



# COVID-19

# Background

Evaluating and Caring for Patients with Post-COVID Conditions: Interim Guidance

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For some patients, recovery from acute SARS-CoV-2 infection may involve continuing, recurrent, or new symptoms and clinical findings that persist for weeks, months, or longer. (1) The term "Post-COVID Conditions," (2) sometimes referred to colloquially as "Long COVID," has been proposed as an umbrella term for the wide range of physical and mental health consequences that are present four or more weeks after SARS-CoV-2 infection. These consequences include both general complications of prolonged illness as well as hospitalization and post-acute sequelae of SARS-CoV-2 infection (PASC), which are more specific to effects of SARS-CoV-2 infection. Persistent symptoms and late sequelae have also been reported among people who were determined to have had asymptomatic infection or who experienced mild acute illness.(3, 4)

At present, robust longitudinal surveillance data on post-COVID conditions are lacking and the prevalence is challenging to estimate. The frequency of long-term symptoms and conditions following SARS-CoV-2 infection varies widely in the literature, ranging from 5% to 80%.(4-11) It has been challenging to create a single universal case definition for post-COVID conditions because studies differ in terms of the symptoms or conditions investigated, the temporal criteria used (three weeks up to many months following SARS-CoV-2 infection), the study settings included (outpatient vs. inpatient), and how symptoms and conditions are assessed (e.g., self-report vs. electronic health record database).(12) Post-COVID conditions have been more commonly reported in women, although it is unknown whether there are biological risk factors for some post-COVID conditions and demographic differences remain unclear. (2) Evidence suggests that post-COVID conditions occur in children and adolescents(13-16) as well as adults, but the true frequency and severity are unknown. Patients with certain characteristics or comorbidities might be at higher risk for post-COVID conditions, although subpopulation prevalence information is limited. Although older patients may have an increased risk for severe acute disease and related ongoing symptoms, younger patients, including those in good health before SARS-CoV-2 infection, have also reported debilitating post-COVID conditions months after acute illness.(1, 17)

Post-COVID conditions are heterogenous and may be attributable to different underlying pathophysiologic processes. Efforts are underway to characterize and differentiate the multiple possible etiologies (for example, organ damage resulting from acute phase infection, complications from a persistent hyperinflammatory state, ongoing viral activity associated with an intra-host viral reservoir, inadequate antibody response, and other potential causes). Factors that may further complicate the presentation of post-COVID conditions include physical deconditioning at baseline or after a

prolonged acute disease course that can be nonspecific to COVID-19, pre-COVID comorbidities (underlying medical conditions), or other physical and mental health consequences of a potentially life-threatening illness with a long or complicated disease course, as well as lifestyle changes due to the COVID-19 pandemic.

Multiple possible onset patterns for post-COVID conditions have been identified that further exemplify their heterogeneity, including, but not limited to: (A) persistent symptoms and conditions that begin at the time of acute COVID-19 illness; (B) new-onset late sequelae following asymptomatic disease or a period of acute symptom relief or remission; or (C) an evolution of symptoms and conditions that include some persistent symptoms (e.g., shortness of breath) with the addition of new symptoms or conditions over time (e.g., cognitive difficulties). Some presentations may share similarities with other post-viral syndromes, such as myalgic encephalomyelitis/chronic fatigue syndrome (ME/CFS), dysautonomia (e.g., postural orthostatic tachycardia syndrome [POTS]), or mast cell activation syndrome (MCAS). Some of these types of conditions were also reported in patients who recovered from severe acute respiratory syndrome (SARS) and Middle East respiratory syndrome (MERS), two other life-threatening illnesses resulting from coronavirus infections.

Post-COVID conditions are associated with a spectrum of physical, social, and psychological consequences, as well as functional limitations that can present substantial challenges to patient wellness and quality of life.<sup>(1, 4, 20)</sup> To inform the interim clinical guidance presented here, the Centers for Disease Control and Prevention (CDC) obtained individual expert perspectives on the evaluation and management of post-COVID conditions. CDC coordinated discussions during March and April 2021 with thirteen U.S. medical professionals with expertise in a range of clinical specialties who care for patients with post-COVID conditions, including:

- Adult and Pediatric Pulmonary Medicine
- Critical Care Medicine
- Infectious Diseases
- Physical Medicine and Rehabilitation
- Neurology
- Psychiatry
- Rheumatology
- Nephrology
- Hematology
- Cardiology

The CDC obtained additional feedback from medical organizations and patient advocacy groups. The information in this interim guidance is based on individual medical expert opinion and the best currently available data. With extensive research underway, evidence-based treatment practices will continue to evolve.

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## References

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- 1. Nalbandian A, Sehgal K, Gupta A, Madhavan MV, et al. Post-acute COVID-19 syndrome. Nat Med. 2021 Apr;27(4):601-615. doi:10.1038/s41591-021-01283-z 
  ☐
- 2. Policy Brief 39 In the Wake of the Pandemic Preparing for Long COVID. Accessed at: https://apps.who.int/iris/bitstream/handle/10665/339629/Policy-brief-39-1997-8073-eng.pdf 
  ☐
- 3. Huang Y, Pinto MD, Borelli JL, et al. COVID Symptoms, Symptom Clusters, and Predictors for Becoming a Long-Hauler: Looking for Clarity in the Haze of the Pandemic. medRxiv. 2021 Mar 5. doi: 10.1101/2021.03.03.21252086 ☑
- 4. Havervall S, Rosell A, Phillipson M, Mangsbo SM, Nilsson P, Hober S, Thålin C. Symptoms and Functional Impairment Assessed 8 Months After Mild COVID-19 Among Health Care Workers. JAMA. 2021 Apr 7. doi:10.1001/jama.2021.5612 ☐
- 5. Office of National Statistics. Prevalence of ongoing symptoms following coronavirus (COVID-19) infection in the UK: 1 April 2021. Accessed at: https://www.ons.gov.uk/peoplepopulationandcommunity/healthandsocialcare/conditionsanddiseases/bulletins/prevalenceofongoingsymptomsfollowingcoronaviruscovid19infectionintheuk/1april2021 
  ☐
- 6. Chevinsky JR, Tao G, Lavery AM, et al. Late conditions diagnosed 1-4 months following an initial COVID-19

- encounter: a matched cohort study using inpatient and outpatient administrative data United States, March 1-June 30, 2020. Clin Infect Dis. 2021 Apr 28. doi: 10.1093/cid/ciab338
- 7. Hernandez-Romieu AC, Leung S, Mbanya A, et al. Health Care Utilization and Clinical Characteristics of Nonhospitalized Adults in an Integrated Health Care System 28-180 Days After COVID-19 Diagnosis Georgia, May 2020-March 2021. MMWR Morb Mortal Wkly Rep. 2021 Apr 30;70(17):644-650. doi: 10.15585/mmwr.mm7017e3 
  ☐
- 8. Lund LC, Hallas J, Nielsen H, Koch A, Mogensen SH, Brun NC, Christiansen CF, Thomsen RW, Pottegård A. Postacute effects of SARS-CoV-2 infection in individuals not requiring hospital admission: a Danish population-based cohort study. Lancet Infect Dis. 2021 May 10. doi:10.1016/S1473-3099(21)00211-5
- 9. Huang C, Huang L, Wang Y, et al. 6-month consequences of COVID-19 in patients discharged from hospital: a cohort study. Lancet. 2021 Jan 16;397(10270):220-232. doi:10.1016/S0140-6736(20)32656-8 ☐
- 10. Pavli A, Theodoridou M, Maltezou HC. Post-COVID syndrome: Incidence, clinical spectrum, and challenges for primary healthcare professionals. Arch Med Res. 2021 May 4. doi:10.1016/j.arcmed.2021.03.010 ☐
- 11. Cabrera Martimbianco AL, Pacheco RL, Bagattini ÂM, et al. Frequency, signs and symptoms, and criteria adopted for long COVID: a systematic review. Int J Clin Pract. 2021 May 11:e14357. doi:10.1111/ijcp.14357 ☑
- 12. Rando HM, Bennett TD, Byrd JB, et al. Challenges in defining Long COVID: Striking differences across literature, Electronic Health Records, and patient-reported information. medRxiv. 2021 Mar 26. doi:10.1101/2021.03.20.21253896 ☑
- 13. Office of National Statistics. Update on long COVID prevalence estimate. Accessed at: https://assets.publishing.service.gov.uk/government/uploads/system/uploads/attachment\_data/file/962830/s10 79-ons-update-on-long-covid-prevalence-estimate.pdf 
  ☐
- 14. Buonsenso D, Munblit D, De Rose C, et al. Preliminary evidence on long COVID in children. Acta Paediatr. 2021 Apr 9. doi:10.1111/apa.15870 ☑
- 15. Say D, Crawford N, McNab S, et al. Post-acute COVID-19 outcomes in children with mild and asymptomatic disease. Lancet Child Adolesc Health. 2021 Apr 20. doi:10.1016/S2352-4642(21)00124-3 ☑
- 16. Osmanov I, Spiridonova E, Bobkova P, et al. Risk factors for long covid in previously hospitalised children using the ISARIC Global follow-up protocol: A prospective cohort study. medRxiv. 2021 Apr 26. doi:10.1101/2021.04.26.21256110 
  ☐
- 17. Assaf G, Davis H, McCorkell L, et al. What does COVID-19 recovery actually look like? An analysis of the prolonged COVID-19 symptoms survey by Patient-Led Research Team. Patient Led Research for COVID-19, 2020. 
  ☐
- 18. Lam MH, Wing YK, Yu MW, et al. Mental morbidities and chronic fatigue in severe acute respiratory syndrome survivors: long-term follow-up. Arch Intern Med. 2009 Dec 14;169(22):2142-7. doi:10.1001/archinternmed.2009.384
- 19. Lee SH, Shin HS, Park HY, et al. Depression as a Mediator of Chronic Fatigue and Post-Traumatic Stress Symptoms in Middle East Respiratory Syndrome Survivors. Psychiatry Investig. 2019 Jan;16(1):59-64. doi:10.30773/pi.2018.10.22.3. ☑
- 20. Lambert N, Survivor Corps, El-Azab SA, et al. COVID-19 Survivors' Reports of the Timing, Duration, and Health Impacts of Post-Acute Sequelae of SARS-CoV-2 (PASC) Infection. medRxiv 2021.03.22.21254026; doi:10.1101/2021.03.22.21254026 ☑
- 21. Colbenson GA, Johnson A, Wilson ME. Post-intensive care syndrome: impact, prevention, and management. Breathe (Sheff). 2019 Jun;15(2):98-101. doi:10.1183/20734735.0013-2019 ☐

- 22. Lavery AM, Preston LE, Ko JY, et al. Characteristics of Hospitalized COVID-19 Patients Discharged and Experiencing Same-Hospital Readmission United States, March-August 2020. MMWR Morb Mortal Wkly Rep. 2020 Nov 13;69(45):1695-1699. doi: 10.15585/mmwr.mm6945e2
- 23. Chopra V, Flanders SA, O'Malley M, et al. Sixty-Day Outcomes Among Patients Hospitalized With COVID-19. Ann Intern Med. 2020 Nov 11. doi: 10.7326/M20-5661 ☐
- 24. Ayoubkhani D, Khunti K, Nafilyan V, et al. Post-covid syndrome in individuals admitted to hospital with covid-19: retrospective cohort study. BMJ. 2021 Mar 31;372:n693. doi:10.1136/bmj.n693 ☑
- 25. Atalla E, Kalligeros M, Giampaolo G, et al. Readmissions among patients with COVID-19. Int J Clin Pract. 2020 Sep 7:e13700. doi:10.1111/jjcp.13700 ☑
- 26. Donnelly JP, Wang XQ, Iwashyna TJ, et al. Readmission and Death After Initial Hospital Discharge Among Patients With COVID-19 in a Large Multihospital System. JAMA. 2021 Jan 19;325(3):304-306. doi:10.1001/jama.2020.21465
- 27. Somani SS, Richter F, Fuster V, et al. Characterization of Patients Who Return to Hospital Following Discharge from Hospitalization for COVID-19. J Gen Intern Med. 2020 Oct;35(10):2838-2844. doi:10.1007/s11606-020-06120-6 ☑
- 28. Jeon WH, Seon JY, Park SY, et al. Analysis of Risk Factors on Readmission Cases of COVID-19 in the Republic of Korea: Using Nationwide Health Claims Data. Int J Environ Res Public Health. 2020 Aug 12;17(16). doi:10.3390/ijerph17165844
- 29. Akinbami LJ, Petersen LR, Sami S, et al. COVID-19 symptoms and SARS-CoV-2 antibody positivity in a large survey of first responders and healthcare personnel, May-July 2020. Clin Infect Dis. 2021 Jan 30. doi:10.1093/cid/ciab080 ☑
- 30. Petersen LR, Sami S, Vuong N, et al. Lack of antibodies to SARS-CoV-2 in a large cohort of previously infected persons. Clin Infect Dis. 2020 Nov 4.
- 31. American Academy of Pediatrics. COVID-19 Interim Guidance: Return to Sports and Physical Activity. Accessed at: https://services.aap.org/en/pages/2019-novel-coronavirus-covid-19-infections/clinical-guidance/covid-19-interim-guidance-return-to-sports/
- 32. Greenhalgh T, Knight M, A'Court C, et al. Management of post-acute covid-19 in primary care. BMJ. 2020;370:m3026. doi:10.1136/bmj.m3026 ☑
- 33. COVID-19 Rapid Guideline: Managing the Long-Term Effects of COVID-19. Accessed at: https://www.nice.org.uk/guidance/NG188 ☑
- 34. Sisó-Almirall A, Brito-Zerón P, Conangla Ferrín L, et al. Long Covid-19: Proposed Primary Care Clinical Guidelines for Diagnosis and Disease Management. Int J Environ Res Public Health. 2021 Apr 20;18(8). doi:10.3390/ijerph18084350
- 35. Parkin A, Davison J, Tarrant R, et al. A Multidisciplinary NHS COVID-19 Service to Manage Post-COVID-19 Syndrome in the Community. J Prim Care Community Health. 2021 Jan-Dec;12:21501327211010994. doi:10.1177/21501327211010994 ☑
- **36.** Brigham E, O'Toole J, Kim SY, et al. The Johns Hopkins Post-Acute COVID-19 Team (PACT): A Multidisciplinary, Collaborative, Ambulatory Framework Supporting COVID-19 Survivors. Am J Med. 2021 Apr;134(4):462-467.e1. doi:10.1016/j.amjmed.2020.12.009 
  ☐
- 37. Santhosh L, Block B, Kim SY, Raju S, Shah RJ, Thakur N, Brigham EP, Parker AM. How I Do It: Rapid Design and Implementation of Post-COVID-19 Clinics. Chest. 2021 Mar 31. doi:10.1016/j.chest.2021.03.044 ☑
- 38. O'Brien H, Tracey MJ, Ottewill C, et al. An integrated multidisciplinary model of COVID-19 recovery care. Ir J Med

- Sci. 2021 May;190(2):461-468. doi:10.1007/s11845-020-02354-9
- 39. Lopez-Leon S, Wegman-Ostrosky T, Perelman C, et al. More Than 50 Long-Term Effects of COVID-19: A Systematic Review and Meta-Analysis. Res Sq. 2021 Mar 1. doi:10.2139/ssrn.3769978 ☐
- **40.** Al-Aly Z, Xie Y, Bowe B. High-dimensional characterization of post-acute sequalae of COVID-19. Nature. 2021 Apr 22. doi:10.1038/s41586-021-03553-9 ☑
- 41. Sudre CH, Murray B, Varsavsky T, et al. Attributes and predictors of long COVID. Nat Med. 2021 Apr;27(4):626-631. doi:10.1038/s41591-021-01292-y ☐
- 42. Lund LC, Hallas J, Nielsen H, et al. Post-acute effects of SARS-CoV-2 infection in individuals not requiring hospital admission: a Danish population-based cohort study. Lancet Infect Dis. 2021 May 10. doi:10.1016/S1473-3099(21)00211-5
- 43. Carfi A, Bernabei R, Landi F, et al. Persistent Symptoms in Patients After Acute COVID-19. JAMA. 2020 Aug 11;324(6):603-605. doi:10.1001/jama.2020.12603 ☑
- 44. Cellai M, O'Keefe JB. Characterization of Prolonged COVID-19 Symptoms in an Outpatient Telemedicine Clinic. Open Forum Infect Dis. 2020 Oct;7(10):ofaa420. doi:10.1093/ofid/ofaa420 
  ☐
- **45.** Logue JK, Franko NM, McCulloch DJ, et al. Sequelae in Adults at 6 Months After COVID-19 Infection. JAMA Netw Open. 2021 Feb 1;4(2):e210830. doi:10.1001/jamanetworkopen.2021.0830 ☐
- **46.** del Rio C, Collins LF, Malani P. Long-term health consequences of COVID-19. JAMA. 2020. doi:10.1001/jama.2020.19719 ☑
- 47. Taquet M, Geddes JR, Husain M, et al. 6-month neurological and psychiatric outcomes in 236379 survivors of COVID-19: a retrospective cohort study using electronic health records. Lancet Psychiatry. 2021 May;8(5):416-427. doi:10.1016/S2215-0366(21)00084-5 ☑
- **48.** Barker-Davies RM, O'Sullivan O, Senaratne KPP, et al. The Stanford Hall consensus statement for post-COVID-19 rehabilitation. Br J Sports Med. 2020 Aug;54(16):949-959. doi:10.1136/bjsports-2020-102596 ☑
- 49. Li Z, Zheng C, Duan C, et al. Rehabilitation needs of the first cohort of post-acute COVID-19 patients in Hubei, China. Eur J Phys Rehabil Med. 2020 Jun;56(3):339-344. doi: 10.23736/S1973-9087.20.06298-X. PMID: 32672029. doi:10.23736/s1973-9087.20.06298-x
- 51. Berger Z, Altiery DE Jesus V, Assoumou SA, et al. Long COVID and Health Inequities: The Role of Primary Care. Milbank Q. 2021 Mar 30. doi:10.1111/1468-0009.12505 ☐
- 52. Waltenburg MA, Victoroff T, Rose CE, et al. Update: COVID-19 Among Workers in Meat and Poultry Processing Facilities United States, April–May 2020. MMWR Morb Mortal Wkly Rep 2020;69:887-892. Accessed at: https://www.cdc.gov/mmwr/volumes/69/wr/mm6927e2.htm
- 53. DT Arnold, A Milne, E Samms, et al. Are vaccines safe in patients with Long COVID? A prospective observational study. medRxiv. 2021 March 11; 21253225. Accessed at: https://www.medrxiv.org/content/10.1101/2021.03.11.21253225v2 
  ☐
- 54. Santoli JM, Lindley MC, DeSilva MB, et al. Effects of the COVID-19 Pandemic on Routine Pediatric Vaccine Ordering and Administration United States, 2020. MMWR Morb Mortal Wkly Rep. 2020 May 15;69(19):591-593.

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