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Antimicrobials: Resistance surveillance

Detection of MICs of fidaxomicin and comparable antibiotics against a collection of toxin-positive *Clostridium difficile* strains in Hungary

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Objective: Fidaxomicin is a new macrocyclic antibiotic, which has a narrow spectrum of activity against gram-positive anaerobes and is bactericidal against *C. difficile*. Fidaxomicin stands out as the first-in-class oral macrocyclic antibiotic with targeted activity against *C. difficile* which has a minimal damage on the normal colonic flora. The aim of this study was to investigate the in vitro activities of fidaxomicin against 188 *Clostridium difficile* strains isolated in different centres of Hungary. The agar dilution method was used to determine the in vitro susceptibility of this new antibiotic.

Methods: Diarrheal stool samples were collected from different areas of Hungary during the routine diagnostic procedure. Faecal samples were plated on CCFA for isolation of *C. difficile*. The determination of MICs of metronidazole, moxifloxacin, rifampicin and vancomycin has already been done previously by E-test method. The isolates were tested for susceptibility to fidaxomicin using CLSI agar dilution methods. The control strain was *C. difficile* 630 (CD 630).

Results: The *C. difficile* isolates displayed minimum inhibitory concentrations (MIC) for fidaxomicin in the range of <0.008-0.5 µg/ml, with a MIC₉₀ of 0.125 µg/ml. Only four isolates (2.1 %) had 0.5 µg/ml MICs to fidaxomicin. One of these strains was moxifloxacin resistant (MIC= 32 mg/L) and there is another strain which had elevated rifampicin resistance (MIC=1 mg/L). The detected MICs displayed an identical distribution with respect to the EUCAST database for wild-type strains. The MICs of fidaxomicin for the control *C. difficile* strain (CD 630) was 0.064 ± 1 dilution. None of the isolates were resistant for metronidazole or vancomycin.

Conclusion: In summary it can be said that fidaxomicin is a therapeutically useful new antibiotic to treat CDI. Strains with elevated MICs values has not been found yet, but required for the screening of numerous clinical isolates to get a wider picture of the current status of *Clostridium difficile* strains for this new antibiotic.