Supplementary Materials for "Volumetric GWAS of Medial Temporal Lobe Structures Identifies an *ERC1* Locus using ADNI High-resolution T2-weighted MRI Data" by Shan Cong, Xiaohui Yao, Zhi Huang, Shannon L. Risacher, Kwangsik Nho, Andrew J. Saykin, Li Shen; UK Brain Expression Consortium, for the Alzheimer's Disease Neuroimaging Initiative

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UKBEC Acknowledgements

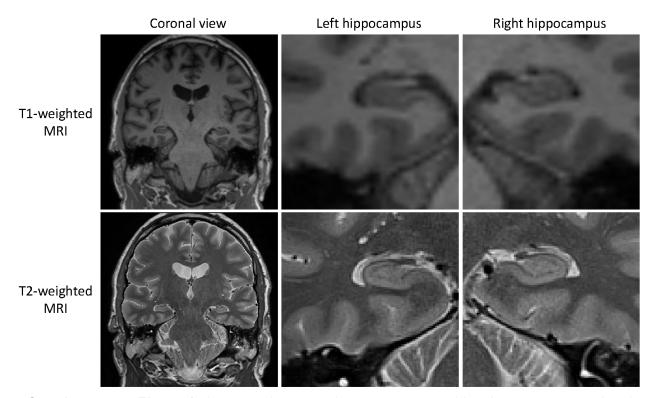
UK Brain Expression Consortium (UKBEC) members and affiliations Mina Ryten^{1,2}
Michael E Weale¹
John Hardy²
Karishma D'Sa^{1,2}
Adaikalavan Ramasamy^{1,2,3}
Daniah Trabzuni^{2,4}

Sebastian Guelfi²
Juan A. Botia²
Jana Vandrovcova²
Colin Smith⁵
Robert Walker⁵

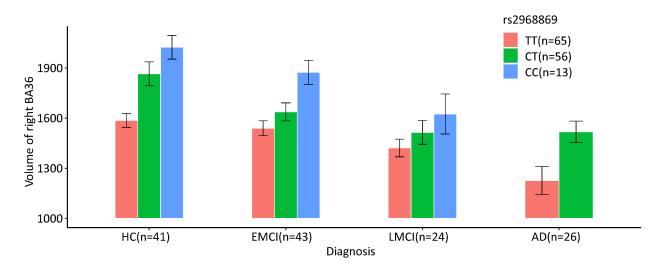
- ¹ Department of Medical & Molecular Genetics, King's College London, Guy's Hospital, London, UK
- ² Reta Lila Weston Research Laboratories, Department of Molecular Neuroscience, University College London (UCL) Institute of Neurology, London, UK
- ³ Jenner Institute, University of Oxford, Oxford, UK
- ⁴ Department of Genetics, King Faisal Specialist Hospital and Research Centre, Riyadh, Saudi Arabia
- ⁵ Department of Neuropathology, MRC Sudden Death Brain Bank Project, University of Edinburgh, Edinburgh, UK

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Supplementary Figure 1. An example comparison on a same subject between conventional 3T T1-weighted MRI (top panel) and 3T T2-weighted high-resolution MRI (bottom panel). The first column shows one coronal slice from each MRI scan, the second and third columns show a zoom-in view of left and right hippocampi respectively.



Supplementary Figure 2. Effect of *ERC1* rs2968869 on the right BA36 volume stratified by diagnostic group. The minor allele (C) of rs2968869 is associated with increased R-BA36 volume for all diagnostic groups. Of note, the R-BA36 volume measures are adjusted by removing the effects of age, gender, education, Intracranial Volume (ICV), and the top four principal components from population stratification analysis.

Supplementary Table 1. Participant characteristics in genetic association study of VBM right hippocampal gray matter density.

| Diagnosis | HC | SMC | EMCI | LMCI | AD | P-value |
|------------------------------|------------|------------|------------|------------|------------|----------|
| Number | 298 | 89 | 237 | 407 | 353 | - |
| Gender (M/F) | 159/139 | 36/53 | 133/104 | 253/154 | 200/153 | 3.23E-03 |
| Age (mean±std) | 76.68±6.05 | 72.35±5.68 | 71.69±7.30 | 75.21±7.98 | 75.81±7.67 | 2.53E-16 |
| Education (mean±std) | 16.23±2.67 | 16.76±2.62 | 15.90±2.57 | 15.91±2.88 | 15.32±3.03 | 1.92E-05 |
| APOE ε4 positive | 27.52% | 33.71% | 42.37% | 53.56% | 66.01% | 3.89E-23 |
| Left hippocampus (mean±std) | 0.48±0.05 | 0.50±0.04 | 0.48±0.05 | 0.45±0.06 | 0.41±0.07 | 6.70E-80 |
| Right hippocampus (mean±std) | 0.45±0.05 | 0.48±0.04 | 0.46±0.05 | 0.42±0.06 | 0.38±0.06 | 2.15E-90 |

Note: p-values were assessed for significant differences among diagnosis groups and were computed using one-way ANOVA (except for gender using chi-square test). The p-values < 0.05 are shown **in bold**. HC=Healthy Control; SMC, significant memory concern; EMCI=Early Mild Cognitive Complaint; LMCI=Late Mild Cognitive Complaint; AD=Alzheimer's disease.

Supplementary Table 2. Participant characteristics in genetic association analysis of FDG right hippocampal glucose metabolism.

| Diagnosis | HC | SMC | EMCI | LMCI | AD | P-value |
|------------------------------|------------|------------|------------|------------|------------|----------|
| Number | 195 | 91 | 241 | 169 | 169 | - |
| Gender (M/F) | 101/94 | 38/53 | 134/107 | 105/64 | 100/69 | 1.78E-02 |
| Age (mean±std) | 76.54±6.64 | 72.54±5.56 | 71.70±7.31 | 74.73±8.32 | 75.39±7.75 | 1.48E-11 |
| Education (mean±std) | 16.29±2.72 | 16.86±2.58 | 15.87±2.58 | 16.16±2.81 | 15.88±2.77 | 2.60E-02 |
| APOE ε4 present | 27.18% | 32.97% | 42.92% | 50.30% | 64.50% | 5.88E-12 |
| Left hippocampus (mean±std) | 1.06±0.09 | 1.09±0.09 | 1.06±0.09 | 1.01±0.11 | 0.92±0.12 | 1.15E-50 |
| Right hippocampus (mean±std) | 1.05±0.10 | 1.07±0.08 | 1.05±0.08 | 1.00±0.11 | 0.93±0.12 | 6.82E-40 |

Note: p-values were assessed for significant differences among diagnosis groups and were computed using one-way ANOVA (except for gender using chi-square test). The p-values < 0.05 are shown **in bold**. HC=Healthy Control; SMC, significant memory concern; EMCI=Early Mild Cognitive Complaint; LMCI=Late Mild Cognitive Complaint; AD=Alzheimer's disease.