

bacteremia (80%, 52/65). Bacteremia was cleared in 90.8% (59/65) of patients and the 30-day mortality rate was 15.4% (10/65). Median time to bacteremia clearance after combination switch was 3 days. Eleven patients received DAP-CPT within 72 hours of index culture. Median time to bacteremia clearance for patients switched to DAP-CPT within 72 hours versus after 72 hours did not differ (2 vs 3 days; $P = 0.526$), however the overall median duration of bacteremia was 4 and 11 days ($P = 0.018$). In a sub analysis, the median time of bacteremia clearance following combination therapy was significantly longer for patients receiving renal replacement therapy (5 vs 2 days; $P = 0.04$).

Conclusion. There were no independent predictors of 30-day mortality identified. DAP-CPT combination therapy resulted in clearance of persistent bacteremia and may serve as an effective salvage therapy.

Disclosures. All Authors: No reported disclosures

1625. Real-World Outpatient Utilization of Ceftolozane/Tazobactam in Physician Office Infusion Centers (OICs)

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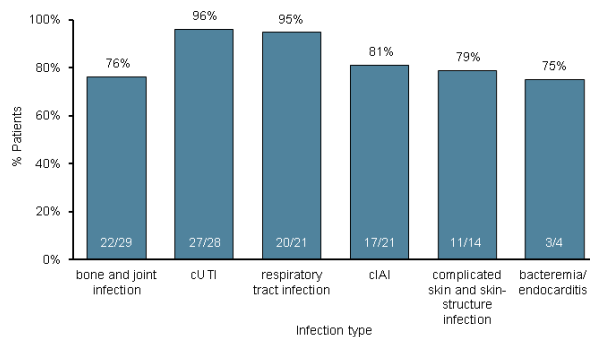
Session: P-71. Treatment of Antimicrobial Resistant Infections

Background. Ceftolozane/tazobactam (C/T) is indicated for the treatment (tx) of complicated Gram-negative infections including urinary tract infection (cUTI), intra-abdominal infection (cIAI), and hospital-acquired/ventilator-associated bacterial pneumonias caused by susceptible bacteria. Real-world data on the use of C/T are limited. We present a multicenter observational review of C/T outpatient utilization in Infectious Disease OICs.

Methods. Medical records of patients (pts) who received C/T for ≥ 3 doses from May 2015 to Sept 2019 were reviewed. Data included demographics, diagnosis, disease history, pathogens, C/T tx hospitalizations, emergency department (ED) visits and clinical outcomes. Clinical success was defined as complete or partial symptom resolution at completion of C/T with oral antibiotics as needed. Persistent infection and early discontinuation (D/C) of C/T were deemed non-successful. Indeterminant outcomes were deemed non-evaluable. Chi Square, Fisher's exact, and t-tests were used to identify characteristics associated with clinical outcome.

Results. 120 pts (mean age: 59 ± 15 years, 60% male) from 33 OICs were identified. Median Charlson score was 5 (IQR, 3-7), with 43% immunocompromised, and 77% refractory/recurrent disease. Primary infections were bone and joint (25%), cUTI, (24%), respiratory tract (18%), cIAI (18%), complicated skin and skin-structure (12%), and bacteremia/endocarditis (3%). Most pts had multi-drug resistant Gram-negative pathogens (80/108; 74%), predominantly *Pseudomonas aeruginosa*. Polymicrobial infections were reported in 44%. Median duration of C/T therapy was 21 days (IQR, 14-34). C/T was initiated in the OIC in 59% of pts. Overall clinical success was 86% (100/117), with rates by infection type in Fig 1. Non-success was reported in 17, 10 due to persistent infection and 7 due to adverse events. The adverse events led to early D/C of C/T, all with resolution. Statistically, infection type did not impact success rate. Hospitalizations and ED visits during tx occurred in 5% of pts with successful outcomes and 35% of pts with non-successful outcomes ($p < 0.001$).

Fig 1. Clinical success rates of C/T by infection type



Conclusion: These real-world results support the effectiveness of C/T in a wide variety of complicated Gram-negative infections treated in the outpatient setting.

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1626. Synergistic Effect of Cefiderocol with Other Antibiotics Against PER-Producing *Acinetobacter baumannii* Isolates from the Multinational SIDERO-WT Studies

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Session: P-71. Treatment of Antimicrobial Resistant Infections

Background. Cefiderocol (CFDC), a novel siderophore cephalosporin, showed potent activity at minimum inhibitory concentrations (MICs) of ≤ 4 $\mu\text{g/mL}$ against $\geq 99\%$ of Gram-negative isolates in the multinational SIDERO-WT studies. PER-producing *Acinetobacter baumannii*, mainly from Russia, showed high CFDC MICs of $8 - >64$ $\mu\text{g/mL}$. This study evaluated the synergistic effects of CFDC combined with other antibiotics against PER-producing *A. baumannii* isolates with high CFDC MICs.

Methods. Two isolates of PER-producing *A. baumannii* with resistance to CFDC (MIC 16 $\mu\text{g/mL}$), meropenem (MEM; MIC 64 $\mu\text{g/mL}$), ceftazidime-avibactam (CZA; MIC 64/4 $\mu\text{g/mL}$), amikacin (AMK; MIC 32 or 64 $\mu\text{g/mL}$), and ciprofloxacin (CIP; MIC ≥ 64 $\mu\text{g/mL}$) were tested. Against ampicillin-sulbactam (SAM), one isolate was resistant (MIC 32/64 $\mu\text{g/mL}$) and another was susceptible (MIC 8/16 $\mu\text{g/mL}$). Effects of CFDC combined with other antibiotics were evaluated by checkerboard assay and chemostat model reproducing humanized antibiotic exposure. The checkerboard assay used a single agent (e.g. ceftazidime [CAZ], avibactam [AVI], ampicillin [AMP] or sulbactam [SUL]). Iron-depleted cation-adjusted Mueller-Hinton broth was used as the standard medium for CFDC, as recommended by the Clinical Laboratory and Standards Institute.

Results. Against both isolates, synergy with CFDC was seen for two β -lactamase inhibitors, AVI and SUL, with a fractional inhibitory concentration (FIC) index of 0.026-0.033 and 0.26-0.27, respectively. A synergistic to additive effect was seen for MEM and AMK, with an FIC index of 0.53-0.75 and 0.25-0.52, respectively. In the chemostat model, regrowth during 24-h treatment was observed with single agents (CFDC 2 g, q8h, 3-h infusion; MEM 2 g, q8h, 1-h infusion; CZA 2 g, q8h, 2-h infusion; SAM 3 g, q8h, 3-h infusion; AMK 15 mg/kg, q8h, 3-h infusion) for both isolates, including the SAM-susceptible isolate. However, no regrowth was seen when CFDC was combined with CZA, MEM, SAM or AMK.

Conclusion. The most potent synergy was seen between CFDC and AVI against PER-producing *A. baumannii* with a decreased MIC to ≤ 1 $\mu\text{g/mL}$ for all isolates, followed by SUL and MEM. Under humanized pharmacokinetic exposure, combination of CFDC and CZA, MEM, SAM or AMK is expected to be effective against PER-producing *A. baumannii* in spite of high CFDC MICs.

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1627. Tedizolid is Well-tolerated Among Patients Receiving Prolonged Treatment Courses, Including Those Who are Intolerant of Alternative Agents

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Session: P-71. Treatment of Antimicrobial Resistant Infections

Background. Tedizolid (TZD) is approved for acute bacterial skin and skin structure infections (ABSSI), but often used for complicated infections to avoid linezolid (LZD) adverse events (AE), particularly when long-term treatment is indicated. This study aimed to characterize the tolerability of TZD, including patients (pts) receiving prolonged treatment.

Methods. Retrospective review of pts who received TZD > 72 hours. Thrombocytopenia was defined as a 50% decrease from baseline platelet count. Favorable clinical outcome was defined as completing therapy without an AE or hospital readmission within 30 days.

Results. 86 pts accounting for 102 courses were included. Median age of pts was 57 years and 43% were immunocompromised. Median duration of TZD therapy was 8 days (range: 4 - 350) and 32% of courses were >14 days. Common indications were ABSSI (n=42), bacteremia (n=15), intra-abdominal infection (n=11), and pneumonia (n=10). 47% and 5% of courses were associated with MRSA or VRE and M. abscessus, respectively. 44% of TZD courses were preceded by treatment failure or AE associated with alternative therapies. AEs attributed to LZD were documented in 13 patients: thrombocytopenia (n=11), lactic acidosis (n=1), or both (n=1). Serotonergic agents were administered during 76% of TZD courses; however, no patient developed serotonin syndrome. 8% of TZD courses were stopped prematurely due to AEs that included thrombocytopenia (n=3), gastrointestinal intolerance (n=2), confusion (n=1), eosinophilia (n=1) and thrombocytopenia with lactic acidosis (n=1). All cases of thrombocytopenia occurred in pts with baseline platelets $< 100,000$ cells/L. 79% of pts receiving > 14 days of TZD completed therapy successfully without AEs. Among pts who failed alternative therapies, 74% were able to tolerate TZD and completed therapy. Overall, 80% of courses were completed with a favorable outcome.