

Supplementary table 1. Information about the different SNPs studied

	Gene polymorphism	rs code	Chromosomal Location
ACE	A239T	ACE rs4291 A/T	17q23.3, promoter region
	A7941G	ACE rs4318 A/G	17q23.3, exon1, codon 1
	A10539G	ACE rs4335 A/G	17q23.3, intron 15
	A11599G	ACE rs4343 A/G	17q23.3, exon 16, codon 3
	A11599G	ACE rs4344 A/G	17q23.3, intron 17
	A15990G	ACE rs4353 A/G	17q23.3, intron 19
	A20060G	ACE rs4363 A/G	17q23.3, intron 13
AGTR1	A-777T	AGTR1 rs275651 A/T	3q24, promoter region
	G-680T	AGTR1 rs275652 G/T	3q24, promoter region
	A-119G	AGTR1 rs275653 A/G	3q24, promoter region
	C43732T	AGTR1 rs5182 T/C	3q24, exon 3, codon 3
	A44221G	AGTR1 rs5183 A/G	3q24, exon 3, codon 3
Endothelin 1	-1394 T/G	Edn1 rs1800541 G/T	6p24.1
	Glu105Glu	Edn1 rs5369 A/G	6p24.1

-974 C/A

Edn1 rs3087459

6p24.1

A/C

Supplementary table 2. Associations of SNPs with presence of systemic sclerosis (SSc)

Gene	SSc	Healthy	P value	OR	95% CI
polymorphism	patients (n=170 patients/340 alleles)	controls (n=199 controls/ 398 alleles)			
ACE rs4291 A/T (A239T)					
Allele	N (a.f.)	N (a.f.)			
A	247 (0.725)	301 (0.757)	0.6	0.82	0.39-1.72
T	93 (0.275)	97 (0.243)	0.4	1.18	0.78-1.78
Genotype	N (g.f.)	N (g.f.)			
AA	93 (0.545)	117 (0.589)	0.42	0.84	0.55-1.27
AT	61 (0.36)	67 (0.335)	0.73	1.1	0.71-1.69
TT	16 (0.096)	15 (0.076)	0.64	1.2	0.61-2.66
ACE rs4318 A/G (A7941G)					
Allele	N (a.f.)	N (a.f.)			
A	337 (0.991)	394 (0.99)	1.0	1.1	0.21-6.45
G	3 (0.009)	4 (0.01)	1.0	0.88	0.16-4.6
Genotype	N (g.f.)	N (g.f.)			
AA	167 (0.982)	195 (0.98)	1.0	1.1	0.21-6.5
AG	3 (0.018)	4 (0.02)	1.0	0.88	0.15-4.7
GG	0	0	-	-	-
ACE rs4335 A/G (A10539G)					

Allele	N (a.f.)	N (a.f.)			
A	213 (0.627)	259 (0.65)	0.66	1.14	0.63-2.03
G	127 (0.373)	139 (0.35)	0.13	1.40	0.9-2.12
Genotype	N (g.f.)	N (g.f.)			
AA	67 (0.394)	91 (0.454)	0.26	0.77	0.5-1.19
AG	79 (0.466)	77 (0.389)	0.16	1.38	0.89-2.13
GG	24 (0.14)	31 (0.157)	0.80	0.89	0.4-1.65
ACE rs4343 A/G (A11599G)					
Allele	N (a.f.)	N (a.f.)			
A	214 (0.629)	261 (0.656)	0.72	1.1	0.62-1.97
G	126 (0.371)	137 (0.344)	0.23	1.28	0.84-1.92
Genotype	N (g.f.)	N (g.f.)			
AA	68 (0.40)	93 (0.464)	0.23	0.76	0.49-1.17
AG	78 (0.46)	75 (0.38)	0.13	1.4	0.90-2.17
GG	24 (0.14)	31 (0.156)	0.80	0.89	0.4-1.6
ACE rs4344 A/G (A11599G)					
Allele	N (a.f.)	N (a.f.)			
A	203 (0.598)	253 (0.635)	0.97	0.99	0.57-1.69
G	137 (0.402)	145 (0.365)	0.23	1.28	0.84-1.95
Genotype	N (g.f.)	N (g.f.)			
AA	64 (0.376)	89 (0.449)	0.38	0.81	0.53-1.26
AG	75 (0.444)	75 (0.373)	0.25	1.31	0.84-2.02
GG	31 (0.18)	35 (0.178)	0.97	1.0	0.59-1.84

ACE rs4353 A/G (A15990G)

Allele	N (a.f.)	N (a.f.)			
A	136 (0.40)	148 (0.372)	0.47	1.13	0.83-1.53
G	204 (0.60)	250 (0.628)	0.47	0.89	0.65-1.21
Genotype	N (g.f.)	N (g.f.)			
AA	31 (0.18)	35 (0.178)	0.97	1.05	0.59-1.84
AG	74 (0.438)	78 (0.39)	0.46	1.20	0.77-1.85
GG	65 (0.382)	86 (0.432)	0.38	0.81	0.52-1.26

ACE rs4363 A/G (A200060G)

Allele	N (a.f.)	N (a.f.)			
A	213 (0.626)	251 (0.63)	0.69	1.15	0.64-1.92
G	127 (0.374)	147 (0.37)	0.67	1.09	0.72-1.66
Genotype	N (g.f.)	N (g.f.)			
AA	71 (0.416)	87 (0.438)	0.78	0.92	0.60-1.43
AG	71 (0.421)	77 (0.387)	0.62	1.14	0.73-1.76
GG	28 (0.163)	35 (0.175)	0.88	0.92	0.52-1.65

AGTR1 rs275651 A/T (A777T)

Allele	N (a.f.)	N (a.f.)			
A	27 (0.078)	32 (0.080)	0.79	0.92	0.51-1.66
T	313 (0.921)	366 (0.919)	0.68	0.65	0.10-3.97
Genotype	N (g.f.)	N (g.f.)			
AA	3 (0.017)	2 (0.011)	0.66	1.77	0.24-15.30
AT	21 (0.124)	28 (0.139)	0.74	0.86	0.45-1.64

TT	146 (0.860)	169 (0.850)	0.91	1.08	0.58-2.01
AGTR1 rs275652 G/T (G-680T)					
Allele	N (a.f.)	N (a.f.)			
G	28 (0.082)	30 (0.074)	0.81	1.07	0.59-1.94
T	312 (0.918)	368 (0.925)	1	0.64	0.10-3.92
Genotype	N (g.f.)	N (g.f.)			
GG	3 (0.017)	2 (0.01)	0.66	1.77	0.24-15.3
GT	22 (0.129)	26 (0.13)	0.90	0.99	0.52-1.89
TT	145 (0.854)	171 (0.86)	0.98	0.95	0.51-1.77
AGTR1 rs275653 A/G (A-119G)					
Allele	N (a.f.)	N (a.f.)			
A	310 (0.913)	365 (0.916)	0.7	1.34	0.29-6.08
G	30 (0.087)	33 (0.084)	0.76	1.09	0.61-1.95
Genotype	N (g.f.)	N (g.f.)			
AA	143 (0.843)	170 (0.854)	0.83	0.9	0.49-1.66
AG	24 (0.14)	25 (0.124)	0.77	1.14	0.6-2.18
GG	3 (0.017)	4 (0.022)	1.0	0.88	0.15-4.70
AGTR1 rs5182 T/C (C43732T)					
Allele	N (a.f.)	N (a.f.)			
T	205 (0.604)	211 (0.53)	0.16	1.45	0.86-2.45
C	135 (0.396)	187 (0.47)	0.07	0.67	0.43-1.04
Genotype	N (g.f.)	N (g.f.)			
TT	64 (0.376)	58 (0.29)	0.10	1.47	0.93-2.32

TC	77 (0.455)	95 (0.48)	0.71	0.91	0.59-1.40
CC	29 (0.169)	46 (0.23)	0.18	0.68	0.39-1.18
AGTR1 rs5183 A/G (A44221G)					
Allele	N (a.f.)	N (a.f.)			
A	333 (0.98)	381 (0.957)	0.13	2.12	0.82-5.71
G	7 (0.02)	17 (0.043)	0.13	0.45	0.18-1.22
Genotype	N (g.f.)	N (g.f.)			
AA	163 (0.961)	182 (0.916)	0.13	2.18	0.82-5.95
AG	7 (0.039)	17 (0.084)	0.13	0.46	0.17-1.21
GG	0	0	-	-	-
Edn1 RS1800541 G/T (-1394 T/G)					
Allele	N (a.f.)	N (a.f.)			
G	34 (0.10)	38 (0.095)	0.79	1.07	0.62-1.86
T	306 (0.90)	360 (0.905)	1.0	1.03	0.06-16.71
Genotype	N (g.f.)	N (g.f.)			
GG	1 (0.006)	1 (0.005)	1.0	1.17	0-43.1
GT	32 (0.188)	36 (0.178)	0.96	1.05	0.6-1.84
TT	137 (0.806)	162 (0.814)	0.94	0.95	0.55-1.65
Edn1 rs5369 A/G (Glu105Glu)					
Allele	N (a.f.)	N (a.f.)			
A	21 (0.0617)	15 (0.037)	0.17	1.68	0.79-3.57
G	319 (0.938)	383 (0.963)	1	0.97	0.94-1.04
Genotype	N (g.f.)	N (g.f.)			

AA	1 (0.006)	0	0.46	-	-
AG	19 (0.112)	15 (0.074)	0.30	1.54	0.72-3.32
GG	150 (0.882)	184 (0.926)	0.22	0.61	0.29-1.3
Edn1 rs3087459 A/C (-974 C/A)					
Allele	N (a.f.)	N (a.f.)			
A	307 (0.903)	355 (0.893)	1	1.03	0.06-16.71
C	33 (0.097)	43 (0.107)	0.68	0.88	0.51-1.52
Genotype	N (g.f.)	N (g.f.)			
AA	138 (0.812)	157 (0.79)	0.67	1.15	0.67-1.99
AC	31 (0.182)	41 (0.205)	0.65	0.86	0.69-1.23
CC	1 (0.006)	1 (0.005)	1.0	1.17	0-43.16

a.f.: allele frequency, g.f.: genotype frequency

Supplementary Table 3. Associations of SNPs with Systemic Sclerosis (SSc) subset

Gene	dcSSc patients (n=77 patients/154 alleles)	lcSSc patients (n=93 patients/ 186 alleles)	P value	OR	95% CI
ACE rs4291 A/T (A239T)					
Allele	N (a.f.)	N (a.f.)			
A	123 (0.799)	125 (0.674)	0.012*	1.94	1.14-3.29
T	31 (0.201)	61 (0.325)	0.012*	0.52	0.30-0.88
Genotype	N (g.f.)	N (g.f.)			
AA	51 (0.657)	42 (0.455)	0.006*	2.48	1.26-4.89
AT	21 (0.284)	41 (0.438)	0.03*	0.48	0.24-0.95
TT	5 (0.06)	10 (0.107)	0.48	0.58	0.16-1.95
ACE rs4318 A/G (A7941G)					
Allele	N (a.f.)	N (a.f.)			
A	152 (0.987)	178 (1.0)	0.21	0	0-3.52
G	2 (0.013)	0	0.21	-	-
Genotype	N (g.f.)	N (g.f.)			
AA	75 (0.974)	89 (1.0)	0.21	0	0-3.50
AG	2 (0.026)	0	0.21	-	-
GG	0	0	-	-	-
ACE rs4335 A/G (A10539G)					

Allele	N (a.f.)	N (a.f.)			
A	99 (0.641)	114 (0.612)	0.50	1.37	0.54-3.5
G	55 (0.358)	72 (0.387)	0.79	0.91	0.47-1.75
Genotype	N (g.f.)	N (g.f.)			
AA	31 (0.403)	36 (0.382)	0.96	1.07	0.55-2.07
AG	37 (0.478)	42 (0.461)	0.82	1.12	0.59-2.15
GG	9 (0.119)	15 (0.157)	0.54	0.69	0.26-1.81
ACE rs4343 A/G (A11599G)					
Allele	N (a.f.)	N (a.f.)			
A	100 (0.65)	114 (0.612)	0.64	1.37	0.54-3.5
G	54 (0.35)	72 (0.388)	0.65	0.86	0.45-1.64
Genotype	N (g.f.)	N (g.f.)			
AA	32 (0.418)	36 (0.382)	0.82	1.13	0.58-2.18
AG	36 (0.463)	42 (0.461)	0.95	1.07	0.56-2.05
GG	9 (0.119)	15 (0.157)	0.54	0.69	0.26-1.81
ACE rs4344 A/G (A11599G)					
Allele	N (a.f.)	N (a.f.)			
A	94 (0.612)	112 (0.601)	0.85	1.08	0.47-2.45
G	60 (0.388)	74 (0.399)	0.98	1.006	0.52-1.92
Genotype	N (g.f.)	N (g.f.)			
AA	31 (0.403)	37 (0.393)	0.92	1.02	0.53-1.98
AG	32 (0.418)	38 (0.412)	0.94	1.03	0.53-1.99
GG	14 (0.179)	18 (0.195)	0.99	0.93	0.40-2.15

ACE rs4353 A/G (A15990G)					
Allele	N (a.f.)	N (a.f.)			
A	62 (0.403)	74 (0.399)	0.98	1.02	0.64-1.61
G	92 (0.597)	112 (0.601)	0.98	0.98	0.62-1.55
Genotype	N (g.f.)	N (g.f.)			
AA	15 (0.194)	17 (0.18)	0.99	1.08	0.47-2.5
AG	32 (0.418)	40 (0.438)	0.97	0.94	0.49-1.82
GG	30 (0.388)	36 (0.382)	0.90	1.01	0.52-1.97
ACE rs4363 A/G (A200060G)					
Allele	N (a.f.)	N (a.f.)			
A	95 (0.62)	116 (0.623)	0.94	1.03	0.44-2.41
G	59 (0.38)	70 (0.377)	0.87	1.05	0.55-2.0
Genotype	N (g.f.)	N (g.f.)			
AA	31 (0.403)	39 (0.42)	0.94	0.93	0.48-1.81
AG	33 (0.433)	38 (0.41)	0.91	1.09	0.56-2.10
GG	13 (0.164)	16 (0.169)	0.88	0.98	0.41-2.34
AGTR1 rs275651 A/T (A777T)					
Allele	N (a.f.)	N (a.f.)			
A	11 (0.07)	15 (0.078)	0.99	0.99	0.39-2.51
T	143 (0.93)	171 (0.921)	0.5	1.77	1.54-2.03
Genotype	N (g.f.)	N (g.f.)			
AA	0	2 (0.022)	0.60	-	-
AT	11 (0.14)	11 (0.115)	0.80	1.24	0.47-3.32

TT	66 (0.86)	80 (0.862)	0.86	0.98	0.38-2.52
AGTR1 rs275652 G/T (G-680T)					
Allele	N (a.f.)	N (a.f.)			
G	10 (0.067)	16 (0.084)	0.83	0.9	0.36-2.26
T	144 (0.933)	170 (0.915)	0.5	1.77	1.54-2.03
Genotype	N (g.f.)	N (g.f.)			
GG	0	2 (0.022)	0.67	-	-
GT	10 (0.134)	12 (0.124)	0.83	1.01	0.37-2.69
TT	67 (0.866)	79 (0.854)	0.86	1.19	0.46-3.11
AGTR1 rs275653 A/G (A-119G)					
Allele	N (a.f.)	N (a.f.)			
A	142 (0.925)	169 (0.91)	0.5	1.77	1.54-2.03
G	12 (0.075)	17 (0.09)	0.89	0.94	0.38-2.26
Genotype	N (g.f.)	N (g.f.)			
AA	65 (0.85)	78 (0.843)	0.90	1.04	0.42-2.58
AG	12 (0.15)	13 (0.135)	0.93	1.14	0.45-2.87
GG	0	2 (0.022)	0.50	-	-
AGTR1 rs5182 T/C (C43732T)					
Allele	N (a.f.)	N (a.f.)			
T	100 (0.65)	110 (0.59)	0.89	1.06	0.44-2.56
C	54 (0.35)	76 (0.41)	0.16	0.62	0.32-1.2
Genotype	N (g.f.)	N (g.f.)			
TT	34 (0.448)	32 (0.35)	0.25	1.51	0.77-2.94

TC	32 (0.403)	46 (0.49)	0.38	0.73	0.38-1.40
CC	11 (0.149)	15 (0.16)	0.90	0.87	0.34-2.17
AGTR1 rs5183 A/G (A44221G)					
Allele	N (a.f.)	N (a.f.)			
A	153 (0.992)	181 (0.972)	0.22	4.23	0.47-96.6
G	1 (0.007)	5 (0.028)	0.23	0.25	0.02-2.23
Genotype	N (g.f.)	N (g.f.)			
AA	76 (98.5)	88 (0.944)	0.23	4.32	0.47-99.8
AG	1 (0.015)	5 (0.056)	0.22	0.23	0.01-2.11
GG	0	0	-	-	-
Edn1 RS1800541 G/T (-1394 T/G)					
Allele	N (a.f.)	N (a.f.)			
G	16 (0.106)	19 (0.101)	1	0.96	0.43-2.16
T	138 (0.894)	167 (0.899)	0.44	2.29	1.91-2.75
Genotype	N (g.f.)	N (g.f.)			
GG	1 (0.015)	0	0.67	-	-
GT	14 (0.182)	19 (0.202)	0.86	0.87	0.37-1.99
TT	62 (0.803)	74 (0.798)	0.96	1.06	0.47-2.42
Edn1 rs5369 A/G (Glu105Glu)					
Allele	N (a.f.)	N (a.f.)			
A	9 (0.06)	9 (0.047)	0.60	1.31	0.46-3.69
G	145 (0.94)	177 (0.953)	0.86	0.82	0.29-2.32
Genotype	N (g.f.)	N (g.f.)			

AA	0	0	-	-	-
AG	9 (0.12)	9 (0.095)	0.86	1.24	0.42-3.62
GG	68 (0.88)	84 (0.905)	0.86	0.81	0.28-2.37
Edn1 rs3087459 A/C (-974 C/A)					
Allele	N (a.f.)	N (a.f.)			
A	139 (0.901)	167 (0.899)	0.44	2.29	1.91-2.75
C	15 (0.098)	19 (0.101)	0.75	0.87	0.38-1.99
Genotype	N (g.f.)	N (g.f.)			
AA	63 (0.818)	74 (79.8)	0.86	1.16	0.5-2.67
AC	13 (0.167)	19 (0.202)	0.69	0.79	0.34-1.84
CC	1 (0.015)	0	0.45	-	-

*p value after logistic regression analysis was non-significant, dcSSc: diffuse cutaneous systemic sclerosis, lcSSc: limited cutaneous systemic sclerosis, a.f.: allele frequency, g.f.: genotype frequency

Supplementary Table 4. Associations of SNPs with severe vascular involvement ("Medsger severity scale" scores 3-4)

Gene	Severe	No severe	P value	OR	95% CI
polymorphism	vascular involvement (n= 79 patients/ 158 alleles)	vascular involvement (n= 80 patients/ 160 alleles)			
ACE rs4291 A/T (A239T)					
Allele	N (a.f.)	N (a.f.)			
A	119 (0.753)	112 (0.70)	0.27	1.87	0.6-5.87
T	39 (0.247)	48 (0.30)	0.37	0.75	0.40-1.41
Genotype	N (g.f.)	N (g.f.)			
AA	46 (0.582)	41 (0.513)	0.46	1.33	0.68-2.60
AT	27 (0.342)	30 (0.375)	0.78	0.87	0.43-1.74
TT	6 (0.076)	9 (0.113)	0.60	0.65	0.19-2.13
ACE rs4318 A/G (A7941G)					
Allele	N (a.f.)	N (a.f.)			
A	156 (0.987)	160 (1.0)	0.24	-	-
G	2 (0.013)	0	-	-	-
Genotype	N (g.f.)	N (g.f.)			
AA	77 (0.975)	80 (1.0)	0.24	-	-
AG	2 (0.025)	0	-	-	-
GG	0	0	-	-	-

ACE rs4335 A/G (A10539G)

Allele	N (a.f.)	N (a.f.)			
A	103 (0.651)	96 (0.60)	0.12	2.04	0.81-5.14
G	55 (0.349)	64 (0.40)	0.82	0.92	0.49-1.75
Genotype	N (g.f.)	N (g.f.)			
AA	32 (0.405)	31 (0.388)	0.94	1.08	0.54-2.14
AG	39 (0.494)	34 (0.425)	0.47	1.32	0.67-2.59
GG	8 (10.1)	15 (0.188)	0.18	0.49	0.18-1.33

ACE rs4343 A/G (A11599G)

Allele	N (a.f.)	N (a.f.)			
A	104 (0.658)	96 (0.60)	0.12	2.04	0.81-5.14
G	54 (0.342)	64 (0.40)	0.69	0.88	0.46-1.66
Genotype	N (g.f.)	N (g.f.)			
AA	33 (0.418)	31 (0.388)	0.82	1.13	0.57-2.25
AG	38 (0.481)	34 (0.425)	0.58	1.25	0.64-2.46
GG	8 (0.101)	15 (0.188)	0.18	0.49	0.18-1.33

ACE rs4344 A/G (A11599G)

Allele	N (a.f.)	N (a.f.)			
A	100 (0.633)	93 (0.582)	0.23	1.62	0.72-3.63
G	58 (0.367)	67 (0.418)	0.58	0.83	0.44-1.57
Genotype	N (g.f.)	N (g.f.)			
AA	33 (0.418)	31 (0.388)	0.82	1.13	0.57-2.25
AG	34 (0.43)	31 (0.388)	0.69	1.19	0.60-2.36

GG	12 (0.152)	18 (0.225)	0.32	0.62	0.25-1.48
ACE rs4353 A/G (A15990G)					
Allele	N (a.f.)	N (a.f.)			
A	61 (0.386)	66 (0.412)	0.71	0.90	0.56-1.44
G	97 (0.614)	94 (0.587)	0.71	1.12	0.69-1.79
Genotype	N (g.f.)	N (g.f.)			
AA	14 (0.177)	16 (0.20)	0.86	0.86	0.36-2.05
AG	33 (0.418)	34 (0.425)	0.94	0.97	0.49-1.91
GG	32 (0.405)	30 (0.375)	0.82	1.13	0.57-2.26
ACE rs4363 A/G (A200060G)					
Allele	N (a.f.)	N (a.f.)			
A	102 (0.645)	96 (0.60)	0.3	1.54	0.66-3.58
G	56 (0.354)	64 (0.40)	0.69	0.88	0.46-1.65
Genotype	N (g.f.)	N (g.f.)			
AA	34 (0.43)	32 (0.40)	0.81	1.13	0.57-2.24
AG	34 (0.43)	32 (0.40)	0.81	1.13	0.27-2.24
GG	11 (0.14)	16 (0.20)	0.41	0.65	0.26-1.61
AGTR1 rs275651 A/T (A777T)					
Allele	N (a.f.)	N (a.f.)			
A	15 (0.095)	9 (0.056)	0.27	1.76	0.70-4.51
T	143 (0.905)	151 (0.943)	0.27	0.51	0.22-1.43
Genotype	N (g.f.)	N (g.f.)			
AA	0	2 (0.025)	0.49	0	0-4.14

AT	15 (0.19)	5 (0.063)	0.02*	3.52	1.11-11.79
TT	64 (0.81)	73 (0.913)	0.10	0.41	0.14-1.16
AGTR1 rs275652 G/T (G-680T)					
Allele	N (a.f.)	N (a.f.)			
G	17 (0.108)	8 (0.050)	0.02**	3.3	1.2-9.1
T	141 (0.892)	152 (0.950)	0.49	2.01	1.72-2.35
Genotype	N (g.f.)	N (g.f.)			
GG	0	2 (0.025)	0.49	-	-
GT	17 (0.220)	4 (0.050)	0.006**	5.2	1.6-16.2
TT	62 (0.780)	74 (0.925)	0.02*	0.3	0.10-0.86
AGTR1 rs275653 A/G (A-119G)					
Allele	N (a.f.)	N (a.f.)			
A	139 (0.880)	152 (0.950)	0.04*	0.39	0.15-0.97
G	19 (0.120)	8 (0.050)	0.008**	3.9	1.4-10.4
Genotype	N (g.f.)	N (g.f.)			
AA	60 (0.759)	74 (0.925)	0.008**	0.26	0.09-0.74
AG	19 (0.241)	4 (0.050)	0.001**	6.0	1.9-18.6
GG	0	2 (0.025)	0.49	-	-
AGTR1 rs5182 T/C (C43732T)					
Allele	N (a.f.)	N (a.f.)			
T	90 (0.570)	106 (0.663)	0.11	0.49	0.2-1.2
C	68 (0.430)	54 (0.337)	0.21	1.49	0.78-2.8
Genotype	N (g.f.)	N (g.f.)			

TT	27 (0.342)	35 (0.438)	0.28	0.67	0.33-1.33
TC	36 (0.456)	36(0.450)	0.93	1.02	0.52-2.01
CC	16 (0.203)	9 (0.113)	0.17	2.0	0.77-5.32
AGTR1 rs5183 A/G (A44221G)					
Allele	N (a.f.)	N (a.f.)			
A	154 (0.975)	158 (0.987)	0.44	0.49	0.06-3.14
G	4 (0.025)	2 (0.012)	0.44	2.0	0.37-11.69
Genotype	N (g.f.)	N (g.f.)			
AA	75 (0.949)	78 (0.975)	0.44	0.48	0.06-3.18
AG	4 (0.051)	2 (0.025)	0.44	2.08	0.31-16.9
GG	0	0	-	-	-
Edn1 RS1800541 G/T (-1394 T/G)					
Allele	N (a.f.)	N (a.f.)			
G	144 (0.913)	139 (0.868)	0.30	1.55	0.72-3.37
T	14 (0.087)	21 (0.132)	0.3	0.64	0.3-1.39
Genotype	N (g.f.)	N (g.f.)			
GG	65 (0.827)	60 (0.75)	0.35	1.55	0.67-3.58
GT	14 (0.173)	19 (0.237)	0.45	0.69	0.3-1.60
TT	0	1 (0.013)	1.0	-	-
Edn1 rs5369 A/G (Glu105Glu)					
Allele	N (a.f.)	N (a.f.)			
A	9 (0.054)	8 (0.053)	0.95	1.03	0.36-2.9
G	149 (0.946)	152 (0.947)	0.97	0.87	0.30-2.54

Genotype	N (g.f.)	N (g.f.)			
AA	0	0	-	-	-
AG	9 (0.108)	8 (0.105)	0.97	1.16	0.38-3.53
GG	70 (0.892)	72 (0.895)	0.97	0.86	0.28-2.62
Edn1 rs3087459 A/C (-974 C/A)					
Allele	N (a.f.)	N (a.f.)			
A	143 (0.907)	141 (0.883)	0.61	1.28	0.59-2.79
C	15 (0.093)	19 (0.117)	0.61	0.81	0.36-1.78
Genotype	N (g.f.)	N (g.f.)			
AA	64 (0.813)	62 (0.779)	0.72	1.24	0.54-2.86
AC	15 (0.187)	17 (0.208)	0.87	0.87	0.37-2.02
CC	0	1 (0.013)	0.49	-	-

*p value after logistic regression analysis was non-significant; **p value after logistic regression analysis was statistically significant and it is shown in Table 2 in the main manuscript, a.f.: allele frequency, g.f.: genotype frequency