

4CPS-031 THERAPEUTIC DRUG MONITORING OF CEFTAZIDIME/ AVIBACTAM ADMINISTERED BY CONTINUOUS INFUSION: PK/PD TARGET ACHIEVEMENT AND CLINICAL OUTCOMES

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Background and Importance Ceftazidime/avibactam (CAZ/AVI) is a novel betalactam antibiotic utilised for multi-drug resistant (MDR) gram-negative bacteria. Therapeutic drug monitoring (TDM) ensures that CAZ/AVI levels achieve the pharmacokinetic/pharmacodynamic (PK/PD) target. Continuous infusion (CI) has been used to optimise CAZ/AVI pharmacodynamics.

Aim and Objectives To analyse the correlation between PK/PD target attainment of CAZ/AVI administered by CI, clinical outcomes and toxicity.

Material and Methods Patients treated with CAZ/AVI administered by CI and undergoing TDM of the CAZ plasma concentrations were included. Definitions:

CAZ/AVI PK/PD target:

- time that CAZ free concentrations remain 4 times above the minimum inhibitory concentration (MIC) of the causative pathogen ($\%fT > 4 \times \text{MIC}$).
- Overexposure: $\%fT > 10 \times \text{MIC}$.
- Clinical cure: disappearance of all signs and symptoms related to the infection and no requirement for additional antibiotic treatment initiation (except as part of de-escalation strategy) for the disease to be investigated within 48h after completion of the study drug.
- Thirty-day all-cause mortality: death from any cause during the 30 days following the end of treatment.

When real MIC was not available, a MIC of 8 mg/L was assumed.

Results Thirty-one patients (28 males, median (range) age of 64 (37-78) years) infected with extensively drug-resistant *Pseudomonas aeruginosa* and extended-spectrum betalactamase-producing *Klebsiella pneumoniae* were included (26 directed treatments and 5 empirical).

Twenty-six (83.9%) achieved the PK/PD target, 15 of which presented overexposure. Only 4 (26.6%) overexposed patients presented adverse reactions (3 increased liver enzymes and 1 thrombocytopenia).

Twenty-one (67.7%) patients achieved clinical cure, 18 (85.7%) of which achieved the PK/PD target. There was a higher frequency of patients with $\%fT > 4 \times \text{MIC}$ that achieved clinical cure (18/26 (69.2%) in patients with clinical cure vs 2/5 (40%) with clinical failure, $p = 0.686$).

The 30-day all-cause mortality was 19.4% (6 patients). A lower mortality rate was observed in patients that did achieve $\%fT > 4 \times \text{MIC}$ (14.8%) in patients who survived vs 50% in those who died, $p = 0.096$.

Conclusion and Relevance CI seems a useful strategy to reach the PK/PD target of CAZ/AVI. Few patients with overexposure presented adverse events. There seems to be a correlation between PK/PD target attainment, clinical cure and 30-day all-

cause mortality but larger studies with bigger samples are needed.

REFERENCES AND/OR ACKNOWLEDGEMENTS

Conflict of Interest No conflict of interest

4CPS-034 ANALYSIS OF THE EFFECTIVENESS OF SOTROVIMAB IN PATIENTS DIAGNOSED WITH COVID-19

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Background and Importance Sotrovimab is indicated in treatment of COVID19 in adults and adolescents who do not require supplemental oxygen and who are at increased risk of progressing to COVID-severe. The drug is administered according to prioritisation criteria published by the Spanish Agency of Medicines and Health Products (AEMPS)1.

Aim and Objectives To analyse the effectiveness of sotrovimab and to know the profile of patients.

Material and Methods Observational, retrospective and descriptive study in a tertiary level hospital. Patients who had received sotrovimab from January/2022-May/2022 were included. Variables: sex, age, mild-moderate/severe disease, vaccination-COVID, risk factors, hospitalisation/death at 29 day. Effectiveness was measured as rate of patients without progression to COVID-severe (defined as hospitalisation/death at 29 days). Variables were collected from digital medical records and in-hospital electronic prescribing.

Results Thirty-seven patients were included, mean age=61 years (21-82), 20 women (54.05%). Twenty-nine patients (78.38%) had mild-moderate COVID. 29 patients (78.38%) had received a complete vaccination regimen (3 doses), 6 patients (16.22%) two doses and 2 patients (5.41%) not vaccinated. Risk factors: 23 hypertension (62.16%), 13 diabetes (35.14%), 5 obesity (13.51%) and 4 asthma (10.81%). All patients were immunosuppressed. 17 patients (45.94%) with 2 risk factors, 9 with 3 risk factors (24.32%), 7 with 1 risk factor (18.91%) and 2 patients (5.40%) with 4 risk factors. According to the AEMPS prioritisation criteria, all belonged to the group of 'Immuno-compromised persons and high-risk conditions, regardless of vaccination status'. The high-risk conditions were: 23 patients (62.16%) had received solid organ transplantation with immunosuppressive treatment, 13 patients (35.14%) had received immunosuppressive treatment with antiCD20 in the previous 6 months (100% rituximab) and 1 patient (2.7%) was receiving active treatment with myelotoxic chemotherapy (inotuzumab) for acute lymphocytic leukaemia. 7 patients (18.9%) were hospitalised/dead at 29 days (3 exitus). All these patients had received rituximab. 30 patients (81.1%) did not progress to severe COVID. During the study period, 6 patients attended the emergency department, without admission.

Conclusion and Relevance Most patients presented good response and tolerance to treatment. This result was independent of previous treatments or risk factors. Previous treatment with anti-CD20 seems to show a tendency to progression to severe COVID. Long-term studies are needed to confirm results