



## Results of the pilot study on molecular characterisation of MDR clinical *Acinetobacter baumannii* isolates from neighbouring countries



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### ECDC

Figure 3.23. Acinetobacter spp. Percentage (%) of invasive isolates with combined resistance to fluoroquinolones, aminoglycosides and carbapenems, by country, EU/EEA countries, 2017



### Published data from Serbia

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**Genome** nnouncements<sup>30</sup> Published as *Microbiology Resource Announcements* starting 12 July 2018

<u>Genome Announc</u>. 2015 Nov-Dec; 3(6): e01390-15. Published online 2015 Dec 10. doi: <u>10.1128/genomeA.01390-15</u> PMCID: PMC4675936 PMID: 26659671

#### Whole-Genome Sequence of a European Clone II and OXA-72-Producing *Acinetobacter baumannii* Strain from Serbia

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Nicolas Fortineau,<sup>a,b,c</sup> Thierry Naas<sup>a,b,c</sup>

Serbia

First Occurrence of OXA-72-Producing Acinetobacter baumannii in

Laurent Dortet, a.b.o <sup>®</sup> Rémy A. Bonnin, <sup>b.o</sup> Sandrine Bernabeu, <sup>a,b,o</sup> Lélia Escaut, <sup>d</sup> Daniel Vittecog, <sup>d</sup> Delphine Girlich, <sup>o</sup> Dilek Imanci, <sup>e</sup>

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Genome Annound

PLoS One. 2015 Mar 30;10(3):e0122793. doi: 10.1371/journal.pone.0122793. eCollection 2015.

Carbapenem-resistant Acinetobacter baumannii from Serbia: revision of CarO classification.

Novovic K<sup>1</sup>, Mihajlovic S<sup>1</sup>, Vasiljevic Z<sup>2</sup>, Filipic B<sup>3</sup>, Begovic J<sup>1</sup>, Jovcic B<sup>4</sup>.

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Antimicrob Agents Chemother. 2018 Jun 26;62(7). pii: e00219-18. doi: 10.1128/AAC.00219-18. Print 2018 Jul.

#### OXA-72-Mediated Carbapenem Resistance in Sequence Type 1 Multidrug (Colistin)-Resistant Acinetobacter baumannii Associated with Urinary Tract Infection in a Dog from Serbia.

Misic D<sup>1</sup>, Asanin J<sup>2</sup>, Spergser J<sup>3</sup>, Szostak M<sup>#3</sup>, Loncaric I<sup>#3</sup>.

### Published data from Bosnia and Herzegovina 2011-2015

Medicinski Glasnik, Volumen 8, Number 2, August 2011

#### LETTER TO THE EDITOR

#### Multidrug-resistant Acinetobacter baumannii- the pathogen with no borders?

#### Ivana Goić-Barišić<sup>1</sup>, Vanja Kaliterna<sup>2</sup>

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**ORIGINAL ARTICLE** 

Emergence of extensive drug-resistant (XDR) *Acinetobacter baumannii*in the Clinical Center University of Sarajevo, Bosnia and Herzegovina

Amela Dedeić-Ljubović<sup>1</sup>, Đana Granov<sup>1</sup>, Mirsada Hukić<sup>2,3</sup>

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368

Letters to the Editor / Journal of H

Outbreak in Croatia caused by a new carbapenem-resistant clone of *Acinetobacter baumannii* producing OXA-72 carbapenemase

### Published data from Bosnia and Herzegovina 2017-2018

Turk J Med Sci. 2017 Apr 18;47(2):715-720. doi: 10.3906/sag-1507-180.

Molecular characteristics and antibiotic resistance of Acinetobacter baumanniibeta-lactamaseproducing isolates, a predominance of intrinsic blaOXA-51, and detection of TEM and CTX-M genes.

Ibrahimagić A<sup>1</sup>, Kamberović F<sup>2</sup>, Uzunović S<sup>1</sup>, Bedenić B<sup>3,4</sup>, Idrizović E<sup>1</sup>.

Infect Genet Evol. 2018 Mar;58:192-198. doi: 10.1016/j.meegid.2017.12.021. Epub 2017 Dec 22.

### Arrival of carbapenem-hydrolyzing-oxacillinases in Acinetobacter baumannii in Bosnia and Herzegovina.

Petrović T<sup>1</sup>, Uzunović S<sup>2</sup>, Barišić I<sup>3</sup>, Luxner J<sup>4</sup>, Grisold A<sup>5</sup>, Zarfel G<sup>6</sup>, Ibrahimagić A<sup>7</sup>, Jakovac S<sup>8</sup>, Slaćanac D<sup>9</sup>, Bedenić B<sup>10</sup>.

Carbapenem resistance of *A. baumannii* in Croatia for the period 2005. - 2008.

### 2002 – 2009 A. baumannii

• IMI R < 10%

•OXA 107 •European clone 1





Croatian Committee for Antibiotic Resistance Surveillance

# Resistance to carbapenems in Croatia 2009-2017



### Croatian Committee for Antibiotic Resistance Surveillance

### Published data from Croatia

#### BRIEF COMMUNICATION

First Report of Molecular Characterization of Carbapenem-Resistant Acinetobacter baumannii in Different Intensive Care Units in University Hospital Split, Croatia

I. GOIC-BARISIC<sup>1</sup> - B. BEDENIC<sup>2</sup> M. TONKIC<sup>1</sup> - S. KATIC<sup>3</sup> - S. KALENIC<sup>3</sup> V. PUNDA-POLIC<sup>1</sup>



Occurrence of OXA-107 and ISAba1 in Carbapenem-Resistant Isolates of Acinetobacter baumannii from Croatia

Ivana Goic-Barisic, Branka Bedenic, Marija Tonkic, Anita Novak, Stjepan Katic, Smilja Kalenic, Volga Punda-Polic, Kevin J. Towner

Outbreak in Croatia caused by a new carbapenem-resistant clone of *Acinetobacter baumannii* producing OXA-72 carbapenemase

I. Goic-Barisic<sup>a</sup>,\* 🗹 🖂, K.J. Towner<sup>b</sup>, A. Kovacic<sup>c</sup>, K. Sisko-Kraljevic<sup>c</sup>, M. Tonkic<sup>a</sup>, A. Novak<sup>a</sup>, V. Punda-Polic<sup>a</sup>

European Journal of Clinical Microbiology & Infectious Diseases

Carbapenem resistance and acquired class D betalactamases in *Acinetobacter baumannii* from Croatia 2009–2010

Authors

Authors and affiliations

M. Vranić-Ladavac, B. Bedenić 🖂 , F. Minandri, M. Ištok, Z. Bošnjak, S. Frančula-Zaninović, R. Ladavac, P. Visca 🖂

### CROCMID 2016 – CROCMID 2019

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Figure 3.19. Acinetobacter spp. Distribution of isolates: fully susceptible and resistant to one, two and three antimicrobial groups (among isolates tested against fluoroquinolone, aminoglycoside and carbapenems), EU/EEA countries, 2016





Fully Resistant to one Resistant to two Resistant to three Resistant to four susceptible antimicrobial group antimicrobial groups antimicrobial groups antimicrobial groups

Resistant to five antimicrobial groups

Figure 3.19. Acinetobacter spp. Distribution of isolates: fully susceptible and resistant to one, two and three antimicrobial groups (among isolates tested against fluoroquinolones, aminoglycosides and carbapenems), EU/EEA countries, 2017



• the aim of this pilot study is to compare the genotype resemblance and resistance mechanism of MDR clinical isolates of *A. baumannii* in region of southeastern Europe

• 12 clinical isolates of carbapenem-resistant *A. baumannii* 

- three different hospitals in neighbouring countries Croatia (UHS), Bosnia and Herzegovina (UHM), and Serbia (IPHV)
- all collected isolates shared high level of resistance to carbapenems with MIC >32mg/L to both imipenem and meropenem
- beside the carbapenem-resistance, isolates were uniformly resistant to gentamicin and ciprofloxacin, but susceptible to colistin

- four isolates originated from University Hospital of Split, Croatia, were isolated from the tracheal and bronchoalveolar aspirates of patients from adult and paediatric Intensive Care Units in different outbreaks periods from 2009-2018
- all isolates belong to IC 2

- two isolates were collected from University Hospital Mostar, Bosnia and Herzegovina in the beginning of 2018, from urine sample and wound swab
- six isolates came from different wards of Clinical Centre of Vojvodina, one hospital in universityaffiliated medical centre Novi Sad, Serbia and were collected from blood cultures during 2017 and 2018

- multiplex polymerase chain reaction (PCR) using specific primers for *bla*OXA-51-like, *bla*OXA-40-like, *bla*OXA-23-like, *bla*OXA-58-like and *bla*OXA-143-like genes was performed to investigate carbapenem resistance
- the relatedness of collected A. baumannii isolates was assessed by using pulsed-field gel electrophoresis (PFGE)

# Multiplex PCR results from 12 clinical isolates of *A. baumannii*



### Results

- the multiplex PCR confirmed the presence of *bla*OXA-40-like genes in half (6/12) of the collected isolates from neighbouring countries, besides the presence of *bla*OXA-23 gene
- all obtained amplicons of *bla*OXA genes were sequenced on both strands (commercial service Macrogen Europe, The Netherlands)

### Neighbour-joining phylogenetic tree inferred on *bla*OXA-72 gene



Brazil Taiwan Croatia Serbia



5

### Results

- identical sequences were obtained from 6 clinical isolates harboured OXA-72 oxacillinases confirming long time (more than a decade) of this mechanism of resistance to carbapenems in south-eastern Europe
- the *bla*OXA-72 gene sequence determined in this study has been marked as Cro1 and deposited in GenBank under number MN366238

### Results

- identical sequences were obtained from 6 clinical isolates harboured OXA-23 oxcillinases, as a second confirmed mechanism of carbapenem resistance in *A. baumannii*
- the relatedness of collected A. baumannii isolates was assessed by using pulsed-field gel electrophoresis (PFGE) and displayed diversity of genotyping profiles
- MLST is going on....

### ECCMID Amsterdam 2019



# Thank you

- Ana Kovačić
- Marijo Pirija
- Deana Medić
- Sanja Jakovac
- Tanja Petrović
- Marija Tonkić
- Jasna Hrenović





University of Split School of Medicine

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