

Facility reported vs. CLSI MIC breakpoint comparison of Carbapenem not-susceptible (*Carb-NS*) *Pseudomonas aeruginosa* (PSA) from 2016-2019: A multicenter evaluation

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Background

- CLSI lowered the *P. aeruginosa* (PSA) Carbapenem (Carb) breakpoint minimum inhibitory concentrations (MICs) in 2012.
- It often takes several years for commercial antimicrobial susceptibility testing (AST) device manufacturers and microbiology labs to incorporate revised breakpoints.
- We used the BD Insights Research Database, which includes ~13% of annual hospital admissions in the US¹, to compare facility-reported rates of Carb-NS PSA to those assigned by applying 2012 CLSI MIC breakpoints for a large nationwide collection of isolates from 2016-2020^{2,3}.

Methods

- All adults with a positive non-contaminant PSA culture (first isolate per 30-day period from blood, respiratory, urine, skin/wound, intra-abdominal, or other) in ambulatory and inpatient settings from 298 US hospitals from Q1 2016 to Q4 2020 were evaluated (BD Insights Research Database [Becton, Dickinson & Company, Franklin Lakes, NJ]).
- Facility-reported antimicrobial susceptibility results were based on lab information system feed designations of susceptible (S) or NS (intermediate [I] or resistant [R]) for PSA as S/NS to imipenem (IPM), meropenem (MEM), and doripenem (DOR) per commercial panels.
 - Where available, MICs were interpreted using CLSI 2012 Carb breakpoints ($\mu\text{g/ml}$) of ≤ 2 (S), 4 (I), and ≥ 8 (R). These breakpoints are identical to those in the current CLSI M100 Ed. 31 (2021) standard^{2,3}.

- For evaluable PSA isolates we compared susceptibility results as reported by the facility to those using CLSI MIC breakpoints.
- Two proportion Z tests were used to assess the difference between facility-reported and revised susceptibility results overall and by hospital demographics. P-values < 0.05 indicate a significant difference between facility-reported and revised CLSI MIC breakpoints in NS determinations.

Results

- Overall, 86.9% (255,844/294,426) of non-duplicate PSA isolates with facility reported IPM/MEM/DOR susceptibility interpretations also had interpretable MIC results. These were termed the evaluable populations.
- S rates were 84.9% and 83.3% as reported by facilities and determined by CLSI criteria, respectively (Figure 2).
- Facilities under-reported Carb-NS by 9.8% compared with CLSI criteria; 10.4% and 7.7% of R and I isolates, respectively, were missed by facility reporting (Table 1).

Results

Figure 1. NS evaluations in PSA for imipenem/meropenem/doripenem.

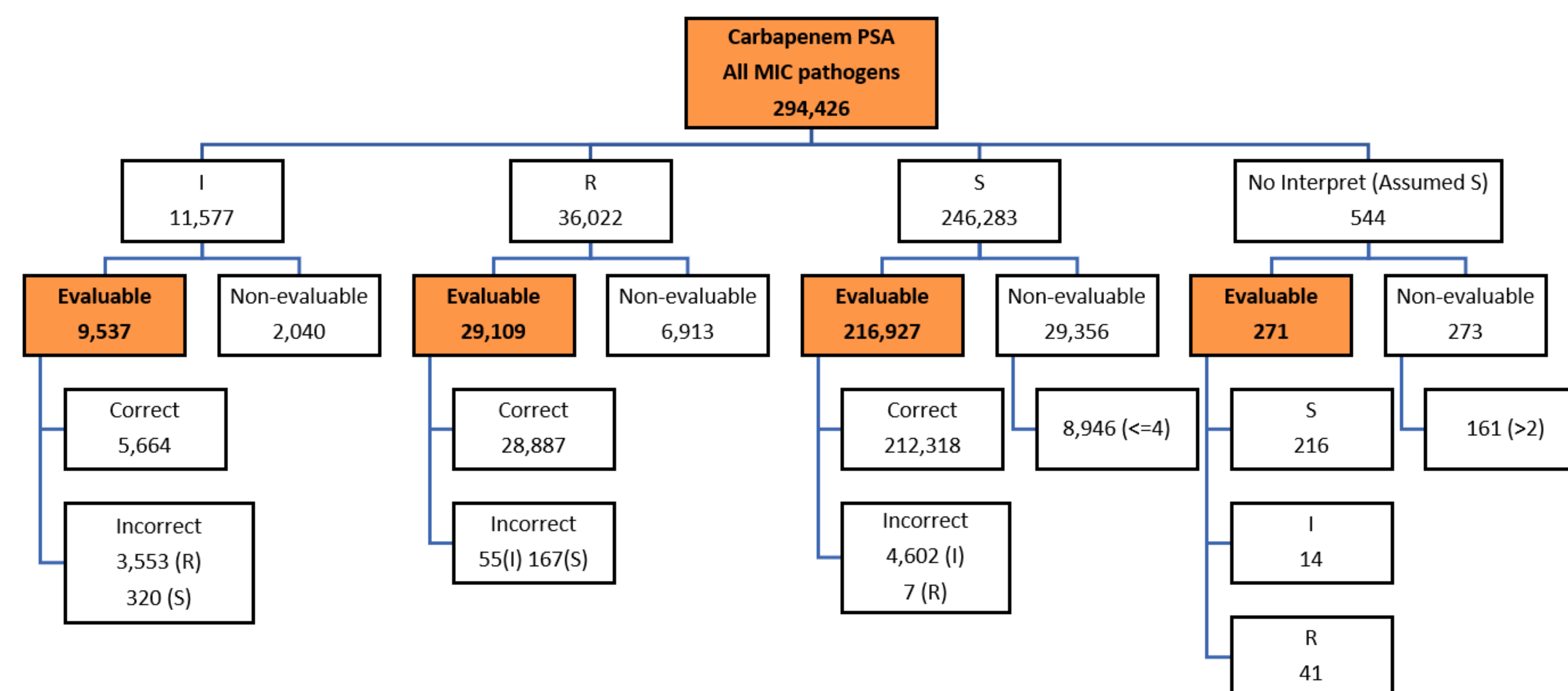


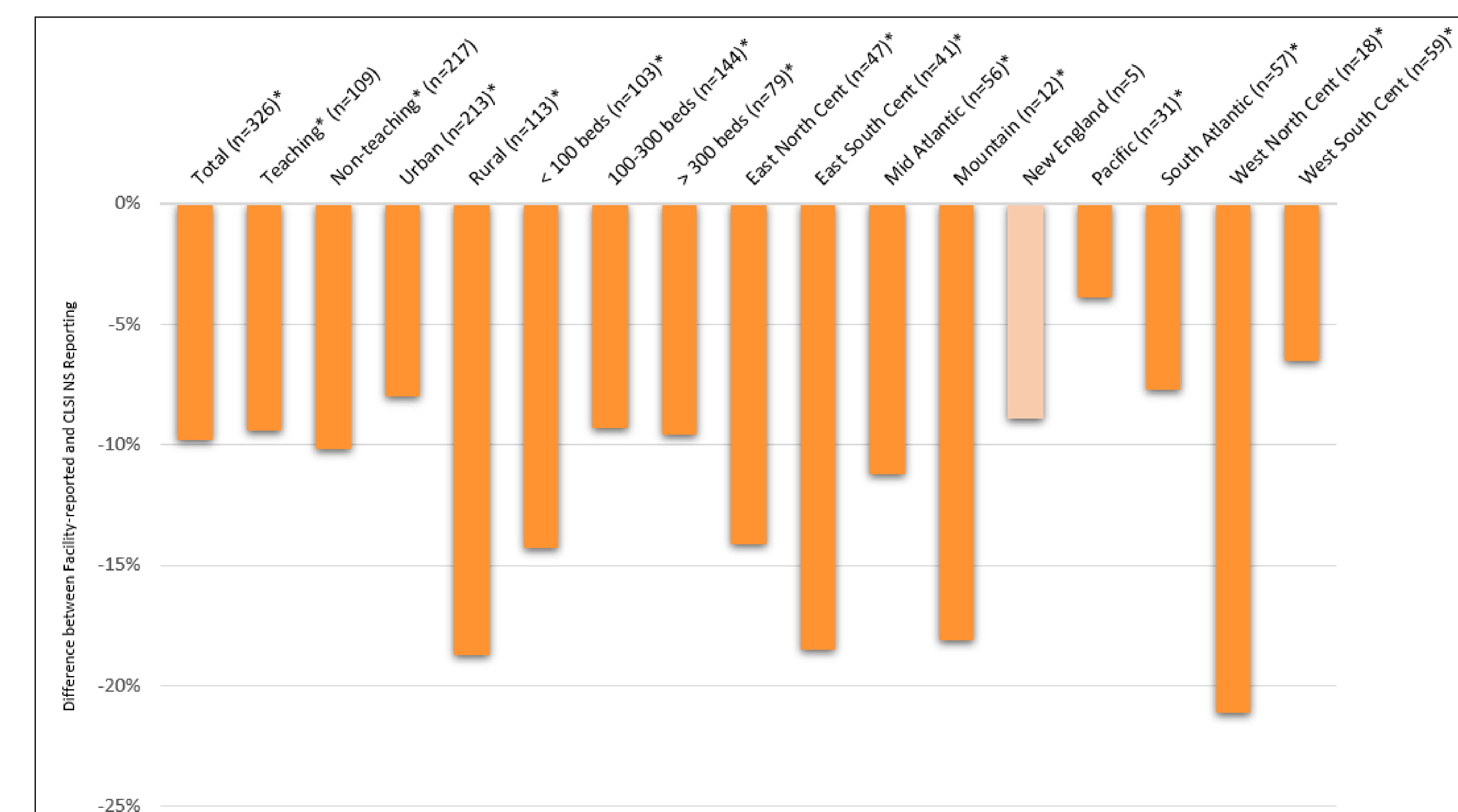
Table 1. Comparison of S/NS for facility-reported and CLSI imipenem/meropenem/doripenem breakpoints in PSA.

IPM/MEM/DOR interpretations for PSA			
Interpretation (MIC; $\mu\text{g/mL}$)	Facility Reported: n (%)	Revised per CLSI: n (%)	Underreporting by Facility vs. Revised per CLSI (%)
I (4)	9,537 (3.7%)	10,335 (4.0%)	↓ 7.7%
R (≥ 8)	29,109 (11.4%)	32,488 (12.7%)	↓ 10.4%
S (≤ 2)	217,198 (84.9%)	213,021 (83.3%)	

References

- Centers for Disease Control and Prevention. Antibiotic resistance threats in the United States, 2019. November 2019. Available at: www.cdc.gov/DrugResistance/Biggest-Threats.html. Accessed September 16, 2021.
- CLSI. Performance Standards for Antimicrobial Susceptibility Testing. 31st ed.
- CLSI supplement M100. Clinical and Laboratory Standards Institute; 2021

Figure 2. Carb NS underreporting by hospital demographics (n = facility count; * p < 0.03 [darker color]) for imipenem/meropenem/doripenem.



Conclusions

- Systematic application of CLSI breakpoints in 2016-20 would have had minimal impact on PSA S rates in the US.
- However, facility reporting failed to identify ~10% of Carb-NS isolates in this study's subset of US hospitals. This discordance was significant across hospital demographics and regions.
 - Smaller hospitals (<100 beds) in rural areas may be particularly at-risk of failure to identify Carb-NS
- Facilities should know their local epidemiology, decide if under-reporting might be an issue, and then assess if there is any impact on their patients.
- Further analyses are required to understand whether the Carb-NS reporting discordances as seen in this analysis also apply more broadly to other hospitals in the US.

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