

Since January 2020 Elsevier has created a COVID-19 resource centre with free information in English and Mandarin on the novel coronavirus COVID-19. The COVID-19 resource centre is hosted on Elsevier Connect, the company's public news and information website.

Elsevier hereby grants permission to make all its COVID-19-related research that is available on the COVID-19 resource centre - including this research content - immediately available in PubMed Central and other publicly funded repositories, such as the WHO COVID database with rights for unrestricted research re-use and analyses in any form or by any means with acknowledgement of the original source. These permissions are granted for free by Elsevier for as long as the COVID-19 resource centre remains active.



Contents lists available at ScienceDirect

Journal of Clinical Anesthesia

journal homepage: www.elsevier.com/locate/jclinane

Correspondence



Anesthesia and the "post-COVID syndrome": Perioperative considerations for patients with prior SARS-CoV-2 infection

To the Editor:

The COVID-19 pandemic continues to present a major global health threat. Although most patients infected with SARS-CoV-2 do not develop fatal or critical illness, even mild cases may have long-term health consequences. A recent study in *The Lancet* by Huang *et al* found that, among 1733 previously-hospitalized COVID-19 patients, 76% experienced at least one residual symptom 6 months after diagnosis [1]. Symptoms ranging from chest tightness and dyspnea to anosomia and headaches have been described as part of a chronic "post-COVID syndrome." [1,2] [Table 1].

This "post-COVID syndrome" may have significant implications for anesthetic and perioperative care, and is likely to affect a large number of patients worldwide. Very little is known about perioperative management, morbidity and mortality in patients who have recovered from COVID-19. In this paper, we summarize what is known regarding the chronic effects of SARS-CoV-2 infection, and raise key considerations for the perioperative evaluation and anesthetic management of previouslyinfected patients [Table 2].

1. Neurologic considerations

Neurologic manifestations of COVID-19 range from dizziness and headache to encephalitis, seizures and stroke and demyelinating polyneuropathy [3]. SARS-CoV-2 is also known to cause loss of smell and taste, which may reflect direct injury to olfactory neurons. Neurologic sequelae may persist beyond acute illness. In the recent Lancet article, Huang *et al* found that 11-13% of previously-hospitalized COVID patients experienced ongoing loss of smell after 6 months, while 7–9% reported loss of taste. In addition, 5–8% experienced dizziness, and 2–3% complained of persistent headaches [1].

Data are limited regarding optimal anesthetic management of this population. If a patient presents with evidence of peripheral neuropathy, it may be prudent to apply management strategies used in patients with other neuromuscular disorders. These include the judicious use of opiates and neuromuscular blockers (NMBs), and quantitative monitoring of NMB reversal. Avoidance of regional anesthesia may also be advisable. In some cases, patients might also be encouraged to consult a Neurologist postoperatively.

2. Cardiovascular considerations

SARS-CoV-2 infection is associated with significant adverse cardiovascular effects. 20–30% of hospitalized COVID patients show evidence of acute myocardial injury, potentially leading to dysrhythmias and ischemic or non-ischemic cardiomyopathy [4]. Acute cardiac involvement is associated with worse clinical outcomes in COVID-19 patients, but chronic cardiovascular effects of COVID may also be significant [4]. Huang *et al* found that 4–10% of patients reported chest pain and 9–11% experienced palpitations after 6 months [1]. These effects may reflect myocardial fibrosis or inflammation [4]. In another study of 26 patients who complained of new cardiac symptoms following COVID-19 infection, 58% demonstrated evidence of myocardial edema and/or decreased perfusion on cardiac MRI, at a median of 47 days after symptom-onset [5].

Anesthesiologists must be vigilant of possible cardiac dysfunction in patients with prior SARS-CoV-2 infection. Surveillance EKG and TTE have been recommended for COVID-19 patients 2–6 months after acute infection [2]. A low threshold for pre-operative transthoracic echocardiography (TTE) or further evaluation may be appropriate.

3. Pulmonary considerations

COVID-19 may cause long-lasting pulmonary injury in patients who initially develop mild respiratory symptoms. Among patients who did not require supplemental oxygen when hospitalized for COVID-19, 22% demonstrated diffusion impairment on pulmonary function testing (PFTs) 6 months after acute infection, and 22–29% of patients performed below normal on a 6-min walk test [1]. Small-airway dysfunction and new-onset restrictive lung disease have also been described in recovered COVID-19 patients, independent of acute pneumonia severity. [6]

Anesthesiologists must remain vigilant regarding possibly compromised pulmonary function in patients with prior COVID-19. Depending on history and symptom severity, surveillance PFTs and lung imaging may be indicated [7]. If general anesthesia is planned, it may be prudent to emphasize the possibility of postoperative mechanical ventilation.

4. Renal considerations

Moderate or severe SARS-CoV-2 infection is frequently associated with acute kidney injury (AKI). Mechanisms of kidney damage include the pre-renal impact of systemic inflammation, as well as direct viral injury to the kidney *via* the angiotensin converting enzyme 2 (ACE2) receptor [1]. Renal injury may persist beyond the acute phases of COVID-19. Huang *et al* found that 35% of patients demonstrated decreased glomerular filtration rate at follow up, as did 10% of patients with no evidence of AKI in the acute illness setting [1],

Further research is needed to determine the duration of COVIDinduced kidney injury, and it has been recommended that patients who develop AKI in the setting of SARS-CoV-2 infection receive medical follow-up for at least 2–3 months to monitor renal function [8]. Although COVID-induced renal injury is unlikely to be diagnosed in the

https://doi.org/10.1016/j.jclinane.2021.110283

Received 14 January 2021; Received in revised form 18 March 2021; Accepted 20 March 2021 Available online 8 April 2021 0952-8180/© 2021 Elsevier Inc. All rights reserved.

Table 1

Symptoms reported by patients recovered from COVID-19 [1,2].

Fatigue or muscle weakness Mvalgias Sleep difficulties Hair loss Smell disorder Palpitations / Tachycardia Joint Pain Decreased appetite Taste disorder Dizziness Diarrhea or vomiting Chest pain Chest tightness Sore throat or dysphagia Rash Headache Anxiety Low grade fever

Table 2

Management considerations for the anesthesia provider when caring for patients with Post-COVID symptoms.

	Symptom or finding	Management considerations and strategies
Neurologic	Peripheral neuropathy	Judicious use of neuromuscular blockade (NMB) Quantitative reversal of NMB Avoidance of regional anesthesia Perioperative Neurology consult
Cardiovascular	Palpitations	Perioperative EKG
	Chest pain	Transthoracic Echocardiogram
	Decreased myocardial perfusion	Cardiology referral/evaluation
Pulmonary	Residual small airway	Avoidance of general endotracheal
	dysfunction	anesthesia
	Restrictive lung	Discussion of possible postoperative
	disease	mechanical ventilation with patients,
	Diffusion impairment	families and proceduralist team
Hematologic	Hypercoagulability	Pre- and intraoperative
	Venous and arterial thromboses	thromboprophylaxis (mechanical and pharmacologic)
		Use of Enhanced Recovery After
		Surgery (ERAS) protocols to facilitate early mobilization and ambulation
Functional	Fatigue	Referral to pre-operative exercise and
status	Muscle weakness	conditioning programs
	Decreased mobility	

perioperative setting, Anesthesiologists should continue to pay careful attention to clinical and biologic markers of renal function when evaluating patients perioperatively. Avoidance of known nephrotoxic agents may be warranted.

5. Hematologic considerations

Critically ill COVID-19 patients are at particular risk of thromboses affecting all major organs, but patients with mild illness may also be affected. Large vessel ischemic strokes and acute embolic limb ischemia have been reported in young and otherwise healthy patients, at a median 78 days following initial COVID diagnosis [9].

The duration of the COVID-induced pro-thrombotic state is unknown. However, perioperative inflammation and immobility are independent risk factors for venous thromboembolism. Anesthesia providers should be aware of the elevated and possibly compounded risk of thrombosis in surgical patients recovering from COVID-19, and should follow best-practice guidelines regarding thromboprophylaxis in the peri-operative period. "Enhanced Recovery After Surgery" (ERAS) protocols, to facilitate early ambulation and reduce venous thromboembolism risk, may be particularly important.

6. Frailty and decreased functional status

Frail patients have decreased physiologic reserve and significantly increased risk of postoperative complications [10]. Huang *et al* reported that 59–81% of previously-hospitalized COVID patients experienced fatigue and muscle weakness 6 months after diagnosis; between 6 and 14% of patients reported decreased mobility [1].

Anesthesiologists should carefully assess each patient's functional status. If a patient remains de-conditioned following COVID illness, it may be prudent to defer non-urgent procedures. Pre-operative exercise programs, shown to reduce complications in frail patients [10], should be considered.

7. Conclusion

Even beyond the acute crises of the COVID-19 pandemic, millions of people worldwide remain at risk for long-term sequelae of SARS-CoV-2 infection. Anesthesiologists must be prepared to recognize and safely manage these residual effects of COVID-19 infection.

Disclosure

This research did not receive any specific grant from funding agencies in the public, commercial, or not-for-profit sectors. The authors report not proprietary or commercial interest in any product mentioned or concept discussed in this article.

References

- Huang C, Huang L, Wang Y, et al. 6-month consequences of COVID-19 in patients discharged from hospital: a cohort study. The Lancet.
- [2] Davido B, Seang S, Tubiana R, de Truchis P. Post-COVID-19 chronic symptoms: a postinfectious entity? Clin Microbiol Infect 2020;26(11):1448–9.
- [3] Aghagoli G, Gallo Marin B, Katchur NJ, Chaves-Sell F, Asaad WF, Murphy SA. Neurological involvement in COVID-19 and potential mechanisms: a review. Neurocrit Care 2020:1–10. https://doi.org/10.1007/s12028-020-01049-4.
- [4] Mitrani RD, Dabas N, Goldberger JJ. COVID-19 cardiac injury: implications for long-term surveillance and outcomes in survivors. Heart Rhythm 2020;17(11): 1984–90.
- [5] Huang L, Zhao P, Tang D, et al. Cardiac involvement in patients recovered from COVID-2019 identified using magnetic resonance imaging. JACC Cardiovasc Imaging 2020;13(11):2330–9.
- [6] Torres-Castro R, Vasconcello-Castillo L, Alsina-Restoy X, et al. Respiratory function in patients post-infection by COVID-19: a systematic review and meta-analysis. Pulmonology. 2020. https://doi.org/10.1016/j.pulmoe.2020.10.013. S2531-0437 (20)30245-2.
- [7] Hull JH, Lloyd JK, Cooper BG. Lung function testing in the COVID-19 endemic. Lancet Respir Med 2020;8(7):666–7.
- [8] Nadim MK, Forni LG, Mehta RL, et al. COVID-19-associated acute kidney injury: consensus report of the 25th acute disease quality initiative (ADQI) workgroup. Nat Rev Nephrol 2020;16(12):747–64.
- [9] Fan BE, Umapathi T, Chua K, et al. Delayed catastrophic thrombotic events in young and asymptomatic post COVID-19 patients. J Thromb Thrombolysis 2020: 1–7.
- [10] Howard R, Yin YS, McCandless L, Wang S, Englesbe M, Machado-Aranda D. Taking control of your surgery: impact of a Prehabilitation program on major abdominal surgery. J Am Coll Surg 2019;228(1):72–80.

Marguerite M. Hoyler, MD^{*}, Robert S. White, MD MS, Christopher W. Tam, MD, Richard Thalappillil, MD Department of Anesthesiology, New York-Presbyterian/Weill Cornell Medical Center, 525 East 68th Street, Box 124, New York, NY 10065, USA

* Corresponding author.

E-mail address: mam9508@nyp.org (M.M. Hoyler).