

**Supplementary Table 4.** Linear predictions of average marginal effects of hePRS-IR and mePRS-IR regions in brain for IGF-1, apolipoprotein A, and triglycerides according to diabetes status in men

Variable	Diabetes status	b (95% CI)	P for ePRS	P for interaction
IGF-1				
mePRS-IR	Normal	-14.6 (-24.1 to -5.1)	0.003	0.01
mePRS-IR	Prediabetes	11 (-2.3 to 24.3)	0.105	
mePRS-IR	Diabetes	-2.8 (-18 to 12.3)	0.712	
Apolipoprotein A				
hePRS-IR	Normal	0.03 (0 to 0.05)	0.07	0.044
hePRS-IR	Prediabetes	-0.02 (-0.05 to 0.004)	0.095	
hePRS-IR	Diabetes	0.01 (-0.03 to 0.05)	0.707	
Triglycerides				
mePRS-IR	Normal	0.02 (-0.04 to 0.08)	0.486	0.056
mePRS-IR	Prediabetes	-0.18 (-0.33 to -0.03)	0.022	
mePRS-IR	Diabetes	0.04 (-0.14 to 0.22)	0.676	

hePRS-IR, polygenic risk score for the hippocampal-insulin receptor; mePRS-IR, polygenic risk score for the mesocorticolimbic-insulin receptor; IGF-1, insulin growth factor 1; CI, confidence interval; ePRS, polygenic risk score for the gene network.