



“Post- COVID-19 syndrome”: Rising from the dead?

Arash Eatemadi ¹, Ramin Jamshidian ², Mostafa Gouya ³, Alireza Moradzadegan ⁴

¹ Infectious disease specialist, Department of General Medicine, Suhar hospital, Suhar, Sultanate of Oman, corresponding author, dr_arash53@yahoo.com

² Infectious disease specialist, Department of Infectious disease, Sina hospital, Iran

³ General practitioner, Al Hayat hospital, Muscat, Sulatante of Oman

⁴ Medicine intern, Jundishapur university of medical sciences, Ahwaz, Iran

Abstract

A significant percentage of critically ill COVID-19 patients after recovery of acute phase of the disease, will face physical, cognitive and psychological consequences which stand for long period and are troublesome. These heterogenous signs and symptoms are constitute “post-COVID syndrome”. This syndrome has no definite pathophysiology and the main compliant is fatigue. The most important differential diagnosis of this syndrome is, “post-intensive care syndrome”. There are increasing reports about “post-COVID-19 syndrome” worldwide which means it is a considerable consequence of COVID-19, especially severe types.

Key words: COVID-19, post-COVID-19 syndrome, post- intensive care syndrome

Irrespective of COVID-19 severity, a large proportion of patients suffer from persistent wide range of symptoms usually up to 6 months after acute phase of infection. Currently, there is no consensus definition for this condition but usually is mentioned as “Post-COVID syndrome” (1-3)

“Post- covid-19 syndrome” sounds to be a multisystem disorder, (4) which presents with one or constellation of symptoms such as, cough, low grade fever, fatigue, shortness of breath, chest pain, headache, neurocognitive difficulties, body pain, fatigue, weakness, gastrointestinal upset, metabolic disruption (such as uncontrolled diabetes), thromboembolic events, depression and other mental health conditions. (5,6)

An important differentiation should be made between symptoms of chronic inflammation (convalescent phase), sequelae of organs damage (such as pulmonary fibrosis and chronic kidney disease), and nonspecific problems about hospitalization and social issues. (7,8) Persistent viremia, reinfection,

chronic inflammation, and mental disorder all could be responsible in the pathogenesis of “Post-COVID-19 syndrome”. (9-14)

For better understanding, it is wise to evaluate only mild COVID patients. because this group is unlikely to have chronic organ dysfunctions. In evaluation of the causes of fatigue in these patients, some correctable etiologies have been found including, anemia, vitamin D deficiency, hypothyroidism and cortisol insufficiency. (15,16)

Conversely, patients with severe forms of COVID-19 often progress toward acute respiratory distress syndrome (ARDS) which cause permanent lung parenchyma damage, including lung fibrosis. (17) In addition, Patients experiencing post intensive care syndrome (PICS) generally report higher incidences of cognitive and physical dysfunctions, which often persist long-term. (18)

Patients with COVID-19 might be not receive the appropriate physiotherapy during admission due to fear about COVID-19 transmission or staff shortage, resulting in following considerable disability. (19) There are a few studies in medical literature about rate of post-COVID-19 symptoms:

Claudia Carvalho-Schneider et al. conducted a descriptive clinical follow-up study of 150 non-critical patients with COVID-19 and watched that two-thirds of adults experienced persistent symptoms up to 2 months after symptom onset, primarily anosmia/ageusia, dyspnea or asthenia. (20)

Similarly, Eve Garrigues et al. studied patients with COVID-19 discharged from the hospital and interviewed over 3 months post diagnosis and showed the majority of patients experienced continued symptoms, most commonly including fatigue and dyspnea. (21)

In another study by Angelo Carfi et al. patients with COVID-19 discharged from the hospital with SARS-CoV-2 RNA clearance by RT-PCR and interviewed approximately 2 months after diagnosis, the majority of patients experienced continued symptoms, with the most common symptoms being fatigue and dyspnea. (22)

Also, Halpin et al. conducted a study on patients with COVID-19 discharged from the hospital at least 4 weeks prior to study enrollment, the majority of patients experienced continued symptoms, with the most common symptoms being fatigue and dyspnea. (23)

Finally, In the study of Tenforde, patients who were diagnosed with COVID-19 in the outpatient setting, primarily had mild disease and were interviewed a median of 16 days post diagnosis, the majority had continued symptoms. The most common symptoms included cough and fatigue. (24)

Survived COVID-19 patients, have just escaped from the first stage (25) and they are predisposed to long-term physical and psychological dysfunctions. Long-term psychological problems and post-traumatic stress disorder (PTSD) can develop in more than half of survived patients. (26) follow up and management of these patients warrants multidisciplinary team at least including, internal medicine physician, physiotherapist, occupational therapist and psychologist (27,28)

In conclusion, more studies are needed to better understand the “Post-COVID syndrome”. Clinicians should update patients about the potential long-term consequences of COVID-19 prior to discharge from the hospital.

Conflicts of interest

The authors declare no competing interests

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References

- 1- Yvonne M.J. Goërtz, et al. Persistent symptoms 3 months after a SARS-CoV-2 infection: the post-COVID-19 syndrome? ERJ Open Research 2020 6: 00542-2020; DOI: 10.1183/23120541.00542-2020
- 2- Itsuro Kazama. Stabilizing mast cells by commonly used drugs: a novel therapeutic target to relieve post-COVID syndrome? Drug Discov Ther. 2020 Nov 4;14(5):259-261. doi: 10.5582/ddt.2020.03095. Epub 2020 Oct 29
- 3- COVID Symptom Study. How long does COVID-19 last? Kings College London, 2020. https://covid19.joinzoe.com/post/covid-long-term?fbclid=IwAR1RxIcmmdL-EFjh_al
- 4- Geddes L. Why strange and debilitating coronavirus symptoms can last for months. New Scientist 2020. <https://www.newscientist.com/article/mg24632881-400-why-strange-and-debilitating-coronavirus-symptoms-can-last-for-months>
- 5- Assaf G, Davis H, McCorkell L, et al. An analysis of the prolonged COVID-19 symptoms survey by Patient-Led Research Team. Patient Led Research, 2020. <https://patientresearchcovid19.com/>.
- 6- Dasgupta A, Kalhan A, Kalra S. Long term complications and rehabilitation of COVID-19 patients. J Pak Med Assoc 2020; 70: S131-5. doi:10.5455/JPMA.32 pmid:32515393
- 7- Trisha Greenhalgh, Matthew Knight, Christine A'Court, Maria Buxton, Laiba Husain. Management of post-acute covid-19 in primary care. BMJ 2020; 370 doi: <https://doi.org/10.1136/bmj.m3026>
- 8- Bernd Lamprecht. Is there a post-COVID syndrome? Pneumologe (Berl). 2020 Oct 8;1-4. doi: 10.1007/s10405-020-00347-0.
- 9- Wu F, Wang A, Liu M, et al. Neutralizing antibody responses to SARS-CoV-2 in a COVID-19 recovered patient cohort and their implications. 2020. <https://www.medrxiv.org/content/medrxiv/early/2020/04/06/2020.03.30.20047365.full.pdf>
- 10- Lan L, Xu D, Ye G, et al. Positive RT-PCR test results in patients recovered from COVID-19. JAMA 2020; 323:1502-3. doi:10.1001/jama.2020.2783 pmid:32105304
- 11 - Colafrancesco S, Alessandri C, Conti F, Priori R. COVID-19 gone bad: A new character in the spectrum of the hyperferritinemic syndrome? Autoimmun Rev 2020;19:102573. doi: 10.1016/j.autrev.2020.102573 pmid:32387470
- 12- Tay MZ, Poh CM, Rénia L, MacAry PA, Ng LFP. The trinity of COVID-19: immunity, inflammation and intervention. Nat Rev Immunol 2020; 20:363-74. doi:10.1038/s41577-020-0311-8 pmid:32346093

- 13-Forte G, Favieri F, Tambelli R, Casagrande M. COVID-19 pandemic in the Italian population: validation of a post-traumatic stress disorder questionnaire and prevalence of PTSD symptomatology. *Int J Environ Res Public Health* 2020; 17:4151. doi:10.3390/ijerph17114151 pmid:32532077
- 14- Jiang H-j, Nan J, Lv Z-y, et al. Psychological impacts of the COVID-19 epidemic on Chinese people: Exposure, post-traumatic stress symptom, and emotion regulation. *Asian Pac J Trop Med* 2020; 13:252.
- 15- Perna Garg, Umang Arora, Arvind Kumar, Naveet Wig. The "post-COVID" syndrome: How deep is the damage? *J Med Virol.* 2020 Aug 27;10.1002/jmv.26465. doi: 10.1002/jmv.26465
- 16- Chen M, Zhou W, Xu W. Thyroid function analysis in 50 patients with COVID-19: a retrospective study [published online ahead of print July 10, 2020]. *Thyroid.* 2020. <https://doi.org/10.1089/thy.2020.0363>
- 17- Thompson BT, Chambers RC, Liu KD Review Acute Respiratory Distress Syndrome. *N Engl J Med.* 2017 Aug 10; 377(6):562-572.)
- 18- Pandharipande PP, Girard TD, Jackson JC, Morandi A, Thompson JL, Pun BT, Brummel NE, Hughes CG, Vasilevskis EE, Shintani AK, Moons KG, Geevarghese SK, Canonico A, Hopkins RO, Bernard GR, Dittus RS, Ely EW, BRAIN-ICU Study Investigators. Long-term cognitive impairment after critical illness. *N Engl J Med.* 2013 Oct 3; 369(14):1306-16.
- 19- David H. Jiang, Rozalina G. McCoy. Planning for the Post-COVID Syndrome: How Payers Can Mitigate Long-Term Complications of the Pandemic. *J Gen Intern Med.* 2020 Oct; 35(10): 3036–3039. doi: 10.1007/s11606-020-06042-3
- 20- Claudia Carvalho-Schneider, et al. Follow-up of adults with noncritical COVID-19 two months after symptom onset. *Clin Microb Inf.* Published: October 05, 2020 DOI: <https://doi.org/10.1016/j.cmi.2020.09.052>
- 21-Eve Garrigues, et al. Post-discharge persistent symptoms and health-related quality of life after hospitalization for COVID-19. *J Inf.* Published: August 24, 2020; DOI: <https://doi.org/10.1016/j.jinf.2020.08.029>
- 22- Angelo Carfi, et al. Persistent Symptoms in Patients After Acute COVID-19. *JAMA.* 2020;324(6):603-605. doi:10.1001/jama.2020.12603
- 23- Stephen J. Halpin, et al. Postdischarge symptoms and rehabilitation needs in survivors of COVID-19 infection: A cross-sectional evaluation. *J Med Virol.* 30 July 2020 <https://doi.org/10.1002/jmv.26368>
- 24- Mark W. Tenforde, et al. Symptom Duration and Risk Factors for Delayed Return to Usual Health Among Outpatients with COVID-19 in a Multistate Health Care Systems Network — United States, March–June 2020. *Morbidity and Mortality Weekly Report (MMWR).* Weekly / July 31, 2020 / 69(30);993-998
- 25- Ohtake PJ, Lee AC, Scott JC, Hinman RS, Ali NA, Hinkson CR, Needham DM, Shutter L, Smith-Gabai H, Spires MC, Thiele A, Wiencek C, Smith JM. Physical Impairments Associated with Post-Intensive Care Syndrome: Systematic Review Based on the World Health Organization's International Classification of Functioning, Disability and Health Framework. *Phys Ther.* 2018 Aug 1; 98(8):631-645.

26- Hatch R, Young D, Barber V, Griffiths J, Harrison DA, Watkinson P. Anxiety, Depression and Post Traumatic Stress Disorder after critical illness: a UK-wide prospective cohort study. *Crit Care*. 2018; 22:310. doi: 10.1186/s13054-018-2223-6

27- Gemelli Against COVID-19 Post-Acute Care Study Group. Post-COVID-19 global health strategies: the need for an interdisciplinary approach. *Aging Clin Exp Res*2020; doi:10.1007/s40520-020-01616-x. pmid:32529595

28- Klok FA, Boon GJAM, Barco S, et al. The Post-COVID-19 Functional Status scale: a tool to measure functional status over time after COVID-19. *Eur Respir J*2020; 56:2001494. doi:10.1183/13993003.01494-2020 pmid:32398306

