

## Supplemental Data

### Gene-centric Association Signals for Lipids

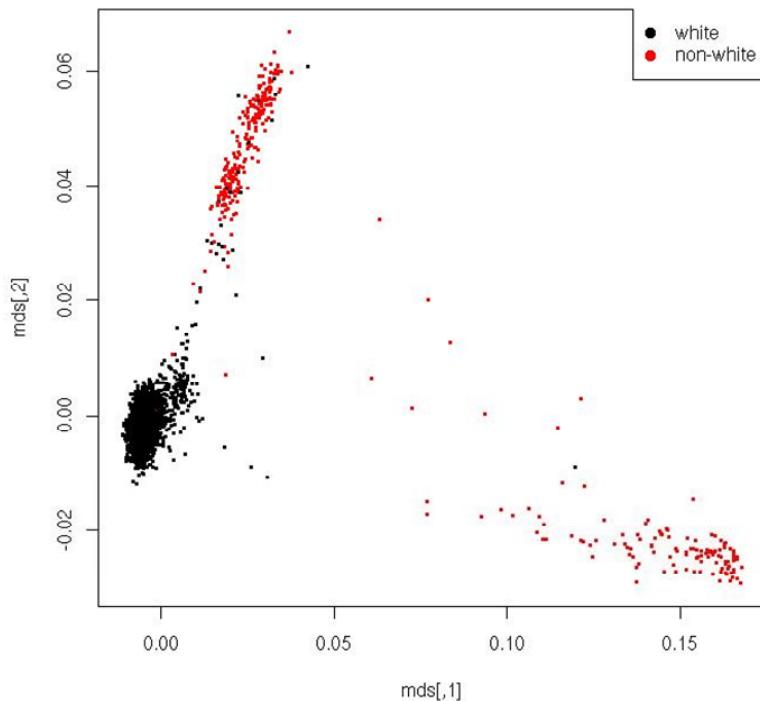
### and Apolipoproteins Identified

### via the HumanCVD BeadChip

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Figure S1. Population Structure

ANALYSIS 1: A multidimensional scaling plot based on the pairwise Identity-By-State distances of all QC-passed samples (5441) and SNPs (48032)



ANALYSIS 2: A multidimensional scaling plot based on the pairwise Identity-By-State distances of Caucasian samples (5067), showing 8 outliers circled in red.

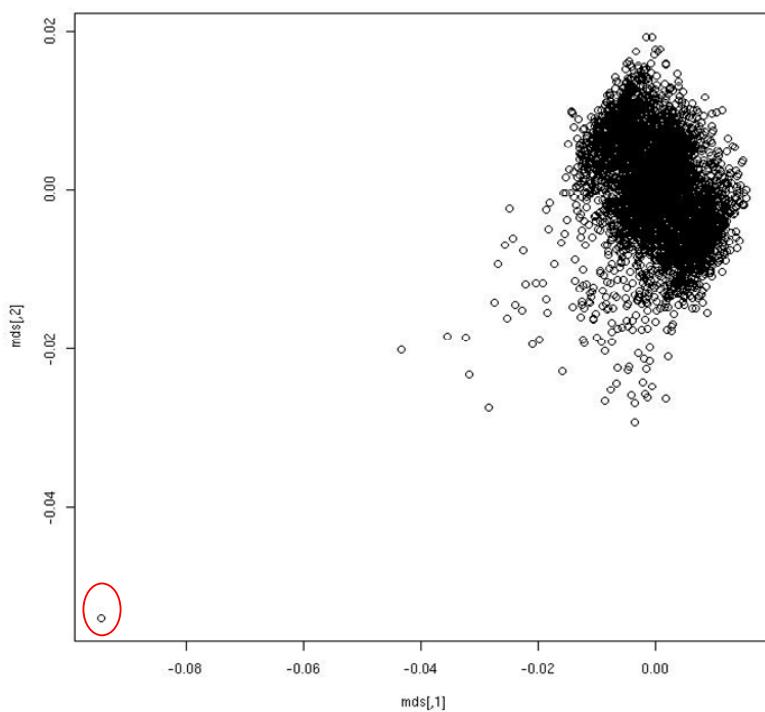
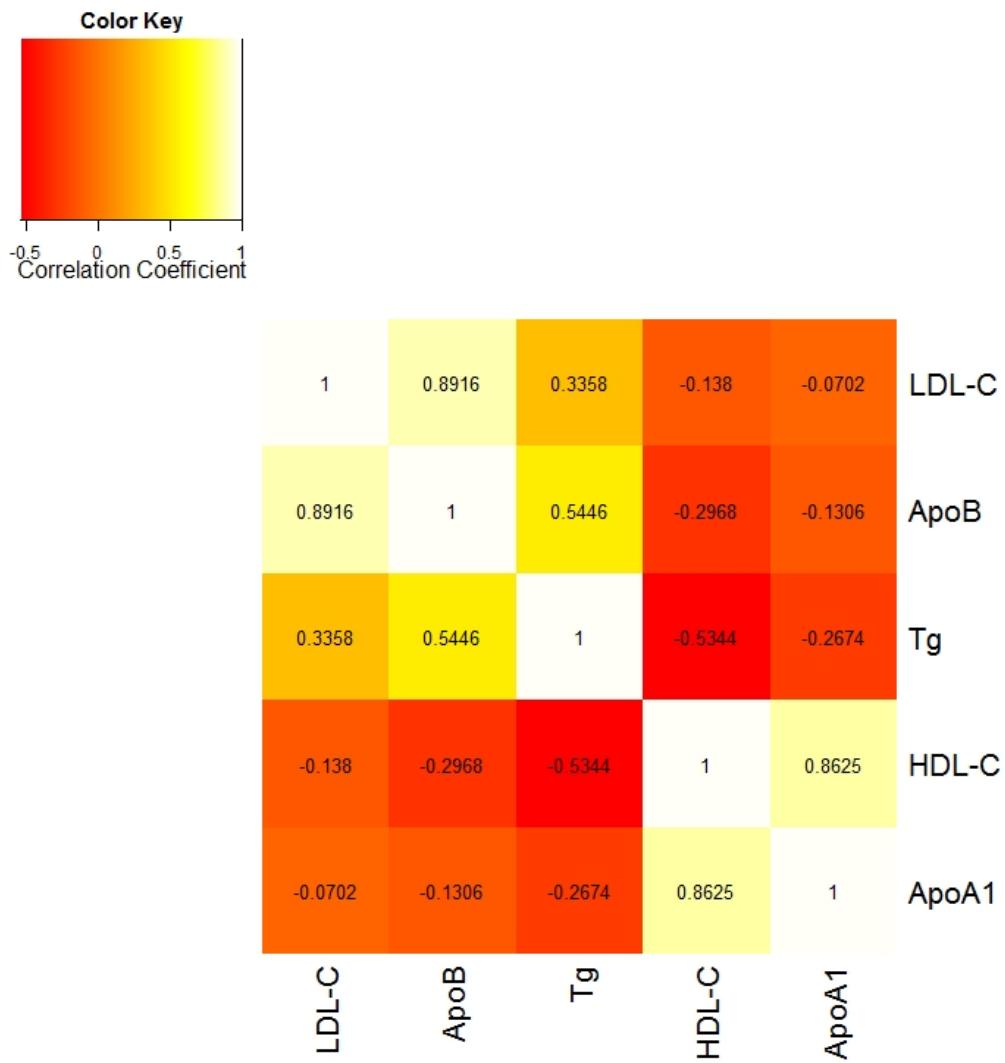


Figure S2 . Heat plot reflecting the relationship between the five traits with SNPs that that were significant at the cut-off of  $10^{-5}$  for at least one of the traits. White shows the strongest association m, red the weakest.



**Figure S3a**

Haplovew plots for ApoB showing the SNPs significant by univariate analysis ( at  $p < 10^{-5}$ ) and those retained after variable selection (red boxes). For the Chromosome 19 APOE locus those SNPs in orange are no longer retained when the APOE rs rs429358 and rs7412 defining the common alleles were included into the analysis

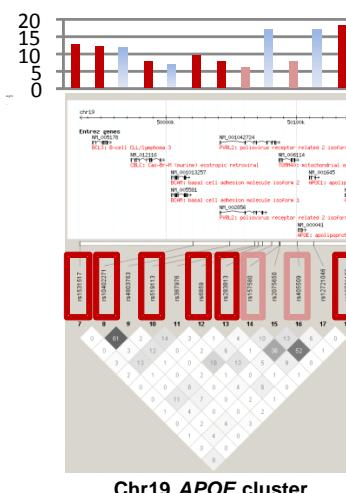
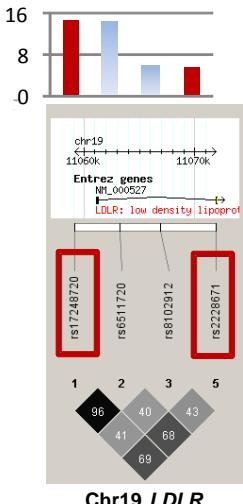
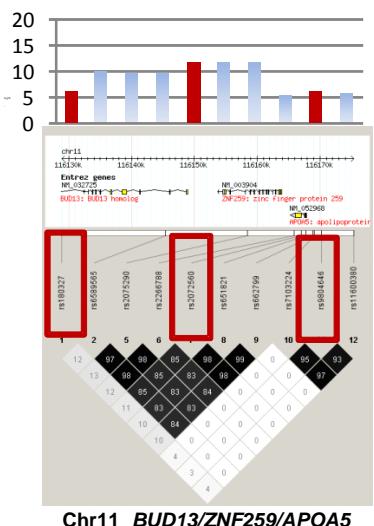
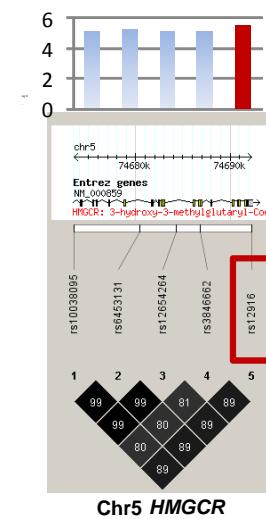
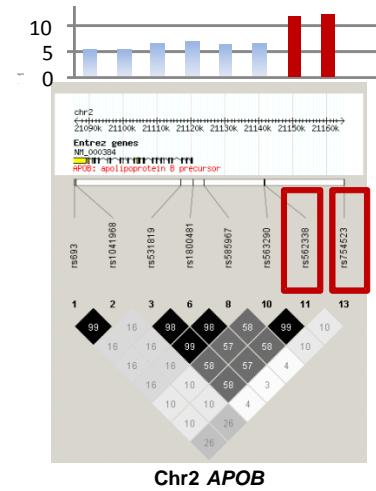
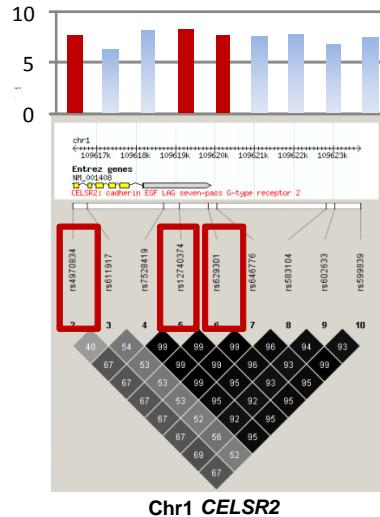


Figure S3b. Haploview plots for ApoAI showing the SNPs significant by univariate analysis (at  $p < 10^{-5}$ ) and those retained after variable selection (red boxes)

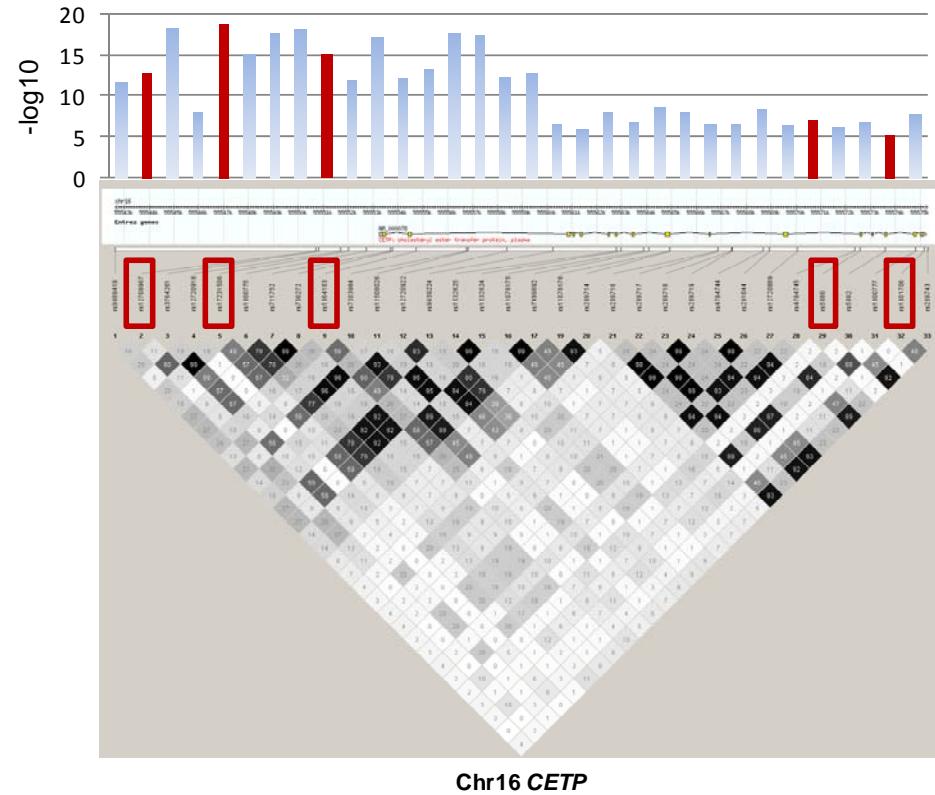
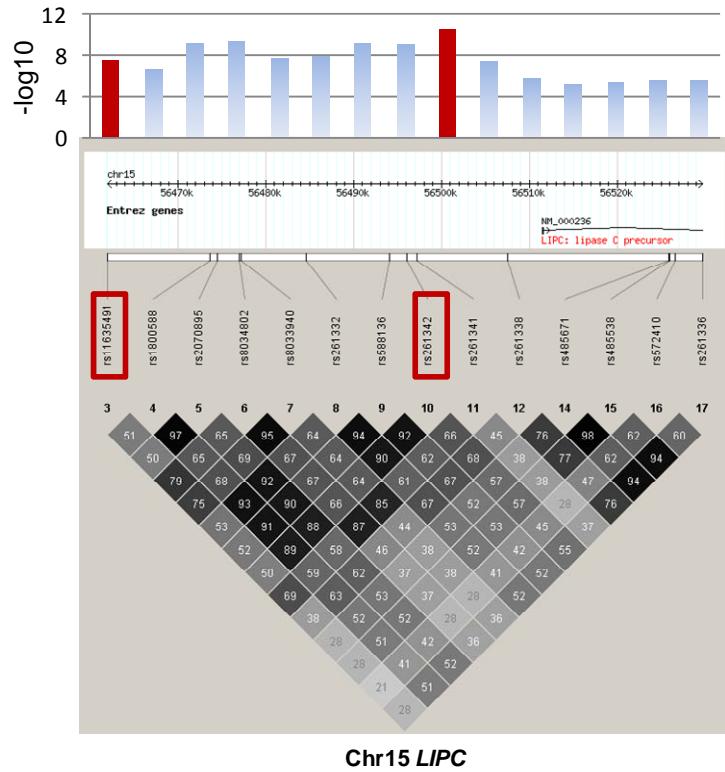


Figure S4. Frequency distribution of the gene count score and the effect of gene score on a) HDL-C, b) ApoAI, c) apoB, d) apoB/apoAI. The odds ratio for occupancy of the extreme 10% of the trait distribution at different cut points of the respective gene score distributions are presented below each histogram. The medians (interquartile ranges) of the gene counts were 28 (26-30) for apoB, 12 (10-14) for HDL-C, 10 (8-11) for apoAI, and 25 (24-26) for apoB/apoAI.

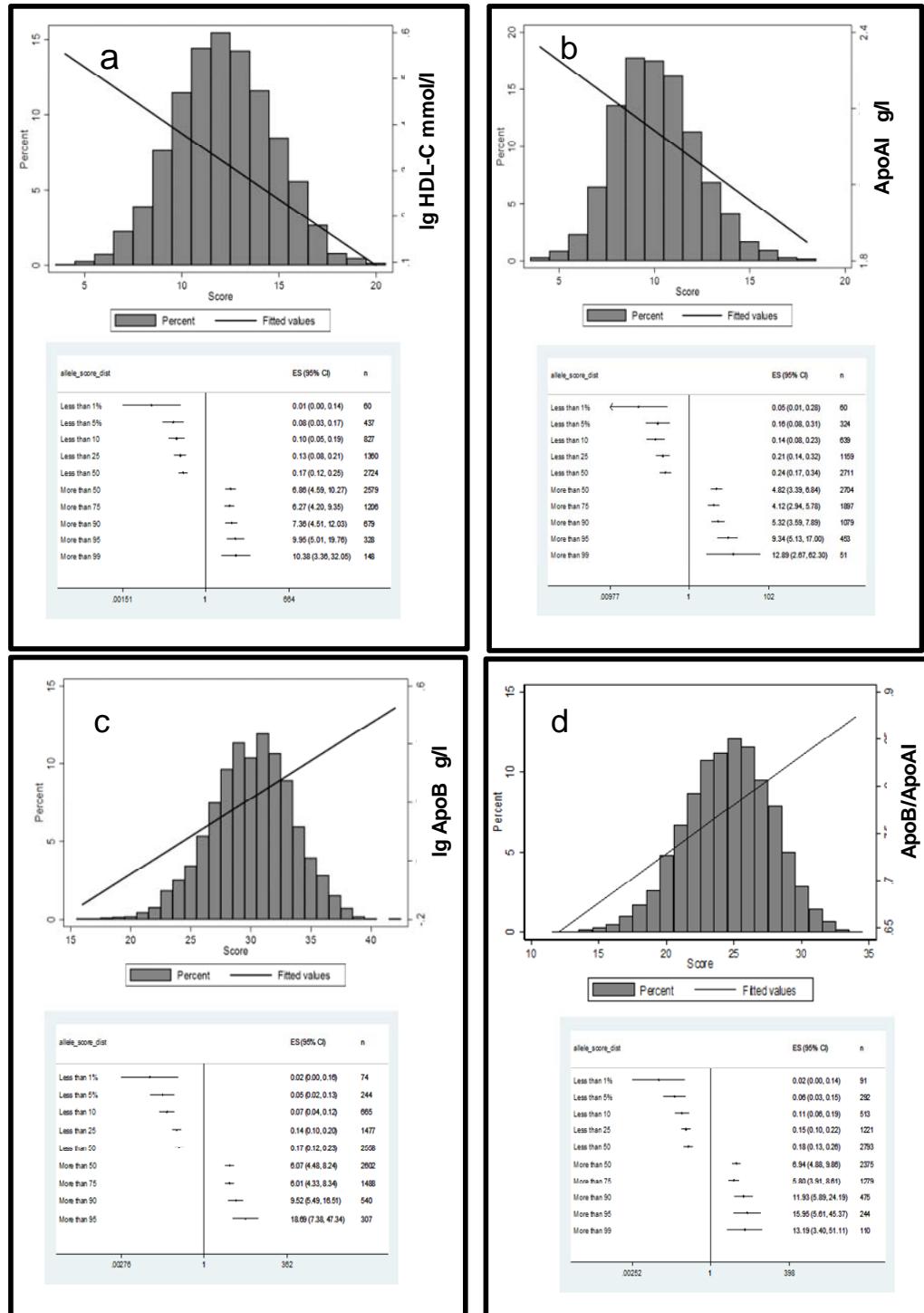
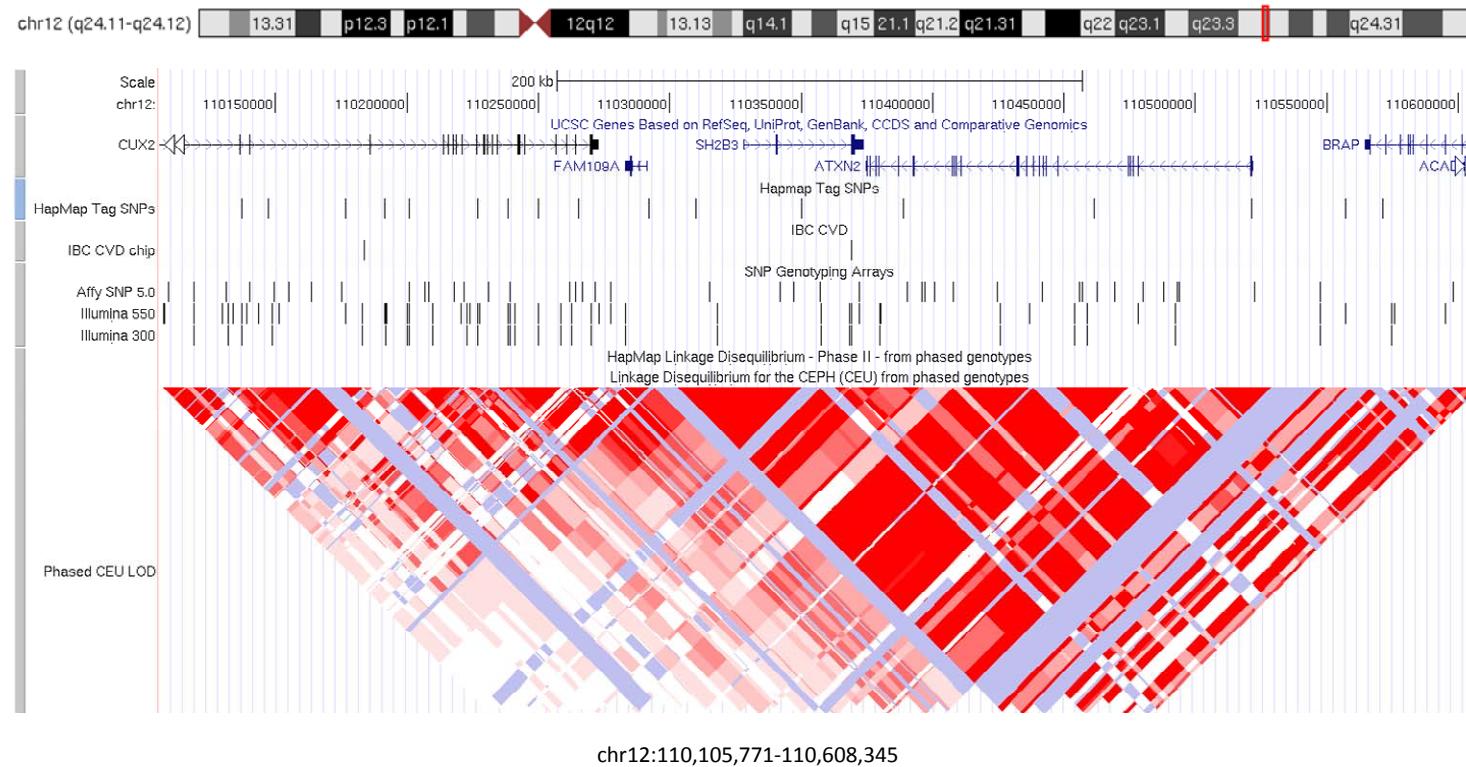
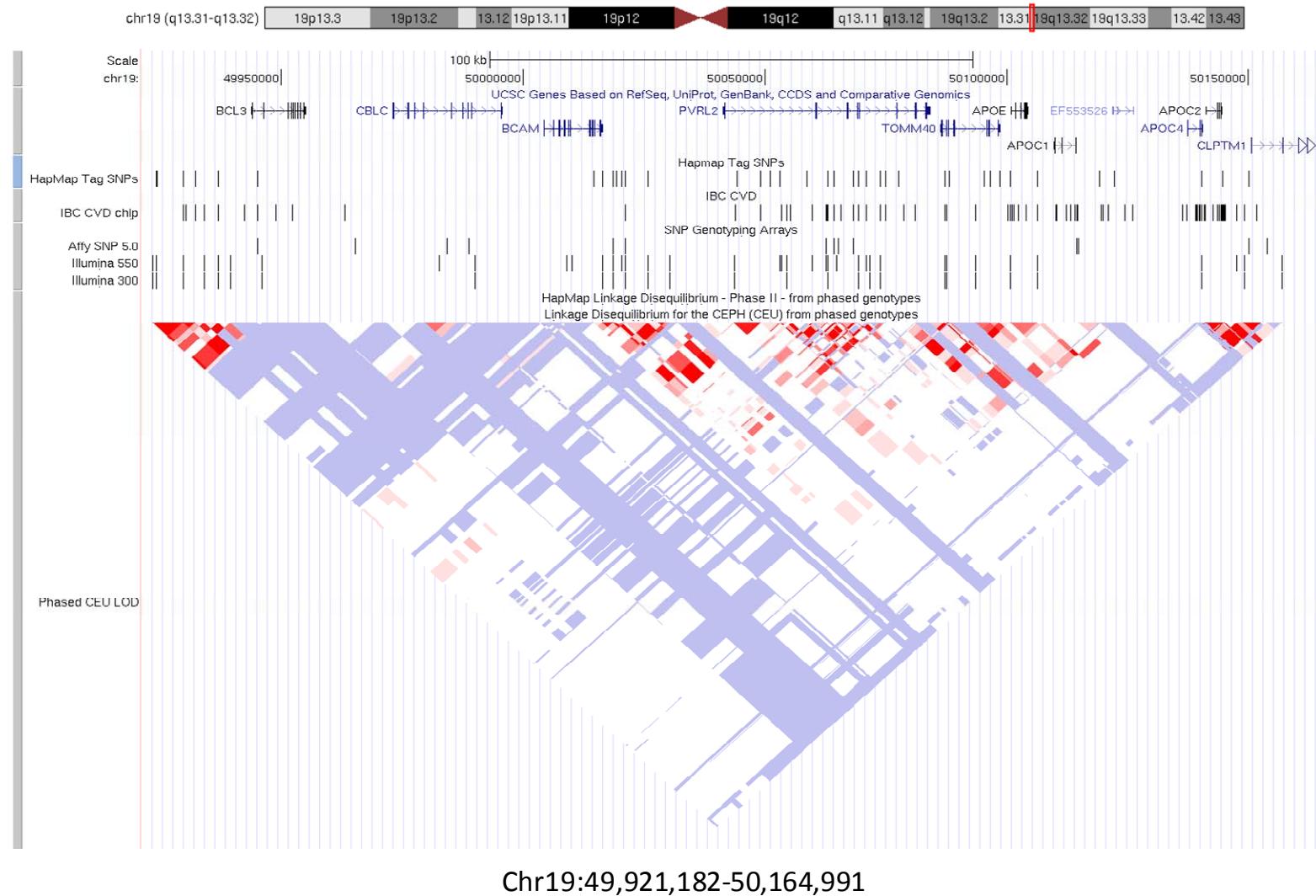


Figure S5a-d The figures shows the SNP coverage of the IBC CVD chip compared to the Illumina 550K, Illumina 300K and Affy5.0 GWAS chips using the UCSC Genome Browser (<http://genome.ucsc.edu/>). The IBC CVD chip has much denser coverage for loci with a high likelihood of functional significance and candidate loci that are potentially involved in phenotypes of interest

a) *SH2B3* locus



b) *APOE* locus



c) *LDLR* locus



d) *BMPR2* locus



Table S1. Details of the five studies used in the meta-analyses and genotyping methodology

	<b>Whitehall II</b> <sup>1</sup>	<b>BWHHS</b> <sup>2</sup>	<b>BRIGHT</b> <sup>3</sup>	<b>ASCOT</b> <sup>4</sup>	<b>NORDIL</b> <sup>5</sup>
Design	Prospective observational	Prospective observational	Cases (from case-control) 1751, but see below	DNA repository from randomized trial 1262, but see below	DNA repository from randomized trial 1935
n	5058	3313			
Males (%)	73.5	0		40	82
Mean age (yrs)	49	69		58	63
Proportion receiving lipid lowering medication (%)	0.8	7		11.3	12.6 not analysed
Genotyping platform	IBC chip	IBC chip	IBC chip	IBC chip	IBC chip
Calling algorithm	Gencall	Gencall	Illuminus	Illuminus	Illuminus
Contribution to current analysis	Index study	Meta-analysis	Meta-analysis	Meta-analysis	Meta-analysis

### **BRIGHT and ASCOT**

Sample sizes for BRIGHT are as follows:

without age/sex covariates: 1646 for LDL-C, and 1751 for other lipid phenotypes

with age/sex covariates: 1599 for LDL-C, and 1700 for other lipid phenotypes

Sample sizes for ASCOT are as follows:

1169 for LDL-C, 1192 for TG, and 1262 for HDLC

For both BRIGHT and ASCOT, % males, mean age and % on LLA were calculated using largest sample size (resp. 1751 and 1262)

## References

1. Marmot,M.G., Smith,G.D., Stansfeld,S., Patel,C., North,F., Head,J., White,I., Brunner,E., and Feeney,A. (1991). Health inequalities among British civil servants: the Whitehall II study. *Lancet* *337*, 1387-1393
2. Lawlor,D.A., Bedford,C., Taylor,M., and Ebrahim,S. (2003). Geographical variation in cardiovascular disease, risk factors, and their control in older women: British Women's Heart and Health Study. *J. Epidemiol. Community Health* *57*, 134-140
3. Caulfield,M., Munroe,P., Pembroke,J., Samani,N., Dominiczak,A., Brown,M., Benjamin,N., Webster,J., Ratcliffe,P., O'Shea,S. et al. (2003). Genome-wide mapping of human loci for essential hypertension. *Lancet* *361*, 2118-2123
4. Sever,P.S., Dahlof,B., Poulter,N.R., Wedel,H., Beevers,G., Caulfield,M., Collins,R., Kjeldsen,S.E., McInnes,G.T., Mehlsen,J. et al. (2001). Rationale, design, methods and baseline demography of participants of the Anglo-Scandinavian Cardiac Outcomes Trial. ASCOT investigators. *J. Hypertens.* *19*, 1139-1147
5. Hansson,L., Hedner,T., Lund-Johansen,P., Kjeldsen,S.E., Lindholm,L.H., Syvertsen,J.O., Lanke,J., de,F.U., Dahlof,B., and Karlberg,B.E. (2000). Randomised trial of effects of calcium antagonists compared with diuretics and beta-blockers on cardiovascular morbidity and mortality in hypertension: the Nordic Diltiazem (NORDIL) study. *Lancet* *356*, 359-365

Table S2. SNP p Values Adjusted for Age and Gender for the 195 SNPs Associated with Lipid Traits

Excel file available online at <http://www.cell.com/AJHG>.

Table S3a. All SNPs significantly associated with LDL-C

SNP	CHR	BP	GENE	Alleles		MAF	Common HMZ mean mmol/l	HTZ mean mmol/l	Rare HMZ mean mmol/l	Beta	P-value	R2 (univariate)	Chr R2
				1	2								
rs11591147	1	55278235	PCSK9	C	A	0.02	4.38	3.84	2.98	-0.55	9.28E-12	6.98E-02	0.07
rs4970834	1	109616403	CELSR2	G	A	0.18	4.42	4.27	4.08	-0.16	5.18E-09		
rs611917	1	109616775	CELSR2	A	G	0.31	4.44	4.31	4.26	-0.11	1.97E-06		
rs7528419	1	109618715	CELSR2	A	G	0.21	4.43	4.27	4.14	-0.15	2.41E-09		
rs12740374	1	109619113	CELSR2	C	A	0.21	4.43	4.28	4.12	-0.15	1.82E-09		
rs629301	1	109619829	CELSR2	A	C	0.21	4.43	4.28	4.14	-0.15	6.50E-09		
rs646776	1	109620053	PSRC1	A	G	0.21	4.43	4.28	4.14	-0.14	1.03E-08		
rs583104	1	109622830	PSRC1	A	C	0.22	4.43	4.28	4.15	-0.14	1.15E-08		
rs602633	1	109623034	PSRC1	C	A	0.20	4.43	4.27	4.17	-0.14	3.15E-08		
rs599839	1	109623689	PSRC1	A	G	0.22	4.42	4.29	4.16	-0.14	4.31E-08		
rs693	2	21085700	APOB	A	G	0.48	4.48	4.34	4.27	-0.11	3.86E-08	7.05E-02	0.07
rs1041968	2	21086309	APOB	A	G	0.48	4.48	4.34	4.27	-0.11	5.32E-08		
rs10199768	2	21097505	APOB	C	A	0.44	4.29	4.36	4.49	0.10	6.58E-07		
rs531819	2	21117144	APOB	C	A	0.15	4.41	4.26	4.11	-0.16	4.91E-08		
rs1367117	2	21117405	APOB	G	A	0.33	4.30	4.39	4.53	0.12	2.63E-08		
rs12714264	2	21119023	APOB	T	A	0.13	4.41	4.26	4.09	-0.16	1.23E-07		
rs1800481	2	21120715	APOB	G	A	0.15	4.41	4.26	4.10	-0.16	4.95E-08		
rs934197	2	21120966	APOB	G	A	0.33	4.30	4.40	4.53	0.12	3.43E-08		
rs585967	2	21124059	APOB	C	A	0.15	4.41	4.26	4.10	-0.16	3.28E-08		
rs7575840	2	21126995	APOB	C	A	0.35	4.30	4.38	4.56	0.12	7.84E-08		
rs563290	2	21141731	APOB	A	G	0.18	4.43	4.23	4.20	-0.18	1.33E-11		
rs562338	2	21141826	APOB	G	A	0.18	4.43	4.24	4.16	-0.18	1.21E-11		
rs754524	2	21165046	APOB	A	C	0.28	4.31	4.40	4.57	0.12	1.69E-07		
rs754523	2	21165196	APOB	A	G	0.33	4.30	4.37	4.60	0.13	2.86E-09		
rs4299376	2	43926080	ABCG8	A	C	0.32	4.28	4.42	4.53	0.13	8.70E-10		
rs10038095	5	74673467	HMGCR	A	T	0.38	4.28	4.40	4.51	0.12	4.29E-08	5.87E-02	0.06
rs10474434	5	74680437	HMGCR	C	A	0.23	4.31	4.42	4.57	0.12	2.05E-06		
rs6453131	5	74680462	HMGCR	A	C	0.38	4.27	4.40	4.50	0.11	4.59E-08		
rs17238484	5	74684252	HMGCR	C	A	0.23	4.31	4.43	4.57	0.12	1.79E-06		
rs12654264	5	74684359	HMGCR	T	A	0.38	4.27	4.40	4.50	0.11	5.76E-08		
rs3846662	5	74686840	HMGCR	A	G	0.43	4.27	4.38	4.49	0.11	5.17E-08		

rs12916	5	74692295	<i>HMGCR</i>	A	G	0.41	4.26	4.39	4.51	0.12	6.66E-09		
rs3804231	5	74732535	<i>COL4A3BP</i>	G	A	0.13	4.33	4.46	4.62	0.14	5.15E-06		
rs4704220	5	74793312	<i>COL4A3BP</i>	G	A	0.39	4.30	4.38	4.49	0.09	8.05E-06		
rs6589565	11	116145447	<i>BUD13</i>	G	A	0.07	4.34	4.50	4.94	0.19	3.33E-06	5.63E-02	0.06
rs2075290	11	116158506	<i>ZNF259</i>	A	G	0.07	4.34	4.49	4.94	0.18	6.05E-06		
rs2266788	11	116165896	<i>APOA5</i>	A	G	0.07	4.34	4.49	4.94	0.18	5.43E-06		
rs2072560	11	116167036	<i>APOA5</i>	G	A	0.06	4.34	4.51	5.33	0.22	2.36E-07		
rs651821	11	116167789	<i>APOA5</i>	A	G	0.06	4.34	4.50	5.33	0.21	6.90E-07		
rs662799	11	116168917	<i>APOA4</i>	A	G	0.06	4.34	4.50	5.33	0.21	7.19E-07		
rs3764261	16	55550825	<i>CETP</i>	C	A	0.33	4.43	4.33	4.23	-0.11	1.06E-06	5.61E-02	0.06
rs17231506	16	55552029	<i>CETP</i>	G	A	0.32	4.44	4.33	4.22	-0.11	5.02E-07		
rs1529729	19	11024562	<i>SMARCA4</i>	A	G	0.45	4.28	4.39	4.44	0.09	6.71E-06	1.11E-01	0.11
rs17248720	19	11059187	<i>LDLR</i>	G	A	0.13	4.45	4.14	3.81	-0.31	7.86E-25		
rs6511720	19	11063306	<i>LDLR</i>	C	A	0.13	4.45	4.14	3.85	-0.30	7.30E-24		
rs8102912	19	11066975	<i>LDLR</i>	G	A	0.23	4.44	4.26	4.19	-0.16	2.24E-11		
rs8110695	19	11067530	<i>LDLR</i>	T	A	0.22	4.43	4.27	4.21	-0.14	1.12E-08		
rs2228671	19	11071912	<i>LDLR</i>	G	A	0.13	4.42	4.21	4.18	-0.18	6.52E-10		
rs2569559	19	11075533	<i>LDLR</i>	A	C	0.23	4.43	4.27	4.22	-0.14	1.74E-08		
rs1531517	19	49934013	<i>BCL3</i>	G	A	0.07	4.40	4.17	4.08	-0.22	6.26E-08		
rs10402271	19	50021054	<i>PVRL2</i>	A	C	0.33	4.26	4.44	4.53	0.15	2.06E-12		
rs4803763	19	50049131	<i>PVRL2</i>	G	C	0.26	4.29	4.43	4.58	0.15	2.20E-10		
rs519113	19	50068124	<i>PVRL2</i>	C	G	0.24	4.42	4.31	4.18	-0.12	9.37E-07		
rs387976	19	50070900	<i>PVRL2</i>	A	C	0.35	4.43	4.34	4.22	-0.10	5.98E-06		
rs6859	19	50073874	<i>PVRL2</i>	G	A	0.41	4.27	4.38	4.52	0.12	3.53E-08		
rs283813	19	50081014	<i>PVRL2</i>	T	A	0.07	4.39	4.23	3.70	-0.19	2.51E-06		
rs157580	19	50087106	<i>TOMM40</i>	A	G	0.40	4.47	4.34	4.21	-0.11	1.03E-07		
rs2075650	19	50087459	<i>TOMM40</i>	A	G	0.14	4.31	4.51	4.87	0.23	1.14E-14		
rs12721046	19	50113094	<i>APOC1</i>	G	A	0.15	4.30	4.51	4.74	0.21	7.58E-14		
rs12721109	19	50139061	<i>APOC2</i>	G	A	0.02	4.39	3.82	4.32	-0.54	5.06E-14		

HMZ, homozygote ; HTZ, heterozygote

Table S3b. All SNPs significantly associated with ApoB

SNP	CHR	BP	GENE	Alleles		MAF	Common HMZ mean g/l	HTZ mean g/l	Rare HMZ mean g/l	Beta	P-value	R2 (univariate)	Chr R2
				1	2								
rs11591147	1	55278235	<i>PCSK9</i>	C	A	0.02	1.28	1.14	0.94	-0.12	1.89E-10	7.91E-03	0.08
rs4970834	1	109616403	<i>CELSR2</i>	G	A	0.18	1.29	1.25	1.18	-0.03	2.00E-08	6.00E-03	
rs611917	1	109616775	<i>CELSR2</i>	A	G	0.31	1.29	1.25	1.23	-0.03	6.69E-07	4.86E-03	
rs7528419	1	109618715	<i>CELSR2</i>	A	G	0.21	1.29	1.25	1.19	-0.03	8.09E-09	6.92E-03	
rs12740374	1	109619113	<i>CELSR2</i>	C	A	0.21	1.29	1.25	1.19	-0.03	4.70E-09	7.18E-03	
rs629301	1	109619829	<i>CELSR2</i>	A	C	0.21	1.29	1.25	1.19	-0.03	1.76E-08	6.67E-03	
rs646776	1	109620053	<i>PSRC1</i>	A	G	0.21	1.29	1.25	1.19	-0.03	2.40E-08	6.55E-03	
rs583104	1	109622830	<i>PSRC1</i>	A	C	0.22	1.29	1.25	1.19	-0.03	2.18E-08	6.37E-03	
rs602633	1	109623034	<i>PSRC1</i>	C	A	0.20	1.29	1.25	1.20	-0.03	2.21E-07	5.64E-03	
rs599839	1	109623689	<i>PSRC1</i>	A	G	0.22	1.29	1.25	1.20	-0.03	4.79E-08	5.96E-03	
rs693	2	21085700	<i>APOB</i>	A	G	0.48	1.29	1.27	1.25	-0.02	5.69E-06	3.06E-03	0.08
rs1041968	2	21086309	<i>APOB</i>	A	G	0.48	1.29	1.27	1.25	-0.02	7.80E-06	2.95E-03	
rs531819	2	21117144	<i>APOB</i>	C	A	0.15	1.28	1.25	1.20	-0.03	3.21E-07	4.55E-03	
rs12714264	2	21119023	<i>APOB</i>	T	A	0.13	1.28	1.24	1.21	-0.04	1.60E-07	4.64E-03	
rs1800481	2	21120715	<i>APOB</i>	G	A	0.15	1.28	1.25	1.19	-0.03	7.41E-07	4.36E-03	
rs585967	2	21124059	<i>APOB</i>	C	A	0.15	1.28	1.24	1.20	-0.03	2.48E-07	4.63E-03	
rs563290	2	21141731	<i>APOB</i>	A	G	0.18	1.29	1.23	1.20	-0.04	7.38E-13	9.29E-03	
rs562338	2	21141826	<i>APOB</i>	G	A	0.18	1.29	1.24	1.19	-0.04	6.00E-13	9.24E-03	
rs754523	2	21165196	<i>APOB</i>	A	G	0.33	1.25	1.27	1.32	0.02	1.11E-06	4.15E-03	
rs4299376	2	43926080	<i>ABCG8</i>	A	C	0.32	1.25	1.28	1.32	0.03	7.60E-08	5.74E-03	
rs10038095	5	74673467	<i>HMGCR</i>	A	T	0.38	1.25	1.28	1.31	0.02	8.05E-06	3.98E-03	0.07
rs6453131	5	74680462	<i>HMGCR</i>	A	C	0.38	1.25	1.28	1.31	0.02	6.34E-06	4.07E-03	
rs12654264	5	74684359	<i>HMGCR</i>	T	A	0.38	1.25	1.28	1.30	0.02	9.25E-06	3.90E-03	
rs3846662	5	74686840	<i>HMGCR</i>	A	G	0.43	1.25	1.27	1.30	0.02	8.57E-06	3.86E-03	
rs12916	5	74692295	<i>HMGCR</i>	A	G	0.41	1.25	1.27	1.31	0.02	3.34E-06	4.56E-03	
rs180327	11	116128869	<i>BUD13</i>	A	G	0.36	1.24	1.29	1.29	0.02	7.70E-07	5.02E-03	0.08
rs6589565	11	116145447	<i>BUD13</i>	G	A	0.07	1.26	1.33	1.46	0.06	2.01E-10	7.59E-03	
rs2075290	11	116158506	<i>ZNF259</i>	A	G	0.07	1.26	1.33	1.46	0.06	4.73E-10	7.23E-03	
rs2266788	11	116165896	<i>APOA5</i>	A	G	0.07	1.26	1.33	1.45	0.06	3.42E-10	7.34E-03	

rs2072560	11	116167036	<i>APOA5</i>	G	A	0.06	1.26	1.34	1.58	0.07	1.40E-12	9.10E-03	
rs651821	11	116167789	<i>APOA5</i>	A	G	0.06	1.26	1.33	1.58	0.07	3.55E-12	8.69E-03	
rs662799	11	116168917	<i>APOA4</i>	A	G	0.06	1.26	1.33	1.58	0.07	5.18E-12	8.77E-03	
rs7103224	11	116169176	<i>APOA4</i>	G	A	0.08	1.28	1.22	1.28	-0.04	7.82E-06	3.98E-03	
rs9804646	11	116170289	<i>APOA4</i>	G	A	0.09	1.28	1.22	1.22	-0.04	1.25E-06	4.49E-03	
rs11600380	11	116175392	<i>APOA4</i>	A	G	0.08	1.28	1.22	1.26	-0.04	5.19E-06	4.02E-03	
rs3764261	16	55550825	<i>CETP</i>	C	A	0.33	1.29	1.26	1.23	-0.02	1.52E-06	4.32E-03	0.07
rs17231506	16	55552029	<i>CETP</i>	G	A	0.32	1.29	1.26	1.23	-0.02	1.36E-06	4.41E-03	
rs17248720	19	11059187	<i>LDLR</i>	G	A	0.13	1.29	1.22	1.14	-0.06	2.10E-15	1.22E-02	0.13
rs6511720	19	11063306	<i>LDLR</i>	C	A	0.13	1.29	1.22	1.15	-0.05	6.77E-15	1.18E-02	
rs8102912	19	11066975	<i>LDLR</i>	G	A	0.23	1.29	1.25	1.24	-0.03	1.26E-06	4.01E-03	
rs2228671	19	11071912	<i>LDLR</i>	G	A	0.13	1.28	1.24	1.23	-0.03	5.00E-06	4.22E-03	
rs1531517	19	49934013	<i>BCL3</i>	G	A	0.07	1.28	1.20	1.15	-0.07	1.96E-13	1.05E-02	
rs10402271	19	50021054	<i>PVRL2</i>	A	C	0.33	1.24	1.29	1.32	0.04	7.05E-13	9.93E-03	
rs4803763	19	50049131	<i>PVRL2</i>	G	C	0.26	1.25	1.29	1.34	0.04	2.75E-12	9.45E-03	
rs519113	19	50068124	<i>PVRL2</i>	C	G	0.24	1.29	1.25	1.22	-0.03	2.14E-08	6.27E-03	
rs387976	19	50070900	<i>PVRL2</i>	A	C	0.35	1.29	1.26	1.22	-0.03	1.54E-07	5.86E-03	
rs6859	19	50073874	<i>PVRL2</i>	G	A	0.41	1.24	1.28	1.32	0.03	1.49E-10	9.21E-03	
rs283813	19	50081014	<i>PVRL2</i>	T	A	0.07	1.28	1.23	1.09	-0.05	2.61E-08	5.41E-03	
rs157580	19	50087106	<i>TOMM40</i>	A	G	0.40	1.30	1.26	1.23	-0.02	6.55E-07	6.92E-03	
rs2075650	19	50087459	<i>TOMM40</i>	A	G	0.14	1.25	1.31	1.45	0.06	8.38E-18	1.50E-02	
rs405509	19	50100676	<i>APOE;PKP2</i>	C	A	0.48	1.25	1.27	1.30	0.03	1.89E-08	5.06E-03	
rs12721046	19	50113094	<i>APOC1</i>	G	A	0.15	1.25	1.32	1.41	0.06	1.01E-17	1.55E-02	
rs12721109	19	50139061	<i>APOC2</i>	G	A	0.02	1.28	1.10	1.05	-0.15	3.88E-19	1.67E-02	

HMZ, homozygote ; HTZ, heterozygote

Table S3c. All SNPs significantly associated with Triglycerides

SNP	CHR	BP	GENE	Alleles		MAF	Common HMZ mean mmol/l	HTZ mean mmol/l	Rare HMZ mean mmol/l	Beta	P-value	R2 (univariate)	Chr R2
				1	2								
rs1260326	2	27584444	GCKR	G	A	0.40	1.38	1.43	1.63	0.06	1.83E-07	5.67E-03	0.06
rs714052	7	72502805	BAZ1B	A	G	0.12	1.48	1.33	1.29	-0.09	4.38E-07	4.88E-03	0.07
rs2074755	7	72515102	BAZ1B	A	G	0.12	1.48	1.33	1.29	-0.09	2.94E-07	4.97E-03	
rs17145713	7	72542746	BAZ1B	G	A	0.20	1.50	1.36	1.25	-0.09	5.29E-10	7.11E-03	
rs12539316	7	72615834	TBL2	A	G	0.29	1.49	1.41	1.33	-0.05	9.39E-06	3.78E-03	
rs2286276	7	72625290	TBL2	G	A	0.29	1.49	1.41	1.33	-0.05	9.48E-06	3.73E-03	
rs11974409	7	72627326	TBL2	A	G	0.20	1.49	1.36	1.27	-0.08	5.66E-09	6.03E-03	
rs1051921	7	72645879	MLXIPL	G	A	0.20	1.49	1.36	1.28	-0.08	1.25E-08	5.60E-03	
rs17145750	7	72664314	MLXIPL	G	A	0.16	1.48	1.35	1.29	-0.07	1.48E-06	4.15E-03	
rs7800944	7	72673793	MLXIPL	A	G	0.29	1.49	1.41	1.33	-0.05	8.59E-06	3.79E-03	
rs253	8	19855697	LPL	G	A	0.45	1.53	1.44	1.34	-0.05	6.48E-06	4.68E-03	0.08
rs255	8	19856181	LPL	A	G	0.15	1.48	1.36	1.23	-0.07	2.46E-06	4.14E-03	
rs256	8	19856247	LPL	G	A	0.15	1.48	1.37	1.15	-0.07	2.07E-06	4.27E-03	
rs258	8	19856532	LPL	C	G	0.45	1.52	1.44	1.33	-0.05	8.69E-06	4.53E-03	
rs263	8	19857092	LPL	G	A	0.18	1.48	1.40	1.12	-0.07	3.09E-06	4.25E-03	
rs264	8	19857460	LPL	G	A	0.15	1.48	1.37	1.17	-0.07	2.51E-06	4.16E-03	
rs271	8	19857982	LPL	G	A	0.15	1.48	1.36	1.25	-0.08	6.59E-07	4.59E-03	
rs285	8	19859469	LPL	G	A	0.47	1.54	1.45	1.31	-0.07	2.65E-09	7.73E-03	
rs301	8	19861214	LPL	A	G	0.25	1.51	1.38	1.21	-0.08	2.17E-11	8.38E-03	
rs320	8	19863357	LPL	A	C	0.28	1.51	1.40	1.26	-0.08	1.79E-10	7.33E-03	
rs327	8	19863816	LPL	A	C	0.29	1.51	1.40	1.25	-0.08	2.40E-10	7.40E-03	
rs328	8	19864004	LPL	C	G	0.11	1.48	1.31	1.07	-0.11	8.42E-10	7.26E-03	
rs331	8	19864685	LPL	G	A	0.27	1.51	1.39	1.24	-0.08	1.73E-11	8.13E-03	
rs12679834	8	19864713	LPL	A	G	0.11	1.48	1.31	1.07	-0.11	1.06E-09	7.30E-03	
rs3289	8	19867472	LPL	A	G	0.03	1.43	1.70	1.04	0.16	3.16E-06	3.79E-03	
rs3208305	8	19867928	LPL	A	T	0.30	1.52	1.40	1.24	-0.08	3.06E-11	8.09E-03	
rs3735964	8	19868325	LPL	C	A	0.12	1.48	1.31	1.17	-0.10	9.09E-09	6.18E-03	
rs13702	8	19868772	LPL	A	G	0.30	1.52	1.40	1.24	-0.08	2.85E-11	8.21E-03	
rs3916027	8	19869148	SLC18A1	G	A	0.27	1.51	1.40	1.24	-0.08	1.04E-10	7.43E-03	

rs2197089	8	19870653	<i>SLC18A1</i>	A	G	0.45	1.35	1.45	1.57	0.06	5.03E-09	7.37E-03	
rs17482753	8	19876926	<i>SLC18A1</i>	C	A	0.11	1.48	1.31	1.07	-0.11	3.20E-09	6.76E-03	
rs10503669	8	19891970	<i>SLC18A1</i>	C	A	0.11	1.48	1.30	1.08	-0.11	1.05E-09	7.28E-03	
rs17410962	8	19892360	<i>SLC18A1</i>	G	A	0.13	1.49	1.31	1.13	-0.10	4.64E-09	7.01E-03	
rs17489268	8	19896325	<i>SLC18A1</i>	A	T	0.27	1.51	1.40	1.24	-0.08	1.78E-10	7.48E-03	
rs17411031	8	19896590	<i>SLC18A1</i>	C	G	0.27	1.51	1.40	1.24	-0.08	2.90E-10	7.27E-03	
rs17489282	8	19896798	<i>SLC18A1</i>	G	A	0.27	1.51	1.40	1.24	-0.08	2.13E-10	7.33E-03	
rs17411126	8	19899552	<i>SLC18A1</i>	A	G	0.27	1.51	1.40	1.24	-0.08	1.74E-10	7.27E-03	
rs765547	8	19910554	<i>SLC18A1</i>	G	A	0.27	1.51	1.40	1.24	-0.08	2.02E-10	7.20E-03	
rs11986942	8	19911725	<i>SLC18A1</i>	G	C	0.31	1.51	1.41	1.28	-0.06	4.79E-08	5.47E-03	
rs1919484	8	19913956	<i>SLC18A1</i>	G	A	0.27	1.51	1.40	1.24	-0.08	1.46E-10	7.33E-03	
rs6586891	8	19958878	<i>SLC18A1</i>	A	C	0.34	1.41	1.45	1.56	0.05	4.17E-06	3.54E-03	
rs2001844	8	126547927	<i>TRIB1</i>	A	G	0.47	1.52	1.43	1.38	-0.05	8.89E-06	3.86E-03	
rs17321515	8	126555591	<i>TRIB1</i>	A	G	0.47	1.53	1.43	1.37	-0.05	3.47E-06	4.25E-03	
rs2980869	8	126557432	<i>TRIB1</i>	G	A	0.47	1.53	1.43	1.38	-0.05	6.88E-06	3.86E-03	
rs17108993	10	95354023	<i>RBP4</i>	C	G	0.03	1.42	1.72	1.53	0.14	8.04E-06	4.22E-03	0.06
rs499790	11	116024949	<i>BUD13</i>	G	A	0.10	1.41	1.55	2.06	0.09	3.24E-07	5.34E-03	0.09
rs480823	11	116030940	<i>BUD13</i>	A	G	0.08	1.41	1.59	2.29	0.11	1.07E-08	6.75E-03	
rs481843	11	116031077	<i>BUD13</i>	G	A	0.08	1.41	1.59	2.32	0.11	1.48E-08	6.57E-03	
rs4938303	11	116090197	<i>BUD13</i>	A	G	0.27	1.39	1.49	1.59	0.06	3.47E-06	4.64E-03	
rs28927680	11	116124283	<i>BUD13</i>	C	G	0.07	1.42	1.62	1.90	0.13	4.24E-09	7.63E-03	
rs180327	11	116128869	<i>BUD13</i>	A	G	0.36	1.31	1.52	1.57	0.08	3.09E-12	1.02E-02	
rs11820589	11	116139072	<i>BUD13</i>	G	A	0.06	1.42	1.62	1.99	0.13	2.58E-09	7.80E-03	
rs6589565	11	116145447	<i>BUD13</i>	G	A	0.07	1.39	1.81	1.92	0.19	4.49E-20	1.63E-02	
rs12286037	11	116157417	<i>ZNF259</i>	G	A	0.06	1.42	1.64	1.83	0.13	1.55E-09	7.96E-03	
rs2075290	11	116158506	<i>ZNF259</i>	A	G	0.07	1.39	1.80	1.92	0.19	6.55E-20	1.60E-02	
rs603446	11	116159645	<i>ZNF259</i>	G	A	0.45	1.52	1.45	1.31	-0.05	8.78E-06	4.26E-03	
rs12285095	11	116163241	<i>ZNF259</i>	A	C	0.06	1.42	1.63	1.85	0.13	7.29E-09	7.18E-03	
rs2266788	11	116165896	<i>APOA5</i>	A	G	0.07	1.39	1.81	1.88	0.19	1.16E-19	1.59E-02	
rs2072560	11	116167036	<i>APOA5</i>	G	A	0.06	1.39	1.81	2.04	0.21	1.92E-20	1.62E-02	
rs12287066	11	116167541	<i>APOA5</i>	C	A	0.06	1.42	1.64	1.85	0.13	3.13E-09	7.70E-03	
rs651821	11	116167789	<i>APOA5</i>	A	G	0.06	1.39	1.81	2.04	0.21	8.89E-21	1.64E-02	
rs662799	11	116168917	<i>APOA4</i>	A	G	0.06	1.39	1.81	2.04	0.21	2.91E-20	1.62E-02	
rs10750097	11	116169250	<i>APOA4</i>	A	G	0.21	1.38	1.53	1.70	0.09	1.26E-10	8.11E-03	
rs7396835	11	116189238	<i>APOA4</i>	G	A	0.13	1.40	1.56	1.79	0.09	1.38E-08	6.27E-03	

rs7396851	11	116189374	<i>APOA4</i>	G	A	0.13	1.40	1.56	1.80	0.09	4.66E-09	6.67E-03	
rs33989105	11	116205352	<i>APOC3</i>	G	A	0.25	1.40	1.47	1.67	0.07	1.75E-07	5.44E-03	
rs5142	11	116207060	<i>APOA1</i>	G	A	0.09	1.41	1.61	1.88	0.11	1.26E-08	6.14E-03	
rs5072	11	116212793	<i>APOA1;NCAN</i>	G	A	0.08	1.41	1.60	1.84	0.11	1.74E-07	5.50E-03	
rs3794991	19	19471596	<i>PBX4</i>	G	A	0.09	1.47	1.32	1.08	-0.09	4.41E-06	4.37E-03	0.06
rs16996148	19	19519472	<i>PBX4</i>	C	A	0.08	1.47	1.31	1.09	-0.09	1.69E-06	4.76E-03	
rs12610185	19	19582722	<i>PBX4</i>	G	A	0.09	1.47	1.31	1.09	-0.10	5.64E-07	5.22E-03	
rs10500212	19	19584215	<i>GMIP</i>	G	A	0.09	1.47	1.31	1.09	-0.10	6.79E-07	5.12E-03	
rs2304128	19	19607151	<i>GMIP</i>	C	A	0.09	1.47	1.31	0.99	-0.10	3.15E-07	5.29E-03	

HMZ, homozygote ; HTZ, heterozygote

Table S3d. All SNPs significantly associated with HDL-C

SNP	CHR	BP	GENE	Alleles		MAF	Common HMZ mean mmol/l	HTZ mean mmol/l	Rare HMZ mean mmol/l	Beta	P-value	R2 (univariate)	Chr R2
				1	2								
rs285	8	19859469	<i>LPL</i>	G	A	0.47	1.41	1.42	1.49	0.03	1.61E-06	4.94E-03	0.17
rs301	8	19861214	<i>LPL</i>	A	G	0.25	1.41	1.46	1.52	0.04	9.25E-11	6.87E-03	
rs320	8	19863357	<i>LPL</i>	A	C	0.28	1.41	1.45	1.50	0.03	3.61E-09	5.21E-03	
rs327	8	19863816	<i>LPL</i>	A	C	0.29	1.41	1.45	1.50	0.03	6.71E-09	5.22E-03	
rs328	8	19864004	<i>LPL</i>	C	G	0.11	1.42	1.49	1.51	0.05	3.17E-08	5.53E-03	
rs331	8	19864685	<i>LPL</i>	G	A	0.27	1.41	1.45	1.50	0.04	2.83E-10	5.89E-03	
rs12679834	8	19864713	<i>LPL</i>	A	G	0.11	1.42	1.49	1.51	0.05	5.39E-08	5.51E-03	
rs3208305	8	19867928	<i>LPL</i>	A	T	0.30	1.41	1.45	1.51	0.04	5.35E-10	5.87E-03	
rs3735964	8	19868325	<i>LPL</i>	C	A	0.12	1.42	1.49	1.47	0.04	1.17E-07	4.80E-03	
rs13702	8	19868772	<i>LPL</i>	A	G	0.30	1.41	1.45	1.51	0.04	7.49E-10	5.87E-03	
rs3916027	8	19869148	<i>SLC18A1</i>	G	A	0.27	1.41	1.45	1.51	0.04	5.39E-10	5.64E-03	
rs2197089	8	19870653	<i>SLC18A1</i>	A	G	0.45	1.48	1.41	1.41	-0.03	2.68E-06	4.58E-03	
rs17482753	8	19876926	<i>SLC18A1</i>	C	A	0.11	1.42	1.49	1.51	0.05	5.31E-08	5.38E-03	
rs10503669	8	19891970	<i>SLC18A1</i>	C	A	0.11	1.42	1.49	1.51	0.05	6.00E-08	5.42E-03	
rs17410962	8	19892360	<i>SLC18A1</i>	G	A	0.13	1.42	1.48	1.52	0.04	1.31E-07	5.52E-03	
rs17489268	8	19896325	<i>SLC18A1</i>	A	T	0.27	1.41	1.45	1.51	0.04	1.19E-09	5.73E-03	
rs17411031	8	19896590	<i>SLC18A1</i>	C	G	0.27	1.41	1.45	1.51	0.04	1.09E-09	5.80E-03	
rs17489282	8	19896798	<i>SLC18A1</i>	G	A	0.27	1.41	1.45	1.51	0.04	7.54E-10	5.75E-03	
rs17411126	8	19899552	<i>SLC18A1</i>	A	G	0.27	1.41	1.45	1.51	0.04	7.77E-10	5.65E-03	
rs765547	8	19910554	<i>SLC18A1</i>	G	A	0.27	1.41	1.45	1.51	0.04	5.55E-10	5.69E-03	
rs11986942	8	19911725	<i>SLC18A1</i>	G	C	0.31	1.41	1.45	1.49	0.03	4.01E-08	4.44E-03	
rs1919484	8	19913956	<i>SLC18A1</i>	G	A	0.27	1.41	1.45	1.51	0.04	9.68E-10	5.59E-03	
rs180327	11	116128869	<i>BUD13</i>	A	G	0.36	1.45	1.43	1.38	-0.03	3.81E-06	4.31E-03	0.17
rs11820589	11	116139072	<i>BUD13</i>	G	A	0.06	1.44	1.39	1.14	-0.05	7.66E-06	4.19E-03	
rs6589565	11	116145447	<i>BUD13</i>	G	A	0.07	1.44	1.37	1.40	-0.05	4.40E-07	4.25E-03	
rs2075290	11	116158506	<i>ZNF259</i>	A	G	0.07	1.44	1.37	1.40	-0.05	4.24E-07	4.11E-03	
rs2266788	11	116165896	<i>APOA5</i>	A	G	0.07	1.44	1.37	1.41	-0.05	4.82E-07	4.19E-03	
rs2072560	11	116167036	<i>APOA5</i>	G	A	0.06	1.44	1.37	1.28	-0.06	7.39E-08	4.63E-03	
rs651821	11	116167789	<i>APOA5</i>	A	G	0.06	1.44	1.38	1.28	-0.06	1.03E-07	4.36E-03	

rs662799	11	116168917	<i>APOA4</i>	A	G	0.06	1.44	1.37	1.28	-0.06	6.52E-08	4.70E-03	
rs10750097	11	116169250	<i>APOA4</i>	A	G	0.21	1.45	1.41	1.37	-0.03	1.33E-07	4.41E-03	
rs4775041	15	56461987	<i>LIPC</i>	C	G	0.30	1.41	1.43	1.53	0.03	2.05E-07	4.51E-03	0.18
rs1800588	15	56510967	<i>LIPC</i>	G	A	0.21	1.41	1.45	1.54	0.03	3.85E-07	4.87E-03	
rs2070895	15	56511231	<i>LIPC</i>	G	A	0.22	1.41	1.45	1.54	0.03	2.38E-07	5.08E-03	
rs261332	15	56514617	<i>LIPC</i>	G	A	0.21	1.41	1.45	1.55	0.03	1.92E-07	5.14E-03	
rs588136	15	56517790	<i>LIPC</i>	A	G	0.21	1.41	1.46	1.53	0.03	7.02E-07	4.94E-03	
rs261342	15	56518445	<i>LIPC</i>	C	G	0.22	1.41	1.45	1.54	0.03	6.28E-08	5.48E-03	
rs9989419	16	55542640	<i>CETP</i>	G	A	0.40	1.49	1.42	1.34	-0.05	9.76E-21	1.58E-02	0.21
rs12708967	16	55550712	<i>CETP</i>	A	G	0.19	1.47	1.38	1.29	-0.06	8.79E-23	1.71E-02	
rs3764261	16	55550825	<i>CETP</i>	C	A	0.33	1.37	1.47	1.56	0.07	6.80E-36	2.60E-02	
rs12720918	16	55551713	<i>CETP</i>	A	G	0.28	1.47	1.40	1.37	-0.05	1.00E-15	1.03E-02	
rs17231506	16	55552029	<i>CETP</i>	G	A	0.32	1.37	1.47	1.56	0.07	2.21E-36	2.62E-02	
rs1800775	16	55552737	<i>CETP</i>	C	A	0.49	1.35	1.44	1.52	0.06	2.63E-29	2.21E-02	
rs711752	16	55553712	<i>CETP</i>	G	A	0.43	1.35	1.45	1.52	0.06	6.48E-32	2.36E-02	
rs708272	16	55553789	<i>CETP</i>	G	A	0.43	1.35	1.45	1.52	0.06	1.87E-32	2.41E-02	
rs1864163	16	55554734	<i>CETP</i>	G	A	0.26	1.48	1.38	1.32	-0.06	2.80E-27	2.07E-02	
rs7203984	16	55556759	<i>CETP</i>	A	C	0.19	1.47	1.37	1.31	-0.06	1.08E-20	1.51E-02	
rs11508026	16	55556829	<i>CETP</i>	G	A	0.43	1.35	1.45	1.52	0.06	9.92E-30	2.15E-02	
rs708273	16	55557450	<i>CETP</i>	G	A	0.30	1.46	1.41	1.37	-0.03	1.00E-06	4.51E-03	
rs12720922	16	55558386	<i>CETP</i>	G	A	0.18	1.47	1.37	1.30	-0.06	1.34E-21	1.60E-02	
rs12597002	16	55559905	<i>CETP</i>	C	A	0.30	1.46	1.41	1.37	-0.03	1.31E-06	4.40E-03	
rs9939224	16	55560233	<i>CETP</i>	C	A	0.20	1.47	1.38	1.30	-0.06	5.81E-23	1.71E-02	
rs1532625	16	55562802	<i>CETP</i>	G	A	0.44	1.35	1.45	1.52	0.06	2.91E-31	2.25E-02	
rs1532624	16	55562980	<i>CETP</i>	C	A	0.44	1.35	1.45	1.52	0.06	4.96E-31	2.23E-02	
rs11076175	16	55563879	<i>CETP</i>	A	G	0.17	1.47	1.37	1.29	-0.07	5.58E-23	1.76E-02	
rs7499892	16	55564091	<i>CETP</i>	G	A	0.17	1.47	1.36	1.30	-0.07	7.14E-24	1.83E-02	
rs5883	16	55564854	<i>CETP</i>	G	A	0.06	1.42	1.51	1.54	0.05	5.92E-06	4.31E-03	
rs11076176	16	55564947	<i>CETP</i>	A	C	0.16	1.45	1.39	1.31	-0.05	4.62E-12	8.18E-03	
rs289714	16	55564952	<i>CETP</i>	A	G	0.17	1.45	1.39	1.35	-0.04	2.56E-10	6.91E-03	
rs289716	16	55566877	<i>CETP</i>	A	T	0.31	1.40	1.45	1.51	0.03	1.22E-09	7.35E-03	
rs289717	16	55566889	<i>CETP</i>	G	A	0.35	1.46	1.42	1.36	-0.03	6.89E-10	6.54E-03	
rs289718	16	55567433	<i>CETP</i>	A	G	0.32	1.40	1.44	1.52	0.04	2.55E-10	8.36E-03	
rs289719	16	55567442	<i>CETP</i>	G	A	0.31	1.40	1.44	1.51	0.03	1.27E-09	7.19E-03	
rs4784744	16	55568686	<i>CETP</i>	G	A	0.35	1.46	1.42	1.36	-0.03	5.37E-10	6.40E-03	

rs291044	16	55568953	<i>CETP</i>	G	A	0.35	1.46	1.42	1.36	-0.03	9.67E-10	6.23E-03	
rs12720889	16	55570064	<i>CETP</i>	T	A	0.30	1.40	1.45	1.51	0.03	1.33E-09	7.23E-03	
rs4784745	16	55572376	<i>CETP</i>	A	G	0.35	1.46	1.43	1.36	-0.03	5.13E-09	5.75E-03	
rs5880	16	55572592	<i>CETP</i>	G	C	0.05	1.44	1.34	1.20	-0.08	6.90E-12	7.02E-03	
rs5882	16	55573593	<i>CETP</i>	A	G	0.32	1.41	1.44	1.50	0.03	2.78E-07	4.91E-03	
rs1800777	16	55574820	<i>CETP</i>	G	A	0.04	1.44	1.33	1.23	-0.09	8.46E-11	6.19E-03	
rs289743	16	55575297	<i>NLRCS5</i>	A	G	0.31	1.41	1.44	1.51	0.03	8.62E-09	6.38E-03	

HMZ, homozygote ; HTZ, heterozygote

Table S3e. All SNPs significantly associated with ApoAI

SNP	CHR	BP	GENE	Alleles		MAF	Common HMZ mean	HTZ mean g/l	Rare HMZ mean g/l	Beta	P-value	R2 (univariate)	Chr R2
				1	2								
rs301	8	19861214	<i>LPL</i>	A	G	0.25	2.12	2.15	2.21	0.04	4.16E-06	3.60E-03	0.14
rs1883025	9	106704122	<i>ABCA1</i>	G	A	0.26	2.16	2.12	2.09	-0.04	4.81E-06	4.29E-03	0.14
rs4775041	15	56461987	<i>LIPC</i>	C	G	0.30	2.12	2.14	2.22	0.04	2.39E-08	5.48E-03	0.15
rs11635491	15	56507033	<i>LIPC</i>	G	A	0.27	2.12	2.16	2.20	0.04	2.00E-07	5.34E-03	
rs1800588	15	56510967	<i>LIPC</i>	G	A	0.21	2.12	2.16	2.26	0.05	5.38E-10	7.47E-03	
rs2070895	15	56511231	<i>LIPC</i>	G	A	0.22	2.12	2.16	2.26	0.05	4.54E-10	7.55E-03	
rs8034802	15	56512084	<i>LIPC</i>	T	A	0.28	2.11	2.16	2.20	0.04	1.50E-08	6.52E-03	
rs8033940	15	56512134	<i>LIPC</i>	G	A	0.29	2.11	2.16	2.20	0.04	1.02E-08	6.60E-03	
rs261332	15	56514617	<i>LIPC</i>	G	A	0.21	2.12	2.16	2.27	0.05	6.48E-10	7.38E-03	
rs588136	15	56517790	<i>LIPC</i>	A	G	0.21	2.12	2.16	2.26	0.05	9.75E-10	7.51E-03	
rs261342	15	56518445	<i>LIPC</i>	C	G	0.22	2.12	2.16	2.26	0.06	2.85E-11	8.44E-03	
rs261341	15	56518859	<i>LIPC</i>	G	A	0.29	2.11	2.15	2.20	0.04	4.05E-08	5.85E-03	
rs261338	15	56522297	<i>LIPC</i>	G	A	0.17	2.12	2.16	2.27	0.04	1.73E-06	4.24E-03	
rs485671	15	56528426	<i>LIPC</i>	C	A	0.17	2.13	2.16	2.23	0.04	7.88E-06	3.51E-03	
rs485538	15	56528469	<i>LIPC</i>	A	G	0.17	2.12	2.16	2.23	0.04	4.32E-06	3.81E-03	
rs572410	15	56528676	<i>LIPC</i>	G	C	0.24	2.12	2.15	2.22	0.04	2.69E-06	4.24E-03	
rs261336	15	56529710	<i>LIPC</i>	A	G	0.16	2.12	2.16	2.24	0.04	2.64E-06	4.02E-03	
rs9989419	16	55542640	<i>CETP</i>	G	A	0.40	2.18	2.13	2.08	-0.05	2.59E-12	9.33E-03	0.16
rs12708967	16	55550712	<i>CETP</i>	A	G	0.19	2.16	2.10	2.02	-0.06	1.60E-13	9.84E-03	
rs3764261	16	55550825	<i>CETP</i>	C	A	0.33	2.10	2.16	2.21	0.06	6.19E-19	1.35E-02	
rs12720918	16	55551713	<i>CETP</i>	A	G	0.28	2.16	2.12	2.09	-0.04	1.15E-08	5.23E-03	
rs17231506	16	55552029	<i>CETP</i>	G	A	0.32	2.09	2.17	2.21	0.07	1.41E-19	1.39E-02	
rs1800775	16	55552737	<i>CETP</i>	C	A	0.49	2.08	2.14	2.19	0.05	8.75E-16	1.15E-02	
rs711752	16	55553712	<i>CETP</i>	G	A	0.43	2.08	2.15	2.20	0.06	2.19E-18	1.33E-02	
rs708272	16	55553789	<i>CETP</i>	G	A	0.43	2.08	2.15	2.20	0.06	1.02E-18	1.37E-02	
rs1864163	16	55554734	<i>CETP</i>	G	A	0.26	2.17	2.10	2.06	-0.06	8.48E-16	1.17E-02	
rs7203984	16	55556759	<i>CETP</i>	A	C	0.19	2.16	2.10	2.05	-0.06	1.19E-12	8.74E-03	

rs11508026	16	55556829	<i>CETP</i>	G	A	0.43	2.08	2.15	2.20	0.06	7.52E-18	1.26E-02	
rs12720922	16	55558386	<i>CETP</i>	G	A	0.18	2.16	2.09	2.05	-0.06	8.69E-13	8.97E-03	
rs9939224	16	55560233	<i>CETP</i>	C	A	0.20	2.16	2.10	2.03	-0.06	5.63E-14	1.01E-02	
rs1532625	16	55562802	<i>CETP</i>	G	A	0.44	2.08	2.15	2.20	0.06	2.61E-18	1.29E-02	
rs1532624	16	55562980	<i>CETP</i>	C	A	0.44	2.08	2.15	2.20	0.06	4.69E-18	1.26E-02	
rs11076175	16	55563879	<i>CETP</i>	A	G	0.17	2.16	2.09	2.04	-0.06	5.66E-13	9.44E-03	
rs7499892	16	55564091	<i>CETP</i>	G	A	0.17	2.16	2.09	2.04	-0.07	1.27E-13	9.97E-03	
rs11076176	16	55564947	<i>CETP</i>	A	C	0.16	2.15	2.11	2.06	-0.05	2.07E-07	4.56E-03	
rs289714	16	55564952	<i>CETP</i>	A	G	0.17	2.15	2.11	2.08	-0.04	1.14E-06	4.05E-03	
rs289716	16	55566877	<i>CETP</i>	A	T	0.31	2.11	2.14	2.22	0.04	1.06E-08	6.69E-03	
rs289717	16	55566889	<i>CETP</i>	G	A	0.35	2.16	2.13	2.09	-0.04	1.65E-07	4.94E-03	
rs289718	16	55567433	<i>CETP</i>	A	G	0.32	2.11	2.14	2.23	0.04	2.45E-09	7.60E-03	
rs289719	16	55567442	<i>CETP</i>	G	A	0.31	2.11	2.14	2.22	0.04	1.01E-08	6.58E-03	
rs4784744	16	55568686	<i>CETP</i>	G	A	0.35	2.16	2.13	2.09	-0.04	2.17E-07	4.69E-03	
rs291044	16	55568953	<i>CETP</i>	G	A	0.35	2.16	2.13	2.09	-0.04	2.62E-07	4.65E-03	
rs12720889	16	55570064	<i>CETP</i>	T	A	0.30	2.11	2.15	2.22	0.04	3.91E-09	6.97E-03	
rs4784745	16	55572376	<i>CETP</i>	A	G	0.35	2.16	2.13	2.08	-0.04	4.07E-07	4.53E-03	
rs5880	16	55572592	<i>CETP</i>	G	C	0.05	2.15	2.07	1.97	-0.08	6.81E-08	4.39E-03	
rs5882	16	55573593	<i>CETP</i>	A	G	0.32	2.12	2.14	2.21	0.04	5.42E-07	4.79E-03	
rs1800777	16	55574820	<i>CETP</i>	G	A	0.04	2.14	2.06	2.00	-0.09	1.79E-07	3.97E-03	
rs1801706	16	55575163	<i>CETP</i>	G	A	0.18	2.12	2.16	2.24	0.04	6.98E-06	4.25E-03	
rs289743	16	55575297	<i>NLRC5</i>	A	G	0.31	2.12	2.14	2.22	0.04	1.54E-08	6.30E-03	

HMZ, homozygote ; HTZ, heterozygote

Table S4. SNPs from published GWASs (Kathiresan, 2008 and Willer, 2008) that did not reach significance in the WHII study.

GWAS	Trait	Chr	Gene	SNP	Reported $\beta$ (sem) (meta-analysis)	Reported p-value (meta-analysis)	Reported $\beta$ (sem) (initial GWAS)	Reported p-value (initial GWAS)	p-value in WHII	
Kathiresan, 2008, Nat Genet	LDL	19q13.32	<i>APOE, APOC1, APOC4, APOC2</i>	rs4420638	0.19 (0.02)	1.00E-60	0.25 (0.03)	3.00E-13	not in chip	
	LDL	1p32.3	<i>PCSK9</i>	rs11591147	-0.47 (0.03)	2.00E-44	not typed		not in chip	
	LDL	19p13	<i>CILP2, PBX4</i>	rs16996148	-0.10 (0.02)	3.00E-08	-0.10 (0.05)	4.00E-02	6.01E-04	
	HDL	1q42.13	<i>GALNT2</i>	rs4846914	-0.07 (0.01)	2.00E-13	-0.10 (0.03)	3.00E-04	not in chip	
	HDL	9q31	<i>ABCA1</i>	rs3890182	-0.10 (0.02)	3.00E-10	-0.17 (0.04)	3.00E-05	1.71E-02	
	HDL	11q23	<i>APOA1-C3-A4-A5, ZNF259, BUD13</i>	rs28927680	-0.13 (0.03)	2.00E-05	-0.06 (0.05)	3.10E-01	1.58E-04	
	HDL	18q21	<i>LIPG, ACAA2</i>	rs2156552	-0.07 (0.01)	2.00E-07	-0.09 (0.03)	2.00E-02	1.44E-02	
	Triglycerides	1q42.13	<i>GALNT2</i>	rs4846914	0.08 (0.01)	7.00E-15	0.11 (0.03)	9.00E-05	not in chip	
	Triglycerides	1p31.3	<i>ANGPTL3, DOCK7, ATG4C</i>	rs12130333	-0.11 (0.02)	2.00E-08	-0.12 (0.03)	6.00E-04	not in chip	
	Triglycerides	7q11	<i>BCL7B, TBL2, MLXIPL</i>	rs17145738	-0.14 (0.02)	7.00E-22	-0.12 (0.03)	3.00E-03	1.04E-05	
	Willer, 2008, Nat Genet*	LDL	19q13.32	<i>APOE, APOC1, APOC4, APOC2</i>	rs4420638	6.61mg/dl	3.00E-43	8.02mg/dl	3.20E-21	not in chip
		LDL	19	<i>NCAN</i>	rs2228603			6.46mg/dl	1.80E-07	1.53E-05
		HDL	9	<i>GRIN3A, PPP3R2</i>	rs1323432	-0.03mg/dl	7.70E-04	1.93mg/dl	2.50E-08	1.98E-01
		HDL	9q31	<i>ABCA1</i>	rs4149274	0.82mg/dl	1.20E-10	1.51mg/dl	7.40E-08	not in chip
		HDL	18q21	<i>LIPG</i>	rs4939883			1.87mg/dl	1.40E-07	not in chip
		HDL	1q42.13	<i>GALNT2</i>	rs4846914			1.15mg/dl	2.90E-07	1.26E-02
		Triglycerides	11q23	<i>APOA5</i>	rs964184			18.12mg/dl	1.50E-16	not in chip
		Triglycerides	8p21	<i>LPL</i>	rs6993414			14.2mg/dl	1.40E-13	not in chip
		Triglycerides	8	<i>TRIB1</i>	rs2954029			6.42mg/dl	2.80E-08	1.54E-05
		Triglycerides	19	<i>NCAN</i>	rs10401969			12.28mg/dl	2.30E-07	not in chip

\*Data reported as per-allele change in mean lipid value (mg/dl).

## References

- (1) Kathiresan S, Melander O, Guiducci C, Surti A, Burtt NP, Rieder MJ et al. Six new loci associated with blood low-density lipoprotein cholesterol, high-density lipoprotein cholesterol or triglycerides in humans. *Nat Genet* 2008; 40:189-197.
- (2) Willer CJ, Sanna S, Jackson AU, Scuteri A, Bonnycastle LL, Clarke R et al. Newly identified loci that influence lipid concentrations and risk of coronary artery disease. *Nat Genet* 2008; 40:161-169

Table S5. Mean lipid values and gene count score for approximate quintiles of WHI individuals.

Traits	Number of individuals	Mean genescore	Trait mean	95CI-	95CI+
<b>LDL-C mmol/l</b>					
1	1,022	22.33	3.96	3.90	4.02
2	842	25.54	4.25	4.18	4.32
3	991	27.50	4.40	4.34	4.46
4	876	29.44	4.59	4.52	4.65
5	798	32.21	4.79	4.72	4.86
<b>ApoB g/l</b>					
1	1,031	23.47	1.12	1.10	1.14
2	975	26.54	1.21	1.19	1.23
3	1,114	28.50	1.27	1.25	1.28
4	948	30.45	1.30	1.28	1.32
5	540	32.96	1.38	1.36	1.41
<b>TG mmol/l</b>					
1	1,046	8.744741	1.05	1.01	1.08
2	1,155	11.5645	1.16	1.12	1.19
3	699	13	1.22	1.17	1.27
4	1,007	14.43694	1.30	1.26	1.35
5	701	17.28103	1.52	1.46	1.59
<b>HDL-C mmol/l</b>					
1	1,360	8.87	1.26	1.25	1.28
2	654	11.00	1.33	1.30	1.36
3	1,373	12.48	1.40	1.38	1.42
4	527	14.00	1.48	1.44	1.51
5	679	15.79	1.54	1.51	1.57
<b>ApoAI g/l</b>					
1	1,159	6.98	2.04	2.02	2.06
2	745	9.00	2.09	2.07	2.11
3	1,625	10.50	2.15	2.14	2.17
4	626	12.00	2.21	2.19	2.24
5	453	13.48	2.29	2.26	2.33
<b>ApoB/ApoAI ratio</b>					
1	1,221	20.39	0.54	0.53	0.55
2	1,012	23.51	0.58	0.57	0.59
3	560	25.00	0.61	0.60	0.62
4	976	26.45	0.64	0.63	0.65
5	839	29.05	0.67	0.66	0.69

Table S6. SNPs independently associating with lipid and apolipoprotein traits in genes/regions previously detected by GWASs or candidate gene studies in the WH-II study.

<i>SNP</i>	<i>Chr</i>	<i>Basepairs</i>	<i>Gene</i>	<i>cSNPs</i>	<i>MAF</i>	<i>Previous GWAS</i>	<i>Common HMZ<sup>c</sup> mean</i>	<i>HTZ<sup>d</sup> mean</i>	<i>Rare HMZ<sup>c</sup> mean</i>	<i>Beta</i>	<i>p-value<sup>a</sup></i>	<i>StepAIC<sup>b</sup>*</i>
<b>LDL-C</b>												
rs11591147	1	55278235	<i>PCSK9</i>	L46R	0.02	<sup>1</sup>	4.38	3.84	2.98	-0.55	9.28E-12	4.95E-12
rs4970834	1	109616403	<i>CELSR2</i>		0.18		4.42	4.27	4.08	-0.16	5.18E-09	1.16E-01
rs12740374	1	109619113	<i>CELSR2</i>		0.21	<sup>2</sup> ; LD with rs646776 from <sup>3</sup>	4.43	4.28	4.12	-0.15	1.82E-09	4.73E-04
rs629301	1	109619829	<i>CELSR2</i>		0.21	LD with rs12740374 from <sup>2</sup>	4.43	4.28	4.14	-0.15	6.50E-09	1.24E-03
rs693	2	21085700	<i>APOB</i>	T2488T	0.48		4.48	4.34	4.27	-0.11	3.86E-08	1.00E-01
rs934197	2	21120966	<i>APOB</i>		0.33		4.30	4.40	4.53	0.12	3.43E-08	1.16E-02
rs562338	2	21141826	<i>APOB</i>		0.18	<sup>4</sup> , <sup>5</sup> ; LD with rs515135 from <sup>2</sup> ; LD with rs506585 from <sup>3</sup>	4.43	4.24	4.16	-0.18	1.21E-11	8.86E-07
rs4299376	2	43926080	<i>ABCG8</i>		0.32	LD with rs6544713 from <sup>2</sup>	4.28	4.42	4.53	0.13	8.70E-10	1.33E-09
rs12916	5	74692295	<i>HMGCR</i>		0.41	LD with rs3846663 from <sup>2</sup> ; LD with rs12654264 <sup>1</sup>	4.26	4.39	4.51	0.12	6.66E-09	2.22E-05
rs3804231	5	74732535	<i>HMGCR</i>		0.13		4.33	4.46	4.62	0.14	5.15E-06	2.59E-02
rs2072560	11	116167036	<i>APOA5</i>		0.06		4.34	4.51	5.33	0.22	2.36E-07	2.39E-07
rs17231506	16	55552029	<i>CETP</i>		0.32		4.44	4.33	4.22	-0.11	5.02E-07	4.63E-07
rs1529729	19	11024562	<i>LDLR</i>		0.45		4.28	4.39	4.44	0.09	6.71E-06	1.13E-01
rs17248720	19	11059187	<i>LDLR</i>		0.13		4.45	4.14	3.81	-0.31	7.86E-25	2.00E-16
rs8110695	19	11067530	<i>LDLR</i>		0.22		4.43	4.27	4.21	-0.14	1.12E-08	1.51E-01
rs2228671	19	11071912	<i>LDLR</i>	C27C	0.13	<sup>6</sup>	4.42	4.21	4.18	-0.18	6.52E-10	1.08E-04
rs1531517	19	49934013	<i>BCL3</i>		0.07		4.40	4.17	4.08	-0.22	6.26E-08	1.23E-02
rs10402271	19	50021054	<i>BCL3/PVRL2</i>		0.33		4.26	4.44	4.53	0.15	2.06E-12	9.72E-06
rs519113	19	50068124	<i>PVRL2</i>		0.24		4.42	4.31	4.18	-0.12	9.37E-07	1.42E-02
rs6859	19	50073874	<i>PVRL2</i>		0.41		4.27	4.38	4.52	0.12	3.53E-08	2.95E-03
rs283813	19	50081014	<i>PVRL2</i>		0.07		4.39	4.23	3.70	-0.19	2.51E-06	6.49E-05
rs2075650	19	50087459	<i>TOMM40</i>		0.14	<sup>6</sup>	4.31	4.51	4.87	0.23	1.14E-14	2.45E-02
rs12721046	19	50113094	<i>APOC1</i>		0.15		4.30	4.51	4.74	0.21	7.58E-14	1.16E-01
rs12721109	19	50139061	<i>APOC4</i>		0.02		4.39	3.82	4.32	-0.54	5.06E-14	9.44E-08
<b>ApoB</b>												
rs11591147	1	55278235	<i>PCSK9</i>	L46R	0.02	<sup>1</sup> , <sup>3</sup>	1.28	1.14	0.94	-0.12	1.89E-10	1.08E-10
rs4970834	1	109616403	<i>CELSR2</i>		0.18		1.29	1.25	1.18	-0.04	2E-08	1.46E-01
rs12740374	1	109619113	<i>CELSR2</i>		0.21	<sup>2</sup>	1.29	1.25	1.19	-0.03	4.7E-09	2.13E-04
rs629301	1	109619829	<i>CELSR2</i>		0.21		1.29	1.25	1.19	-0.03	1.76E-08	6.14E-04
rs562338	2	21141826	<i>APOB</i>		0.18	<sup>5</sup> , <sup>4</sup>	1.29	1.24	1.19	-0.05	6E-13	4.77E-09
rs754523	2	21165196	<i>APOB</i>		0.33		1.25	1.27	1.32	0.02	1.11E-06	1.22E-02
rs4299376	2	43926080	<i>ABCG8</i>		0.32		1.25	1.28	1.32	0.03	7.6E-08	1.03E-07



rs1260326	2	27584444	GCKR	P446L	0.40	<sup>2, 7, 3</sup>		1.38	1.43	1.63	0.06	1.83E-07	2.21E-07
rs17145713	7	72542746	BAZ1B		0.20			1.50	1.36	1.25	-0.09	5.29E-10	6.05E-10
rs285	8	19859469	LPL		0.47			1.54	1.45	1.31	-0.07	2.65E-09	6.15E-05
rs331	8	19864685	LPL		0.27	LD with rs326 from <sup>8</sup>		1.51	1.39	1.24	-0.08	1.73E-11	1.40E-02
rs3289	8	19867472	LPL		0.03			1.43	1.70	1.04	0.16	3.16E-06	5.58E-06
rs3916027	8	19869148	LPL		0.27	LD with rs326 from <sup>8</sup>		1.51	1.40	1.24	-0.08	1.04E-10	2.55E-02
rs10503669	8	19891970	LPL		0.11	<sup>4</sup> , LD with rs10096633 from <sup>6</sup> , LD with rs12678919 from <sup>2</sup> , LD with rs328 from <sup>3</sup>		1.48	1.30	1.08	-0.11	1.05E-09	1.61E-02
rs17321515	8	126555591	TRIB1		0.47			1.53	1.43	1.37	-0.05	3.47E-06	4.62E-06
rs17108993	10	95354023	RBP4		0.03			1.42	1.72	1.53	0.14	8.04E-06	8.04E-06
rs6589565	11	116145447	BUD13		0.07			1.39	1.81	1.92	0.19	4.49E-20	5.73E-02
rs12286037	11	116157417	ZNF259		0.06	<sup>4</sup> ; LD with rs12272004 from <sup>6</sup> ; LD with rs28927680 from <sup>1</sup>		1.42	1.64	1.83	0.13	1.55E-09	1.01E-12
rs651821	11	116167789	APOA5		0.06	LD with rs662799 from <sup>3</sup>		1.39	1.81	2.04	0.21	8.89E-21	3.35E-03
rs10750097	11	116169250	APOA4		0.21			1.38	1.53	1.70	0.09	1.26E-10	5.46E-03
rs33989105	11	116205352	APOC3		0.25			1.40	1.47	1.67	0.07	1.75E-07	1.72E-03
rs5072	11	116212793	APOA1/ NCAN		0.08			1.41	1.60	1.84	0.11	1.74E-07	1.21E-01
rs2304128	19	19607151	GMIP		0.09			1.47	1.31	0.99	-0.10	3.15E-07	3.74E-07

<sup>a</sup>P-value for association adjusted for age and gender.

<sup>b</sup>P-value for the specified SNP in the final stepwise regression model, i.e. adjusted for age, gender and all other SNPs in this final model

<sup>c</sup>HMZ, homozygote ; <sup>d</sup> HTZ, heterozygote

LDL-C, low density lipoprotein –cholesterol; ApoB, apolipoprotein B; HDL-C, high density lipoprotein –cholesterol; ApoAI, apolipoprotein AI;

LD, linkage disequilibrium; StepAI.C, stepwise selection scheme using the Akaike's Information Criterion (AIC)<sup>9</sup>.

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Table S7. Results from the meta-analysis of SNPs with  $\text{MAF} > 0.01$  and p-value cut off of  $10^{-5}$  and their association with lipid and apolipoprotein traits



rs1049817	2	27404471	<i>GTF3C2</i>	A	G	0.39	1.51	1.43	1.33	-0.05	5.67E-05 <sup>#</sup>	0.004	1.45E-06
rs3134947	6	32253183	<i>RNF5</i>	G	A	0.22	1.47	1.40	1.35	-0.05	5.08E-05	0.003	6.21E-03
rs2854275	6	32736406	<i>HLA-DQB1</i>	C	A	0.14	1.47	1.38	1.33	-0.07	4.24E-05	0.004	8.33E-04
rs2781659	6	131933513	<i>ARG1;MED3</i>	A	G	0.32	1.49	1.43	1.30	-0.05	4.40E-05	0.003	3.81E-03
rs17108993	10	95354023	<i>RBP4</i>	C	G	0.03	1.42	1.72	1.53	0.14	8.04E-06	0.004	1.62E-03
rs163182	11	2800792	<i>KCNQ1</i>	C	G	0.45	1.38	1.43	1.56	0.05	4.32E-05	0.004	7.37E-03
rs163184	11	2803645	<i>KCNQ1</i>	A	C	0.47	1.37	1.44	1.55	0.05	2.91E-05	0.004	1.13E-02
rs2228603	19	19190924	<i>GATAD2A</i>	G	A	0.08	1.47	1.33	1.00	-0.09	1.53E-05	0.004	-
rs3794991	19	19471596	<i>PBX4</i>	G	A	0.09	1.47	1.32	1.08	-0.09	4.41E-06	0.004	8.48E-05
rs16996148	19	19519472	<i>PBX4</i>	C	A	0.08	1.47	1.31	1.09	-0.09	1.69E-06	0.005	4.92E-05
rs12610185	19	19582722	<i>PBX4</i>	G	A	0.09	1.47	1.31	1.09	-0.10	5.64E-07	0.005	2.31E-05
rs10500212	19	19584215	<i>GMIP</i>	G	A	0.09	1.47	1.31	1.09	-0.10	6.79E-07	0.005	2.53E-05
rs2304128	19	19607151	<i>GMIP</i>	C	A	0.09	1.47	1.31	0.99	-0.10	3.15E-07	0.005	4.23E-06

<sup>a</sup> P-value in replication study <0.05    <sup>b</sup> P-value in replication study <1x10<sup>-3</sup>    <sup>#</sup> P-value in replication study <1x10<sup>-5</sup>

<sup>c</sup> HMZ, homozygote ;<sup>d</sup> HTZ, heterozygote

Highlighted in bold are those SNPs which showed significant replication in the meta-analyses

LDL-C, low density lipoprotein –cholesterol; ApoB, apolipoprotein B, HDL-C, high density lipoprotein –cholesterol, ApoAI, apolipoprotein AI;

Chr, chromosome , BP, base pair; MAF, minor allele frequency