S Table 1. Effect of GWAS sample size on PRS performance. Difference in R<sup>2</sup> for different groupings of individuals between the UKBB GWAS down-sampled to that of the sample size used in GIANT (N=226,960), and the full UKBB European sample size (N=377,921). Test African ancestry individuals and 5,000 random European ancestry UKBB individuals were used as LD panels for all African and European ancestry test sets, respectively.

Age group	Ancestry	N	PRS R <sup>2</sup> UKBB Down- sampled sample size	PRS R <sup>2</sup> UKBB Full sample size	P-value
Adults	European	55418	0.0582	0.0734	<4.94x10 <sup>-324</sup>
Teenagers		3114	0.0588	0.0667	3.75x10 <sup>-9</sup>
Children		5943	0.0274	0.0347	6.51x10 <sup>-47</sup>
Adults		5912	0.0140	0.0197	1.85x10 <sup>-46</sup>
Teenagers	African	1606	0.0179	0.0274	1.37x10 <sup>-29</sup>
Children		3668	0.00612	0.00715	4.55x10 <sup>-9</sup>

S Table 2. Selected Phecodes in eMERGE which were tested for their association to the  $\mathsf{PRS}_\mathsf{BMI}$ .

Phecode	Description	Cases/Controls (Adults)		
		EUR	AFR	
401.1	Essential hypertension	38181/17166	4317/1588	
272.1	Hyperlipidemia	32823/22524	2625/3280	
272.11	Hypercholesterolemia	22537/32810	1912/3993	
278.1	Obesity (positive control)	16348/38999	2282/3623	
411.4	Coronary atherosclerosis	18548/36799	1449/4456	
250.2	Type 2 diabetes	16342/39005	2525/3380	
296.2	Depression	14651/40696	1630/4275	
208	Benign neoplasm of colon	16254/39093	1205/4700	
562.1	Diverticulosis	15656/39691	1068/4837	
495	Asthma	9391/45956	1253/4652	

S Figure 1. Differences in PRS R<sup>2</sup> across different age groups (the best performing LD panel was used for each run, p-values above each bar). We consistently observed more significant differences in PRS R<sup>2</sup> in children versus adults and teenagers. The downsampled UKBB summary statistics were used for these comparisons.

