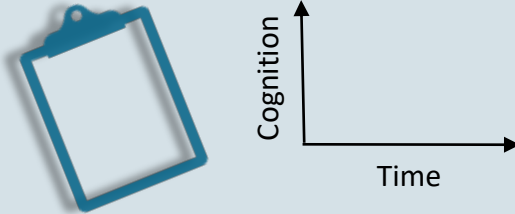# Cohort

Late middle-aged cognitively normal individuals

n=167 (WRAP cohort)
- Baseline age 59±6 years
- Age at PET 67.5±6.2 years
- 115 females
- 73% with family history of dementia
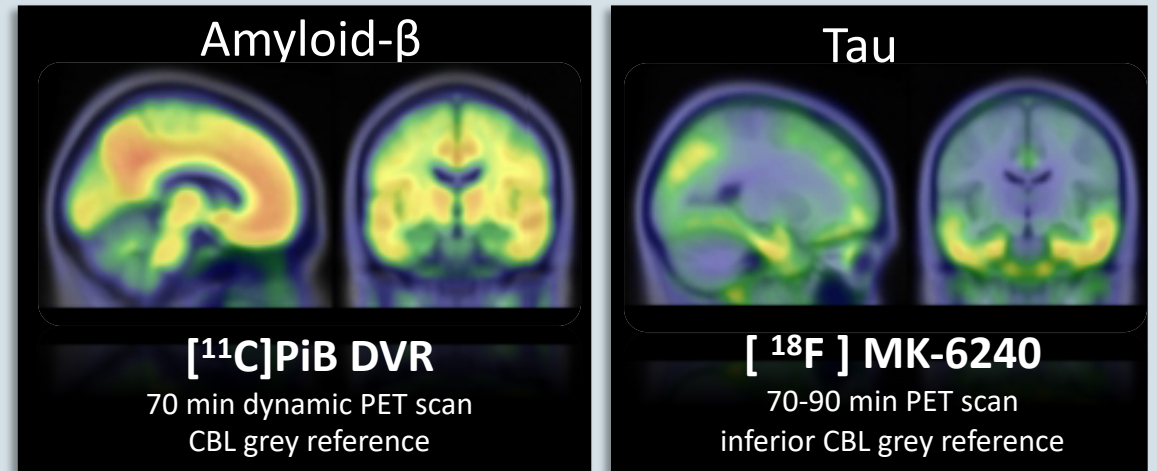
Retrospective cognition

Longitudinal trajectories

7.8 ± 2.1 years → 6 MCI

Cognitive composite
- PACC-3: verbal learning, logical memory, digit symbol
- measured approx. every 2 years

PET imaging of pathology

	Amyloid-β	Tau
	[^{11}C]PiB DVR	[^{18}F] MK-6240
	70 min dynamic PET scan CBL grey reference	70-90 min PET scan inferior CBL grey reference

A+:
PiB DVR >1.19

	T-	T+
A-	74% *n*=124	3% *n*=5
A+	14% *n*=23	9% *n*=15

T+:
entorhinal SUVR >1.27
(+2 SD of A- group)

Key findings:

- Approximately 3x faster cognitive decline in A+/T+ group compared to other subgroups

- No age association with regional [^{18}F]MK-6240 PET retention when accounting for global [^{11}C]PiB signal

- Strongest associations between global [^{11}C]PiB signal with [^{18}F]MK-6240 signal in medial temporal lobe