

## Clinical outcomes of ESBL producing Enterobacteriaceae bacteremia with non-carbapenem antibiotics

제주대학교병원 감염내과<sup>1</sup>, 제주대학교 의과대학 내과학교실<sup>2</sup>

오현주<sup>1</sup>, 김미선<sup>1</sup>, 허상택<sup>1,2</sup>, 유정래<sup>1,2</sup>

**Background/Aims:** Carbapenem antibiotics are commonly recommended for treating extended-spectrum  $\beta$ -lactamase producing Enterobacteriaceae (ESBL-E) bacteremia however, the threat of carbapenem-resistant Enterobacteriaceae continues to be raised. In addition, studies on the clinical outcomes of treatment with non-carbapenem antibiotics on patients with ESBL-E bacteremia are still ongoing.

**Methods:** We conducted a retrospective propensity score-matched (PSM) case-control study to evaluate the effect of definitive treatment with carbapenem vs. non-carbapenem on the outcome of patients with bacteremia due to ESBL-E. Using electronic medical records data, we screened blood cultures positive for Enterobacteriaceae from January 1, 2021, to December 31, 2021, in a single academic hospital and included ESBL-E bacteremia cases. Antimicrobial susceptibility and phenotypic ESBL-E identification were tested by the Vitek 2 (BioMérieux).

**Results:** The baseline characteristics including gender, age, residents in long-term care facilities, the number of cases of prior antibiotic uses, primary infection source, and isolated strains were similar between the two groups after PSM. The most frequently isolated strain was *Escherichia coli* (38 [67.9%]) in the carbapenem group vs. 41 [73.2%] in the non-carbapenem group). The most common primary infection source was urinary tract infection in both groups. The composite treatment failure was 9 (16.4%) in the carbapenem group and 14 (25.9%) in the non-carbapenem group ( $p=0.323$ ). There were no significant differences between the carbapenem group and the non-carbapenem group in terms of 30-day all-cause mortality (4 [7.7%] vs. 6 [11.8%]). In addition, the two groups of microbiologic failure, clinical failure, acute kidney injury, and duration of antibiotics were similar.

**Conclusions:** In this study, we used the propensity-score matching method to compensate for the limitation of the retrospective observational study. In summary, the current study showed no significant differences between the carbapenem group and the non-carbapenem group regarding the clinical outcomes for treating ESBL-E bacteremia.

Table 1. Baseline characteristics of ESBL-E bacteremia cases during the study period

Characteristics	Unmatched		Propensity score Matched	
	Carbapenem (n=62)	Non-carbapenem (n=56)	Carbapenem (n=56)	Non-carbapenem (n=56)
Male, n (%)	34 (54.8%)	20 (35.7%)	30 (53.6%)	20 (35.7%)
Age, median (IQR), yr	79.0 [68.0;84.0]	83.0 [71.0;87.0]	79.5 [67.0;84.5]	83.0 [71.0;87.0]
BMI, mean $\pm$ SD, kg/m <sup>2</sup>	22.8 $\pm$ 4.7	22.1 $\pm$ 4.4	22.6 $\pm$ 4.7	22.1 $\pm$ 4.4
Long term care facility, n (%)	18 (29.0%)	26 (46.4%)	9 (34.6%)	8 (47.1%)
Prior antibiotics use, n (%)	24 (38.7%)	11 (19.6%)	18 (32.1%)	11 (19.6%)
Prior hospital stay, n (%)	23 (37.1%)	12 (21.4%)	17 (30.4%)	12 (21.4%)
HAI, n (%)	16 (25.8%)	14 (25.0%)	13 (23.2%)	14 (25.0%)
Underlying diseases, n (%)				
Diabetes mellitus	23 (37.1%)	22 (39.3%)	20 (35.7%)	22 (39.3%)
Immunocompromised	3 (4.8%)	1 (1.8%)	1 (1.8%)	1 (1.8%)
Chronic kidney disease	4 (6.5%)	4 (7.1%)	3 (5.4%)	4 (7.1%)
Indwelling urinary catheter	9 (14.5%)	5 (8.9%)	9 (16.1%)	5 (8.9%)
Biliary drainage	1 (1.6%)	6 (10.7%)	1 (1.8%)	6 (10.7%)
Central venous catheter	2 (3.2%)	3 (5.4%)	2 (3.6%)	3 (5.4%)
Malignancy	22 (35.5%)	17 (30.4%)	20 (35.7%)	17 (30.4%)
Pitt bacteria score	1.0 [0.0; 2.0]	1.0 [0.0; 2.0]	1.0 [0.0; 2.0]	1.0 [0.0; 2.0]
ICU admission	20 (32.3%)	16 (28.6%)	17 (30.4%)	16 (28.6%)
Length of hospital stay, median (IQR), days	16.0 [11.0;28.0]	13.0 [7.5;17.5]	16.0 [11.0;26.5]	13.0 [7.5;17.5]
Isolated strain				
<i>Escherichia coli</i>	43 (69.4%)	41 (73.2%)	38 (67.9%)	41 (73.2%)
<i>Klebsiella pneumoniae</i>	16 (25.8%)	9 (16.1%)	15 (26.8%)	9 (16.1%)
<i>Enterobacter cloacae</i>	3 (4.8%)	2 (3.6%)	3 (5.4%)	2 (3.6%)
<i>Proteus mirabilis</i>	0 (0.0%)	3 (5.4%)	0 (0.0%)	3 (5.4%)
<i>Morganella morganii</i>	0 (0.0%)	1 (1.8%)	0 (0.0%)	1 (1.8%)
Primary infection sources				
Urinary tract infection	34 (54.8%)	35 (62.5%)	34 (60.7%)	35 (62.5%)
Biliary infection	11 (17.7%)	9 (16.1%)	10 (17.9%)	9 (16.1%)
Intra-abdominal infection	7 (11.3%)	7 (12.5%)	6 (10.7%)	7 (12.5%)
Bone and soft tissue infection	3 (4.8%)	2 (3.6%)	2 (3.6%)	2 (3.6%)
Infective endocarditis	2 (3.2%)	1 (1.8%)	1 (1.8%)	1 (1.8%)
Others	5 (8.1%)	2 (3.6%)	3 (5.4%)	2 (3.6%)

Others includes catheter relate blood stream infection, pneumonia, and unknown source of infection

Prior hospital stay: within 30-day prior to study period

BMI, body mass index; HAI, hospital acquired infection; ICU, intensive care unit; IQR, interquartile range; SD, standard deviation

Table 2. Outcomes according to the definitive antibiotics

Variables	Unmatched		p-value	Propensity score-matched		p-value
	Carbapenem (n=62)	Non-carbapenem (n=56)		Carbapenem (n=56)	Carbapenem (n=62)	
Primary outcome						
Composite treatment failure, n (%)	11 (18.0%)	14 (25.9%)	0.457	9 (16.4%)	14 (25.9%)	0.323
Secondary outcomes						
30-day all-cause mortality, n(%)	5 (8.8%)	6 (11.8%)	0.846	4 (7.7%)	6 (11.8%)	0.526
Microbiologic failure, n(%)	3 (5.8%)	2 (4.7%)	1.000	3 (6.1%)	2 (4.7%)	1.000
Clinical failure, n(%)	2 (7.1%)	3 (13.0%)	0.647	2 (7.7%)	6 (10.7%)	0.271
AKI, n(%)	13 (22.0%)	11 (20.8%)	1.000	11 (20.8%)	11 (20.8%)	1.000
Duration of antibiotics, median (IQR)	14.0 [12.0;18.0]	14.0 [12.0;16.0]	0.491	14.0 [12.5;18.0]	14.0 [12.0;16.0]	0.564

AKI, acute kidney injury; IQR, interquartile range