Amy Attaway, MD Respiratory Institute, Cleveland Clinic

Management of patients with COPD during the COVID-19 pandemic

Posted April 13, 2020

ABSTRACT

Patients with COPD have an increased risk for severity of COVID-19. Flu-like symptoms with dyspnea may distinguish COVID-19 infection from dyspnea due to a COPD-related exacerbation. Management of COPD with COVID-19 warrants standard-of-care treatment with antimicrobial agents and corticosteroids, though limiting the dose and duration of steroids use may be warranted due to the risk of increased viral shedding. Modalities to treat acute respiratory failure can be used with some caveats. Patients with COPD and COVID-19 infection who treat their illness at home should self-isolate, use nebulizers in ways that avoid viral aerosolization, and frequently disinfect room surfaces.

COPD AND COVID-19

Chronic obstructive pulmonary disease (COPD) represents a large population of patients in the United States¹ and is the fourth most common cause of death.² Coexisting illnesses such as COPD increase the risk for severity of COVID-19,^{3,4} and a better understanding of the therapeutic implications for COVID-19 in the COPD population is needed.

Ways to distinguish early COVID-19-related viral dyspnea from a COPD exacerbation include the coexistence of flu-like symptoms not typically seen with COPD flare-ups, such as fever (89%),³ anorexia (40%), myalgia (35%), and gastrointestinal symptoms (40%).⁵

Steroids have been used with caution during COVID-19 infection given the increased viral shedding seen with the related coronaviruses MERS-CoV and SARS-CoV.^{6,7} There may be indications for treatment with steroids (such as in "cytokine storm") in

doi:10.3949/ccjm.87a.ccc007

COVID-19 as these recommendations evolve.⁸ To date, no evidence of harm caused by the concurrent use of inhaled corticosteroids exists, and we recommend continuing these therapies for patients with COPD.

While previous coronaviruses have been associated with COPD exacerbations,⁹ MERS-CoV, SARS-CoV, and COVID-19 have not been shown to cause COPD exacerbations. If a patient presents with a COPD exacerbation with concurrent COVID-19 infection, we recommend standard-of-care treatment with corticosteroids,¹⁰ but at the same time trying to limit the dose and duration given the concern for increased viral shedding. Comorbid diabetes and hyperglycemia have also been associated with increased risk related to COVID-19,^{4,11} and glycemic control may be an additional challenge when using corticosteroids.

In addition to steroids, azithromycin is a common antimicrobial that is used for COPD exacerbations but has emerged as a possible adjunct therapy with hydroxychloroquine for COVID-19.¹² As such, azithromycin may be in short supply. Alternative antimicrobial therapies may be used including clarithromycin, or secondor third-generation cephalosporins (eg, cefuroxime, cefpodoxime, cefinir) for those without risk factors for Pseudomonas infection. For those at risk of *Pseudomonas* infection, treatment with a fluoroquinolone (eg, ciprofloxacin, levofloxacin) may be considered.¹³

The usual standard of care for acute respiratory failure in COPD involves nebulized therapy, noninvasive positive pressure ventilation (NIPPV), and sometimes high-flow nasal cannula (HFNC) prior to committing to invasive mechanical ventilation. However, use of these modalities has the potential to cause aerosolization of COVID-19, thus increasing the risk of exposure of healthcare workers.¹⁴ We recommend continued use of these modalities with the following caveats:

• Patients should be switched to bronchodilator therapy through metered-dose inhalers or drypowder inhalers whenever possible. Metered

The statements and opinions expressed in COVID-19 Curbside Consults are based on experience and the available literature as of the date posted. While we try to regularly update this content, any offered recommendations cannot be substituted for the clinical judgment of clinicians caring for individual patients.

dose inhalers should be utilized with a spacer.

- When using nebulized therapy, HFNC, and NIPPV, healthcare workers in the room should wear personal protective equipment appropriate for airborne precautions. NIPPV use should include viral filters.
- For HFNC, we recommend limiting the flow to less than 30 L/min.

While the threshold for endotracheal intubation and mechanical ventilation is considered lower for COVID-19, data from China^{4,6,15} show that HFNC and NIPPV were utilized often and that not all episodes of acute respiratory failure were severe enough to require mechanical ventilation. Therefore, we believe there is a role for these modalities and they must be weighed with the risk of COVID-19 aerosolization. While NIPPV use for acute respiratory distress syndrome (ARDS) is not considered standard of care,¹⁶ it is well recognized as an important aspect of treatment for acute respiratory failure secondary to COPD exacerbations and is associated with improvements in mortality rates.¹⁷

In terms of HFNC use in COPD, there are few studies that include use of HFNC in COPD patients with acute respiratory failure, thus further studies are needed. We believe that utilizing HFNC when a patient has hypercapnia (a common occurrence in COPD) should be avoided when possible because the data are limited or have failed to show benefit.^{16,18}

COPD patients are particularly at risk for poor nutritional status¹⁹ and skeletal muscle loss, or sarcopenia.²⁰ Those who develop ARDS from COVID-19 often require mechanical ventilation, and adjunct therapies for ARDS including neuromuscular blockade are both associated with intensive care unitacquired weakness.²¹ Physical therapy and aggressive nutritional supplementation are extremely important to aid COPD patients in recovering from severe illness. The risk of prolonged mechanical ventilation in those with severe underlying COPD is high, and candid discussions with patients and families about difficulty weaning from ventilation and advancedcare planning should be performed early.

A significant proportion of COPD patients with COVID-19 will treat their illness at home. For patients who remain home, it is very important to self-isolate in a room and ideally have their own bathroom. Using a nebulizer in the home has the potential to aerosolize COVID-19. Viral particles can persistent in the air for 1 to 3 hours,²² The nebulizer should be used in a room with the doors closed and that remain closed for several hours afterwards. Also,

cleaning and disinfecting surfaces in the room should be done often. Patients with COPD can consider increasing their bronchodilator use with nebulizers or inhalers while infected with COVID-19 but should only do so after discussion with their provider.

REFERENCES

- Sullivan J, Pravosud V, Mannino DM, Siegel K, Choate R, Sullivan T. National and state estimates of COPD morbidity and mortality – United States, 2014–2015. Chronic Obstr Pulm Dis 2018; 5(4):324– 333. doi:10.15326/jcopdf.5.4.2018.0157
- 2. Heron M. Deaths: leading causes for 2017. Natl Vital Stat Rep 2017; 66(5):1–76. pmid:29235984
- Guan WJ, Ni ZY, Hu Y, et al; China Medical Treatment Expert Group for Covid-19. Clinical characteristics of coronavirus disease 2019 in China. N Engl J Med 2020; Feb 28. doi:10.1056/NEJMoa2002032
- 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; Feb 24. doi:10.1001/jama.2020.2648
- Zhang JJ, Dong X, Cao YY, et al. Clinical characteristics of 140 patients infected with SARS-CoV-2 in Wuhan, China. Allergy 2020; Feb 19. doi:10.1111/all.14238
- Zhu N, Zhang D, Wang W, et al; China Novel Coronavirus Investigating and Research Team. A novel coronavirus from patients with pneumonia in China, 2019. N Engl J Med 2020; 382(8):727–733. doi:10.1056/NEJMoa2001017
- Lee N, Allen Chan KC, Hui DS, et al. Effects of early corticosteroid treatment on plasma SARS-associated coronavirus RNA concentrations in adult patients. J Clin Virol 2004; 31(4):304–309. doi:10.1016/j.jcv.2004.07.006
- Russell CD, Millar JE, Baillie JK. Clinical evidence does not support corticosteroid treatment for 2019-nCoV lung injury. Lancet 2020; 395(10223):473–475. doi:10.1016/S0140-6736(20)30317-2
- Kurai D, Saraya T, Ishii H, Takizawa H. Virus-induced exacerbations in asthma and COPD. Front Microbiol 2013; 4:293. doi:10.3389/ fmicb.2013.00293
- Leuppi JD, Schuetz P, Bingisser R, et al. Short-term vs conventional glucocorticoid therapy in acute exacerbations of chronic obstructive pulmonary disease: the REDUCE randomized clinical trial. JAMA 2013; 309(21):2223–2231. doi:10.1001/jama.2013.5023
- Yang X, Yu Y, Xu J, et al. Clinical course and outcomes of critically ill patients with SARS-CoV-2 pneumonia in Wuhan, China: a singlecentered, retrospective, observational study. Lancet Respir Med 2020; Feb 24. doi:10.1016/S2213-2600(20)30079-5
- 12. Gautret P, Lagier JC, Parola P, et al. Hydroxychloroquine and azithromycin as a treatment of COVID-19: results of an open-label non-randomized clinical trial. Int J Antimicrob Agents 2020; Mar 20. doi:10.1016/j.ijantimicag.2020.105949
- Siddiqi A, Sethi S. Optimizing antibiotic selection in treating COPD exacerbations. Int J Chron Obstruct Pulmon Dis 2008; 3(1):31–44. doi:10.2147/copd.s1089
- Tran K, Cimon K, Severn M, Pessoa-Silva CL, Conly J. Aerosol generating procedures and risk of transmission of acute respiratory infections to healthcare workers: a systematic review. PLoS One 2012; 7(4):e35797. doi:10.1371/journal.pone.0035797
- Wang D, Hu B, Hu C, et al. Clinical characteristics of 138 hospitalized patients with 2019 novel coronavirus-infected pneumonia in Wuhan, China. JAMA 2020; Feb 7. doi:10.1001/jama.2020.1585
- Frat JP, Thille AW, Mercat A, et al; FLORALI Study Group; REVA Network. High-flow oxygen through nasal cannula in acute hypoxemic respiratory failure. N Engl J Med 2015; 372(23):2185–2196. doi:10.1056/NEJMoa1503326
- Lightowler JV, Wedzicha JA, Elliott MW, Ram FS. Non-invasive positive pressure ventilation to treat respiratory failure resulting from exacerbations of chronic obstructive pulmonary disease: Cochrane systematic review and meta-analysis. BMJ 2003; 326(7382):185. doi:10.1136/bmj.326.7382.185

2 CLEVELAND CLINIC JOURNAL OF MEDICINE

- Thille AW, Muller G, Gacouin A, et al; HIGH-WEAN Study Group and REVA Research Network. Effect of postextubation high-flow nasal oxygen with noninvasive ventilation vs high-flow nasal oxygen alone on reintubation among patients at high risk of extubation failure: a randomized clinical trial. JAMA 2019; 322(15):1465–1475. doi:10.1001/jama.2019.14901
- Nguyen HT, Collins PF, Pavey TG, Nguyen NV, Pham TD, Gallegos DL. Nutritional status, dietary intake, and health-related quality of life in outpatients with COPD. Int J Chron Obstruct Pulmon Dis 2019; 14:215–226. doi:10.2147/COPD.S181322
- Schols AM, Broekhuizen R, Weling-Scheepers CA, Wouters EF. Body composition and mortality in chronic obstructive pulmonary disease. Am J Clin Nutr 2005; 82(1):53–59. doi:10.1093/ajcn.82.1.53
- 21. Price DR, Mikkelsen ME, Umscheid CA, Armstrong EJ. Neuromuscular blocking agents and neuromuscular dysfunction acquired in critical illness: a systematic review and meta-analysis. Crit Care Med 2016; 44(11):2070–2078. doi:10.1097/CCM.00000000001839
- van Doremalen N, Bushmaker T, Morris DH, et al. Aerosol and surface stability of SARS-CoV-2 as compared with SARS-CoV-1. N Engl J Med 2020; Mar 17. doi:10.1056/NEJMc2004973