

## Supplementary Tables

**Table S1.** Linkage disequilibrium between pairs of variation sites observed along the coding region of the *HFE* gene in patients and healthy control individuals.

POPULATION	SNP 1	SNP 2	<i>P</i> value
<b>HH</b>	rs1799945 (H63DC>G)	rs2071303 (IVS2(+4)T>C)	0.010
	rs1799945 (H63DC>G)	rs1800562 (C282YG>A)	0.041
	rs2071303 (IVS2(+4)T>C)	rs1800562 (C282YG>A)	0.034
<b>HCV-IO<sup>+</sup></b>	rs1799945 (H63DC>G)	rs2071303 (IVS2(+4)T>C)	< 0.001
	rs2071303 (IVS2(+4)T>C)	rs1800708 (IVS4(−44)T>C)	< 0.001
<b>HCV-IO<sup>-</sup></b>	rs1799945 (H63DC>G)	rs2071303 (IVS2(+4)T>C)	< 0.001
	rs2071303 (IVS2(+4)T>C)	rs1800708 (IVS4(−44)T>C)	< 0.001
<b>HCC HCV-IO<sup>+</sup></b>	rs1799945 (H63DC>G)	rs2071303 (IVS2(+4)T>C)	0.001
	rs2071303 (IVS2(+4)T>C)	rs1800708 (IVS4(−44)T>C)	0.001
<b>HCC-IO<sup>?</sup></b>	rs2071303 (IVS2(+4)T>C)	rs1800708 (IVS4(−44)T>C)	0.015
<b>CTL</b>	rs1799945 (H63DC>G)	rs2071303 (IVS2(+4)T>C)	< 0.001
	rs2071303 (IVS2(+4)T>C)	rs1800708 (IVS4(−44)T>C)	< 0.001
<b>GLOBAL TEST</b>	rs1799945 (H63DC>G)	rs2071303 (IVS2(+4)T>C)	< 0.001
	rs807209 (G>C)	rs1800562 (C282YG>A)	< 0.001

Patients with hereditary hemochromatosis (HH), hepatitis C presenting (HCV-IO<sup>+</sup>) or not (HCV-IO<sup>-</sup>) iron overload (IO), hepatocellular carcinoma (HCC) associated with HCV-IO<sup>+</sup> (HCC HCV-IO<sup>+</sup>) and associated with other causes and presenting no information regarding iron overload (HCC-IO<sup>?</sup>), and healthy control individuals (CTL).

**Table S2.** Linkage disequilibrium among the H63DC>G (rs1799945), IVS2(+4)T>C (rs2071303) and C282YG>A (rs1800562) variations sites observed along the *HFE* gene in hereditary hemochromatosis (HH) and healthy control (CTL) individuals.

		OBSERVED FREQUENCY	EXPECTED FREQUENCY	STANDARDIZED VALUE OF DISEQUILIBRIUM ( <i>D'</i> )		STANDARDIZED VALUE OF CORRELATION ( <i>r</i> <sup>2</sup> )		QUI <sup>2</sup> VALUE	<i>P</i> VALUE OF QUI <sup>2</sup>	
HH	H63DC>G	IVS2(+4)T>C								
		T	C	T	C	T	C			
		24	1	21.43	3.57	1.0000	-1.0000	0.7200	20.1600	0.0000
	G	0	3	2.57	0.43	-1.0000	1.0000	0.7200	20.1600	0.0000
	H63DC>G	C282YG>A								
		G	A	G	A	G	A			
		5	20	7.14	17.86	-1.0000	1.0000	0.3000	8.4000	0.0038
	G	3	0	0.86	2.14	1.0000	-1.0000	0.3000	8.4000	0.0038
	IVS2(+4)T>C	C282YG>A								
		G	A	G	A	G	A			
4		20	6.86	17.14	-1.0000	1.0000	0.4267	11.6667	0.0006	
C	4	0	1.14	2.86	1.0000	-1.0000	0.4267	11.6667	0.0006	
CTL	H63DC>G	IVS2(+4)T>C								
		T	C	T	C	T	C			
		122	43	100.65	64.35	1.0000	-1.0000	0.3318	66.3559	0.0000
	G	0	35	21.35	13.65	-1.0000	1.0000	0.3318	66.3559	0.0000
	H63DC>G	C282YG>A								
		G	A	G	A	G	A			
		156	9	157.57	7.42	-1.000	1.000	0.0100	1.9990	0.1574
	G	35	0	33.42	1.57	1.000	-1.000	0.0100	1.9990	0.1574
	IVS2(+4)T>C	C282YG>A								
		G	A	G	A	G	A			
T	114	8	116.51	5.49	-0.7151	0.7151	0.0154	3.0811	0.0792	

	C	77	1	74.49	3.51	0.7151	-0.7151	0.0154	3.0811	0.0792
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Polymorphic sites were defined according to NCBI (<http://www.ncbi.nlm.nih.gov>) and Ensembl (<http://www.ensembl.org>) databases. Significant *P* values are shown in bold. Shaded cells are showing the pairs with higher LD.

**Table S3.** Linkage disequilibrium analyses encompassing the *HFE* gene (*HFE* allele, H63DC>G and C282YG>A) and classical and non-classical HLA loci.

<i>LOCUS 1</i>	<i>LOCUS 2</i>	<i>P</i> VALUE	STANDARD DEVIATION
<i>HLA-A</i>	<i>HFE</i>	0.22	0.03
<i>HLA-B</i>	<i>HFE</i>	0.59	0.04
<i>HLA-C</i>	<i>HFE</i>	0.33	0.04
<i>HLA-DRB1</i>	<i>HFE</i>	0.11	0.02
<i>HLA-DQB1</i>	<i>HFE</i>	0.51	0.03
<i>TNFA</i>	<i>HFE</i>	0.22	0.03
<i>TNFB</i>	<i>HFE</i>	0.49	0.03
<i>TNFC</i>	<i>HFE</i>	0.65	0.01
<i>TNFD</i>	<i>HFE</i>	0.35	0.03
<i>HLA-G 14pb</i>	<i>HFE</i>	0.54	0.01
<i>HLA-A</i>	H63D	0.09	0.01
<b><i>HLA-B</i></b>	<b>H63D</b>	<b>0.03</b>	<b>0.00</b>
<i>HLA-C</i>	H63D	0.15	0.01
<i>DRB1</i>	H63D	0.51	0.02
<i>DQB1</i>	H63D	0.31	0.01
<i>TNFA</i>	H63D	0.26	0.01
<i>TNFB</i>	H63D	0.90	0.00
<i>TNFC</i>	H63D	0.27	0.00
<i>TNFD</i>	H63D	0.67	0.01
<i>HLA-A</i>	C282Y	0.31	0.01
<i>HLA-B</i>	C282Y	0.57	0.01
<i>HLA-C</i>	C282Y	0.91	0.00
<i>DRB1</i>	C282Y	0.96	0.00
<i>DQB1</i>	C282Y	0.82	0.00
<i>TNFA</i>	C282Y	0.80	0.01
<i>TNFB</i>	C282Y	0.66	0.00
<i>TNFC</i>	C282Y	0.26	0.00
<i>TNFD</i>	C282Y	0.89	0.00
H63D	C282Y	0.79	0.00

Alleles (ensemble of coding region single nucleotide polymorphisms); SNPs (H63DC>G and C282YG>A); Classical HLA (*HLA-A*, *-B*, *-C*) loci; Non-classical HLA class I (*HLA-G 14bp* INDEL), class III (*TNFA-e* microsatellites), and class II (*HLA-DRB1* and *-DQB1*) loci. Significant results are shown in bold.