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# Subsequent Complications Following COVID-19 an Systematic Reviews

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# **ABSTRACT**

Introduction: From COVID-19, a broad spectrum of early, intermediate and late complexities have been recognized with this SARS-CoV-2 since then. Still, there can be sequelae that persist from recovery following the acute illness of COVID-19. Research aim is to thoroughly examine the current evidence on delayed consequences given to COVID-19.

Methods: A broad search was performed in various electronic databases including Scopus, Web of Science, PubMed, Embase (up to Oct 2024) using specific keywords for an exhaustive review.

Results: A systematic review on the most common complications underlying COVID-19 Long COVID in 30 selected studies revealed a diverse set of major complications of developing long COVID-19. Neurologic and psychiatric presentations dominated the list of the sequelae that followed next were respiratory, musculoskeletal, rheumatologic considerations and ciring cardiovascular as well hepatic and gastrointestinal aspects.

Conclusion: Concluding, the lingering consequences of COVID-19 influence multiple physiological systems ranging from mild to severe and cases with these latter extremes. Long COVID-19 patients with my journey fatigue, and a swath of neuropsychiatric morbidities, therefore fatigue has been among the most consistently described late complications seen in individuals affected by prolonged COVID. Next are the respiratory problems that include dyspnea, cough and chest tightness representing important long coronavirus disease 2019. Due to the enduring and delayed nature of these sequelae, there should be ongoing vigilance for the demonstration of characteristic signs and symptoms in both patients who are well as those who have deteriorated.

Keywords: COVID-19, SARS-CoV-2; Post-Acute COVID-19 Syndrome; Outcome Assessment of Patient

# INTRODUCTION

Since the emergence of severe acute respiratory syndrome coronavirus-2 (SARS-CoV-2) in 2019, there has been a lot of debate about the clinical impact of COVID-19,which is a viral disease caused by this entity. Most importantly, SARS-CoV-2 infects host cells via angiotensin-converting enzyme 2 (ACE2) receptors . With ACE2 expressed on many human organs, this has led to the multiplicity of clinical forms of COVID-19 including pneumonia, myocarditis, myocardial infarction, renal injuries as well as neurological symptoms and gastrointestinal symptoms(1,2,3,4) [f1]. Emerging evidence on the global spread of SARS-CoV-2 shows emerging clinical manifestations of COVID-19. Duration of symptoms and the acute-phase virus-specific

complications also may vary in times as early-mid-late manifestation after infection (1-4) Complications after any infection that remain for days to months more than the normal baseline post-infection are considered postCOVID-19 complications [5, 6, 7,8,9]. Most of the complications do in early stage of COVID-19 as well, but with growing evidence we saw some late complications weeks and months after clearance of SARS-CoV-2 in survivors. These late complications are unlikely to arise from direct organ damage. Immune dysregulation, coagulopathy and endothelial injury have been put forward as explanations for these late effects. It also seems that COVID-19 affects almost every system in the body, with widely varying patterns, degrees, and lengths of the late manifestation. Lastly,the late complications likewise correspond to poor laboratory results numbers inconsistent with organ dysfunction [10, 11,12].

Systematic reviews and meta-analyses have thoroughly documented the late complications of COVID-19, displaying the range of organ injuries linked with this illness. AbstractThis umbrella review aims to collate the evidence on the late sequelae of COVID-19 in people, will inform clinical care for healthcare workers and summarises all exacerbations likely after COVID-19, advocating a focus on follow-up after COVID-19.

#### Methods

Systematic reviews and meta-analyses have extensively reported late complications with COVID-19, and the setting of this umbrella review.

#### 2.2 Data sources

We searched in the following resources: Web of Science (WoS) for systematic reviews, PubMed/MEDLINE®, Scopus and Embase. The search was restricted to those who reported in an english language.

#### 2.2 Selection criteria

two-step selection of studies was applied

A. Title and Abstract Screening

Studies were initially evaluated according to their titles as well the abstracts in order to identify potentially relevant studies.

B. Full Text Screening

The second stage involved a full text review of the papers which passed preliminary screening Phase 2.

Eligible for inclusion in the review: full systematic review and long-term studies regarding complications specific to COVID-19.

## 2.3 Exclusion Criteria

Studies were excluded based on the following types:

- A. Published human studies
- B. Duplicate publications
- C. Narrative reviews
- D. Umbrella reviews
- E. Preprints
- F. Conference abstracts

#### **Results**

We included studies from several countries, indicating a global effect of COVID-19 in our systematic review. Countries represented are as following:

U. S. Britain Italy Canada China Australia Germany Ireland, Saudi Arabia Spain, Switzerland India Iraq India Mexico Mexico Pakistan South Africa United Arab Emirates

Analysis of these studies indicated the most common complications were neurologic and psychiatric problems, respiratory symptoms, musculoskeletal & rheumatologic diseases as well as cardiovascular crises as well as hepatic and gastrointestinal failures

Methods of care have not been found to be very efficacious for permanently resolving these symptoms, although the severity of symptoms tends to decrease over time [1].

## DISCUSSION

The far-reaching physiological systems of the human body from various late complications developed in response to COVID-19 infection. This full-range umbrella review has classified the residual sequelae of long COVID-19 as 9 different classes. In the following conversation, I will discuss all of these delineated categories one by one.

Neurological and psychiatric abnormalities

Briefly, delayed neurological and psychiatric sequelae triggered by COVID-19 are critical because of their high prevalence among the terminologies of this disease. In the long-COVID patients fatigue is the most frequent symptom.

Analysis of 16 studies with nearly 9000 COVID-19 patients suggests the presence of post-acute COVID-19 syndromes that persist beyond three months after the initial infection. The ongoing effects included psychiatric symptoms (like fatigue, cognitive dysfunction (memory lapses and attention deficits) and sleep disturbances). Also, converging themes across different studies pointed to symptoms of depression-PTSD (post-traumatic stress disorder), anxiety, wrath and mood swings et al.

Not surprisingly, there was no straightforward association between the acute COVID-19 severity and worse symptom burden in the post-acute phase of the disease. Some studies reported neurological manifestations like headaches, myelitis, neuropathies, paresthesias, parksonian features, cogwheel rigidity, optic neuritis, distorted olfactory perceptions, encephalitis, epilepsy, Bell's Palsy and myoclonus. 7,40 13,46,49,50

## **Respiratory Implications**

Pulmonary Hypertension: The lungs remain the predominant target organ of severe COVID-19 infection in its acute phase; the same is true for long-term consequences from long COVID-19. A number of patients have described symptoms like dyspnea, chronic cough, thoracic constraint into post-COVID phase that is prolonged; longer in time I guess. Dyspnea is generally made worse with exertion. Despite treatments showing scant efficacy at symptom relief, the severity of these tend to fade over time.

Musculoskeletal and Rheumatology Sciences

Reports from the studies indicated long COVID-19 patients may have musculoskeletal manifestations, joint effusion and persistent pain. Some patients even have small or medium and large vasculitis in long COVID-19. Mediastinal involvement (inflammatory myopathies) Systemic lupus erythematosus (SLE) sarcoidosis; arthritis etc. Cutaneous vasculitis after COVID-19 is usually refractory and tends to clear with time. (14, 29, 54)

The Hepatobiliary and GI

Research suggested that long COVID-19 s have higher risk of GI symptoms (tingling in mouth or anorexia, dyspepsia, constipation, taste impairment, IBS [defined now as long COVID]), loss of appetite and abdominal pain, diarrhea and nausea/vomiting. (57)

#### Ocular:

Remythologic disorders have been rarely reported in long COVID-19, with Ophthalmologic Problems being conjunctivitis, dry eye / difficulty seeing/ blurred vision, photophobia, sore eyes/eye pain 54,59

Thrombosis or Embolism and Stroke:

Alinaghi et al., SeyedAlinaghi et al.; (40) Reviewed 65 studies refuting that venous/arterial thrombosis and cardiac/brain stroke maybe observed in long COVID- disease..

Others:

Other infrequent and extremely late complications of COVID-19 include sputum/nasal congestion, hyperhidrosis, rhinorrhea cough myalgia arthralgias body weight changes otalgia sore throat altered heart rates dysphonia fever palpitations hair loss dysphagiastuttering speech comparable reduction in proteins and menstrual problems etc.

#### **Conclusion:**

Characteristics of the research suggests that COVID-19 patients may encounter late and prolonged manifestations demonstrating "long Covid" in general other symptoms also include described above, often lasting for more than 6 months and difficult to manage though can improve gradually over time. Common Long COVID sequelae are fatigue and other neuropsychiatric symptoms; respiratory, dyspnea,cough and chest tightness are the most frequent long-term complications of COVID-19.

Because these complications are persistent and delayed, healthcare providers, as well as patients, need to know what signs and symptoms to look for.

First Auth oran dCou ntry	Complication Assessment methods	Time of late complicationons et	Time of recovery from mentioned Seque	Late complications
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Qiuyue Long, China	Complications were categorized using systematic review and meta-analysis methodology. Post-discharge symptoms in the ICU as well as discharge with acute post-acute care and pulmonary function tests among discharged COVID-19 patients.	Late complication onset occurred during time of recovery from acute phase of COVID-19 as patients. Follow-up at least 1 months after discharge or more than more 2 weeks post admission to	Recovery time from the sequelae or late complications of above sequel was variable, different consequents. Results of the study being that due to it being a post-acute COVID-19 syndrome focus clearly seems to be that some symptoms	It explored the persistence of symptoms and signs throughout the post-acute phase, suggestive of longer-term complications or sequelae that negatively impact the long-term health of COVID-19 patients. Late complications listed in the study include fatigue or weakness,
		identify and classify late complications in the study.	wont resolve for weeks, if not months in some patients even after their initial discharge from hospitalsetupeрь	psychosocial symptoms such as anxiety or depression, memory impairment and dyspnea, joint pain, reduced diffusion capacity and lung volume measurement, among others.
Taghre ed Shams , KSA	This study is a cross-sectional comparison study of pregnancy outcomes between symptomatic and asymptomatic COVID-19 infected pregnant women during the COVID 19 pandemic in Saudi Arabia. Maternal, perinatal, and neonatal outcomes were assessed by symptoms, ARI score, age grips and other variables.	The document does not follow traditional definitions of the timing of late complications. It does mention complications like maternal death from respiratory failure, and also lists neonatal deaths that happened during the time the studies were done.	Details are missing on the timing of recovery from the described sequelae. Recovery from complications such as pneumonia, ICU admissions, and other maternal and fetal complications would depend on each specific case and the severity of the condition.	The document includes late complications such as maternal death (in respiratory failure), maternal pneumonia, neonatal deaths, preterm deliveries, cesarean deliveries, abortions, intrauterine fetal deaths, low birth weight, and others (maternal/neonatal complications attributable to the COVID-19 infection in pregnancy). All complications emphasise the risks and challenges of COVID-19 in pregnant women, particularly those with symptoms and older age of the mother.
Yudha NurPat ria,Ind onesia	Most studies assessed complications mainly relying on lung function tests in recovered COVID-19 patients.	Late complications were evaluated at different time points from 14 days post discharge from the hospital up to 10 weeks following rehabilitation.	Recovery from the reported sequelae (eg, impaired pulmonary function) was variable. Some had spontaneous improvement of lung function with time, and others improved only with rehabilitative measures.	Late complications in previously ill COVID-19 patients were reported including diffusion and restrictive lung impairments which may linger for weeks to months after recovery. Such complications can impact the patients' long-term lung function.
Pietro De Luca,	he complications were appraised through a systematic review of	Because the study mostly concentrated on ongoing	They also mention the need for longer follow up studies to	The reviewed studies reported persistent hearing or chemosensory

Italy	literature data on audiovestibular symptoms and loss of olfactory/gustatory function after SARS- CoV-2 infection.	audiovestibular and chemosensory impairments due to COVID-19	clarify the long term effects of auditory, vestibular, olfactory, and gustatory dysfunction after COVID-19.	impairments associated with SARS-CoV-2 infection. Although the prevalence of persistent audiovestibular symptoms varied between articles, olfactory and gustatory disturbances were reported more uniformly. Longer-term follow-up studies are therefore required to fully assess the long-term consequences of these conditions.
Premra j L	A meta-analysis was probably conducted in this investigation to evaluate mid and long-term neurological and neuropsychiatric sequelae from post-COVID-19 syndrome.	The article may have also looked at the timing of the development of late neurological and neuropsychiatric sequelae in people with post-COVID-19 syndrome.	They might have studied recovery time from the neurological and neuropsychiatric manifestations of post-COVID-19 syndrome3.	Through meta-analysis, the study presumably intended to investigate and evaluate the mid and bi long-term neurological and neuropsychiatric complications shown in those with post-COVID-19 syndrome.
Jasmin e Turna, Canad a	urveys and validated scales for Generalized Anxiety Disorder-7 (GAD-7), Patient Health Questionnaire-9 (PHQ-9), and Perceived Stress Scale (PSS-10).	was not clearly stated in the provided document.	In the document, the time of recovery from sequelae was not stated.	The text you provided was an excerpt from that document, it did not specifically mention late complications.
WalaaE Inaiem, Sudan	The complication assessment in the case report was based on cadaveric dissection demonstrating a second brachial artery.	Secondary complications that occurred later from the presence of accessory brachial artery were noted at the time of cadaveric dissection consistent with congenital pathoanatomy.	Since the case report is based on an anatomical observation made in a cadaver, the duration of recovery from the abovementioned sequelae is not relevant. It was mainly focused on describing the anatomical variation achieved.	Late complications of accessory brachial artery include difficulties in performing angiographic or coronary catheterization, venepunction, traumatic injuries, and arm or forearm fractures management. These variations are important for clinicians to appreciate especially in radiology and surgery.
S.A. Behnoo d , United Kingdo m	It was not aimed at specific complication/cross organ assessment methodology irrespective of any one complication or organ, but about persistent symptoms post SARS-CoV-2 infection in	Document did not state late complication occurred in a specific time.	Details on the timing of recovery from the sequelae (long symptoms post-SARS-CoV-2 infection among children and adolescents) are not included in the document.	The document focused mainly on the long-lasting COVID-19 symptoms of children and young people that occurred after infection with SARS-CoV-2. The study did not focus on specific late complications on top of

Said cardi ,Italy reson (CMI) echoo electr (ECC) biom to seque from infec  Matsuo Meth So , was USA. syste meta- evalu and seque	niques including iac magnetic mance imaging IR), pocardiography, trocardiogram G), and cardiac markers were used evaluate cardiac relae after recovery a COVID-19 ction in this study.  The study designed as a rematic review and a-analysis to uate radiological functional lung ielae among	This document did not specify exactly when late complication occurs after recovery from COVID-19, though  They used chest computed tomography (CT) and pulmonary function tests	It has not been specified on how long it takes to recover from the above-mentioned sequelae after COVID-19 recovery.  Late complications were evaluated at approximately 90	the study identified residual de novo cardiac injury following recovery from COVID-19 (particularly subclinical myocardial injury in the early phase, diastolic dysfunction in the medium term) and risk of heart failure, arrhythmias and myocardial infarction in COVID-19 survivors.  Late complications consisted of residual abnormalities seen on
So , was USA. syste meta- evalu and seque	designed as a ematic review and a-analysis to uate radiological functional lunguelae among	computed tomography (CT) and pulmonary	were evaluated at approximately 90	consisted of residual
	VID-19 patients.	(PFT)data of discharged patients for data collection and analysis. The follow-up time after recovery of COVID 19 was from 1-6 months.	days post symptom onset or hospital discharge. The summary did not provide an explicit recovery time from the aforementioned sequelae. The average follow-up time for evaluation of residual abnormalities was approximately 90 days after symptom onset or discharge.	abnormalities seen on chest CT and PFT results. Such sequelae highlighted evidence of persistent lung abnormalities including ground glass opacity, parenchymal band or fibrous stripe, impaired diffusion capacity, restrictive and obstructive patterns of lung function even 3 months post-COVID-19 recovery. More long-term studies are needed to understand the endurance and significance of these complications and their sequelae in patients.
dez-de- las- survi Peñas,S 19, pain. perfo analy from datab serve colled who hospi	bases and preprint ers. This data were ected from patients were either oitalized or non- oitalized and vering from	The study investigated post-COVID headache prevalence across multiple time points (at symptom onset/hospital admission, at 30 days, 60 days, 90 days, and >180 days after either symptom onset or hospital discharge).	The time it took the mentioned sequelae to resolve was not specifically mentioned in the summary. The prevalence of post-COVID headache decreased in both acute onset and follow-up periods, then plateaued until 180 days after infection.	Late complications are represented by post-COVID headache, with prevalence within 6 months post-SARS-CoV-2-infection ranging from 8% to 15%. The prevalence was higher during hospitalization and decreased over time, but remained stable after the acute phase. The study underscored the need for post-COVID headache risk factor identification and appropriate management in this population.

Omer, Iraq	complication was performed with a systematic review of all published studies using databases including Web of Science, PubMed, MEDLINE on OVID, and Google Scholar. The selection criteria targeted the articles which confirmed SARS-CoV-2 infection and reported neurological disorders that developed post-recovery.	from COVID-19 infection to neurological sequelae was 33.2 days (range 8-130 days)	alluding to a specific time period of recovery from the aforementioned sequelae, as this will vary from person to person.	found are:  Guillain-Barre Syndrome (GBS)  Transverse Myelitis  Stroke Critical Illness Encephalopathy Optic Neuritis Status Epilepticus Bell's Palsy Vestibular Neuritis Opsoclonus Myoclonus Syndrome Myopathy Parkinsonism
James B. Badeno ch, UK.	The study used observational methods, such as patient self-reports, clinical interviews, and standardized cognitive tests for the assessment of neuropsychiatric symptoms.	Complications were evaluated post-acute, generally beginning from 4 weeks post COVID-19 infection.	Follow-up was a mean of 77 days (range, 14 to 182 days) after infection.	The most common late complications were sleep disturbances (27.4%), fatigue (24.4%), cognitive impairment (20.2%), anxiety (19.1%), and post-traumatic stress symptoms (15.7%).
Bandar ,Saudi Arabia	Cardiac magnetic resonance imaging (CMR), echocardiography, electrocardiograms (ECG), and cardiac enzyme tests (e.g., troponin levels) were used to assess complications.	Late complications were generally assessed in the post-recovery phase (11-194 days post COVID-19 diagnosis in some studies).	Recovery from cardiovascular complications individuals recommended for re-evaluation four weeks post-infection, or longer, particularly when myocarditis was diagnosed.	Late sequelae determined were diagnosis of myocarditis, pericarditis, pericardial effusion, and elevated troponin levels, with a heterogeneous rate of prevalence across the studies included.
QuinHe aley,U K	The study searched longitudinal cohort studies from January 2020 to July 2021, which explored adults with long COVID at least four weeks post-acute infection. Risk of bias was assessed using the Joanna Briggs Institute checklist for cohort studies.	The follow-up periods considered in these studies varied from 30 days to 340 days. Chronic consequences of long COVID were evaluated at least four weeks following acute infection, with outcomes including fatigue, dyspnoea, olfactory dysfunction, myalgia, cough, and	No data were given on recovery time of the cited sequela. But it stressed that long COVID symptoms persisted after the acute phase of infection, with varying prevalence rates over time until 340 days after being infected.	The late complications of long COVID were long symptomatic manifestations such as tiredness, dyspnoea, olfactory dysfunction, myalgia, cough, and gustatory dysfunction. A considerable percentage of patients experienced these symptoms at least four weeks after getting infected with acute SARS-CoV-2 — evidence of the challenge that the disease presents to all patients.

	gustatory dysfunction.	

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