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Article

# Resistance Exercise Program Is Feasible and Effective in Improving Functional Strength in Post-Covid Survivors

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**Abstract:** Evidence suggests that COVID-19 infection can cause lasting health consequences. Multidisciplinary rehabilitation services have been recommended to reduce the sequelae. However, the effectiveness of physical exercise interventions remains insufficiently documented. The aim of this study was to develop and implement a specific and well-tolerated protocol-based, intervention to reduce muscle weakness in older adults impacted by COVID-19. Forty six older adults randomized into Intervention and Control Groups. Isometric and isokinetic strength assessments were conducted for selected muscle groups using JBA Staniak® torquemeter and Biodex System 3 dynamometer. Functional abilities were evaluated with the Time Up and Go test and Chair Stand Tests. Men in intervention group demonstrated a significant improvement in static conditions for knee flexors (KF), trunk extensors (TE) and trunk flexors (TF) and in dynamic conditions for knee extensors (KE). Women in intervention group showed significant improvement in static condition for EF, KF, TE and TF and in dynamic for a KE and for a KF. The interaction GROUP x TESTING SESSION was significant for Chair test (s) and Chair test (n). Our results demonstrate the effectiveness of a well-tolerated, protocol-based approach that can be used to diminish long lasting functional deficits in post-covid survivors.

**Keywords:** protocol; weakness; older adults; function; muscle

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## 1. Introduction

COVID-19 has affected more than 237 countries, and more than 517 million people have been infected across the globe, causing the deaths of more than 6 million people [1,2]. Vaccinations and the number of recovered individuals has weakened the virus however, new variants continue to emerge. The understanding of acute symptoms and complications related to COVID-19 has been at the center of attention amongst the scientific community; however, emergent evidence has also shown persistent or new symptoms in survivors of COVID-19 [3]. WHO European Regional data suggests 17 million people are affected [4]. Definitions of Long Covid or Post-acute COVID-19 syndrome (PACS) or Post-acute Sequelae of SARS CoV-w (PASC) vary but are characterized by persistent symptoms and/or delayed or long-term complications persisting or beginning beyond four weeks from the symptom onset of SARS-CoV-2 infection [5–7].

WHO identifies over 200 PACS/PASC related symptoms that impact everyday activities. The five most common long-term symptoms are fatigue (58%), headache (44%), attention disorder (27%), hair loss (25%), and dyspnea (24%) [8]. Other persistent symptoms have been reported, including cough, chest

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pain, myalgia, joint pain, impaired mobility, cognitive impairment, olfactory and gustatory dysfunction, sleep disorders, depression, anxiety, post-traumatic stress disorder, gastrointestinal upset, rashes, and palpitations [9–14]. As an international priority, the US has dedicated \$1.15 B for diagnosis and treatment research [15].

Muscle damage resulting from COVID-19 infection has been reported. Specifically, both cardiac and skeletal muscle tissues exhibit robust ACE2 (angiotensin-converting enzyme 2) and TMPRSS2 (transmembrane serine protease 2) expression, indicating a potential vulnerability to SARS-CoV-2 infection in muscle. This damage results from the inflammatory effects, cytokine storm, and muscle catabolism [16–18]. Consequently, damage to muscles will result in a decrease in muscle strength and may contribute to pain. In previous studies, we demonstrated a significantly lower level of muscle strength in women after recovering from COVID-19 compared to age-matched healthy individuals [19].

Moreover, even without COVID-19 infection, in the 6<sup>th</sup> decade of life, an accelerated, non-linear decrease in muscle strength has been reported (up to 15%), and by the 8<sup>th</sup> decade, this strength loss may be up to 30% in healthy elderly [20]. In people with less active lifestyle, these age-related changes manifest earlier [21]. This natural age-related loss is compounded by COVID-19 infection and pandemic protocols causing decreased socialization and work/exercise outside of the home. Once infected with COVID 19, prescribed rest to minimize the metabolic demand and channel resources towards the recovery process may have accelerated a functional decline in elderly patients. As a point of reference, Kortebein et al. (2008) found a substantial loss in muscle strength and power (knee extension  $p = 0.004$ , knee flexion  $p = 0.003$  and stair ascent power  $p = 0.01$ ) after 10 days of bed rest in healthy elderly people (60–85 years old) [22].

To reduce the consequences of COVID-19 infection, the NICE guideline [23] and American consensus [24] recommend integrated multidisciplinary rehabilitation services for individuals with long-term effects of COVID-19. However, the effectiveness of exercise interventions for post-COVID patients remains insufficiently documented. Most reports concerned the effects of pulmonary rehabilitation, where the study groups were compared to no rehabilitation/drug-only intervention [25–27].

Previous studies have suggested that resistance training is both safe and highly effective in combatting strength loss, as well as declining functional capacity in the healthy elderly [28,29]. It has not been established that these treatments are safe to implement or effective in individuals with PACS/PASC. Furthermore, the high volume of affected individuals exceeds the availability of rehabilitation specialists to create individual programs.

Therefore, the aim of this study was to develop and implement a specific, early, and well tolerated protocol-based intervention to reduce muscle weakness in older adults impacted by COVID-19. The secondary aim was to test the hypothesis that resistance training will improve muscle strength in post-COVID survivors and that the strength gains will be accompanied by improved function.

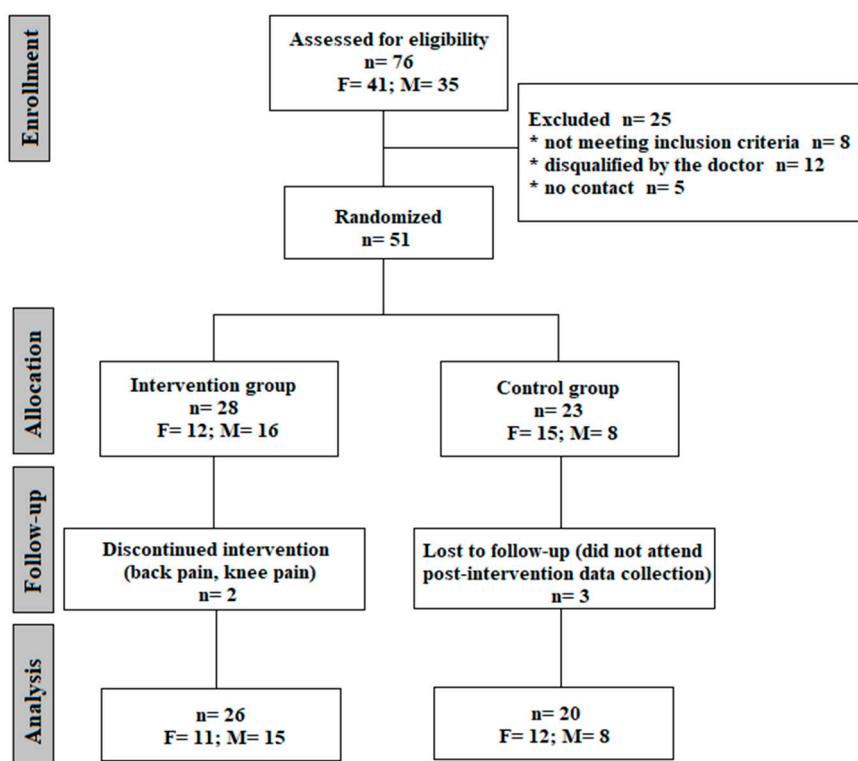
## 2. Materials and Methods

### 2.1. Subjects

Participants were recruited from nursing homes, primary health care facilities, a University of the Third Age program, social media of local University, and from surrounding communities using posters, leaflets, and videos from January to March 2023. The inclusion criteria were as follows: both sexes, age 65 and older, positive RT-PCR test and/or positive result in test for antibodies against the SARS-CoV-2 coronavirus 3–12 months prior to commencement of the study and reporting one or more of post-covid signs and symptoms, such as fatigue, muscle weakness, dizziness, headache, memory and concentration disorders, exercise intolerance, depression. Before initiating the rehabilitation program, participants were screened by physician, and exacerbation of post-exercise symptoms was assessed based on a questionnaire [30] and an orthostatic test (Freeman et al. 2011). People aged less than 65, with active cardiac disease, oxygen desaturation below 95% for more than 1 minute, dysfunctions of the autonomic nervous system (orthostatic intolerance), and serious health conditions such as cancer were excluded from this study.

It's worth mentioning that 92% of respondents were vaccinated with at least one dose of the anti-SARS-CoV-2 vaccine, and only 27% of study participants got sick before the vaccine. The average time

from the onset of the disease in people qualified according to the inclusion criteria was 9 months, and 33% described the infection as mild, 51% as moderate, 10% as severe and 6% as very severe. Post-infection interview regarding recall of signs and symptoms was conducted. The symptoms reported by all participants included: dizziness and equilibrium disorders (55% of responded), perceived muscle weakness (35,3%), exercise intolerance (31,3%) and 19.6%- memory and concentration symptoms. Other, less frequently reported symptoms included: cough, dyspnea, libido deterioration, insomnia, and deterioration of taste and smell. After meeting inclusion criteria and passing medical screening, participants were allocated into one of two groups: an intervention group, that received resistance training, and a control group, advised to retain their activity level as usual. Allocation to groups was done randomly using an Excel random number generator. Finally, 46 people completed the study protocol, including pre- and post- testing conducted in a Central Laboratory at the Józef Pilsudski University of Physical Education in Warsaw. The consort diagram is presented in Figure 1.



**Figure 1.** CONSORT flow diagram of the progress through the phases of parallel trial in two groups.

F- females; M- males.

The data from 46 post-COVID Seniors were analyzed. Anthropometric characteristics of the tested groups at baseline are shown in Table 1. No significant differences were found between the intervention and control groups in anthropometric parameters, except the age in men groups ( $p<0.05$ ). The necessary minimum total number of subjects ( $n=40$ ) was obtained using the G\*Power program assuming detection of medium-sized effects ( $\eta^2=0.06$ ) at a significance level of  $\alpha=0.05$  and statistical power of 0.85.

**Table 1.** Anthropometric characteristics of the tested groups at baseline.

	Intervention (f=11, m=15)	Control (f=12, m=8)
<b>Female</b>		
Age (years)	69.3±5.2	73.3±7.4
Body mass (kg)	65.3±10.5	72.6±12.1
Height (m)	163.1±7.6	160.9±5.7

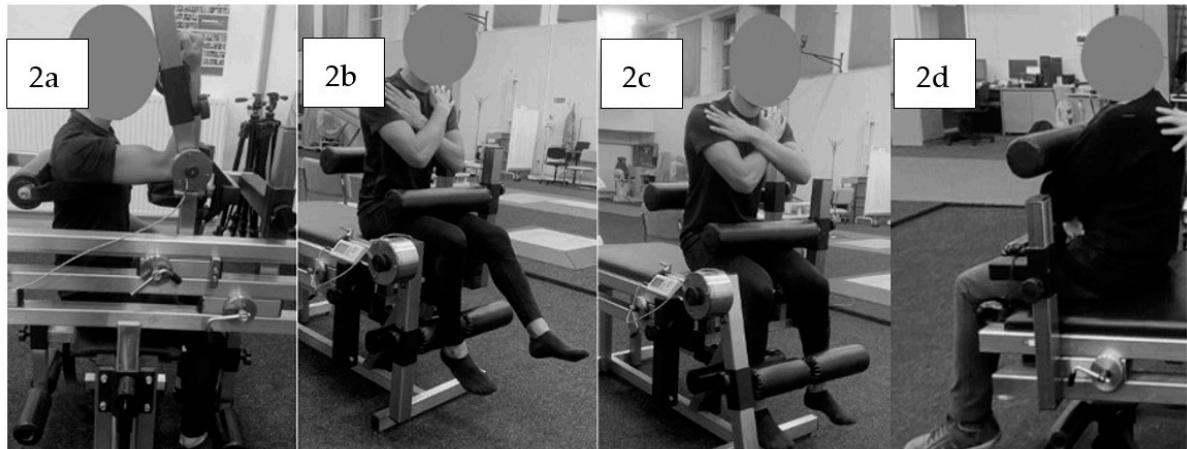
BMI (kg/m <sup>2</sup> )	24.5±4.8	28.1±3.3
<b>Male</b>		
Age (years)	69.5±4.8	75.6±7.1*
Body mass (kg)	87.7±15.1	89.5±20.3
Height (m)	176.7±6.9	179.0±6.1
BMI (kg/m <sup>2</sup> )	27.86±3.4	27.07±5.4

## 2.2. Methods

### 2.2.1. Muscle strength tested in a static condition.

Strength, defined as the ability to produce force, was measured in two conditions: static and dynamic. The results of the test in static condition will be referred to as the moment of force. All measurements were performed after the subject was familiarized with the measurement protocol and after a warm-up. Participants were instructed to exert maximal effort while pushing the bar or the testing apparatus for 3 seconds. The subjects performed 3 repetitions with one minute of rest between measurements. The highest value (peak) was used for the statistical analysis. Finally, the absolute [N·m] and relative to body mass [N·m·kg<sup>-1</sup>] values of the following parameters of the strength were taken for the analysis: *elbow flexors (EF)*, *knee flexors (KF)*, *knee extensors (KE)*, *trunk flexors (TF)* and *trunk extensors (TE)*.

The highest value (peak moment of force) muscle groups were extracted for further analysis: flexors and extensors of the knee, trunk and only flexors of the elbow. Measurements were taken on a custom-made device ("JBA", Zbigniew Staniak, Poland)– type LR1-P (upper extremities) and TBK2-PM (lower extremities and trunk). Technical specifications of the measuring device are as follows: • torque meter: strain gauge head – measuring range for 0 to 500 N·m for LR1-P and 0–2000 N·m for TBK2-PM, • relative error in strain-gauge bridge amounts to 1.0%. Measurements of muscle torque were performed according to the generally accepted principles [30]. The dominant upper extremity's forearm flexors were tested seated, the angle at the shoulder joint was 90 degrees of flexion, the supported forearm was perpendicular to the arm. The trunk was stabilized by a belt (Figure 2a).



**Figure 2.** Muscle strength testing position in a static condition, (2a) elbow flexors (2b) knee extensors (2c) knee flexors (2d) trunk flexors.

The knee flexor and extensor torque of the dominant limb were also tested in the sitting position, with the hip and knee joints at 90° of flexion. The subjects were stabilized by a close-fitting roller at the level of the anterior iliac spines at proximal part of the tight and posteriorly at lumbar spine. The upper limbs were crossed over the chest (Figure 2b, 2c). Trunk flexors and extensors were tested in the sitting position and the estimated axis of rotation at the hip, coincided with the axis of rotation of the torque meter lever. The

participants were also stabilized by a roller at the level of the proximal part of the tight with the upper limbs crossed over the chest (Figure 2d).

### 2.2.2. Isokinetic strength evaluation

The dynamic strength evaluation was conducted using an isokinetic dynamometer. The BiodeX System 3 dynamometer (BiodeX Medical Systems, Inc., Shirley, NY, USA) was used for the isokinetic strength assessments. Only the dominant leg was tested. Isokinetic knee extension and flexion pattern at 60°/s (5 repetitions) and 180°/s (10 repetitions) were tested. A 90-second-rest time was provided between sets. Subjects were seated in the chair of the dynamometer and stabilized by belts around their trunk, pelvis, and mid-thigh of the test leg. Hip flexion was set at 85° and the dynamometer axis was aligned to the knee's anatomical axis of rotation. The ankle pad was positioned just above the medial malleolus (Figure 3).



**Figure 3.** Muscle strength testing position in a dynamic condition - isokinetic knee extension and flexion at 60°/s (5 repetitions) and 180°/s (10 repetitions).

Range of motion (ROM) was limited to between 90° and 0° of flexion. Extension ROM for each participant was defined in accordance with their individual limits. Participants were instructed to exert maximal effort during each test while verbal encouragement and visual feedback were provided. The following parameters were analyzed: *Peak Torque E60 [N·m], Peak TQ/BW(%) E60 [N·m·kg-1], Peak Torque F60 [N·m], Peak TQ/BW(%) F60 [N·m·kg-1], Peak Torque E180 [N·m], Peak TQ/BW(%) E180 [N·m·kg-1], Peak Torque F180 [N·m], Peak TQ/BW(%) F180 [N·m·kg-1]*.

### 2.2.3. Functional tests

#### 2.2.3.1. Time Up and Go test (TUG).

The TUG required participants to start from a seated position in standardized chair, stand up and walk 2.44 m, walk around a cone, return to the chair, and get back into the starting position. The time taken was measured to the millisecond.

#### 2.2.3.2. Chair Stand Test

- CS-30 scores - participants were instructed to complete sit-to-stand trials using a 40-cm high seat without using their arms as many times as possible in 30 s [33]. The number of stands was recorded.
- 5STS - participants were asked to sit on a 40-cm high seat without using their arms and then stand repeatedly for five times as quickly as possible [34]. The time was recorded.

### 2.3. Intervention

Resistive Training (RT) aimed at improving muscle strength was conducted in the Zdrofit Gym Warsaw-Bielany twice a week, 60 min per session for 8 weeks according to the recommendations of World Physiotherapy (World Physiotherapy, 2021 and NICE, 2020) [23]. Before every session heart rate, blood pressure and oxygen saturation were measured. If the blood pressure was  $>160/100$  mmHg, heart rate (HR)  $>100$  or  $<50$  beats per minute, participants were not allowed to do exercises during that session.

The first training session included the determination of 1 Repetition Maximum (1RM) on each exercise. Therefore, all subjects performed 4-5 trials with increasing load and rest periods between trials were 3 min of passive recovery. The goal was to perform 3-5 repetitions with maximum load. The subjects were instructed to perform the exercise at a comfortable pace. The 1RM was calculated according to the formula developed by Brzycki (1993) [35].

Each training session aimed to reach an exercise intensity of 70% of 1RM and consisted of three sets of 12 repetitions on each exercise (incline bench press, 45 degrees leg press, latissimus pull down, trunk crunch, T-Bar row, leg extension, and leg curl) (Figure 4).



**Figure 4.** Exercises in each session: (4a) leg extension, (4b) leg curl, (4c) trunk crunch, (4d) incline bench press, (4e) 45 degrees leg press, (4f) latissimus pull down, (4g) T-Bar row.

The rest periods between sets included a 2 min passive recovery. Before performing each training session subjects performed a 15 min general warm-up on an orbitrec or treadmill with individual intensity (60-65% HRmax). The training loads during the training sessions were increased individually by 5 kg when a subject completed all repetitions during an exercise.

### 2.3. Statistical analysis

Statistical analysis was performed using STATISTICA 14.0 (TIBCO Software Inc. (2020). Data Science Workbench, version 14. <http://tibco.com>). The normality of the distributions of the study variables in the groups was tested using the Shapiro-Wilk test. Comparisons of means were made using analysis of variance for repeated measures. GROUP (Control, Intervention) and SEX (Men, Women) were used as fixed factors. The repeated-measures factor TESTING SESSION took two values: Before, After. Due to the primary aim of the study, it was assumed that the different response of the study groups would be described by the GROUP x TESTING SESSION interaction. Detailed comparisons were done using the Tukey post-hoc test. In the case of variables not meeting the condition of normality of distributions, the Mann-Whitney U test was used for comparisons between groups. Changes in variables before and after

the intervention were assessed using the Wilcoxon test. For these variables, the response to the intervention was assessed by comparing the increments of the study variables in the two groups (Intervention, Control) using the Mann-Whitney U test.

Effect sizes were assessed using partial eta square (ANOVA), Glass's rank-biserial correlation coefficient (Mann-Whitney test) and equivalent correlation coefficient (Wilcoxon test). A significance level  $\alpha=0.05$  was adopted.

### 3. Results

The results of Seniors in the intervention group, who did not miss more than 3 sessions, were taken for analysis. Two subjects were lost due to unrelated musculoskeletal pain (low back, knee pain). The average attendance rate was 93% (80%-100%).

Post-infection interview regarding recall of signs and symptoms were conducted in the intervention group for comparison with the results before training. The frequency of post-COVID-19 symptoms before and after RT are shown in the table below (Table 2).

**Table 2.** Self-reported, percentage of common post-COVID-19 symptoms in an intervention group before and after training. .

Post-COVID-19 symptom	Dizziness and equilibrium disorders	Perceived muscle weakness	Exercise intolerance	Memory and concentration symptoms
Before intervention	46.2%	38.5%	38.5%	15.3%
After intervention	15.4%	7.6%	3.8%	7.6%

The results of two testing sessions (pre- and post-test) of maximal muscle torque relative to body mass values [ $\text{N}\cdot\text{m}\cdot\text{kg}^{-1}$ ] in static conditions are shown in Table 3. In the intervention group, both in female and men, an increase in all parameters was observed. The largest increase was found in a trunk flexion (18.9%) and extension (22%) in a group of active men ( $p<0.001$ ). In turn, in the group of active women, the most significant improvement was noticed in the elbow flexion (17%) knee flexion (20.2%) and a trunk extension (23.3) ( $p<0.01$ ). Absolute values of peak torque at the elbow, knee and trunk in static conditions are attached in Supplement files.

In the control group, a decrease in most parameters values expressed relative to body mass [ $\text{N}\cdot\text{m}\cdot\text{kg}^{-1}$ ] was observed in a group of men, in contrast to female that improved the results but not significantly. To assess the effect of intervention, GROUP x TESTING SESSION (pre-post-test) interactions were considered. The interaction was significant for relative maximal muscle torque of all measured muscle groups ( $p<0.001$ ) (Table 3).

Statistical analysis revealed significant improvement in a knee extension isokinetic torque ( $60^\circ/\text{s}$ ) in absolute values and normalized to body mass ( $p<0.05$  and  $p<0.01$ , respectively) and in a knee flexion isokinetic torque ( $60^\circ/\text{s}$ ) ( $p<0.05$ ) in a women group after intervention. In an intervention group of men, the significant improvement was observed in a knee extension isokinetic torque ( $180^\circ/\text{s}$ ) normalized to body mass ( $p<0.05$ ). However, due to the increase in isokinetic torque in both groups (control and intervention), the effect of intervention GROUP x TESTING SESSION (pre-post-test) interactions were not found (Tables 4 and 5). The results of flexors/extensors ratio are presented in Table 6. The values of the F/E ratio increased in the intervention group, however, the effect of intervention GROUP x TESTING SESSION (pre-post-test) interactions were not observed.

The results of three functional tests are presented in Table 7. In the Intervention group, both in men and female, a significant improvement was observed in Chair test (s) ( $p<0.001$  and  $p<0.05$ , respectively) and Chair test (n) ( $p<0.01$  in both groups). The interaction GROUP x TESTING SESSION (pre-post-test) was significant for Chair test (s) ( $F_{1,42}=8.49$ ,  $p<0.01$ ,  $\eta^2=0.68$ ) and Chair test (n) ( $Z=4.65$ ,  $p<0.001$ ,  $R=-0.806$ ).

**Table 3.** Absolute values of maximal muscle torque at the elbow, knee and trunk in static conditions normalized to body mass.

TESTING SESSION		GROUP				GROUP X		
		Control		Intervention		TESTING SESSION		
		Male (n = 8)	Female (n = 12)	Male (n = 15)	Female (n = 11)	F <sub>1,42</sub>	p	η <sup>2</sup>
EF/m (N·m·kg <sup>-1</sup> )	Before	0.71 ± 0.19	0.42 ± 0.10	0.75 ± 0.14	0.49 ± 0.16	17.41	0.0001	0.293
	After	0.62 ± 0.14	0.45 ± 0.09	0.78 ± 0.13	0.59 ± 0.13**			
KF/m (N·m·kg <sup>-1</sup> )	Before	0.94 ± 0.22	0.69 ± 0.21	1.08 ± 0.31	0.75 ± 0.22	10.15	0.0027	0.195
	After	0.97 ± 0.27	0.69 ± 0.18	1.21 ± 0.28*	0.94 ± 0.28**			
KE/m (N·m·kg <sup>-1</sup> )	Before	1.55 ± 0.39	1.23 ± 0.27	1.84 ± 0.54	1.43 ± 0.43	11.35	0.0016	0.213
	After	1.26 ± 0.38	1.28 ± 0.22	2.00 ± 0.46	1.57 ± 0.45			
TF/m (N·m·kg <sup>-1</sup> )	Before	1.51 ± 0.40	1.05 ± 0.31	1.84 ± 0.52	1.14 ± 0.33	14.21	0.0005	0.253
	After	1.44 ± 0.38	1.13 ± 0.42	2.27 ± 0.67***	1.46 ± 0.34*			
TE/m (N·m·kg <sup>-1</sup> )	Before	2.65 ± 0.77	1.64 ± 0.69	3.20 ± 1.08	2.64 ± 1.23	15.75	0.0003	0.273
	After	2.48 ± 0.76	1.84 ± 0.82	4.10 ± 0.81***	3.44 ± 1.30**			

Difference pre-post: \* - p < 0.05, \*\* - p < 0.01, \*\*\* - p < 0.001. EF - elbow flexors, KF - knee flexors, KE - knee extensors, TF - trunk flexors, and TE - trunk extensors, m - body mass.

**Table 4.** Knee extension and flexion isokinetic torque (60°/s).

TESTING SESSION		GROUP				GROUP x TESTING SESSION		
		Control		Intervention		TESTING SESSION		
		Male (n = 8)	Female (n = 12)	Male (n = 15)	Female (n = 11)	F <sub>1,42</sub>	p	η <sup>2</sup>
Peak Torque E60 (N·m)	Before	109.3 ± 34.8	78.5 ± 20.0	143.5 ± 39.4	84.1 ± 15.8	1.06	0.3095	0.025

	After	120.5 ± 24.5	80.0 ± 19.0	160.0 ± 37.2*	90.9 ± 18.6			
Peak Torque/m E60 (N·m·kg <sup>-1</sup> )	Before	1.29 ± 0.59	1.10 ± 0.30	1.65 ± 0.36	1.31 ± 0.28	1.11	0.2982	0.026
	After	1.47 ± 0.55	1.11 ± 0.29	1.86 ± 0.40**	1.43 ± 0.31			
Peak Torque F60 (N·m)	Before	68.1 ± 29.4	45.4 ± 19.6	76.4 ± 31.1	46.4 ± 13.5	3.73	0.0601	0.082
	After	69.2 ± 23.0	47.2 ± 14.1	88.3 ± 27.2*	54.3 ± 13.6			
Peak Torque/m F60 (N·m·kg <sup>-1</sup> )	Before	0.78 ± 0.36	0.64 ± 0.30	0.88 ± 0.34	0.74 ± 0.26	3.30	0.0766	0.073
	After	0.83 ± 0.37	0.65 ± 0.20	1.02 ± 0.28	0.86 ± 0.25			

Difference pre-post: \* - p < 0.05, \*\* - p < 0.01, \*\*\* - p < 0.001. Peak Torque E60- peak torque of knee extension at 60°/s, Peak Torque/m E60- peak torque of knee extension at 60°/s normalized to body mass, Peak Torque F60 - peak torque of knee flexion at 60°/s, Peak Torque/m F60 - peak torque of knee flexion at 60°/s normalized to body mass.

**Table 5.** Knee extension and flexion isokinetic torque (180°/s).

		GROUP				F <sub>1,42</sub>	p	η <sup>2</sup>			
		Control		Intervention							
		TESTING SESSION									
		Male (n = 8)	Female (n = 12)	Male (n = 15)	Female (n = 11)						
Peak Torque E180 (N·m)	Before	65.08 ± 13.36	45.00 ± 19.29	88.32 ± 25.72	48.33 ± 14.62	0.16	0.6870	0.004			
	After	78.15 ± 14.81	53.16 ± 14.07	96.62 ± 20.05	58.12 ± 12.33						
Peak Torque /m E180 (N·m·kg <sup>-1</sup> )	Before	0.76 ± 0.26	0.63 ± 0.28	1.02 ± 0.27	0.76 ± 0.24	0.09	0.7705	0.002			
	After	0.94 ± 0.30*	0.74 ± 0.21	1.12 ± 0.24	0.91 ± 0.21*						
Peak Torque F180 (N·m)	Before	47.08 ± 16.67	34.14 ± 16.70	61.03 ± 27.26	35.13 ± 10.89	0.10	0.7573	0.002			
	After	57.86 ± 14.98	36.90 ± 14.05	67.02 ± 20.62	45.16 ± 14.68						
Peak Torque /m F180 (N·m·kg <sup>-1</sup> )	Before	0.56 ± 0.25	0.48 ± 0.24	0.70 ± 0.28	0.56 ± 0.20	0.27	0.6086	0.006			
	After	0.70 ± 0.28	0.52 ± 0.23	0.78 ± 0.23	0.71 ± 0.23						

Difference pre-post: \* - p < 0.05, \*\* - p < 0.01, \*\*\* - p < 0.001. Peak Torque E180- peak torque of knee extension at 180°/s, Peak Torque E180/m- peak torque of knee extension at 180°/s normalized to body mass, Peak Torque F180 - peak torque of knee flexion at 180°/s, Peak Torque F180 - peak torque of knee flexion at 180°/s normalized to body mass. .

**Table 6.** Knee flexors/extensors peak torque ratio at 60°/s and 180°/s.

		GROUP						
		Control		Intervention				
		TESTING SESSION				GROUP x TESTIN SESSION		
		Male (n = 8)	Female (n = 12)	Male (n = 15)	Female (n = 11)	F <sub>1,42</sub>	p	η <sup>2</sup>
Flexors/Extensors ratio (%) 60	Before	62.20 ± 20.79	56.30 ± 18.10	51.95 ± 10.80	54.82 ± 11.06	3.44	0.0707	0.076
	After	55.08 ± 14.85	56.85 ± 8.75	55.01 ± 8.28	59.90 ± 11.76			
Flexors/Extensors ratio (%) 180	Before	73.16 ± 23.67	73.48 ± 12.36	67.42 ± 15.50	72.91 ± 14.11	0.48	0.4925	0.011
	After	74.29 ± 14.75	68.58 ± 13.87	69.06 ± 13.51	74.49 ± 15.74			

**Table 7.** Functional tests; Time Up and Go (TUG) and Chair Tests pre and post intervention.

		GROUP						
		Control		Intervention		GROUP x TESTING SESSION		
		Male (n = 8)	Female (n = 12)	Male (n = 15)	Female (n = 11)	F <sub>1,42</sub>	p	η <sup>2</sup>
TUG (s)	Before	6.16 ± 0.91	6.83 ± 1.57	5.71 ± 0.89	5.59 ± 0.69	3.06	0.0876	0.068
	After	6.21 ± 1.18	6.65 ± 1.34	5.17 ± 0.63	5.21 ± 0.77			
Chair test 5STS (s)	Before	9.10 ± 2.56	10.78 ± 4.54	9.48 ± 2.00	8.78 ± 1.98	8.49	0.0057	0.168
	After	8.72 ± 2.47	9.60 ± 3.22	6.65 ± 1.15 <sup>##</sup>	7.01 ± 1.35 <sup>#</sup>			
						Z	p	R
Chair test CS-30 (n)	Before	17.5 (13.0 - 22.5)	19.0 (13.0 - 19.5)	15.0 (14.0 - 20.0)	18.0 (14.0 - 21.0)	4.65	0.0001	0.806
	After	17.0 (14.0 - 24.0)	16.0 (13.0 - 19.0)	24.0 (21.0-28.0) <sup>##</sup>	25.0 (19.0-28.0) <sup>**##</sup>			

Difference intervention-control: \* - p < 0.05, \*\* - p < 0.01, \*\*\* - p < 0.001,. Difference pre-post: # - p < 0.05, ## - p < 0.01, ### - p < 0.001, n - number of repetitions, s - seconds.

#### 4. Discussion

Our results show that there was improvement in strength and function utilizing a protocol- based intervention of resistance training in elderly adults post-COVID-19 infection. The specific protocol used was short enough to be plausible in a clinical setting, was well tolerated, and patient was able to adhere to the program as evidenced by the low drop out rate.

The seven equipment-based progressive resistive exercises targeted three body regions, namely the upper extremity, lower extremity, and trunk. For testing purposes sagittal plane isometric strength of the elbow, knee and trunk was selected. Additionally, knee extensors and flexors were tested isokinetically at 60 degrees/seconds and 180 degrees/seconds.

*Elbow flexor strength* is among the most tested muscle groups of the upper extremity. This is partially due to the ease and reliability of the testing procedure. A recent study by Rodriguez-Rodriguez determined that elbow flexor power, tested in a seated position is strongly related to well-established whole body functional test used with older adults [36]. In our study, *elbow flexor strength* performance, tested *isometrically*, was a proxy for overall upper extremity physical performance. At baseline the average, normalized to body weight, *elbow flexor strength* was  $0.75 \pm 0.14$  for men, and  $0.49 \pm 0.16$  for women. These values were higher than those estimated from a subgroup of 178 healthy adults for male subjects 0.59 and female =0.38. [37]. In our study, a post-hoc analysis determined an improved strength in women (from  $0.49 \pm 0.16$  to  $0.59 \pm 0.13$ ), but in men. These comparisons suggest that the subjects in our study, especially women, were not overly weak and were able to improve their elbow flexor strength.

*Knee flexors and extensors* are the primary muscle groups selected for strength testing in the lower extremity. Seated position, ease of thigh and trunk stabilization in addition to a known susceptibility to age related muscle loss make this test the first choice for lower extremity strength testing [38]. At entry into this study, the *normalized isometric knee flexor strength* (Nm/kg) in men and women was higher than that reported by Sarabon et al., 1.08 versus 0.79 for men and 0.75 versus 0.64 for women [39]. Our intervention improved knee flexor strength in men from  $1.08 \pm 0.31$  to  $1.21 \pm 0.28$  and  $0.75 \pm 0.22$  to  $0.94 \pm 0.28$  for women. It is worth noting the surprising similarity of the pre- and post- 8 weeks data for the control group (mean difference for men 0.03 and null difference for women). Indirectly, the consistent data for the control group speaks to the reliability of knee flexor strength measurements in our study. *Isometric knee extensor strength* was lower than in a group of healthy older adults [39] and higher than in a group of institutionalized but independent women older than 70 years of age [40]. Although the averaged bodyweight-normalized values improved in the intervention group, our Group x Testing Session design did not identify differences in a post-hoc analysis.

*Since inception in 1960-th, isokinetic knee torque performance* has been used extensively in characterization and intervention studies providing wide-ranging comparative data. The touted benefits of isokinetic strength testing are dynamic testing of strength at a pre-set constant velocity allowing for the torque to be measured reliably throughout the range of the joint's motion. In this study, we selected the two most tested velocities, 60 and 180 degrees/second. The body weight normalized peak torque at 60 degrees/second was within the values reported testing older adults [41,42]. Our 8-week strengthening intervention did not result in altered peak torques at either velocity. This is intriguing, since the exercise program was equipment-based and dynamic, and the isometric strength discussed above improved. It is possible that the moving resistance arm during isokinetic testing provides for a secondary cognitive demand, or perhaps fear, resulting in lower torque output.

Although trunk strength testing is less commonly implemented than elbow and knee, that metric provides a valuable quantitative characterization of muscle performance in the largest segment of body mass. At entry into this study, our cohort's average isometric trunk extensor strength, normalized to body weight was within the range of values reported by Keinbacher et al. on population of older adults aged 50 to 90 years (average age 67.2 years old) with history of low back pain [43]. In the current study, the 8 weeks of exercise program significantly increased the normalized trunk flexor strength from 1.84 Nm/kg to 2.27 Nm/kg for men and from 1.14 Nm/kg to 1.46 Nm/kg for women.

Common daily tasks such as stand-to-sit, sit-to-stand as well as walking initiation and termination, have become the basis of widely used standardized functional tests in clinical-trial and in the clinic. This study chose to use three functional tests: time up and go, repeated sit-to-stand, number of sit to stand performed in 30 seconds.

The *TUG test* is performed in the following sequence: start in seated position, stand up, walk 2.44 meters, and return to a seated position as fast as able. The results are reported in seconds. In addition to functional mobility and gait, the *Time Up and Go (TUG)* test assesses non-vestibular and vestibular aspects of balance relevant for the post-COVID symptoms. This test requires controlling whole body acceleration in the vertical and fore-aft directions. The older adults in our study performed the test within the ranges reported in the literature (5.1 to 9.0 seconds) [44–48]. There was no statistically significant improvement in performance, since results at entry into the study were typical for this age group. The *repeated sit-to-stand test* is performed 5 times. The results are reported in seconds. The test requires controlling vertical acceleration of the center of mass. The average baseline values in the current study ranged between  $8.78 \pm 1.98$  and  $10.78 \pm 4.45$  seconds, which was shorter than average values for older adults in a large study (5352 participants) by Ostchega et al.  $13.11 \pm 0.19$  for male and  $14.05 \pm 0.2$  for women [49]. Both male and female subject in our study performed the test faster, but the participants in the intervention study showed significant improvement. Finally, to gain insight into fast transition movements, this study chose to use the 30 seconds sit-to-stand test. The outcome of this test is the number of completed repetitions. At the entry into this study the number of repetitions ranged between 13 and 22. Both intervention groups improved significantly male from 15 to 24 and female from 18 to 25 repetitions of sit-to-stand in 30 seconds. This improvement is similar the one reported by Sato et al. testing the effects of exercise games (from 17 to 24 repetition in a group of women aged  $70 \pm 5.4$  years [50].

There are several limitations within this study. First is the small number of subjects in the study. While larger numbers would be preferred, the size is appropriate considering that this is one of the first studies utilizing an active exercise program following the end of the public health emergency. It was important to establish the safety and effectiveness of the protocol before increasing the size of the cohort. The second limitation is that the participants in the intervention group were already functioning at a reasonably high level. Once again, establishing efficacy in a higher functioning group was an important first step. It is likely that a group functioning at a lower level would have achieved even greater gains. Another limitation is that we were not able to systematically track symptom improvement within the control or intervention group. Future studies will expand on this work with larger cohort and assessing impact on a cluster of symptoms.

## 5. Conclusions

The COVID-19 pandemic had an immediate and profound economic and health impact across the world. Obvious impacts on health included illness, hospitalization, and death. However, we are now realizing that the disease also created a long-term impact that will last for generations. Many individuals are suffering from poorly understood symptoms of “long covid” and many of our society have not recovered from the effect of decreased physical activity and increased sitting time necessitated by online meeting and work.

The pandemic was an insult/injury to body systems that requires rehabilitation. While we cannot yet address all problems, we can address the strength deficits that impact function and long-term health. This will lower the burden of care, lower risk of falls, reduce assistance needed, increase independence, and improve quality of life.

As generalized fitness programs may not be appropriate for people who have been infected with COVID-19 or suffer from other chronic conditions, there has been a significant demand on medical professionals. Unfortunately, these specialists are unavailable because of the extreme backlog of deferred care that will last for many years. It is critical that a program that can be individualized based on criteria is developed and demonstrated to be effective and well tolerated.

Our results demonstrate the effectiveness of a well-tolerated, protocol-based approach that can be used by professionals to diminish long lasting functional deficits following COVID-19 infection in post-covid survivors.

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**Institutional Review Board Statement:** The study was conducted in accordance with the Declaration of Helsinki, and approved by the Ethical clearance for the study was granted from the local Ethics Committee (SKE 01-41/2022) (21.XII.2022). The study protocol was registered on clinicaltrials.org (NCT05934279).

**Informed Consent Statement:** Informed consent was obtained from all subjects involved in the study. Written informed consent has been obtained from the patients to publish this paper.

**Data Availability Statement:** The datasets generated and analyzed during the current study are not publicly available due to the restrictions involved when obtaining ethical approval for our study, which commit us to share the data only with members of the research team but allow data to be made available from the corresponding author upon reasonable request.

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