# Assessment of short- and long-term functionality and quality of life in patients with post-acute COVID-19 syndrome

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#### Abstract.

**BACKGROUND:** Although the number of new cases of coronavirus 2019 (COVID-19) has been drastically reduced worldwide, patients who demonstrate long-term symptoms need more attention from health systems, as these symptoms can negatively affect functionality and quality of life.

**OBJECTIVE:** To evaluate muscle function and quality of life at 3, 6, 9 and 12 months in patients with post-acute COVID-19 syndrome and to assess their associations with general fatigue and lung function.

**METHODS:** This observational and longitudinal study evaluated patients with post-acute COVID-19 syndrome. Participants were subjected to the following evaluations: Short Form-36; handgrip strength; Functional Assessment of Chronic Illness Therapy-Fatigue scale; and spirometry.

**RESULTS:** Among the 350 participants who were evaluated in the third month, 74.6%, 61.4% and 45.4% reported general fatigue, dyspnoea and cough, respectively. In the comparisons between the third month and the sixth month, there were significant increases in Functional Assessment of Chronic Illness Therapy-Fatigue scale, pulmonary function and several Short Form-36 domains. In the comparisons between the sixth month and the ninth month, there was a significant increase only in the social functioning domain of the Short Form-36. In the comparisons between the ninth month and the twelfth month, there was an increase only in some Short Form-36 domains. Significant correlations were observed between the Short Form-36 domains with Functional Assessment of Chronic Illness Therapy-Fatigue scale, handgrip strength and pulmonary function.

**CONCLUSION:** In patients with post-acute COVID-19 syndrome, there was a progressive improvement in quality of life, general fatigue and pulmonary function during the 12 months of follow-up, with this improvement being more pronounced in the first 6 months. There was a relationship between functionality and quality of life in these patients.

Keywords: SARS-CoV-2, muscles, fatigue, respiratory function tests, rehabilitation

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#### 1. Introduction

According to the World Health Organization (WHO), viral diseases continue to emerge and represent a serious public health problem. More recently, we experienced the coronavirus 2019 (COVID-19) pandemic, caused by severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) [1]. Among the most severe clinical manifestations of acute disease are severe pneumonia, sepsis and septic shock. However, COVID-19 has much greater consequences than generating an acute disease and may cause long-term symptoms or even permanent sequelae that can significantly impact functionality and quality of life (QoL) [2–4].

Maximum forces exerted by humans are important in many aspects of life, and its measurements are often used not only in ergonomics but also in rehabilitation [5]. Applying force manually is the most common method of operating hand tools and moving objects [6]. Handgrip strength (HGS) is not only an indicator of the individual's performance, but it can also be used to reduce persistent diseases, as muscle difficulty is strongly linked to practical restrictions and physical disability [7]. In patients with post-acute COVID-19 syndrome (PACS), the worse their HGS and general fatigue is, the more impaired their physical function [8]. In addition, a history of previous hospitalization for COVID-19 can cause subsequent worsening of physical function. In this sense, the evaluation of functional limitations including muscular dysfunctions of the hands is necessary in COVID-19 survivors to estimate the long-term burden of the disease [4].

Although it is believed that the vast majority of patients recover from their functional limitations, muscle involvement in COVID-19 can be problematic, producing myalgia and muscle weakness with the potential to further deteriorate QoL [9]. Assessing non-hospitalized adults with PACS, a Brazilian study showed worse functionality and postural balance than controls, which was associated with lower HGS, general fatigue and poor QoL [10]. Still in the Brazilian population, another study showed that the reduced functional exercise capacity in women with PACS was strongly explained by the deterioration of muscle strength and pulmonary function [11]. In a large cohort of hospitalized patients in Brazil, the authors showed that general fatigue and arthralgia were among the most prevalent symptoms in PACS [12]. However, these studies did not assess the longitudinal changes in functionality and QoL of these patients, which are important to allow the design of new rehabilitation strategies.

There is evidence that SARS-CoV-2 can impair physical function, deteriorate lung function and significantly reduce QoL [13,14]. Several studies have evaluated the QoL of patients with PACS in the short- and mediumterm, although they have not focused on the long-term changes and the impact of abnormalities on functionality and QoL [15-18]. Since muscles and lungs are often affected in the acute phase of COVID-19 [4], understanding how PACS-affected individuals react to physical, muscular and respiratory dysfunction over time will allow tracing the development profile of PACS for these abnormalities. With this, it will be possible to identify how such limitations will affect the functionality and QoL of these subjects and to promote monitoring to prevent the development of significant changes in physical, muscle and respiratory functions. The objectives of the present study were to evaluate muscle function and QoL at 3, 6, 9 and 12 months in patients with PACS and to assess their associations with general fatigue and pulmonary function.

### 2. Materials and methods

#### 2.1. Participants

From October 2020 to July 2022, an observational and longitudinal study was conducted with 350 survivors of COVID-19 (of 315 eligible) aged  $\ge$  18 years treated at the Piquet Carneiro Policlinic, State University of Rio de Janeiro, Rio de Janeiro, Brazil. These patients were cared for in triage tents or were admitted during the acute phase of COVID-19 at our institution, and were followed up by telemonitoring for the first 12 weeks; after this period, they were recontacted about their clinical manifestations for face-toface evaluation. The study included patients with persistent symptoms or development of sequelae at least 12 weeks after the onset of acute symptoms of COVID-19, characterizing PACS [19]. Patients without a previous diagnosis of COVID-19 confirmed by reversetranscription polymerase chain reaction (RT-PCR) and those who were unable to perform acceptable manoeuvres in spirometry were excluded.

In addition to the assessment in the third month (T1), these participants were subsequently evaluated in case of persistent symptoms (diagnosis of PACS) in the sixth month (T2), ninth month (T3) and twelfth month (T4). Patient appointments were confirmed the day before with a phone call.

The project was approved by the Brazilian National Research Ethics Committee (number CAAE- 30135320.0.0000.5259) and was conducted in accordance with the principles of the Declaration of Helsinki. All participants signed an informed consent form.

#### 2.2. Instruments and measurements

QoL was assessed using the Short Form-36 (SF-36), which is a multidimensional instrument with questions grouped into two major components (physical and mental) that allow measurement of the health state transition. Scores can vary between 0 and 100, and higher values indicate better QoL. The 36 questions that make up the SF-36 are grouped into 8 domains, as follows: physical functioning, physical role limitations, bodily pain, general health perceptions, vitality, social functioning, emotional role limitations and mental health [20]. The Portuguese translation of the SF-36 is adequate, given the completeness of responses and its internal consistency [21]. In Portuguese version, the Cronbach's alpha was 0.82 for the physical and 0.87 for the mental dimension [22].

Handgrip strength (HGS) was assessed using a digital handheld dynamometer (SH5001, Saehan Corporation, Yangsan, Korea). Participants were instructed to sit with neutral shoulders, elbows bent at 90° and neutral forearms and wrists. Then, they were instructed to perform a maximum contraction for 3 seconds in each test. Three measurements with intervals of 30 seconds were performed. The highest value was considered for analysis [23].

General fatigue was assessed using the Functional Assessment of Chronic Illness Therapy-Fatigue (FACIT-F) scale. It is a specific fatigue assessment scale that has good representativeness of the individual's condition and is considered an easy-to-apply instrument. The FACIT-F scale is a 13-item scale that aims to evaluate the overall fatigue of the individual correlated with his or her daily activities. These items provide a score ranging from 0 to 52. The lower the score is, the lower the fatigue index [24,25]. The FACIT-F scale has been used in clinical populations and is valid for use in the Portuguese language. Internal consistency reliability is good, with Cronbach's alpha of the 0.93 [26].

Pulmonary function was assessed using a volume spirometer (Vitatrace VT 130 SL, Codax Ltda, Rio de Janeiro, Brazil) following the recommendation of the American Thoracic Society/European Respiratory Society [27]. The following variables were evaluated: forced vital capacity (FVC), forced expiratory volume in one second (FEV<sub>1</sub>) and the ratio  $FEV_1/FVC$ . The reference values were obtained using national prediction equations [28], and the results were expressed as values in percentages of the predicted value.

#### 2.3. Statistical analysis

Statistical analysis was performed using IBM SPSS Statistics version 26.0 software (IBM Corp., Armonk, NY, USA). To verify the homogeneity of the sample, the Shapiro-Wilk test was used. The results are expressed as the median (interquartile range) for numerical data and as the frequency and percentage for categorical data. In the longitudinal analysis, the comparison of physical function, pulmonary function and QoL parameters over time (T1, T2, T3 and T4) was performed by the Wilcoxon signed-rank test. The correlation between the absolute deltas of the SF-36 domains and the parameters of physical, muscular and respiratory function was analysed using the Spearman correlation coefficient. The significance level adopted was 5%.

To provide insight into the clinical significance of the results, we calculated the effect size in Jeffreys's Amazing Statistics Program version 0.10.2 (https://jaspstats.org). To provide context for interpreting the null findings, a *post hoc* power analysis was performed using GPower 3.1.1 software based on the differences for the three evaluation moments (T1-T2, T2-T3 and T3-T4).

# 3. Results

Among the 365 patients who were evaluated for inclusion in the study, 15 were excluded for the following reasons: diagnosis without confirmation by RT-PCR (n = 10) and failure to perform spirometry (n = 5). In the sample initially evaluated at T1, 214 (61.1%)were women, while the median age and body mass index was 55.5 (45-64) years and 28.7 (25-33) kg/m<sup>2</sup>, respectively. The most frequent comorbidities were hypertension, diabetes, heart disease, asthma and chronic obstructive pulmonary disease, which were observed in 168 (48%), 88 (25.1%), 30 (8.6%), 30 (8.6%) and 22 (6.3%) participants, respectively. One hundred fortysix (41.7%) participants reported previous hospitalization, while 76 (21.7%) were admitted to the intensive care unit. The median time between the onset of acute COVID-19 symptoms and the assessments was 96 (92-99) days.

In the sixth month after COVID-19, 111 participants returned due to the permanence of symptoms, 29 (26.1%) with general fatigue, 26 (23.4%) with dyspnoea and 21 (18.9%) with cough. Thus, 31.7% of the initial sample still met the criteria for the diagnosis of PACS. In the comparisons between T1 and T2 (Table 1), there were significant increases in FACIT-F (p = 0.0001),

Variable	Compariand 6th	Comparison between the 3rd and 6th month $(n = 111)$		compar and 9t	comparison between the out and 9th month $(n = 46)$		and 12	and 12th month $(n = 32)$	
1	T1	T2	<i>P</i> -value	T2	T3	<i>P</i> -value	T3	T4	P-value
Physical function									
HGS (kgf)	25.9 (18.1-33)	25 (19–35.4)	0.081	28 (19-35.3)	26 (20–34)	0.33	27 (20-35.8)	27.5 (18.5–38.8)	0.99
FACIT-F (points)	28 (20-36)	32 (24-41.2)	0.0001	33 (25–42)	35.5 (21.8-43.3)	0.56	33.8 (21.3-43.8)	39 (24.8–42)	0.36
Spirometry									
FVC (% predicted)	80.2 (69.6–92.6)	84.2 (75.9–91.7)	0.001	83 (74.5–90.1)	85.3 (74.9–92.9)	0.17	86.8 (76.8–94.3)	84.4 (74.1–94.8)	0.55
FEV <sub>1</sub> (% predicted)	82.8 (70.7–93.8)	84.9 (75.7–93.1)	0.006	83.2 (72.3–91.1)	83.2 (76.7–94.7)	0.30	82.2 (77.2–95)	83.2 (72.2–92.4)	0.13
$FEV_1/FVC(\%)$	80.8 (75.1-85.3)	81.5 (74.6-85.3)	0.40	81.4 (74.9-85.9)	80.3 (76.7-85.3)	0.51	78.8 (76.1-84.1)	79.1 (74.2-84.3)	0.077
SF-36 (points)									
Physical functioning	35 (17.5–50)	40 (25–60)	0.003	40 (23.8-56.3)	45 (28.8–66.3)	0.14	42.5 (25-63.8)	45 (25–65)	0.66
Physical role limitations	0(0-25)	25 (0-50)	0.001	25 (0-25)	25 (0-50)	0.31	30 (0-68.8)	35 (0-75)	0.42
Bodily pain	40 (20-62)	41 (31-63)	0.015	41 (31–72.5)	41 (31-54.5)	0.47	41 (31-59.3)	52 (31-54)	0.060
General health perceptions	40 (30-57)	47 (30–62)	0.29	50 (32-67)	45 (32-58.3)	0.21	40 (30.5-60.8)	52 (40-69.5)	0.003
Vitality	35 (25–50)	45 (30-60)	0.0001	45 (25–70)	50 (30-65)	0.68	55 (22.5–65)	62.5 (35-70)	0.037
Social functioning	50 (25-62.5)	50 (25–75)	0.001	50 (25-65.6)	56.3 (37.5-75)	0.019	56.3 (37.5–75)	75 (50–100)	0.022
Emotional role limitations	0 (0-66.7)	33.3 (0-83.4)	0.028	33.3 (0-66.7)	33.3 (0-66.7)	0.58	33.3 (0-33.3)	33.3 (0-91.7)	0.10
Mental health	56 (40–76)	60(40-80)	0.020	62 (40-81)	64 (43–77)	0.72	66 (41–79)	74 (49–80)	0.048

Table 1

J.E. de Azevedo Vieira et al. / Functionality in post COVID-19 syndrome

FVC (p = 0.001) and FEV<sub>1</sub> (p = 0.006), with no significant improvement in HGS. Regarding the QoL, there was an increase in the following SF-36 domains: physical functioning (p = 0.003), physical role limitations (p = 0.001), bodily pain (p = 0.015), vitality (p =0.0001), social functioning (p = 0.001), emotional role limitations (p = 0.028) and mental health (p = 0.020).

In the ninth month after COVID-19, 46 participants returned due to continuing symptoms, with 9 (19.6%) experiencing general fatigue, 8 (17.4%) experiencing dyspnoea and 4 (8.7%) experiencing cough. In the comparisons between T2 and T3 (Table 1), no significant increase was observed for HGS, FACIT-F or pulmonary function. Regarding the QoL, there was a significant increase only in the social functioning domain of the SF-36 (p = 0.019).

In the twelfth month after COVID-19, 32 participants returned due to the continuing symptoms, 6 (18.8%) with general fatigue, 5 (15.6%) with dyspnoea and 2 (6.2%) with cough. In the comparisons between T2 and T3 (Table 1), no significant increase was observed for HGS, FACIT-F or pulmonary function. Regarding the QoL, there was an increase in the following SF-36 domains: general health perceptions (p = 0.003), vitality (p = 0.037), social functioning (p = 0.022) and mental health (p = 0.048).

Additionally, we evaluated the associations between changes in T1, T2, T3 and T4 (Table 2). When the absolute deltas between T1 and T2 were evaluated, we observed significant correlations between the FACIT-F and all domains of the SF-36 (except bodily pain) (Fig. 1). When the absolute deltas between T1 and T3 were evaluated, we observed significant correlations between FACIT-F and physical functioning, general health perceptions, vitality and mental health domains of SF-36. Significant correlations were observed between HGS and physical functioning, vitality and social functioning domains of SF-36. Additionally, significant correlations were observed between the FVC and the SF-36 social functioning domain. Finally, when the absolute deltas between T1 and T4 were evaluated, we observed significant correlations between the FACIT-F and the SF-36 physical functioning, physical role limitations, bodily pain, vitality and mental health domains. Significant correlations were observed between the HGS and the physical functioning, physical role limitations, general health perceptions, vitality and social functioning domains of the SF-36. Additionally, significant correlations were observed between the FVC and the SF-36 physical functioning and physical role limitations domains.

Based on an a priori type I error  $\alpha = 0.05$  (twotailed), the power analysis showed that significant effects were detected in the comparisons for the three evaluation moments (T1-T2, T2-T3 and T3-T4), as follows: HGS: effect size = range of 0.46 to 0.67, range of power = 50% to 72%; FACIT-F: range of 0.68 to 1.35, range of power = 63% to 98%; FVC: range of 0.72 to 1.30, range of power = 66% to 96%; FEV<sub>1</sub>: range of 0.70 to 1.22, range of power = 65% to 95%; FEV<sub>1</sub>/FVC: range of 0.69 to 0.72, range of power = 64% to 74%; physical functioning: range of 0.48 to 1.18, range of power = 52% to 92%; physical role limitations: range of 0.70 to 1.30, range of power = 65% to 96%; bodily pain: range of 0.71 to 1.17, range of power = 66% to 91%; general health perceptions: range of 0.68 to 1.18, range of power = 64% to 92%; vitality: range of 0.50 to 1.35, range of power = 56% to 98%; social functioning: range of 0.16 to 1.30, range of power = 91% to 96%; emotional role limitations: range of 0.73 to 1.22, range of power = 66% to 93%; and mental health: range of 0.53 to 1.23, range of power = 60% to 94%.

# 4. Discussion

There is growing interest in long-term sequelae after recovery from the acute phase of COVID-19. In the 1-year follow-up of a sample of patients with PACS, the main finding of the present study was the improvement of QoL, general fatigue and pulmonary function in the first 6 months of follow-up without improvement in peripheral muscle strength (HGS). In these patients, the improvement after 6 months is slower, occurring especially in the components that involve QoL (both physical and mental). In addition, there was an association between general fatigue and the domains of the physical and mental components of QoL in the first 6 months after acute infection. After this period, there was an association of QoL with muscle function and lung function in the subsequent months until the end of follow-up at 1 year. To our knowledge, this is the first study that evaluated the associations of QoL with functionality in the follow-up of patients with PACS in the course of 1 year.

In patients with PACS, the persistence of symptoms negatively affects QoL, making it necessary to monitor symptoms in the medium and long term. In our sample, all SF-36 domains were below the expected values for the Brazilian population from the third to the twelfth month after acute infection of COVID-19 [29]. More-

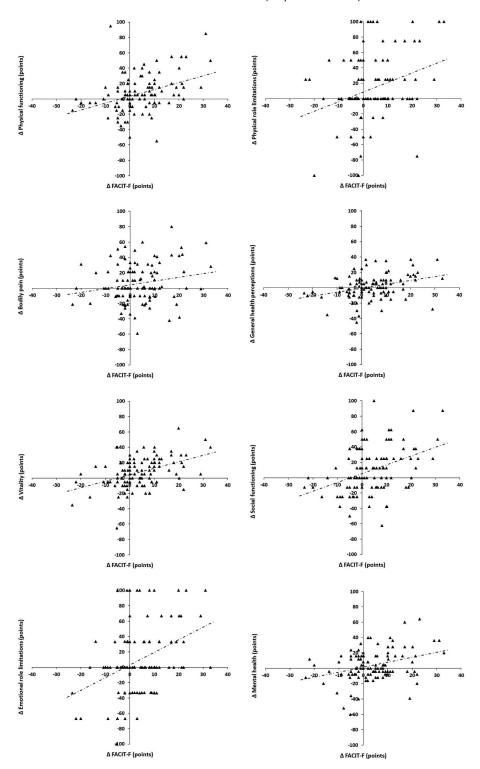


Fig. 1. Relationships of the Functional Assessment of Chronic Fatigue Illness Therapy (FACIT-F) with the domains of the Short Form-36 (SF-36) for 3rd to 6th month absolute deltas: physical functioning ( $r_s = 0.388$ , p < 0.0001) (a), physical role limitations ( $r_s = 0.282$ , p = 0.003) (b), bodily pain ( $r_s = 0.168$ , p < 0.081) (c), general health perceptions ( $r_s = 0.352$ , p = 0.0002) (d), vitality ( $r_s = 0.427$ , p < 0.0001) (e), social functioning ( $r_s = 0.379$ , p < 0.0001) (f), emotional role limitations ( $r_s = 0.307$ , p = 0.001) (g), and mental health ( $r_s = 0.271$ , p = 0.004) (h).

	SF-36								
	Physical functioning	Physical role limitations	Bodily pain	General health perceptions	Vitality	Social functioning	Emotional role limitations	Mental health	
3rd to 6th mo	onth								
HGS	0.069	-0.019	0.029	0.030	0.030	0.111	0.047	0.031	
FACIT-F	<b>0.388</b> §	<b>0.282</b> <sup>‡</sup>	0.168	0.351§	<b>0.427</b> §	<b>0.379</b> §	<b>0.307</b> <sup>‡</sup>	<b>0.271</b> <sup>‡</sup>	
FVC	0.127	0.125	-0.129	-0.031	0.073	0.032	0.090	-0.064	
$FEV_1$	0.134	0.172	-0.059	0.023	0.066	0.042	0.043	-0.123	
FEV <sub>1</sub> /FVC	0.098	0.078	0.039	0.101	0.120	0.147	0.016	-0.021	
3rd to 9th mc	onth								
HGS	<b>0.394</b> <sup>†</sup>	0.090	0.181	-0.003	0.368*	0.298*	0.129	0.077	
FACIT-F	<b>0.440</b> ‡	0.110	0.112	0.375*	<b>0.600</b> §	0.280	0.132	<b>0.471</b> ‡	
FVC	-0.056	0.192	0.088	0.195	0.055	0.318*	0.197	0.096	
$FEV_1$	0.022	0.274	0.082	0.205	0.147	0.211	0.216	0.018	
FEV <sub>1</sub> /FVC	0.082	0.034	0.047	-0.011	-0.018	-0.150	-0.065	-0.206	
3rd to 12th m	onth								
HGS	0.459 <sup>†</sup>	0.386*	0.214	0.366*	0.403*	<b>0.491</b> <sup>†</sup>	0.139	0.214	
FACIT-F	<b>0.615</b> §	0.361*	0.453*	0.224	<b>0.568</b> §	0.128	0.247	0.455*	
FVC	0.462*	0.420*	0.165	0.063	0.169	0.326	0.035	-0.014	
$FEV_1$	0.579‡	0.348	0.167	0.177	0.198	0.253	0.121	0.000	
FEV <sub>1</sub> /FVC	0.233	-0.065	0.016	0.091	0.096	-0.135	0.027	0.002	

 Table 2

 Spearman's correlation coefficients between quality of life, physical function and lung function

The values in bold refer to significant differences. \*p-value < 0.05; †p-value < 0.01; ‡p-value < 0.005; %p-value < 0.001. List of abbreviations: SF-36 – Short Form-36, HGS – handgrip strength, FACIT-F – Functional Assessment of Chronic Illness Therapy-Fatigue, FVC – forced vital capacity; FEV<sub>1</sub> – forced expiratory volume in one second.

over, there was an increase in almost all dimensions of the SF-36 over the 12 months, with values higher than the minimal clinically important difference (MCID) of 5 points described in the SF-36 [30]. Contrary to our results, O'Brien et al. [31] did not notice significant changes in the mean scores of the eight SF-36 domains during the 1-year period, although these authors noted values below the normative data of the population and evaluated only patients with previous hospitalizations. Although our sample was not included in any physical reconditioning program, the improvement in functionality and OoL, especially in the first 6 months, suggests that therapeutic strategies may be more beneficial in this initial period of PACS, especially in those who have a significant deterioration of physical capacity and QoL [32].

General fatigue has been described as one of the main symptoms of PACS, occurring in 34.8% to 90% of cases depending on the evaluation tool and the time in which it is evaluated after the acute phase of the disease [14,33]. In fact, approximately three-quarters of our sample reported general fatigue in the third month, and the presence of this symptom progressively decreased during follow-up. Considering the median value of the FACIT-F scale, there was a variation from 28 to 39 points between the third and twelfth months; this variation is greater than the MCID described for the FACIT-F of 4 points [34], although still below the nor-

mative value of 40.1 points [35]. Throughout the 12month follow-up, we observed significant correlations between changes in the FACIT-F scale and the SF-36 domains, especially with the vitality domain, which best reflects the general fatigue in patients with PACS [36]. In agreement with our findings, Rass et al. [37] observed that impairments in the physical health component of the SF-36 were strongly related to general fatigue in a sample of patients with PACS. In fact, the significant load of general fatigue in patients with PACS is related to a high degree of difficulty in performing activities of daily living, deterioration of physical function and, therefore, worsening of QoL [38].

HGS evaluates the static forces compressing around the dynamometer and is a practical, fast and viable tool for the study of vitality and physical fitness [7]. HGS can be used as a reference to establish reaction time, and problems with HGS are indicators of difficulties in performing everyday tasks and worse performance at work [6]. In our sample, the median HGS value was 27.2 kgf in the third month, which is well below the value described in healthy Brazilian adults (38.0  $\pm$ 10.7 kgf) [39]. However, no MCID was observed in the longitudinal evaluation between the third and twelfth months [40], which suggests that the recovery of muscle function occurs slowly over the course of PACS. It is important to highlight the correlations between changes in HGS and changes in QoL (SF-36 domains), especially those related to the physical component. These correlations occurred for the absolute deltas from the third to the twelfth month, which suggests that, to some degree, muscle dysfunction negatively affects the QoL of patients with PACS.

Individuals with PACS should be carefully monitored for respiratory dysfunction as a sequel of SARS-CoV-2 acute lung injury [41]. In the present study, we observed a gradual improvement in pulmonary function throughout follow-up, mainly between 3 and 6 months. Following a cohort of patients with PACS for 1 year, Zhou et al. [42] observed mean values of pulmonary function higher than those of our sample, which can be explained at least in part by the predominance of nonsevere cases in their study. During the pandemic, it is important to evaluate the relationship between lung function and QoL in COVID-19 survivors. We observed a correlation between the physical functioning domain of SF-36 and pulmonary function, which is in agreement with the study by Bardakci et al. [16]. Notably, we evaluated the correlations between the absolute deltas of the QoL time intervals and the absolute deltas of the time intervals of the functionality variables. This provided a more reliable idea of the improvements over time, rather than the strictly cross-sectional analysis performed by most studies [16,41,42].

Our study has several limitations. First, we did not use a control group that could assist in the interpretation of our results. Also, the initial sample was not followed up during the 12 months, although we defined the persistence of symptoms in the subsequent months (diagnosis of PACS) as a follow-up criterion, and this definition is more compatible with the "real world". Third, because the results of the baseline data are unknown, our findings cannot be attributed exclusively to COVID-19. This is because previous musculoskeletal and cardiopulmonary diseases may have affected our results. Despite these limitations, our results may contribute to the design of studies focused on muscle and cardiopulmonary reconditioning, especially those focused on the first 6 months of PACS. This could contribute to a faster recovery of patients' physical capacity and consequently QoL, in addition to reducing public spending on social programs aimed at this population that may affect work activities.

# 5. Conclusions

Our results show that in patients with PACS, there is progressive improvement in QoL, general fatigue and pulmonary function during the 12 months of follow-up, and this improvement is more pronounced in the first 6 months after the acute phase of COVID-19. There is a relationship between general fatigue and QoL in the first 6 months and between functionality and QoL in the subsequent period until completing 1 year. Thus, we believe that assessment of muscle function and QoL can improve the understanding of the medium- and long-term consequences of COVID-19 and thus provide opportunities to apply tailored interventions in patients with PACS.

# **Ethical approval**

The project was approved by the Brazilian National Research Ethics Committee (number CAAE-30135320.0.0000.5259).

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## **Informed consent**

All participants signed an informed consent form.

## **Conflict of interest**

None of the authors have any conflict of interest to declare.

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#### **Author contributions**

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550