

Fig. S1. Correlation between the body weight and the muscle weight, and between the body weight and the physiological cross-sectional area (pCSA) of the extensor carpi ulnaris (ECU) muscle in normal (n=15) and affected (n=15) dogs.

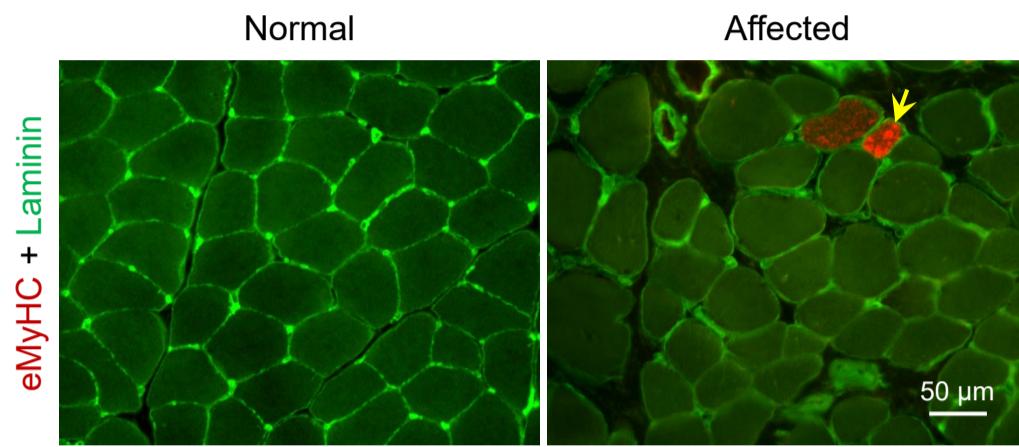


Fig. S2. Affected dog extensor carp ulnaris muscle shows low-level fresh regeneration.
Representative embryonic myosin heavy chain (eMyHC) immunofluorescence staining images from normal and affected ECU muscles. Individual myofibers were outlined by laminin immunostaining. Yellow arrow, a myofiber containing eMyHC.

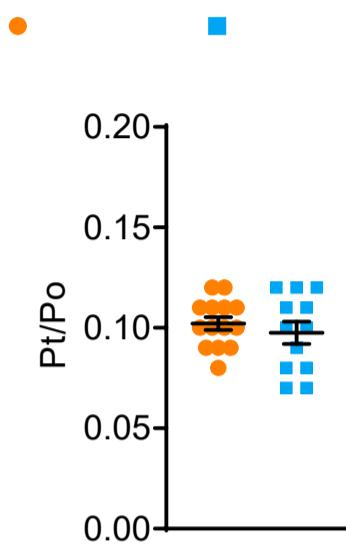


Fig. S3. The twitch (Pt) to tetanic (Po) force ratio in normal (n=14) and affected (n=12) dogs.

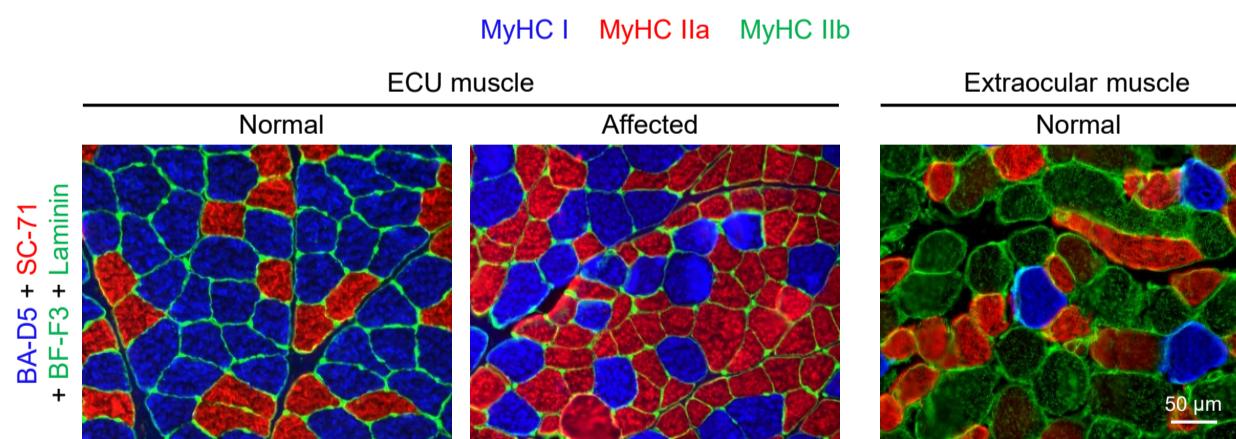


Fig. S4. Type IIb myofiber is absent in the dog ECU muscle. Representative myosin heavy chain isoform (MyHC) immunostaining photomicrographs from normal and affected ECU muscles, and a normal extraocular muscle. Type I myofibers (blue) were stained with antibody BA-D5. Type IIa myofibers (red) were stained with antibody SC-71. Type IIb myofibers (green) were stained with antibody BF-F3. The muscle cell membrane (green) was stained with a polyclonal antibody against laminin.

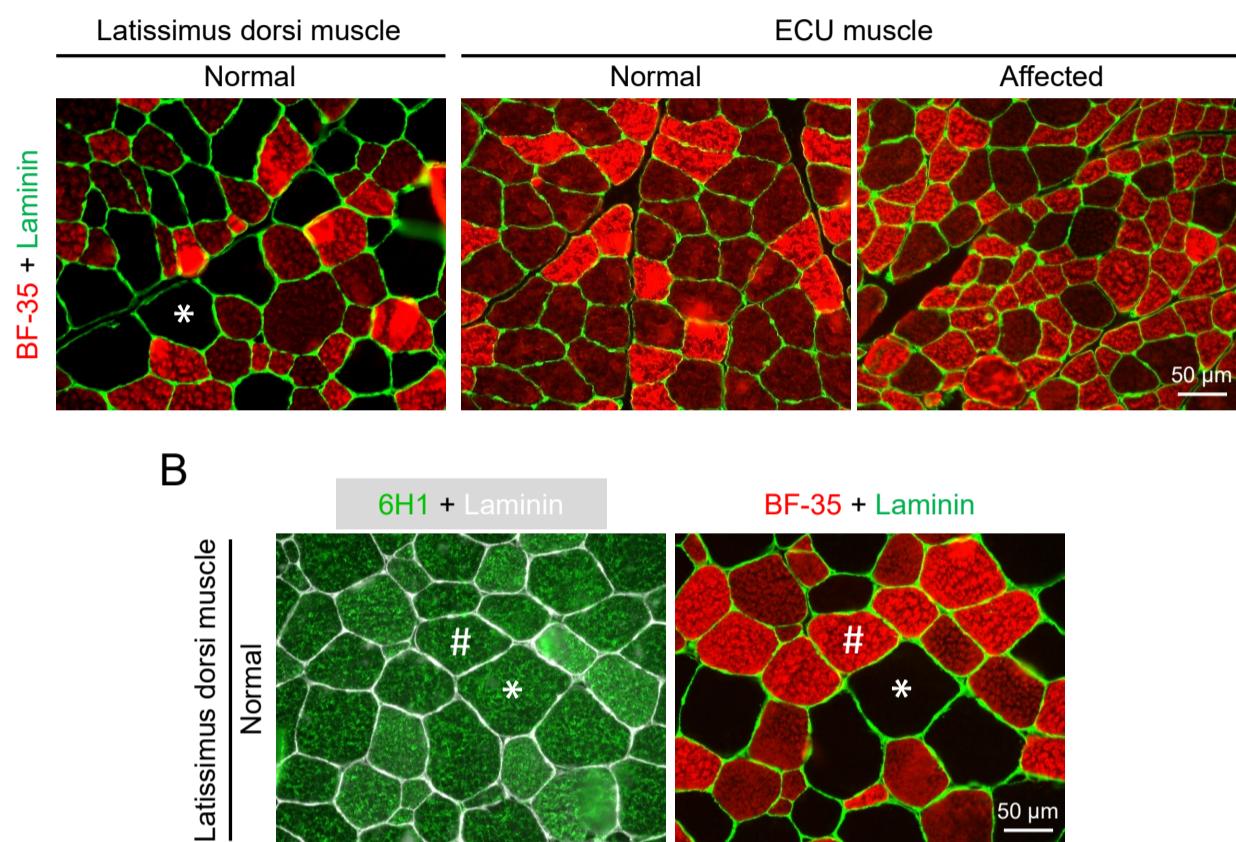


Fig. S5. Evaluation of type IIx fibers in dog muscle. (A) Representative myosin heavy chain isoform immunostaining using antibody BF-35. The BF-35 antibody stains all myosin heavy chain isoforms except type IIx. Type IIx fibers can be readily identified in the latissimus dorsi muscle. No type IIx fiber was detected in the dog ECU muscle. Asterisk (*), a type IIx myofiber. **(B)** Representative myosin heavy chain isoform immunostaining of the latissimus dorsi muscle using antibodies 6H1 and BF-35. The 6H1 antibody cannot identify type IIx myofibers in dog muscle. Asterisk (*), a type IIx myofiber. Pound sign (#), a non-type IIx myofiber.

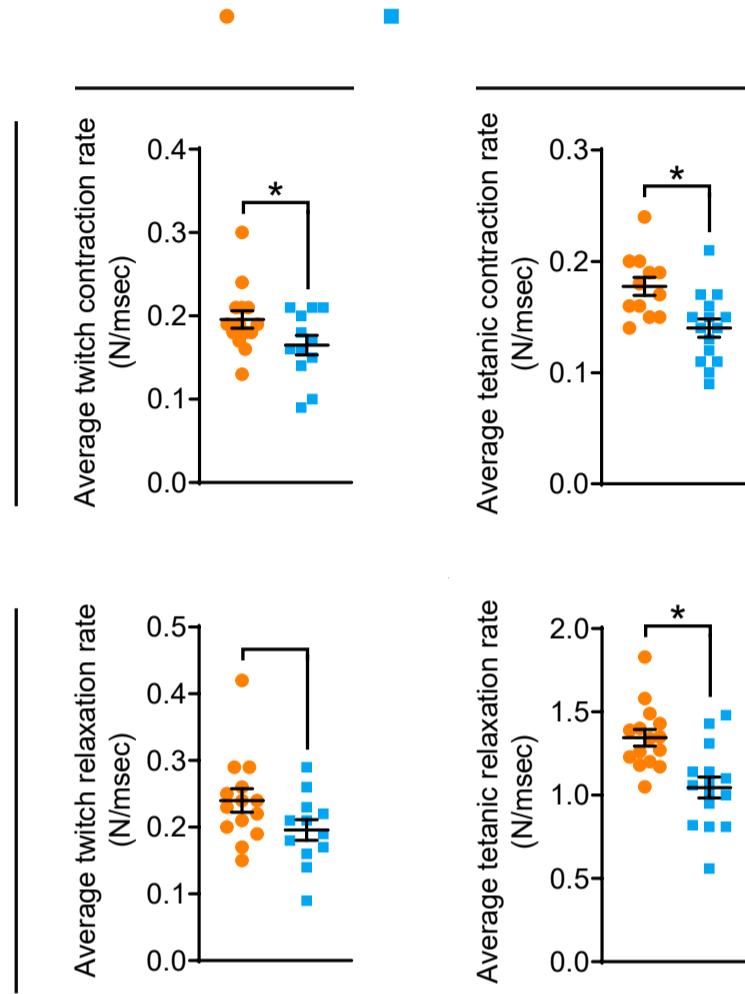


Fig. S6. Quantification of the average contraction and relaxation rate. **(A)** The average contraction rate of twitch contraction in the normal ($n=14$) and affected ($n=12$) ECU muscle. The rate is calculated using the equation P_t/T_{PT} . **(B)** The average contraction rate of tetanic contraction in the normal ($n=15$) and affected ($n=15$) ECU muscle. The rate is calculated using the equation P_o/T_{PT} . **(C)** The average relaxation rate (based on $\frac{1}{2} RT$) of twitch contraction in the normal ($n=14$) and affected ($n=12$) ECU muscle. The rate is calculated using equation $(P_t) / (\frac{1}{2} RT)$. **(D)** The average relaxation rate (based on $\frac{1}{2} RT$) of tetanic contraction in the normal ($n=15$) and affected ($n=15$) ECU muscle. The rate is calculated using equation $(P_o) / (\frac{1}{2} RT)$. Asterisk (*), significantly different from each other. $P < 0.05$, unless noted on the graph.

Table S1. Optimized stimulation parameters of the ECU muscle

	Normal	Affected	
Sample size	15	15	
Gender	M	M	
Optimal resting tension (g)	Min Max Average Majority ^a	700 2000 1257.53 ± 352.49 1000 to 2000 (80%)	300 1600 $728.57 \pm 329.17^*$ 600 to 1600 (73%)
Optimal stimulation current (mA)	Min Max Average Majority ^a	20 600 100.67 ± 144.54 40 to 80 (73%)	20 300 77.33 ± 77.04 20 to 80 (80%)
Optimal stimulation duration (msec)	Min Max Average Majority ^a	500 800 760.00 ± 105.56 800 (87%)	500 800 706.67 ± 122.28 800 (60%), 600 (27%)
Optimal stimulation frequency (Hz)	Min Max Average Majority ^a	80 120 101.33 ± 14.07 100 (53%)	80 120 94.67 ± 14.07 100 (47%), 80 (40%)

ECU, extensor carpi ulnaris muscle; Min, minimum value; Max, maximum value.

*, Significantly different from normal dogs.

a, Optimal condition(s) in most of the dogs in the same column.

Data are presented as mean \pm SD

Table S2. Antibodies used in myofiber type immunofluorescence staining

Antibody name	Isotype	MyHC isoform specificity	Dilution	Figure
BA-D5	IgG2b	MyHC I	1:20	Figs 7, S4
SC-71	IgG1	MyHC IIa	1:100	Figs 7, S4
BF-F3	IgM	MyHC IIb	1:40	Figs 7, S4
6H1	IgM	MyHC IIx	1:50	Fig. S5B
BF-35	IgG1	All isoforms except IIx	1:50	Fig. S5
F1.652	IgG1	eMyHC	1:250	Figs 3D, S2

Table S3. Primers and probes used in droplet digital PCR

Gene	NCBI reference sequence #	MyHC isoform	Forward primer (5' - 3')	Reverse primer (5' - 3')	Probe (5' - 3')
MYH7	NM_001113711.1	I	AGAATGACCTCCAGCTTCAAGTG	AGCGCTCCTCTGCATCAG	AAGCGGAACAAGAAC
MYH2	NM_001076795.1	IIa	TCAGTGTGGCTAGCATTGATGATC	GTGAGCTTGTAGATGGAGACTTTCT	ACAGATAGTGCTATTGATATTT
MYH1	NM_001113717.1	IIx	ACATTGCCGGCTTGTAGATCTT	GTTTCTCATTTGGTAAGTTGATGCA	CAGGCTGTTAAAATCA
MYH4	NM_001076794.1	IIb	CTGATGAGGGTGGATTCAAGAAGA	CGGACATTGACTGGATGCAGAAAA	ATGGAGAGGAGAGGTCC

Table S4. Comparison of the current study and our previous study

	Current study (Hakim <i>et al.</i>)		Our previous study (Yang <i>et al.</i>)	
	Normal	Affected	Normal	Affected
Sample size	15	15	7	7
Sex	M	M	M	M & F
Age (year)	Min	2.16	2.78	0.74
	Max	3.54	3.65	2.55
	Average	2.96 ± 0.43	3.13 ± 0.41	1.64 ± 0.79
Body Weight (Kg)	Min	15.38	14.74	13.00
	Max	34.47	29.84	32.20
	Average	21.23 ± 4.65	19.28 ± 4.05	20.34 ± 7.29
ECU muscle weight (g)	Min	6.09	4.44	3.60
	Max	12.93	11.72	15.46
	Average	9.49 ± 2.48	7.49 ± 1.95*	8.37 ± 4.84
ECU weight / Body weight ratio (g/kg)	Min	0.32	0.28	0.25
	Max	0.60	0.49	0.54
	Average	0.45 ± 0.08	0.39 ± 0.05*	0.39 ± 0.11
ECU muscle length (cm)	Min	11.34	11.76	9.70
	Max	16.38	16.38	19.50
	Average	14.14 ± 1.54	14.14 ± 1.31	14.48 ± 3.71
pCSA (cm ²)	Min	10.10	7.86	7.19
	Max	17.07	14.89	19.65
	Average	13.78 ± 2.35	10.91 ± 2.15*	11.37 ± 4.51
Po (N)	Min	113.02	56.23	52.80
	Max	193.38	130.71	159.70
	Average	147.05 ± 24.11	97.63 ± 21.13*	97.15 ± 43.38
Po / muscle weight (N/g)	Min	12.75	8.86	10.33
	Max	19.86	16.82	15.54
	Average	15.95 ± 2.12	13.30 ± 2.41*	12.45 ± 2.16
Po / pCSA (N/cm ²)	Min	9.26	6.08	6.84
	Max	12.27	11.97	9.81
	Average	10.72 ± 0.84	9.04 ± 1.82*	8.38 ± 1.08
				7.04 ± 0.96*

ECU, extensor carpi ulnaris muscle; M, Male; F, Female; N, newton; Po, tetanic force; pCSA, physiological cross-sectional area; Min, minimum value; Max, maximum value.

*, Significantly different from normal dogs.

Data are presented as mean ± SD

Table S5. Comparison of the parameters used in eccentric contraction in the current study and our previous study

	Current study (Hakim <i>et al.</i>)		Our previous study (Yang <i>et al.</i>)	
	Normal	Affected	Normal	Affected
% Length stretched	Min	5.00	5.00	4.99
	Max	5.00	5.00	5.00
	Average	5.00 ± 0.00	5.00 ± 0.00	5.00 ± 0.00
Stretch rate (mm/s)	Min	5.67	5.88	6.58
	Max	8.19	8.19	10.19
	Average	7.07 ± 0.77	7.07 ± 0.65	7.90 ± 2.00
	Variance	0.59	0.43	3.99

Min, minimum value; Max, maximum value.

Data are presented as mean ± SD

Dataset 1. Raw data tables

Fig. 2c	
% of Centrally nucleated fibers	
Normal	Affected
0.32	47.26
0.00	31.63
2.81	19.82
0.86	18.98

Fig. 2D	
eMyHC (% Area)	
Normal	Affected
0.40	1.78
0.22	1.38
0.18	2.38
0.25	1.22
0.26	0.92
0.13	2.56
0.05	1.31
0.21	1.83
0.19	

Average: 1.00 29.42
SD: 1.26 13.22
Sample size (n): 4 4

Average: 0.21 1.67
SD: 0.10 0.57
Sample size (n): 9 8

Statistics Analysis
Test: unpaired *t*-test
P value: 0.0052

Statistics Analysis
Test: unpaired *t*-test
P value: <0.0001

Mini-Feret diameter (μm)	% of myofibers									
	Normal					Affected				
20	0	0	0	0	0	5	1	2	1	2
30	0	7	1	12	2	14	15	16	10	14
40	16	38	13	55	28	31	17	34	28	34
50	50	34	41	29	48	50	17	31	31	27
60	28	16	33	4	20	15	20	10	15	17
70	5	5	11	0	2	2	11	6	5	7
80	0	0	1	0	0	0	10	2	2	3
90	0	0	0	0	0	0	3	0	0	1
100	0	0	0	0	0	0	2	0	0	0

Mini-Feret diameter	Normal		Normal	
	Average	SD	Average	SD
20	0.00	0.00	2.57	1.59
30	4.00	4.20	17.29	8.24
40	30.17	14.02	30.43	5.88
50	42.00	8.14	25.86	6.06
60	19.33	9.37	13.71	4.23
70	4.17	3.53	5.57	2.66
80	0.17	0.37	2.86	3.04
90	0.00	0.00	0.86	0.99
100	0.00	0.00	0.29	0.70

Sample size (n): 6 7

Fig. 2D	
Fibrosis (%) Area)	
Normal	Affected
7.62	18.65
11.30	15.15
7.25	32.30
12.86	31.66
13.76	19.22
23.07	22.33
11.30	18.54
12.82	27.18
9.78	30.34
11.28	

Average: 12.10 23.93
SD: 4.41 6.52
Sample size (n): 10 9

Statistics Analysis
Test: unpaired *t*-test
P value: 0.0002

Fig. 3B		Fig. 3C		Fig. 3D		Fig. 3E		Fig. 3F		Fig. 3G		Fig. 3H		Fig. 3I		Fig. 3J	
Pt (N)	Affected	sPt (N/cm ²)		TPT (msec)		1/RRT (msec)		Max df/dt (N/msec)		Max -df/dt (N/msec)		Time to max df/dt (msec)		Time to max -df/dt (msec)			
Normal	Affected	Normal	Affected	Normal	Affected	Normal	Affected	Normal	Affected	Normal	Affected	Normal	Affected	Normal	Affected	Normal	Affected
19.47	5.65	1.49	0.67	93.70	53.87	66.80	41.41	0.48	0.17	-0.21	-0.09	22.10	23.97	39.40	21.27		
12.35	12.77	1.11	1.09	76.10	62.27	63.12	49.22	0.34	0.36	-0.14	-0.17	19.85	21.10	35.80	25.40		
15.73	10.91	1.23	0.89	75.40	52.90	73.99	48.14	0.45	0.36	-0.14	-0.16	20.50	21.73	31.20	22.20		
14.24	5.53	1.07	0.70	75.27	62.43	60.59	64.68	0.38	0.15	-0.16	-0.06	20.40	24.40	27.37	25.30		
13.32	9.18	1.05	1.16	76.70	65.40	71.49	58.16	0.35	0.23	-0.13	-0.10	21.20	25.60	27.10	24.47		
11.93	10.14	0.96	0.75	90.90	62.40	80.91	55.27	0.28	0.32	-0.10	-0.12	23.03	20.80	32.93	27.87		
12.76	7.31	1.06	0.77	69.93	47.07	54.38	34.78	0.34	0.26	-0.16	-0.14	21.00	21.10	32.70	22.10		
16.83	13.04	1.18	1.23	70.17	60.77	58.28	44.61	0.45	0.38	-0.19	-0.19	21.70	21.37	28.60	24.07		
22.06	8.65	1.29	0.79	72.47	59.33	52.38	50.70	0.60	0.27	-0.29	-0.12	20.47	21.50	34.70	23.30		
12.57	9.13	1.10	0.73	66.17	50.97	52.06	44.04	0.36	0.31	-0.16	-0.14	20.50	19.97	32.83	25.33		
13.71	9.86	0.81	0.66	71.80	57.63	53.24	45.66	0.35	0.30	-0.18	-0.15	22.53	21.30	27.60	21.47		
15.92	11.54	0.96	1.30	86.37	57.87	73.59	62.26	0.38	0.36	-0.15	-0.11	21.83	20.73	40.47	26.20		
14.85		0.92		80.87		90.00		0.36		-0.11		22.80		35.50			
12.74		1.26		59.27		50.61		0.42		-0.16		19.97		28.97			

Average: 14.89 9.48 Average: 1.11 0.90 Average: 76.08 57.74 Average: 64.39 49.91 Average: 0.40 0.29 Average: -0.16 -0.13 Average: 21.28 21.96 Average: 32.51 24.08
SD: 2.94 2.45 SD: 0.17 0.23 SD: 9.39 5.50 SD: 12.17 8.81 SD: 0.08 0.08 SD: 0.05 0.04 SD: 1.06 1.72 SD: 4.34 2.06
Sample size (n): 14 12 Sample size (n): 14 12

Statistics Analysis
Test: unpaired *t*-test
P value: <0.0001

Statistics Analysis
Test: unpaired *t*-test
P value: 0.0146

Statistics Analysis
Test: unpaired *t*-test
P value: <0.0001

Statistics Analysis
Test: unpaired *t*-test
P value: 0.0022

Statistics Analysis
Test: unpaired *t*-test
P value: 0.0019

Statistics Analysis
Test: unpaired *t*-test
P value: 0.0548

Statistics Analysis
Test: unpaired *t*-test
P value: 0.2249

Statistics Analysis
Test: unpaired *t*-test
P value: <0.0001

Fig. 5B

% of contraction force	Normal															Affected														
	Average df/dt (N/msec)																													
	Dog # 1	Dog # 2	Dog # 3	Dog # 4	Dog # 5	Dog # 6	Dog # 7	Dog # 8	Dog # 9	Dog # 10	Dog # 11	Dog # 12	Dog # 13	Dog # 14	Dog # 15	Dog # 1	Dog # 2	Dog # 3	Dog # 4	Dog # 5	Dog # 6	Dog # 7	Dog # 8	Dog # 9	Dog # 10	Dog # 11	Dog # 12	Dog # 13	Dog # 14	Dog # 15
0-10	0.50	0.56	0.40	0.48	0.37	0.44	0.41	0.39	0.48	0.65	0.39	0.46	0.44	0.49	0.48	0.24	0.36	0.35	0.45	0.17	0.25	0.43	0.33	0.39	0.19	0.27	0.35	0.18	0.41	0.18
10-20	0.70	0.67	0.52	0.57	0.45	0.55	0.50	0.53	0.68	0.87	0.56	0.64	0.63	0.74	0.60	0.40	0.56	0.57	0.69	0.28	0.43	0.57	0.55	0.60	0.29	0.45	0.54	0.27	0.68	0.28
20-30	0.68	0.58	0.47	0.54	0.47	0.52	0.48	0.50	0.63	0.86	0.52	0.65	0.66	0.83	0.50	0.39	0.52	0.54	0.65	0.28	0.42	0.56	0.53	0.58	0.28	0.48	0.43	0.27	0.66	0.27
30-40	0.57	0.50	0.42	0.46	0.44	0.42	0.37	0.47	0.52	0.75	0.41	0.58	0.60	0.75	0.47	0.35	0.43	0.50	0.59	0.24	0.38	0.49	0.49	0.50	0.25	0.43	0.35	0.24	0.59	0.24
40-50	0.69	0.42	0.57	0.56	0.35	0.42	0.45	0.36	0.47	0.66	0.37	0.47	0.53	0.61	0.44	0.28	0.37	0.44	0.55	0.20	0.35	0.43	0.40	0.43	0.22	0.38	0.33	0.20	0.48	0.20
50-60	0.53	0.34	0.29	0.34	0.30	0.34	0.31	0.27	0.38	0.52	0.34	0.39	0.38	0.51	0.37	0.24	0.31	0.38	0.42	0.16	0.29	0.39	0.31	0.34	0.19	0.34	0.24	0.18	0.39	0.17
60-70	0.39	0.27	0.23	0.31	0.25	0.29	0.28	0.23	0.29	0.37	0.27	0.30	0.28	0.40	0.23	0.19	0.24	0.28	0.31	0.12	0.22	0.30	0.26	0.25	0.15	0.26	0.19	0.14	0.29	0.13
70-80	0.30	0.19	0.17	0.23	0.19	0.22	0.21	0.18	0.23	0.28	0.17	0.23	0.20	0.28	0.19	0.14	0.19	0.21	0.20	0.09	0.16	0.23	0.19	0.11	0.20	0.15	0.11	0.19	0.10	
80-90	0.21	0.13	0.11	0.17	0.13	0.15	0.16	0.12	0.15	0.18	0.12	0.15	0.13	0.19	0.13	0.09	0.13	0.14	0.15	0.06	0.11	0.14	0.12	0.13	0.07	0.13	0.10	0.07	0.13	0.06
90-100	0.11	0.05	0.03	0.08	0.04	0.04	0.08	0.03	0.05	0.03	0.04	0.04	0.03	0.02	0.04	0.03	0.02	0.04	0.03	0.05	0.02	0.04	0.03	0.02	0.01	0.02	0.01	0.02	0.01	0.01

Sample size (n):

15

15

Statistics Analysis

Test: unpaired t-test

% of contraction force	Average	SD	Average	SD	P Value
0-10	0.46	0.07	0.30	0.10	
10-20	0.61	0.11	0.48	0.15	
20-30	0.59	0.13	0.46	0.14	
30-40	0.52	0.12	0.41	0.12	
40-50	0.49	0.11	0.35	0.11	
50-60	0.37	0.08	0.29	0.09	
60-70	0.29	0.05	0.22	0.07	
70-80	0.22	0.04	0.16	0.05	
80-90	0.15	0.03	0.11	0.03	
90-100	0.05	0.02	0.02	0.01	

15

15

Fig. 5D

% of relaxation force	Normal															Affected														
	Average -df/dt (N/msec)																													
	Dog # 1	Dog # 2	Dog # 3	Dog # 4	Dog # 5	Dog # 6	Dog # 7	Dog # 8	Dog # 9	Dog # 10	Dog # 11	Dog # 12	Dog # 13	Dog # 14	Dog # 15	Dog # 1	Dog # 2	Dog # 3	Dog # 4	Dog # 5	Dog # 6	Dog # 7	Dog # 8	Dog # 9	Dog # 10	Dog # 11	Dog # 12	Dog # 13	Dog # 14	Dog # 15
0-10	0.00	-0.02	-0.01	-0.01	-0.01	-0.01	-0.04	0.00	-0.01	0.00	-0.01	-0.01	-0.01	-0.01	0.00	0.00	-0.01	0.00	0.00	0.00	0.00	-0.01	-0.01	-0.01	0.00	0.00	0.00	0.00	0.00	0.00
10-20	-0.59	-0.64	-0.45	-0.18	-0.16	-0.18	-0.41	-0.44	-0.67	-0.94	-0.64	-0.32	-0.65	-0.57	-0.54	-0.28	-0.42	-0.45	-0.36	-0.18	-0.25	-0.42	-0.40	-0.40	-0.17	-0.41	-0.31	-0.24	-0.41	-0.20
20-30	-1.11	-1.19	-0.87	-0.79	-0.76	-0.77	-0.85	-0.94	-1.18	-1.55	-1.05	-1.06	-1.14	-1.29	-0.96	-0.64	-0.79	-0.82	-0.75	-0.41	-0.61	-0.90	-0.68	-0.86	-0.53	-0.92	-0.63	-0.51	-0.74	-0.43
30-40	-1.45	-1.50	-1.06	-1.17	-1.19	-1.24	-1.10	-1.37	-1.60	-2.12	-1.47	-1.55	-1.49	-1.92	-1.29	-0.89	-1.13	-1.16	-1.23	-0.60	-0.90	-1.23	-0.88	-1.21	-0.84	-1.37	-1.02	-0.78	-1.09	-0.61
40-50	-1.84	-1.81	-1.34	-1.55	-1.62	-1.70	-1.35	-1.81	-2.07	-2.72	-1.86	-1.98	-1.66	-2.46	-1.62	-1.10	-1.51	-1.46	-1.72	-0.77	-1.12	-1.55	-1.09	-1.54	-1.11	-1.76	-1.38	-1.04	-1.45	-0.77
50-60	-2.24	-2.16	-2.16	-1.72	-1.90	-2.09	-1.66	-2.12	-2.46	-3.15	-2.13	-2.30	-1.64	-2.78	-1.89	-1.27	-1.81	-1.66	-2.11	-0.89	-1.29	-1.89	-1.27	-1.81	-1.29	-2.02	-1.62	-1.22	-1.71	-0.91
60-70	-2.45	-2.36	-1.96	-2.06	-2.40	-2.41	-1.83	-2.21	-2.58	-3.23	-2.18	-2.40	-1.42	-2.77	-2.02	-1.36	-1.94	-1.73	-2.32	-0.94	-1.36	-2.10	-1.37	-1.93	-1.36	-2.10	-1.71	-1.29	-1.81	-0.97
70-80	-2.32	-2.22	-1.93	-1.91	-2.42	-2.32	-2.01	-2.32	-2.85	-3.99	-2.16	-2.38	-1.94	-3.12	-1.84	-1.63	-2.27	-0.88	-1.29	-2.08	-1.32	-1.83	-1.28	-1.98	-1.62	-1.22	-1.70	-0.94		
80-90	-1.71	-1.59	-1.51	-1.33	-2.00	-1.72	-1.25	-1.42	-1.89	-1.46	-0.92	-1.56	-1.53	-1.08	-1.42	-1.31	-1.86	-0.67	-1.02	-1.69	-1.06	-1.41	-1.01	-1.58	-1.29	-0.96	-1.32	-0.74		
90-100	-0.19	-0.18	-0.16	-0.17	-0.19	-0.12	-0.17	-0.19	-0.25	-0.16	-0.19	-0.20	-0.22	-0.16	-0.11	-0.14	-0.13	-0.20	-0.07	-0.11	-0.15	-0.11	-0.16	-0.11	-0.15	-0.12	-0.10	-0.13	-0.08	

Sample size (n):

15

15

Statistics Analysis

Test: unpaired t-test

% of contraction force	Average	SD	Average	SD	P Value
0-10	-0.01	0.01	0.00	0.00	
10-20	-0.49	0.22	-0.33	0.10	
20-30	-1.04	0.22	-0.68	0.16	
30-40	-1.43	0.29	-1.00	0.23	
40-50	-1.83	0.37	-1.29	0.31	
50-60	-2.16	0.41	-1.52	0.38	
60-70	-2.29	0.42	-1.62	0.42	
70-80	-2.11	0.39	-1.55	0.41	
80-90	-1.52	0.26	-1.23	0.34	
90-100	-0.18	0.03	-0.12	0.03	

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15

Fig. 5F	
FWHM at contraction (msec)	
Normal	Affected
150.50	170.80
127.70	142.70
141.90	104.40
260.30	100.70
231.30	152.90
155.00	100.00
135.80	121.00
191.40	108.10
174.20	128.30
165.90	109.00
168.10	135.80
153.70	132.60
174.10	151.50
155.00	141.10
175.10	141.00

Average: 170.70 129.30

SD: 35.14 21.56

Sample size (n): 15 15

Statistics Analysis

Test: unpaired *t*-test

P value: 0.0006

Fig. 5H	
FWHM at relaxation (msec)	
Normal	Affected
49.00	45.70
55.90	35.30
46.70	38.20
40.70	36.30
44.10	48.60
57.00	34.30
45.60	46.30
55.60	40.10
47.00	41.30
52.40	45.30
88.50	47.30
45.50	45.60
40.80	51.50
41.10	41.00
46.40	48.00

Average: 50.42 42.99

SD: 11.83 5.32

Sample size (n): 15 15

Statistics Analysis

Test: unpaired *t*-test

P value: 0.0347

Stimulation frequency (Hz)	Normal														Affected														
	Dog # 1	Dog # 2	Dog # 3	Dog # 4	Dog # 5	Dog # 6	Dog # 7	Dog # 8	Dog # 9	Dog # 10	Dog # 11	Dog # 12	Dog # 13	Dog # 14	Dog # 1	Dog # 2	Dog # 3	Dog # 4	Dog # 5	Dog # 6	Dog # 7	Dog # 8	Dog # 9	Dog # 10	Dog # 11	Dog # 12	Dog # 13	Dog # 14	
1	25.00	21.71	17.73	16.52	16.55	17.50	16.69	16.69	15.31	16.00	15.84	15.84	15.84	15.84	17.27	17.13	17.00	17.00	17.00	17.00	17.00	17.00	17.00	17.00	17.00	17.00	17.00	17.00	
20	102.73	175.68	103.80	72.83	88.57	93.31	74.31	73.87	90.07	103.33	63.98	92.45	88.79	91.01	59.98	77.59	59.39	55.04	26.87	50.10	66.33	78.40	64.55	57.17	39.16	53.04	57.44	51.05	
40	150.09	146.44	115.42	111.69	125.25	133.69	105.33	107.39	139.54	163.42	99.66	141.89	134.86	146.52	93.69	99.34	100.61	45.79	74.00	101.18	56.83	94.79	102.54	87.18	65.74	102.85	81.48	87.03	
60	169.09	154.48	118.22	125.97	141.04	146.29	114.43	119.98	153.84	184.54	109.69	157.06	148.60	164.65	108.86	72.09	101.28	119.55	55.17	83.27	114.58	67.12	108.20	122.60	94.85	75.81	120.25	89.32	103.72
80	175.11	158.54	119.83	130.69	149.78	151.76	118.85	124.81	157.80	191.39	111.28	162.37	152.97	171.45	113.53	76.15	103.00	125.77	56.23	85.63	119.73	71.10	112.45	129.57	97.75	79.70	124.30	93.53	110.46
100	175.07	160.13	120.93	132.15	140.32	152.40	123.33	123.88	156.63	193.38	111.02	163.49	152.18	172.56	117.35	77.57	98.95	124.93	54.03	82.27	121.43	71.80	110.40	130.76	97.63	80.62	122.30	93.76	111.13
120	166.62	158.60	120.43	129.53	146.06	152.91	127.66	118.95	149.78	193.25	110.33	162.13	145.97	173.49	117.35	76.84	92.03	122.00	50.51	76.75	121.56	71.92	105.50	130.30	97.99	80.19	116.13	92.69	109.50

Sample size (n):

15

14

Statistics Analysis				
Test: unpaired t-test				
% of contraction force	Normal	Affected		
Average	SD	Average	SD	
5	17.57	3.52	11.07	2.53
20	88.71	16.97	49.47	13.89
40	127.66	21.01	82.03	18.90
60	141.11	23.67	94.84	21.66
80	155.42	24.32	98.41	19.98
100	146.42	24.32	98.41	19.98
120	144.47	23.87	95.99	22.96

15

14

Statistics Analysis				
Test: unpaired t-test				
% of contraction force	Normal	Affected		
Average	SD	Average	SD	
5	1.53	1.59	1.23	1.45
20	6.06	9.63	9.29	6.10
40	8.90	11.22	10.33	8.75
60	10.00	11.22	10.33	8.75
80	10.38	12.14	10.72	10.24
100	10.00	12.27	10.82	10.35
120	9.88	12.15	10.77	10.15

15

14

Statistics Analysis				
Test: unpaired t-test				
% of contraction force	Normal	Affected		
Average	SD	Average	SD	
5	1.28	0.17	1.03	0.25
20	6.52	1.29	5.49	1.14
40	9.28	1.04	8.26	1.39
60	10.28	0.82	8.82	1.58
80	10.60	0.84	9.20	1.69
100	10.67	0.86	9.14	1.69
120	10.54	0.94	8.91	1.68

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14

Statistics Analysis				
Test: unpaired t-test				
% of contraction force	Normal	Affected		
Average	SD	Average	SD	
5	1.28	0.17	1.03	0.25
20	6.52	1.29	5.49	1.14
40	8.85	1.34	7.82	1.38
60	9.15	0.84	8.52	0.88
80	9.67	0.97	9.77	0.92
100	9.90	0.99	9.98	0.99
120	9.51	0.94	9.81	0.99

15

14

Statistics Analysis				
Test: unpaired t-test				
% of contraction force	Normal	Affected		
Average	SD	Average	SD	
5	1.24	1.12	1.24	1.24
20	60.59	9.34	69.69	9.19
40	86.82	3.70	82.45	3.85
60	95.89	2.37	95.41	1.83
80	98.88	1.74	99.47	0.61
100	99.52	0.88	98.83	1.55
120	98.23	2.02	96.34	4.22

15

14

Statistics Analysis				
Test: unpaired t-test				
ECC cycle	Normal	Affected		
Average	SD	Average	SD	
1	100.00	100.00	100.00	100.00
2	100.51	87.39	99.36	100.12
3	98.90	86.56	97.73	96.71
4	99.24	86.63	98.07	96.93
5	101.34	86.78	98.28	96.54
6	99.87	86.26	98.23	96.88
7	98.30	91.68	95.66	92.45
8	100.31	90.72	94.23	92.00
9	99.35	89.03	94.58	90.19
10	97.81	87.36	94.32	87.67

15

14

Statistics Analysis				
Test: unpaired t-test				
ECC cycle	Normal	Affected		
Average	SD	Average	SD	
1	100.00	0.00	100.00	0.00
2	98.30	4.40	97.00	3.98
3	94.60	7.65	79.80	9.56
4	93.94	7.20	71.63	11.31
5	94.23	5.72	65.26	12.98
6	93.44	5.63	60.53	14.12
7	92.80	4.40	59.00	11.43
8	93.63	5.37	52.45	18.88
9	92.97	5.83	49.69	18.35
10	91.93	6.19	47.75	17.74

15

14

Statistics Analysis				
Test: unpaired t-test				
ECC cycle	Normal	Affected		
Average	SD	Average	SD	
1	100.00	0.00	100.00	0.00
2	100.51	87.39	99.36	100.12
3	98.90	86.56	97.73	96.71
4	99.24	86.63	98.07	96.93
5	101.34	86.78	98.28	96.54
6	99.87	86.26	98.23	96.88
7	98.30	91.68	95.66	92.45
8	100.31	90.72	94.23	92.00
9	99.35	89.03	94.58	90.19
10	97.81	87.36	94.32	87.67

15

14

Fig. 7B

% of myofibers

MyHC isoforms	Normal						Affected							
	Dog # 1	Dog # 2	Dog # 3	Dog # 4	Dog # 5	Dog # 6	Dog # 1	Dog # 2	Dog # 3	Dog # 4	Dog # 5	Dog # 6	Dog # 7	Dog # 8
I	71.68	65.74	76.16	61.86	68.30	57.81	28.37	34.90	37.87	25.92	27.95	24.94	23.71	15.83
I/Ila	0.48	0.00	1.02	0.00	0.94	0.48	19.61	12.89	6.84	14.95	20.07	2.68	2.85	14.29
IIa	27.84	34.26	22.82	38.14	30.76	41.71	52.02	52.21	55.28	59.13	51.98	72.38	73.44	69.88

Sample size (n):

6

8

Statistics Analysis

Test: unpaired *t*-test

MyHC isoforms	Normal		Affected		MyHC isoforms	P Value
	Average	SD	Average	SD		
I	66.93	6.63	27.44	6.79	I	0.0000
I/Ila	0.49	0.44	11.77	6.91	I/Ila	0.0019
IIa	32.59	6.90	60.79	9.55	IIa	0.0001

Fig. 7C

Relative percentage (%)

MyHC isoforms	Normal			Affected		
	Dog # 1	Dog # 2	Dog # 3	Dog # 1	Dog # 2	Dog # 3
I	55.55	55.54	47.69	28.60	33.97	44.62
IIa	44.45	44.46	52.31	71.40	66.03	55.38

Sample size (n):

3

Statistics Analysis

Test: unpaired *t*-test

MyHC isoforms	Normal		Affected		MyHC isoforms	P Value
	Average	SD	Average	SD		
I	52.93	4.54	35.73	8.15	I	0.0331
IIa	47.07	4.54	64.27	8.15	IIa	0.0331

Fig. 7B

Transcript copy (copy/ng)

Transcript	Normal						Affected						
	Dog # 1	Dog # 2	Dog # 3	Dog # 4	Dog # 5	Dog # 6	Dog # 1	Dog # 2	Dog # 3	Dog # 4	Dog # 5	Dog # 6	Dog # 7
MyHC7	1.4E+05	7.5E+04	1.3E+05	5.6E+04	1.1E+05	8.3E+04	4.8E+04	4.0E+04	2.9E+04	3.4E+04	7.7E+04	5.0E+04	9.0E+04
MyHC2	9.0E+04	5.9E+04	1.0E+05	5.4E+04	4.6E+04	4.6E+04	1.1E+05	6.4E+04	2.7E+04	7.1E+04	1.2E+05	1.5E+05	1.3E+05
MyHC1	8.4E+02	5.2E+02	1.3E+03	4.3E+02	4.8E+02	3.5E+02	4.6E+02	4.6E+03	9.9E+02	3.1E+03	2.0E+03	—	1.4E+03
MyHC4	1.0E+00	1.0E+00	0.0E+00	0.0E+00	0.0E+00	0.0E+00	2.0E+01	2.3E+01	1.8E+01	9.0E+00	4.0E+00	1.6E+01	3.1E+01

Sample size (n):

6

8

Statistics Analysis

Test: unpaired *t*-test

Transcript	Normal		Affected		Transcript	P Value
	Average	SD	Average	SD		
MyHC7	1.0E+05	3.5E+04	5.2E+04	2.3E+04	MyHC7	0.0208
MyHC2	6.7E+04	2.4E+04	9.5E+04	4.2E+04	MyHC2	0.2547
MyHC1	6.6E+02	3.7E+02	2.1E+03	1.5E+03	MyHC1	0.0177
MyHC4	3.3E-01	5.2E-01	1.7E+01	8.9E+00	MyHC4	0.0021

Table 1

Dog name	Age (Year)	Body weigh (kg)	ECU muscle weight (g)	ECU weight/body weight (g/kg)	ECU muscle lenght (cm)	pCSA (cm ²)
Normal	2.16	34.47	12.93	0.38	15.96	16.86
	2.28	19.50	9.22	0.47	14.70	13.06
	2.50	17.24	6.09	0.35	11.34	11.18
	2.62	18.51	8.50	0.46	13.86	12.77
	2.82	20.41	9.40	0.46	14.70	13.31
	2.82	20.41	8.94	0.44	14.70	12.66
	2.82	15.38	7.26	0.47	12.18	12.41
	3.07	16.92	6.80	0.40	11.76	12.04
	3.09	22.72	10.36	0.46	15.12	14.26
	3.19	23.59	12.40	0.53	15.12	17.07
	3.20	21.45	7.13	0.33	13.02	11.40
	3.40	20.41	12.23	0.60	15.12	16.84
	3.42	20.46	12.00	0.59	15.12	16.52
	3.47	27.22	12.72	0.47	16.38	16.16
	3.54	19.73	6.32	0.32	13.02	10.10
	Min	2.16	15.38	6.09	0.32	11.34
	Max	3.54	34.47	12.93	0.60	16.38
	Average	2.96	21.23	9.49	0.45	14.14
	SD	0.43	4.65	2.48	0.08	2.35
Affected	2.48	15.24	5.63	0.37	13.86	8.46
	2.55	21.59	7.32	0.34	13.02	11.70
	2.75	20.09	8.13	0.40	14.28	11.85
	2.78	20.09	9.44	0.47	15.96	12.31
	2.86	15.83	4.44	0.28	11.76	7.86
	2.90	15.20	5.09	0.33	13.44	7.88
	3.02	19.87	9.78	0.49	15.12	13.46
	3.02	14.88	6.09	0.41	13.44	9.43
	3.22	20.41	7.50	0.37	14.70	10.62
	3.47	22.90	8.37	0.37	15.96	10.92
	3.48	22.13	8.63	0.39	13.86	12.96
	3.49	17.41	7.59	0.44	12.60	12.54
	3.50	29.84	11.72	0.39	16.38	14.89
	3.65	14.74	5.72	0.39	13.44	8.86
	3.73	19.00	6.84	0.36	14.28	9.97
	Min	2.78	14.74	4.44	0.28	11.76
	Max	3.65	29.84	11.72	0.49	16.38
	Average	3.13	19.28	7.49	0.39	14.14
	SD	0.41	4.05	1.95	0.05	2.15
P value	0.2913	0.2318	0.0206	0.0225	1.0000	0.0016

Fig. S1

Dog name	Body weight (kg)	ECU muscle weight (g)	pCSA (cm ²)
Normal	34.47	12.93	16.86
	19.50	9.22	13.06
	17.24	6.09	11.18
	18.51	8.50	12.77
	20.41	9.40	13.31
	20.41	8.94	12.66
	15.38	7.26	12.41
	16.92	6.80	12.04
	22.72	10.36	14.26
	23.59	12.40	17.07
	21.45	7.13	11.40
	20.41	12.23	16.84
	20.46	12.00	16.52
	27.22	12.72	16.16
	19.73	6.32	10.10
Affected	15.24	5.63	8.46
	21.59	7.32	11.70
	20.09	8.13	11.85
	20.09	9.44	12.31
	15.83	4.44	7.86
	15.20	5.09	7.88
	19.87	9.78	13.46
	14.88	6.09	9.43
	20.41	7.50	10.62
	22.90	8.37	10.92
	22.13	8.63	12.96
	17.41	7.59	12.54
	29.84	11.72	14.89
	14.74	5.72	8.86
	19.00	6.84	9.97

Fig. S3

Pt/Po	
Normal	Affected
0.12	0.07
0.10	0.12
0.12	0.09
0.10	0.10
0.09	0.11
0.09	0.08
0.10	0.10
0.11	0.12
0.11	0.07
0.11	0.11
0.08	0.08
0.10	0.12
0.09	
0.11	

Average: 0.10 0.10

SD: 0.01 0.02

Sample size (n): 14 12

Statistics Analysis

Test: unpaired t -test

P value: 0.4575

Fig. S6A	
Pt/TPT	
Normal	Affected
0.21	0.10
0.16	0.21
0.21	0.21
0.19	0.09
0.17	0.14
0.13	0.16
0.18	0.16
0.24	0.21
0.30	0.15
0.19	0.18
0.19	0.17
0.18	0.20
0.18	
0.21	

Average: 0.20 0.17
SD: 0.04 0.04
Sample size (n): 14 12

Fig. S6B	
Pt/1/2RT	
Normal	Affected
0.29	0.14
0.20	0.26
0.21	0.23
0.24	0.09
0.19	0.16
0.15	0.18
0.23	0.21
0.29	0.29
0.42	0.17
0.24	0.21
0.26	0.22
0.22	0.19
0.17	
0.25	

Average: 0.24 0.20
SD: 0.07 0.05
Sample size (n): 14 12

Fig. S6C	
Po/TPT	
Normal	Affected
0.20	0.10
0.15	0.17
0.17	0.15
0.19	0.21
0.16	0.09
0.16	0.17
0.24	0.15
0.15	0.11
0.20	0.14
0.19	0.16
0.18	0.12
0.14	0.11
	0.15
	0.13
	0.14

Average: 0.18 0.14
SD: 0.03 0.03
Sample size (n): 12 15

Fig. S6D	
Po/1/2RT	
Normal	Affected
1.49	0.81
1.35	1.06
1.18	1.00
1.27	1.48
1.34	0.56
1.39	0.82
1.05	1.14
1.26	0.81
1.43	1.14
1.83	1.43
1.20	1.07
1.40	0.95
1.23	1.31
1.58	1.01
1.17	1.10

Average: 1.35 1.05
SD: 0.19 0.24
Sample size (n): 15 15

Statistics Analysis

Test: unpaired *t*-test
P value: 0.0470

Statistics Analysis

Test: unpaired *t*-test
P value: 0.0785

Statistics Analysis

Test: unpaired *t*-test
P value: 0.0036

Statistics Analysis

Test: unpaired *t*-test
P value: 0.0009

Table S1

Dog name	Optimal resting tension (g)	Optimal stimulation current (mA)	Optimal Stimulation duration (msec)	Optimal stimulation frequency (Hz)
Normal	1100	20	800	80
	1100	60	800	80
	1300	600	500	80
	1500	40	800	100
	1600	40	800	100
	1400	50	500	100
	900	60	800	100
	1463	60	800	100
	700	80	800	100
	800	100	800	100
	1000	200	800	100
	1000	40	800	120
	1500	40	800	120
	2000	40	800	120
	1500	80	800	120
Min	700.00	20.00	500.00	80.00
Max	2000.00	600.00	800.00	120.00
Average	1257.53	100.67	760.00	101.33
SD	352.49	144.54	105.56	14.07
Affected	600	60	500	80
	1000	20	600	80
	600	60	600	80
	600	80	600	80
	600	100	600	80
	1600	20	800	80
	400g	40	500	100
	1100	20	800	100
	700	20	800	100
	300	40	800	100
	500	80	800	100
	500	200	800	100
	500	300	800	100
	900	40	800	120
	700	80	800	120
Min	300.00	20.00	500.00	80.00
Max	1600.00	300.00	800.00	120.00
Average	728.57	77.33	706.67	94.67
SD	329.17	77.04	122.28	14.07
P value	0.0003	0.5855	0.2115	0.2052

Table S4

Dog name	Age (Year)	Body weigh (kg)	ECU muscle weight (g)	ECU weight/body weight (g/kg)	ECU muscle lenght (cm)	pCSA (cm ²)	Po (N)	Po/ECU weight (N/g)	Po/pCSA (N/cm ²)
Normal	2.16	34.47	12.93	0.38	15.96	16.86	175.11	13.54	10.38
	2.28	19.50	9.22	0.47	14.70	13.06	160.13	17.37	12.27
	2.50	17.24	6.09	0.35	11.34	11.18	120.93	19.86	10.82
	2.62	18.51	8.50	0.46	13.86	12.77	132.15	15.55	10.35
	2.82	20.41	9.40	0.46	14.70	13.31	141.04	15.00	10.60
	2.82	20.41	8.94	0.44	14.70	12.66	152.91	17.10	12.08
	2.82	15.38	7.26	0.47	12.18	12.41	128.56	17.71	10.36
	3.07	16.92	6.80	0.40	11.76	12.04	123.38	18.14	10.25
	3.09	22.72	10.36	0.46	15.12	14.26	157.80	15.23	11.06
	3.19	23.59	12.40	0.53	15.12	17.07	193.38	15.60	11.33
	3.20	21.45	7.13	0.33	13.02	11.40	113.02	15.85	9.92
	3.40	20.41	12.23	0.60	15.12	16.84	163.49	13.37	9.71
	3.42	20.46	12.00	0.59	15.12	16.52	152.97	12.75	9.26
	3.47	27.22	12.72	0.47	16.38	16.16	173.49	13.64	10.73
	3.54	19.73	6.32	0.32	13.02	10.10	117.35	18.57	11.62
Min	2.16	15.38	6.09	0.32	11.34	10.10	113.02	12.75	9.26
Max	3.54	34.47	12.93	0.60	16.38	17.07	193.38	19.86	12.27
Average	2.96	21.23	9.49	0.45	14.14	13.78	147.05	15.95	10.72
SD	0.43	4.65	2.48	0.08	1.54	2.35	24.11	2.12	0.84
Affected	2.48	15.24	5.63	0.37	13.86	8.46	77.37	13.74	9.15
	2.55	21.59	7.32	0.34	13.02	11.70	103.00	14.07	8.80
	2.75	20.09	8.13	0.40	14.28	11.85	72.02	8.86	6.08
	2.78	20.09	9.44	0.47	15.96	12.31	125.77	13.32	10.22
	2.86	15.83	4.44	0.28	11.76	7.86	56.23	12.66	7.15
	2.90	15.20	5.09	0.33	13.44	7.88	85.63	16.82	10.86
	3.02	19.87	9.78	0.49	15.12	13.46	121.56	12.43	9.03
	3.02	14.88	6.09	0.41	13.44	9.43	71.92	11.81	7.63
	3.22	20.41	7.50	0.37	14.70	10.62	112.45	14.99	10.59
	3.47	22.90	8.37	0.37	15.96	10.92	130.71	15.62	11.97
	3.48	22.13	8.63	0.39	13.86	12.96	97.99	11.35	7.56
	3.49	17.41	7.59	0.44	12.60	12.54	80.62	10.62	6.43
	3.50	29.84	11.72	0.39	16.38	14.89	124.30	10.61	8.35
	3.65	14.74	5.72	0.39	13.44	8.86	93.76	16.39	10.58
	3.73	19.00	6.84	0.36	14.28	9.97	111.13	16.25	11.15
Min	2.78	14.74	4.44	0.28	11.76	7.86	56.23	8.86	6.08
Max	3.65	29.84	11.72	0.49	16.38	14.89	130.71	16.82	11.97
Average	3.13	19.28	7.49	0.39	14.14	10.91	97.63	13.30	9.04
SD	0.41	4.05	1.95	0.05	1.31	2.15	23.13	2.41	1.82
P value	0.2913	0.2318	0.0206	0.0225	1.0000	0.0016	0.0000	0.0034	0.0031

Table S5

Dog name	% of Length stretch	Stretch rate
Normal	5.00	7.98
	5.00	7.35
	5.00	5.67
	5.00	6.93
	5.00	7.35
	5.00	7.35
	5.00	6.09
	5.00	5.88
	5.00	7.56
	5.00	7.56
	5.00	6.51
	5.00	7.56
	5.00	7.56
	5.00	8.19
	5.00	6.51
	Min	5.00
	Max	8.19
Average	5.00	7.07
SD	0.00	0.77

Affected	5.00	6.93
	5.00	6.51
	5.00	7.14
	5.00	7.98
	5.00	5.88
	5.00	6.72
	5.00	7.56
	5.00	6.72
	5.00	7.35
	5.00	7.98
	5.00	6.93
	5.00	6.30
	5.00	8.19
	5.00	6.72
	5.00	7.14

Min	5.00	5.88
Max	5.00	8.19
Average	5.00	7.07
SD	0.00	0.65

P value 1.00