

# EFFICACY OF FOCAL MECHANIC VIBRATION TREATMENT ON BALANCE IN CHARCOT- MARIE-TOOTH 1A DISEASE: A PILOT STUDY.



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BACKGROUND AND AIM

ORIGINAL COMMU	NICATION	
	l mechanic vibration treat rie-Tooth 1A disease: a pilo	

The cause of postural instability in Charcot - Marie-Tooth (CMT) is still a matter of debate. Different factors contribute in CMT imbalance and a major challenge to maintain postural control has been attributed not only to the progressive loss of proprioception,<sup>[1,2]</sup> caused by both altered large myelinated fibers<sup>[2,3]</sup> and small fibers<sup>[4]</sup> but also to the reduced muscle strength and ankle instability.<sup>[5]</sup>

In the last years, a non-invasive approach based on a mechanical vibratory stimulation has been proposed to enhance balance and muscle performance.<sup>[6]</sup> Focal muscle vibration (fMV) was demonstrated as a highly selective stimulus for Ia spindle afferents<sup>[6-11]</sup>. In fact, vibratory stimuli with specific parameters (i.e., frequency of 100 Hz, peak-to-peak amplitude of 0.20-0.50 mm) may activate different mechanoceptors, in particular spindle afferents and Golgi tendon organs. Activation of peripheral contractile elements strongly influences the activity of the motoneuron system, and therefore the muscle spindle in providing afferent information. [6] This activation is able to determine a long-term reorganization of central nervous system both at spinal and cortical level <sup>[12]</sup> determining an improvement on several function such as motor performance and postural stability. <sup>[9,13]</sup>

The aim of our study was to evaluate the effects of focal mechanical vibration (fMV) on the balance of Charcot-Marie-Tooth (CMT) 1A patients.

#### MATERIAL AND METHODS

We enrolled 14 genetically confirmed CMT1A patients (8 female and 6 male, mean age 49.2 yrs, range 32-74, mean BMI: 25.2, range: 18.78-33.98, mean duration of disease: 13 years, range 1-30), regularly attending to our neurophysiological laboratory for periodic controls. Patients were excluded if they had other causes of peripheral neuropathy, previous chemotherapy or diseases that may cause or contribute to peripheral nerve damage (e.g., diabetes, renal insufficiency, alcohol abuse, vitamin B12 deficiency).

#### **Outcome Meaures:**

The primary outcome measure was the Berg Balance Scale (BBS) and the secondary were the dynamic gait index (DGI), the 6 Minutes Walking Test (6MWT), the muscular strength of lower limbs, the Quality of Life (QoL) guestionnaire and the stabilometric variables.

#### fMV Protocol:

Patients underwent a 3-days fMV treatment on guadriceps and triceps surae and were evaluated before the treatment (T0), 1 week after the end of the treatment (T1), and then after 1 month (T2). All patients received fMV over two muscular groups: 1) the quadriceps to improve stance control and muscle performance[8] and 2) triceps surae muscles to enhance plantar-flexor muscles force production by increasing muscle activation and proprioceptive messages. Patients were first asked to lie supine to treat the quadriceps and then prone during the application of fMV in the triceps surae muscles. During the application on both muscles, patients were asked to perform an isometric muscle contraction; in the first case they were asked to keep the popliteal cavum in contact with the bed, while for the second muscle, they were asked to keep the plantar flexion; the examiners clinically verified the muscle contraction throughout the whole fMV application.

fMV was delivered by using a specific device consisting of an electromechanical transducer, a mechanical support, and an electronic control device.

The transducer was positioned bilaterally on the quadriceps tendon close to the rectus femoris insertion at about 2 cm from the medial edge of the patella and on the triceps surae muscles at the myotendinous junction. For each muscle group, fMV was applied for 3 sessions of 10 minutes each, with an inter-session interval of 1-minute. The same protocol was repeated for 3 consecutive days in order to elicit "cumulative after-effects" and according to previously reported techniques.

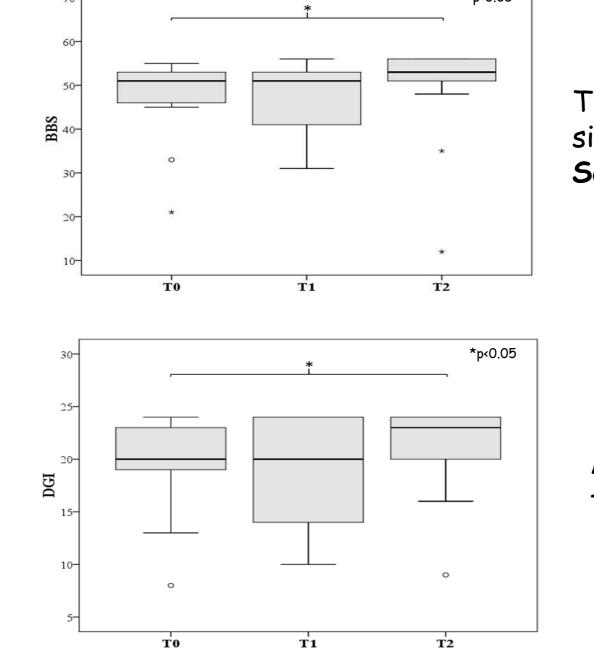
The mechanical support for delivering fMV allowed the compression of soft tissues overlying the muscle-tendon complex with low amplitude (200-500 µm) and high frequency (100 Hz) sinusoidal displacement.

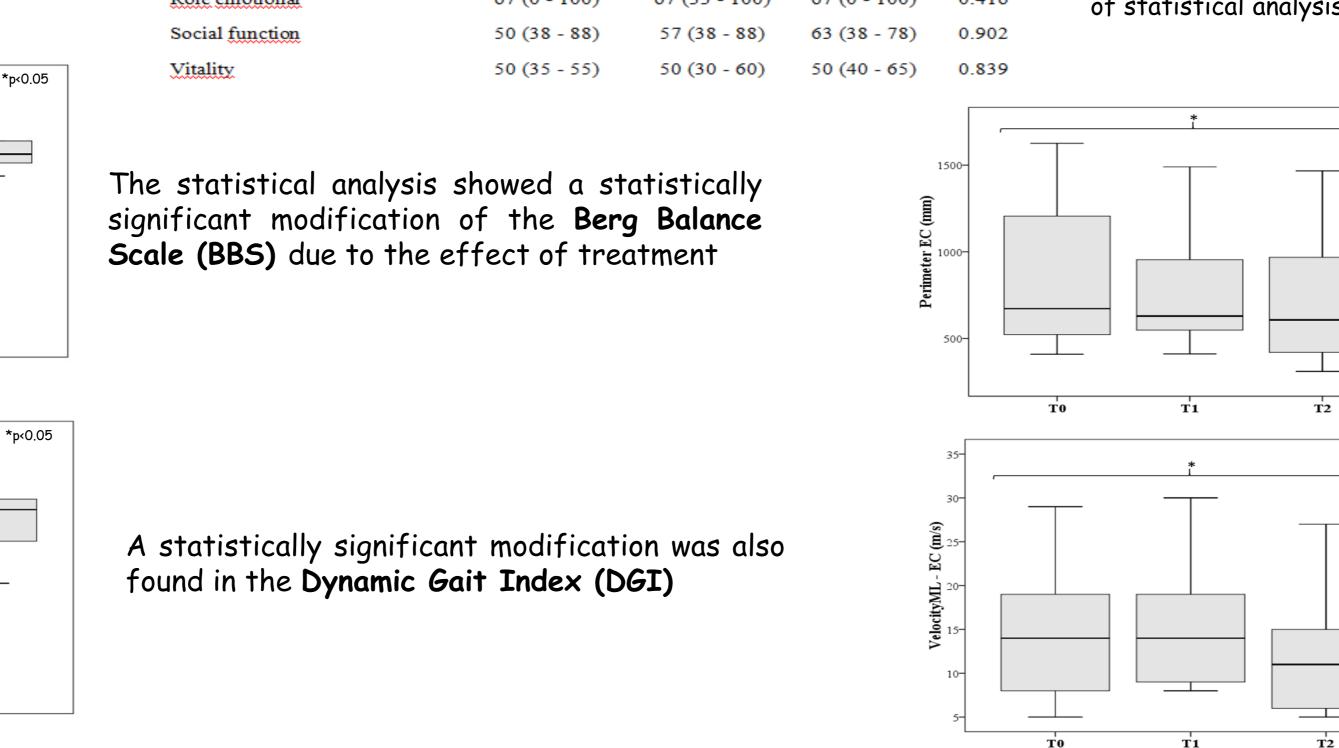
Note that the baseline and the follow up evaluations were performed by a blind examiner, this means that the examiner did not know the kind of treatment the patients underwent or whether the whole sample would receive an effective treatment.

RESULTS			TO	Tl	T2								
	Patient Age Gender CMTES		Variables	Median (IQR)	Median (IQR)	Median (IQR)	P <u>value</u>						
Pt Pt Pt Pt Pt Pt Pt Pt Pt Pt Pt	Pt 1 36 M	M 4 -   M 6   F 6   M 7   F 2   M 5   F 1   M 8   F 9   F 1   F 8   F 2	Berg Balance Scale	49 (45 - 53)	51 (41 - 53)	53 (50 - 56)	0.024	Variables	T0	T1	T2	Bushus	
	Pt 2     50     M       Pt 4     46     F		Dynamic Gait Index	20 (18 - 23)	20 (14 - 24)	23 (20 - 24)	0.018		Median (IQR)	Median (IQR)	Median (IQR)	P value	
	Pt 5     74     M       Pt 6     56     F		6 Minute Walking Test	349 (282 - 381)	342 (303 - 384)	363 (337 - 402)	0.583	<u>Eves</u> open					
	Pt 7 46 M		MRC					Swayap (mm)	4.5 (4 - 5)	4 (3 - 5)	4 (3 - 6)	0.539	
	Pt 8     37     F       Pt 9     44     M		Hip Flexion (LS)	5 (4 - 5)	5 (5 - 5)	5 (5 - 5)	0.064	Swayng (mm)	4 (2 - 4)	3.5 (3 - 5)	3 (2 - 4)	0.122	
	Pt 10 65 F Pt 11 40 F		Hip Flexion (RS)	5 (3.5 - 5)	5 (5 - 5)	5 (4 - 5)	0.161	VelocityAp(mm/s)	12 (7 - 15)	10 (7 - 13)	10 (7 - 14)	0.063	
	Pt 1265FPt 1339F		Knee Extension (LS)	5 (4 - 5)	5 (5 - 5)	5 (4.8 - 5)	0.504	Velocitym_ (mm/s)	9 (5 - 11)	10 (6 - 13)	7 (4 - 9)	0.083	
	Pt 14 32 M		Knee Extension (RS)	5 (4.5 - 5)	5 (5 - 5)	5 (4 - 5)	0.192	Sway pathlength	497 (330 - 596)	514 (318 - 650)	391 (320 - 515)	0.465	
			Knee Flexion (LS)	5 (5 - 5)	5 (5 - 5)	5 (5 - 5)	0.071	(mm)					
			Knee Flexion (RS)	5 (5 - 5)	5 (5 - 5)	5 (4.8 - 5)	0.173	Area (mm <sup>2</sup> )	247 (202 - 341)	264 (213 - 299)	204 (124 - 306)	0.238	
	-	nows the demographical and ta of CMT 1 A patients	Foot Dorsiflexion (LS)	2 (1 - 4)	2.5 (2 - 4)	2 (0 - 4)	0.507	Exes closed					
	al data of CMTTA pa		Foot Dorsiflexion (RS)	2 (0 - 3)	2.5 (1 - 4)	2 (0 - 4)	0.223	Swayar (mm)	6.5 (5 - 8)	5 (5 - 9)	5 (4 - 7.3)	0.159	
			Foot Plantarflexion (LS)	3 (2 - 4)	4.5 (3 - 5)	4 (2 - 5)	0.131	Swayng (mm)	6 (3 - 9)	6 (4 - 7)	4 (3 - 6.2)	0.138	
			Foot Plantarflexion (RS)	3 (2 - 4)	4 (3 - 5)	3 (2 - 5)	0.131	Velocity <sub>AP</sub> (mm/s)	17 (13 - 32)	17 (13 - 22)	13 (10 - 24)	0.441	
			Hallux Ext Long (LS)	1.5 (0 - 3)	0.5 (0 - 4)	0 (0 - 3)	0.630	VelocitymL (mm/s)	15 (8 - 19)	15 (9 - 19)	11 (5.8 - 16)	0.038	
		ole 2 shows the values (median and	Haller Fret Lana (DS)					Sway pathlength		667 (549 - 948)		0.043	
			N#-10	1 (0 - 3)	1 (0 - 4)	0 (0 - 4)	0.630	(mm)					
interquartile range) of the clinical scales		D 1'1 '	62 (41 - 84)	52 (41 - 100)	84 (47 - 100)	0.641	Area (mm²)	570 (272 – 1044)	571 (356 - 647)	318 (201 - 544)	0.198		
aft med		baseline (TO), after 1 week (T1) and ter 1 month (T2) from the focal		55 (30 - 67)	52 (30 - 72)	50 (38 - 70)	0.678	RombergLength	1.6 (1.4 - 2.3)	1.7 (1.4 - 2)	1.7 (1.3 - 2)	0.397	
	•	n treatment and the		47 (36 - 53)	46 (34 - 53)	47 (35 - 52)	0.861	Rombergaces	2.3 (1.3 - 4.9)	2.1 (1.1 - 3.9)	1.6 (1.2 - 3.5)	0.583	
	results of statistical		Mental health	68 (56 - 80)	68 (52 - 80)	64 (59 - 80)	0.716						
			Physical composite score	43 (35 - 49)	44 (34 - 51)	44 (35 - 52)	0.099						
			Physical function	63 (50 - 85)	73 (50 - 85)	80 (39 - 90)	0.281	Table 3 shows the values (median and interquartile range					
		Role physical	63 (25 - 100)	75 (0 - 100)	75 (0 - 100)	0.771	stabilometric assessment at baseline (TO), after 1 week (T1) and month (T2) from the focal mechanical vibration treatment and the						
		Role emotional	67 (0 - 100)	67 (33 - 100)	67 (0 - 100)	0.416							
			Social function	50 (38 - 88)	57 (38 - 88)	63 (38 - 78)	0.902	of statistical analy					
			Vitality	50 (35 - 55)	50 (30 - 60)	50 (40 - 65)	0.839						
70-		*p<0.05	Y ILLILLY	50 (55 - 55)	50 (50 - 00)	50 (40 - 05)	0.039						

of the after 1 results

\*p<0.05





stabilometric Concerning the variables found significant we changes only for the eyes closed condition;

particular, statistically in ۵ significant decrease was found in Velocity<sub>MI</sub> and Perimeter.

### CONCLUSIONS

Our pilot study, although based on a small sample, showed that fMV is able to determine an improvement of balance in CMT 1 A patients. More studies, conducted on a larger sample of CMT patients of different genetic mutations and with different scores of disability, are needed to confirm this preliminary data.

REFERENCES

1) Vinci P, Perelli S, Esposito C (2001) Charcot-Marie-Tooth Disease: poor balance and rehabilitation. J Peripher Nerv Syst 6:58-58; 2) van der Linden MH, van der Linden SC, Hendricks HT, et al. (2010) Postural instability in Charcot-Marie-Tooth type 1A patients is strongly associated with reduced somatosensation. Gait Posture 31:483-8; 3) Lencioni T, Piscosquito G, Rabuffetti M, et al. (2015) The influence of somatosensory and muscular deficits on postural stabilization: Insights from an instrumented analysis of subjects affected by different types of Charcot-Marie-Tooth disease. Neuromuscul Disord 25:640-5; 4) Nardone A, Tarantola J, Miscio G, et al. (2000) Loss of large-diameter spindle afferent fibres is not detrimental to the control of body sway during upright stance: evidence from neuropathy. Exp brain Res 135:155-62; 5) Lencioni T, Rabuffetti M, Piscosquito G, et al. (2014) Postural stabilization and balance assessment in Charcot-Marie-Tooth 1A subjects. Gait Posture 40:481-6; 6) Filippi GM, Brunetti O, Botti FM, et al. (2009) Improvement of stance control and muscle performance induced by focal muscle spindles. J Bianconi R, van der Meulen J (1963) The response to vibration of the end organs of mammalian muscle spindles. J Neurophysiol 26:177-90; 8) Matthews PB, Watson JD (1981) Action of vibration on the response of cat muscle spindle Ia afferents to low frequency sinusoidal stretching. J Physiol 317:365-81; 9) Fattorini L, Ferraresi A, Rodio A, et al. (2006) Motor performance changes induced by muscle vibration. Eur J Appl Physiol 98:79-87; 10) Fallon JB, Macefield VG (2007) Vibration sensitivity of human muscle spindles and Golgi tendon organs. Muscle Nerve 36:21-9; 11) Sorensen KL, Hollands MA, Patla E (2002) The effects of human ankle muscle vibration on posture and balance during adaptive locomotion. Exp brain Res 143:24-34; 12) Murillo N, Valls-Sole J, Vidal J, et al. (2014) Focal vibration in neurorehabilitation. Eur J Phys Rehabil Med 50:231-42; 13) Brunetti O, Filippi GM, Lorenzini M, et al. (2006) Improvement of posture stability by vibratory stimulation following anterior cruciate ligament reconstruction. Knee Surg Sports Traumatol Arthrosc 14:1180-7.