intervention*

# PiB-amyloid imaging on PET

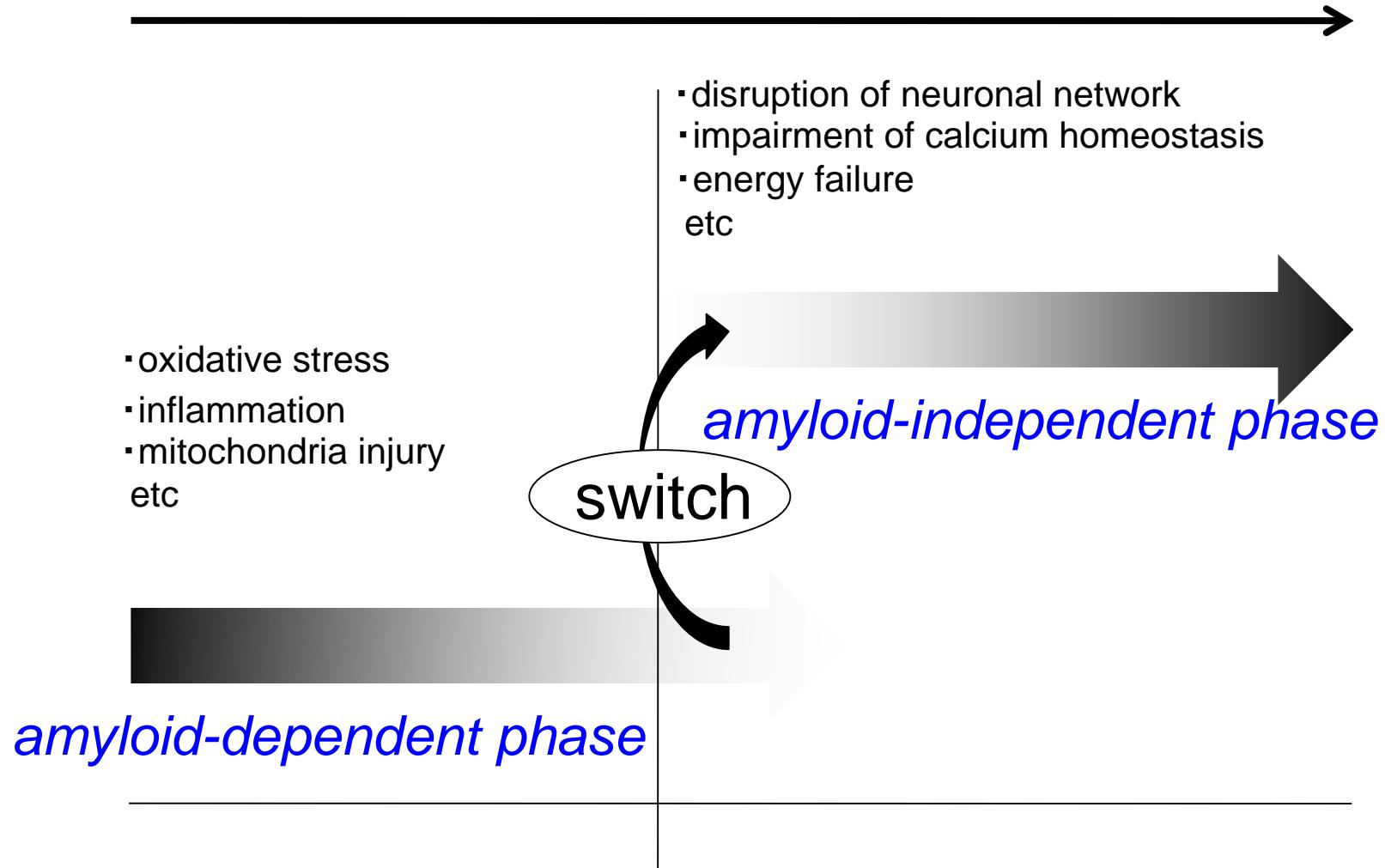
- Study on familial Alzheimer disease -



*Bateman et al, New Engl J Med, 367: 795-804, 2012*

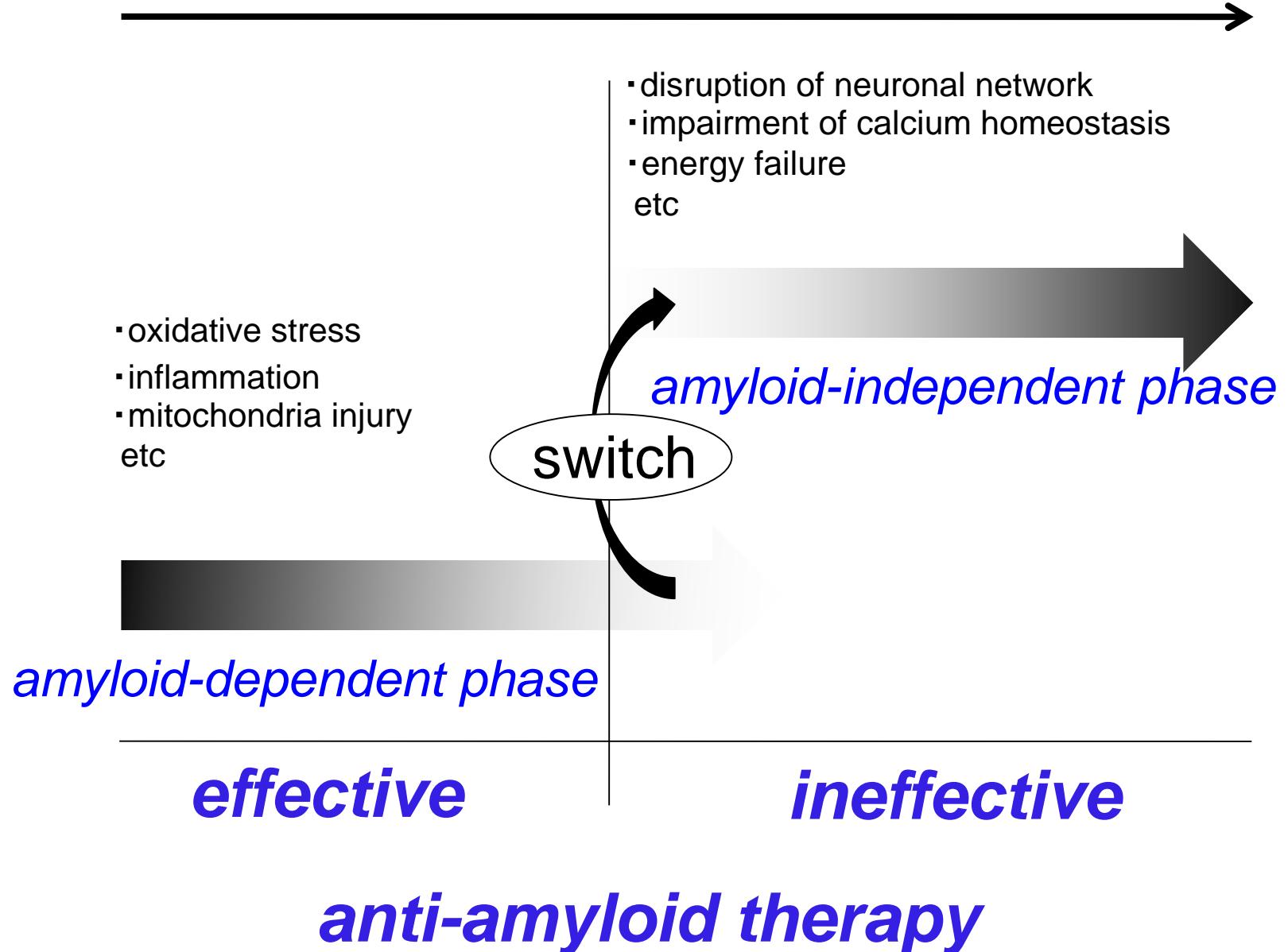


# Alzheimer Disease

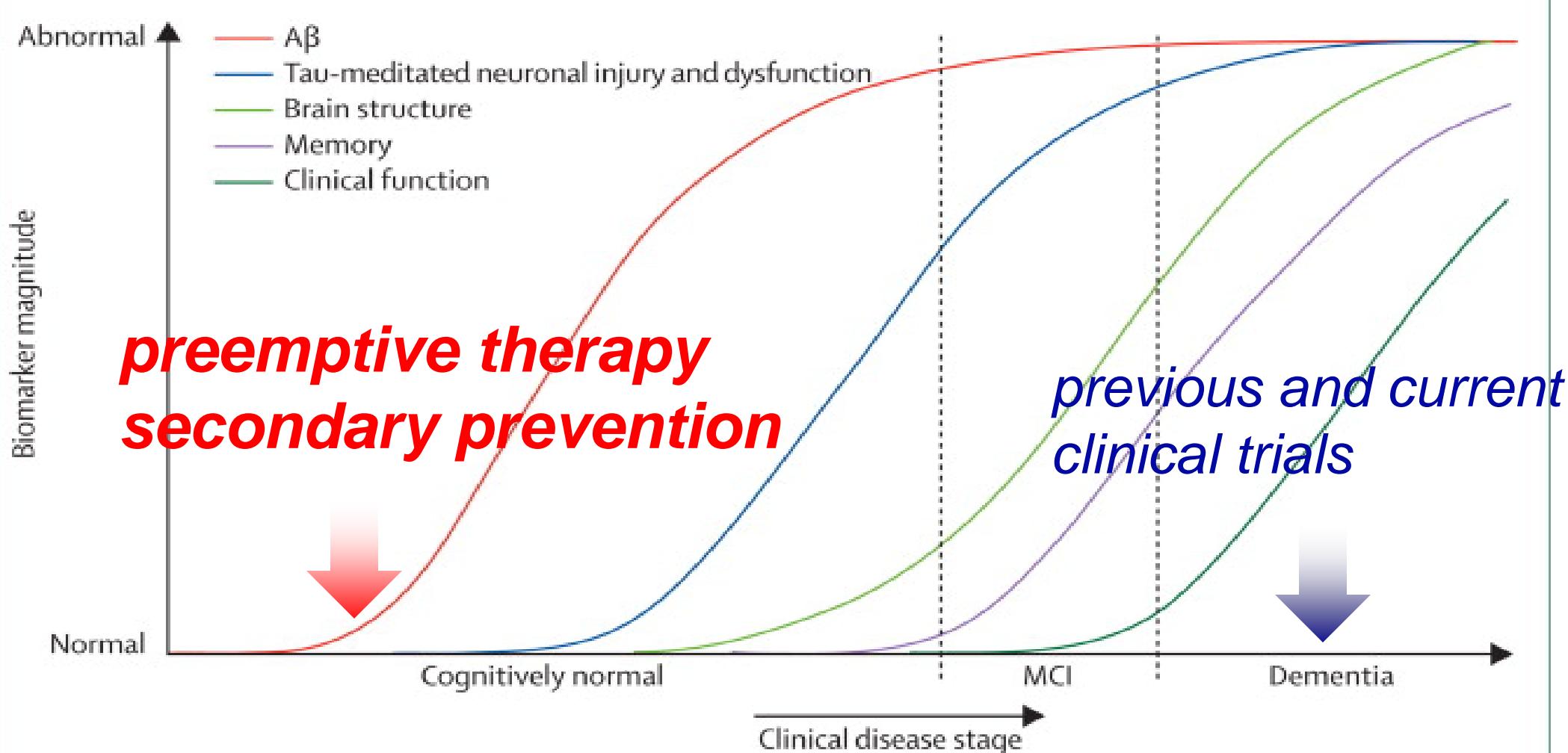


*modified from Yanagisawa. Igaku-no-Ayumi 239: 379-382, 2011*

# Alzheimer Disease



# Pathological stage of Alzheimer Disease



Modified Jack et al. *The Lancet Neurology*, 2010

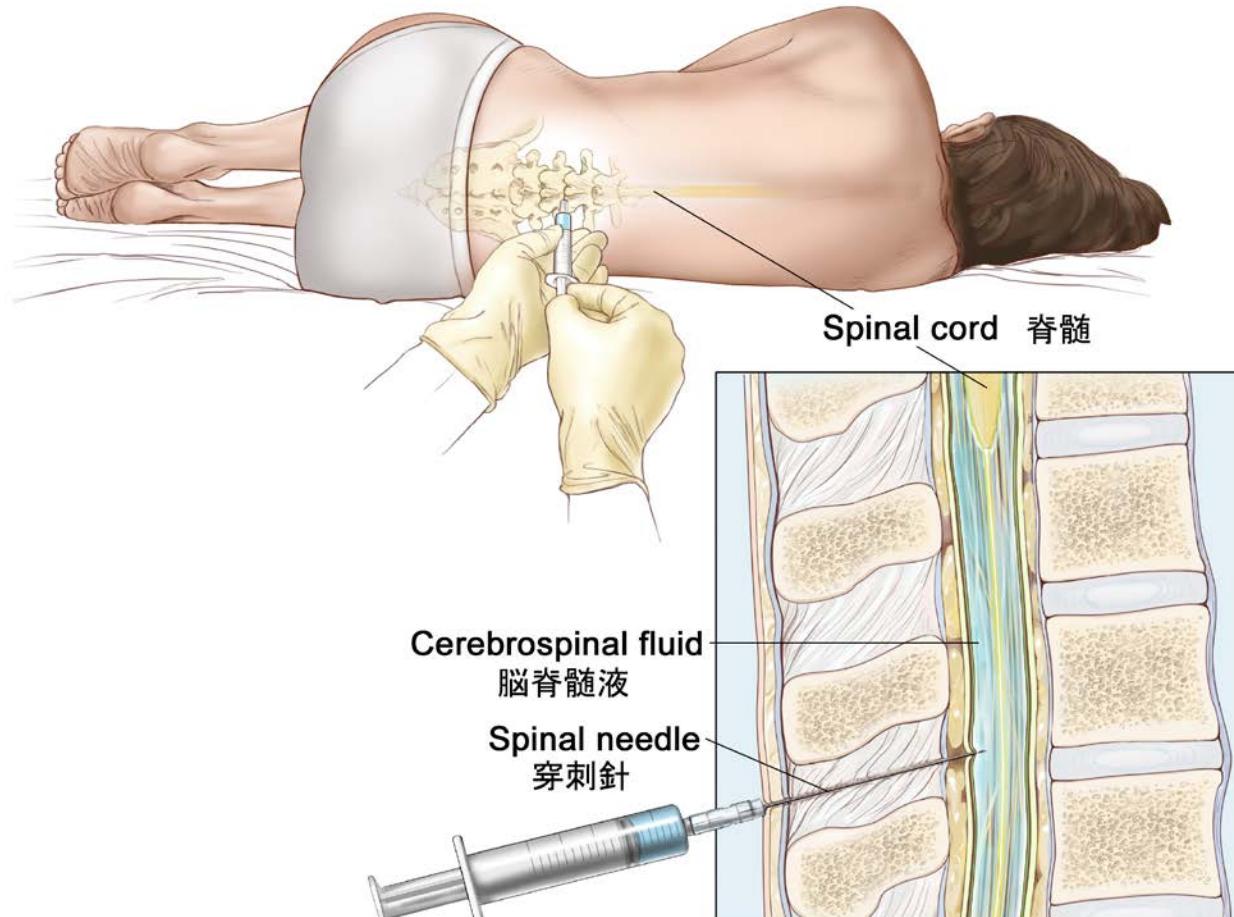


Alzheimer

## *Alzheimer Disease*

*How should you know the pathological change  
before clinical onset?*

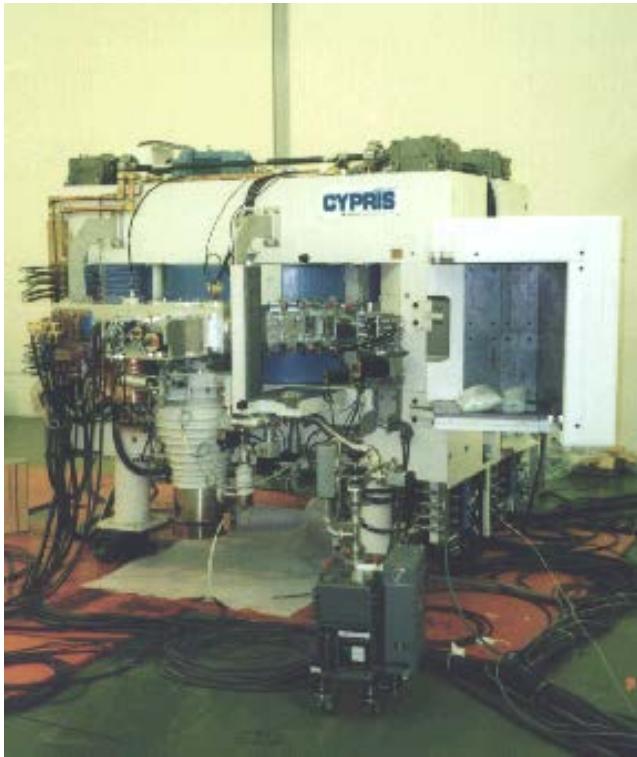
# Cerebrospinal fluid examination



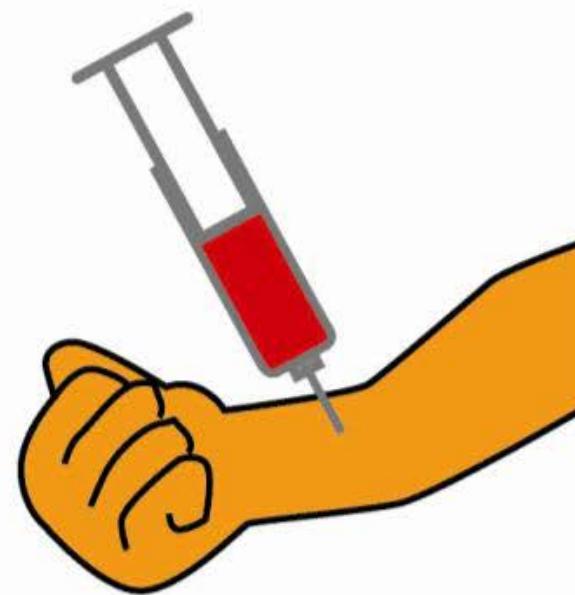
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(Yahoo Japanより取得)

# Amyloid imaging on PET



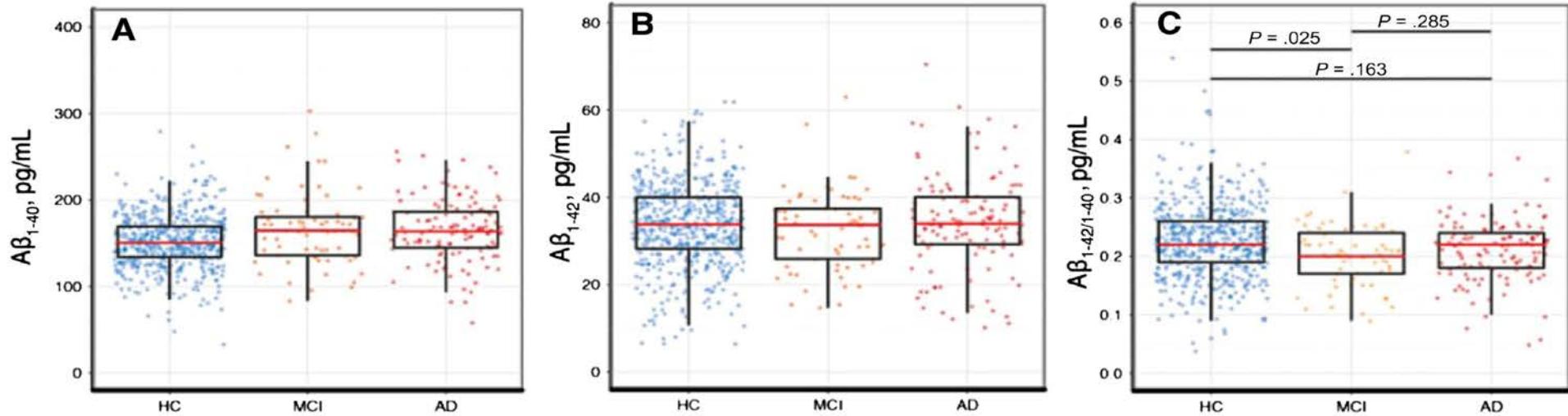
# Plasma!



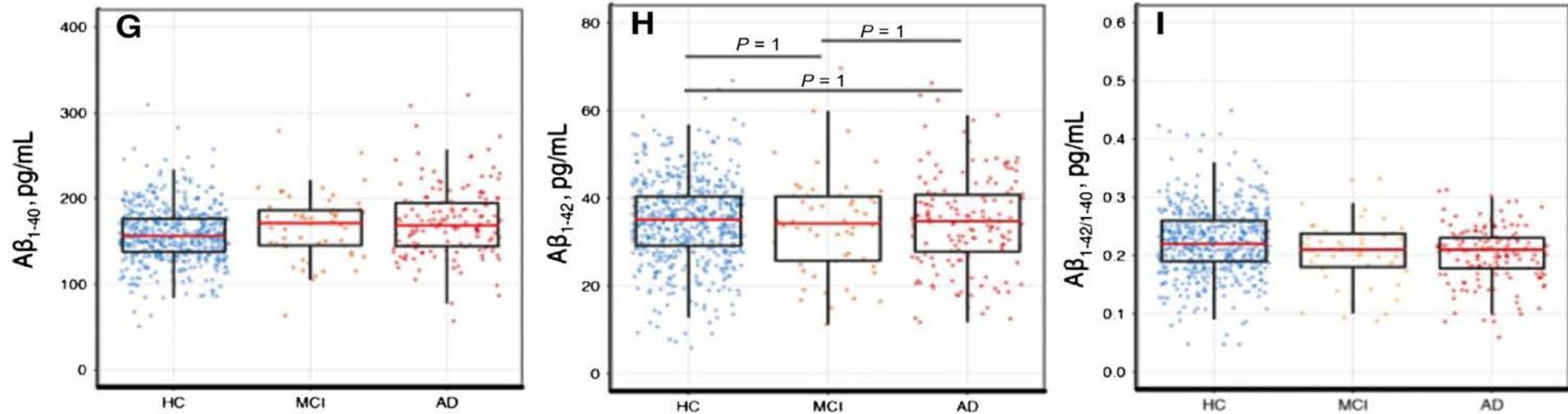
*obtained from Yahoo Japan*

# Plasma amyloid $\beta$ -proteins

Baseline



18-months later



# Novel procedure to detect A $\beta$ from plasma using mass spectrometry

Kaneko et al., Proc. Jpn. Acad. Ser. B,  
104-117, 2014

