

P1852 Cefiderocol *in vitro* activity against Gram-negative clinical isolates collected in Europe: result from three SIDERO-WT surveillance studies between 2014-2017

Masakatsu Tsuji*¹, Meredith Hackel^{2,3}, Yoshinori Yamano¹, Roger Echols⁴, Christopher Longshaw⁵, Davide Manissero⁵, Dan Sahn²

¹ SHIONOGI & CO., Ltd, OSAKA, Japan, ² IHMA, Inc., Schaumburg, United States, ³ IHMA, Inc., Schaumburg, United States, ⁴ ID3C, Easton, United States, ⁵ Shionogi Ltd., London, United Kingdom

Background: The emergence and dissemination of potent resistance mechanisms among Gram-negative species underscores the need for the development of new and effective therapeutic choices to treat the infections caused by these challenging organisms. Cefiderocol (CFDC) is a novel parenteral siderophore cephalosporin with potent activity against a wide variety of Gram-negative pathogens including carbapenem-resistant strains. This study evaluated the *in vitro* activity of CFDC and comparator agents against resistant clinical isolates collected in 2014-2017 from Europe based on the information on the site of infection and geography.

Materials/methods: A total of 10,551 Enterobacteriaceae, 2,126 *Acinetobacter spp.*, 2,767 *Pseudomonas aeruginosa*, 627 *Stenotrophomonas maltophilia*, and 81 *Burkholderia cepacia* complex collected from 11 European countries in 2014 - 2017 were tested. MICs were determined for CFDC, cefepime (FEP), ceftazidime-avibactam (CZA), ceftolozane-tazobactam (C/T), ciprofloxacin (CIP), colistin (CST), and meropenem (MEM) by broth microdilution according to CLSI guidelines. As recommended by CLSI, CFDC was tested in iron-depleted cation-adjusted Mueller Hinton broth. Carbapenem non-susceptible (CarbNS) strain of all bacterial species was defined as meropenem MIC \geq 4 mg/L. Quality control testing was performed on each day of testing.

Results: CFDC exhibited potent *in vitro* activity against a total of 16,154 strains of Gram-negative bacteria with MIC₉₀ of 1 mg/L. As shown in the following table, MIC₉₀ of CFDC against Enterobacteriaceae, *P. aeruginosa*, *A. baumannii*, and *S. maltophilia* were \leq 1 mg/L. Even against CarbNS isolates, MIC₉₀ of CFDC was \leq 4 mg/L irrespective of bacterial species. The *in vitro* activity of CFDC did not vary by infection sites (RTI, BSI, SSTI, and UTI). CFDC MIC₉₀ is almost similar among EU countries (Czech Republic, France, Germany, Greece, Hungary, Italy, Spain, Sweden, Turkey, United Kingdom) except for *A. baumannii* originating from Russia.

Conclusions: The potent *in vitro* activity of CFDC was demonstrated against CarbNS isolates of Enterobacteriaceae, *A. baumannii*, and *P. aeruginosa* collected from Europe, with greater than 97% of isolates having MIC values \leq 4 mg/L. These results indicate this agent has high potential for treating infections caused by these problematic organisms.