

The Post-acute COVID-19 Syndrome (Long COVID)

Issa Al-Jahdhami^{1*}, Khalid Al-Naamani¹ and Adhra Al-Mawali²

¹Department of Medicine, Armed Forces Hospital, Muscat, Oman

²Centre of Studies and Research, Ministry of Health, Muscat, Oman

ARTICLE INFO

Article history:

Received: 20 November 2020

Accepted: 22 November 2020

Online:

DOI 10.5001/omj.2021.91

The first case of COVID-19 was reported in Wuhan, China, in December 2019.¹ COVID-19 is caused by a novel coronavirus, named severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2 or 2019-nCoV). As of 15 November 2020, more than 54 million people have been infected, and more than one million deaths have been reported.² In Oman, as of 15 November 2020, 118 000 cases have been reported, with 1338 deaths.² The first two cases were diagnosed on 24 February 2020, in Muscat governorate.³ The acute presentation of a COVID-19 infected patient has been well described in various studies.⁴ The majority of patients presented with a fever, sore throat, cough, shortness of breath, and chest pain. Many papers have described multi-organ involvement.⁵ The acute illness is mild in the majority of the patients. Even so, around 20% of those infected need hospitalization, and around 5% require critical care with non-invasive or mechanical ventilation.⁶

There is a misconception that all patients with COVID-19 may recover within two weeks; this is not always the case. The long-term consequences of COVID-19 infection are not well understood. In addition, prolonged recovery of symptoms has been described even in patients who had mild symptoms and did not require hospitalization.^{7,8} This manifestation was termed post-acute COVID-19 syndrome or 'long COVID'.⁹ This editorial aims to explore post-acute COVID-19 syndrome or long COVID.

There is no clear definition of post-acute COVID-19 syndrome. In general, it is an illness described among patients who have recovered from COVID-19 but still have ongoing symptoms or among those who continued to have symptoms

for longer than normally expected.¹⁰ Some authors have suggested the presence of symptoms beyond 12 weeks from the onset of illness as a description of post-acute COVID-19 syndrome.⁹ Other studies have divided these patients into three groups: those who had severe manifestations such as acute respiratory distress syndrome (ARDS), requiring intensive care unit (ICU) admission; those who were not admitted during the acute illness but later presented with symptoms and signs of end-organ damage, such as cardiac or respiratory disease; and those who did not require hospitalization but presented with prolonged symptoms without evidence of end-organ damage.¹¹ It is interesting to note that post-acute COVID-19 syndrome is more common in women.

The exact mechanism of this post-COVID-19 presentation is obscure. Previous investigators suggested low antibody response to SARS-CoV-2 infection, prolonged inflammatory response to the SARS-CoV-2 infection, deconditioning, and reinfection with SARS-CoV-2 as possible mechanisms that might explain post-COVID-19 presentation.⁷

There is a marked variation in the presentation of post-acute COVID-19 syndrome. Patients may present with non-specific symptoms such as fatigue, muscle aches and pains, poor sleep, cough, and breathlessness, to more specific organ-related symptoms, such as orthopnea, leg swelling, and exercise intolerance due to COVID-19 induced heart failure.⁷ Furthermore, chest pain and significant breathlessness might be due to pulmonary embolism.¹² Autonomic symptoms such as palpitations with mild exertion, night sweats, and poor temperature control were also described.¹² The symptoms might be cyclical in some patients.⁸

Based on our experience of following-up COVID-19 patients discharged from the hospital, fatigue appears as a prominent feature, even at 12 weeks post-discharge. Patients assert that they were unable to return to their baseline activity level. They find trivial daily activities fatigue-inducing. This is a major complaint that has been acknowledged by the literature.⁴ Another prominent feature of long COVID experienced by our patients is shortness of breath. This was more significant among patients with severe COVID-19 who were admitted to ICU and required either non-invasive or mechanical ventilation. Further investigation revealed mild anemia in some patients, evidence of pulmonary embolism in some, and changes suggestive of pulmonary fibrosis in computer tomography of the chest among others. Cardiac causes such as heart failure with preserved ejection fraction or impaired ventricular function were found in a small number of patients. However, there was no cause identifiable in the majority of patients.

Post-acute COVID-19 syndrome management remains a clinical challenge as there are no evidence-based international guidelines to follow at the time of writing. Pulmonary embolism is managed in the usual way with anticoagulation for at least three months. There is no consensus on the benefit, or duration, of prolonged prophylaxis, with low molecular weight heparin, post-discharge. Management of post-COVID-19 pulmonary fibrosis is also unclear. Clinical, radiological, and biochemical markers are required to help clinicians anticipate which patients with COVID-19-related ARDS are susceptible to developing pulmonary fibrosis following the resolution of COVID-19.¹³ We have used systemic steroids in some patients with good results. The role of the antifibrotic drugs, pirfenidone and nintedanib, is awaiting clinical trial evidence.

A unified definition of long COVID and characterization of its manifestation are important for early detection. In addition, more research should be directed to identify risk factors and exact mechanisms leading to the development of long COVID syndrome. Such knowledge may enhance further research aimed at the prevention of such a complication.

In conclusion, a large number of patients have been affected by COVID-19, and as physicians, we are going to face many patients with symptoms suggestive of long COVID. The entire spectrum of long COVID is not well characterized. Further research looking at risk factors, mechanisms explaining development of long COVID, and preventative measures are imperative to prevent such a complication.

REFERENCES

1. Zhou P, Yang X-L, Wang X-G, Hu B, Zhang L, Zhang W, et al. A pneumonia outbreak associated with a new coronavirus of probable bat origin. *nature*. 2020;579(7798):270-273.
2. WHO Coronavirus Disease (COVID-19) Dashboard [Internet]. [cited 2020 November 16]. Available from: <https://covid19.who.int>.
3. Khamis F, Al Rashidi B, Al-Zakwani I, Al Wahaibi AH, Al Awaidy ST. Epidemiology of COVID-19 infection in Oman: analysis of the first 1304 cases. *Oman Med J* 2020 Jun;35(3):e145.
4. Shi S, Qin M, Shen B, Cai Y, Liu T, Yang F, et al. Association of cardiac injury with mortality in hospitalized patients with COVID-19 in Wuhan, China. *JAMA Cardiol* 2020; 5(7):802-810
5. Gavriatopoulou M, Korompoki E, Fotiou D, Ntanasis-Stathopoulos I, Psaltopoulou T, Kastritis E, et al. Organ-specific manifestations of COVID-19 infection. *Clin Exp Med* 2020 Nov;20(4):493-506.
6. Wu Z, McGoogan JM. Characteristics of and important lessons from the coronavirus disease 2019 (COVID-19) outbreak in China: Summary of a report of 72 314 cases from the chinese center for disease control and prevention. *JAMA* 2020 Apr;323(13):1239-1242.
7. Outhoff K. Sick and tired of COVID-19: long haulers and post viral (fatigue) syndromes. *South Afr Gen Pract J*. 2020;1(4):132-133.
8. Nath A. Long-Haul COVID. *AAN Enterprises*; 2020.
9. Greenhalgh T, Knight M, A'Court C, Buxton M, Husain L. Management of post-acute COVID-19 in primary care. *BMJ* 2020;370:m3026.
10. Mahase E. Covid-19: What do we know about "long COVID"? *BMJ* 2020;370:m2815.
11. Ladds E, Rushforth A, Wieringa S, Taylor S, Rayner C, Husain L, et al. Persistent symptoms after COVID-19: qualitative study of 114 long COVID patients and draft quality criteria for services. *BMC Health Services Research* (2020) 20:1144.
12. Butler M, Pollak TA, Rooney AG, Michael BD, Nicholson TR. Neuropsychiatric complications of COVID-19. *British Medical Journal Publishing Group*; 2020;7(10):875-882.
13. Vasarmidi E, Tsitoura E, Spandidos DA, Tzanakis N, Antoniou KM. Pulmonary fibrosis in the aftermath of the COVID-19 era (Review). *Exp Ther Med* 2020 Sep;20(3):2557-2560.