

Supplementary Materials: The following supporting information can be downloaded at: www.mdpi.com/xxx/s1, Table S1: Characteristics; Table S2: Distribution of *STAT4* rs10181656, rs7574865, rs7601754, rs10168266 genotypes in patients with optic neuritis and controls; Table S3: *STAT4* (rs10181656, rs7574865, rs7601754, rs10168266) binary logistic regression analysis of genotypes; Table S4: *STAT4* (rs10181656, rs7574865, rs7601754, rs10168266) distribution of genotypes in patients with optic neuritis and controls according to the age of the subjects; Table S5: Distribution of *STAT4* (rs10181656, rs7574865, rs7601754, rs10168266) genotypes in patients with optic neuritis and controls according to gender; Table S6: Binary logistic regression analysis of *STAT4* (rs10181656, rs7574865, rs7601754, rs10168266) genotypes by age of the subjects; Table S7: Binary logistic regression analysis of *STAT4* (rs10181656, rs7574865, rs7601754, rs10168266) genotypes by gender; Table S8: Binary logistic regression analysis of *STAT4* (rs10181656, rs7574865, rs7601754, rs10168266) genotypes in patients with multiple sclerosis (with MS) and without multiple sclerosis (without MS); Table S9: Haplotype association with the predisposition to optical neuritis with MS occurrence;

Table S1. Characteristics.

Characteristics	Group		p-value
	ON group (n=81)	Control group (n=158)	
Male	28 (34.56%)	37 (23.41%)	0.067
Female	53 (65.44%)	121 (76.59%)	
Age median (IQR)	33 (16.5)	29.5 (14)	0.325
Multiple sclerosis:			
-With	42 (51.85%)	-	-
-Without	32 (39.5%)	-	-
-NA	7 (8.64%)	-	-

IQR – interquartile range

Table S2. Distribution of *STAT4* rs10181656, rs7574865, rs7601754, rs10168266 genotypes in patients with optic neuritis and controls.

Gene	Genotype/Aallele	ON group (n=81) N (%)	Control group (n=158) N (%)	p-value	P-value HWE	
<i>STAT4</i> (rs10181656)	CC	50 (61.73)	90 (56.96)	0.621	0.950	
	CG	28 (34.57)	58 (36.71)			
	GG	3 (3.70)	10 (6.33)			
	In total:	81 (100)	158 (100)			
	Allele:			0.366		
	C	128 (79.01)	238 (75.32)			
<i>STAT4</i> (rs7574865)	G	34 (20.99)	78 (24.68)			
	GG	52 (64.2)	91 (57.6)	0.585	0.677	
	GT	25 (30.86)	56 (35.44)			
	TT	4 (4.94)	11 (6.96)			
	In total:	81 (100)	158 (100)			
	Allele:			0.751		
	G	129 (79.63)	238 (75.32)			
<i>STAT4</i> (rs7601754)	T	33 (20.37)	78 (24.68)			
	AA	59 (72.84)	121 (76.58)	0.810	0.736	
	GA	20 (24.69)	34 (21.52)			
	GG	2 (2.47)	3 (1.9)			
	In total:	81 (100)	157 (100)			

	Allele: A G	138 (85.19) 24 (14.81)	276 (87.34) 40 (12.66)	0.511	
STAT4 (rs10168266)	CC	57 (70.37)	105 (66.46)	0.655	0.539
	CT	21 (25.93)	49 (31.01)		
	TT	3 (3.70)	4 (2.53)		
	In total:	81 (100)	158 (100)		
	Allele: C T	135 (83.33) 27 (16.67)	259 (81.96) 57 (18.04)	0.709	

p-value – level of significance; p-value HWE - significance level according to the Hardy-Weinberg equilibrium, differences were considered statistically significant when p<0.05.

Table S3. *STAT4* (rs10181656, rs7574865, rs7601754, rs10168266) binary logistic regression analysis of genotypes.

***STAT4* (rs10181656):**

Model	Genotype/Aallele	OR (95% CI)	p-value	AIC
Codominant	CG vs. CC	0.869 (0.492-1.534)	0.628	309.070
	GG vs. CC	0.540 (0.142-2.054)	0.366	
Dominant	CG+GG vs. CC	0.821 (0.474-1.419)	0.479	307.565
Recessive	GG vs. CC+CG	0.569 (0.152-2.129)	0.402	307.306
Overdominant	CG vs. CC+GG	0.911 (0.52-1.562)	0.744	307.962
Additive	C	0.810 (0.512-1.28)	0.367	307.240

***STAT4* (rs7574865):**

Model	Genotype/Aallele	OR (95% CI)	p-value	AIC
Codominant	GT vs. GG	0.781 (0.437-1.398)	0.406	308.983
	TT vs. GG	0.638 (0.193-2.1)	0.458	
Dominant	GT+TT vs. GG	0.757 (0.436-1.317)	0.325	307.091
Recessive	TT vs. GG+GT	0.694 (0.214-2.253)	0.543	307.682
Overdominant	GT vs. GG+TT	0.813 (0.458-1.442)	0.479	307.564
Additive	G	0.789 (0.503-1.239)	0.303	306.984

***STAT4* (rs7601754):**

Model	Genotype/Aallele	OR (95% CI)	p-value	AIC
Codominant	GA vs. AA	1.206 (0.64-2.274)	0.562	309.652
	GG vs. AA	1.367 (0.222-8.405)	0.736	
Dominant	GA+GG vs. AA	1.219 (0.661-2.25)	0.526	308.669
Recessive	GG vs. AA+GA	1.308 (0.214-7.989)	0.771	307.986
Overdominant	GA vs. AA+GG	1.196 (0.636-2.249)	0.579	307.764
Additive	A	1.195 (0.697-2.049)	0.518	307.656

***STAT4* (rs10168266):**

Model	Genotype/Aallele	OR (95% CI)	p-value	AIC
Codominant	CT vs. CC	0.789 (0.431-1.445)	0.443	309.223
	TT vs. CC	1.382 (0.299-6.389)	0.679	
Dominant	CT+TT vs. CC	0.834 (0.467-1.49)	0.540	307.690
Recessive	TT vs. CC+CT	1.481 (0.323- 6.781)	0.613	307.819
Overdominant	CT vs. CC+CT	0.779 (0.427-1.419)	0.414	307.392
Additive	C	0.908 (0.547-1.506)	0.708	307.927

p-value – level of significance; differences were considered statistically significant when p<0.05, OR – capability ratio; AIC – Akaike.

Table S4. *STAT4* (rs10181656, rs7574865, rs7601754, rs10168266) distribution of genotypes in patients with optic neuritis and controls according to the age of the subjects.

Gene	Genotype	≤ 30 y.		P-value	>30 y.		P-value
		ON group (n = 38) N (%)	Control group (n = 88) N (%)		ON group (n = 43) N (%)	Control group (n = 70) N (%)	
<i>STAT4</i> (rs1018 1656)	CC	24 (63.16)	46 (52.27)	0.425	26 (60.47)	44 (62.86)	0.915
	CG	13 (34.2)	36 (40.91)		15 (34.88)	22 (31.43)	
	GG	1 (2.63)	6 (6.82)		2 (4.65)	4 (5.71)	
	Allele:			0.205			0.906
	C	61 (80.26)	128 (72.73)		67 (77.91)	110 (78.57)	
	G	15 (19.74)	48 (27.27)	0.432	19 (22.09)	30 (21.43)	0.967
	GG	25 (65.79)	47 (53.41)		27 (62.79)	44 (62.86)	
<i>STAT4</i> (rs7574 865)	GT	11 (28.95)	34 (38.64)	0.432	14 (32.56)	22 (31.43)	0.967
	TT	2 (5.26)	7 (7.95)		2 (4.65)	4 (5.71)	
	Allele:						
	G	61 (80.26)	128 (72.73)	0.205	68 (79.07)	110 (78.57)	0.929
	T	15 (19.74)	48 (27.27)		18 (20.93)	30 (21.43)	
<i>STAT4</i> (rs7601 754)	AA	25 (65.79)	66 (75.0)	0.564	34 (79.07)	55 (78.57)	0.929
	GA	12 (31.58)	20 (22.73)		8 (18.60)	14 (20.0)	
	GG	1 (2.63)	2 (2.27)		1 (2.33)	1 (1.43)	
	Allele:			0.330			0.964
	A	62 (81.58)	152 (86.36)		76 (88.37)	124 (88.57)	
<i>STAT4</i> (rs1016 8266)	G	14 (18.42)	24 (13.64)	0.328	10 (11.63)	16 (11.43)	0.873
	CC	28 (73.68)	55 (62.5)		29 (67.44)	50 (71.43)	
	CT	8 (21.05)	30 (34.09)		13 (30.23)	19 (27.14)	
	TT	2 (5.26)	3 (3.41)	0.387	1 (2.33)	1 (1.43)	0.626
	Allele:						
	C	64 (84.2)	140 (79.55)	0.329	71 (82.56)	119 (85.0)	0.747
	T	12 (15.79)	36 (20.45)		15 (17.44)	21 (15.0)	

p-value – level of significance; differences were considered statistically significant when $p < 0.05$.

Table S5. Distribution of *STAT4* (rs10181656, rs7574865, rs7601754, rs10168266) genotypes in patients with optic neuritis and controls according to gender.

Gene	Genotype	Males			Females		
		ON group (n = 28) N (%)	Control group (n = 37) N (%)	P-value	ON group (n = 53) N (%)	Control group (n = 121) N (%)	P-value
<i>STAT4</i> (rs10181656)	CC	20 (71.43)	23 (62.16)	0.647	30 (56.60)	67 (55.37)	0.858
	CG	7 (25)	11 (29.73)		21 (39.62)	47 (38.84)	
	GG	1 (3.57)	3 (8.11)		2 (3.77)	7 (5.79)	
	Allele:				0.329		

	C	47 (83.93)	57 (77.03)		81 (76.42)	181 (74.79)	
	G	9 (16.07)	17 (22.97)		25 (23.58)	61 (25.21)	
STAT4 (rs7574865)	GG	20 (71.43)	25 (67.57)	0.945	32 (60.38)	66 (54.55)	0.661
	GT	6 (21.43)	9 (24.32)		19 (35.85)	47 (38.84)	
	TT	2 (7.14)	3 (8.11)		2 (3.77)	8 (6.61)	
	Allele:						0.388
STAT4 (rs7601754)	G	46 (82.14)	59 (79.73)	0.729	83 (78.30)	179 (73.97)	0.832
	T	10 (17.86)	15 (20.27)		23 (21.7)	63 (26.03)	
	AA	19 (67.86)	29 (78.38)	0.475	40 (75.47)	92 (76.03)	0.832
	GA	8 (28.57)	6 (16.22)		12 (22.64)	28 (23.14)	
	GG	1 (3.57)	2 (5.41)		1 (1.89)	1 (0.83)	
	Allele:						0.833
STAT4 (rs10168266)	A	46 (82.14)	64 (86.49)	0.496	92 (86.8)	212 (87.6)	0.896
	G	10 (17.86)	10 (13.51)		14 (13.21)	30 (12.4)	
	CC	21 (75)	27 (72.97)	0.606	36 (67.92)	78 (64.46)	0.896
	CT	5 (17.86)	9 (24.32)		16 (30.2)	40 (33.06)	
	TT	2 (7.14)	1 (2.70)		1 (1.89)	3 (2.48)	
	Allele:						0.653
	C	47 (83.93)	63 (85.14)	0.849	88 (83.02)	196 (80.99)	0.653
	T	9 (16.07)	11 (14.86)		18 (16.98)	46 (19.01)	

p-value - level of significance; differences were considered statistically significant when p<0.05.

Table S6. Binary logistic regression analysis of *STAT4* (rs10181656, rs7574865, rs7601754, rs10168266) genotypes by age of the subjects.

Age ≤30 years old				
STAT4 (rs10181656):				
Model	Genotype/Aallele	OR (95% CI)	p-value	AIC
Codominant	CG vs. CC	0.841 (0.409-1.73)	0.638	189.610
	GG vs. CC	0.491 (0.097-2.473)	0.388	
Dominant	CG+GG vs. CC	0.785 (0.392-1.574)	0.496	188.056
Recessive	GG vs. CC+CG	0.528 (0.108-2.587)	0.431	187.832
Overdominant	CG vs. CC+GG	0.901 (0.444-1.829)	0.774	188.440
Additive	C	0.775 (0.436-1.376)	0.384	187.748
STAT4 (rs7574865):				
Model	Genotype/Aallele	OR (95% CI)	p-value	AIC
Codominant	GT vs. GG	0.888 (0.427-1.847)	0.750	190.168
	TT vs. GG	0.679 (0.17-2.714)	0.679	
Dominant	GT+TT vs. GG	0.849 (0.423-1.702)	0.644	188.309
Recessive	TT vs. GG+GT	0.712 (0.184-2.76)	0.623	188.270
Overdominant	GT vs. GG+TT	0.930 (0.454-1.903)	0.842	188.483
Additive	G	0.853 (0.492-1.477)	0.570	188.195
STAT4 (rs7601754):				
Model	Genotype/Aallele	OR (95% CI)	p-value	AIC
Codominant	GA vs. AA	1.501 (0.673-3.364)	0.321	189.549
	GG vs. AA	1.212 (0.106-13.830)	0.877	
Dominant	GA+GG vs. AA	1.476 (0.678-3.214)	0.327	187.577

Recessive	GG vs. AA+GA	1.098 (0.097-12.416)	0.940	188.517
Overdominant	GA vs. AA+GG	1.493 (0.672-3.319)	0.325	187.572
Additive	A	1.367 (0.687-2.722)	0.373	187.743

STAT4 (rs10168266):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	CT vs. CC	0.754 (0.351-1.62)	0.470	189.819
	TT vs. CC	1.354 (0.215-8.514)	0.747	
Dominant	CT+TT vs. CC	0.802 (0.385-1.668)	0.554	188.169
Recessive	TT vs. CC+CT	1.481 (0.239-9.175)	0.673	188.349
Overdominant	CT vs. CC+CT	0.743 (0.348-1.585)	0.442	187.921
Additive	C	0.886 (0.469-1.672)	0.708	188.381

Age >30 years old

STAT4 (rs10181656):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	CG vs. CC	1.005 (0.39-2.59)	0.991	122.346
	GG vs. CC	0.804 (0.069-9.379)	0.862	
Dominant	CG+GG vs. CC	0.983 (0.395-2.45)	0.971	120.377
Recessive	GG vs. CC+CG	0.803 (0.07-9.208)	0.860	120.346
Overdominant	CG vs. CC+GG	1.016 (0.397-2.599)	0.974	120.377
Additive	C	0.965 (0.44-2.117)	0.928	120.370

STAT4 (rs7574865):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	GT vs. GG	0.678 (0.254-1.811)	0.438	121.732
	TT vs. GG	0.720 (0.062-8.378)	0.793	
Dominant	GT+TT vs. GG	0.682 (0.266-1.752)	0.427	119.734
Recessive	TT vs. GG+GT	0.803 (0.7-9.208)	0.860	120.364
Overdominant	GT vs. GG+TT	0.688 (0.259-1.828)	0.453	119.803
Additive	G	0.735 (0.325-1.664)	0.461	119.818

STAT4 (rs7601754):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	GA vs. AA	0.849 (0.3-2.407)	0.758	122.165
	GG vs. AA	1.577 (0.94-26.324)	0.751	
Dominant	GA+GG vs. AA	0.901 (0.332-2.445)	0.838	120.336
Recessive	GG vs. AA+GA	1.636 (0.099-27.056)	0.731	120.261
Overdominant	GA vs. AA+GG	0.838 (0.297-2.366)	0.738	120.265
Additive	A	0.967 (0.404-2.314)	0.940	120.373

STAT4 (rs10168266):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	CT vs. CC	0.914 (0.336-2.491)	0.744	122.230
	TT vs. CC	1.600 (0.096-26.749)		
Dominant	CT+TT vs. CC	0.960 (0.365-2.522)	0.934	120.371
Recessive	TT vs. CC+CT	1.636 (0.99-27.056)	0.731	120.261
Overdominant	CT vs. CC+CT	0.901 (0.332-2.445)	0.838	120.336
Additive	C	1.013 (0.432-2.373)	0.977	120.377

p-value – level of significance; differences were considered as statistically significant when p<0.05, OR – capability ratio; AIC – Akaike.

Table S7. Binary logistic regression analysis of STAT4 (rs10181656, rs7574865,

rs7601754, rs10168266) genotypes by gender.

Males				
<i>STAT4</i> (rs10181656):				
Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	CG vs. CC	0.732 (0.238-2.246)	0.585	91.957
	GG vs. CC	0.383 (0.037-3.984)	0.422	
Dominant	CG+GG vs. CC	0.657 (0.229-1.888)	0.436	90.242
Recessive	GG vs. CC+CG	0.420 (0.041-4.266)	0.463	90.258
Overdominant	CG vs. CC+GG	0.788 (0.26-2.387)	0.673	90.680
Additive	C	0.674 (0.288-1.579)	0.364	90.005
<i>STAT4</i> (rs7574865):				
Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	GT vs. GG	0.833 (0.254-2.735)	0.764	92.747
	TT vs. GG	0.833 (0.127-5.479)	0.850	
Dominant	GT+TT vs. GG	0.833 (0.286-2.431)	0.739	90.747
Recessive	TT vs. GG+GT	0.872 (0.136-5.604)	0.885	90.836
Overdominant	GT vs. GG+TT	0.848 (0.262-2.745)	0.784	90.783
Additive	G	0.883 (0.399-1.952)	0.758	90.763
<i>STAT4</i> (rs7601754):				
Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	GA vs. AA	2.035 (0.609-6.799)	0.248	91.384
	GG vs. AA	0.763 (0.065-9.015)	0.830	
Dominant	GA+GG vs. AA	1.717 (0.513-5.233)	0.342	89.952
Recessive	GG vs. AA+GA	0.648 (0.056-7.529)	0.729	90.734
Overdominant	GA vs. AA+GG	2.067 (0.623-6.851)	0.235	89.431
Additive	A	1.326 (0.547-3.215)	0.532	90.468
<i>STAT4</i> (rs10168266):				
Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	CT vs. CC	0.714 (0.208-2.451)	0.593	91.858
	TT vs. CC	2.571 (0.218-30.318)	0.453	
Dominant	CT+TT vs. CC	0.9 (0.293-2.763)	0.854	90.825
Recessive	TT vs. CC+CT	2.769 (0.238-32.184)	0.416	90.149
Overdominant	CT vs. CC+CT	0.676 (0.199-2.301)	0.531	90.459
Additive	C	1.082 (0.446-2.622)	0.862	90.829
Females				
<i>STAT4</i> (rs10181656):				
Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	CG vs. CC	0.998 (0.51-1.952)	0.995	217.597
	GG vs. CC	0.638 (0.125-3.255)	0.589	
Dominant	CG+GG vs. CC	0.951 (0.469-1.824)	0.880	215.896
Recessive	GG vs. CC+CG	0.639 (0.128-3.182)	0.584	215.597
Overdominant	CG vs. CC+GG	1.033 (0.534-2.001)	0.923	215.910
Additive	C	0.912 (0.527-1.577)	0.741	215.809
<i>STAT4</i> (rs7574865):				
Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	GT vs. GG	0.834 (0.422-1.645)	0.600	217.052
	TT vs. GG	0.516 (0.103-2.569)	0.419	
Dominant	GT+TT vs. GG	0.788 (0.408-1.518)	0.476	215.407

Recessive	TT vs. GG+GT	0.554 (0.114-2.701)	0.465	215.328
Overdominant	GT vs. GG+TT	0.880 (0.45-1.719)	0.708	215.778
Additive	G	0.783 (0.452-1.358)	0.385	215.146

STAT4 (rs7601754):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	GA vs. AA	0.986 (0.456-2.132)	0.971	217.582
	GG vs. AA	2.300 (0.14-37.693)	0.559	
Dominant	GA+GG vs. AA	1.031 (0.486-2.187)	0.937	215.913
Recessive	GG vs. AA+GA	2.308 (0.142-37.603)	0.557	215.583
Overdominant	GA vs. AA+GG	0.972 (0.45-2.099)	0.943	215.914
Additive	A	1.079 (0.539-2.161)	0.831	215.874

STAT4 (rs10168266):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	CT vs. CC	0.876 (0.43-1.748)	0.689	217.698
	TT vs. CC	0.722 (0.073-7.185)	0.781	
Dominant	CT+TT vs. CC	0.857 (0.431-1.702)	0.659	215.722
Recessive	TT vs. CC+CT	0.756 (0.077-7.444)	0.811	215.859
Overdominant	CT vs. CC+CT	0.876 (0.436-1.76)	0.709	215.779
Additive	C	0.862 (0.463-1.607)	0.641	215.699

p-value – level of significance; differences were considered as statistically significant when p<0.05, OR – capability ratio; AIC – Akaike.

Table S8. Binary logistic regression analysis of *STAT4* (rs10181656, rs7574865, rs7601754, rs10168266) genotypes in patients with multiple sclerosis (with MS) and without multiple sclerosis (without MS).

With MS				
STAT4 (rs10181656):				
Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	CG vs. CC	0.853 (0.381-1.911)	0.700	175.554
	GG vs. CC	0.450 (0.054-3.719)	0.459	
Dominant	CG+GG vs. CC	0.794 (0.363-1.736)	0.563	173.944
Recessive	GG vs. CC+CG	0.477 (0.059-3.867)	0.488	173.704
Overdominant	CG vs. CC+GG	0.903 (0.407-2.006)	0.802	174.219
Additive	C	0.778 (0.402-1.507)	0.457	173.708
STAT4 (rs7574865):				
Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	GT vs. GG	0.565 (0.237-1.35)	0.199	173.755
	TT vs. GG	0.360 (0.044-2.93)	0.339	
Dominant	GT+TT vs. GG	0.531 (0.231-1.222)	0.137	171.937
Recessive	TT vs. GG+GT	0.431 (0.054-3.463)	0.429	173.505
Overdominant	GT vs. GG+TT	0.607 (0.256-1.441)	0.258	172.929
Additive	G	0.578 (0.284-1.179)	0.132	171.769
STAT4 (rs7601754):				
Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	GA vs. AA	1.456 (0.614-3.454)	0.394	175.406
	GG vs. AA	1.833 (0.182-18.493)	0.607	
Dominant	GA+GG vs. AA	1.486 (0.646-3.42)	0.351	173.441
Recessive	GG vs. AA+GA	1.667 (0.168-16.554)	0.663	174.107

Overdominant	GA vs. AA+GG	1.427 (0.605-3.369)	0.417	173.645
Additive	A	1.419 (0.69-2.92)	0.341	173.417

STAT4 (rs10168266):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	CT vs. CC	0.412 (0.149-1.138)	0.087	172.871
	TT vs. CC	1.010 (0.108-9.417)	0.993	
Dominant	CT+TT vs. CC	0.457 (0.177-1.179)	0.105	171.369
Recessive	TT vs. CC+CT	1.424 (0.134-11.492)	0.849	174.247
Overdominant	CT vs. CC+CT	0.412 (0.15-1.133)	0.086	170.871
Additive	C	0.552 (0.237-1.286)	0.169	172.156

Without MS

STAT4 (rs10181656):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	CG vs. CC	0.836 (0.403-1.732)	0.629	209.195
	GG vs. CC	0.692 (0.143-3.36)	0.648	
Dominant	CG+GG vs. CC	0.814 (0.405-1.637)	0.564	207.247
Recessive	GG vs. CC+CG	0.740 (0.156-3.514)	0.705	207.430
Overdominant	CG vs. CC+GG	0.862 (0.42-1.768)	0.686	207.417
Additive	C	0.834 (0.468-1.468)	0.538	207.195

STAT4 (rs7574865):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	GT vs. GG	0.910 (0.437-1.896)	0.801	209.517
	TT vs. GG	0.993 (0.257-3.834)	0.992	
Dominant	GT+TT vs. GG	0.924 (0.462-1.845)	0.822	207.536
Recessive	TT vs. GG+GT	1.028 (0.273-3.866)	0.967	207.581
Overdominant	GT vs. GG+TT	0.911 (0.443-1.87)	0.799	207.517
Additive	G	0.956 (0.552-1.665)	0.872	207.557

STAT4 (rs7601754):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	GA vs. AA	1.148 (0.512-2.575)	0.738	209.434
	GG vs. AA	1.301 (0.131-12.942)	0.822	
Dominant	GA+GG vs. AA	1.160 (0.532-2.532)	0.709	207.445
Recessive	GG vs. AA+GA	1.260 (0.128-12.434)	0.843	207.545
Overdominant	GA vs. AA+GG	1.140 (0.51-2.549)	0.750	207.483
Additive	A	1.146 (0.577-2.275)	0.698	207.434

STAT4 (rs10168266):

Model	Genotype/Allele	OR (95% CI)	p-value	AIC
Codominant	CT vs. CC	1.032 (0.491-2.17)	0.934	209.069
	TT vs. CC	1.944 (0.338-11.181)	0.456	
Dominant	CT+TT vs. CC	1.101 (0.54-2.244)	0.792	207.513
Recessive	TT vs. CC+CT	1.925 (0.34-10.887)	0.459	207.075
Overdominant	CT vs. CC+CT	0.997 (0.478-2.082)	0.994	207.583
Additive	C	1.158 (0.628-2.14)	0.640	207.367

p-value – level of significance; differences were considered as statistically significant when p<0.05, OR – capability ratio; AIC – Akaike.

Table S9. Haplotype association with the predisposition to optical neuritis with MS occurrence.

Haplotype	<i>STAT4</i>	<i>STAT4</i>	<i>STAT4</i>	<i>STAT4</i>	Frequency		OR (95 % CI)	p-value
	rs10181656	rs7574865	rs7601754	rs10168266	Control	ON		
1	C	G	A	C	59.43	61.82	1.00	—
2	G	T	A	T	14.57	10.06	0.61 (0.26 - 1.43)	0.26
3	C	G	G	C	11.78	14.93	1.34 (0.63 – 2.81)	0.45
4	G	T	A	C	9.16	3.41	0.46 (0.13 - 1.59)	0.22
5	C	G	A	T	2.59	-	-	-

OR: odds ratio; CI: confidence interval; p-value: significance level (after Bonferroni correction statistically significant when p<0.0125).