



ß-amyloid PET imaging:

improving understanding, diagnosis and management of Alzheimer's disease

the AMYPAD Consortium

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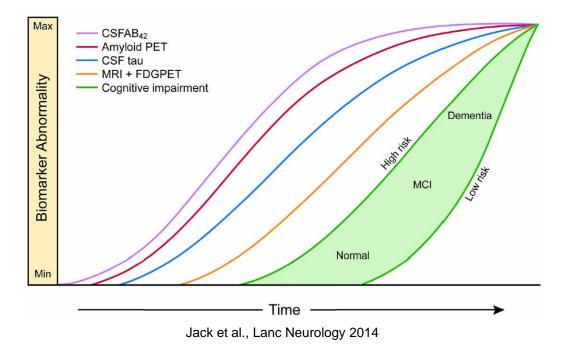
Alzheimer Europe 2018, Barcelona







β -amyloid imaging – the added value

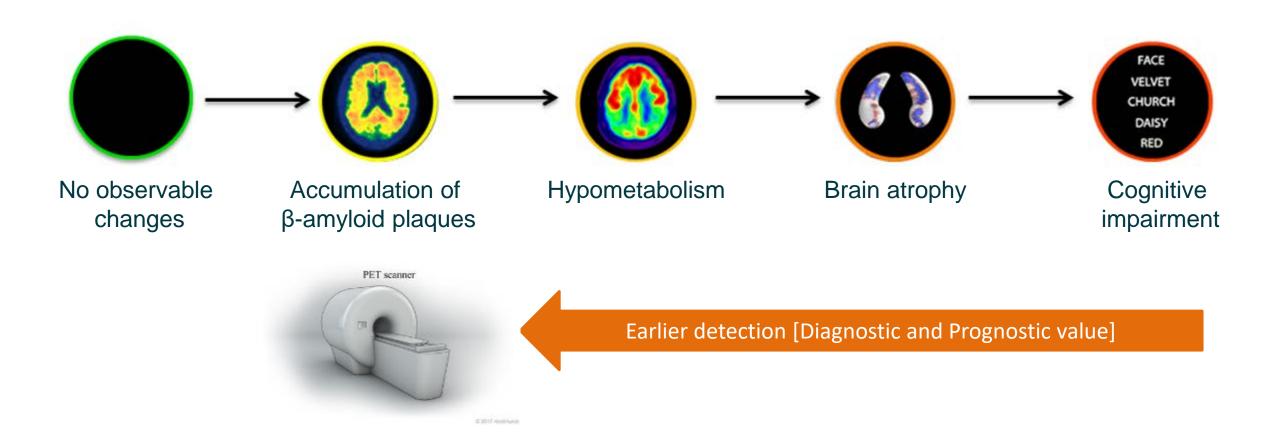


Amyloid is an **early and necessary** step in the development of AD dementia

- Clinical practice -> supporting physicians in diagnostic confidence / exclusion of AD
- Clinical trials -> slowing or modifying the course of the disease

What about prevention?

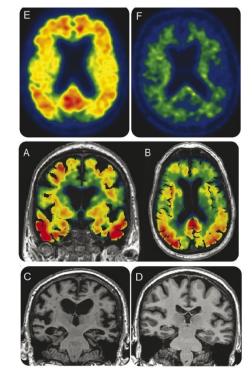






Biomarker-based prevention

Clinical routine diagnosing AD disease to prevent AD dementia



Clinical trials of biomarker-modifying therapies

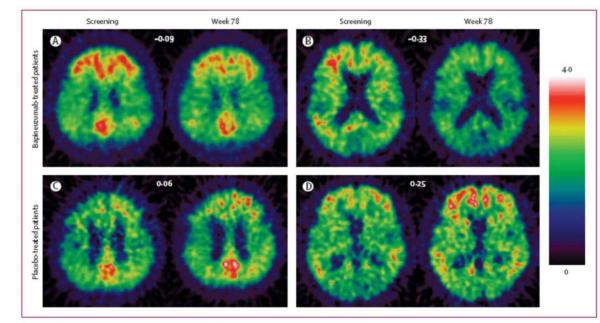


Figure 4: 11C-PiB PET images from patients treated with bapineuzumab and those given placebo

Changes from screening to week 78 in patients treated with bapineuzumab (A, B) and in patients treated with placebo (C, D). Mean "C-PiB PET changes are shown at the top centre of each panel for each patient. The scale bar shows the PiB uptake ratios relative to cerebellum by colour. The scans before and after treatment are from MRI co-registered images in the same plane. PiB= Pittsburgh compound B.



How AMYPAD?

Study 1: Diagnostic and Patient Management Study

- Usefulness of amyloid PET imaging in the clinic
- Impact of this technique in patient management
- Crucial information to payers

Study 2: Prognostic and Natural History Study

- Natural history Alzheimer's disease
- Enrich and empower secondary prevention trials
- Complement characterization of EPAD LCS

| | Cohort | Baseline PET | Repeat PET (2 yr) | Total scans |
|---|----------------------------|--------------|-------------------|-------------|
| Study 1: Diagnostic and Patient Management Study Study 2: Longitudinal Cohort Study (EPAD) Supports PoC studies | Memory clinics | 900 | 300 | 1200 |
| | Natural history cohorts | 2000 | 1000 | 3000 |
| | Total subjects | 2900 | 1300 | 4200 |







Diagnostic and Patient Management Study (DPMS)





Current knowledge





- Medical Management Changes post-PET:
 - 67.8% of MCI patients (47.8% change in AD drugs, 36.0% change in other drugs, 23.9% changes in counseling)
 - 65.9% of dementia patients (47.7% change in AD drugs, 32.2% change in other drugs, 15.3% change in counseling);
- Need for additional testing post-PET:

Domontia

Reduced need when amyloid PET scans were made from 26.3% to 11.0% in neuropsychological testing and from 10.5% to 1.0% in spinal fluid testing

MCI

SCD

| Deltaplan Dementie | |
|-----------------------|--|
| | |

ABIDE Alzheimer Biomarkers in Daily Practice

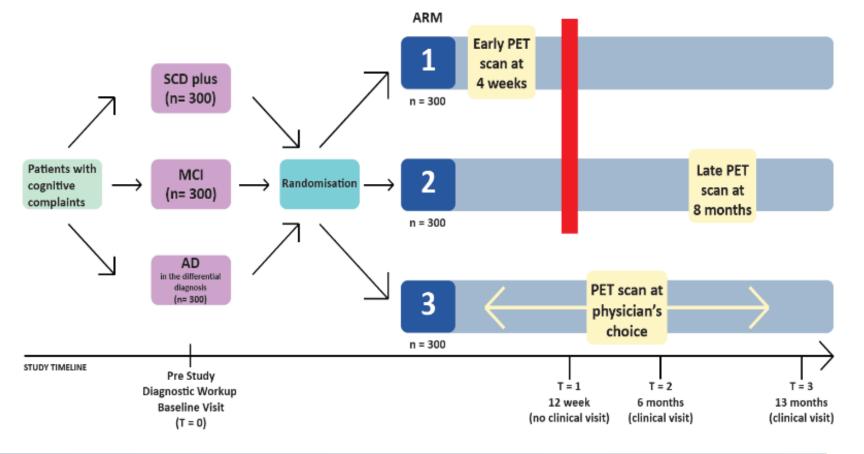
| | Dementia | | | | 300 | |
|---------------------------|-----------|----------|----------|----------|----------|----------|
| | AD | non-AD | AD | non-AD | AD | non-AD |
| n | 164 | 70 | 72 | 42 | 16 | 143 |
| Amyloid PET, positive (%) | 128 (78%) | 23 (33%) | 45 (63%) | 10 (24%) | 6 (38%) | 30 (21%) |
| Change in syndrome | 4 (2%) | 0 (0%) | 2 (3%) | 6 (14%) | 1 (6%) | 4 (3%) |
| Change in etiology | 36 (22%) | 14 (20%) | 27 (38%) | 11 (26%) | 10 (63%) | 27 (19%) |

Overall, 25% change in diagnosis. Due to negative amyloid PET in 66% Demented: 21% change. Non-demented: 27% change.



What is the effect of using amyloid PET imaging in the clinic for:

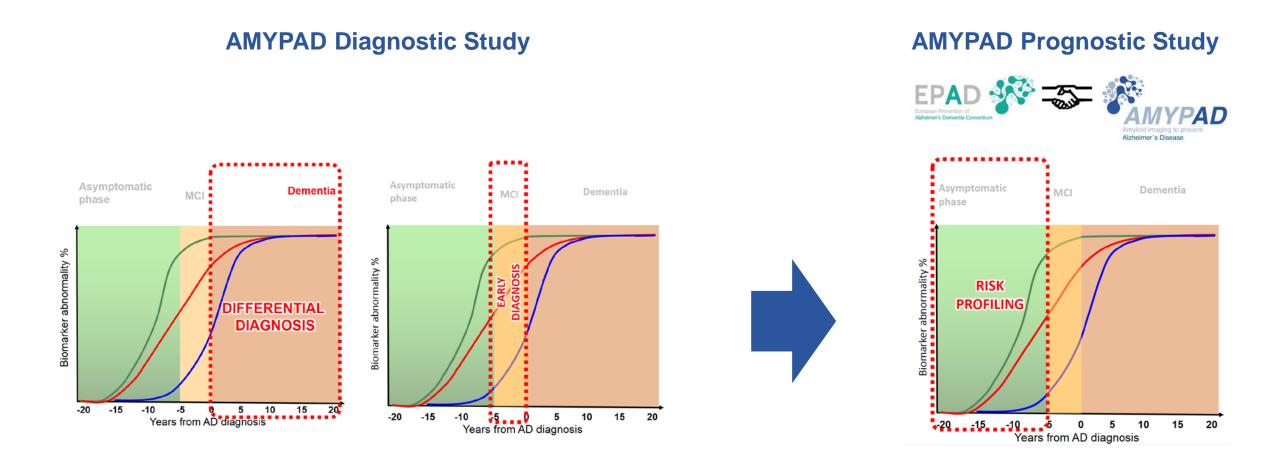
- Physicians confidence in etiological diagnosis
- Patient management
- ✤ Health-care resource utilization
- Patient-related outcomes





Prognostic and Natural History Study (PNHS)



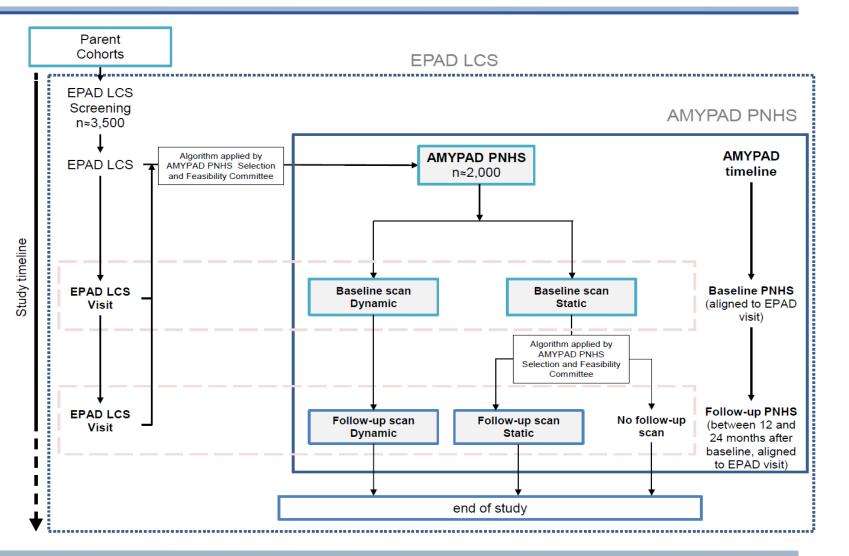




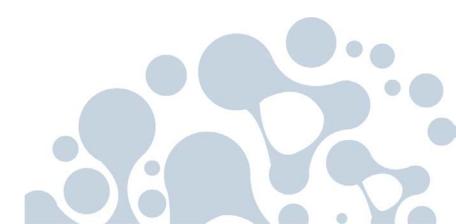
Study design and objectives

What is the value of amyloid PET imaging to:

- Determine a risk of progression to AD dementia
- Individualize that risk
- Understand the natural history of the disease
- Support preventive trials in accurate participant selection and measurement of treatment effects

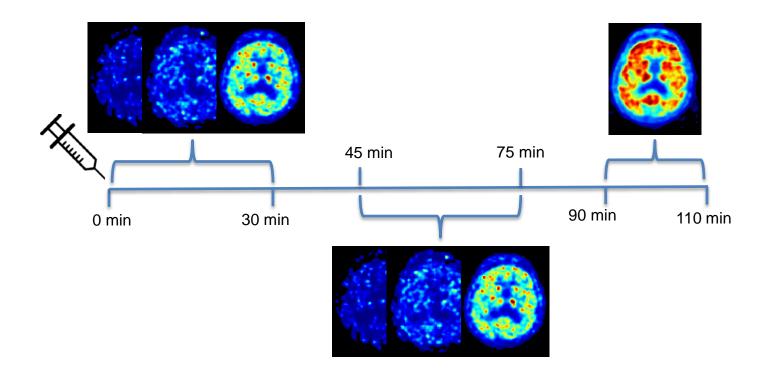


AMYPAD today



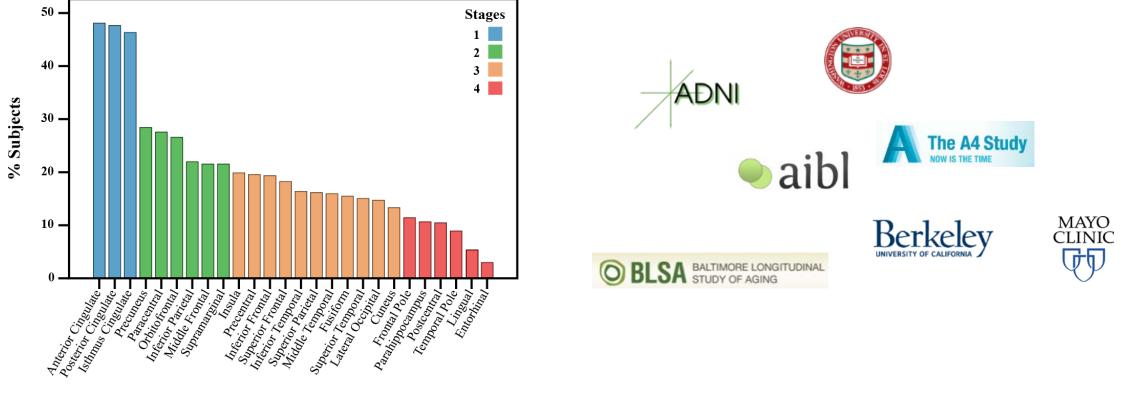


An imaging protocol to deliver **high-quality** efficiently and with <u>comfort to the patient/participant</u>





Collection of external data sets (RE-USE) to understand earliest stages of AD



Regions of interest



Collection of external data sets (RE-USE) to understand earliest stages of AD





...both studies ongoing to deliver much more

Diagnostic and Patient Management Study (DPMS)





Prognostic and Natural History Study (PNHS)

58 patients randomized, 19 scanned

3 sites recruiting, 5 more on board before spring

6 participants included, 1 scanned

1 site recruiting, 4 more on board before spring and 6 more on board by summer

The addition of TWO sub-studies to understand the biomarker disclosure process and its impact on patients



Thank you!

