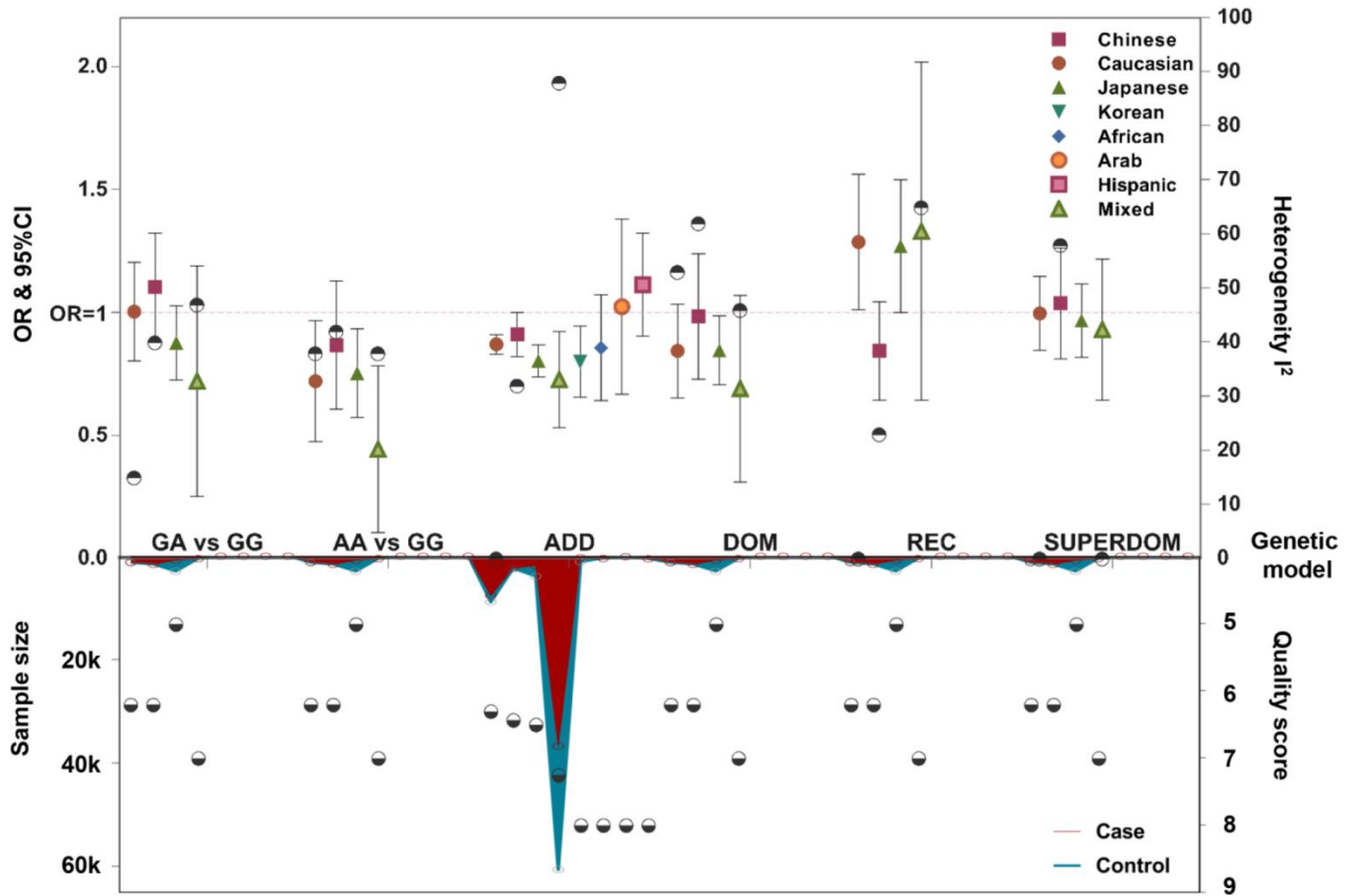
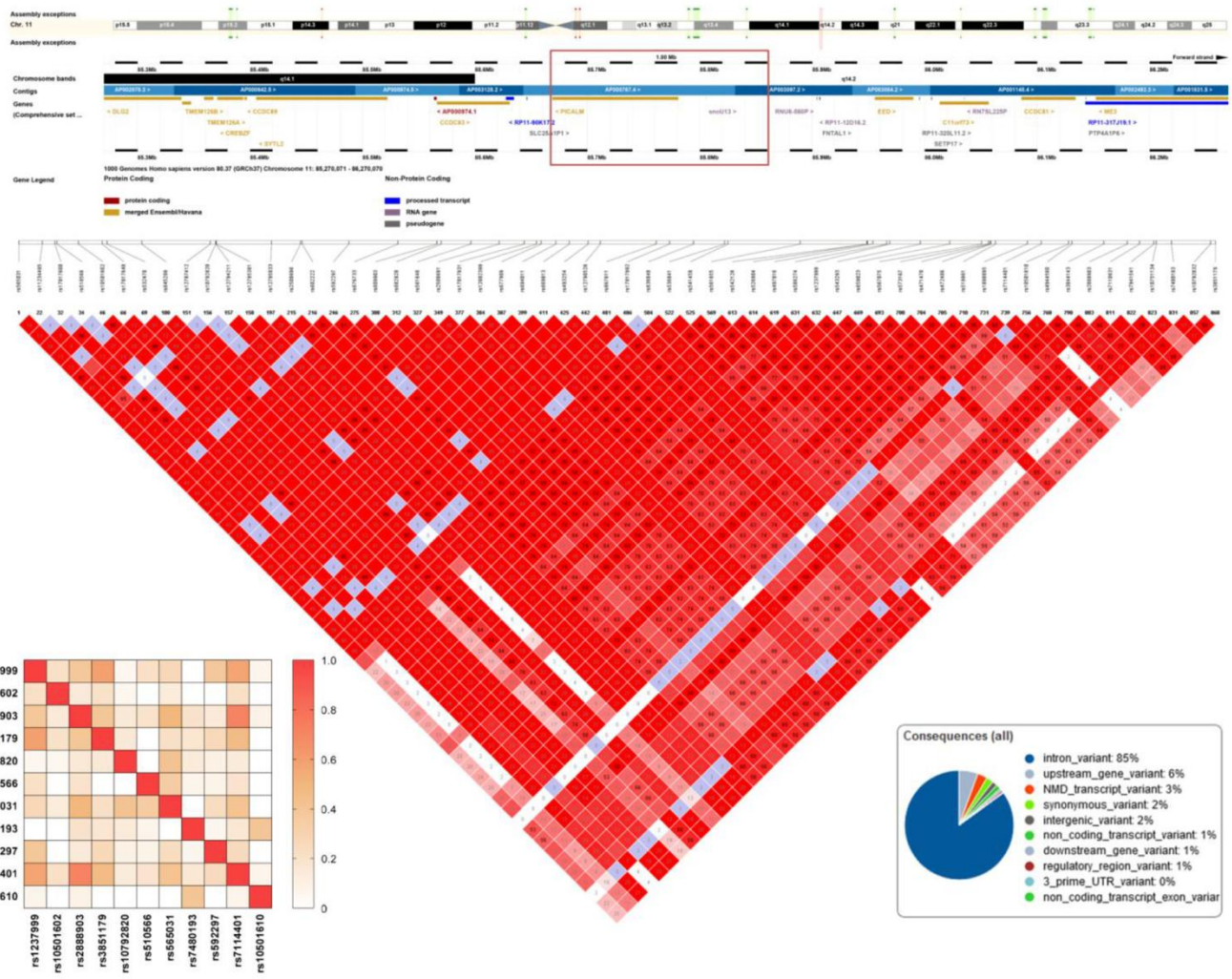


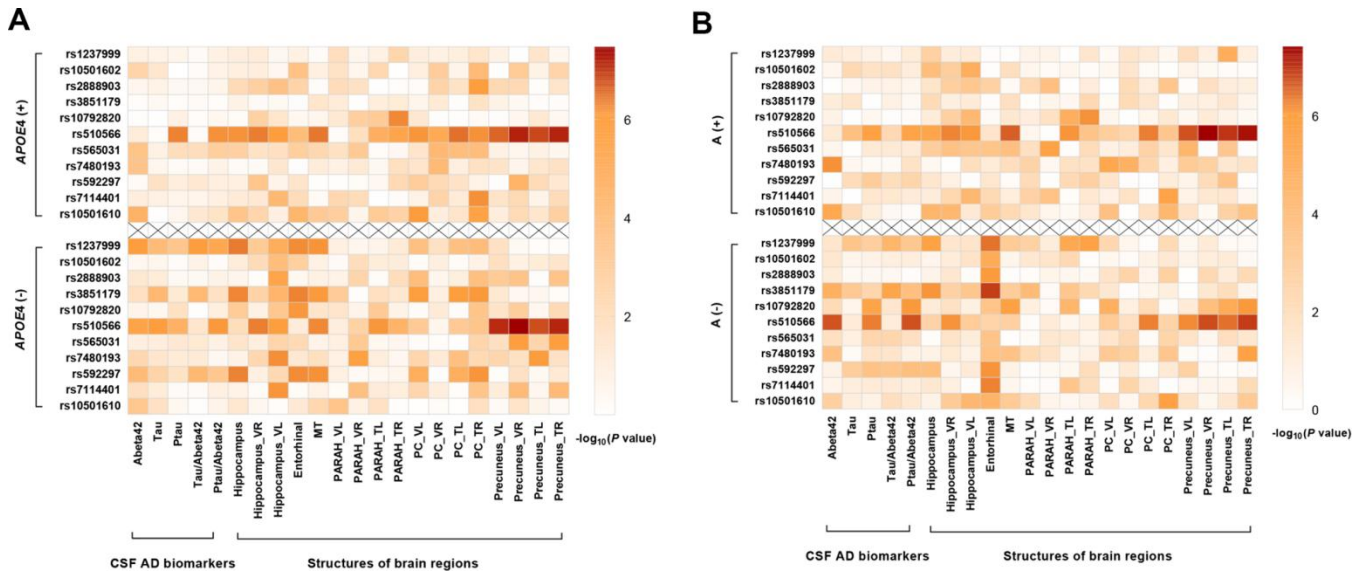
SUPPLEMENTARY FIGURES



Supplementary Figure 1. Association of rs3851179 with AD risk with different genetic models in different ethnicities. rs3851179 (allele A) was associated with lower AD risk, with the effect size ranging from 9% to 29% in Caucasian ($I^2 = 38\%$), Chinese ($I^2 = 42\%$), Japanese, Korean, and population of mixed races ($I^2 = 38\%$), but not in other races. The association remained significant in Caucasian population for other genetic models, such as REC (AA vs GG+GA), and genotype (AA vs GG).



Supplementary Figure 2. Linkage Disequilibrium Analysis revealed eleven tag SNPs. Eleven SNPs were selected by LD analysis, such that these 11 loci could independently capture 100% of all alleles at $r^2 \geq 0.8$.



Supplementary Figure 3. Association results of *PICALM* tag loci with AD CSF biomarkers and feature neurodegeneration, stratified by *APOE4* and amyloid status. The association for rs510566 was not influenced by subgrouping according to *APOE4*, but remained significant only in A (-) subgroup. The associations with specific loci showed significant trends in *APOE4* (-) subgroup, including rs1237999 ($p = 0.002$ for HIPPO, $p = 0.009$ for ENTOR, and $p = 0.018$ for MT), rs592297 ($p = 0.003$ for HIPPO, $p = 0.008$ for ENTOR, and $p = 0.014$ for MT), and rs3851179 ($p = 0.005$ for HIPPO, $p = 0.0037$ for ENTOR, $p < 0.05$ for MT, and $p < 0.05$ for PC).