

Supplementary Table 1. Clinical data, results of questionnaire assessment, and small fiber tests in men with FMS with normal and impaired glucose tolerance.

	Normal glucose tolerance (n = 32)	Impaired glucose tolerance (n = 10)
Age (years)	55 (33 – 75)	61 (31 – 74)
BMI (kg/m ²)	28.9 (21.3 – 40.6)	30.1 (23.0 – 45.6)
Pain duration (years)	10 (2 – 60)	20 (4 – 50)
Current pain intensity on NRS	5 (1 – 8)	5 (2 – 8)
NPSI sum score	3.7 (0 – 7.4)	2.6 (0 – 8.0)
NPSI discriminative score	66.7 (42.4 – 84.3)	54.5 (42.0 – 79.6)
GCPS		
– Pain intensity	65 (37 – 93)	62 (42 – 83)
– Disability due to pain	49 (0 – 83)	62 (8 – 80)
PCS sum score	20 (2 – 41)	24 (2 – 42)
ADS sum score	18 (7 – 39)	24 (8 – 48)
FIQ sum score	45 (27 – 67)	48 (24 – 66)
STAI-S	43 (32 – 73)	38 (30 – 48)*
STAI-T	43 (31 – 73)	42 (34 – 55)
MPSS Gerbershagen Grad	3 (2 – 3)	3 (3 – 3)
WPI	13.5 (8 – 19)	13.5 (8 – 18)
SSS	8 (3 – 11)	6.5 (5 – 12)
Small fiber tests		
IENFD (fibers/mm)		
– distal	4.3 (1.0 – 8.3)	4.2 (1.6 – 14.4)

- proximal	7.6 (0.6 – 15.5)	6.4 (2.5 – 16.0)
IENFD subgroups		
- Normal IENFD	6/32 (19%)	1/10 (10%)
- Proximally reduced	1/32 (3%)	1/10 (10%)
- Distally reduced	9/32 (28%)	2/10 (20%)
- Generalized reduction	16/32 (50%)	6/10 (60%)
CCM		
- NFD (no/mm ²)	22.4 (4.2 – 35.4)	19.8 (5.2 – 33.3)
- NBD (no/mm ²)	56.8 (19.8 – 114.6)	63.5 (13.5 – 140.8)
- NFL (mm/mm ²)	12.8 (5.0 – 18.4)	12.4 (4.8 – 18.3)
QST		
- CDT	-0.29 (-2.26 – 1.32)	-0.11 (-1.54 – 0.78)
- WDT	-0.81 (-1.75 – 1.29)	-0.87 (-1.52 – 0.50)
- TSL	-0.43 (-1.65 – 1.32)	-0.33 (-1.3 – 0.60)
- PHS	-0.67 (-2.98 – 0.48)	-1.83 (-2.98 – 0.48)
- CPT	-0.49 (-0.56 – 3.24)	0.88 (-0.56 – 2.88)
- HPT	-0.41 (-0.97 – 1.64)	-0.24 (-0.97 – 2.92)
- MDT	-0.47 (-2.47 – 1.26)	0.38 (-1.18 – 1.26)*
- MPT	-0.57 (-2.07 – 1.09)	-0.33 (-1.91 – 1.09)
- MPS	0.47 (-1.46 – 2.61)	0.58 (-0.82 – 2.46)
- VDT	0.18 (-2.27 – 1.14)	0.26 (-2.55 – 1.14)
- PPT	0.48 (-2.16 – 2.61)	0.13 (-0.71 – 1.49)
Pleasant touch	8 (0 – 10)	7 (-5 – 10)

Data are given as median with range in brackets. *p<0.05.

Abbreviations: ADS = Allgemeine Depressionsskala; CCM = corneal confocal microscopy; FIQ = Fibromyalgia Impact Questionnaire; CDT = cold detection threshold; CPT = cold pain threshold; GCPS = Graded Chronic Pain Scale; HPT = heat pain threshold; IENFD = intraepidermal nerve fiber density; MDT = mechanical detection threshold; MPS = mechanical pain sensitivity; MPT= mechanical pain threshold; NBD = nerve branching density; NFD = nerve fiber density; NFL = nerve fiber length; NPSI = Neuropathic Pain Symptom Inventory; PCS = Pain Catastrophizing Scale; PHS = paradoxical heat sensation; PPT = pressure pain threshold; QST = Quantitative Sensory Testing; SSS = symptom severity score; STAI = State-Trait Anxiety Inventory; TSL = thermal sensory limen; VDT = vibration detection threshold; WDT = warm detection threshold; WPI = widespread pain index.

Supplementary Table 2. Laboratory normative values for intraepidermal nerve fiber density (normative values are not age-adjusted).

Distal IENFD (fibers/mm)	9 +/- 3
Proximal IENFD (fibers/mm)	12 +/- 4

Data are given as median with range in brackets.

Abbreviations: IENFD = intraepidermal nerve fiber density.

Supplementary Table 3. Correlation of cutaneous and corneal innervation with age, BMI, and pain duration in men with FMS and male healthy controls.

FMS			
	Age	BMI	Pain duration
IENFD distal (n = 42)	-0.045, p = 0.779	-0.217, p = 0.168	-0.172, p = 0.275
IENFD proximal (n = 42)	-0.042, p = 0.790	-0.125, p = 0.429	-0.179, p = 0.256
NFD (n = 41)	-0.108, p = 0.501	-0.133, p = 0.409	0.004, p = 0.979
NBD (n = 41)	-0.173, p = 0.278	0.196, p = 0.219	-0.228, p = 0.151
NFL (n = 41)	-0.227, p = 0.153	-0.163, p = 0.308	-0.162, p = 0.312
Healthy controls			
	Age		
IENFD distal (n = 55)	-0.244, p = 0.073		
IENFD proximal (n = 40)	-0.233, p = 0.148		
NFD (n = 28)	-0.276, p = 0.155		
NBD (n = 28)	-0.086, p = 0.664		
NFL (n = 28)	-0.130, p = 0.511		

The bivariate Spearman correlation was performed for correlation analysis.

Abbreviations: BMI = Body mass index; FMS = fibromyalgia syndrome; IENFD = intraepidermal nerve fiber density; NBD = nerve branch density; NFD = nerve fiber density; NFL = nerve fiber length.

Supplementary Table 4. Results of questionnaire assessment and small fiber tests in the FMS subgroups classified according to the skin innervation pattern.

	Normal IENFD (n = 7)	Proximally reduced IENFD (n = 2)	Distally reduced IENFD (n = 11)	Generalized reduction of IENFD (n = 22)	Pathological IENFD (n=35)
NPSI sum score	3.3 (0 – 6.4)	5.4 (2.7 – 8.0)	3.7 (1.1 – 5.9)	3.7 (1.6 – 7.4)	3.7 (1.1 – 8.0)
NPSI discriminative score	59.5 (42.0 – 71.3)	74.6 (74.4 – 74.7)	62.7 (44.8 – 77.3)	66.5 (44.9 – 84.3)	66.5 (44.8 – 84.3)
GCPS					
– Pain intensity	57 (42 – 93)	72 (60 – 83)	67 (50 – 80)	62 (37 – 90)	63 (37 – 90)
– Disability due to pain	40 (7 – 83)	63 (50 – 77)	47 (0 – 73)	59 (2 – 83)	53 (0 – 83)
PCS sum score	26 (11 – 41)	4 (2 – 6)	20 (3- 38)	22 (2 – 42)	20 (2 – 42)
ADS sum score	15 (11 – 39)	20 (7 – 33)	18 (8 – 38)	24 (8 – 48)	21 (7 – 48)
FIQ sum score	34 (28 – 58)	39 (29 – 48)	38 (27 – 67)	48 (24 – 66)	46 (24 – 67)
STAI-S	38 (37 – 63)	41 (39 – 43)	41 (32 – 73)	43 (30 – 51)	41 (30 – 73)
STAI-T	41 (36 – 59)	38 (37 – 39)	44 (33 – 73)	44 (31 – 55)	43 (31 – 73)

MPSS Gerbershagen	3 (2 – 3)	3 (3 – 3)	3 (2 – 3)	3 (3 – 3)	3 (2 – 3)
Grad					
WPI	10 (8 – 16)	13.5 (13 – 14)	15 (12 – 17)	13.5 (8 – 19)	14 (8 – 19)
SSS	8 (5 – 10)	6.5 (5 – 8)	7 (4 – 12)	8 (3 – 11)	8 (3 – 12)
Small fiber tests					
IENFD (fibers/mm)					
– distal	7.3 (6.7 – 14.4)	7.4 (6.5 – 8.3)	4.1 (1.9 – 5.9)	3.4 (1.0 – 5.9)	4.0 (1.0 – 8.3)
– proximal	12.0 (8.2 – 16.0)	3.4 (2.5 – 4.3)	9.8 (8.1 – 13.2)	6.1 (0.6 – 7.7)	6.7 (0.6 – 13.2)
CCM					
– NFD (no/mm ²)	22.9 (4.2 – 27.1)	29.2 (29.2 – 29.2)	22.9 (16.7 – 33.3)	19.3 (5.2 – 35.4)	21.4 (5.2 – 35.4)
– NBD (no/mm ²)	66.4 (28.1 – 104.2)	50.0 (50.0 – 50.0)	80.2 (21.9 – 140.8)	52.6 (13.5 – 101.6)	57.8 (13.5 – 140.8)
– NFL (mm/mm ²)	14.2 (5.0 – 15.6)	14.0 (14.0 – 14.0)	14.1 (8.4 – 18.4)	10.9 (4.8 – 17.1)	12.4 (4.8 – 18.4)
QST					
– CDT	-0.46 (-2.26 – 0.67)	0.11 (-0.59 – 0.8)	-0.04 (-1.89 – 1.32)	-0.28 (-2.26 – 1.27)	-0.26 (-2.26 – 1.32)
– WDT	-0.49 (-1.75 – 0.37)	0.35 (0-19 – 0.5)	-1.13 (-1.4 – 0.74)	-0.86 (-1.73 – 1.29)	-0.88 (-1.73 – 1.29)
– TSL	-0.17 (-1.5 – 0.6)	0.31 (0.23 – 0.38)	-0.51 (-1.65 – 0.89)	-0.43 (-1.62 – 1.32)	-0.44 (-1.65 – 1.32)

– PHS	-0.67 (-2.98 – 0.48)	-0.10 (-0.67 – 0.48)	-2.98 (-2.98 – 0.48)	-0.10 (-2.98 – 0.48)	-0.67 (-2.98 – 0.48)
– CPT	0.23 (-0.56 – 2.96)	0.69 (-0.56 – 1.94)	-0.41 (-0.56 – 2.98)	-0.14 (-0.56 – 3.24)	-0.41 (-0.56 – 3.24)
– HPT	0.26 (-0.88 – 1.64)	0.98 (-0.97 – 2.92)	-0.43 (-0.97 – 0.73)	-0.45 (-0.97 – 1.39)	-0.43 (-0.97 – 2.92)
– MDT	0.1 (-1.65 – 0.77)	0.64 (0.05 – 1.22)	-0.47 (-1.54 – 1.26)	-0.44 (-2.47 – 1.26)	-0.26 (-2.47 – 1.26)
– MPT	-0.8 (-1.96 – -0.33)	1.04 (0.99 – 1.09)	-0.8 (-1.96 – 1.09)	-0.41 (-2.07 – 0.83)	-0.33 (-2.07 – 1.09)
– MPS	0.57 (-1.46 – 1.65)	0.73 (-1.01 – 2.46)	0.58 (-0.77 – 1.66)	0.35 (-0.82 – 2.61)	0.48 (-1.01 – 2.61)
– VDT	0.18 (-1.05 – 1.14)	-0.82 (-2.55 – 0.91)	0.01 (-1.7 – 1.14)	0.36 (-2.27 – 1.14)	0.18 (-2.55 – 1.14)
– PPT	0.71 (0.03 – 1.94)	-0.24 (-1.97 – 1.49)	1.03 (-2.16 – 2.0)	0.29 (-0.83 – 2.61)	0.42 (-2.16 – 2.61)
Pleasant touch	8 (0 – 10)	1.5 (-5 – 8)	8 (0 – 10)	10 (0 – 10)	8 (-5 – 10)

Data are given as median with range in brackets.

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severity score; STAI = State-Trait Anxiety Inventory; TSL = thermal sensory limen; VDT = vibration detection threshold; WDT = warm detection threshold; WPI = widespread pain index.