

Table S1. Genes and SNPs characteristics.

Gene	SNP ID	ChrPosition ^a	SNP	SNP Type
	rs11065987	12:111634620	A/G	N/A ^b
<i>SH2B3</i>	rs17696736	12:112049014	A/G	Intron Variant
	rs3184504	12:111446804	T/C	Missense Variant
	rs491552	6:150989919	C/G	Intron Variant
<i>MTHFD1L</i>	rs6922269	6:150931849	G/A	Intron Variant
	rs803422	6:150894458	A/G	Intron Variant
	rs803455	6:150926303	A/G	Intron Variant
<i>GGCX</i>	rs28928872	2:85551919	C/G	Missense Variant
<i>ITGB3</i>	rs398122372	17:47307581	G/A G/C	Missense Variant
	rs398122374	17:47307567	T/A T/C T/G	Missense Variant

^a. Chromosome positions are based on NCBI Human Genome Assembly Build.

^bN/A: Not Available.

Table S2. List of SNPs, their minor allele frequencies, and HWE p-values.

Gene	SNP ID	MA ^a	Patients MAF ^b	Controls MAF ^b	HWE ^c <i>P</i> -value
	rs11065987	G	0.39	0.37	0.03
<i>SH2B3</i>	rs17696736	G	0.39	0.36	0.05
	rs3184504	T	0.4	0.37	0.03
	rs491552	C	0.47	0.46	0.08
<i>MTHFD1L</i>	rs6922269	A	0.27	0.25	0.62
	rs803422	A	0.29	0.23	0.05
	rs803455	A	0.07	0.11	0.36

^a. MA: Minor allele.

^b. MAF: Minor allele frequency.

^c. HWE: Hardy-Weinberg equilibrium.

Table S3. The distributions of *SH2B3* and *MTHFD1L* haplotypes and 211 cardiovascular patients in compare to 213 healthy controls.

Gene	Haplotypes	Patients (%)	Controls (%)	Odds ratio (95% CI)	<i>P</i> -value*
<i>SH2B3</i>	AAC	0.59	0.62	1.00	---
	GGT	0.39	0.35	1.13 (0.86 - 1.48)	0.37
	AAT	0.01	0.007	2.05 (0.50 - 8.33)	0.32
<i>MTHFD1L</i>	GGGG	0.23	0.30	1.00	---
	CGGG	0.21	0.19	1.26 (0.78 - 2.04)	0.35
	CAGG	0.11	0.09	1.35 (0.68 - 2.68)	0.4
	GAGG	0.11	0.09	1.39 (0.76 - 2.53)	0.29
	GGAG	0.13	0.07	1.98 (1.04 - 3.78)	0.04
	CGAG	0.01	0.09	1.41 (0.77 - 2.56)	0.26
	CGGA	0.03	0.04	1.16 (0.46 - 2.91)	0.75
	GAAG	0.04	0.02	3.50 (0.79 - 15.53)	0.1
	GGGA	0.03	0.03	0.71 (0.12 - 4.28)	0.71
	CAAG	0.01	0.03	0.42 (0.05 - 3.33)	0.41
	GGAA	0.0	0.02	0.44 (0.01 - 12.88)	0.63

*Chi-Square Test with $p < 0.05$ is considered significant.

Table S4. The distributions of *SH2B3* and *MTHFD1L* haplotypes among 212 warfarin sensitive patients.

Gene	Haplotypes	Frequency (%)	Odds ratio (95% CI)	P-value*
<i>SH2B3</i>	AAC	0.47	0.00	---
	GGT	0.26	0.02 (-0.11 - 0.14)	0.77
	AGC	0.13	0.04 (-0.14 - 0.21)	0.67
	GAT	0.13	0.06 (-0.12 - 0.24)	0.52
	AAT	0.01	-0.01 (-0.47 - 0.45)	0.98
<i>MTHFD1L</i>	GGGG	0.26	0.00	---
	CGGG	0.17	-0.15 (-0.36 - 0.06)	0.16
	CAGG	0.13	0.14 (-0.09 - 0.36)	0.23
	GGAG	0.13	-0.09 (-0.33 - 0.16)	0.5
	CGAG	0.10	-0.09 (-0.3 - 0.12)	0.4
	GAGG	0.1	-0.18 (-0.47 - 0.11)	0.22
	CGGA	0.03	-0.22 (-0.62 - 0.17)	0.27
	CAAG	0.03	-0.05 (-0.49 - 0.38)	0.81
	GAAG	0.02	0.05 (-0.53 - 0.62)	0.87
	GGGA	0.02	-0.2 (-0.68 - 0.27)	0.4
	CGAA	0.01	-0.13 (-0.65 - 0.39)	0.62

*Chi-Square Test with $p < 0.05$ is considered significant.

Table S5. Post Hoc tests for the association of *SH2B3* and *MTHFD1L* SNPs with variability on warfarin required doses.

Gene	SNP ID	Genotype		Initiation Dose <i>P</i> -value*	Maintenance Dose <i>P</i> -value*
<i>SH2B3</i>	rs11065987	AA	GA	0.88	0.70
			GG	0.09	1
		GA	AA	0.88	0.70
			GG	0.18	0.83
	rs17696736	GG	AA	0.09	1
			GA	0.18	0.83
		AA	AG	0.98	0.99
			GG	0.1	1
		AG	AA	0.98	0.99
			GG	0.13	0.99
		GG	AA	0.1	1
			AG	0.13	0.99
	rs3184504	CC	TC	0.75	0.39
			TT	0.14	1
		TC	CC	0.75	0.39
			TT	0.37	0.52
		TT	TC	0.14	1
			TT	0.37	0.52
	rs491552	CC	CG	0.75	0.99
			GG	0.60	0.96
		CG	CC	0.75	0.99
			GG	0.92	0.86
		GG	CC	0.60	0.96
			CG	0.92	0.86
		AA	AG	0.28	0.24
			GG	1	0.98
<i>MTHFD1L</i>	rs6922269	AG	AA	0.28	0.24
			GG	0.01	0.003
		GG	AA	1	0.98
			AG	0.01	0.003
		AA	GA	0.93	0.91
			GG	1	0.86
	rs803422	GA	AA	0.93	0.91
			GG	0.77	0.98
		GG	AA	1	0.86
			GA	0.77	0.98

*Post-Hoc Multiple comparisons Test with $p < 0.05$ is considered significant. Compare means of the initiation and maintenance dose among all genotypes.

Post hoc tests are not performed for rs803455 because at least one group has fewer than two cases.

Table S6.The distributions of *SH2B3* and *MTHFD1L* haplotypes among 212 warfarin responsiveness patients.

Gene	Haplotypes	Frequency (%)	Odds ratio (95% CI)	P-value*
<i>SH2B3</i>	AAC	0.47	0.00	---
	GGT	0.26	0 (-0.1 - 0.1)	0.99
	AGC	0.13	0.22 (0.08 - 0.36)	0.002
	GAT	0.13	0.23 (0.09 - 0.38)	0.002
<i>MTHFD1L</i>	GGGG	0.26	0.00	---
	CGGG	0.19	-0.14 (-0.29 - 0.02)	0.085
	GGAG	0.12	-0.09 (-0.26 - 0.07)	0.28
	GAGG	0.12	-0.02 (-0.23 - 0.18)	0.83
	CAGG	0.10	-0.05 (-0.23 - 0.14)	0.61
	CGAG	0.10	-0.18 (-0.38 - 0.02)	0.08
	CGGA	0.03	0.09 (-0.16 - 0.35)	0.48
	CAAG	0.03	-0.05 (-0.35 - 0.24)	0.73
	GAAG	0.02	0.1 (-0.28 - 0.47)	0.61
	GGGA	0.02	0.08 (-0.27 - 0.42)	0.67
	CGAA	0.01	-0.38 (-0.92 - 0.16)	0.17

*Chi-Square Test with p<0.05 is considered significant.

Table S7.Post Hoc Tests for the Association of *SH2B3* and *MTHFD1L* SNPs with INR Treatment Outcome.

Gene	SNP ID	Genotype		Initiation INR <i>P</i> -value*	Maintenance INR <i>P</i> -value*
<i>SH2B3</i>	rs11065987	AA	GA	0.68	1
			GG	0.96	0.82
		GA	AA	0.68	1
			GG	0.92	0.81
		GG	AA	0.96	0.82
			GA	0.92	0.81
	rs17696736	AA	AG	0.50	1
			GG	0.94	0.89
		AG	AA	0.50	1
			GG	0.85	0.85
		GG	AA	0.94	0.89
			AG	0.85	0.85
	rs3184504	CC	TC	0.70	1
			TT	0.94	0.76
		TC	CC	0.70	1
			TT	0.95	0.77
		TT	CC	0.94	0.76
			TC	0.95	0.77
<i>MTHFD1L</i>	rs491552	CC	CG	0.86	0.75
			GG	0.17	0.24
		CG	CC	0.86	0.75
			GG	0.24	0.01
		GG	CC	0.17	0.24
			CG	0.24	0.01
		AA	AG	0.85	1
			GG	0.72	0.88
	rs6922269	AG	AA	0.85	1
			GG	0.92	0.65
		GG	AA	0.72	0.88
			AG	0.92	0.65
		AA	GA	0/34	0.75
			GG	0.75	0.95
		GA	AA	0.34	0.75
			GG	0.50	0.75
	rs803422	GG	AA	0.75	0.95
			GA	0.50	0.75
			AA	0.75	0.95
			GA	0.50	0.75

*Post-Hock Multiple comparisons Test with $p < 0.05$ is considered significant. Compare initiation and maintenance dose among all genotypes.
Post hoc tests are not performed for rs803455 because at least one group has fewer than two cases.