Species belonging to the Hijikata Morita, including 24 SUD-resistant strains, and one CLSI QC strain. BMD was MD® are a major Nakao Yam at e The panel consisted of 204 1, MD, PhD, Tomohide Iizuka, Fukuoka, Japan; Yuji Osato, Osaka, Japan; and Ohnishi Komatsugawa, Tokyo, Japan S430 • OFID 2021:8 (Suppl 1) • Abstracts

656. Sulbactam-Durlobactam MIC Determination: Comparative Evaluation of the New ETEST SUD to the CLSI 2021 Broth Microdilution Method Veronique Sauvonnet; Elodie Escotter; n/a; Christine Franceschi, n/a; Diane Halimi, MA; Roland Martelin, MS; Laetitia Pellet, n/a; Gilles Zambardi; Biomerieux SA, La Balme Les Grottes, Rhone-Alpes, France Session: P-29. Diagnostics: Bacteriology/mycobiology Background. Species belonging to the Acinetobacter baumannii-calcoaceticus (ABC) complex, such as A. baumannii, A. pittii, and A. nosocomialis, are a major cause of hospital acquired infections and outbreaks with increasing occurrence of multidrug-resistance. Sulbactam-durlobactam (SUD), a combination of one active β-lactam antibiotic (sulbactam) with a new β-lactamase inhibitor (durlobactam), is currently being tested in a phase 3 clinical trial by Entasis Therapeutics for the treatment of serious infections caused by ABC, including multidrug-resistant strains. At the same time, an ETEST SUD (sulbactam-durlobactam - MIC range 0.08/4-64/4 µg/mL) has been developed and calibrated versus the broth microdilution reference method (BMD) as described by the Clinical and Laboratory Standards Institute (CLSI). This test is intended to determine the MIC of sulbactam-durlobactam for species of the ABC complex. The aim of this study was to perform a first comparative study of ETEST SUD with the CLSI BMD method on a panel of 263 isolates.

Methods. The panel consisted of 204 A. baumannii, 29 A. pittii, 30 A. nosocomialis, including 24 SUD-resistant strains, and one CLSI QC strain. BMD was performed using the 2021 CLSI guidelines. ETEST SUD was evaluated using the standard ETEST procedure for Acinetobacter baumannii spp. (inoculum 0.5 McFarland, Mueller Hinton medium, incubation at 35°C for 20-24h). For each method, the MIC was read at complete inhibition of visible growth. To determine category agreement (CA) and error rates, the sulbactam-durlobactam provisional breakpoint point of 4 µg/mL was applied.

Results. The QC strain MICs were in the expected range with reproducible results. The essential MIC agreement [EA, ±1 dilution] was 97.7% without any tendency to over- or underestimate the MIC when compared to BMD. The CA was 98.5%. Two Very Major Errors, both within the EA, and two Major Errors, one within the EA, were observed.

Conclusion. In this study, the ETEST SUD was found to be equivalent to the CLSI reference method. MIC end points were easy to read. With a 15-dilution range and simplicity of use, ETEST SUD could represent a valuable tool for MIC determination and could be an alternative to BMD.

Disclosures. Gabriella S. Lamb, MD, MPH, Nothing to disclose

657. Genomic Insights into Virulence Factors Affecting a Tissue-invasive Klebsiella pneumoniae Infection Takashi Matono, MD, PhD; Daisuke Masaoka, PhD; Ryosuke Kikukawa, MD; Yuji Teshima, MD; Masato Nakao, MT; Ryououke Yamate, MD; Takamichi Hikijita, MD; Kosuke Hoashi, MD; Yuichi Hasegawa, MD; Tomohide Iizuka, MD; Makoto Ohnishi, MD, PhD; Aso Iizuka Hospital, Iizuka, Fukuoka, Japan; National Institute of Infectious Diseases, Shinyaku, Tokyo, Japan Session: P-29. Diagnostics: Bacteriology/mycobacteriology Background. Japan is one of the hypervirulent Klebsiella pneumoniae (hvKp) endemic areas, resulting in an alarming issue in actual clinical settings. However, little is known regarding key virulence factors responsible for hvKp infection.

Methods. We analyzed K. pneumoniae isolates collected between 2017 and 2019, and defined hvKp as a pyrogenic infection. Classical K. pneumoniae (cKp) involved a non-invasive infection or uncomplicated bacteremia. Isolates belonging to the K. pneumoniae species complex were excluded.

Results. We analyzed 112 isolates, including 19 hvKp, 67 cKp, and 26 colonizers, by whole-genome sequencing. Population genomics revealed that the K1-strain genotype (ST) 82 clade was distinct from that of K1-ST23 clone (Figure 1). The virulence gene profiles also differed between K1-ST82 (aerobactin and rmpA) and K1-ST23 (aero- bactin, yersiniabactin, salmochelin, colibactin, and rmpA/rmpA2). The K2 genotype was more diverse than that of K1. A neighboring subclade of K1-ST23 (comprising ST29, ST412, ST36, and ST268) showed multidrug-resistance and hypervirulence potentials. Logistic-regression analysis revealed that diabetes mellitus was associated with K. pneumoniae infection (odds ratio [OR]: 4.11; 95% confidence interval [CI]: 1.14–14.8). No significant association was found between hvKp diagnosis and clinical characteristics, such as diabetes mellitus or community acquisition (Table 1). The K1 genotype (OR: 9.02; 95% CI: 2.49–32.7; positive-likelihood ratio [LR]: 4.08), rmpA (OR: 8.26; 95% CI: 1.77–38.5; positive LR: 5.83), and aerobactin (OR: 4.59; 95% CI: 1.22–17.2; positive LR: 3.49) were substantial diagnostic predictors of hvKp (Table 2).

Figure 1. Phylogenetic distribution of genetic virulence factors in 112 K. pneumoniae isolates

Table 1. Variables analyzed for predicting hvKp infection

| Variables                      | OR (95% CI) | p value |
|--------------------------------|-------------|---------|
| Diabetes mellitus              | 1.49 (0.52–4.23) | 0.46    |
| Liver cirrhosis                | 12.4 (1.21–127) | 0.034   |
| Community-acquired positive    | 1.34 (0.48–3.73) | 0.58    |
| string test                    | 4.07 (1.08–15.3) | 0.038   |
| K1                             | 9.02 (2.49–32.7) | 0.001   |
| K2                             | 0.94 (0.32–2.82) | 0.92    |
| Aerobactin                     | 4.59 (1.17–17.2) | 0.024   |
| Yersiniabactin                 | 2.11 (0.74–6.04) | 0.16    |
| Salmochelin                    | 2.56 (0.83–7.91) | 0.11    |
| Colibactin                     | 1.86 (0.63–5.52) | 0.26    |
| rmpA                           | 8.20 (1.77–38.5) | 0.007   |
| rmpA2                          | 1.26 (0.44–3.37) | 0.71    |

hvKp, hypervirulent K. pneumoniae; OR, odds ratio; CI, confidence interval

Table 2. Microbiological diagnostic predictive values for hvKp

| Characteristics | Sensitivity (%) | Specificity (%) | PPV (%) | NPV (%) | LR+ | LR- |
|----------------|----------------|-----------------|---------|---------|-----|-----|
| Positive string test | 84.2 | 43.3 | 29.6 | 90.6 | 1.49 | 0.37 |
| K1 genotype | 61.5 | 84.9 | 42.1 | 92.5 | 4.08 | 0.45 |
| Aerobactin | 30.8 | 91.2 | 84.2 | 46.3 | 3.49 | 0.76 |
| rmpA | 33.3 | 94.3 | 89.5 | 49.3 | 5.83 | 0.71 |

hvKp, hypervirulent K. pneumoniae; PPV, positive predictive value; NPV, negative predictive value; LR, likelihood ratio

Disclosures. All Authors: No reported disclosures