

Supplementary Table S2: Clinical characteristics of pure small fiber neuropathy patients with a (potentially) pathogenic *SCN10A* variant

c position	Gender	Age	Age debut	INFD (5th percentile)	INFD	TTT	Associated condition(s)	Erythromalgia	Ich	Cramps	Family history	Actual VAS	VASmin	NPS Pain Severity	Sharpness	Hottness	Dullness	Coldness	Sensitivity	Itchiness	Unpleasantness	Intensity deep pain	Intensity surface pain	SPN-SIG No. autonomic	SPN-SIG No. somatic	Neuropathic pain medication at moment hospital visit		
c416G>T	F	59	55	5.5 (4.3)	Normal	Abnormal	IGT, autoimmune disease, ↑ vitamin B6	No	No	No	Negative	71	10	91	8	7	8	3	4	3	5	5	8	3	7	4	None	
c416G>T	F	64	53	2.6 (3.2)	Abnormal	Abnormal	Idiopathic	No	Yes	No	Negative	97	88	94	9	9	9	8	0	10	10	9	8	9	6	4	None	
c416G>T	F	58	56	8.1 (4.3)	Normal	Abnormal	IGT, ↓ vitamin B12	No	No	No	Negative	49	36	68	6	5	7	NA	0	0	7	7	6	7	3	1	Pregabalin 75mg BID, duloxetine 30mg daily	
c416G>T	F	49	47	1.1 (5.7)	Abnormal	Abnormal	DM, alcohol abuse	No	No	No	Positive	28	22	87	5	8	5	3	9	9	4	8	9	6	8	3	Pregabalin 500mg BID	
c4266A>A	F	66	65	4.2 (3.2)	Normal	Abnormal	Idiopathic	No	No	No	Negative	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	None	
c1141A>G	F	58	53	3.1 (4.3)	Abnormal	Abnormal	Idiopathic	No	Yes	No	Positive	88	76	85	8	8	8	8	9	8	8	8	8	8	9	4	None	
c1168A>T	F	47	45	7.6 (5.7)	Normal	Abnormal	Idiopathic	No	No	No	Negative	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	None	
c1879>C	M	55	49	1.2 (3.5)	Abnormal	Normal	Autoimmune disease	Yes	No	No	Negative	83	62	88	8	8	3	2	4	8	8	8	2	2	0	Gabapentin 1200mg TID		
c2212C>G	F	73	70	8 (2.2)	Normal	Abnormal	Idiopathic	No	No	No	Negative	25	13	61	3	4	5	3	6	6	2	6	2	5	8	4	None	
c2972C>T	F	50	49	14.2 (3.8)	Normal	Abnormal	Autoimmune disease	No	No	Yes	NA	72	72	84	7	7	9	7	0	1	0	8	8	1	6	1	Duloxetine 30mg BID	
c2972C>T	M	33	16	5.8 (5.2)	Normal	Abnormal	Autoimmune disease	No	Yes	Yes	Negative	NA	NA	NA	7	9	9	6	9	5	2	9	8	7	6	3	None	
c2972C>T	F	20	15	10.5 (8.4)	Normal	Abnormal	Idiopathic	No	No	No	Positive	21	14	77	NA	8	8	1	1	NA	5	NA	7	9	6	3	None	
c2972C>T	F	37	36	7.4 (7.1)	Normal	Abnormal	Idiopathic	Yes	No	No	Positive	NA	NA	9	9	8	7	8	8	8	8	9	8	9	4	None		
c3482>C	F	60	55	5.1 (3.2)	Normal	Abnormal	↓ vitamin B12	No	No	No	Negative	73	32	97	9	9	9	8	7	8	6	9	8	10	9	4	Oxycodone 25mg extended release BID, oxycodone 10mg on occasion	
c.3607C>T+ c.4568G>A	M	63	62	1.9 (2.8)	Abnormal	Abnormal	Idiopathic	No	No	No	Negative	NA	NA	NA	8	8	4	7	0	6	0	8	7	8	4	3	Pregabalin 300mg BID	
c.3674>C	F	48	42	4.5 (5.7)	Abnormal	Abnormal	Idiopathic	No	Yes	No	Positive	21	11	79	6	3	0	7	7	3	9	5	NA	NA	6	4	Pregabalin 75mg daily	
c.3786>T	M	50	46	0 (5.5)	Abnormal	Abnormal	Idiopathic	Yes	No	No	Negative	36	13	51	3	3	1	4	5	1	3	NA	NA	2	4	None		
c.3803G>A	M	62	59	5 (6.8)	Normal	Abnormal	Autoimmune disease	No	No	No	Negative	57	39	77	6	9	7	6	1	8	8	7	6	5	9	4	None	
c.3803G>A	M	49	48	9.5 (4.4)	Normal	Abnormal	Idiopathic	No	No	No	Negative	NA	NA	NA	3	7	1	NA	0	1	0	4	1	5	2	2	None	
c.3803G>A	M	67	57	6.9 (2.9)	Normal	Abnormal	↓ vitamin B12	No	No	No	Negative	57	13	78	9	9	1	2	9	5	1	9	8	4	2	0	None	
c.3803G>A	M	58	54	0.1 (3.5)	Abnormal	Abnormal	Autoimmune disease, IGT, ↓ vitamin B12	No	No	No	Negative	NA	NA	NA	7	9	9	8	9	2	9	8	8	6	8	4	None	
c.3910G>A	F	57	47	9.3 (3.3)	Normal	Abnormal	Idiopathic	No	No	Yes	Positive	28	30	78	NA	NA	8	1	7	3	0	NA	8	4	6	4	None	
c.4139G>A	F	80	76	3 (5.0)	Normal	Abnormal	Idiopathic	No	No	No	Positive	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	None	
c.4378C>T	F	45	43	7.8 (5.7)	Normal	Abnormal	Autoimmune disease + IGT	Yes	No	No	Positive	93	54	100	9	9	10	9	8	9	9	9	10	9	9	4	Pregabalin 75mg BID	
c.4378C>T	M	52	51	10.6 (3.5)	Normal	Abnormal	Idiopathic	Yes	No	No	Negative	68	20	94	9	9	10	4	7	7	6	9	9	4	3	3	Pregabalin 75mg BID	
c.4568G>A	M	62	58	9.3 (2.8)	Normal	Abnormal	Idiopathic	Yes	No	No	Negative	85	46	94	9	9	9	9	10	10	8	10	9	9	8	3	Pregabalin 75mg BID, tramadol on occasion	
c.4568G>A	M	57	46	0.8 (3.5)	Abnormal	Abnormal	Idiopathic	No	No	No	NA	55	15	70	5	5	6	8	8	8	2	4	4	4	9	4	Gabapentin 600mg TID, amitriptyline 10mg BID, tramadol hydrochloride/paracetamol 37.5/325mg TID	
c.4568G>A	F	39	32	7 (3.1)	Normal	Abnormal	Autoimmune disease	Yes	No	No	Negative	75	36	87	8	8	6	7	6	6	7	7	7	7	7	4	Tramadol 50mg daily	
c.4568G>A	M	58	57	4.9 (3.5)	Normal	Abnormal	Alcohol abuse	No	No	No	NA	38	9	39	5	6	4	3	2	2	5	6	5	3	5	4	None	
c.4568G>A	F	57	55	2.9 (4.9)	Abnormal	Abnormal	Idiopathic	No	No	No	Positive	63	NA	88	7	9	7	6	5	9	5	8	9	7	6	4	Pregabalin 225mg BID	
c.4568G>A	F	49	40	0 (5.7)	Abnormal	Abnormal	Idiopathic	No	No	Yes	Positive	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	NA	Amitriptyline 25mg daily, gabapentin 600mg daily, oxycodone 5mg on occasion
c.4568G>A	M	26	24	1.5 (6.1)	Abnormal	Abnormal	Idiopathic	No	No	No	Negative	28	0	68	7	1	7	7	8	0	0	7	7	4	5	1	None	
c.4568G>A	F	66	65	4.8 (3.2)	Normal	Abnormal	Idiopathic	No	No	No	NA	NA	NA	NA	7	7	7	7	0	1	0	7	7	7	4	4	Pregabalin 300mg daily	
c.4724>C	F	49	37	4.6 (5.7)	Abnormal	Abnormal	Autoimmune disease	No	No	Yes	Positive	36	22	72	5	7	NA	NA	9	NA	1	NA	7	2	9	4	Amitriptyline 50mg daily	
c.4878G>A	M	58	52	0.8 (3.5)	Normal	Abnormal	Autoimmune disease	No	No	No	Negative	17	20	33	3	3	5	5	0	0	0	5	7	5	5	2	Nortriptyline 25mg BID	
c.4984G>A*	F	28	25	17.3 (8.4)	Normal	Abnormal	Idiopathic	No	No	No	Negative	85	36	100	9	9	10	9	0	10	0	9	9	9	2	4	None	
c.4984G>A*	F	24	22	10.7 (8.4)	Normal	Abnormal	Idiopathic	No	No	Yes	Negative	57	NA	81	NA	7	9	6	8	5	4	7	7	NA	7	3	Pregabalin 75mg BID	
c.4984G>A*	F	61	45	3.3 (3.2)	Normal	Abnormal	Chronicitripsy	No	No	No	Negative	49	20	86	8	8	7	6	5	8	6	8	9	9	9	4	Duloxetine 30mg daily, oxycodone extended release 10mg BID	
c.5116A>G	M	62	61	0.4 (3.8)	Abnormal	Abnormal	Idiopathic	No	No	Yes	Negative	7	17	91	2	7	7	2	3	8	8	8	4	7	3	3	Amitriptyline 25mg daily	
c.5474>C	M	63	58	3 (2.8)	Normal	Abnormal	Autoimmune disease + IGT	No	No	No	Negative	9	8	33	3	5	6	2	0	5	0	3	1	3	6	3	None	
c.5539C>T	F	68	65	1.8 (3.2)	Abnormal	Abnormal	Idiopathic	No	No	Yes	Positive	NA	NA	NA	8	8	9	10	9	7	8	7	3	8	7	4	None	
c.5606G>A	F	59	56	3.1 (4.3)	Abnormal	Abnormal	Idiopathic	No	No	No	Positive	30	15	67	4	3	5	3	0	2	0	5	4	3	2	3	None	

F: female; M: male; INFD: intraepidermal nerve fibre density; TTT: temperature threshold testing; VAS: visual analogue scale (VAS is ranging from 0-100 [0 = no pain and 100 = the worst possible pain]); min: minimum; max: maximum; NPS: neuropathic pain scale (0 = not applicable to the experienced pain and 10 = in the most severe form applicable to the experienced pain); SPN-SIG: small fiber neuropathy inventory questionnaire; No. autonomic: number of positive autonomic symptoms; No. somatic: number of positive somatic symptoms; NA: not available; DM: diabetes mellitus; IGT: impaired glucose tolerance; ↓ vitamin B12: vitamin B12 deficiency; † vitamin B6: vitamin B6 intoxication; BID: twice daily; TID: three times daily; QID: four times daily. *Variant classified as pathogenic according to the Waxman recommendations.