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The impact of lockdown on Functional Motor Disorders patients during the first COVID-19 outbreak: acase-control study

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Functional Motor Disorders (FMDs) are frequent and disabling neurological disorders characterized by motor abnormalities that are inconsistent over time, altered by distraction and clinically incongruent with classical movement disorders [1]. Non-motor symptoms (NMS), including anxiety, pain, fatigue and so forth, common to the general population, are often part of the clinical spectrum [1]. Traditionally viewed as "psychogenic", FMDs raise important concerns in terms of increased susceptibility to stressful conditions and environmental dramatic events [2]. The 2019-Coronavirus disease (COVID-19) pandemic and the severe restrictions adopted during lockdown to limit virus diffusion, might have been expected to exacerbate functional symptoms, also given the increased psychological distress, depression and stress reported in the general population [3]. Interestingly, despite increased levels of NMSs in FMDs, particularly anxiety [4-6], functional motor symptoms have not been clearly reported to worsen during lockdown [4-6]. However, the available studies were limited by the lack of a control group [4-6] or by a small sample size [4-6].

In Italy, the worst-hit European Country by the first outbreak of COVID-19, a strict lockdown of all unnecessary activities was imposed from March to the end of April 2020, with radical measures for social distancing and limitations of all outdoor activities, including physical activity. In our study, we explored the impact of lockdown on motor, numerous NMS and global health status (GHS) in a large cohort of FMDs patients and compared the results with those obtained in healthy controls (HC), matched for age and sex. Patients with an established diagnosis of FMDs followed at our specialized FMDs clinic in Verona and HC, who were friends or relatives of researchers, matched for age and sex, were contacted in October 2020, via phone or mail and given the link for a semi-anonymous Survey and two weeks to complete it. All provided informed consent, and the local Ethics Committee approved the study. We employed a 39-item Survey divided into three sections: i) sociodemographics ii) patient-estimated severity of symptoms during lockdown: motor and NMS, assessed through a 5-point Likert scale (1 = none to 5 = severe) and GHS, assessed using a 5-point scale (1 = excellent to 5 = poor) iii) changes in symptoms severity compared to pre-lockdown (February 2020), assessed through a 3-point Likert scale (improved/ unchanged/worsened) and changes in adherence to home based physical exercise, if prescribed.

Results are shown in Table 1 and main demographics and clinical data are reported in the Supplementary material. We recruited 73 out of 96 contacted patients (response rate 76%) and 101 HC.

During lockdown, motor symptoms were minimal to mild in 62%. Compared to controls, patients showed comparable levels of anxiety, depression, apathy, physical and mental fatigue and suffered from more severe levels of pain and sleep disturbances (Mann-Whitney, p < 0.005). Compared to pre-lockdown, most patients reported stability of motor symptoms (severity and frequency) and NMS, except for mental fatigue that worsened in \sim 50% of patients. In particular, motor symptoms remained stable in ${\sim}60\%$ of patients, worsened in about one third and 18% of patients reported new-onset motor symptoms. Worsening in severity of motor symptoms was significantly predicted by mental fatigue in a multiple regression analysis (Table 1). Compared to prelockdown, pain worsened significantly more in patients than controls, while anxiety and depression worsened significantly more in controls than in patients (all, Fisher test p < 0.001). Patients reported a significantly worse GHS than controls (Mann-Whitney, p < 0.001), but it did not change due to lockdown, as in HC. Almost 70% of patients who were prescribed with home based physical exercise plan continued exercising, although 36% of them less frequently than the pre-lockdown period.

In line with previous studies, our results confirm that motor symptoms remained stable in the majority of patients [4–6]. In contrast with previous findings, our cohort of patients showed stable NMS [4–6]. Moreover, compared to controls, only pain worsened significantly more in patients, possibly representing a specific manifestation of the stress related to COVID-19 in FMD patients, while anxiety and depression worsened significantly more in controls. Although with inevitable recall bias, this study suggests that patients with an established diagnosis of FMDs have shown a substantial stability of functional motor, MNMS and

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[;] FMDs, Functional Motor Disorders; NMS, Non-motor symotoms; GHS, Global Health Status.

Table 1

Clinical variables in FMD and HC and their associations with worsening in severity of motor symptoms.

Table 1 (continued)

Aotor symptoms, NMS and GHS during lockdown						FMD
	FMD	НС	Test	P-value Z value (Mann- Whitney Test) X-squared (Chi-squared Test) OR (Fisher Test)	Improved	14% (n 10)
Motor		-	-	-	Frequency of Motor	
symptoms (5- point Likert)					symptoms n	
Mean \pm SD	$2.90 \pm 1.15;$				(%)	
Median (IQR)	3.00 (2–4)				Worsened	27% (n
None	11% (n = 8)				Unchanged	20) 63% (n
Minimal	26% (n = 19)				Unchanged	46)
Mild	36% (n =				Improved	10% (n
	26)				Anxiety n (%)	
Moderate	16% (n =				Worsened	34% (n
	12)				Unchanged	25) 50% (p
Severe	11% (n = 8)				Unchanged	59% (n 43)
I do not Know I felt anxious (5-	-		Mann-		Improved	7% (n =
point Likert)			Whitney		Depression n	
Mean \pm SD	2.53 ± 1.14	2.54 ± 0.95		p = 0.863	(%)	
Median (IQR)	3.00 (2-3)	3.00 (2–3)		Z = -0.172	Worsened	41% (n
I felt depressed			Mann-			30)
(5-point			Whitney		Unchanged	48% (n 35)
Likert)	2.72 ± 1.20	2.64 ± 1.02		- 0.975	Improved	33) 11% (n
Mean ± SD Median (IQR)	2.73 ± 1.29 3.00 (2-4)	2.64 ± 1.02 3.00 (2-3)		p = 0.875 Z = -0.157	Physical Fatigue	1170 (ii
I felt physically	3.00 (2-4)	3.00 (2-3)	Mann-	$\Sigma = -0.137$	n (%)	
fatigued (5-			Whitney		Worsened	45% (n
point Likert)			-			33)
Mean \pm SD	$\textbf{3.04} \pm \textbf{1.21}$	$\textbf{2.36} \pm \textbf{1.20}$		p = 0.017	Unchanged	47% (n
Median (IQR)	3.00 (2–4)	2.00 (1–3)		Z = -2.379	Improved	34) 8% (n =
I felt mentally fatigued (5-			Mann- Whitney		Mental Fatigue	0%) (II -
point Likert)			winney		n (%)	
Mean \pm SD	2.90 ± 1.22	2.56 ± 1.13		p = 0.075	Worsened	49% (n
Median (IQR)	3.00 (2-4)	3.00 (2–3)		Z = -2.379		36)
I felt			Mann-		Unchanged	45% (n
demotivated			Whitney		Improved	33) 6% (n =
(5-point Likert)					Apathy n (%)	0%0 (II -
Mean \pm SD	2.46 ± 1.31	2.34 ± 1.16		p = 0.728	Worsened	38% (n
Median (IQR)	2.40 ± 1.31 2.00 (1-3)	2.34 ± 1.10 2.00 (1-3)		p = 0.728 Z = -0.347		28)
I felt pain (5-			Mann-		Unchanged	50% (n
point Likert)			Whitney			37)
Mean \pm SD	$\textbf{2.76} \pm \textbf{1.29}$	1.50 ± 0.78		p < 0.001	Improved	12% (n
Median (IQR)	3.00 (2–4)	1.00 (1–2)		Z = -6.783	Pain n (%)	
I had trouble sleeping (5-			Mann- Whitney		Worsened	33% (n
point Likert)			winney			24)
Mean \pm SD	2.64 ± 1.45	1.99 ± 1.09		p = 0.004	Unchanged	57% (n
Median (IQR)	3.00 (1-3)	2.00 (1-3)		Z = -2.893		42)
Global Health		$\textbf{3.02} \pm \textbf{0.81}$	Mann-		Improved	10% (n
Status (5-		3.00 (3-3)	Whitney		Quality of Sleep n (%)	
point Likert) Mean ± SD	2.19 ± 0.97			n < 0.001	Worsened	32% (n
Median (IQR)	2.19 ± 0.97 2.00 (1–3)			p < 0.001 Z = -5.416		23)
				2 0.110	Unchanged	60% (n
Changes in motor Symptoms	symptoms, NM FMD	S and GHS HC	Test	P value X-		44)
symptoms	FWD	нс	Test	squared (Chi- squared test)	Improved General Health Status n (%)	8% (n =
Severity of		-	-	-	Worsened	32% (n
Motor						23)
symptoms n					Unchanged	58% (n
(%) Worsened	30% (n =				Turn 1	42)
worsened	30% (fi = 22)				Improved	11% (n
Unchanged	56% (n =				Adherence to Physical	
-	41)				Exercise	

Motor symptoms, N	MS and GHS du	ring lockdown		
	FMD	HC	Test	P-value Z value (Mann- Whitney Test) X-squared (Chi-squared Test) OR (Fisher Test)
Improved	14% (n = 10)			
Frequency of Motor symptoms n (%)	10)	-		
Worsened	27% (n = 20)			
Unchanged	63% (n = 46)			
Improved	10% (n = 7)			
Anxiety n (%)	240/ (-	E00/ (Fisher Test	p = 0.002
Worsened	34% (n = 25)	58% (n = 59)		
Unchanged	59% (n =	41% (n =		
<u> </u>	43)	41)		
Improved	7% (n = 5)	1% (n = 1)	mit 1	0.000
Depression n			Fisher Test	p < 0.001
(%) Worsened	41% (n =	72% (n =		
	30)	73)		
Unchanged	48% (n =	25% (n =		
Improved	35) $11\% (n - 8)$	25) $3\% (n - 3)$		
Improved Physical Fatigue n (%)	11% (n = 8)	3% (n = 3)	Chi-squared	p = 0.96
Worsened	45% (n =	47% (n =		X-squared =
	33)	48)		0.092
Unchanged	47% (n = 34)	45% (n = 45)		
Improved Mental Fatigue n (%)	8% (n = 6)	8% (n = 8)	Fisher Test	p = 0.47
Worsened	49% (n =	58% (n =		
	36)	59)		
Unchanged	45% (n = 33)	36% (n = 36)		
Improved	6% (n = 4)	6% (n = 6)	Chicanord	
Apathy n (%) Worsened	38% (n =	55% (n =	Chi-squared	
	28)	56)		
Unchanged	50% (n =	40% (n =		p = 0.055
Improved	37) 12% (n = 9)	40) 5% (n = 5)		X-squared =
	1=/0 (n -))	575 (n = 5)		5.79
Pain n (%)			Fisher Test	p < 0.001
Worsened	33% (n =	15% (n =		
Unchanged	24) 57% (n =	15) 84% (n =		
	42)	85)		
Improved	10% (n = 7)	1% (n = 1)		
Quality of Sleep			Fisher Test	p = 0.085
n (%) Worsened	32% (n =	42% (n =		
	23)	42)		
Unchanged	60% (n = 44)	56% (n = 57)		
Improved	8% (n = 6)	2% (n = 2)		
General Health			Fisher Test	0.089
Status n (%) Worsened	32% (n =	30% (n =		
	23)	30)		
Unchanged	58% (n =	67% (n =		
Improved	42) 11% $(n - 8)$	68) $3\% (n - 3)$		
Improved Adherence to	11% (n = 8)	3% (n = 3)		
Physical				
Production of the second				

(continued on next page)

Table 1 (continued)

Higher frequency

Motor symptoms, N	NMS and GHS d	uring lockdown		
	FMD	НС	Test	P-value Z value (Mann- Whitney Test) X-squared (Chi-squared Test) OR (Fisher Test)
(prescribed in $n = 55$) n (%) Not performed	31% (n =	_		
	17)			
	Lower	36% (n =		
	frequency -	20)		
	Same	26% (n =		
	frequency	14)		
	-			

Associations of NMS and adherence to rehabilitation with worsening in severity of motor symptoms in FMDs

7% (n = 4)

of motor symptoms in FMDs						
	Single comparison		Multivariable Logistic model			
NMS Changes	Odds ratio	p-value	Odds ratio	p-value		
during						
lockdown in						
FMDs						
Adherence to	3.81	0.061	1.98	0.52		
rehabilitation	[0.94–15.4]		[0.24–16.3]			
Anxiety	3.51	0.019*	1.21	0.86		
	[1.23-10]		[0.16–9.13]			
Depression	5.14	0.003**	0.63	0.66		
	[1.75–15.2]		[0.08–5.03]			
Physical Fatigue	29.23	< 0.001***	2.16	0.57		
	[6–142]		[0.15–30.7]			
Mental Fatigue	50.4	<0.001***	30.82	0.012*		
	[6.21-409]		[2.11-450]			
Apathy	3.47	0.019*	2.38	0.37		
	[1.22–9.82]		[0.35–16.1]			
Pain	14.33	< 0.001***	1.46	0.72		
	[4.3–47.8]		[0.18 - 11.8]			
Quality of sleep	4.36	0.007**	1.36	0.74		
	[1.49–12.7]		[0.22-8.56]			

Abreviations: FMD, Functional Motor Disorders; HC, Healthy Controls. NMS, Non-motor Symptoms; GHS, General Health Status.

Legend: the Table shows motor symptoms, non-motor symptoms (NMS), general health status during lockdown (March–April 2020), their changes compared to pre-lockdown (February 2020) and regression analyses between changes in NMS in FMD and worsening in motor symptoms severity. All variables were classified as 0 or 1 (0 = unchanged, improved; 1 = worsened).

*p value adjusted for multiple comparisons (Bonferroni correction) p < 0.007 was statistically significant. Statistically significant results are given in bold.

GHS during lockdown. In the context of social isolation and home confinement, decreased operational demands and lower psychosocial strains, compared to everyday life, could have prevented worsening of symptoms. Moreover, FMDs are known to be affected by self-attention/monitoring [3] and the diverted attention toward external events due to a global pandemic might have further influenced

our findings. We are now facing unpredictable phases of a longstanding pandemic and future studies are needed to elucidate long-term effects of COVID-19 pandemic on FMDs.

Appendix A. Supplementary data

Supplementary data to this article can be found online at https://doi.org/10.1016/j.parkreldis.2021.11.007.

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