

## Background

Vancomycin has been the mainstay of treatment for methicillin-resistant Staphylococcus aureus (MRSA) infective endocarditis (IE). MRSA reduced susceptibility to vancomycin is a growing threat. Data assessing the effect of vancomycin reduced susceptibility on outcomes of MRSA related IE is limited. Our study aimed to evaluate characteristics and outcomes of MRSA-related IE based on the minimum inhibitory concentration (MIC) to vancomycin

# Methods

We conducted a retrospective cohort study at a tertiary care center. Records of hospitalized adults with a diagnosis of IE by ICD-9/ICD-10 CM codes were identified from 2011 to 2018. Patients with "definitive" or "possible" IE based on the modified Duke criteria were included. 51 patients had MRSA-related IE and were selected for the analysis. Demographic, microbiologic, Imaging and outcome variables were obtained. Characteristics and outcomes of patients with MRSA-related IE according to the MIC to vancomycin ( $\leq$  vs. > 1 mcg/mL) were compared. IRB approval was obtained.

### Results

51 patients (20.9%) had IE due to MRSA, among which 18 (35.3%) had MRSA with a MIC to vancomycin > 1 mcg/mL. 59% were men and mean age was 46±3 years old. 65% of patients with MRSA-related IE acquired the infection through injection drug use. Only 3.9% of patients had prosthetic valve

# 8 Years of Characteristics and Outcomes of Patients with MRSA Endocarditis Based on Vancomycin Minimum Inhibitory Concentration: Experience at a Tertiary Care Hospital

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> **Table 1**. Characteristics and outcomes of patients with MRSA-related
>  endocarditis according to the minimal inhibitory concentration (MIC) to vancomycin

### Variable

Mean age, years Men, n (%) Intravenous drug use, n (%) **Co-infections** HIV, n (%) HCV, n (%) HBV, n (%) **Presence of vegetations** Tricuspid valve, n (%) Mitral valve, n (%) Aortic valve, n (%) Intra-cardiac complications Valvular abscess, n (%) Valvular perforation, n (%) Intra-cardiac fistula, n (%) Presence of central venous catheters, n (%) Port-A-Cath, n (%) Tunneled catheter, n (%) **Peripherally inserted center** catheter, n (%) **Cardiac surgery** During index hospitalization, n (%) Within 6 months, n (%) Death During index hospitalization, n (%) Within 90 days, n (%)

| Vancomycin   | Vancomycin  | Ρ     |
|--------------|-------------|-------|
| $MIC \leq 1$ | MIC > 1     | value |
| mcg/mL       | mcg/mL      |       |
| N=33 (65%)   | N=18 (35%)  | 0.40  |
| $48 \pm 19$  | $44 \pm 19$ | 0.49  |
| 20 (60.6)    | 10 (55.6)   | 0.73  |
| 21 (63.6)    | 12 (66.7)   | 0.83  |
|              |             |       |
| 4 (19.0)     | 0 (0.0)     | 0.14  |
| 11 (68.8)    | 6 (54.5)    | 0.45  |
| 0 (0.0)      | 1 (7.1)     | 0.19  |
|              |             |       |
| 10 (30.3)    | 8 (44.4)    | 0.31  |
| 8 (24.2)     | 5 (27.8)    | 0.78  |
| 9 (27.3)     | 2 (11.1)    | 0.18  |
|              |             |       |
| 2 (6.1)      | 3 (16.7)    | 0.22  |
| 1 (3.0)      | 2 (11.1)    | 0.24  |
| 1 (3.0)      | 2 (11.1)    | 0.24  |
| 2 (6.1)      | 3 (16.7)    | 0.22  |
|              |             |       |
| 1 (3.0)      | 0 (0.0)     | 0.46  |
| 1 (3.0)      | 0 (0.0)     | 0.46  |
| 0 (0.0)      | 2 (11.1)    | 0.05  |
|              |             |       |
|              |             |       |
| 5 (15.2)     | 2 (11.1)    | 0.69  |
| 4 (12.1)     | 2 (11.1)    | 0.92  |
|              |             |       |
| 5 (15.2)     | 2 (11.1)    | 0.69  |
| 4 (12.5)     | 2 (11.1)    | 0.88  |

33.3% had positive serology for hepatitis C infection and 13% were HIV positive. 35.3% of patients had tricuspid valve vegetations, 25.5% had mitral valve vegetations, and 21.6% had aortic valve vegetations. Two patients had IE possibly related to a PICC-line infection; both patients had MIC to vancomycin >1 mcg/mL, suggestive of prolonged antibiotic therapy. All patients with MRSA-related IE were started empirically on vancomycin. Patients with a MIC to vancomycin > 1 mcg/mL were more likely to be switched to a combination of daptomycin and ceftaroline, compared to those with a MIC  $\leq$  1 mcg/mL (44.4% vs. 6.1%; P=0.001). Thirteen patients (25.4%) underwent valvular replacement within 6 months, 7 of which had the surgery during the index hospitalization. Six patients (12%) died within 90 days. MRSA-related IE with a MIC to vancomycin > 1mcg/mL did not confer and an increase risk in in-hospital mortality (11.1%) vs. 15.2%; P=0.67) or mortality at 90 days (11.1% vs. 12.5%; P=0.89).

In this single-center experience conducted in a modern era, 35% of patients with MRSA-related IE had MIC>1 mcg/mL. Although this is an alarming finding, we didn't find an increase in valvular surgery requirement or mortality among those with MIC > 1mcg/d as compared to those with a more sensitive MRSA strain. A study with more power or a meta-analysis will be required to better answer this question.

#### **ABBREVIATIONS:**



# Conclusion