Extended spectrum beta-lactamase-producing *Escherichia coli* from human and animal sources in Italy: a "One Health" approach.

Maria Giufre'¹, Elena Mazzolini², Fabrizio Agnoletti³, Antonella Agodi⁴, Giovanni Loris Alborali⁵, Milena Arghittu⁶, Francesco Auxilia⁶, Martina Barchitta⁴, Valentina Baldo⁵, Anna Bertoncelli⁷, Natasha Bosco², Alessandro Camporese⁸, Virginia Carfora⁹, Pierlanfranco D'Agaro¹⁰, Rita De Rosa⁸, Laura D'Este², Alessia Franco⁹, Raffaella Koncan¹⁰, Paolo Lanzafame¹¹, Claudia La Mastra⁴, Annarita Mazzariol⁷, Chiara Moschioni³, Stefania Pane¹², Alessandra Pitozzi⁵, Lorenza Putignani¹², Antonio Teri⁶, Claudia Thoma⁷, Elena Tonon³, Marina Cerquetti¹, Silvio Brusaferro^{13,14}

¹ Department of Infectious Disease, Istituto Superiore di Sanità, Roma, Italy, ² Department of Epidemiology, Istituto Zooprofilattico Sperimentale delle Venezie - Sede Centrale, Legnaro, Italy, ³ Treviso's Diagnostic Department, Istituto Zooprofilattico Sperimentale delle Venezie - Sezione di Treviso, Villorba, Italy, ⁴ Dipartimento di "Scienze Mediche, Chirurgiche e Tecnologie avanzate - G.F. Ingrassia", LaPoSS - Azienda Ospedaliera Universitaria "Policlinico -Vittorio Emanuele", Università degli Studi di Catania, Catania, Italy, 5 Istituto Zooprofilattico Sperimentale della Lombardia e dell'Emilia Romagna "Bruno Ubertini", Brescia, Italy, ⁶ Fondazione IRCCS Ca' Granda Ospedale Maggiore Policlinico di Milano, Milano, Italy, 7 Diagnostic and Public Health Department, Università degli Studi di Verona, Verona, Italy, Microbiologia e Virologia, Azienda per l'Assistenza Sanitaria N.5 "Friuli Occidentale" Presidio Ospedaliero S. Maria degli Angeli, Pordenone, Italy, 9 National Reference Laboratory for Antimicrobial Resistance, Istituto Zooprofilattico Sperimentale del Lazio e della Toscana "M. Aleandri", Rome, Italy, 10 Dipartimento di Scienze Mediche Chirurgiche e della Salute, Università degli Studi di Trieste, Trieste, Italy, 11 Microbiologia e Virologia, Azienda Provinciale per i Servizi Sanitari Provincia Autonoma di Trento Ospedale S.Chiara, Trento, Italy, 12 Unità di Microbiologia, Parassitologia, Virologia e Unità di Ricerca Metagenomica, Ospedale Pediatrico Bambino Gesù, Roma, Italy, 13 Department of Medicine - University of Udine & Clinical Risk Management and Performance Assessment Unit, Udine Healthcare and University Integrated Trust, Udine, Italy, 14Istituto Superiore di Sanità, Roma, Italy

Introduction. Escherichia coli producing extended spectrum beta-lactamase (ESBL) is a serious public health concern. Food-producing animals have been suggested as a potential source for ESBL-producing E. coli affecting humans, thus surveillance requires a One Health approach. This study aimed to compare ESBL-producing E. coli isolates from humans and food-producing animals with respect to ESBL gene, phylogenetic group and sequence type (ST). Materials and Methods. Overall, 919 ESBL-producing E. coli isolates from humans (n. 474) and food-producing animals (n. 445) were collected in six Italian Regions (2016-2017). Human clinical isolates were collected from urine (n. 375) or blood (n. 99) while indicator isolates from foodproducing animals were recovered from the intestinal content or faecal samples of bovines (n. 131) swine (n. 120) and poultry (n. 194). Phenotypically confirmed ESBL-producing E. coli isolates were screened for the presence of the blactx-m-, blashy- and blacmy-2-like genes by PCR and sequencing and classified according to phylogenetic typing and MLST genotyping. Results. CTX-M was the most frequent ESBL type in both human and animal isolates with CTX-M-15 predominant in humans (74.8%) and bovine (51.1%) but not in poultry (36.6%) and swine (31.7%). CTX-M-1 was common (58%) in swine. SHV type and and CMY-2-like were found mainly in animal isolates, especially in poultry (28.9% for CMY-2-like and 17.0% for SHV12). Human ESBL-producing E. coli isolates mostly (76.8%) belonged to phylogroup B2, while animal isolates were distributed among groups A (35.7%), B1 (26.1%) and C (12.4%). Only a few animal isolates (almost all recovered from poultry) were classified into group B2 (4.3%). Most human isolates (81.6%) belonged to the pandemic ST131 clone and frequently carried CTX-M-15 (66.3%). Among animal isolates, ST131 was rarely detected (n. 3 isolates from poultry) and never carried CTX-M-15. Other than ST131 isolates were disseminated among several different STs. Twelve STs were shared by human and animal isolates with ST10, ST410 and ST38 more frequently detected. Of note, ST410 isolates of both human and animal origin shared the same ESBL profiles including either CTX-M-15 or CTX-M-1.

Discussion and Conclusions. Different subgroups of ESBL-producing *E. coli* isolates from both human and animal source may share: i) ESBL genes but carried by different ST clones, ii) ST clones but containing distinct ESBL genes, iii) both ESBL genes and ST clones. According to our results the potential exchange of ESBL genes through plasmids or isolates from animal to humans and vice versa is feasible, underlying the need for a strict monitoring based on an "One Health" approach.

2009