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Abstract (poster session)

Emergence of carbapenem-non-susceptible extended-spectrum beta-lactamase-producing *Klebsiella pneumoniae* isolate after meropenem therapy

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Objectives: To investigate the resistance mechanisms of carbapenem susceptibility isolate recovered from one patient after treatment with meropenem. **Methods:** Since November 2010, an infection control program for ESBL-producing Enterobacteriaceae was introduced in an acute-care private hospital in Lisboa, Portugal. Seven *Klebsiella pneumoniae* isolates were recovered from one ICU inpatient. Cultures of rectal swabs and clinical samples were performed on a chromogenic medium (chromIDTM ESBL; bioMérieux). The isolates were studied by Etest MBL, PCR for blaCTX-M, blaIMP, blaVIM, blaKPC, blaOXA-48 genes and ERIC fingerprinting. Experiments were performed to detect synergy between meropenem or other antimicrobials and the efflux pump inhibitor reserpine. **Results:** A 79 years old female patient was admitted to H.SAMS with large bowel angiodysplasia leading to hemicolectomy. She had a long recovery period in ICU with prolonged ventilatory support. She was colonized with *K. pneumoniae* K137 recovered from a rectal swab prior to meropenem therapy. After twenty seven days in ICU the patient had ventilator-associated pneumonia and urinary tract infection and two *K. pneumoniae* strains K138, K139 were identified from bronchial secretions and urine, respectively. The patient was subsequently treated with meropenem, after which, during one month, three *K. pneumoniae* isolates were once again isolated from rectal swab, bronchial secretions and urine. Thirty seven days after meropenem therapy one *K. pneumoniae* K168 isolate was recovered from a rectal swab with different antibiotics susceptibility, for imipenem and meropenem MIC values of 4mL/L and 1.5 mL/L, while MIC values to previous strains were 0.25 and 0.125mL/L, respectively. From ERIC electrophoresis profiling, all strains exhibited identical banding patterns and all were ESBL producing CTX-M-15 enzyme. They gave negative results in the MBL Etest and lacked genes encoding carbapenemases. Only the *K. pneumoniae* K168 exhibited synergy between meropenem, ertapenem and reserpine, indicating that efflux pump activity could contribute to less susceptibility to carbapenem. **Conclusions:** The carbapenem phenotype observed in *K. pneumoniae* K168 was attributable to a combination of ESBL CTX-M-15 enzyme and an up-regulated efflux pump. The clonal relationship observed between the initial and subsequent *K. pneumoniae* strains may explain the emergence of resistance under meropenem selective pressure.