SURGE IN VENTILATOR-ASSOCIATED PNEUMONIAS AND BLOODSTREAM INFECTIONS IN AN ACADEMIC REFERRAL CENTER CONVERTED TO TREAT COVID-19 PATIENTS

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ABSTRACT

Background: Healthcare-associated infections (HAIs) are important adverse events that must be prevented. **Objective:** The objective of the study was to report and study possible changes in HAI rates as well as their causes after the COVID-19 hospital surge capacity response (HSCR) in an academic referral center. **Methods:** This was a before-after observational study. The Infection Prevention and Control (IPC) program (prospective surveillance, prevention bundles, antibiotic stewardship, continuing education, and feedback) was transiently disrupted after the start of HSCR (March 2020). HAI rates were compared before (January 2019-February 2020) and after (April-July 2020) HSCR, and plausible predisposing factors in affected patients were compared. **Results:** An increase in the HAI rate from 6.2 to 11.8 cases/1000 patient-days was noted between periods due to increases in ventilator-associated pneumonia and bloodstream infection (BSI) rates. More critically ill patients were admitted during HSCR, and use of invasive devices increased. Prone positioning and infusion of muscle relaxants became commonplace. The nurse-to-patient ratio in the intensive care unit decreased, and 4 h shifts were introduced to avoid fatigue. The BSI rate decreased after the IPC program with additional measures was reintroduced in May 2020. **Conclusions:** The strain on the workforce and modifications to the IPC program very possibly underlay the findings. IPC programs continue to be essential during the pandemic. (REV INVEST CLIN. 2021;73(4):210-15)

Key words: Healthcare-associated infection. Ventilator-associated pneumonia. Prevention and control. COVID-19.

INTRODUCTION

Healthcare-associated infections (HAIs) are the most frequent adverse events related to patient care¹, but they can be avoided when adhering to prevention

practices^{2,3}. Use of prevention bundles⁴ in our hospital led to a gradual improvement in the prevention of ventilator-associated pneumonias (VAPs)⁵ and bloodstream infections (BSIs) (not published), achieving nearly zero rates in months previous to the severe

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acute respiratory syndrome coronavirus 2 pandemic. Full COVID-19 hospital surge capacity response (HSCR) started on March 16, 2020, and with it came the following changes in clinical workflow: (1) the nurse-to-patient ratio in the intensive care unit (ICU) declined from 1:1 to 1:2 during HSCR due to bed expansion; (2) <10% of staff was specialized in critical patient care; (3) 4 h shifts were introduced to avoid fatigue related to the use of personal protective equipment; and (4) prone positioning and infusion of muscle relaxants were fully implemented during HSCR, and daily awakening protocols were reduced as a consequence. In addition, epidemiological surveillance rounds were temporarily restricted by heads of different hospital areas in an attempt to lower overall risk of contagion among staff.

It was unknown if these changes had an impact on HAI rates. Therefore, the objective of this research was to calculate HAI rates before and after HSCR, as well as to study the most likely causes of any differences.

METHODS

A before-after observational study was conducted. Active search and report of HAIs as well as patient predisposing factors are routinely done in a prospective, standardized manner and did not undergo any changes during the study period. Mainly due to this reason, the institutional review board granted a waiver for informed consent; furthermore, the study was deemed to pose no risk to patients due to the use of deidentified data in the database.

Our institutional Infection Prevention and Control (IPC) program relies on the following: (1) daily prospective surveillance, (2) HAI prevention bundles, (3) antibiotic stewardship, (4) continuing education, and (5) performance feedback on a monthly basis. Objective evidence on the conduct of these activities is routinely recorded on standardized sheets and reported monthly, as per hospital policies.

Starting in May 2020, additional IPC measures instituted were (1) retraining in prevention bundles, (2) retraining in safe central venous catheter insertion practices, (3) halting of double gloving, (4) emphasis on hand hygiene, and (5) weekly meetings between hospital authorities and heads of clinical areas.

HAIs were identified in all hospitalization areas (emergency department, ICU, accessory critical area, and hospital wards) and defined according to the NHSN standardized criteria current in 2019 and 2020⁶. HAI events and the corresponding denominator data were recorded daily. Monthly HAI rates were calculated dividing the number of events by the number of patient-days and then multiplying by 1000; patient-days was used to simplify comparisons between different types of HAIs (rates adjusted to device-days and patient-days gave similar trends).

The following variables were compared among patients affected by HAIs: sex, age, length of stay before diagnosis of HAI, hospital area where HAI was identified, rate of in-hospital death, risk factors for hospital-acquired pneumonias and VAPs (endotracheal intubation, tracheostomy, use of humidifier, use of feeding tubes, use of proton-pump inhibitors, or H2 receptor antagonists), risk factors for BSIs (use of any type of venous or arterial catheter, parenteral nutrition), risk factors for catheter-associated urinary tract infection (use of urinary catheter), and risk factors for *Clostridioides difficile* infection (previous use of antibiotics, proton-pump inhibitors, and H2 receptor antagonists). Absolute and relative frequencies were used to summarize data.

Two time periods were compared: before HSCR (January 2019–February 2020) and during HSCR (April–July 2020). March 2020 was left out due to active HSCR. Gradual resumption of routine hospital activities began in July 2020.

Statistical Analysis

Statistical analysis by the Chi-square test or Fisher's exact test (for categorical variables) or the Wilcoxon rank-sum test (for numerical variables) was done to contrast differences between the two time periods, considering a p cutoff value of ≤ 0.05 as statistically significant. Stata[®] version 14.0 (StataCorp, College Station, TX, USA) was used.

RESULTS

Before HSCR, 449 HAIs in 72,832 patient-days were identified (HAI rate = 6.2/per 1000 patient-days). During HSCR, 170 HAIs in 14,372 patient-days occurred (HAI rate = 11.8 cases per 1000 patient-days).

Figure 1. Healthcare-associated infection rates stratified by type of infection. Catheter-associated urinary tract infection and surgical site infection were left out for clarity; no changes were noted in rates of the former and no surgeries related to non-COVID-19 conditions were done during hospital surge capacity response.

The increase in the total HAI rate during HSCR was due to increases in VAP and BSI rates (Fig. 1).

A greater proportion of critically ill patients was admitted during HSCR, leading to an increase in the prevalence of risk factors related to insertion and use of invasive devices (Table 1).

Due to the workflow modifications during HSCR mentioned above, surveillance and estimation of adherence to the various measures of the IPC program were non-existent during HSCR.

The types of microorganisms causing HAIs differed between the two periods (Table 1). During HSCR, an increase in the proportion of Gram-negative bacilli and fungi was noted; in contrast, cases of *C. difficile* infection diminished.

DISCUSSION

An increase in BSI and VAP rates in our hospital was temporally associated with the interruption of our IPC program and changes in work standards due to HSCR.

Our findings are in agreement with another study that reported a notorious increase in the BSI rate in COVID-19 patients, mostly due to Gram-negative bacteria⁷; among the causes, a higher number of critical patients admitted to strained ICUs, reduced staff-to-patient ratios, and increased length of stay of patients were found. As noticed in that report, we also observed a reduction in our BSI rate after adoption of preventive actions. Although we are not able to provide a statistical value to this observation due to the limited HSCR time period during which BSI rates increased, there was a notorious reduction after the IPC program was resumed.

Regarding the VAP rate, our findings were similar to those of another study⁸: VAP in COVID-19 patients was a very frequent occurrence when mechanical ventilation was required. The rise in the endotracheal intubation rate related to the pandemic was a contributory factor⁵; however, prone positioning, use of muscle relaxants, a decreased rate of daily awakening protocols, and decreased surveillance of HAIs could also be associated. Of the potential associated factors mentioned previously, only resumption of surveillance was possible since modification of the others

Table 1. Comparison of patient characteristics, risk factors, and microorganisms isolated from healthcare-associated infections before and during HSCR

Variable	Before HSCR n (%) ^a	During HSCR n (%) ^a	p
HAI rate (×1000 patient-days)	6.2	11.8	0.023
Male sex	213 (47.4)	117 (68.8)	<0.001
Age, years (median, IQR)	53 (41-68)	53 (43-60)	0.150
LoS before HAI, days (median, IQR)	13 (7-23)	10.5 (8-17)	0.010
In-hospital death	24 (5.3)	55 (32.4)	<0.001
Types of HAIs			
VAP	45 (10.0)	93 (54.7)	<0.001
Hospital-acquired pneumonia	121 (26.9)	31 (18.2)	0.025
BSI	6 (1.3)	35 (20.6)	<0.001
Candidemia	0 (0)	14 (8.2)	<0.001
Catheter-associated urinary tract infection	37 (8.3)	6 (3.5)	0.039
Surgical site infection	113 (25.2)	0 (0) ^b	NA
<i>C. difficile</i> infection	68 (15.2)	3 (1.8)	<0.001
Other	59 (13.1)	2 (1.2)	<0.001
Risk factors for various HAIs			
Arterial catheter	2 (0.4)	52 (30.6)	<0.001
Hemodialysis catheter	11 (2.4)	12 (7.1)	0.007
Central venous catheter	306 (68.2)	158 (92.9)	<0.001
Venous peripheral catheter	232 (51.7)	86 (50.6)	0.806
Parenteral nutrition	3 (0.7)	3 (1.8)	0.354
Endotracheal intubation	51 (11.4)	154 (90.6)	<0.001
Time from endotracheal intubation to VAP, days (median, IQR)	9 (6-14)	10.5 (8-16)	0.087
Use of humidifier	19 (4.2)	5 (2.9)	0.458
Tracheostomy	11 (2.4)	5 (2.9)	0.778
Nasogastric tube	5 (1.1)	4 (2.4)	0.268
Nasojejunal tube	0 (0)	27 (15.9)	<0.001
Use of PPI or H2 receptor antagonists	311 (69.3)	116 (68.2)	0.806
Urinary catheter	186 (41.4)	155 (91.2)	<0.001
Use of antibiotics before HAI	381 (84.9)	121 (71.2)	<0.001
Hospital area where HAI was identified			
Wards	365 (81.3)	9 (5.3)	<0.001
Emergency department	44 (9.8)	108 (63.5)	
ICU	39 (8.7)	38 (22.4)	
Accessory critical area	NA	15 (8.8)	
Other area	1 (0.2)	0 (0)	

(Continues)

Table 1. Comparison of patient characteristics, risk factors, and microorganisms isolated from healthcare-associated infections before and during HSCR (*continued*)

Variable	Before HSCR n (%) ^a	During HSCR n (%) ^a	p
Hospital area where HAI was identified			
Microorganisms isolated	288	207	
Gram-negative bacilli	140 (48.6)	141 (68.1)	<0.001
<i>Enterobacter cloacae</i>	4	33	
<i>Escherichia coli</i>	34	22	
<i>Klebsiella pneumoniae</i>	16	18	
Other <i>Enterobacteriaceae</i>	28	28	
<i>Pseudomonas aeruginosa</i>	27	24	
Other non-fermenting Gram-negative bacilli	31	16	
Gram-positive cocci	42 (14.6)	27 (13.0)	0.624
Enterococci	24	5	
<i>Staphylococcus aureus</i>	9	9	
Coagulase-negative staphylococci	1	9	
Other cocci	8	4	
Fungi	23 (8.0)	35 (17.0)	0.002
<i>Candida albicans</i>	9	9	
Non-albicans <i>Candida</i> spp.	6	14	
<i>Aspergillus</i> spp.	5	9	
Other fungi	3	3	
<i>C. difficile</i>	68 (23.6)	3 (1.4)	<0.001
Other microorganism	15 (5.2)	1 (0.5)	0.003
Resistant microorganisms of epidemiological importance			
ESBL producers	4/8 (50.0)	12/29 (41.4)	0.705
AmpC producers	6/14 (42.9)	52/82 (63.4)	0.146
Carbapenem-resistant <i>Enterobacteriaceae</i>	2/12 (16.7)	9/79 (11.4)	0.635
Methicillin-resistant <i>S. aureus</i>	2/6 (33.3)	0/5 (0)	0.455
Multidrug-resistant <i>P. aeruginosa</i>	5/7 (71.4)	2/21 (9.5)	0.004
Carbapenem-resistant <i>Acinetobacter baumannii</i>	4/4 (100)	—	NA
Azole-resistant <i>Candida</i> species	—	8/13 (61.5)	NA

^aUnless otherwise stated.^bNon-COVID-19 surgical procedures were halted during HSCR.HSCR: hospital surge capacity response; HAI: healthcare-associated infection; IQR: interquartile range; LoS: length of stay; VAP: ventilator-associated pneumonia; BSI: bloodstream infection; PPI: proton-pump inhibitor; NA: not applicable; ESBL: extended-spectrum beta-lactamase; AmpC: ampC beta-lactamase. ICU: intensive care unit, *C. difficile*: *Clostridioides difficile*.

was unfortunately restricted to patients without severe disease. This could explain the lack of reduction in VAP rates.

The rates of hospital-acquired pneumonia in wards remained stable, possibly due to the admission of non-intubated and less severely ill patients in such areas.

Finally, two circumstances that may account for reduced rates of *C. difficile* infection were (1) not admitting patients with active infection and (2) increased adherence of hand hygiene and contact precautions by staff⁹.

Pending further studies, the present one suggests that HAI preventive tasks may be compromised during the COVID-19 pandemic. A previous survey found that the majority of the personnel dedicated to IPC duties spent most of the time responding to the COVID-19 emergency, undermining important activities such as surveillance of adherence to hand hygiene, chlorhexidine bathing, antimicrobial stewardship programs, and provider feedback¹⁰.

We acknowledge limitations. Because interruption of the IPC program coincided with changes in workflow and in medical procedures (especially prone positioning), we were not able to estimate the relative contribution of each separate event to the increase in HAIs during HSCR. However, the decrease in BSI rates after resumption of the IPC program in May 2020 (and accompanying additional measures specific to the prevention of BSIs) plus no further changes in the other likely associated factors previously mentioned, support the observation that interruption of the IPC program was very possibly related to the surge in HAIs. Although the use of dexamethasone could be theoretically related to increases in HAI rates elsewhere, we consider this highly unlikely in our setting since dexamethasone use was established after June 30, 2020; therefore, this variable was not analyzed in our study.

In summary, an increase in the incidence of VAPs and BSIs was noted during HSCR in a COVID-19 setting. Resumption of prevention activities allowed a reduction

of BSI rates. IPC programs must remain an essential activity during the COVID-19 pandemic.

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