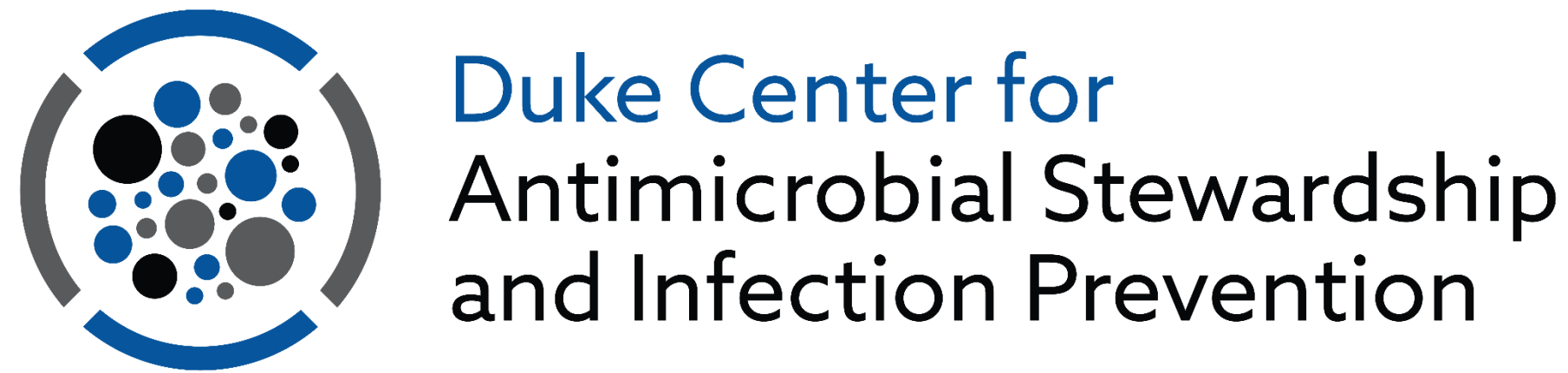


Beta is Better: Impact of a Multifaceted Stewardship Initiative on the Sequential Timing of Beta-Lactam Administration among patients with Bacteremia



Deri CR^{1,2}, Schultheis J¹, Shroba J^{1,2}, Boreyko J³, Wrenn RH^{1,2}, Keener C⁴, Moehring RW^{2,5}
1- Duke University Hospital Department of Pharmacy, Durham, NC, USA; 2- Duke Center for Antimicrobial Stewardship and Infection Prevention, Durham, NC, USA; 3- Duke Regional Hospital Department of Pharmacy, Durham, NC, USA; 4- Clinical Education and Professional Development, Duke University Medical Center, Durham, NC, USA; 5- Division Of Infectious Diseases, Duke University Medical Center, Durham, NC, USA



Background

- Beta-lactams and anti-methicillin-resistant *Staphylococcus aureus* (MRSA) agents are often empiric regimens for patients with sepsis or septic shock
- Appropriate antibiotic sequence (*i.e.* beta-lactam before vancomycin) may reduce early mortality in patients with bloodstream infections (BSIs) (Amoah, et al. CID 2022)
- AIM:** Describe the impact of a multifaceted stewardship initiative on the sequence of antibiotic administration in adults with at least one positive blood culture

Methods

Study Design: Pre-post implementation analysis
Study Period and Comparison Groups:
PRE: March 1, 2021 – May 22, 2022 (~15 months)
POST: May 31, 2022 – Feb 12, 2023 (~8 months)
Setting: Duke University Health System (3 hospitals: 1 university, 2 community)
Primary Outcome: Percent of beta-lactam first administration

Inclusion:
Adult: ≥ 18 years
Concurrent vancomycin and beta-lactam: antibiotic administration within 8 hours of each other
Beta-lactam: cefepime, piperacillin-tazobactam, or meropenem

Statistical Analysis:
Chi-square tests were used to compare rates between groups

Intervention:
1) System-wide adult beta-lactam order panel combining load and maintenance doses for select beta-lactams (Figure 1)
2) Nursing administration instructions facilitating the appropriate sequential order of antibiotic administration (Figure 2)
3) System-wide education to physicians, pharmacists, and nurses

Results

Figure 1. Beta-Lactam Order Panel, piperacillin-tazobactam

ADULT piperacillin-tazobactam (ZOSYN) panel (PREFERRED) Accept

Most patients should be loaded to achieve faster time to adequate serum drug concentrations followed by an extended infusion (EI) maintenance order.

If patients already received loading dose over 30 min in ED, select 'NO loading dose' below and start 4 hours after load if CrCl > 20 mL/min or 8 hours after load if CrCl ≤ 20 mL/min.

☒ Loading dose + maintenance

☐ CrCL > 20 mL/min or CRRT

☐ CrCL ≤ 20 mL/min, peritoneal, or IHD

☐ NO loading dose (ALREADY ADMINISTERED); extended infusion maintenance ONLY

Figure 2. Beta-Lactam First: Nursing Administration Instructions, cefepime

Admin Instructions: 100%

If antibiotics may NOT be administered simultaneously (e.g. IV incompatibility), the initial cefepime dose should be administered before vancomycin unless otherwise specified by provider. Infuse bag over 30 minutes for Traditional Infusion Therapy.

Table 1. Characteristics of Adult Encounters with Positive Blood Cultures Initiated on Combination Therapy

	Pre-intervention (n=224)	Post-intervention (n=137)
Median Age (IQR)	65 (52.8-74.3)	65 (52-74)
Beta-lactam received		
FEP	70 (31.3)	28 (20.4)
MEM	5 (2.2)	0 (0)
TZP	149 (65.5)	109 (79.6)
Blood culture gram-stain		
Gram-positive	147 (65.6)	96 (70.1)
Gram-negative	61 (27.2)	34 (24.8)
Polymicrobial	10 (4.5)	6 (4.4)
Fungus or AFB	6 (2.7)	1 (0.7)

Data reported as n (%) unless otherwise specified
AFB: acid-fast bacilli, FEP: cefepime, MEM: meropenem, TZP: piperacillin-tazobactam

Table 2. Sequence of Antibiotic Administration

	Vancomycin first (n=19)	Beta-lactam first (n=319)	Same time (n=23)	Overall (n=361)
PRE	14 (6.3%)	198 (88.4%)	12 (5.4%)	224
FEP	5	61	4	70
MEM	0	5	0	5
TZP	9	132	8	149
POST	5 (3.6%)	121 (88.3%)	11 (8.0%)	137
FEP	1	27	0	28
MEM	0	0	0	0
TZP	4	94	11	109
OVERALL	19/361 (5.3%)	319/361 (88.4%)	23/361 (6.4%)	

FEP: cefepime, MEM: meropenem, TZP: piperacillin-tazobactam

Figure 3. Timing of Beta-Lactam Administration to Vancomycin by Administration Location

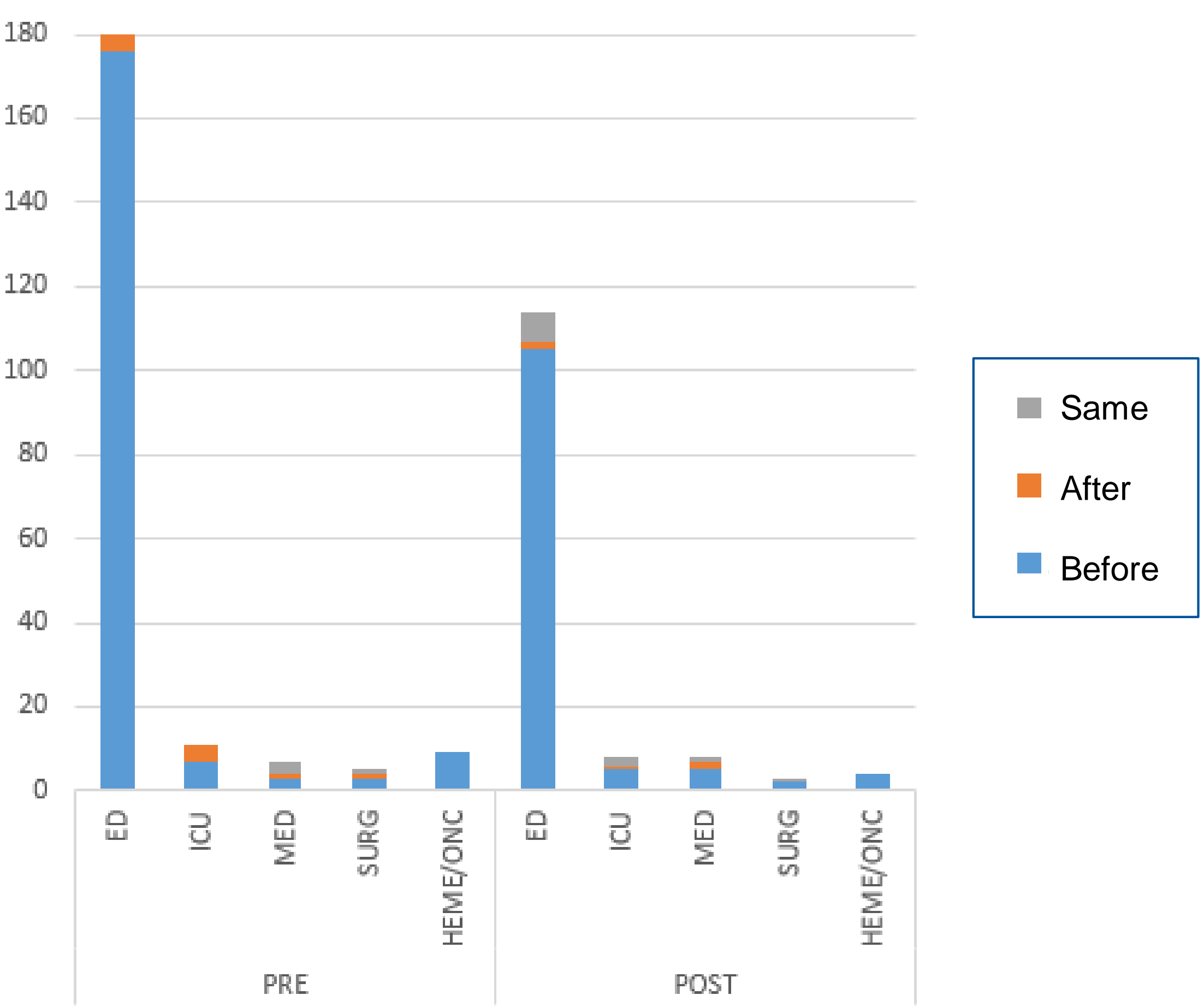
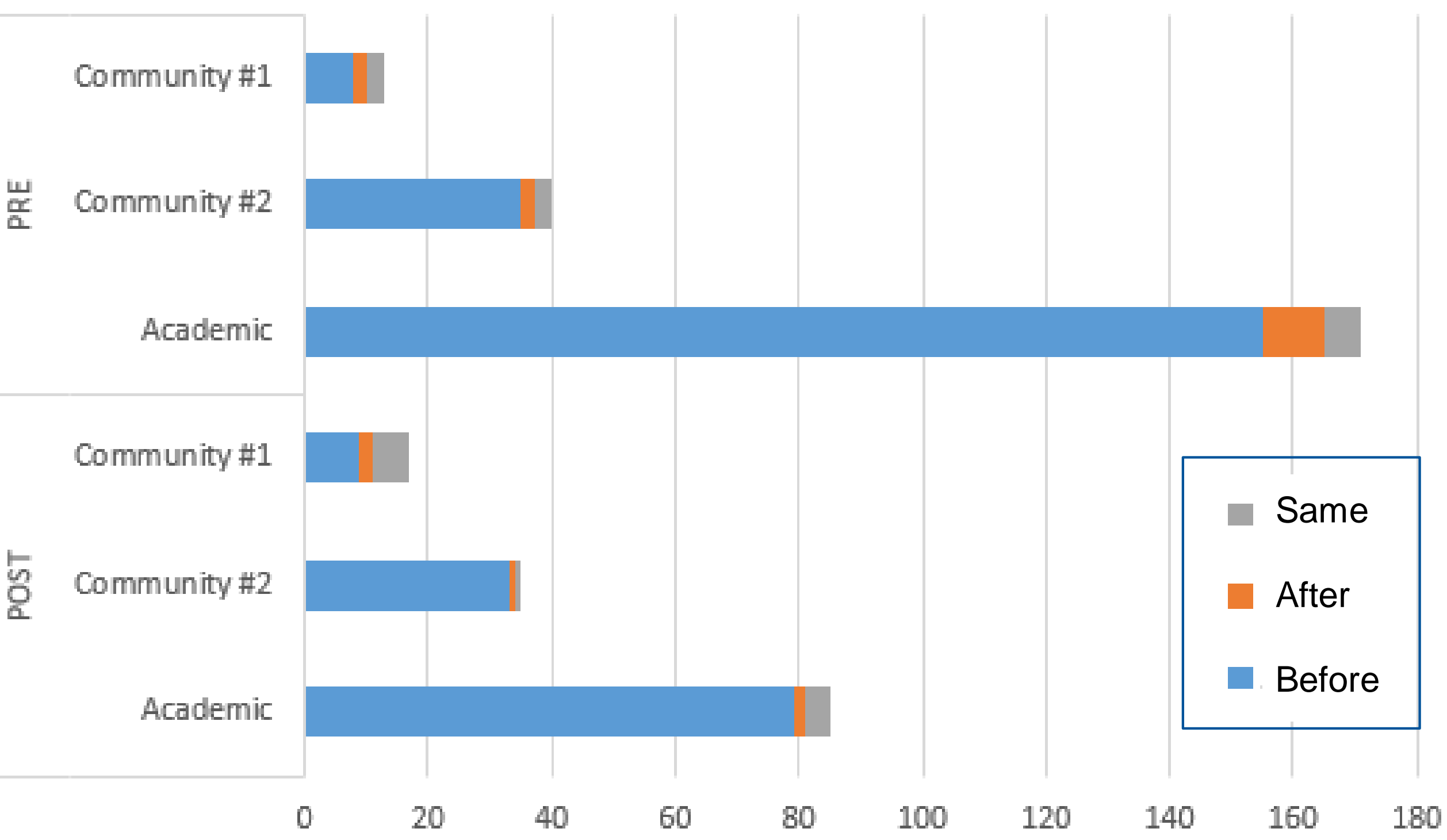


Figure 4. Timing of Beta-Lactam Administration to Vancomycin by Hospital



- Piperacillin-tazobactam (TZP) + vancomycin was the most common combination (71.5%, Table 1)
- Most of the eligible encounters occurred at the academic hospital (70.9%) and the emergency department (ED) (84.8%)
- Receipt of beta-lactam first was high at baseline, then higher post-intervention compared to pre-intervention (96.4% vs 93.8%, p = 0.283) (Table 2)
- Beta-lactam first improvement was most notable in the ICU (63.6% vs 87.5%) and ED (95.8% vs 98.2%)

Conclusion

- A multifaceted stewardship initiative numerically increased the percent of beta-lactam first administrations among patients with positive blood cultures, in the setting of a baseline high beta-lactam first administration rate.
- System change and education regarding antibiotic administration best practices impacted nursing workflow.