

- 678 **Fig. S1. Types of DXA images acquired from the UKB.** (Left) Image of patient imaged on
- 679 white background. (Right) Image of patient imaged on black background. Sizes of images are
- true to scale.



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742 Fig. S2. A comparison of HRNet and ResNet deep learning architectures. (A) High-

743 Resolution Network (HRNet) architecture maintains parallel high to low resolution subnetworks.

744 (B) Simple Baseline deep learning architecture (ResNet) which relies on a high-to-low and low-

to-high framework. Both images are taken directly from Sun et al., (35) to illustrate the

- architectural differences between HRNet and a standard architecture for this prediction task.
- 747



Pixel height: 665 px Pixel height: 720 px

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**Fig. S3. DXA images from the UKB that have undergone different image scaling.** Example

of two individuals who were measured to be the same height in the FID 50 in the UKB (overall

height) but pixel-based measurements of one image were considerably smaller than the other due

- to image scaling/resolution differences.
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778 Fig. S4. A linear regression of image-measured height against UKB-measured height. For

each image pixel-ratio, we regressed height measured in the UKB with height we calculated in

pixels from the DXA scan. This provided a conversion from pixels to cm that we used as a

781 normalization factor to correct for differences in resolution.



- 805 Fig. S5. Examples of individuals who were outliers on our measurement and were removed
- 806 **from analysis**. (Left) Individual with femur deformity and metal implants. (Right) Individual
- 807 with missing forearm.



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927 Fig. S6. A heatmap comparison of genotype and phenotype correlations between ratios and

928 residuals. (A) Matrix of genotype and phenotype correlation with each phenotype computed as a

929 ratio of height. (B) Matrix of genotype and phenotype correlation with each phenotype computed

930 by regressing the phenotype with height and then obtaining residuals.





Fig. S7. Correlation of genotype and phenotype correlations across skeletal traits, computedusing ratios with height and second residualizing for height



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946 Fig. S8. Comparison of effect estimates of independent genome-wide significant SNPs

947 **across different phenotypes.** Effect estimates of genome-wide significant SNPs for each

948 phenotype (p < 5e-08) showing same effect directionality for skeletal proportions and raw

949 measurements.



- 1126
- 1127 Fig. S9. Heatmap of genetic correlations and LDSC cross-trait intercepts across skeletal
- 1128 proportion phenotypes within odd-numbered chromosomes
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Fig. S10. Scatterplot of GCTA and LDSC genetic correlation estimates across skeletal ratios.



1133 Fig. S11. Screeplots of PCA and difference in PCA from LDSC Parallel Analysis









- 1139 **diseases and skeletal endophenotypes**. Red lines represent best fitting regression lines. Blue
- 1140 dashed lines represent perfect fit (Observed effects = Model-implied effects). Labeled traits are
- 1141 outliers detected based on standardized differences between the observed and the common factor
- 1142 model-implied effects for the skeletal traits > 2.
- 1143
- 1144







- 1147 and FinnGen. Red lines represent best fitting regression lines. Blue dashed lines represent
- 1148 perfect correspondence (intercept = 0, slope = 1).
- 1149

 $\begin{array}{l} \mbox{Model D (traits residualized by height)}\\ \chi^2 \ (13) = 197.76, \ p < 0.001 \\ \mbox{AIC} = 227.762 \\ \ CFI = 0.93 \\ \ SRMR = 0.06 \end{array}$ 



- 1150
- 1151 Fig. S15. Confirmatory Factor Model D applied to residuals. Preferred model D fully
- 1152 standardized parameter estimates fitted on height-residualized skeletal traits as well as excluding
- 1153 overall arms and leg length
- 1154





1183 Fig. S16. Genetic correlations between males and females, estimated using bi-variate LD Score



1185 estimated in the sample with both sexes combined (x-axis) for all traits



## 1186

Polygenic Score, Estimated in Males (SDs from mean)



- 1188 polygenic score estimated in an independent sample of females. Points show mean values in one
- 1189 decile of the polygenic score; the fitted line and associated effect estimate and  $R^2$  correspond to
- 1190 regressions on the raw, non-binned data.







accelerated regions (HARs) and phenotype-associated genes. Blue bars are background

1362 distributions generated from 5,000 simulations of matched element sets. An example is shown

1363 here for skin pigmentation.