OBSERVATIONAL STUDY

OPEN

Characteristics and Outcomes of Tracheostomized Patients With and Without COVID-19

IMPORTANCE: Outcomes of tracheostomized patients with COVID-19 are seldomly investigated with conflicting evidence from the existing literature.

OBJECTIVES: To create a study evaluating the impact of COVID-19 on tracheostomized patients by comparing clinical outcomes and weaning parameters in COVID-19 positive and negative cohorts.

DESIGN, SETTING, AND PARTICIPANTS: A retrospective observational cohort study of 604 tracheostomized patients hospitalized in 16 ICUs in New York City between March 9, 2020, and September 8, 2021.

MAIN OUTCOMES AND MEASURES: Patients were stratified into two cohorts: 398 COVID-19 negative (COVID-ve) and 206 COVID-19 positive (COVID+ve) patients. Clinical characteristics, outcomes, and weaning parameters (first pressure support [PS], tracheostomy collar [TC], speech valve placement, and decannulation) were analyzed.

RESULTS: COVID+ve had fewer comorbidities including coronary artery disease, congestive heart failure, malignancy, chronic kidney disease, liver disease, and HIV (p < 0.05). Higher Fio $_2$ (53% vs 44%), positive end-expiratory pressure (PEEP) (7.15 vs 5.69), Pco $_2$ (45.8 vs 38.2), and lower pH (7.41 vs 7.43) were observed at the time of tracheostomy in COVID+ve (p < 0.005). There was no statistical difference in post-tracheostomy complication rates. Longer time from intubation to tracheostomy (15.90 vs 13.60 d; p = 0.002), tracheostomy to first PS (2.87 vs 1.80 d; p = 0.005), and TC placement (11.07 vs 4.46 d; p < 0.001) were seen in COVID+ve. However, similar time to speech valve placement, decannulation, and significantly lower 1-year mortality (23.3% vs 36.7%; p = 0.001) with higher number of discharges to long-term acute care hospital (LTACH) (23.8% vs 13.6%; p = 0.015) were seen in COVID+ve.

CONCLUSIONS AND RELEVANCE: Patients with COVID-19 required higher Fio₂ and PEEP ventilatory support at the time of tracheostomy, with no observed change in complication rates. Despite longer initial weaning period with PS or TC, similar time to speech valve placement or decannulation with significantly lower mortality and higher LTACH discharges suggest favorable outcome in COVID-19 positive patients. Higher ventilatory support requirements and prolonged weaning should not be a deterrent to pursuing a tracheostomy.

KEY WORDS: COVID-19; decannulation; tracheostomy; ventilator weaning; weaning parameters

n 2019, a novel coronavirus was identified in Wuhan, China, leading to the global pandemic of COVID-19 disease. The virus that causes COVID-19 is designated severe acute respiratory syndrome coronavirus 2. Compared with other viral infections such as influenza, increased rates of mechanical ventilation, and higher rates of mortality were reported in patients with COVID-19 (1). Increasing severity of acute respiratory distress syndrome (ARDS) in COVID-19 frequently leads to prolonged weaning (2).

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KEY POINTS

Question: We studied the impact of COVID-19 on tracheostomized patients by analyzing clinical outcomes and weaning parameters in COVID-19 positive and negative cohorts.

Findings: This retrospective, observational, cohort study showed that COVID-19 patients required higher Fio₂ and positive end-expiratory pressure ventilatory support at the time of tracheostomy, with no observed change in complication rates. Despite longer initial weaning period with pressure support or tracheostomy collar, similar time to speech valve placement or decannulation with significantly lower mortality and higher long-term acute care hospital discharges were observed.

Meaning: Higher ventilatory support and prolonged weaning should not be a deterrent to pursuing a tracheostomy.

Per 2001 guidelines by The American College of Chest Physicians, the readiness for Spontaneous Breathing Trials and ultimately the ability to be weaned off a ventilator includes the improvement of the cause of the respiratory failure, Pao,/Fio, greater than or equal to 150 or oxygen saturation greater than or equal to 89% on Fio, less than or equal to 40% and positive end-expiratory pressure (PEEP) less than or equal to 5, pH greater than 7.25 with little to no vasopressor support with hemodynamic stability and inspiratory drive (3). In patients not meeting the above criteria with projected prolonged wean, tracheostomies are recommended. Benefits of tracheostomies include less need for deep sedation, shorter weaning time, therefore, a shorter intensive care unit (ICU) and hospital stay (4). Early tracheostomy is typically defined as less than 14 days and is reported to shorten the duration of artificial ventilation and ICU stay, although does not significantly alter mortality (5). Percutaneous technique is usually the procedure of choice over open due to lower risks of surgical site infections and stomatitis (6-9). During the pandemic, this technique was also preferred for minimization of hypoxia and aerosolization (10).

Early literature on tracheostomized COVID-19 patients focused on risks of transmission to health-care professionals. This was later followed by several

reviews analyzing outcomes of weaning, decannulation, and survival, but conflicting evidence still exists surrounding the outcomes of tracheostomized COVID-19 patients. We present a unique cohort study comparing the clinical characteristics, outcomes, and weaning parameters of tracheostomized patients with and without COVID-19, with the aim to evaluate the clinical impact of COVID-19 and provide further guidance on management of tracheostomized patients.

METHODS

Study Design, Setting, and Population

This retrospective, observational, cohort study included all consecutive tracheostomized adult patients (> 18 yr) hospitalized in 16 ICUs within the Mount Sinai Health System in New York City including the Mount Sinai Hospital, Mount Sinai Morningside, West, and Beth Israel, between March 9, 2020, and September 8, 2021. Exclusion criteria included patients who underwent a tracheostomy in an elective setting, had an existing tracheostomy requiring an exchange, and who remained hospitalized at the point of data collection. The selection process is demonstrated in **Figure 1**.

A total of 604 patients were identified and included in the study. The diagnosis of COVID-19 was confirmed by reverse transcriptase-polymerase chain reaction of nasopharyngeal or oropharyngeal specimens. The institutional review board of Mount Sinai Health System initially approved this STUDY-21-01159 on August 17, 2021. As no direct patient contact or intervention from the study group was needed, informed consent was waived. Researchers exclusively used deidentified data. The procedures were followed in accordance with the ethical standards of the responsible committee on human experimentation and with the Helsinki Declaration of 1975.

Data Collection

Clinical data was accessed via the electronic medical record system, Epic, and relevant de-identified data were extracted following review of patient medical charts. Patient demographics, coexisting medical conditions, and clinical data including medications, oxygen requirements, vital signs, and laboratory data were collected. Coexisting medical conditions and presenting symptoms were obtained from physician

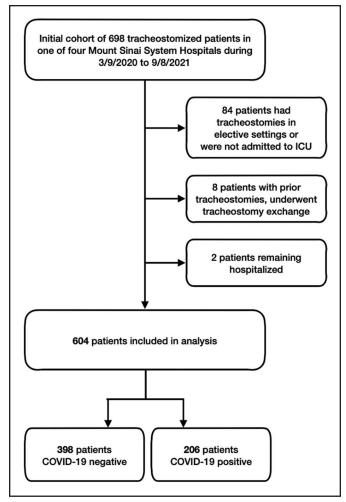


Figure 1. A flow chart of patients included into the study.

documentation. One-year mortality was gathered via Epic chart review.

Statistical Analysis

All analyses were performed with R software (Version 3.6.1; R Foundation for Statistical Computing, Vienna, Austria). Continuous variables are presented as means and sps for normally distributed data or as medians and interquartile ranges for nonparametric data. Categorical variables are summarized as frequencies and percentages. Differences in distributions of characteristics of those with and without COVID-19 were examined using Student *t* test for continuous variables and chi-square test or Fisher exact test (for samples with < 15 observations) for categorical variables. *p* values were calculated with the use of two-sided exact tests and *p* value of less than or equal to 0.05 was considered to indicate statistical significance.

RESULTS

Baseline Characteristics

A total of 604 patients were identified and included in the study. Patients were stratified into two cohorts: 398 patients were COVID-19 negative (COVID-ve) and 206 patients were COVID-19 positive (COVID+ve). The primary reason for admission to ICU for COVID+ve was respiratory failure. Indications of tracheostomy in COVID-ve included pneumonia (n = 66, 16.6%), ARDS (n = 45, 11.3%), other pulmonary disease (n = 32, 8.0%), neurologic disease (neuromuscular/neurovascular disease including anoxic brain injury from cardiac arrest) (n = 176, 44.2%), and other causes (n = 79, 19.8%).

The baseline characteristics of both groups are summarized in **Table 1**. Both groups had a similar mean age (62.08 vs 61.55 yr) and gender distribution (35.7% vs 36.4% females). A larger Hispanic population was seen in COVID+ve (9.5% vs 18.4%). COVID+ve had fewer comorbidities including coronary artery disease, congestive heart failure, malignancy, chronic kidney disease, liver disease, and HIV (p < 0.05).

Ventilator Settings at the Time of Tracheostomy

As demonstrated in Table 2, the mean time from intubation to tracheostomy for COVID+ve was longer with 15.90 days, compared with that of COVID-ve of 13.60 days (p = 0.002). The mean time from last day of continuous sedation to tracheostomy was shorter in COVID+ve with 0.95 versus 3.05 days in COVID-ve (p < 0.001). More sedative agents were used in COVID+ve compared with COVID-ve (4.39 vs 3.22; p < 0.001). The mean F10, and PEEP at the time of tracheostomy for COVID+ve was 53% and 7.15, compared with 44% and 5.69 for COVID-ve (p <0.001). The mean tidal volume was statistically similar in both groups (412.40 mL in COVID-ve, 399.24 mL in COVID+ve; p = 0.093). The mean pH and Pco₂ at the time of tracheostomy for COVID+ve was 7.41 and 45.8, compared with 7.43 and 38.2 for COVID-ve (p =0.004, p < 0.001, respectively).

Clinical Outcomes and Weaning Parameters

Time from tracheostomy to first pressure support (PS) (2.87 vs 1.80 d; p = 0.005) and tracheostomy collar (TC)

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TABLE 1.

Comparison of Clinical Characteristics of Tracheostomized Patients With and Without COVID-19

Characteristics	COVID-19 Negative (<i>n</i> = 398)	COVID-19 Positive (<i>n</i> = 206)	p
Demographics			
Age, yr, mean (sd)	62.08 (14.84)	61.55 (13.16)	0.665
Females	142 (35.7)	75 (36.4)	0.93
Body mass index, kg/m², mean (sp)	26.40 (9.06)	27.93 (7.93)	0.042
Race			< 0.001
White	124 (31.2)	61 (29.6)	
Hispanic	38 (9.5)	38 (18.4)	
African-American	102 (25.6)	29 (14.1)	
Asian	26 (6.5)	6 (2.9)	
Other	108 (27.1)	72 (35.0)	
Comorbidities			
Hypertension	210 (52.8)	113 (54.9)	0.687
Hyperlipidemia	115 (28.9)	64 (31.1)	0.645
Diabetes	130 (32.7)	79 (38.3)	0.193
Coronary artery disease	102 (25.6)	29 (14.1)	0.002
Congestive heart failure	76 (19.1)	13 (6.3)	< 0.001
Chronic obstructive pulmonary disease/asthma	44 (11.1)	34 (16.5)	0.078
Interstitial lung disease	5 (1.3)	4 (1.9)	0.499
Other lung disease	31 (7.8)	10 (4.9)	0.229
Connective tissue disease	10 (2.5)	2 (1.0)	0.237
Malignancy	61 (15.3)	19 (9.2)	0.049
Chronic kidney disease	62 (15.6)	17 (8.3)	0.016
End-stage renal disease	39 (9.8)	11 (5.3)	0.063
Liver disease	48 (12.1)	10 (4.9)	0.004
HIV	16 (4.0)	2 (1.0)	0.042
Cerebrovascular accident	43 (10.8)	16 (7.8)	0.295
Dementia	18 (4.5)	8 (3.9)	0.834

Note—except where indicated, data are number of patients, with percentages in parentheses. Continuous variables are presented as means and SDS for normally distributed data. Categorical variables are summarized as frequencies and percentages. Differences in distributions of characteristics of those with and those without COVID-19 were analyzed using Student t test for continuous variables and χ^2 test or Fisher exact test (for samples with < 15 observations) for categorical variables. Boldface values indicate statistical significance (p < 0.05).

placement (11.07 vs 4.46 d; p < 0.001) were longer in COVID+ve. However, there was no statistically significant difference in the days to first speech valve placement (23.09 vs 29.91 d; p = 0.062) and decannulation post-tracheostomy (49.2 vs 54.40 d; p = 0.474) between the two groups. Similar proportion of patients were eventually decannulated during the hospital stay:

28.1% in COVID-ve (n=112), 30.6% in COVID+ve (n=63; p=0.594). Hospital mortality rates were similar in both groups: 36.4% (n=145) versus 35.9% (n=74; p=0.973). One-year mortality was significantly lower in COVID+ve of 23.3% (n=48) compared with COVID-ve of 36.7% (n=146) with p value of 0.001. More COVID-ve were discharged to a skilled

TABLE 2.Comparison of ICU Therapies in Patients With and Without COVID-19

Characteristics	COVID-19 Negative (n = 398)	COVID-19 Positive (n = 206)	p
Time from intubation to tracheostomy (d)	13.60 (8.66)	15.90 (8.60)	0.002
Time from continuous sedation to tracheostomy (d)	3.05 (5.23)	0.95 (2.69)	< 0.001
Time from antibiotics to tracheostomy (d)	1.97 (3.71)	2.45 (4.92)	0.185
Number of sedative agents since admission, mean (SD)	3.22 (1.23)	4.39 (1.18)	< 0.001
Ventilator settings, mean (SD)			
Fio ₂ at the time of tracheostomy (%)	44.24 (12.96)	53.28 (16.79)	< 0.001
Positive end-expiratory pressure at the time of tracheostomy (mm Hg)	5.69 (1.87)	7.15 (2.71)	< 0.001
Tidal volume at the time of tracheostomy (mL)	412.40 (93.51)	399.24 (83.38)	0.093
Blood gas, mean (SD)			
Potential of hydrogen at the time of tracheostomy	7.43 (0.07)	7.41 (0.07)	0.004
Pco ₂ at the time of tracheostomy	38.17 (10.24)	45.81 (13.14)	< 0.001

Note—except where indicated, data are number of patients, with percentages in parentheses. Continuous variables are presented as means and sps for normally distributed data. Differences in distributions of characteristics of those with and those without COVID-19 were analyzed using Student t test. Boldface values indicate statistical significance (p < 0.05).

nursing facility or a rehabilitation center (41.2% vs 32.5%), whereas more COVID+ve were discharged to a long-term acute care hospital (LTACH) (23.8% vs 13.6%) (p = 0.015).

Last, there was no statistical difference in the rate of complications post-tracheostomy between COVID-ve and COVID+ve. One hundred twenty-nine patients (21%) in total had complications post-tracheostomy across the cohorts: pneumonia (n = 40, 31%), tracheitis (n = 37, 28.7%), minor bleeding (n = 25, 19.4%), major bleeding requiring transfusion (n = 13, 10.1%), stenosis (n = 7, 5.4%), and other (n = 7, 5.4%). Most tracheostomies were performed percutaneously (n = 525, 86.9%). While many tracheostomies in COVID-ve were performed by an ICU team, more tracheostomies were performed by other specialties including general surgery and cardiothoracic surgery for COVID+ve.

There was a statistically significant decrease in number of complications associated with percutaneous tracheostomies (p < 0.01) and with tracheostomies by ICU team (p < 0.005) on regression analyses. There were no statistically significant associations between ventilator settings, recent sedation, antibiotics,

and steroid use prior to the tracheostomy with the rate of complications.

Comparison of outcomes and weaning parameters are summarized in **Table 3**. Tracheostomy characteristics are summarized in **Supplementary Data Table 1** (http://links.lww.com/CCX/B227).

DISCUSSION

Tracheostomy is one of the most frequently performed surgical procedures in the critically ill patient, with known benefits of reduction in days of mechanical ventilation, ICU, and hospital stays (11). The optimal timing and technique of tracheostomy have remained controversial even before the COVID-19 era. Some reported no impact of these on mortality and time to decannulation, whereas others claimed that an early tracheostomy helps reduce ventilator dependence and length of stay (LOS), but with lower 30-day survival (12–14).

Increased rates of mechanical ventilation and increased severity of ARDS with prolonged weaning have been reported to be associated with COVID-19 (1, 2). A higher survival rate in tracheostomized

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TABLE 3.Comparison of Clinical Outcomes and Weaning Parameters in Tracheostomized Patients With and Without COVID-19

Outcomes	COVID-19 Negative (n = 398)	COVID-19 Positive (n = 206)	p
Ventilation weaning			
Time from tracheostomy to first pressure support placement (d)	1.80 (3.43)	2.87 (5.28)	0.005
Time from tracheostomy to first trach collar placement (d)	4.46 (5.87)	11.07 (14.29)	< 0.001
Time from tracheostomy to speech valve placement (d)	23.09 (21.84)	29.91 (22.08)	0.062
Time from tracheostomy to decannulation (d)	49.32 (43.70)	54.50 (46.11)	0.474
Eventual decannulation	112 (28.1)	63 (30.6)	0.594
LOS, d			
Hospital LOS	60.14 (56.40)	54.29 (38.61)	0.182
ICU LOS	40.22 (46.42)	39.20 (24.27)	0.769
Mortality			
Hospital mortality	145 (36.4)	74 (35.9)	0.973
One-yr mortality	146 (36.7)	48 (23.3)	0.001
Disposition			
Home	23 (5.8)	14 (6.8)	0.015
Skilled nursing facility/rehabilitation	164 (41.2)	67 (32.5)	
Long-term acute care hospital	54 (13.6)	49 (23.8)	
Expired	145 (36.4)	73 (35.4)	
Others	12 (3.0)	3 (1.5)	
Complications from tracheostomy			0.695
Yes	65 (16.3)	37 (18.0)	
No	333 (83.7)	169 (82.0)	

LOS = length of stay.

Note—except where indicated, data are number of patients, with percentages in parentheses. Continuous variables are presented as means and sps for normally distributed data. Categorical variables are summarized as frequencies and percentages. Differences in distributions of characteristics of those with and those without COVID-19 were analyzed using Student t test for continuous variables and χ^2 test or Fisher exact test (for samples with < 15 observations) for categorical variables. Boldface values indicate statistical significance (p < 0.05).

COVID-19 cohort when compared with the non-tracheostomized has been described (15). Our large cohort study of tracheostomized COVID-19 and non-COVID-19 patients focused on comparison of clinical settings at the time of tracheostomy, clinical outcomes, and weaning parameters following the tracheostomy.

Clinical Settings: The Timing of Tracheostomy

Optimal timing of tracheostomy has remained controversial, even before the COVID-19 era. An average of

16.5 days from intubation to tracheostomy have been described in two reviews involving 47 studies (n = 5,268) and 37 studies (n = 3,876) of tracheostomized COVID-19 positive patients (12, 13). Owing to the nature of the disease of slow recovery and longer ventilator dependency of COVID ARDS, early tracheostomy was not an option for many due to inability to tolerate a loss of positive airway pressure during the tracheostomy procedure (16). A recent multicenter study of 549 patients describes association of early tracheostomies (defined as < 14 d from intubation) with shorter

duration of ventilation and ICU stay (14, 17–19) but also with increased mortality (20, 21). Controversially, a study argues that an early tracheostomy is noninferior to late tracheostomy with improvement in LOS with no increased infections in clinicians (22).

Our study reports longer time from intubation to tracheostomy of 15.90 days in COVID+ve compared with 13.60 days of COVID-ve (p = 0.002). This can be explained by longer duration to medical stabilization in COVID+ve, earlier tracheostomy in non-COVID neurologic patients, and the availability of staffs and resources due to overburden on the health system during the pandemic.

Clinical Settings: Ventilator Settings

We report that tracheostomies were performed in COVID+ve when they were sicker with lower pH (p = 0.004), higher Pco₂, and required higher ventilator settings than COVID-ve (p < 0.001). Our finding can be explained in two ways. ARDS is described in 42% of COVID+ve patients, with 61-81% of those requiring intensive care (23, 24). ARDS typically involves diffuse alveolar damage with increased epithelial barrier permeability, leading to reduced compliance, compromise of gas exchange and eventual hypoxemia (25). However, COVID ARDS differs from typical ARDS in its severity, longer onset time of the disease (8–12 d), and relatively normal lung compliance in some patients (26). Nearnormal lung compliance and severe hypoxemia due to ventilation/perfusion mismatch is described in type-L patients of COVID ARDS, as opposed to more serious hypoxemia in the setting of low compliance in type-H patients, latter resembling classic ARDS (27). Such patients with good compliance may have been able to undergo the tracheostomy on high ventilator settings. Patients with COVID ARDS are reported to require higher ventilator settings such as PEEP, Pao,, and paralytic agents (2, 24). A review of 26 studies analyzing COVID-19 patients showed average PEEP ranging from 9 to 16.5 cm H₂O, which was higher than mean PEEP of our study of 7.15 cm H₂O (2), in contrast to 5.69 cm H₂O of COVID-ve.

Second, it is also possible that more providers were willing to perform tracheostomy with higher ventilator settings than usual as they witnessed COVID+ve patients' prolonged ventilator requirements and slow recovery.

Significantly longer duration of continuous sedation and greater use of sedative agents (p < 0.001) observed in ventilated COVID+ve patients are consistent with the current literature. Moderate to deep level of sedation is often needed to achieve ventilator synchrony in severe COVID-19 pneumonia and ARDS (28, 29). However, it should be recognized that prolonged and high sedation requirement by COVID+ patients not only creates pressure on the supply chain resulting in shortages of critical medications but also increases the rates of delirium, LOS, and mortality (28, 30).

Clinical Outcomes and Weaning Parameters

Longer time to first PS and TC placement were observed in COVID+ve, compared with COVID-ve (2.87 vs 1.80 d; p = 0.005 and 11.07 vs 4.46 d; p < 0.001, respectively). These findings were expected as COVID+ve were sicker with higher ventilator settings during tracheostomy.

Despite later timing of tracheostomy and higher ventilator settings during tracheostomy; hence, delayed PS placement for COVID+ve, both groups had similar recovery path thereafter with no significant difference in the timing of placement of speech valves and decannulation of tracheostomies. Eventually, both groups had similar ICU and hospital LOS.

Hospital mortality was also similar in both groups during the hospitalization but COVID+ve had better outcome upon discharge with significantly lower 1-year mortality compared with COVID-ve (23.3% vs 36.7%; p < 0.001). The lower 1-year mortality rate we deduced demonstrates initial clinical vulnerability and slow improvement of COVID+ve patients during an early to mid-phase of the disease process, followed by relatively stable and rapid recovery in late phase. It should be noted, however, that the 1-year mortality rates were obtained via chart review and not all deaths may have been recorded.

Furthermore, more tracheostomized COVID+ve were able to be discharged to a LTACH compared with COVID-ve (23.8% vs 13.6%; p = 0.015). This finding shows that COVID+ve were able to demonstrate their potential of ventilator weaning and tolerability of aggressive rehabilitation at the time of discharge (31). As a post-acute care facility, the care at LTACHs is driven by patients' continued acute medical needs with its focus on facilitation of functional recovery and

optimization of respiratory status, including liberation from prolonged mechanical ventilation (32). LTACHs have been proved to be an optimal facility for post-ICU care during COVID-19 pandemic with the rate of successful wean of 70.9% from prolonged mechanical ventilation (33, 34). The overall eventual favorable outcome in COVID+ve patients can be explained by the observed healthier population with less baseline comorbidities as well.

Our study reports no difference in the rate and type of complications between the two cohorts (Table 3). Tracheostomies performed by the ICU team were equally as safe as those performed by others (Supplementary Data Table 1, http://links.lww.com/CCX/B227). Tracheostomies in patients with difficult anatomy are typically deferred to surgical specialties, which likely explains the higher rates of post-operative complications by these specialties.

Limitations

Despite the patients coming from a unique large urban population, given the variability inpatient demographics in the region, our findings can be considered generalizable. However, several limitations are acknowledged.

Our article included a cohort of patients undergoing tracheostomies of different etiologies including nonpulmonary indications with the aim to determine the effect of COVID-19. Direct comparisons between COVID+ve and COVID-ve with ARDS were not performed as many COVID+ve ARDS patients did not survive or were too unstable to undergo a tracheostomy. We acknowledge that comparing patients who underwent a tracheostomy due to severe respiratory failure to those with a neurologic injury could limit valid comparability between the cohorts. This also likely explains the difference in comorbidities of the patients. However, this approach allowed our unique comparison study of a large cohort which we believe can provide useful information to clinicians when managing tracheostomized patients.

Our study as a retrospective, observational study was exposed to possible confounding and selection bias. Furthermore, this study was performed during the peak of the pandemic, which consisted of different strains of COVID-19 with unvaccinated cohort. Further studies need to be performed for applicability to the newer COVID-19 strains and vaccinated population.

CONCLUSIONS

Patients with COVID-19 required higher Fio₂ and PEEP ventilatory support at the time of tracheostomy, with no observed change in complication rates. Although initial weaning period with PS or TC were longer, similar time to speech valve placement or decannulation with significantly lower mortality and higher discharges to LTACH suggest favorable outcome in COVID-19 positive patients. Higher ventilatory support and prolonged weaning should not be a deterrent to pursuing a tracheostomy.

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Dr. Bahk was involved in conceptualization, data curation, formal analysis, investigation, methodology, project administration, resources, supervision, validation, visualization, and reviewing and editing the original draft and writing. Drs. Dolan and Sharma were involved in data curation, investigation, reviewing and editing the original draft and writing. Dr. Sehmbhi was involved in formal analysis, methodology, and validation. Drs. Fung and Lee were involved in conceptualization, project administration, resources, supervision, validation, and reviewing and editing the writing.

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