

the low-probability was positive. The syndromic diagnoses that most discrepancies were found were: amnesic MCI (37% had negative PIB), amnesic and other domains (40%, negative PIB) into high probability group; non-amnesic MCI (33% positive PIB), and non-logopenic PPA and FTD (33% and 45% positive PIB) into low probability group. Normal controls and AD patients (typical and atypical presentation) were the most consistent across clinical and molecular diagnostics. **Conclusions:** There were discrepancies in molecular diagnosis in both high and low probability groups. The implications of these inconsistencies were different between each clinical category. The most important contribution to the diagnosis of 11C-PIB-PET is really significant in cases of early-onset AD, and atypical presentation (PPA, FTD and PCA).

IC-P-006 HIPPOCAMPAL ATROPHY: AUTOMATED VOLUMETRY VS VISUAL EXAMINATION—AN ARG ADNI COHORT ANALYSIS

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Background: Hippocampal atrophy is one of the earliest findings in Alzheimer Disease (AD). MRI volumetry is a low-cost in vivo study to quantify brain atrophy. Our goal is to analyze MRI brain volumetry in Arg. ADNI cohort in order to determine the clinical usefulness. As a secondary objective a comparison between visual and automated analysis of the hippocampal atrophy was performed. **Methods:** 45 patients of the Argentina-ADNI database were included (12 controls, 12 early mild cognitive impairment (eMCI), 13 late MCI and 8 dementia AD). Neuropsychological assessment, labs, CSF biomarkers, fdg-PET and PiB-PET were performed. All of them had a MRI Brain Volumetry including Absolute Hippocampal volume, whole brain volume and a hippocampal/whole brain ratio (Hi/WBV). Study of correlation between hippocampal visual and automated examination was performed on 49 patients between cognitive neurologist and neuroradiologist. **Results:** Whole brain volume was similar between groups, but hippocampal volume was inversely related to clinical impairment. Specially Hi/WBV ratio. 88% of AD, 46% of LMCI and 42% of eMCI were below a cut-off point of 0.0025. In addition, 92% of control patients had a Hi/WBV superior to 0.0025 value. On the other hand, 68.4% of patients who had Hi/WBV < 0.0025 had another positive AD biomarker (FDG-PET, PiB-PET or CSF profile) versus 34.6% of those > 0.0025. With a specificity of 0.72, sensitivity 0.61, PPV 0.68 and NPV 0.66. A visual examination of hippocampal atrophy showed low concordance in the control group and a high influence of the years of training. **Conclusions:** A reliable cutoff point to measure hippocampal atrophy could be determined using MRI volumetry, this finding correlates with clinical impairment and the presence of other AD biomarkers. Visual examination of MRI seems to be insufficient to assess hippocampal atrophy in the early stages of the disease even for a specialist in neuroradiology or cognitive neurology.

IC-P-007 PATTERNS OF [C-11]PIB AMYLOID BURDEN THAT ARE ASSOCIATED WITH CEREBROSPINAL FLUID AD BIOMARKERS IN PEOPLE AT RISK FOR ALZHEIMER'S DISEASE: FINDINGS FROM THE WRAP STUDY

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Background: Further research is needed on the full spectrum of cerebrospinal fluid (CSF) markers and their relationships to amyloid imaging in all stages of Alzheimer's disease (AD). This study examines a wide range of CSF markers in a relatively young, preclinical cohort enriched with risk factors (APOE4 and AD parental history) for AD in conjunction with [C-11]Pittsburgh Compound B (PiB) positron emission tomography (PET). **Methods:** N=106 Participants (mean age 60.43, SD 5.66) enriched with genetic and familial risk for AD from the Wisconsin Registry for Alzheimer's Prevention (WRAP) underwent PET-PiB imaging and lumbar puncture and were grouped as amyloid-positive (PiB+), amyloid-indeterminate (PiBi), or amyloid-negative (PiB-) based on the amount and pattern of amyloid deposition. ANOVAs were conducted on CSF Aβ, tau, and CSF markers of inflammation (MCP-1, YKL-40) and axonal injury (NFL). Regional voxel-wise regression analyses of PiB scans were performed to examine the spatial relationship between CSF measures and amyloid load. **Results:** ANOVA indicated a pattern of lower Aβ42 in PiB+ compared to PiBi or PiB- while the ratios of Aβ40, p-tau, t-tau, MCP-1, YKL40, and NFL to Aβ42 were all higher in PiB+ compared to PiBi or PiB-. Correlations with extracted PiB distribution volume ratio from precuneus, and voxel-wise regressions, p(FWE) < 0.05, showed a negative relationship between Aβ42 and PiB while the ratios to Aβ42 all showed a positive relationship to PiB. A common voxel-wise pattern was observed for all CSF, but most notably in p-tau/Aβ42 and t-tau/Aβ42 in regions previously implicated in symptomatic AD. **Conclusions:** Group comparisons and voxel-based tests demonstrated that CSF ratios to Aβ42 accounted for more variance than any single CSF measure alone. The positive relationship with PET-PiB likely reflects simultaneous amyloid deposition and early neurodegenerative processes. Voxel-based results are consistent with previous findings that the regions commonly labeled as the default mode network are functionally and anatomically affected in AD, even before symptom onset. An overlay of voxel-based results of Aβ42, 1/Aβ42 and p-tau/Aβ42 indicate that Aβ42 is likely driving the overall pattern, while tau provides additional sensitivity. This study shows that, together, CSF and PET-PiB provide critical and striking information about this important preclinical disease stage.

Table
CSF Statistical Contrasts by Amyloid Grouping

CSF marker	ANOVA	Group Contrast	Post-hoc T-test (Tukey-corrected)
Aβ42	P=0.014	PiBi > PiB+	P=0.040
		PiB- > PiB+	P=0.013
Aβ40/42	P=0.000	PiB+ > PiB-	P=0.000
		PiB+ > PiBi	P=0.000
t-tau/Aβ42	P=0.000	PiB+ > PiB-	P=0.000
		PiB+ > PiBi	P=0.000
p-tau/Aβ42	P=0.000	PiB+ > PiB-	P=0.000
		PiB+ > PiBi	P=0.000
MCP-1/Aβ42	P=0.003	PiB+ > PiB-	P=0.002
		PiB+ > PiBi	P=0.045
YKL40/Aβ42	P=0.000	PiB+ > PiB-	P=0.000
		PiB+ > PiBi	P=0.000
NFL/Aβ42	P=0.000	PiB+ > PiB-	P=0.000
		PiB+ > PiBi	P=0.001
1/Aβ42	P=0.000	PiB+ > PiB-	P=0.000
		PiB+ > PiBi	P=0.004