## SUPPLEMENTARY FIGURE LEGEND

#### **Supplementary Figure 1**

#### Quantile-quantile plots for the GWAS of (a) EOC and (b) HGSOC.

Data are shown as observed and expected negative-log p-values.

#### **Supplementary Figure 2**

#### Manhattan plots for the GWAS of (a) EOC and (b) HGSOC.

Data are shown as negative-log p-values.

#### **Supplementary Figure 3**

Local association plots for genetic loci reported for EOC in Table 2 near the genes: (a) *AKR1C3*, (b) *LOC101927394*, (c) *UGT2A2*, (d) *WWC1*.

Data are shown as negative-log p-values.

#### **Supplementary Figure 4**

Local association plots for genetic loci reported for HGSOC in Table 2 near the genes: (a) *FST*, (b) *MAGEC1*, (c) *LOC105376360*, (d) *PRPSAP1*, (e) *GABRG3*, (f) *LOC105377300 / GK2*.

Data are shown as negative-log p-values.

#### **Supplementary Figure 5**

# Results of eQTL analysis in 260 ovarian tissues from AACES participants for selected SNPs from the GWAS of EOC and HGSOC\*

Listed below the figure: \*Boxplots represent the distribution of measured expression versus genotype (rounded to the nearest whole number for imputed dosage variables). P-values are reported from linear models with covariate adjustment for age and two principal components of ancestry.

### Supplementary Table 1: Descriptive statistics for study participants

		EOC			HGSOC		
	Cases	Controls	Total	Cases	Controls	Total	
Study Sites	n (%)	n (%)	n (%)	n (%)	n (%)	n (%)	
AAS – African American Cancer Epidemiology Study	467 (61.9)	563 (45.6)	1030 (51.8)	349 (65.0)	563 (45.6)	912 (51.5)	
BEL – Belgium Ovarian Cancer Study	0 (0.0)	2 (0.2)	2 (0.1)	0 (0.0)	2 (0.2)	2 (0.1)	
BVU – The BioVU DNA Repository	8 (1.1)	98 (7.9)	106 (5.3)	4 (0.7)	98 (7.9)	102 (5.8)	
CAM – Cancer Research UK, Cambridge Research Institute	1 (0.1)	0 (0.0)	1 (0.1)	0 (0.0)	0 (0.0)	0 (0.0)	
DKE – Duke University Clinic	7 (0.9)	0 (0.0)	7 (0.4)	3 (0.6)	0 (0.0)	3 (0.2)	
DOV – Diseases of the Ovary and their Evaluation	7 (0.9)	27 (2.2)	34 (1.7)	4 (0.7)	27 (2.2)	31 (1.8)	
HAW – Hawaii Ovarian Cancer Study	0 (0.0)	3 (0.2)	3 (0.2)	0 (0.0)	3 (0.2)	3 (0.2)	
HOP – Hormones and Ovarian Cancer Prediction	16 (2.1)	21 (1.7)	37 (1.9)	10 (1.9)	21 (1.7)	31 (1.8)	
LAX – Women's Cancer Program at the Samuel Oschin	18 (2.4)	0 (0.0)	18 (0.9)	15 (2.8)	0 (0.0)	15 (0.9)	
Comprehensive Cancer Institute							
MAY – Mayo Clinic Ovarian Cancer Case-Control Study	3 (0.4)	5 (0.4)	8 (0.4)	2 (0.4)	5 (0.4)	7 (0.4)	
MEC – Multiethnic Cohort Study	10 (1.3)	13 (1.1)	23 (1.2)	6 (1.1)	13 (1.1)	19 (1.1)	
MOF – Moffitt Cancer Center Ovarian Cancer Study	16 (2.1)	14 (1.1)	30 (1.5)	13 (2.4)	14 (1.1)	27 (1.5)	
MSK – Memorial Sloan Kettering Cancer Center	8 (1.1)	20 (1.6)	28 (1.4)	8 (1.5)	20 (1.6)	28 (1.6)	
NCO – North Carolina Ovarian Cancer Study	108 (14.3)	160 (13.0)	268 (13.5)	76 (14.2)	160 (13.0)	236 (13.3)	
NEC – New England Case-Control Study of Ovarian Cancer	6 (0.8)	5 (0.4)	11 (0.6)	6 (1.1)	5 (0.4)	11 (0.6)	
NHS – Nurses' Health Study I and II	3 (0.4)	0 (0.0)	3 (0.2)	3 (0.6)	0 (0.0)	3 (0.2)	
NOR – University of Bergen, Haukeland University Hospital, Norway	1 (0.1)	0 (0.0)	1 (0.1)	0 (0.0)	0 (0.0)	0 (0.0)	
NTH – Nijmegen Ovarian Cancer Study	1 (0.1)	0 (0.0)	1 (0.1)	0 (0.0)	0 (0.0)	0 (0.0)	
ORE – Oregon Ovarian Cancer Registry	1 (0.1)	0 (0.0)	1 (0.1)	1 (0.2)	0 (0.0)	1 (0.1)	
OVA – Ovarian Cancer in Alberta and British Columbia	1 (0.1)	0 (0.0)	1 (0.1)	0 (0.0)	0 (0.0)	0 (0.0)	
PLC – Prostate-Lung-Colorectal and Ovarian Cancer Screening Trial	5 (0.7)	85 (6.9)	90 (4.6)	2 (0.4)	85 (6.9)	87 (4.9)	
RMH – Royal Marsden Hospital Ovarian Cancer Study	1 (0.1)	0 (0.0)	1 (0.1)	1 (0.2)	0 (0.0)	1 (0.1)	
RPC – Roswell Park Cancer Institute Ovarian Cancer Cohort	4 (0.5)	0 (0.0)	4 (0.2)	3 (0.6)	0 (0.0)	3 (0.2)	
SEA – UK Studies of Epidemiology and Risk Factors in Cancer	2 (0.3)	1 (0.1)	3 (0.2)	1 (0.2)	1 (0.1)	2 (0.1)	
Heredity (SEARCH) Ovarian Cancer Study							
SIS – The Sister Study	8 (1.1)	131 (10.6)	139 (7.0)	3 (0.6)	131 (10.6)	134 (7.6)	
SOC – Southampton Ovarian Cancer Study	1 (0.1)	0 (0.0)	1 (0.1)	0 (0.0)	0 (0.0)	0 (0.0)	
STA – Genetic Epidemiology of Ovarian Cancer	12 (1.6)	45 (3.6)	57 (2.9)	4 (0.7)	45 (3.6)	49 (2.8)	
UCI – UC Irvine Ovarian Cancer Study	0 (0.0)	2 (0.2)	2 (0.1)	0 (0.0)	2 (0.2)	2 (0.1)	
UHN – Princess Margaret Cancer Centre	4 (0.5)	0 (0.0)	4 (0.2)	2 (0.4)	0 (0.0)	2 (0.1)	
UKO – UK Ovarian Cancer Population Study	4 (0.5)	10 (0.8)	14 (0.7)	2 (0.4)	10 (0.8)	12 (0.7)	
USC – Los Angeles County Case-Control Studies of Ovarian Cancer	31 (4.1)	30 (2.4)	61 (3.1)	18 (3.4)	30 (2.4)	48 (2.7)	
WMH – Westmead Institute for Cancer Research – Westmead	1 (0.1)	0 (0.0)	1 (0.1)	1 (0.1)	0 (0.0)	1 (0.1)	
Hospital							
TOTAL	755	1235	1 <b>99</b> 0	537	1235	1772	

Subtype	Nearest Gene	SNP ID (Effect / other allele)	OR (95% CI)	P-value	BFDP <sub>1/1K</sub>	BFDP <sub>1/10K</sub>
EOC	AKR1C3	rs4525119 (T/C)	0.70 (0.61-0.81)	4.9 x 10 <sup>-7</sup>	0.08	0.49
	LOC101927394	rs7643459* (T/G)	1.40 (1.22-1.60)	8.4 x 10 <sup>-7</sup>	0.05	0.33
	UGT2A2	rs4286604 (A/G)	0.69 (0.59-0.80)	8.5 x 10 <sup>-7</sup>	0.05	0.34
	WWC1	rs142091544 (T/C)	3.22 (2.02-5.13)	9.4 x 10 <sup>-7</sup>	0.08	0.47
HGSOC	FST	rs37792 (G/A)	0.65 (0.55-0.76)	6.0 x 10 <sup>-8</sup>	0.004	0.04
	MAGEC1	rs57403204 (G/A)	2.62 (1.83-3.76)	1.7 x 10 <sup>-7</sup>	0.01	0.13
	LOC105376360	rs79079890 (G/T)	3.20 (2.05-4.99)	3.0 x 10 <sup>-7</sup>	0.03	0.25
	PRPSAP1	rs66459581 (A/AC)	1.63 (1.35-1.97)	5.1 x 10 <sup>-7</sup>	0.02	0.19
	GABRG3	rs116046250 (G/T)	2.95 (1.92-4.54)	8.7 x 10 <sup>-7</sup>	0.07	0.44
	LOC105377300 / GK2	rs192876988 (C/T)	3.01 (1.94-4.68)	9.2 x 10 <sup>-7</sup>	0.08	0.47

BFDP=Bayesian false-discovery probability; the subscripted number is the level of prior probability

Build 37 **SNP ID** Chr:Pos Subtype Nearest gene Genes within 100kb Genes at 100-500kb<sup>a</sup> rs4525119 10:5091954 AKR1C3 AKR1C3, AKR1C1, AKR1C2 rs7643459 3:8004828 LOC101927394 LOC101927394 EOC rs4286604 4:70442165 UGT2A2 UGT2B4, UGT2A2, UGT2A1 rs142091544 5,167714000 WWC1 WWC1, TENM2 ITGA1, ITGA2, FST, FST rs37792 5:52644647 NDUFS4 rs57403204 X:141078552 MAGEC1 MAGEC1, MAGEC3 rs79079890 10:3684148 LOC105376360 LOC105376360 HGSOC PRPSAP1, UBALD2, 17:74355264 PRPSAP1 QRICH2, U6, SPHK1, UBE2D, rs66459581 AANAT GABRG3, GABRB3, rs116046250 15:27231950 GABRG3 GABRA6, AK124673 GK2 rs192876988 4:80297251 GK2, NAA11, LOC100505875

Supplementary Table 3: Genes targeted for eQTL analysis within the region of ten EOC- and HGSOC-associated SNPs

<sup>a</sup>Genes in the 100-500kb range are reported and considered in eQTL analysis only for variants with no genes within 100kb.

**Supplementary Table 4:** Summary of genetic association for selected SNPs identified in GWAS of EOC and HGSOC with breast cancer in African American individuals from the AABC study

	SNP ID	Build 37	_	Breast cancer - all		ER posit	ive	ER negative	
Nearest Gene	(Effect / Other allele)	Chr:Pos	EAF	OR ( 95% Cl)	P-value	OR (95% CI)	<i>P</i> -value	OR (95% CI)	<i>P</i> -value
AKR1C3	rs4525119 (T/C)	10:5091954	0.328	1.07 (0.99,1.16)	0.098	1.02 (0.93,1.13)	0.621	1.11 (1.00,1.24)	0.058
LOC101927394	rs7643459 (T/G)	3:8004828	0.377	1.05 (0.97,1.13)	0.232	1.02 (0.93,1.12)	0.622	1.13 (1.01,1.26)	0.029
UGT2A2	rs4286604 (A/G)	4:70442165	0.279	0.94 (0.86,1.02)	0.142	0.93 (0.84, 1.04)	0.205	0.96 (0.85,1.09)	0.551
WWC1	rs142091544 (T/C)	5:167714000	0.040	1.07 (0.87, 1.31)	0.530	0.87 (0.67, 1.14)	0.314	1.55 (1.19, 2.02)	0.001
FST	rs37792 (G/A)	5:52644647	0.341	0.96 (0.89, 1.04)	0.424	0.97 (0.88,1.07)	0.578	0.97 (0.87, 1.09)	0.621
MAGEC1	rs57403204 (G/A)	X:141078552	0.051	0.93 (0.77, 1.12)	0.424	0.89 (0.71, 1.12)	0.329	0.97 (0.74, 1.25)	0.813
LOC105376360	rs79079890 (G/T)	10:3684148	0.035	1.22 (0.99,1.50)	0.066	1.21 (0.95,1.55)	0.125	1.12 (0.83,1.52)	0.449
PRPSAP1	rs66459581 (A/AC)	17:74355264	0.228	1.01 (0.92,1.10)	0.890	0.89 (0.80,1.00)	0.054	1.12 (0.99,1.28)	0.081
GABRG3	rs116046250 (G/T)	15:27231950	0.045	0.87 (0.71,1.06)	0.176	0.88 (0.69,1.12)	0.306	0.83 (0.62,1.10)	0.195
LOC105377300 / GK2	rs192876988 (C/T)	4:80297251	0.040	1.19 (0.96,1.47)	0.105	1.32 (1.03,1.69)	0.027	1.02 (0.75,1.39)	0.883

SNP: single nucleotide polymorphism; EAF: effect allele frequency; OR: odds ratio; CI: confidence interval

**Supplementary Table 5:** Summary of genetic association for selected SNPs identified in GWAS of EOC and HGSOC with prostate cancer in African American individuals from the AAPC study

Nearest Gene	SNP ID (Effect / Other allele)	Build 37 Chr:Pos	EAF	OR (95% CI)	<i>P</i> -value
AKR1C3	rs4525119 (T/C)	10:5091954	0.328	1.03 (0.97, 1.10)	0.304
LOC101927394	rs7643459 (T/G)	3:8004828	0.371	1.07 (1.01, 1.13)	0.034
UGT2A2	rs4286604 (A/G)	4:70442165	0.264	1.08 (1.01, 1.15)	0.025
WWC1	rs142091544 (T/C)	5:167714000	0.036	1.11 (0.94, 1.32)	0.197
FST	rs37792 (G/A)	5:52644647	0.325	0.99 (0.93, 1.03)	0.685
MAGEC1	rs57403204 (G/A)	X:141078552	0.056	0.98 (0.89, 1.09)	0.754
LOC105376360	rs79079890 (G/T)	10:3684148	0.037	0.96 (0.82, 1.13)	0.646
PRPSAP1	rs66459581 (A/AC)	17:74355264	0.230	1.03 (0.96, 1.11)	0.363
GABRG3	rs116046250 (G/T)	15:27231950	0.042	0.93 (0.80, 1.08)	0.365
LOC105377300 / GK2	rs192876988 (C/T)	4:80297251	0.043	0.95 (0.81, 1.12)	0.567

SNP: single nucleotide polymorphism; EAF: effect allele frequency; OR: odds ratio; CI: confidence interval

Supplementary Table 6: Summary of results for fine mapping in African ancestry women of loci previously identified in GWAS of European ancestry women

					EOC				HGSC <sup>a</sup>			
Locus	SNP ID	Build 37 Chr:Pos	Nearest Gene	Phenotype	Number of SNPs Plotted	Minimum SNP position in region	Minimum SNP P-value	Number of SNPs Plotted	Minimum SNP position in region	Minimum SNP P-value		
Confirmed	SNPs in Europ	ean ancestry On	coArray meta-analysis									
1p34.3	rs58722170	1:38096421	RSP01	Serous	3078	chr1:37917469	2.91E-04	2685	chr1:37917469	3.38E-04		
2q14.1	rs752590	2:113972945	PAX8	Mucinous	3615	chr2:114298849	2.31E-03					
2q31.1	rs711830	2:177037311	HOXD3	Mucinous	3171	chr2:1177037311	2.23E-03					
2q31.1	rs6755777	2:177043226	HAGLR	Serous	3175	chr2:177326682	2.23E-03	2775	chr2:177442640	2.49E-03		
3q25.31	rs62274041	3:156435640	TIPARP	HGSC	3204	chr3:156252180	2.01E-03	2863	chr3:156204177	2.37E-03		
5p15.33	rs10069690	5:1279790	TERT	Serous	5037	chr5:949207	9.57E-04	4497	chr5:892204	7.97E-04		
5p15.33	rs7705526	5:1285974	TERT	Serous borderline	5005	chr5:949207	9.57E-04	4464	chr5:892204	7.97E-04		
8q21.13	rs76837345	8:82668818	CHMP4C	HGSC	4045	chr8:82866267	1.44E-05 <sup>c</sup>	3523	chr8:82866267	3.98E-06 <sup>b</sup>		
8q24.21	rs1400482	8:128529685	LINC00824	Serous	4414	chr8:128619531	3.51E-04	3897	chr8:128762529	1.86E-03		
9p22.2	rs10962692	9:16915874	BNC2	HGSC	5248	chr9:16978052	2.67E-05°	4746	chr9:16986321	5.57E-05		
9q34.2	rs8176685	9:136138766	ABO	HGSC	3968	chr9:136075407	5.07E-04	3472	chr9:136267149	3.17E-04		
10p12.31	rs144962376	10:21878832	MLLT10	Serous	2514	chr10:21978590	4.13E-03	2162	chr10:22035324	7.73E-03		
17q12	rs7405776	17:36093022	HNF1B	Serous	2787	chr17:36195231	5.44E-03	2324	chr17:36113300	7.82E-04		
17q12	rs11651755	17:36099840	HNF1B	Clear cell	2778	chr17:36195231	5.44E-03					
17q21.31	rs7207826	17:46500673	SKAP1	Serous	2827	chr17:46807761	6.43E-04	2572	chr17:46215732	8.18E-04		
17q21.32	rs1879586	17:43567337	PLEKHM1	HGSC	3735	chr17:43954416	1.31E-04	3370	chr17:43945726	1.44E-04		
19p13.11	rs4808075	19:17390291	BABAM1	HGSC	9100	chr19:17395213	1.77E-04	4077	chr19:17088458	3.07E-04		
19q11.21	rs688187	19:39732752	IFNL3	Mucinous	8360	chr19:39684764	1.21E-04					
Newly ider	ntified SNPs in E	European ancesti	ry OncoArray meta-analysi	is								
2q13	rs2165109	2:111818658	ACOXL	HGSC	2172	chr2:111595642	6.87E-03	1951	chr2:111832215	1.47E-02		
3q22.3	rs112071820	3:138849110	BPESC1	Mucinous	2922	chr3:138839642	3.34E-05°					
3q28	rs9870207	3:190525516	GMNC	Serous borderline, LGSC	4647	chr3:190657915	3.73E-03	4081	chr3:190839089	3.34E-03		
4q32.2	rs13113999	4:167187046	TLL1	Serous borderline	4879	chr4:167492545	4.71E-03	4252	chr4:166876033	8.01E-04		
5q12.3	rs555025179	5:66121089	MAST4	Endometrioid	3720	chr5:65907851	5.91E-03					
8q21.11	rs150293538	8:77320354	LINC01111	Serous borderline, LGSC	3305	chr8:77354021	1.17E-04	2883	chr8:77320354	2.01E-04		
8q24.21	rs9886651	8:128817883	PVT1	HGSC	4449	chr8:129036159	2.67E-04	3962	chr8:129093793	1.52E-03		
9q31.1	rs320203	9:104943226	LOC105376188	Mucinous	4927	chr9:105237170	1.49E-03					
10q24.33	rs7902587	10:105694301	LOC102724351	Serous borderline, LGSC	3192	chr10:105375295	3.40E-05 <sup>c</sup>	2852	chr10:105300054	1.03E-03		
12q24.31	rs7953249	12:121403724	HNF1A-AS1	HGSC	3680	chr12:121113096	6.90E-05	3272	chr12:121113096	2.51E-05 <sup>c</sup>		
18q11.2	rs8098244	18:21405553	LAMA3	Serous borderline, LGSC	2685	chr18:21555816	1.84E-05 <sup>♭</sup>	2431	chr18:21555816	6.19E-05		
22q12.1	rs6005807	22:28934313	TTC28/ LOC101929594	HGSC	2513	chr22:29126617	1.27E-02	2066	chr22:29073209	1.56E-02		

EOC: epithelial ovarian cancer; HGSC: high-grade serous ovarian cancer; SNP: single nucleotide polymorphism; Chr: chromosome; Pos: position; LGSC: low-grade serous ovarian cancer. <sup>a</sup>Fine mapping among HGSC was completed only for those SNPs associated with serous ovarian cancer.

<sup>b</sup>Significant at the Bonferroni threshold (0.05/number of SNPs plotted).

<sup>o</sup>Significant at the suggestive threshold (0.05/(number of SNPs plotted/3)).

Supplementary Table 7: Summary of results for fine mapping in European ancestry women of loci identified in our GWAS of African ancestry women

			_	EOC			HGSOC	
Gene	SNP ID	Build 37 Chr:Pos	Number of SNPs plotted	Minimum SNP position in region	Min SNP p-value	Number of SNPs plotted	Minimum SNP position in region	Min SNP p-value
AKR1C3	rs4525119	10:5091954	6105	chr10:5341275	2.44E-03			
LOC101927394	rs7643459	3:8004828	5466	chr3:8268351	1.12E-03			
UGT2A2	rs4286604	4:70442165	5542	chr4:70592790	1.14E-05 <sup>a</sup>			
WWC1	rs142091544	5:167714000	3761	chr5:167857657	1.74E-04			
FST	rs37792	5:52644647				4416	chr5:52332701	3.94E-05
MAGEC1	rs57403204	X:141078552				3389	chrX:141078552	1.78E-04
LOC105376360	rs79079890	10:3684148				4899	chr10:3463140	2.24E-04
PRPSAP1	rs66459581	17:74355264				4025	chr17:74420211	3.74E-04
GABRG3	rs116046250	15:27231950				3885	chr15:27240688	5.16E-04
LOC105377300/GK2	rs192876988	4:80297251				3443	chr4:80297251	5.76E-04

<sup>a</sup>Significant at the suggestive threshold (0.05/(number of SNPs plotted/3)).







Manichaikul et al., Supplementary Figure 1



(b)



Manichaikul et al., Supplementary Figure 2

(a)



(b)

Epithelial Ovarian Cancer







(d)



Manichaikul et al., Supplementary Figure 3

(c)

(a)



(b)





(d)





(f)



Manichaikul et al., Supplementary Figure 4

(e)



Manichaikul et al., Supplementary Figure 5