#### SUPPLEMENTAL MATERIAL

#### **FIGURE LEGENDS**

Supplemental Figure 1. Plot of the principal component analysis of CARDIA AA cases along with individuals from the 1,000 Genomes Project.

AMR-Ad Mixed American; SAS-South Asian; EAS-East Asian; EUR-European; ACB-African Caribbeans in Barbados; ASW-Americans of African Ancestry in SW USA; AFR-African.

Supplemental Figure 2. Plot of the principal component analysis of BioVU AA cases along with individuals from the 1,000 Genomes Project.

AMR-Ad Mixed American; SAS-South Asian; EAS-East Asian; EUR-European; ACB-African Caribbeans in Barbados; ASW-Americans of African Ancestry in SW USA; AFR-African.

Supplemental Figure 3. Plot of the principal component analysis of CARDIA AA cases along with individuals from the 1,000 Genomes Project.

AMR-Ad Mixed American; SAS-South Asian; EAS-East Asian; AFR-African; EUR-European.

Supplemental Figure 4. Plot of the principal component analysis of BioVU AA cases along with individuals from the 1,000 Genomes Project.

AMR-Ad Mixed American; SAS-South Asian; EAS-East Asian; AFR-African; EUR-European. Supplemental Figure 5. The QQ plot for the meta-analysis results from the GWAS for

number of fibroids (single vs multiple) for BioVU and CARDIA AAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 6. The QQ plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed largest dimension of all fibroids for BioVU and CARDIA AAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

### Supplemental Figure 7. The QQ plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed fibroid volume for BioVU and CARDIA AAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

#### Supplemental Figure 8. The QQ plot for the meta-analysis results from the GWAS for number of fibroids (single vs multiple) for BioVU and CARDIA EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

#### Supplemental Figure 9. The QQ plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed largest dimension of all fibroids for BioVU and CARDIA EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

## Supplemental Figure 10. The QQ plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed fibroid volume for BioVU and CARDIA EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 11. The QQ plot for the transethnic meta-analysis results from the GWAS for number of fibroids (single vs multiple) for BioVU and CARDIA AAs and EAs. The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 12. The QQ plot for the transethnic meta-analysis results from the GWAS for log<sub>10</sub> transformed largest dimension of all fibroids for BioVU and CARDIA AAs and EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

#### Supplemental Figure 13. The QQ plot for the transethnic meta-analysis results from the GWAS for log<sub>10</sub> transformed fibroid volume for BioVU and CARDIA AAs and EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

#### Supplemental Figure 14. The manhattan plot for the meta-analysis results from the GWAS for number of fibroids (single vs multiple) for BioVU and CARDIA AAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

### Supplemental Figure 15. The manhattan plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed largest dimension of all fibroids for BioVU and CARDIA AAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

#### Supplemental Figure 16. The manhattan plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed fibroid volume for BioVU and CARDIA AAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

### Supplemental Figure 17. The manhattan plot for the meta-analysis results from the GWAS for number of fibroids (single vs multiple) for BioVU and CARDIA EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

#### Supplemental Figure 18. The manhattan plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed largest dimension of all fibroids for BioVU and CARDIA EAs. The regression analyses were adjusted for age, BMI, and five principal components. Markers

with a MAF of at least 5% were used in the analyses.

Supplemental Figure 19. The manhattan plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed fibroid volume for BioVU and CARDIA EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 20. The manhattan plot for the transethnic meta-analysis results from the GWAS for number of fibroids (single vs multiple) for BioVU and CARDIA AAs and EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 21. The manhattan plot for the transethnic meta-analysis results from the GWAS for log<sub>10</sub> transformed largest dimension of all fibroids for BioVU and CARDIA AAs and EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 22. The manhattan plot for the transethnic meta-analysis results from the GWAS for log<sub>10</sub> transformed fibroid volume for BioVU and CARDIA AAs and EAs. The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

#### Supplemental Figure 23. The MetaXcan plot for the meta-analysis results from the GWAS for number of fibroids (single vs multiple) for BioVU and CARDIA AAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 24. The MetaXcan plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed largest dimension of all fibroids for BioVU and CARDIA AAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

#### Supplemental Figure 25. The MetaXcan plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed fibroid volume for BioVU and CARDIA AAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 26. The MetaXcan plot for the meta-analysis results from the GWAS for number of fibroids (single vs multiple) for BioVU and CARDIA EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 27. The MetaXcan plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed largest dimension of all fibroids for BioVU and CARDIA EAs. The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 28. The MetaXcan plot for the meta-analysis results from the GWAS for log<sub>10</sub> transformed fibroid volume for BioVU and CARDIA EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 29. The MetaXcan plot for the transethnic meta-analysis results from the GWAS for number of fibroids (single vs multiple) for BioVU and CARDIA AAs and EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 30. The MetaXcan plot for the transethnic meta-analysis results from the GWAS for log<sub>10</sub> transformed largest dimension of all fibroids for BioVU and CARDIA AAs and EAs.

The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Supplemental Figure 31. The MetaXcan plot for the transethnic meta-analysis results from

the GWAS for log<sub>10</sub> transformed fibroid volume for BioVU and CARDIA AAs and EAs. The regression analyses were adjusted for age, BMI, and five principal components. Markers with a MAF of at least 5% were used in the analyses.

Figure 32. Regional association plot of the most significant marker from the EA metaanalyses.

The index SNP from this regional plot is rs141745233 from the EA meta-analysis of fibroid volume. European individuals from the 1000 genomes was used to estimate linkage disequilibrium (LD) for this plot. The X-axis represents the genomic position along each chromosome in megabases (Mb). The Y-axis represents both the  $-\log_{10}$  P-values for each SNP as

well as the recombination rate. The color of each SNP represents the strength of the correlation  $(r^2)$  to the index SNP (purple). Nearby genes are listed below the regional plots.

# Figure 33. Regional association plot of the most significant marker from the transethnic meta-analyses using African individuals from the 1000 genomes to estimate linkage disequilibrium (LD).

(a) The index SNP from this regional plot is rs6605005 from the transethnic meta-analysis of fibroid volume. The X-axis represents the genomic position along each chromosome in megabases (Mb). The Y-axis represents both the  $-\log_{10}$  P-values for each SNP as well as the recombination rate. The color of each SNP represents the strength of the correlation (r<sup>2</sup>) to the index SNP (purple). Nearby genes are listed below the regional plots.

Figure 34. Regional association plot of the most significant marker from the transethnic meta-analyses using European individuals from the 1000 genomes to estimate linkage disequilibrium (LD).

The index SNP from this regional plot is rs6605005 from the transethnic meta-analysis of fibroid volume. The X-axis represents the genomic position along each chromosome in megabases (Mb). The Y-axis represents both the  $-\log_{10}$  P-values for each SNP as well as the recombination rate. The color of each SNP represents the strength of the correlation (r<sup>2</sup>) to the index SNP (purple). Nearby genes are listed below the regional plots.

Cohort	Race	Genotyping Platform	Post-QC, Pre- imputation SNPs	Outcome	Post Imputation SNPs
BioVU	AA	Affymetrix Axiom	1,345,099	Number	9,530,809
		Biobank Array & Axiom		Max Dimension	9,526, 536
		World Array 3 Platform		Volume	9,540,688
	EA	Affymetrix Axiom	602,522	Number	6,942,486
		Biobank Array		Max Dimension	6,945,186
				Volume	6,949,297
CARDIA	AA	Affymetrix 6.0 Array	784,270	Number	9,627,525
				Max Dimension	9,596,033
				Volume	9,597,202
	EA	Affymetrix 6.0 Array	562,358	Number	6,938,376
				Max Dimension	6,953,394
				Volume	6,956,705

Supplemental Table 1. Quality control (QC) details for each study

AA - African American; EA - European American

Number										
rs35322806 in 4q32.1										
Cohort	OR [95% CI]	<b>P-value</b>	EAF	Info						
BioVU AA	0.49 [0.34, 0.70]	9.85x10 <sup>-5</sup>	0.21	0.97						
CARDIA AA	0.31 [0.17, 0.59]	$3.31 \times 10^{-4}$	0.20	0.99						
rs735658 in 17q25.3										
Cohort	OR [95% CI]	<b>P-value</b>	EAF	Info						
BioVU AA	0.39 [0.23, 0.67]	$7.23 \times 10^{-4}$	0.09	0.83						
CARDIA AA	0.09 [0.03, 0.27]	$1.29 \times 10^{-5}$	0.08	0.67						
	rs7625080 in 3 <sub>1</sub>	p <b>26.1</b>								
Cohort	OR [95% CI]	<b>P-value</b>	EAF	Info						
BioVU AA	0.52 [0.38, 0.69]	1.11x10 <sup>-5</sup>	0.34	0.98						
CARDIA AA	0.56 [0.34, 0.93]	0.0257	0.34	0.97						
	rs62203867 in 20	)p12.3								
Cohort	OR [95% CI]	<b>P-value</b>	EAF	Info						
BioVU AA	0.48 [0.34, 0.67]	$1.82 \times 10^{-5}$	0.24	0.97						
CARDIA AA	0.41 [0.20, 0.85]	0.0169	0.22	0.81						
	Max Dimens	ion								
	rs57542984 in	5q34								
Cohort	Beta [95% CI]	<b>P-value</b>	EAF	Info						
BioVU AA	0.13 [0.08, 0.19]	1.66x10 <sup>-6</sup>	0.80	0.91						
CARDIA AA	0.12 [0.05, 0.18]	$3.37 \times 10^{-4}$	0.83	0.91						
	rs62484733 in 7	q22.3								
Cohort	Beta [95% CI]	<b>P-value</b>	EAF	Info						
BioVU AA	0.11 [0.04, 0.18]	$1.94 \times 10^{-3}$	0.10	0.92						
CARDIA AA	0.17 [0.09, 0.25]	$3.14 \times 10^{-5}$	0.09	0.88						
	rs13183849 in 5	q14.3								
Cohort	Beta [95% CI]	<b>P-value</b>	EAF	Info						
BioVU AA	-0.10 [-0.15, -0.06]	1.59x10 <sup>-5</sup>	0.31	0.97						
CARDIA AA	-0.07 [-0.12, -0.02]	0.0105	0.27	0.99						
	Volume									
rs6938199 in 6q22.33										
Cohort	Beta [95% CI]	<b>P-value</b>	EAF	Info						
BioVU AA	-0.35 [-0.55, -0.14]	$8.39 \times 10^{-4}$	0.10	1.00						
CARDIA AA	-0.49 [-0.74, -0.25]	8.86x10 <sup>-5</sup>	0.11	1.00						

Supplemental Table 2. Summary of cohort-specific single SNP signals from the metaanalyses between BioVU and CARDIA blacks for each fibroid outcome

OR-odds ratio; CI-confidence interval; EAF-effect allele frequency.

Number									
SNP	Region	BP	Genes	A2/A1	EAF	OR [95% CI]	<b>P-value</b>	Het. P	Dir.
rs35071394	14q21.1	41,837,615	FBXO33//LRFN5	C/CCT	0.92	2.93 [1.92, 4.48]	$6.56 \times 10^{-7}$	0.308	++
rs1392505	16q21	59,746,130	GOT2//CDH8	A/G	0.80	1.89 [1.46,2.44]	$1.00 \times 10^{-6}$	0.291	++
				Volume					
SNP	Region	BP	Genes	A2/A1	EAF	Beta [95% CI]	<b>P-value</b>	Het. P	Dir.
rs141745233	7q36.1	148,269,774	CNTNAP2//CUL1	T/TG	0.09	-0.46 [-0.64, -0.29]	$3.97 \times 10^{-7}$	0.227	
rs17506409	5q22.1	110,089,908	SLC25A46	G/A	0.08	0.46 [0.28, 0.64]	8.10x10 <sup>-7</sup>	0.339	++

Supplemental Table 3. Summary of most significant SNPs from the meta-analyses between BioVU and CARDIA EAs for each fibroid outcome

SNP-single nucleotide polymorphism; BP-base pairs; A2-effect allele; A1-reference allele; OR-odds ratio; CI-confidence interval; Het. P-heterogeneity p-value; Dir.-direction.

Number									
rs35071394 in 14q21.1									
Cohort	OR [95% CI]	<b>P-value</b>	EAF	Info					
<b>BioVU EA</b>	3.17 [2.02, 4.97]	4.93x10 <sup>-7</sup>	0.91	0.74					
CARDIA EA	1.57 [0.44, 5.60]	0.484	0.94	0.66					
	rs1392505 in 1	6q21							
Cohort	OR [95% CI]	<b>P-value</b>	EAF	Info					
BioVU EA	1.78 [1.35, 2.35]	4.55x10 <sup>-5</sup>	0.80	0.95					
CARDIA EA	2.60 [1.36, 4.97]	3.73x10 <sup>-3</sup>	0.80	0.94					
	Volume								
	rs141745233 in '	7q36.1							
Cohort	Beta [95% CI]	<b>P-value</b>	EAF	Info					
BioVU EA	-0.54 [-0.75, -0.32]	9.60x10 <sup>-7</sup>	0.09	1.00					
CARDIA EA	-0.30 [-0.62, 0.03]	0.0755	0.09	0.66					
rs17506409 in 5q22.1									
Cohort Beta [95% CI] P-value EAF									
<b>BioVU EA</b>	0.53 [0.30, 0.77]	$1.04 \text{x} 10^{-5}$	0.07	0.99					
CARDIA EA	0.35 [0.07, 0.64]	0.0159	0.08	1.00					

Supplemental Table 4. Summary of cohort-specific single SNP signals from the metaanalyses between BioVU and CARDIA whites for each fibroid outcome

OR-odds ratio; CI-confidence interval; EAF-effect allele frequency.

Number									
rs55964111 in 15q15.3									
Cohort	OR [95% CI]	<b>P-value</b>	EAF	Info					
BioVU AA	1.88 [1.39, 2.55]	4.14x10 <sup>-5</sup>	0.60	0.98					
CARDIA AA	1.99 [1.18, 3.37]	0.0104	0.63	0.98					
BioVU EA	1.32 [0.85, 2.04]	0.215	0.92	0.85					
CARDIA EA	3.20 [1.23, 8.35]	0.0173	0.91	0.80					
	rs1419784 in 7 <sub>1</sub>	p14.3							
Cohort	OR [95% CI]	<b>P-value</b>	EAF	Info					
BioVU AA	1.50 [1.11, 2.02]	8.44x10 <sup>-3</sup>	0.68	0.99					
CARDIA AA	1.06 [0.61, 1.82]	0.847	0.67	0.97					
BioVU EA	1.57 [1.23, 2.01]	$2.73 \times 10^{-4}$	0.77	0.99					
CARDIA EA	2.31 [1.29, 4.11]	4.61x10 <sup>-3</sup>	0.77	0.97					
	Max Dimens	ion							
	rs7751006 in (	6q27							
Cohort	Beta [95% CI]	<b>P-value</b>	EAF	Info					
BioVU AA	0.06 [0.00, 0.12]	0.0467	0.84	0.89					
CARDIA AA	0.07 [0.00, 0.15]	0.0622	0.87	0.73					
BioVU EA	0.07 [0.02, 0.11]	2.31x10 <sup>-3</sup>	0.66	0.95					
CARDIA EA	0.09 [0.03, 0.14]	$1.47 \text{x} 10^{-3}$	0.68	0.94					
	rs111855550 in 4	4p16.1							
Cohort	Beta [95% CI]	<b>P-value</b>	EAF	Info					
BioVU AA	-0.04 [-0.10, 0.01]	0.138	0.13	0.98					
CARDIA AA	-0.10 [-0.17, -0.03]	$2.73 \times 10^{-3}$	0.12	0.99					
BioVU EA	-0.06 [-0.12, 0.00]	0.0605	0.13	0.95					
CARDIA EA	-0.14 [-0.21, -0.07]	$1.54 \mathrm{x} 10^{-4}$	0.13	1.00					
	Volume								
	rs6605005 in 1	q31.1							
Cohort	Beta [95% CI]	P-value	EAF	Info					
BioVU AA	0.30 [0.16, 0.44]	1.96x10 <sup>-6</sup>	0.75	1.00					
CARDIA AA	0.17 [0.01, 0.33]	0.0419	0.74	0.95					
BioVU EA	0.40 [0.19, 0.62]	$2.76 \times 10^{-4}$	0.91	1.00					
CARDIA EA	0.02 [-0.23, 0.27]	0.886	0.87	0.75					
	rs10024805 in 4	lq35.1							
Cohort	Beta [95% CI]	P-value	EAF	Info					
BioVU AA	0.22 [0.09, 0.35]	$9.12 \times 10^{-4}$	0.36	1.00					
CARDIA AA	0.16 [0.02, 0.29]	0.0222	0.39	1.00					
BioVU EA	0.12 [-0.03, 0.26]	0.113	0.29	0.99					
CARDIA EA	0.29 [0.13, 0.46]	$3.52 \times 10^{-4}$	0.26	1.00					
	rs7968890 in 12	2p12.1							
Cohort	Beta [95% CI]	<b>P-value</b>	EAF	Info					
BioVU AA	0.12 [-0.01, 0.24]	0.0743	0.44	1.00					
CARDIA AA	0.29 [0.15, 0.44]	7.46x10 <sup>-5</sup>	0.43	0.90					
BioVU EA	0.16 [0.04, 0.29]	0.0120	0.44	1.00					
CARDIA EA	0.14 [-0.01, 0.30]	0.0710	0.45	1.00					

 Table 5. Summary of cohort-specific single SNP signals from the transethnic meta-analyses

 between all BioVU and CARDIA women for each fibroid outcome

OR-odds ratio; CI-confidence interval; EAF-effect allele frequency.

Number									
Meta-analysis	Tissue	Gene	Chr	<b>Z-Score</b>	<b>P-value</b>	$\mathbf{R}^{2a}$	Variance <sup>b</sup>	N SNPs <sup>c</sup>	Model N <sup>d</sup>
Transethnic	Thyroid	EDEM2	20	-4.51	6.34x10 <sup>-6</sup>	0.148	0.053	14	24
EAs	Pituitary	C2orf73	2	4.45	8.48x10 <sup>-6</sup>	0.342	0.229	24	25

Supplemental Table 6. Predicted gene expression summary from MetaXcan

<sup>a</sup> Predicted performance R<sup>2</sup>.
 <sup>b</sup> Variance of predicted gene expression.
 <sup>c</sup> Number of SNPs in the prediction model that were present in the GWAS summary statistics.
 <sup>d</sup> Number of SNPs in the tissue-specific gene expression prediction model using GTEx data.

S <sub>Hom</sub>									
SNP	Region	BP	Genes	A2/A1	P-value	Number Z-Score	Max Dimension Z-Score	Volume Z-Score	
rs200348	14q32.2	99,380,544	VRK1  BCL11B	G/A	$7.13 \times 10^{-7}$	-4.26	-3.12	-1.69	
rs11691049	2q24.3	168,615,362	B3GALT1	C/A	$7.48 \times 10^{-7}$	-3.83	-3.61	-3.38	

Supplemental Table 7. Examining cross phenotype associations for transethnic meta-analyses between all BioVU and CARDIA women

SNP-single nucleotide polymorphism; BP-base pairs; A2-effect allele; A1-reference allele.