Linking structural neuroimaging with transcriptomic profiling of white matter hyperintensities and microvasculature in Alzheimer's disease

<u>Annie G. Bryant^{1,2}, Sulochan Malla¹, Rojashree Jayakumar¹, Benjamin Woost¹, Nina Wolf¹, </u> Andrew Li¹, Sudeshna Das¹, Susanne J. van Veluw¹, Rachel E. Bennett¹

¹Department of Neurology, Massachusetts General Hospital, Charlestown, MA, USA; ²School of Physics, The University of Sydney, Camperdown, NSW, Australia

Background

White matter hyperintensities (WMHs), visible with T2magnetic resonance weighted imaging (MRI), are common to normative aging and to both disease small vessel and Alzheimer's disease (AD). An estimated 90% of individuals older than 65 develop WMHs WMH burden is [1,2], and associated with inversely cognitive performance in AD [3]. prevalent Despite these associations, the etiology and implications of WMHs for AD pathogenesis remain unclear [4].





Conclusions

We deeply characterize the region-specific WM transcriptome in the context of WMH formation in AD, with a focus on cerebral microvasculature. Our findings underscore diverse associations between the WM transcriptome and both AD pathology and WMH burden, with heterogeneity between frontal vs. occipital WM. Our results align with prior work showing upregulated heat shock protein pathway members in AD brain tissue.

Further info

Check out our preprint (Malla S, Bryant AG, et al., biorXiv 2024) or email me with any follow-up Q's:



annie.bryant@sydney.edu.au

Selected References

- 1. De Leeuw FE, et al. J Neurol Neurosurg Psych (2001) 2. Launer LJ, et al. Neuroepidemiology (2006) 3. Garnier-Crussard A, et al. Alzheimers Dement (2022) 4. Wardlaw JM, et al. J Am Heart Assoc (2015) 5. Mitroi DN, et al. J Cell Molec Med (2022)
- 6. Bryant AG, et al. J Neurosci (2023)

Acknowledgements

This work was supported by the NIH National Institute on Aging and the Massachusetts Alzheimer's Disease Research Center. We thank the donors and their families who have generously contributed to this invaluable resource.





MASSACHUSETTS Alzheimer's Disease **Research Center**