SEX DIFFERENCES IN NEUROPSYCHOLOGICAL PROFILE AND BRAIN HYPOMETABOLISM IN BEHAVIOURAL-VARIANT FRONTOTEMPORAL DEMENTIA

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Introduction and aim

Behavioral variant frontotemporal dementia (bvFTD) is a form of early-onset dementia manifesting with executive deficits and behavioral changes. Sex differences in prevalence, clinical characteristics, and biomarkers, have been described in several neurodegenerative diseases, including FTD (1). However, these findings are still sparse and need further characterization.

Our study aimed to investigate sex differences in neuropsychological profile and brain hypometabolism pattern, evaluated with FDG-PET.

Materials and methods

- Patients included: 20, 12 female (F), 8 males (M), diagnosed with bvFTD (2) at the Memory Clinics of Novara and Vercelli, Piedmont, Italy.
- **Investigations:** neurological and ullet**neuropsychological** evaluation at baseline, to characterize the clinical presentation; a brain MRI and a lumbar puncture to support clinical diagnosis; a brain FDG-PET scan, analyzed with a validated SPM method (3) to obtain brain hypometabolism single-subject maps using a large dataset of controls for comparison (4). In familial cases, mutations in genes ulletfrequently associated with FTD were investigated.

Results

- Sex did not influence global cognition, evaluated with the Mini-Mental State Examination and the Montreal Cognitive Assessment.
- Specific cognitive domains: F ulletperformed better than M in working **memory** (i.e., corrected Reverse Digit Span, p=0.01), and worse in language (i.e., MOCA denomination subscore, p=0.03). Specific brain hypometabolic patterns: FDG-PET showed predominant а frontotemporal hypometabolism in both groups. In M, compared with F, the insula and the orbitofrontal cortex were involved with widespread hypometabolism.



TABLE: NEUROPSYCHOLOGICAL DIFFERENCES BETWEEN BVFTD FEMALES AND MALES All results shown are mean values. FAB: Frontal Assessment Battery; MMSE: Mini-Mental State Examination; MOCA: Montreal Cognitive Assessment; RAVLT: Rey Auditory Verbal Learning Test; ROCF: Rey-Osterrieth complex figure. **FIGURE: PATTERN OF BRAIN HYPOMETABOLISM IN FEMALES (LEFT) AND MALES (RIGHT)** Results of the group analysis.

Conclusion and discussion

From our preliminary data, sex might have a role in determining the type of neuropsychological impairment and the brain hypometabolism pattern in bvFTD, with males performing worse than females in working memory tests and better in language, with a broader hypometabolism pattern possibly related with neuropsychiatric burden. These results confirm that sex may play a role in influencing clinical phenotype in bvFTD, supporting the hypothesis of sex as a pivotal variable for biological and clinical differences in neurodegenerative diseases.

Bibliography

 Pengo M., 2022, Sex influences clinical phenotype in frontotemporal dementia, Neurol Sci.
Rascovsky K. et al., 2011, Sensitivity of revised diagnostic criteria for the behavioral variant of frontotemporal dementia. Brain.
Perani D. et al., 2014, Validation of an optimized SPM procedure for FDG-PET in dementia diagnosis in a clinical setting. Neur Clin.
Caminiti S.P. et al., 2021, Validation of FDG-PET datasets of normal controls for the extraction of SPM-based brain metabolism maps. Eur J Nucl Med Mol Imaging.

