

CSF tau biomarkers and structural brain MRI measures in frontotemporal lobar degeneration

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Introduction: In recent years, in the field neurodegenerative diseases, increasing attention has been pointed to CSF biomarkers and their integration with neuroimaging [1]. Frontotemporal lobar degeneration (FTLD) refers to a heterogeneous group of clinical syndromes with different underlying proteinopathies including tau pathology [2]. CSF biomarkers have been proposed as diagnostic and prognostic factors [3].

Objective: Aim of our study was to evaluate the relationship between CSF tau biomarkers and structural MRI brain measures in FTLD.

Materials and Methods: We included early FTLD patient. All included patients underwent lumbar puncture to evaluate amyloid, total-tau (t-tau), phospho-tau 181 (p-tau); p-tau/t-tau ratio was also calculated; brain MRI was performed to estimate whole brain volume, volume of principal deep grey matter structures and regional cortical thickness using FreeSurfer software version 7.1.1 (<http://surfer.nmr.mgh.harvard.edu>). The principal clinical and demographic features were also recorded.

Results: Demographic characteristics of the 28 included patients were as follows: female/male: 9/19; mean±SD age: 67.9±7.7 years. The p-tau/t-tau ratio was significantly correlated with whole brain volume ($r=0.77$; $p < 0.001$), brain-stem volume ($r=0.41$; $p: 0.04$), left putamen volume ($r=0.57$ $p: 0.006$) left pallidum volume ($r=0.41$; $p: 0.04$), right accumbens volume ($r=0.47$; $p: 0.02$). P-tau/t-tau ratio showed also a significant correlation with cortical thickness of left temporal lobe ($r=0.74$; $p < 0.001$) and left caudal middle frontal cortex ($r=0.45$; $p: 0.03$). Linear regression showed a significant relationship between p-tau/t-tau ratio and left temporal pole ($p = 0.001$; $r^2: 0.60$) after controlling for age and gender.

Conclusions: Our data suggest that CSF biomarkers, especially p-tau/t-tau ratio, could play a role as prognostic factor in FTLD. Further longitudinal investigations are needed to confirm these findings.

References:

[1] LH Meeter, LD Kaat, JD Rohrer, JC van Swieten. Imaging and fluid biomarkers in frontotemporal dementia. *Nat Rev Neurol.* 2017 Jul;13(7):406-419.

[2] S Abu-Rumeileh, N Mometto, A Bartoletti-Stella, B Polischi, F Oppi, R. Poda, M Stanzani-Maserati, P Cortelli, R. Liguori, S Capellari, P Parchi. Cerebrospinal Fluid Biomarkers in Patients with Frontotemporal Dementia Spectrum: A Single-Center Study. *J Alzheimers Dis.*;66: 551-563. doi: 10.3233/JAD-180409 (2018).

[3] YA Pijnenburg, NA. Verwey, WM van der Flier, P Scheltens, CE Teunissen. Discriminative and prognostic potential of cerebrospinal fluid phosphoTau/tau ratio and neurofilaments for frontotemporal dementia subtypes. *Alzheimers Dement (Amst).* 14;1: 505-12 (2015).