

Advanced perspectives for the diagnosis of Parkinson's and Alzheimer's disease through machine learning techniques

Malvaso Antonio^{1,2,3,5}, Panarese Silvia^{5,6}, Catalano Mario^{5,7}, Migliore Michele^{6,8}, Caligiore Daniele^{3,4,5}

¹IRCCS Mondino Foundation – National Neurological Institute, Department of Brain and Behavioral Sciences, University of Pavia, Pavia, Italy; ²Neuroimmunology Laboratory and Research Unit, IRCCS Mondino Foundation – National Neurological Institute, Pavia, Italy; ³Computational and Translational Neuroscience Laboratory, Institute of Cognitive Sciences and Technologies, National Research Council (CTNLab-ISTC-CNR), Rome, Italy; ⁴AI2Life s.r.l., Innovative Start-Up, ISTC-CNR Spin-Off, Rome, Italy; ⁵Advanced School in Artificial Intelligence (AS-AI), Rome, Italy; ⁶Dipartimento di Biologia e Biotecnologie, Sapienza Università di Roma, Rome, Italy; ⁷Dipartimento di Biologia, Università degli studi di Roma Tor Vergata Rome, Italy; ⁸Institute of Biophysics, National Research Council, Palermo, Italy

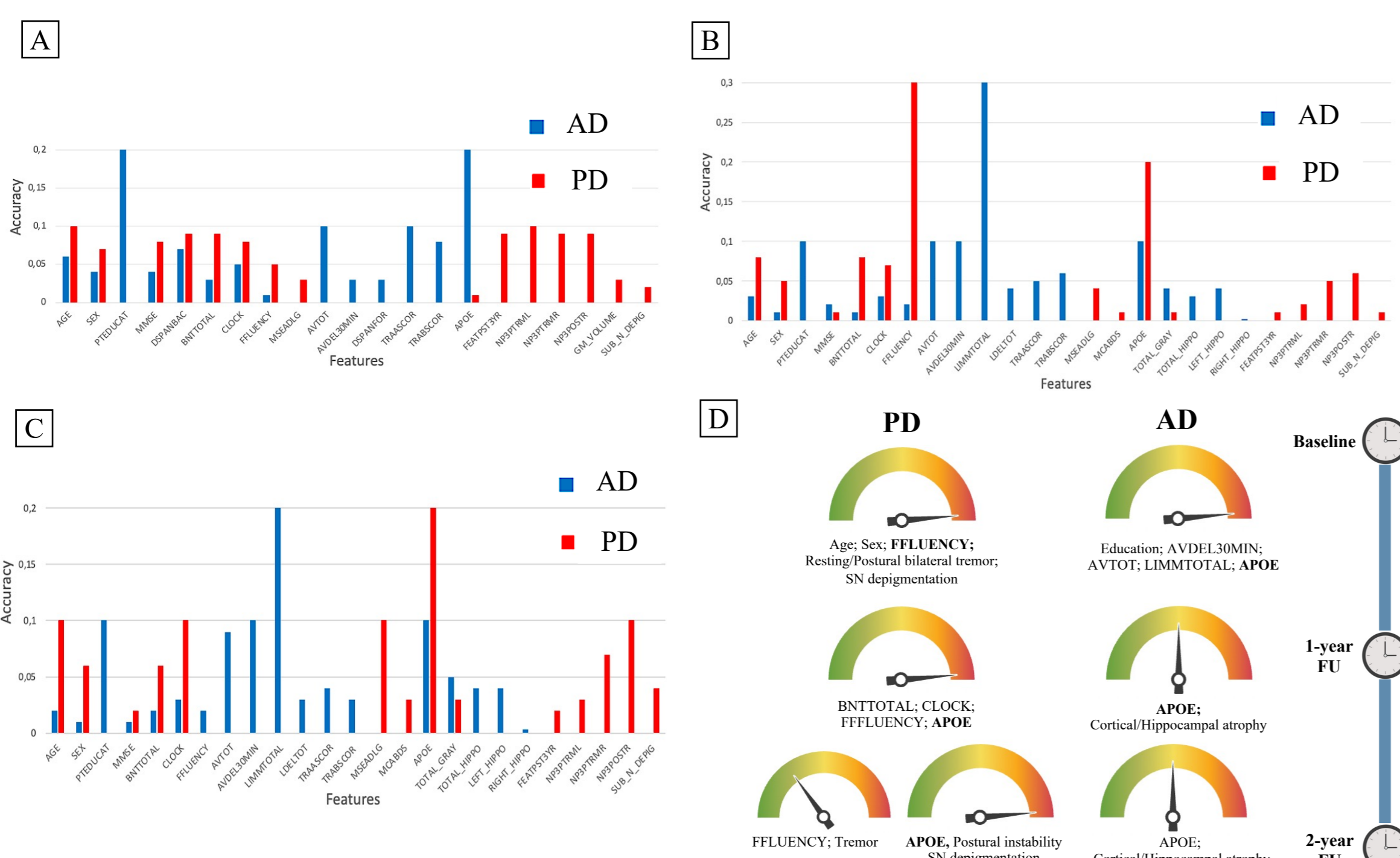
Introduction

A new paradigm called **Neurodegenerative Elderly Syndrome (NES)** conceives Parkinson's disease (PD) and Alzheimer's disease (AD) as different manifestations of a single disease at very early stages [1]. No one has yet obtained a predictive model for both pathologies [2].

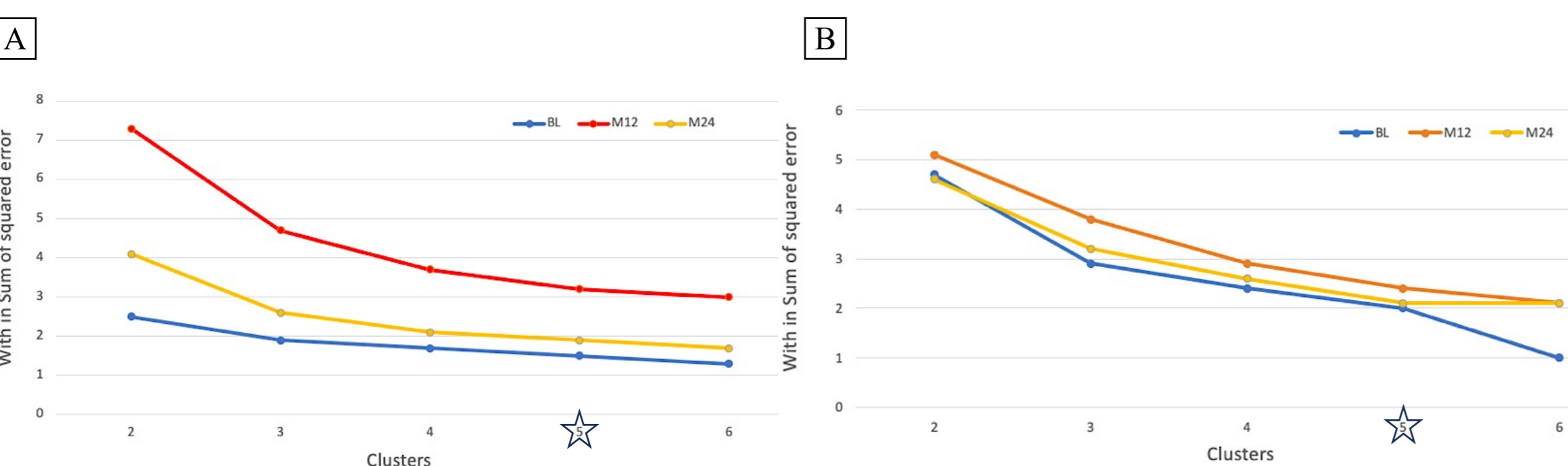
Aims

- To **observe and compare** PDs and ADs common and not features importance at baseline, 12-months and 24-months of follow-up.
- To **predict** disease conversion and **differentiate** or **classify** both pathologies at different time steps.

Results (I)

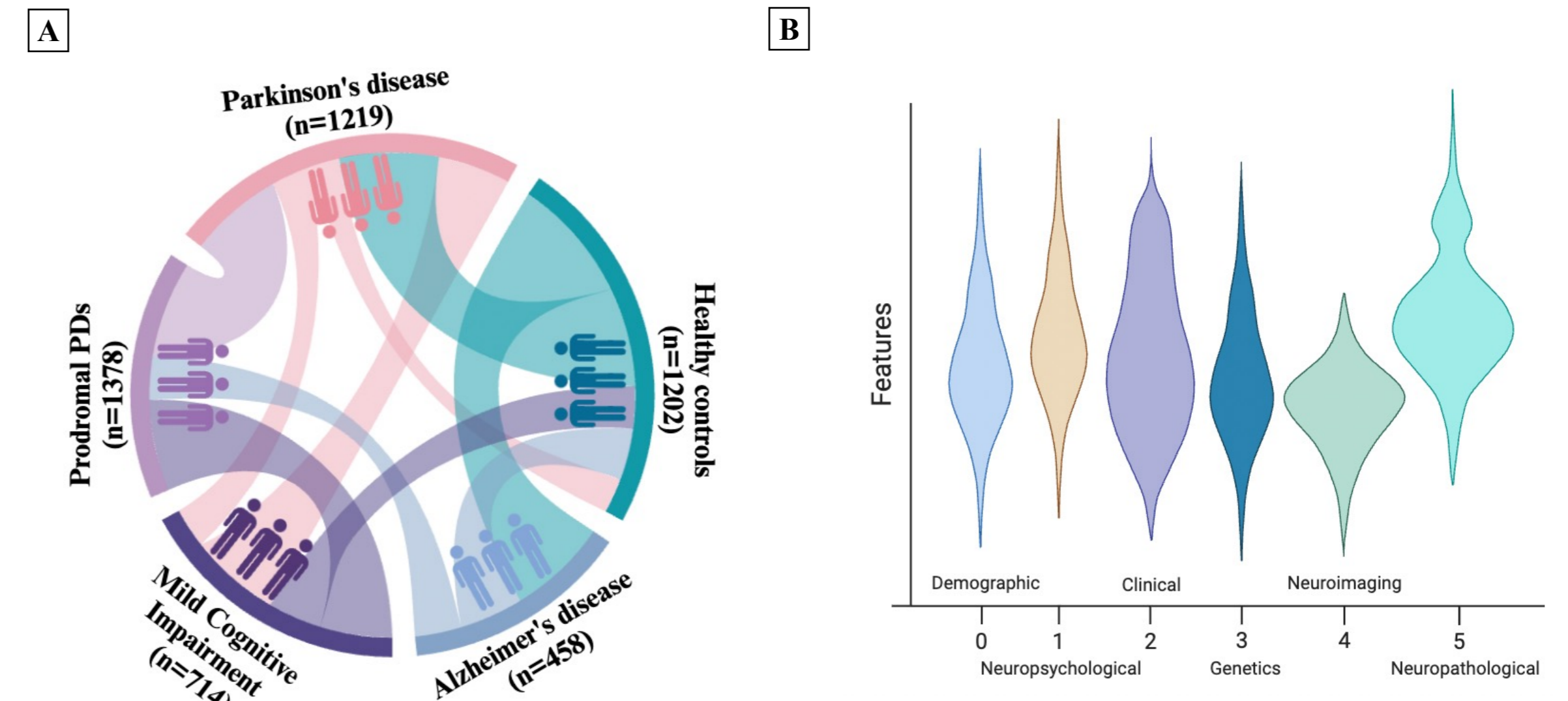


Panel 2. Random-Forest Classifier: Classification accuracy of 96%/86% and Precision of 92%/87% were obtained for PDs/ADs respectively; **Features importance analysis results** at (A) Baseline, (B) m-12 of FU; (C) m-24 of FU considering both PD and AD; (D) Significant differences between PDs and ADs, after features importance analysis, at baseline, m-12 and m-24 of FU respectively (from top to bottom); *p*-value < 0.05.



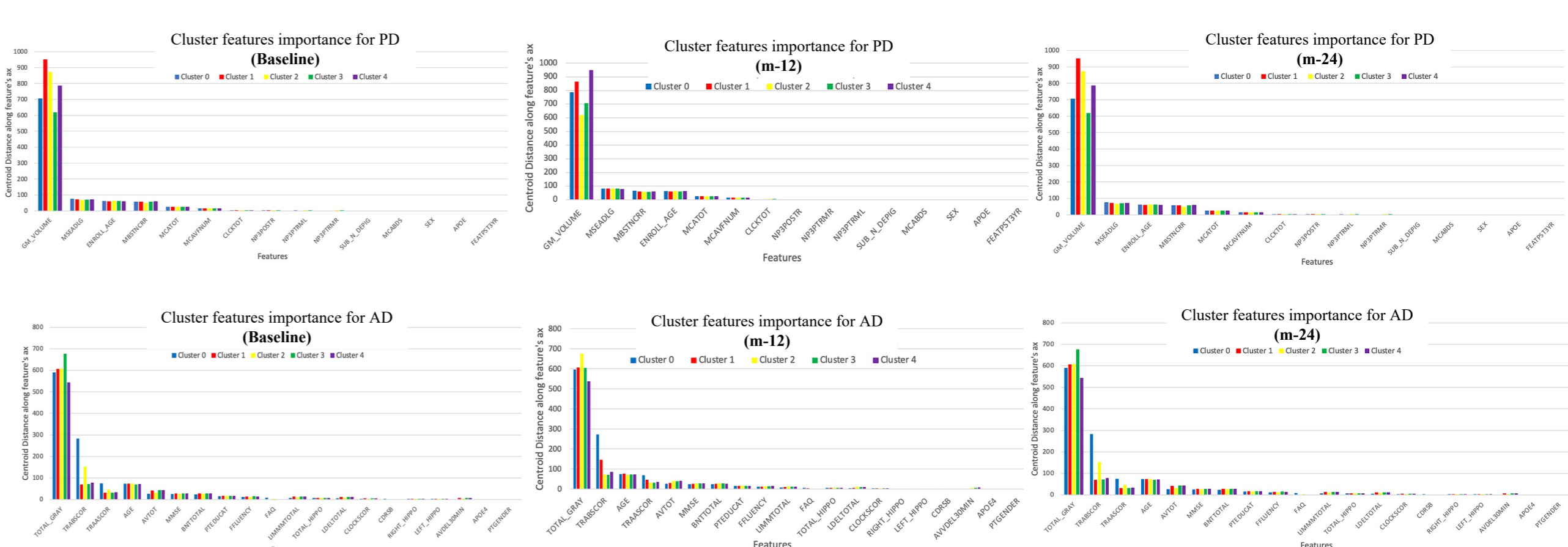
Panel 3. Cluster analysis results: (A) Baseline, m-12 and m-24 for PD; (B) Baseline, m-12 and m-24 for AD. *Y*-axis: Inertia or Sum of squared error; *X*-axis: Numbers of different clusters that could be obtained from the dataset. A total of 5 clusters (Cluster0/Cluster1/Cluster2/Cluster3/Cluster4) were selected.

Materials and Methods

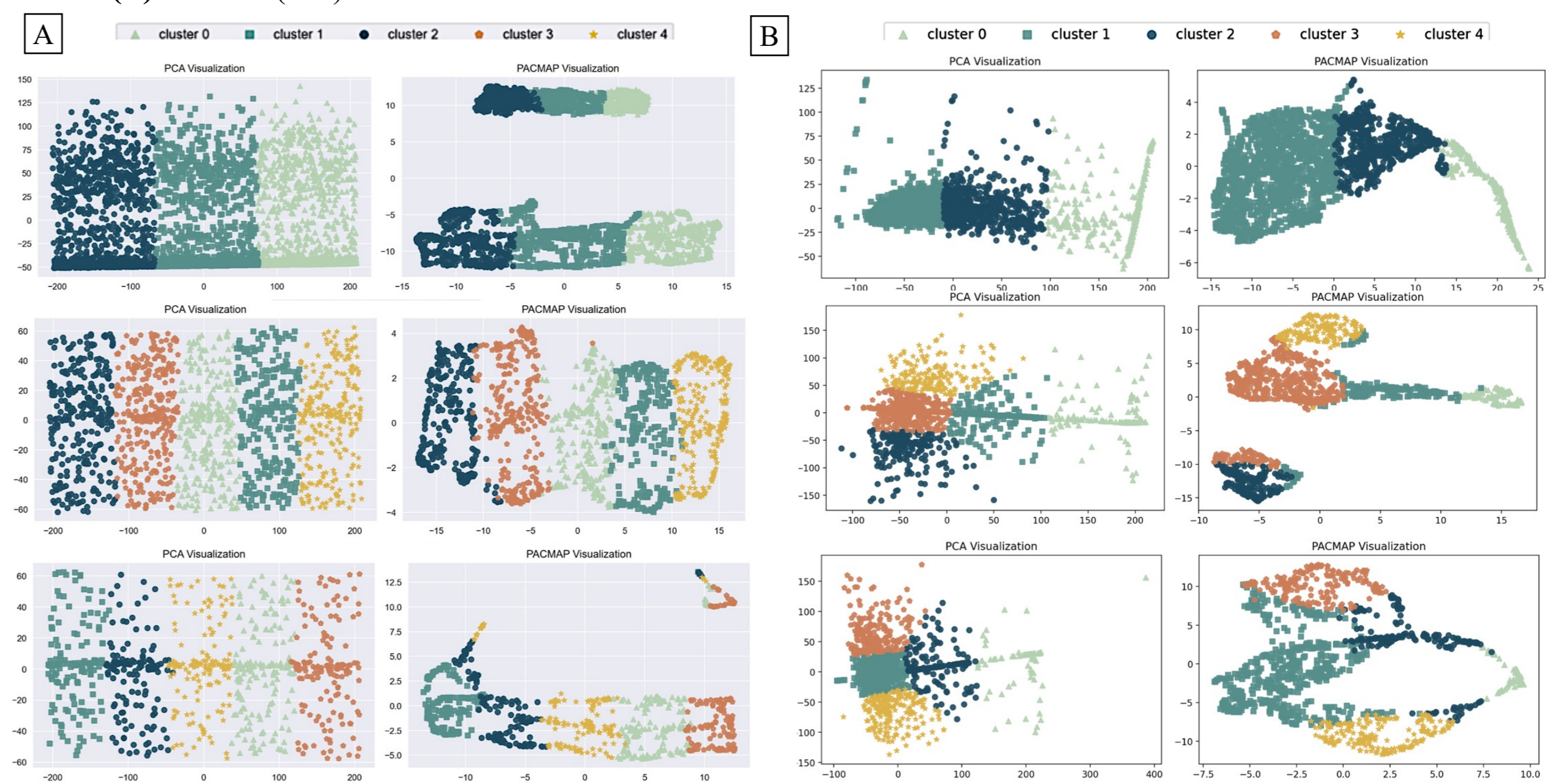


Panel 1. (A) Population was analyzed from Alzheimer's Disease Neuroimaging Initiative (ADNI) and Parkinson's Progression Markers Initiative (PPMI) online databases. Patients and Healthy Controls were aged from 19 to 89, according to female/male differentiation. **(B)** A combination of six groups of features was selected only if possible to match them for both AD and PD: *demographic, cognitive/neuropsychological, clinical, genetic, neuroimaging and neuropathological*. Machine Learning (ML) approaches like **Random-Forest Classifier, Features Importance Analysis and K-means Clustering Analysis** were applied. **Principal Component Analysis (PCA)** and **PCA-MAP** were used for data visualization.

Results (II)



Panel 4. Cluster feature importance analysis results at baseline, m-12 and m-24 for PD (top row of graphs) and for AD (bottom row of graphs), respectively. Generally, **Total gray matter volume** is the best variable for predicting patient classification (PD or AD conversion), followed by **Age** and **MSEADLG** (PD); **TRAA(B)SCORE** (AD).



Panel 5. PCA and PCA-MAP results: The two axes (*x*, *y*) of the graphs represent the two main components (PC1 and PC2) obtained from dimensionality reduction, for both PDs at baseline, m-12 and m-24, from top to bottom (A) and ADs at baseline, m-12 and m-24, from top to bottom (B). **PACMAP** on the other hand uses PCA to reduce the size of the data and then maps them into one two-dimensional (or three-dimensional) space, the two axes of the graph represent the new dimensions obtained from PCA.

Conclusions

We obtained a new **predictive model** able to **compare and classify common features** in ADs and PDs at **baseline** and during **disease progression**. Further studies are needed to validate our findings by testing and refining our predictive models on different multi and monocentric cohorts of patients in a real-life clinical setting.

References

- [1] Caligiore, D., Giocondo, F., & Silveti, M. (2022). The Neurodegenerative Elderly Syndrome (NES) hypothesis: Alzheimer and Parkinson are two faces of the same disease. *IBRO Neuroscience Reports*, 13, 330-343.
[2] Joshi, S., et al. (2010). Classification of Alzheimer's disease and Parkinson's disease by using machine learning and neural network methods. In *2010 Second International Conference on Machine Learning and Computing* (pp. 218-222). IEEE

Contacts

- Antonio Malvaso, MD, Neurology Resident
- IRCCS Mondino, University of Pavia
- antonio.malvaso01@universitadipavia.it

Acknowledgments

We are grateful to S. Torsello, F. Giocondo and M. Silveti for kindly support on selection of filtered data.



Università degli studi di Pavia
Dipartimento di Scienze del Sistema Nervoso e del Comportamento



Parkinson's Progression Markers Initiative