The Disproportionate Impact of COVID-19 Pandemic on Healthcare-Associated
 Infections in Community Hospitals: Need for Expanding the Infectious Disease
 Workforce

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1 Abstract:

Background: The COVID-19 pandemic had a considerable impact on US healthcare
systems, straining hospital resources, staff, and operations. However, a comprehensive
assessment of the impact on healthcare associated infections (HAIs) across different
hospitals with varying level of infectious disease (ID) physician expertise, resources,
and infrastructure is lacking.

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Methods: This retrospective longitudinal multi-center cohort study included central-line-8 associated bloodstream infections (CLABSIs), catheter-associated urinary tract 9 infections (CAUTIs), C. difficile infections (CDIs), and ventilator-associated events 10 (VAEs) from 53 hospitals (academic and community) in Southeastern United States 11 from January 1, 2018 to March 31, 2021. Segmented negative binomial regression 12 generalized estimating equations models estimated changes in monthly incidence rates 13 in the baseline (01/2018 - 02/2020) compared to the pandemic period (03/2020 -14 03/2021, further divided into three pandemic phases). 15

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Results: CLABSIs and VAEs increased by 24% and 34% respectively during the pandemic period. VAEs increased in all phases of the pandemic, while CLABSIs increased in later phases of the pandemic. *CDI* trend increased by 4.2% per month in the pandemic period. On stratifying the analysis by hospital characteristics, the impact of the pandemic on healthcare-associated infections was more significant in smaller sized and community hospitals. CAUTIs did not change significantly during the pandemic across all hospital types.

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Conclusions: CLABSIs, VAEs, and *CDI*s increased significantly during the pandemic,
especially in smaller community hospitals, most of which lack ID physician expertise.
Future efforts should focus on better understanding challenges faced by community
hospitals, strengthening infection prevention infrastructure, and expanding the ID
workforce, particularly to community hospitals.

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8 Keywords: SARS-COV-2, pandemic, infection prevention, healthcare associated

9 infections, CLABSIs, CAUTI, Cdifficile, VAE

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Prior Presentation: These data were presented as an <u>oral presentation</u> #172 "Impact
 of COVID-19 pandemic on healthcare-associated infections in a large network of
 hospitals" at IDWeek 2021 virtually.

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1 Background:

Healthcare associated infections (HAIs) are key quality and safety indicators for US
acute care hospitals. HAI rates are publicly reported and impact hospital reimbursement
by the Centers for Medicare and Medicaid Services (CMS).¹ National efforts led to
reduction of central line associated bloodstream infection (CLABSI), catheter-associated
urinary tract infection (CAUTI), *C. difficile* (CDI) and ventilator associated events (VAEs)
over the past decade.²

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The COVID-19 pandemic had a considerable impact on US healthcare systems, 9 straining hospital resources, staff, and operations, and further exacerbated the shortage 10 of infectious disease (ID) physicians.³⁻⁷ Nearly two-thirds of all Americans are primarily 11 served by community hospitals and live in counties with a below-average density of ID 12 physicians or no ID physician access.⁸ Even before the pandemic, surveys highlighted 13 that less than a guarter of community hospitals had ID specialists on staff, of which only 14 a minority received training in in hospital epidemiology and infection prevention.⁹ Robust 15 evidence supports the association between ID physician intervention and improved 16 outcomes.¹⁰ 17

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By early December 2020, COVID-19 hospitalizations accounted for about 15% of occupied inpatient beds.¹¹ Patients with COVID-19 required higher acuity of care with more diagnostic, therapeutic, and safety measures, which placed tremendous pressure on healthcare workers, from staffing challenges to risk of exposure and infection.^{4,12-14} In addition, there was a precipitous drop in elective surgical cases, and a decline in

admissions for other medical conditions, resulting in a higher case mix index among
hospitalized patient populations.¹¹ Furthermore, COVID-19 patients needed close
monitoring and higher levels of care, including ventilatory and critical care support.¹⁵⁻¹⁷
In summary, hospitals faced dynamic and different challenges to healthcare delivery
during each pandemic surge.

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Prior data from other health systems and national surveillance provided some insights 7 on the impact of the COVID-19 pandemic on HAIs.^{3,5,18} However, these data did not 8 assess impact across hospitals of different types (academic vs. community), were 9 limited to assessment of early pandemic timelines, and did not explore effects in 10 different phases of the pandemic. Our objective was to evaluate the impact of COVID-11 19 pandemic on incidence and trends of HAIs in a large network of academic and 12 community hospitals across different phases of the pandemic, and better understand 13 the impact on under-resourced community hospitals. 14

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16 Methods:

Study Design: We performed a retrospective, longitudinal study of CLABSIs, CAUTIS, CDI and VAEs in a large network of hospitals from January 1, 2018 to March 31, 2021. The study included a "baseline period" from January 1, 2018 – February 29, 2020 and "pandemic period" from March 1, 2020 to March 31, 2021. The "pandemic period" was further divided into three "pandemic phases" in order to match COVID-19 surge patterns observed in study sites: Pandemic Phase 1 (03/2020 – 06/2020), Pandemic Phase 2 (07/2020-10/2020) and Pandemic Phase 3 (11/2020-3/2021) as shown in Figure 1.

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Setting: This study included data from 53 hospitals, which included 2 large academic medical centers: Duke University Hospital, an acute-care, academic, tertiary-care facility in Durham, North Carolina, and the University of North Carolina Medical Center, an academic medical center in Chapel Hill, North Carolina, and 51 community hospitals in the Duke Infection Control Outreach Network (DICON) in the southeastern United States.¹⁹

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Definitions: CLABSI, CAUTI, CDI and VAEs were defined based on Centers for 9 Disease Control and Prevention's (CDC's) National Health Safety Network (NHSN) 10 surveillance definitions.²⁰ There were no major HAI definition changes during the study 11 period. Hospital characteristics included hospital type and size. Hospitals were 12 categorized by type into academic medical centers and community hospitals, and 13 stratified by size into large (≥425 beds), medium (250-424 beds), and small (≤249 14 beds) hospitals, based on the Agency for Healthcare Research and Quality's Healthcare 15 Cost and Utilization Project National Inpatient Sample Description of Data Elements.²¹ 16 Pandemic phases were defined based on COVID-19 case burden (using COVID19 17 admissions), representing the strain on the hospital resources and frontline workers 18 (Figure 1). COVID-19 admission was defined as an inpatient hospitalization with an 19 ICD10 diagnosis code of U07.1 (after April 2020) or B97.29 (March 2020-April 2020). 20 COVID-19 admissions were assigned to the month of the admission date. 21

1 Data Collection: The study design was approved by the institutional review board of Duke University Health System (Pro00107094). Informed consent for data collection 2 was waived because all relevant epidemiologic data were abstracted by each 3 4 participating site without transfer of any protected health information. Trained infection preventionists at each hospital collected surveillance data using a standardized 5 database and NHSN definitions. DICON liaison infection preventionists validated a 6 subset of surveillance data each month. The study database included variables 7 measured per hospital-month on the facility-wide level. The outcomes were reported as 8 CLABSIs per 1000 central catheter days, CAUTIs per 1000 urinary catheter days, CDI 9 per 10,000 CDI patient days, and VAE per 1000 ventilator days. Secondary outcomes 10 included central catheter days, urinary catheter days, and ventilator days. We followed 11 the Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) 12 reporting guideline. 13

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Analysis: Outcomes were reported descriptively as mean (standard deviation, SD) 15 hospital-level monthly incidence rates (IR) by period – "baseline period", "pandemic 16 period" or "pandemic phases". Mean hospital-level monthly IR and device days were 17 compared using negative binomial regression GEE models with period as the only 18 covariate. Segmented regression (SR) analyses using generalized estimating equations 19 20 (GEE) models were performed to estimate changes in monthly IR of CAUTIS, CLABSIS, CDI and VAE. Poisson distribution was assumed for VAE and negative binomial 21 distribution was assumed for all other HAIs. The working correlation matrix assumption 22 23 was compound symmetry for CLABSI, CDI and VAE, and independent for CAUTI

models. Unit of analysis was hospital-month. The data was analyzed in the following
ways to provide summary estimates of interest and then to adjust for hospital
characteristics (type and size) and COVID-19 pandemic phases:

4 1) Pre- and post-comparison of HAI IRs and device days in baseline and entire5 pandemic period.

2) Unadjusted analysis showing overall trends of HAIs in baseline and pandemic
periods: All models included variables for time since the beginning of the observation
period (starting 12/2017), time since the beginning of the pandemic (starting 03/2020),
and type of period (baseline vs. the pandemic).

3) Unadjusted analysis showing HAI trends in baseline and pandemic period stratified by hospital type and size: Stratified analyses included subgroup variable for hospital type (academic vs. community), or hospital size (small, medium or large), and interactions between subgroup and all other variables in the model.

4) Analysis adjusted for hospital characteristics showing HAI trends in baseline period
and three pandemic phases to evaluate the impact of different pandemic phases on
infection trends: All models included variables for time since the beginning of the
observation period (starting 12/2017), pandemic phases (baseline period, pandemic
phase 1, 2, or 3), time since the beginning of each pandemic phase, number of hospital
beds, hospital type (academic vs. community).

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All statistical tests were two-sided with alpha level of 0.05. All analyses were performed
using SAS v.9.4 (SAS Institute, Cary NC).

1 Results:

In our large network of 53 hospitals over a 39-month study period, there were 4060
COVID-19 admissions in pandemic phase 1, 9593 COVID-19 admissions in pandemic
phase 2, and 17,586 COVID-19 admissions in pandemic phase 3 (Figure 1).

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6 Hospital characteristics:

The two large academic medical centers included in the study have 803 and 957 beds, while the community hospitals range in size from 38 to 721 beds (median, 162 beds) in 6 states: North Carolina, South Carolina, Virginia, Florida, Georgia, and West Virginia.¹⁹ Thirty-nine community hospitals were classified as small in size, while 12 community hospitals were medium and large. Seventy five percent of our community hospitals are located in counties without an ID physician, or in counties where ID physician density is lower than national average (1.76 per 100,000 US population, Supplemental Figure 1). ⁸

Pre and post comparison of HAI incidence rates and device days in baseline and pandemic period:

Mean monthly CLABSI IR increased significantly from 0.6 to 0.9 per 1000 catheter days (P=0.0023), while VAE IR increased from 6.1 to 10.9 per 1000 ventilator days (P<0.001) during the entire pandemic period. In contrast, CAUTI IRs (P=0.81) did not change significantly during the pandemic (eTable 1 in the Supplement). Overall mean monthly CDI IR decreased significantly from 3.6 to 2.6 per 10,000 patient days in the pandemic period (P< 0.001). Compared to baseline period, mean central line days (937 vs 969, P=0.033) and mean ventilator days (210 vs 281, P< 0.001) increased, but mean urinary</p> catheter days (798 vs 817, P=0.16) or mean CDI patient days (4877 vs 4765, P=0.21)
remained unchanged during the pandemic period.

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4 Unadjusted analysis examining HAI trends in baseline and pandemic periods:

In the unadjusted analysis, CLABSI and VAE trends were stable in the baseline period. 5 However, CLABSI and VAE IRs increased by 24% (95%CI:1.2% to 51.4%, P=0.038) 6 and 34% (95% CI:7% to 67.8%, P=0.011) respectively, at the start of the pandemic 7 period (Supplemental Figure 2, Table 1: "Pandemic Level Change"). CDI IR was 8 decreasing by 2.1% per month during the baseline period (95% CI: -1.1%- to -3.1%, 9 P<0.001), but increased by 4.2% per month in the pandemic period (95% CI: 1.7% to 10 6.8%, P= 0.001; Table 1: Pandemic Trend Change"). CAUTI IR did not change 11 significantly during the baseline and pandemic periods. (Supplemental Figure 2, Table 12 1: Pandemic Level and Trend Changes). 13

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15 Analysis examining HAI trends further stratified by hospital type and size:

On stratifying the analysis by hospital type, CLABSI and VAE IRs increased significantly in community hospitals by 48% and 41.4% respectively, but did not change significantly in academic hospitals at the start of the pandemic. (eTable 2 in the Supplement). CDI IR increased by 4.5% per month in community hospitals during the pandemic period, but decreased by 43% at the start of the pandemic period in academic hospitals (eTable 2 in the Supplement). CAUTI trends remained unchanged across all hospital types in the pandemic period (eTable 2 in the Supplement). However, the differences between academic and community hospitals were not statistically significant (interaction p-values
 >0.05).

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4 On stratifying the analysis by hospital size, CLABSI IR increased by 82.1% across small hospitals at the start of the pandemic and by 6.3% per month across medium sized 5 hospitals (eTable 3 in the Supplement). VAE IR increased by 48.7% and 104% across 6 small and medium sized hospitals respectively, at the start of the pandemic. (eTable 3 7 in the Supplement). CLABSI and VAE IRs remained relatively stable in large hospitals 8 during the pandemic. CDIs increased in small and medium hospitals by 4.5% and 9 12.5% per month. On the contrary, CDI IR decreased in large hospitals by 27% at the 10 start of the pandemic (eTable 3 in the Supplement). Pandemic level changes for 11 CLABSI IR (interaction p-value = 0.024), and baseline trend changes for CDI and 12 CAUTI IR (interaction p-values 0.042 and 0.023, respectively) were significantly 13 different by hospital size. 14

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Analysis examining HAI trends adjusted for hospital characteristics in baseline and pandemic phases:

After adjusting for hospital characteristics and including pandemic phases, VAE IR increased significantly in all three pandemic phases: 33.9% (95% CI: 4.5% to 71.4%, P=0.020) in phase 1; 42.9% (95% CI: 0.3% to 103.5%, P=0.048) in phase 2, and 53.3% (95%CI: 12.3% to 109.4%, P=0.007) in phase 3 (Figure 2, Table 1: Pandemic Level Change) compared to baseline period. VAE IR trend started to decline in pandemic phase 3 (-7.3%, (95% CI: -12.3% to -2.0%, P=0.007). CLABSI IR was stable in pandemic phase 1, but increased by 44.3% at the start of pandemic phase 2 (95% CI:
6% to 96.2%, P=0.019) and 48.7% at the start of pandemic phase 3 (95% CI: 9.5% to
102%, P=0.011) compared to the baseline period. CDI and CAUTIs IRs did not change
significantly during any pandemic phases (Figure 2, Table 1: Pandemic Level and Trend
Changes).

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7 Discussion:

In this large multistate cohort of hospitals, overall rates of CLABSIs, CDIs, and VAEs increased significantly during the COVID-19 pandemic. In contrast, we observed no change in CAUTI rates during the pandemic. Furthermore, VAE rates increased across all phases of the pandemic, while CLABSIs increased during later phases of the pandemic. Our data also highlight that the COVID-19 pandemic exerted a disproportionately larger impact on smaller community hospitals compared to large academic medical systems.

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To our knowledge, our study is the first to addresses disparities among larger academic 16 hospitals and smaller community hospitals with respect to HAI trends during the 17 pandemic. The differences in impact of the COVID-19 pandemic across hospital types 18 could be secondary to differences in resource allocation, availability of ID expertise, and 19 catchment area served by smaller community hospitals.²² Despite access to remote 20 infection prevention expertise through DICON, these community hospitals struggled to 21 manage complex COVID-19 patients, to advocate for resources, staff, and 22 23 infrastructure, and retain focus on patient safety in the absence of an onsite ID

physician champion.²³⁻²⁵ Additionally, most of the smaller community hospitals in our 1 network are located in rural areas¹⁹ and primarily cater to an underinsured older 2 population with several comorbidities.²⁶ Furthermore, these hospitals struggled with 3 staffing shortages, financial challenges, and operational limitations for years, only to be 4 exacerbated by the pandemic.^{27,9} Even prior to the pandemic, 172 rural hospitals closed 5 according to the North Carolina Rural Health Research Program.²⁸ During the 6 pandemic, the volume and acuity of COVID-19 cases guickly overwhelmed the very 7 limited resources in community hospitals in these areas.¹² 8

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Our analysis also highlights changes in HAI trends during different phases of the 10 pandemic: VAE rates increased across all phases of the pandemic, while CLABSIs 11 increased during later phases of the pandemic. Furthermore, our survey data from early 12 and late stages in the pandemic suggest that specific risk factors were more prominent 13 during different phases of the pandemic.^{4,12} Our initial survey highlighted resource 14 shortages, use of contingency strategies, decrease in elective procedures, and 15 inconsistencies in protocols early in the pandemic.¹² Additionally, in-person patient care 16 was primarily performed only by essential team members, while other staff and 17 consultants performed 'electronic medical record rounding".²⁹ Intravenous pumps were 18 placed in hallways to avoid frequent re-entry of nurses to rooms of patients infected with 19 COVID-19.30 These changes in workflow likely led to lapses in insertion and 20 maintenance practices, and delayed removal of devices. Later in the pandemic, elective 21 procedures were restarted, testing capacity increased, and resource shortages 22 23 improved. However, the increasing COVID-19 burden exacerbated staff fatigue, staffing

shortages, and led to increase in resignations, which further adversely impact patient
care.^{4,31} In addition, hospitals attempted to mitigate staffing issues by recruiting
travelling nurses or reassignment of infection prevention and antibiotic stewardship staff
to cover other critical duties,^{4,32} likely further contributing to lapses in infection
prevention and antibiotic stewardship.¹⁵

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Interestingly, CAUTI rates remained unchanged during the pandemic in our network of 7 hospitals. This is possibly, in part, due to lack of change in urinary catheter use during 8 the pandemic, and partly due to the limitations of the NHSN CAUTI definition.³³ We 9 hypothesize that there may have been an increase in culture negative CAUTIs (not 10 captured by current NHSN definition), as many COVID-19 patients were receiving 11 antimicrobials³⁴ at the time of urine culture collection likely leading to sterilization of 12 urine. There may have also been a shift in testing practices from overuse of urine 13 cultures to focusing on repeat SARS-COV-2 testing for febrile episodes in critically ill 14 patients.³⁵ 15

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Several health systems reported an increase in CLABSIs during the COVID-19 pandemic.^{3,5} However, there were mixed results in trends of CAUTIs and CDIs, and limited data on VAEs.⁵ These mixed results can partly be explained by design of the study, e.g. comparing time-trended analysis with pre-post comparisons. Additionally, prior studies were also limited by shorter time periods of analysis, use of simple prepost comparisons, and lack of adjustment for COVID-19 burden and hospital characteristics.^{3,5,18} In contrast, our data accounts for over 3 years of trended infection

data, includes pandemic phases based on COVID-19 burden and accounts for the
differential impact on under-resourced hospitals that lack ID expertise. Additionally, our
analysis of the impact of COVID-19 on antibiotic use and surgical site infections has
been previously reported.^{36,37} Overall, we believe our methodological improvements
have led to novel insights on the impact of the COVID-19 pandemic on HAIs.

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Our study has limitations. HAI outcomes were based on the use of surveillance 7 definitions. However, surveillance methods were applied systematically across all 8 network hospitals throughout the pandemic. We only included data until March 2021. 9 However, our study reports over 3 years of longitudinal data. We did not include 10 microbiology data, specific data on ID physicians, and severity of illness in this analysis, 11 as our goal was to evaluate the overall impact of pandemic phases and hospital 12 characteristics. While we have adjusted the models for hospital characteristics and 13 pandemic phases, we cannot fully exclude the possibility of unmeasured confounding 14 influencing the results. Lastly, while our hospitals are located across multiple states, our 15 experience may not be fully generalizable to all settings. 16

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18 **Conclusions:**

In this large multistate cohort of hospitals, the COVID-19 pandemic led to notable changes in HAI rates: VAEs increased consistently in all phases of the pandemic, while CLABSIs increased later in the pandemic. This impact on HAIs was disproportionately higher in smaller community hospitals. Future efforts should focus on better understanding staffing and resource challenges faced by community hospitals, and

1 exploring opportunities to build partnerships with public health, government, local entities, and health systems. For example, expansion of tele-ID service and remote 2 infection prevention services to rural and community hospitals will improve access and 3 availability of expertise.³⁸⁻⁴¹ Expanding telehealth alone will not be enough, ideally, it 4 should be accompanied by appropriate resources, infrastructure, and reimbursement 5 models. Medicare's Physician Fee Schedule must also be rebalanced to appropriately 6 compensate ID physicians for value-based care. Additionally, a national plan is needed 7 to strengthen ID workforce in community hospitals to cope with increased complexity of 8 patients.^{3,15} In order to prevent ongoing HAI escalations, health care facilities must 9 invest resources in staff retention and wellbeing, and focus on "recruiting, training, and 10 retaining" our ID workforce. Finally, hospital leaders should further evaluate capacity 11 and infrastructure needed to address COVID-19 burden, invest in ID workforce, and 12 collaborate with policymakers to support enhanced hospital preparedness. 13

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15 NOTES

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- 1 Figure 1: Trend of COVID-19 admissions over baseline (01/01/18 02/29/20) and
- 2 pandemic period (03/01/20-03/31/21) (stratified into three pandemic phases)
- 3
- 4 Figure 2: Segmented Regression Models adjusted for hospital characteristics
- 5 showing CLABSI, CAUTI, CDI and VAE trends in baseline and three pandemic
- 6 phases (blue lines)

Table 1: Impact of COVID-19 pandemic on Incidence Rates of CLABSIS, CAUTIS, VAE, and CDIS

Unadjusted Segmented Regression Analysis showing Level and Trend Changes for baseline and pandemic period

Baseline Trend RR (95% CI)		Pandemic Level Change RR (95% CI)	Pandemic Trend Change RR (95% CI)	
CLABSI	0.996 (0.989-1.003)	1.238* (1.012-1.514)	1.017 (0.993-1.042)	
CAUTI	1.001 (0.991-1.011)	1.030 (0.857-1.238)	0.997 (0.97-1.024)	
VAE	1.007 (0.991-1.022)	1.340* (1.07-1.678)	0.997 (0.976-1.018)	
CDI	0.979* (0.969-0.989)	0.863* (0.742-1.004)	1.042* (1.017-1.068)	

¥ Y		Pandemic Phase 1		Pandemic Phase 2		Pandemic Phase 3	
	Baseline Trend	Level	Trend	Level	Trend	Level	Trend
	RR (95% CI)	Change	Change	Change	Change	Change	Change
		RR (95% CI)	RR (95% CI)	RR (95% CI)	RR (95% CI)	RR (95% CI)	RR (95%
							CI)

					X (
				2			
CLABSI	0.996	0.955	1.162	1.443*	1.001	1.487*	0.975
CLADOI	(0.989-1.003)	(0.732-	(0.993-	(1.060-	(0.906-	(1.095-2.020)	(0.899-
		1.246)	1.361)	1.962)	1.105)		1.057)
CAUTI	1.001	1.086	0.862	1.195	0.998	1.145	0.913
	(0.991-1.011)	(0.79-1.494)	(0.721-	(0.881-	(0.831-	(0.794-1.653)	(0.808-
			1.032)	1.621)	1.198)		1.031)
VAE	1.007	1.339*	0.972	1.429*	0.936	1.533*	0.927*
	(0.991-1.023)	(1.045-	(0.874-	(1.003-	(0.854-	(1.123-2.094)	(0.877-
		1.714)	1.081)	2.035)	1.027)		0.980)
CDI	0.977*	0.956	1.061	0.912	1.032	1.29	1.068
	(0.967-0.987)	(0.719-	(0.926-	(0.715-	(0.906-	(0.998-1.669)	(0.990-
		1.270)	1.215)	1.164)	1.174)		1.151)

*Statistically significant, CLABSI: central-line-associated bloodstream infections, CAUTI: catheter-associated urinary tract infections, CDI: C. difficile infections, VAE: ventilator-associated events, RR: rate ratio, CI: confidence interval



