ORIGINAL ARTICLE



Long COVID: rheumatologic/musculoskeletal symptoms in hospitalized COVID-19 survivors at 3 and 6 months

Fatih Karaarslan¹ · Fulya Demircioğlu Güneri¹ · Sinan Kardeş²

Received: 7 August 2021 / Revised: 22 September 2021 / Accepted: 23 September 2021 / Published online: 29 October 2021 © International League of Associations for Rheumatology (ILAR) 2021

Abstract

Objective To document the detailed characteristics including severity, type, and locations of rheumatic and musculoskeletal symptoms along with other COVID-19 persistent symptoms in hospitalized COVID-19 survivors at 3 and 6 months.

Methods In this extension cohort study, two telephone surveys at 3 and 6 months following the hospitalization were carried out. In these telephone surveys, participants were asked regarding their symptoms through a previously designed standard questionnaire. **Results** At 3 months, 89.0% of survivors had at least one symptom, 74.6% had at least one rheumatic and musculoskeletal symptom, and 82.1% had at least one other COVID-19 symptom. At 6 months, 59.6% of survivors had at least one symptom, 43.2% had at least one rheumatic and musculoskeletal symptom, and 51.2% had at least one other COVID-19 symptom. Regarding the rheumatic and musculoskeletal symptoms, 31.6% had fatigue, 18.6% had joint pain, and 15.1% had myalgia; and regarding the other-COVID-19-symptoms, 25.3% had dyspnea, 20.0% had hair loss, and 17.2% sweat at 6 months. In an adjusted model, female patients were more likely to have fatigue (OR: 1.99, 95% CI: 1.18–3.34), myalgia (3.00, 1.51–5.98), and joint pain (3.39, 1.78–6.50) at 6 months.

Conclusion Approximately 3 in 5 patients had at least one symptom with ≈ 2 in 5 patients had at least one rheumatic and musculoskeletal symptom. Fatigue, joint pain, and myalgia were the most frequent rheumatic and musculoskeletal symptoms. Joint pain and myalgia were mostly widespread. This information guide rheumatologists to understand the nature and features of persistent rheumatic and musculoskeletal symptoms in hospitalized COVID-19 survivors and may contribute to better management of these individuals.

Key Points

- Approximately 3 in 5 patients had at least one symptom with ≈ 2 in 5 patients had at least one rheumatic and musculoskeletal symptom at 6 months
- Fatigue, joint pain, and myalgia were the most frequent rheumatic and musculoskeletal symptoms followed by back pain, low back pain, and neck pain
- Dyspnea, hair loss, and sweat were the most frequent other-COVID-19-symptoms

Keywords Fatigue · Joint pain · Long-haul COVID · Muscle pain · Post-acute COVID-19 syndrome · SARS-CoV-2

Fatih Karaarslan fatih.karaarslan@sbu.edu.tr

> Fulya Demircioğlu Güneri fuliad@hotmail.com

Sinan Kardeş sinan.kardes@istanbul.edu.tr

¹ Department of Medical Ecology and Hydroclimatology, Gulhane Training and Research Hospital, University of Health Sciences, Ankara, Turkey

² Department of Medical Ecology and Hydroclimatology, Istanbul Faculty of Medicine, Istanbul University, Istanbul, Turkey

Introduction

Since the emergence in the late of 2019, severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) has spread across the world leading an unexpected, large, and serious global pandemic [1–5]. By the 5th of July 2021, 182,319,261 confirmed cases of coronavirus disease 2019 (COVID-19), involving 3,954,324 deaths, have been identified worldwide according to the World Health Organization (WHO) [6]. Even though as COVID-19 vaccination becomes more available throughout the world and the COVID-19 infection/death

Clinical Rheumatology (2022) 41:289-296

rates are dropping, clinicians have still caring the COVID-19 survivors who were previously infected and recovered from acute infection and now experience a broad range of persistent symptoms, which are called as post-acute COVID-19 syndrome, long COVID, long-haul COVID, post-COVID condition, or post-acute sequelae of SARS-CoV-2 infection (PASC) [7, 8].

Several observational studies have recently evaluated the persistent symptoms in hospitalized COVID-19 survivors beyond 3 months [9-13]. Although some of these studies have partly described the persistent fatigue, myalgia, and joint pain in hospitalized COVID-19 survivors, they only reported their prevalence. In other words, the severity, type (local or widespread), and locations of persistent rheumatic and musculoskeletal symptoms have not been documented in detail.

Thus, we aimed to document the detailed characteristics including severity, type, and locations of rheumatic and musculoskeletal symptoms along with other COVID-19 persistent symptoms in hospitalized COVID-19 survivors at 3 and 6 months. We also sought to evaluate whether an association exists between the presence of symptoms (i.e., fatigue, myalgia, joint pain) and age, sex, body mass index (BMI), and duration of hospital stay.

Materials and methods

Study design, setting, and ethical approval

We conducted an extension cohort study of our previously published investigation reporting 1-month results [14]. We carried out this study in a tertiary hospital, namely Gülhane Training and Research Hospital. We obtained an ethical approval for this extension study by the Ethic Committee of the Gülhane Scientific Researches, University of Health Sciences (2021/187). We obtained informed verbal consent to participate in the study at the beginning of the telephone interviews. We informed by Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) [15] in reporting our study and by standard recommendations [16] in reporting our results.

Study cohort

COVID-19 survivors who aged 18 to 70 years old and who had been discharged after hospital stay due to the acute COVID-19 infection anytime during the November 18, 2020, through January 30, 2021, were included. We included COVID-19 survivors who had been only treated in hospital ward unit; in other words, we excluded survivors who admitted/or were transferred to intensive care unit during their hospital stay.

Data gathering

One author (FK) carried out all telephone surveys at 3 and 6 months following the hospitalization. In these telephone surveys, participants were asked regarding their symptoms through a previously designed standard questionnaire. The following rheumatic and musculoskeletal symptoms were systematically gathered: fatigue, myalgia, joint pain, low back pain, back pain, and neck pain. The type (local, widespread) and locations of myalgia, and joint pain were asked as well. Additionally, the other COVID-19 symptoms were gathered: fever, cough, lack of appetite, dyspnea, diarrhea, sore throat, headache, dizziness, absence of taste, and absence of smell. We evaluated the severity of all these symptoms through a 5-point Likert-type scale.

Statistical analyses

We conducted the analysis with the use of the SPSS v. 21.0 (IBM, Armonk, NY). We analyzed the data and variables through descriptive statistical analyses and expressed the results either as mean \pm standard deviation (SD) or number (%). We also tested whether there is an association between the presence of symptoms (i.e., fatigue, myalgia, joint pain) and age, sex, BMI, and duration of hospital stay at 6 months using generalized estimating equations by a selection of binary logistic regression models. The 95% confidence interval (CI) values that did not include 1 indicated statistical significance.

Results

Baseline characteristics

We included a total of 300 COVID-19 survivors in our previous publication presenting 1-month results [14]. At 3 months, there were 9 missing, and at 6 months, there were 6 additional missing survivors. The reasons of these 15 missings were dead (n=2), could not be reached through telephone (n=4), did not answer to telephone (n=7), staying in intensive care unit at the time of assessment (n=1), and abdominal hysterectomy with bilateral salpingo-oophorectomy (n=1). We presented the baseline demographic and clinical features of the study cohort at 3 (n=291) and 6 (n=285) months in Table 1.

Symptoms at 3 months

At 3 months, 89.0% of survivors had at least one symptom (in other words, 11.0% had no symptoms), 74.6% had at least one rheumatic and musculoskeletal symptom, and 82.1% had at least one other-COVID-19-symptom. Regarding

Table 1Baseline characteristicsof study population, whichhave data for 3- and 6-monthfollow-up

Parameter	3 months $(n=291)$	6 months $(n=285)$	
Age, yrs	52.54 ± 12.03	52.32 ± 12.05	
Male	173 (59.5)	172 (60.4)	
Female	118 (40.5)	113 (39.6)	
BMI, kg/m ²	28.96 ± 4.76	28.92 ± 4.77	
Schooling grade			
Illiterate	8 (2.7)	8 (2.8)	
Primary	101 (34.7)	97 (34.0)	
Junior high	31 (10.7)	31 (10.9)	
High	51 (17.5)	51 (17.9)	
University	100 (34.4)	98 (34.4)	
Employed	126 (43.3)	125 (43.9)	
Alcohol usage			
Current users	15 (5.2)	15 (5.3)	
None	276 (94.8)	270 (94.7)	
Smoking status			
Current smoker	22 (7.6)	22 (7.7)	
Nonsmoker	220 (75.6)	214 (75.1)	
Ex-smoker	49 (16.8)	49 (17.2)	
Smoking, pack-years	26.70 ± 15.97	26.70 ± 15.97	
Comorbidities			
At least one comorbidity	190 (65.3)	185 (64.9)	
Hypertension	93 (32.0)	88 (30.9)	
Diabetes mellitus	83 (28.5)	80 (28.1)	
Hyperlipidemia	26 (8.9)	25 (8.8)	
Coronary artery disease	42 (14.4)	40 (14.0)	
Thyroid diseases	12 (4.1)	12 (4.2)	
Asthma	19 (6.5)	19 (6.7)	
COPD	6 (2.1)	6 (2.1)	
Osteoarthritis	8 (2.7)	8 (2.8)	
Familial Mediterranean fever	4 (1.4)	4 (1.4)	
Rheumatoid arthritis	5 (1.7)	5 (1.7)	
Scleroderma	1 (.4)	1 (.4)	
Ankylosing spondylitis	1 (.4)	1 (.4)	
Psoriatic arthritis	1 (.4)	1 (.4)	
Sjogren syndrome	1 (.4)	1 (.4)	
SARS-CoV-2 RT-PCR positive	255 (87.6)	249 (87.3)	
SARS-CoV-2 infection chest CT scan findings	263 (90.4)	257 (90.1)	
Duration from PCR test to hospitalization, days	6.43 ± 3.45	6.42 ± 3.45	
Duration of hospital stay, days	7.63 ± 3.96	7.58 ± 3.95	

Data are prevalence (%) or mean \pm standard deviation. *BMI* body mass index, *COPD* chronic obstructive pulmonary disease, *CT* computed tomography, *RT-PCR* reverse transcriptase–polymerase chain reaction, *SARS-CoV-2* severe acute respiratory syndrome coronavirus 2

the rheumatic and musculoskeletal symptoms, 59.5% had fatigue, 40.6% had myalgia, and 39.2% had joint pain. Regarding the other COVID-19 symptoms, 46.1% had hair loss, 45.0% had dyspnea, and 26.8% had sweat (Tables 2 and 3, Fig. 1).

Symptoms at 6 months

At 6 months, 59.6% of survivors had at least one symptom (in other words, 40.4% had no symptoms), 43.2% had at least one rheumatic and musculoskeletal symptom, and 51.2% had at least one other COVID-19 symptom. Regarding the rheumatic and musculoskeletal symptoms, 31.6% had fatigue,

 Table 2
 Prevalence of symptoms at 3 and 6 months

months

Parameter	3 months	6 months	
At least one symptom	259 (89.0)	170 (59.6)	
At least one rheumatic/musculo- skeletal symptom	217 (74.6)	123 (43.2)	
Fatigue	173 (59.45)	90 (31.58)	
Myalgia	118 (40.55)	43 (15.09)	
Joint pain	114 (39.18)	53 (18.59)	
Low back pain	72 (24.74)	32 (11.23)	
Back pain	92 (31.62)	41 (14.39)	
Neck pain	60 (20.62)	27 (9.47)	
At least one other COVID-19 symptom	239 (82.1)	146 (51.2)	
Fever	8 (2.75)	1 (0.35)	
Cough	40 (13.75)	16 (5.61)	
Lack of appetite	19 (6.53)	6 (2.11)	
Dyspnea	131 (45.02)	72 (25.26)	
Diarrhea	18 (6.19)	8 (2.81)	
Sore throat	38 (13.06)	19 (6.67)	
Headache	71 (24.40)	27 (9.47)	
Dizziness	55 (18.90)	16 (5.61)	
Absence of taste	24 (8.25)	11 (3.86)	
Absence of smell	37 (12.71)	15 (5.26)	
Sweat	78 (26.80)	49 (17.19)	
Hair loss	134 (46.05)	57 (20.00)	

Data are prevalence (%)

18.6% had joint pain, and 15.1% had myalgia. Regarding the other COVID-19 symptoms, 25.3% had dyspnea, 20.0% had hair loss, and 17.2% sweat (Tables 2 and 3, Fig. 1).

Location of arthralgia and myalgia at 6 months

Joint pain and myalgia were widespread (64.2% and 69.8%, respectively); if regional, joint pain was mostly in the knee, foot–ankle, and shoulder, and myalgia was mostly in the lower leg, arm, and shoulder girdle (Table 4).

Regression analyses results

In an adjusted model, female patients were more likely to have fatigue (OR: 1.99, 95% CI: 1.18–3.34), myalgia (3.00, 1.51–5.98), and joint pain (3.39, 1.78–6.50) at 6 months, whereas no association was observed between age/BMI/duration of hospital stay and fatigue/myalgia/joint pain (Table 5).

Table 3 Severity of s	ymptoms at 3 and 6 months	
Parameter	3 months	6
Fotiguo		

Fatigue		
None	118 (40.5)	195 (68.4)
Mild	62 (21.3)	60 (21.1)
Moderate	80 (27.5)	21 (7.4)
Severe	25 (8.6)	8 (2.8)
Very severe	6 (2.1)	1 (.4)
Myalgia		
None	173 (59.5)	242 (84.9)
Mild	35 (12.0)	22 (7.7)
Moderate	55 (18.9)	15 (5.3)
Severe	24 (8.2)	6 (2.1)
Very severe	4 (1.4)	0
Joint pain		
None	177 (60.8)	232 (81.4)
Mild	34 (11.7)	25 (8.8)
Moderate	57 (19.6)	22 (7.7)
Severe	21 (7.2)	6 (2.1)
Very severe	2 (.7)	0
Low back pain		
None	219 (75.3)	253 (88.8)
Mild	27 (9.3)	14 (4.9)
Moderate	34 (11.7)	15 (5.3)
Severe	9 (3.1)	1 (.4)
Very severe	2 (.7)	2 (.7)
Back pain		
None	199 (68.4)	244 (85.6)
Mild	36 (12.4)	23 (8.1)
Moderate	40 (13.7)	17 (6.0)
Severe	14 (4.8)	1 (.4)
Very severe	2 (.7)	0
Neck pain		
None	231 (79.4)	258 (90.5)
Mild	25 (8.6)	17 (6.0)
Moderate	20 (6.9)	7 (2.5)
Severe	14 (4.8)	3 (1.1)
Very severe	1 (.3)	0
Fever		
None	283 (97.3)	284 (99.6)
Mild	6 (2.1)	1 (.4)
Moderate	1 (.3)	0
Severe	1 (.3)	0
Verv severe		
Cough		
None	251 (86.3)	269 (94.4)
Mild	26 (8.9)	12 (4.2)
Moderate	10(34)	4 (1 4)
Severe	4 (1 4)	0
Very severe	- (1. -)	ů 0
Lack of appetite	v	v
None	272 (93 5)	279 (97 9)
1 tone	212 (75.5)	217 (71.3)

Table 3 (continued)

Parameter	3 months	6 months
Mild	7 (2.4)	3 (1.1)
Moderate	10 (3.4)	3 (1.1)
Severe	2 (.7)	0
Very severe	0	0
Dyspnea		
None	160 (55.0)	213 (74.7)
Mild	58 (19.9)	52 (18.2)
Moderate	58 (19.9)	17 (6.0)
Severe	14 (4.8)	3 (1.1)
Very severe	1 (.3)	0
Diarrhea		
None	273 (93.8)	277 (97.2)
Mild	6 (2.1)	7 (2.5)
Moderate	10 (3.4)	1 (.4)
Severe	2 (.7)	0
Very severe	0	0
Sore throat		
None	253 (86.9)	266 (93.7)
Mild	30 (10.3)	14 (4.9)
Moderate	5 (1.7)	4 (1.4)
Severe	2 (.7)	0
Very severe	1 (.3)	0
Headache		
None	220 (75.6)	258 (90.5)
Mild	31 (10.7)	22 (7.7)
Moderate	23 (7.9)	3 (1.1)
Severe	13 (4.5)	2 (.7)
Very severe	4 (1.4)	0
Dizziness		
None	236 (81.1)	269 (94.4)
Mild	26 (8.9)	14 (4.9)
Moderate	23 (7.9)	1 (.4)
Severe	5 (1.7)	1 (.4)
Very severe	1 (.3)	0
Absence of taste		
None	267 (91.8)	274 (96.1)
Mild	3 (1.0)	5 (1.8)
Moderate	12 (4.1)	4 (1.4)
Severe	7 (2.4)	1 (.4)
Very severe	2 (.7)	1 (.4)
Absence of smell		
None	254 (87.3)	270 (94.7)
Mild	9 (3.1)	6 (2.1)
Moderate	14 (4.8)	5 (1.8)
Severe	8 (2.7)	3 (1.1)
Very severe	6 (2.1)	1 (.4)
Sweat	. /	
None	213 (73.2)	236 (82.8)
Mild	20 (6.9)	19 (6.7)
Moderate	22 (7.6)	11 (3.9)

Table 3 (continued)				
Parameter	3 months	6 months		
Severe	20 (6.9)	19 (6.7)		
Very severe	16 (5.5)	0		
Hair loss				
None	157 (54.0)	228 (80.0)		
Mild	17 (5.8)	38 (13.3)		
Moderate	26 (8.9)	7 (2.5)		
Severe	25 (8.6)	10 (3.5)		
Very severe	66 (22.7)	2 (.7)		

Data are prevalence (%)

Discussion

We showed that approximately 3 in 5 patients had at least one symptom with approximately 2 in 5 patients had at least one rheumatic and musculoskeletal symptom and just over than half of the patients had at least one other COVID-19 symptom at 6 months. Fatigue (approximately 1 in 3), joint pain (approximately 1 in 5), and myalgia (approximately 1 in 7 patients) were the most frequent rheumatic and musculoskeletal symptoms. Joint pain and myalgia were mostly widespread. Dyspnea (approximately 1 in 4), hair loss (approximately 1 in 5), and sweat (approximately 1 in 6) were the most frequent other COVID-19 symptoms. Furthermore, female patients were more likely to have fatigue, myalgia, and joint pain at 6 months.

In the literature, some observational studies investigated the persistent symptoms in hospitalized COVID-19 survivors beyond 3 months [9–13]. Ghosn and colleagues analyzed 1137 hospitalized survivors (of which 288 admitted to the intensive care unit) and found that 60% had at least one symptom at 6 months, most frequently fatigue, dyspnea, joint pain, and muscle pain [11]. Peghin and colleagues evaluated 599 survivors (outpatients, n = 442; hospital ward unit, n = 134; the intensive care unit, n = 23) and reported that 52% of hospital ward unit survivors had post-COVID-19 syndrome at 6 months [12]. Our data at 6 months showed that 59.6% of survivors had at least one symptom, in consistent with previous two studies [11, 12]. Garrigues and colleagues assessed 120 survivors through telephone surveys after a mean of 111 days after admission either to the hospital ward unit (n=96) or to the intensive care unit (n=24) and documented that 54.2% had fatigue, 39.6% had dyspnea, 14.6% had cough, 14.6% had anosmia, and 9.4% had ageusia in the hospital ward unit group [9]. González-Hermosillo and colleagues assessed 130 survivors through telephone surveys and reported that 46.9% had fatigue, 43.8% joint pain, 42.3% had dyspnea, 36.2% muscle pain, 6.9% had anosmia, and 5.4% had ageusia at 6 months [10]. Fortini and colleagues assessed 59 survivors after a median

su	Fatigue 3m	41 21 28 8 2
oto	Fatigue 6m	68 21 7 3 1
hml	Myalgia_3m	59 12 19 8 2
al S	Myalgia_6m	85 85 2
elet	Joint pain_3m	61 19 7 1
oska	Joint pain_6m	81 81 8 2
culo	Low back pain_3m	75 12 3 1
Aus	Low back pain_6m	88 5 11
c/ N	Back pain_3m	68 12 14 5 1
nati	Back pain_6m	85 86 1
eun	Neck pain_3m	79 8 7 5 1
Rh	Neck pain_6m	90 6 3 1
	Fever_3m	96 2.11
	Fever_6m	99
	Cough_3m	86 9 4 1
	Cough_6m	95 4 1
	Lack of appetite_3m	94 21 3 1
	Lack of appetite_6m	98 11
	Dyspnea_3m	55 20 20 4 1
s	Dyspnea_6m	75 18 6 1
tom	Diarrhea_3m	94 21 3 1
mp1	Diarrhea_6m	97 24
Sy	Sore throat_3m	86 2 11
-19	Sore throat_6m	93 5 2
9	Headache_3m	75 11 8 4 2
0	Headache_6m	90 90 2 1
er-(Dizziness_3m	81 7 2 1
Oth	Dizziness_6m	94 4 11
	Absence of taste_3m	92 92 1
	Absence of taste_6m	95 2111
	Absence of smell_3m	87 87 3 2
	Absence of smell_6m	94 21211
	Sweat_3m	73 73 7 5
	Sweat_6m	83 7 4 6
	Hair loss_3m	54 6 9 8 23
	Hair loss_6m	80 13 3 3 1
		None Mild Moderate Very severe

Fig. 1 Severity of rheumatic/musculoskeletal symptoms and other COVID-19 symptoms at 3 and 6 months. Data are percentage

of 123 days after discharge from the hospital ward unit and observed that 42.4% had fatigue, 37.3% had dyspnea, 16.9% had ageusia, 15.2% had anosmia, 11.9% had cough, 8.5% joint pain, and 8.5% myalgia [13]. In those earlier investigations, generally fatigue and dyspnea were the most frequent symptoms. In our study, fatigue (approximately 1 in 3) and dyspnea (approximately 1 in 4) were the two most common symptoms as well. Also, we showed that joint pain and myalgia were each observed in approximately one-sixth of patients, and back pain, low back pain, and neck pain were each observed in approximately one-tenth of patients. With regard to the other COVID-19 symptoms, hair loss and sweat were each observed in approximately one-fifth of patients with less frequently, headache, sore throat, dizziness, cough, absence of smell, lack of appetite, absence of taste, diarrhea, lack of appetite, and fever. Considering the broad range of symptoms, multidisciplinary teams involving rheumatologists should provide a care for COVID-19 survivors.

In a study evaluating severe fatigue on 239 COVID-19 patients confirmed by PCR/CT, it was shown that severe fatigue lasted 12–23 weeks after the initial symptoms of the disease. It has been stated that the prevalence of long-term persistence of severe fatigue is high [17]. In another study, excessive fatigue persisted in 26 (33.3%) of 78 patients at 3-month follow-up and in 9 (39.1%) of 23 patients at 6-month follow-up [18]. In our study, severe/very severe fatigue was present in 10.7% of patients at 3 months and 3.2% at 6 months.

Table 4Location of myalgia and arthralgia symptoms at 3 and6 months

Parameter	3 months	6 months
Myalgia		
Present	118 (100.0)	43 (100.0)
Widespread	74 (62.7)	30 (69.8)
Regional	44 (37.3)	13 (30.2)
Shoulder girdle	6 (13.6)	2 (15.4)
Arm	13 (29.5)	2 (15.4)
Upper leg	0	0
Lower leg	25 (56.8)	9 (69.2)
Joint pain		
Present	114 (100.0)	53 (100.0)
Widespread	68 (59.6)	34 (64.2)
Regional	46 (40.4)	19 (35.8)
Shoulder	8 (17.0)	4 (21.1)
Elbow	1 (2.1)	0
Hand-wrist	3 (6.4)	1 (5.3)
Hip	8 (17.0)	2 (10.5)
Knee	15 (31.9)	6 (31.6)
Foot-ankle	11 (23.4)	6 (31.6)

Data are frequency (percentage)

We showed that female patients were more likely to have fatigue, myalgia, and joint pain at 6 months. This finding was consistent with two previous studies [11, 12], which showed an association of female sex and post-COVID symptoms. Our study provides information on association of female sex with rheumatic/musculoskeletal symptoms, therefore, extends the results of previous studies.

In the recent review, possible mechanisms that predominantly contribute to post-acute COVID-19 symptoms were listed as cellular invasion by SARS-COV-2, inflammatory and the immune response, and sequelae of post-critical illness [19]. Also, transforming growth factor beta (TGF- β) overexpression causing a prolonged state of immunosuppression and fibrosis was proposed as a unifying hypothesis mechanism for persistent post-COVID syndrome [20]. However, further research is warranted to elaborate the pathophysiologic mechanisms of wide spectrum of manifestations including rheumatologic/musculoskeletal involvement of post-COVID syndrome.

Limitations and strengths

We must acknowledge the limitations of our study. As we only included COVID-19 survivors who had been treated in hospital ward unit, our results are not generalizable to outpatient survivors or to those who admitted to intensive care unit. Also, it was an uncontrolled cohort study, the results would have been better interpreted if a comparative group who were hospitalized for other reasons than COVID-19 could exist. Furthermore, some factors such as medicines patients used might contribute to developing symptoms; however, the study did not investigate these factors. On the other hand, our study has several strengths. It was a prospective study with a relatively long-term follow-up period.

Conclusion

Approximately 3 in 5 patients had at least one symptom with ≈ 2 in 5 patients had at least one rheumatic and musculoskeletal symptom and just over than half of the patients had at least one other COVID-19 symptom at 6 months. Fatigue, joint pain, and myalgia were the most frequent rheumatic and musculoskeletal symptoms. Joint pain and myalgia were mostly widespread. Dyspnea, hair loss, and sweat were the most frequent other COVID-19 symptoms. Furthermore, female patients were more likely

 Table 5
 Association between the presence of symptoms (i.e., fatigue, myalgia, joint pain) and age, sex, BMI, and duration of hospital stay at 6 months

Parameter	Fatigue		Myalgia		Joint pain	
	Crude OR	Adjusted OR	Crude OR	Adjusted OR	Crude OR	Adjusted OR
Age	1.02 (0.99–1.04)	1.02 (0.99–1.04)	1.03 (0.99–1.06)	1.02 (0.99–1.05)	1.05 (1.02–1.08)	1.04 (1.01–1.08)
	<i>P</i> : 0.104	<i>P</i> : 0.186	<i>P</i> : 0.081	<i>P</i> : 0.223	<i>P</i> : 0.001	<i>P</i> : 0.004
Female sex	2.14 (1.28–3.55)	1.99 (1.18–3.34)	3.06 (1.56–5.99)	3.00 (1.51–5.98)	3.47 (1.86–6.47)	3.39 (1.78–6.50)
	<i>P</i> : 0.003	<i>P</i> : 0.010	<i>P</i> : 0.001	<i>P</i> : 0.002	<i>P</i> :<0.001	<i>P</i> : < 0.001
BMI	1.06 (1.01–1.12)	1.05 (0.99–1.11)	1.04 (0.97–1.11)	1.02 (0.95–1.08)	1.04 (0.98–1.11)	1.02 (0.96–1.08)
	<i>P</i> : 0.035	<i>P</i> : 0.109	<i>P</i> : 0.277	<i>P</i> : 0.616	<i>P</i> : 0.161	<i>P</i> : 0.584
Duration of	1.00 (0.94–1.07)	0.99 (0.93–1.07)	1.05 (0.97–1.13)	1.04 (0.96–1.13)	1.05 (0.98–1.13)	1.03 (0.94–1.12)
hospital stay	<i>P</i> : 0.986	<i>P</i> : 0.87	<i>P</i> : 0.219	<i>P</i> : 0.325	<i>P</i> : 0.217	<i>P</i> : 0.566

We performed generalized estimating equations by a selection of binary logistic regression models. In adjusted analyses of age, sex, BMI, and duration of hospital stay, the remaining 3 parameters were controlled in the models. *BMI* body mass index, *OR* Odds ratio

to have fatigue, myalgia, and joint pain at 6 months. This information guide rheumatologists to understand the nature and features of persistent rheumatic and musculoskeletal symptoms in hospitalized COVID-19 survivors and may translate into improved management of such individuals with persistent symptoms who had recovered from

Declarations

acute COVID-19 infection.

Ethical approval We obtained an ethical approval by the Ethic Committee of the Gülhane Scientific Researches, University of Health Sciences (decision no: 2021/187).

Conflict of interest We declare that we have no conflicts of interest regarding the submitted manuscript. Outside of the submitted manuscript SK has received congress travel, accommodation, and participation fee support (12th Anatolian Rheumatology Days) from AbbVie.

References

- Akintayo RO, Bahiri R, El Miedany Y, Olaosebikan H, Kalla AA, Adebajo AO, Migowa AN, Slimani S, Koussougbo OD, Kawther BA, Akpabio AA, Ghozlani I, Dey D, Hassan WA, Govind N, Makan K, Mohamed A, Genga EK, Ghassem MKA, Mortada M, Hamdi W, Wabi MO, Tikly M, Ngandeu-Singwe M, Scott C (2020) African League Against Rheumatism (AFLAR) preliminary recommendations on the management of rheumatic diseases during the COVID-19 pandemic. Clin Rheumatol. https://doi.org/10.1007/s10067-020-05355-2
- Li J, Ringold S, Curtis JR, Michaud K, Johansson T, Yun H, Yazdany J, Schmajuk G (2021) Effects of the SARS-CoV-2 global pandemic on U.S. rheumatology outpatient care delivery and use of telemedicine: an analysis of data from the RISE registry. Rheumatol Int. https://doi.org/10.1007/s00296-021-04960-x
- Kardeş S, Erdem A, Gürdal H (2021) Public interest in musculoskeletal symptoms and disorders during the COVID-19 pandemic: Infodemiology study. Z Rheumatol. https://doi.org/10. 1007/s00393-021-00989-2
- Günendi Z, Yurdakul FG, Bodur H, Cengiz AK, Uçar Ü, Çay HF, Şen N, Keskin Y, Gürer G, Melikoğlu MA, Altıntaş D, Deveci H, Baykul M, Nas K, Çevik R, Karahan AY, Toprak M, Ketenci S, Nayimoğlu M, Sezer İ, Demir AN, Ecesoy H, Duruöz MT, Yurdakul OV, Sarıfakıoğlu AB, Ataman Ş (2021) The impact of COVID-19 on familial Mediterranean fever: a nationwide study. Rheumatol Int 41:1447–1455. https://doi.org/ 10.1007/s00296-021-04892-6
- Amengual O, Atsumi T (2021) COVID-19 pandemic in Japan. Rheumatol Int 41(1):1-5. https://doi.org/10.1007/ s00296-020-04744-9
- 6. https://covid19.who.int/. Accessed on 5th of July 2021
- Lerner AM, Robinson DA, Yang L, Williams CF, Newman LM, Breen JJ, Eisinger RW, Schneider JS, Adimora AA, Erbelding EJ (2021) Toward understanding COVID-19 recovery: National Institutes of Health Workshop on Postacute COVID-19. Ann Intern Med M21–1043.https://doi.org/10.7326/M21-1043
- https://meshb.nlm.nih.gov/record/ui?ui=C000711409/. Accessed on 12th of July 2021
- Garrigues E, Janvier P, Kherabi Y, Le Bot A, Hamon A, Gouze H, Doucet L, Berkani S, Oliosi E, Mallart E, Corre F, Zarrouk V,

Moyer JD, Galy A, Honsel V, Fantin B, Nguyen Y (2020) Postdischarge persistent symptoms and health-related quality of life after hospitalization for COVID-19. J Infect 81:e4–e6. https://doi. org/10.1016/j.jinf.2020.08.029

- González-Hermosillo JA, Martínez-López JP, Carrillo-Lampón SA, Ruiz-Ojeda D, Herrera-Ramírez S, Amezcua-Guerra LM, Martínez-Alvarado MDR (2021) Post-Acute COVID-19 symptoms, a potential link with myalgic encephalomyelitis/chronic fatigue syndrome: a 6-month survey in a Mexican Cohort. Brain Sci 11(6):760. https://doi.org/10.3390/brainsci11060760
- Ghosn J, Piroth L, Epaulard O, Le Turnier P, Mentré F, Bachelet D, Laouénan C; French COVID cohort study and investigators groups (2021) Persistent COVID-19 symptoms are highly prevalent 6 months after hospitalization: results from a large prospective cohort. Clin Microbiol Infect :S1198–743X(21)00147–6. https://doi.org/10.1016/j.cmi.2021.03.012
- Peghin M, Palese A, Venturini M, De Martino M, Gerussi V, Graziano E, Bontempo G, Marrella F, Tommasini A, Fabris M, Curcio F, Isola M, Tascini C (2021) Post-COVID-19 symptoms 6 months after acute infection among hospitalized and non-hospitalized patients. Clin Microbiol Infect:S1198–743X(21)00281–0. https://doi.org/10.1016/j.cmi.2021.05.033.
- Fortini A, Torrigiani A, Sbaragli S, Lo Forte A, Crociani A, Cecchini P, InnocentiBruni G, Faraone A (2021) COVID-19: persistence of symptoms and lung alterations after 3–6 months from hospital discharge. Infection. https://doi.org/10.1007/ s15010-021-01638-1
- Karaarslan F, DemircioğluGüneri F, Kardeş S (2021) Postdischarge rheumatic and musculoskeletal symptoms following hospitalization for COVID-19: prospective follow-up by phone interviews. Rheumatol Int 41(7):1263–1271. https://doi.org/10. 1007/s00296-021-04882-8
- 15 Vandenbroucke JP, von Elm E, Altman DG, Gøtzsche PC, Mulrow CD, Pocock SJ, Poole C, Schlesselman JJ, Egger M, STROBE Initiative (2007) Strengthening the Reporting of Observational Studies in Epidemiology (STROBE): explanation and elaboration. Epidemiology 18(6):805–835. https://doi.org/10.1097/EDE.0b013e3181577511
- Misra DP, Zimba O, Gasparyan AY (2021) Statistical data presentation: a primer for rheumatology researchers. Rheumatol Int 41(1):43–55. https://doi.org/10.1007/s00296-020-04740-z
- Van Herck M, Goertz YMJ, Houben-Wilke S, Machado FVC, Meys R (2021) Delbressine JM et al Severe fatigue in long COVID: follow-up study in members of online long COVID support groups. J Med Internet Res. https://doi.org/10.2196/30274
- Sykes DL, Holdsworh L, Jawad N, Gunasekera P, Morice AH (2021) Crooks MG Post-COVID-19 symptom burden: what is long-COVID and how should we manage it? Lung 199(2):113– 119. https://doi.org/10.1007/s00408-021-00423-z
- Nalbandian A, Sehgal K, Gupta A, Madhavan MV, McGroder C, Stevens JS, Cook JR, Nordvig AS, Shalev D, Sehrawat TS, Ahluwalia N, Bikdeli B, Dietz D, Der-Nigoghossian C, Liyanage-Don N, Rosner GF, Bernstein EJ, Mohan S, Beckley AA, Seres DS, Choueiri TK, Uriel N, Ausiello JC, Accili D, Freedberg DE, Baldwin M, Schwartz A, Brodie D, Garcia CK, Elkind MSV, Connors JM, Bilezikian JP, Landry DW, Wan EY (2021) Post-acute COVID-19 syndrome. Nat Med 27(4):601–615. https://doi.org/ 10.1038/s41591-021-01283-z
- Oronsky B, Larson C, Hammond TC, Oronsky A, Kesari S, Lybeck M, Reid TR (2021) A review of persistent post-COVID syndrome (PPCS). Clin Rev Allergy Immunol. https://doi.org/10. 1007/s12016-021-08848-3

Publisher's note Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.