

# Multidrug resistant *Acinetobacter baumannii* inside and outside hospital setting



SVEUČILIŠTE  
U SPLITU  
MEDICINSKI  
FAKULTET

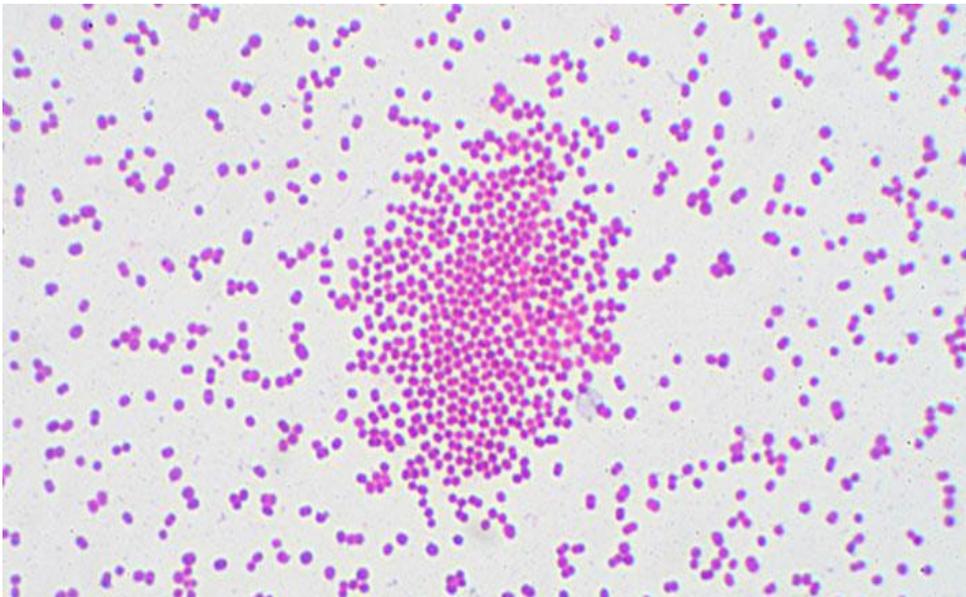
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University of Split School of Medicine



**KBC SPLIT**  
Klinički bolnički centar Split

**Multi-Drug Resistant (MDR) *A. baumannii***  
**are among the most “problematic pathogens”**  
**encountered by clinicians**

Infectious Diseases Society of America:  
*A. baumannii* is one of the “**Red Alert**” pathogens



# CDC's Report: Antibiotic Resistance Threats in the United States, 2015

## Urgent Threats

- *Clostridium difficile*
- Carbapenem-resistant Enterobacteriaceae (CRE)
- Drug-resistant *Neisseria gonorrhoeae*

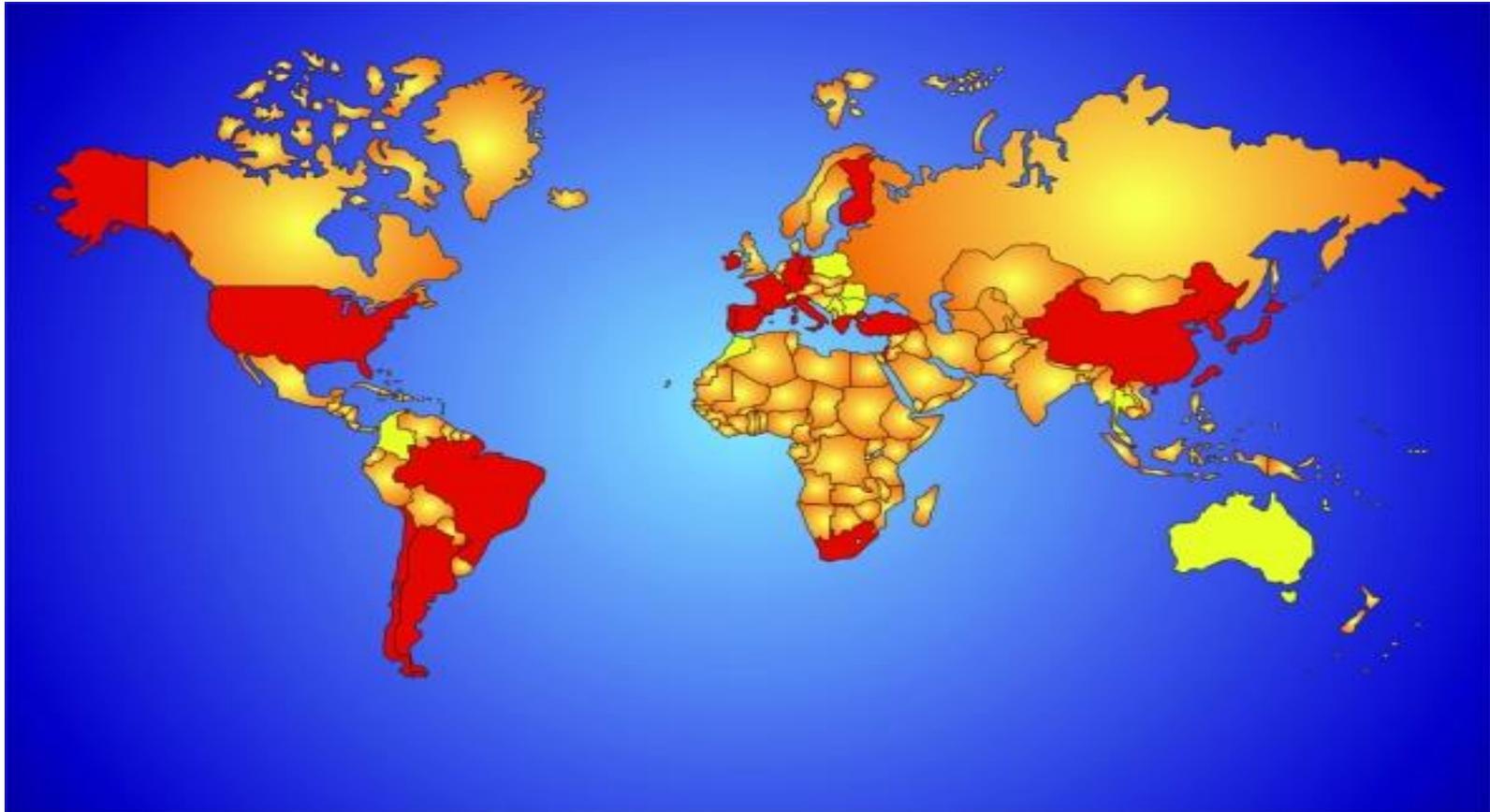
## Serious Threats

- Multidrug-resistant *Acinetobacter*
- Drug-resistant *Campylobacter*
- Fluconazole-resistant *Candida* (a fungus)
- Extended spectrum  $\beta$ -lactamase producing Enterobacteriaceae (ESBLs)
- Vancomycin-resistant *Enterococcus* (VRE)
- Multidrug-resistant *Pseudomonas aeruginosa*
- Drug-resistant Non-typhoidal *Salmonella*
- Drug-resistant *Salmonella* Typhi
- Drug-resistant *Shigella*
- Methicillin-resistant *Staphylococcus aureus* (MRSA)
- Drug-resistant *Streptococcus pneumoniae*
- Drug-resistant tuberculosis

## Concerning Threats

- Vancomycin-resistant *Staphylococcus aureus* (VRSA)
- Erythromycin-resistant Group A *Streptococcus*
- Clindamycin-resistant Group B *Streptococcus*

# *Acinetobacter baumannii*: Emergence of a Successful Pathogen



Countries that have reported an outbreak of carbapenem-resistant *Acinetobacter baumannii*. Red signifies outbreaks reported before 2006, and yellow signifies outbreaks reported since 2006.

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

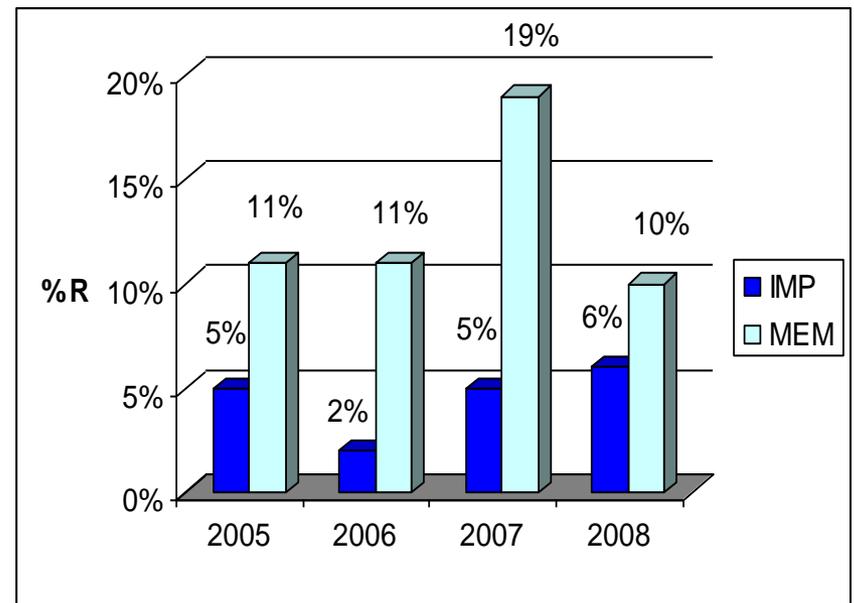
2002 – 2009

*A. baumannii*

• IMI R < 10%

• OXA 107

• European clone 1

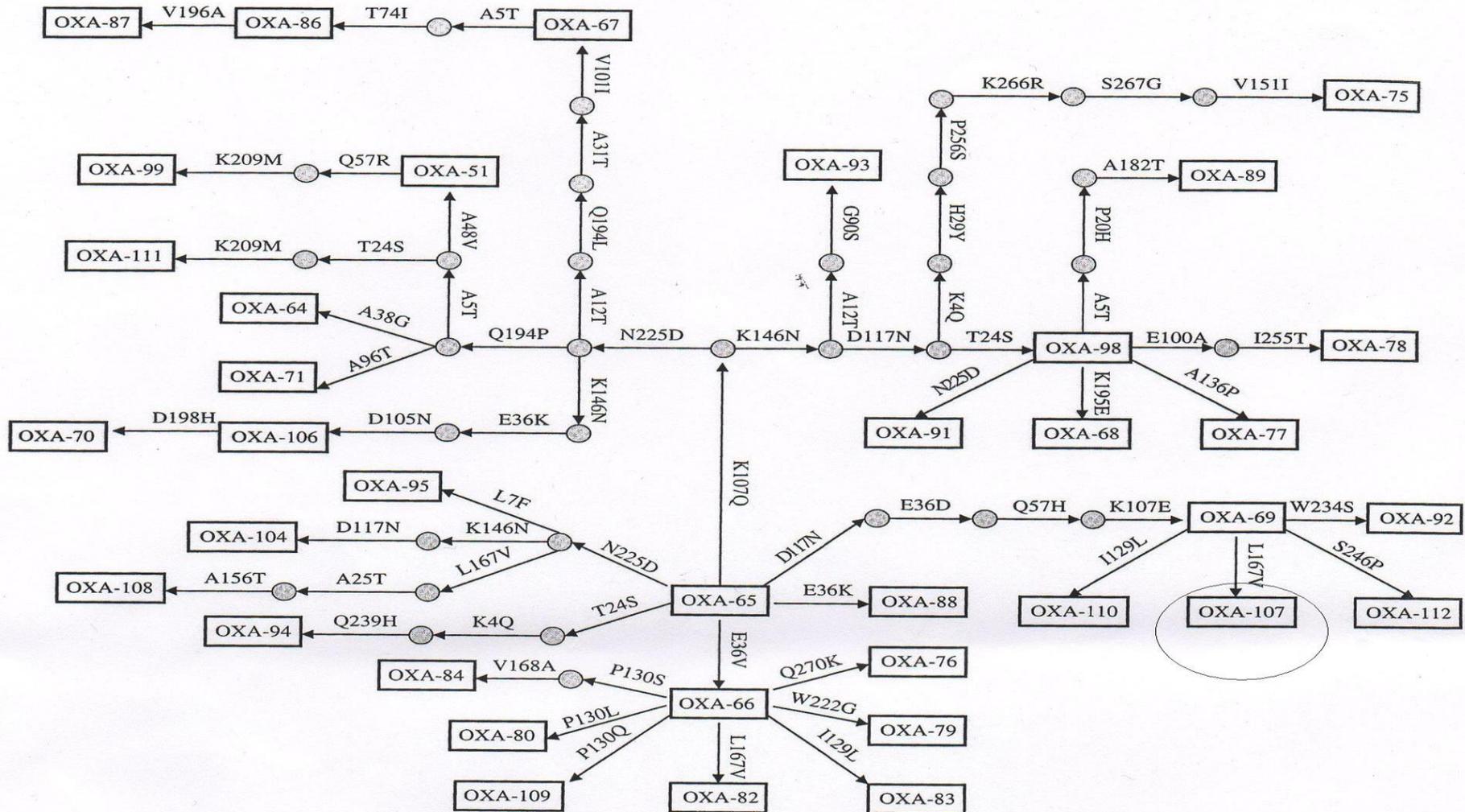


Journal of  
**Chemotherapy**

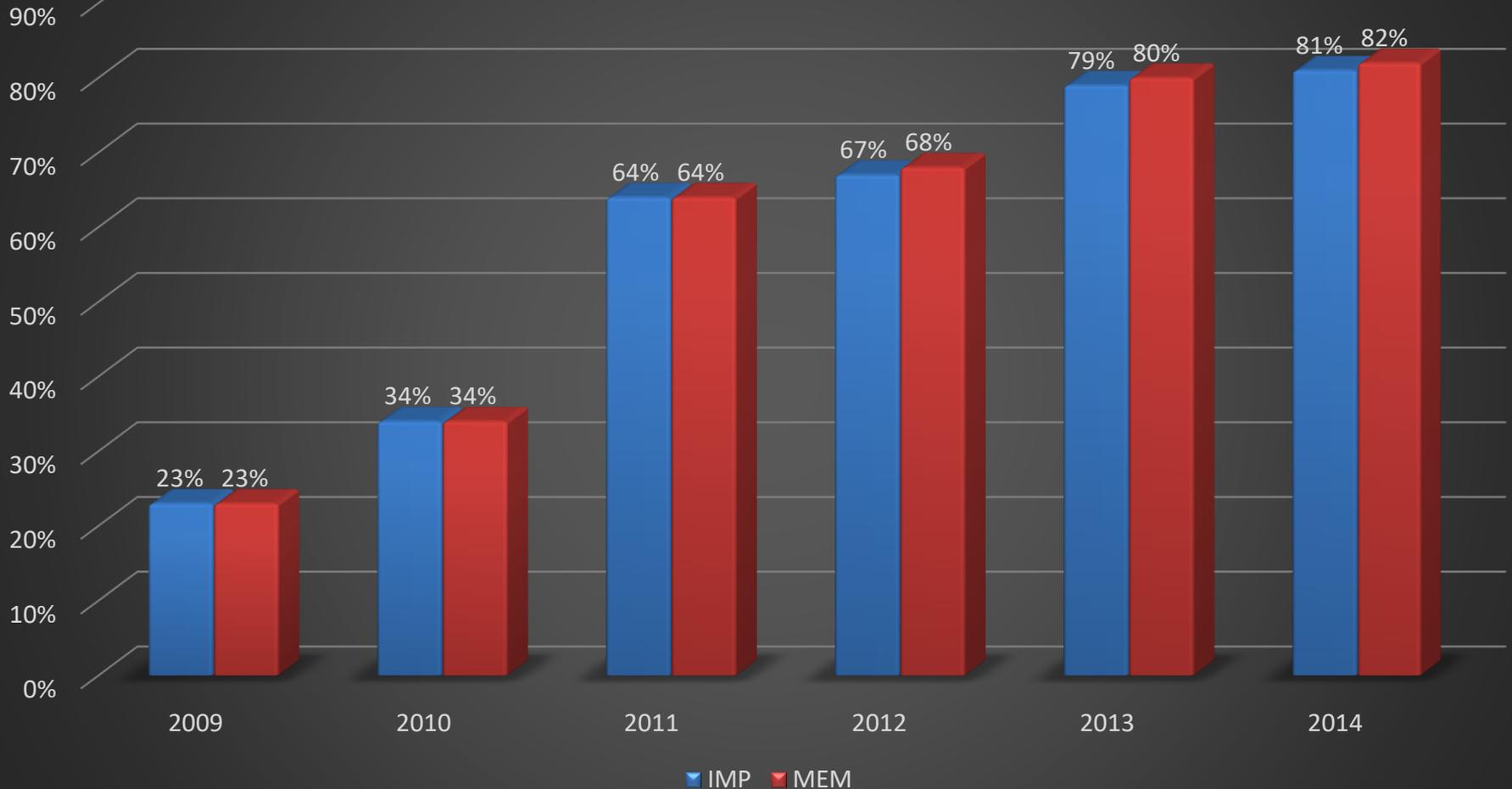


Croatian Committee for  
Antibiotic Resistance Surveillance

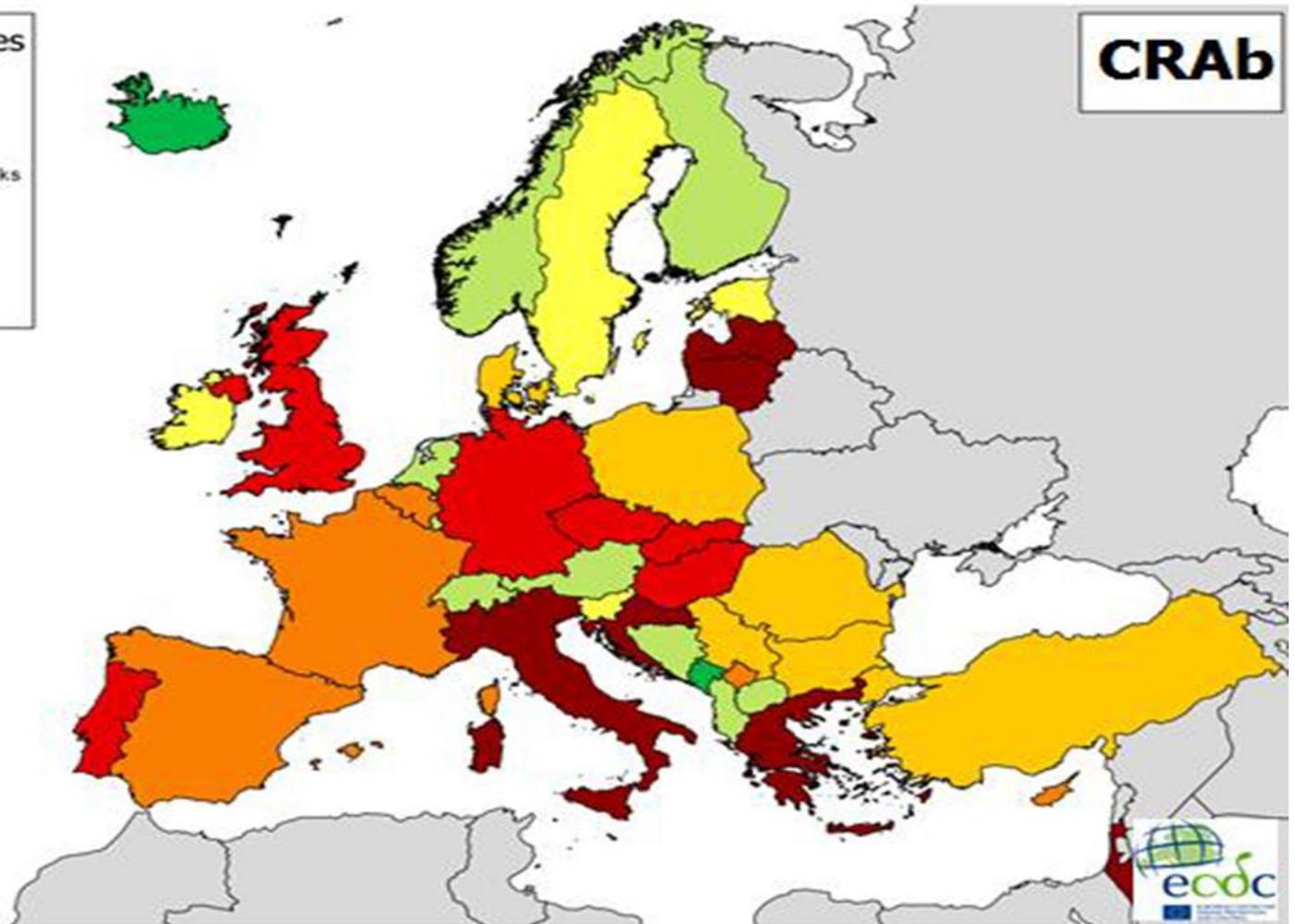
# Mechanism of resistance – hyperproduction of OXA-107 due to the IS<sub>Aba1</sub> location upstream of the gene



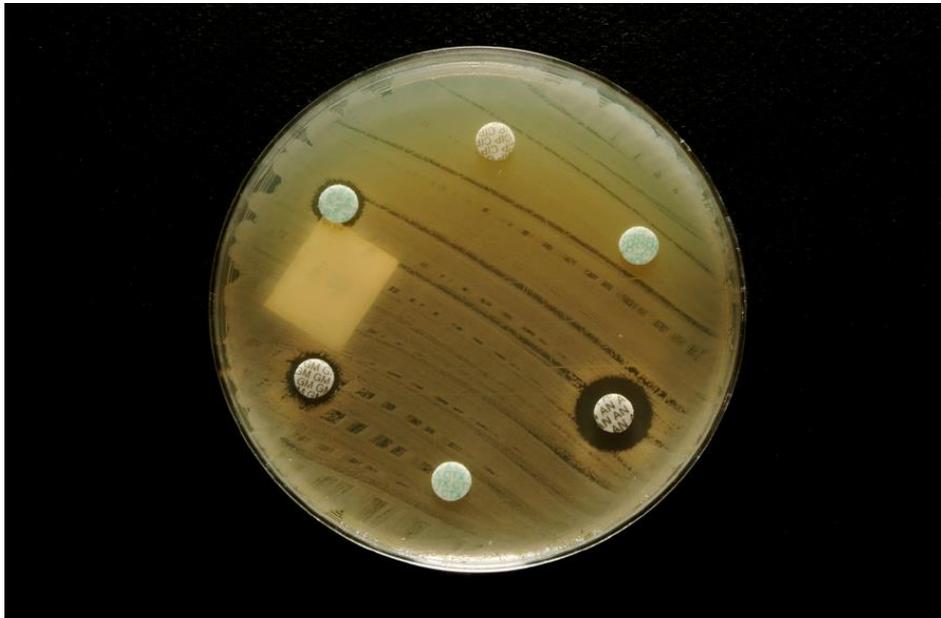
# Carbapenem resistance of *A. baumannii* in Croatia 2009-2014



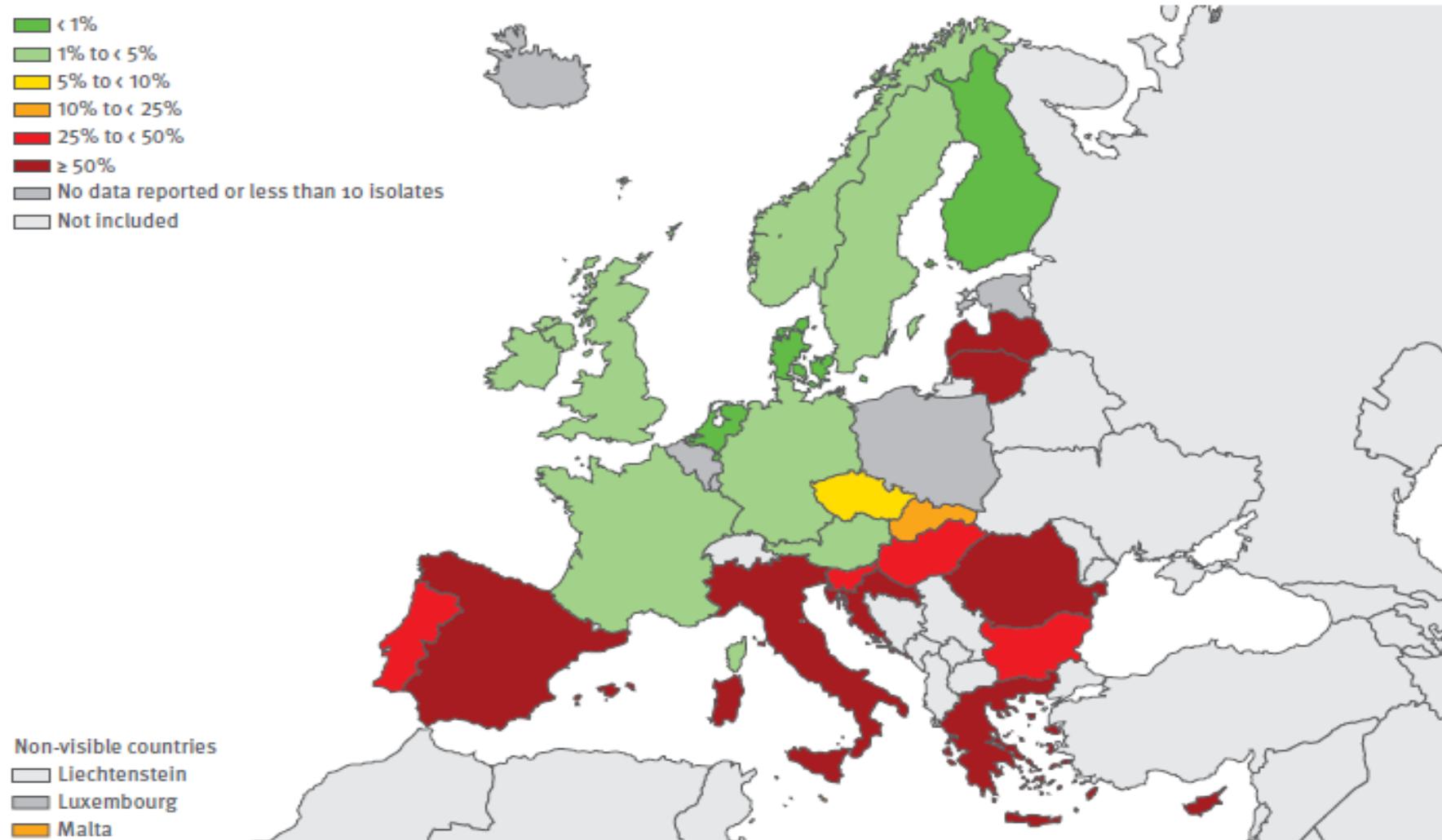
# Occurrence of carbapenem-resistant *Acinetobacter baumannii* (CRAB) 2013-14



**“Mostar clone” = European clone 2  
(GC 2) with OXA-72 carbapenemase**



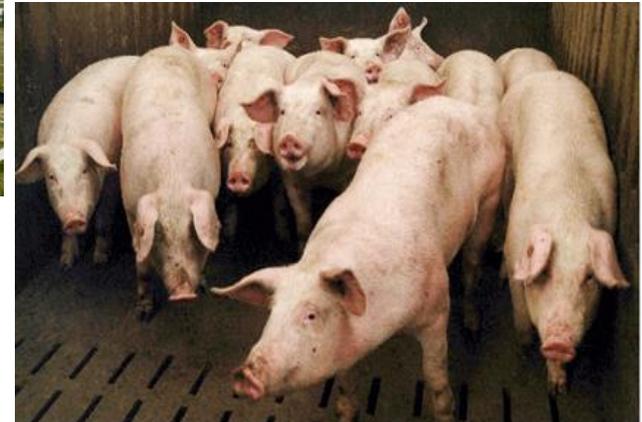
**Figure 3.20.** *Acinetobacter* spp. Percentage (%) of Invasive Isolates with combined resistance to fluoroquinolones, aminoglycosides and carbapenems, by country, EU/EEA countries, 2014



# Where does *A. baumannii* come from?



?

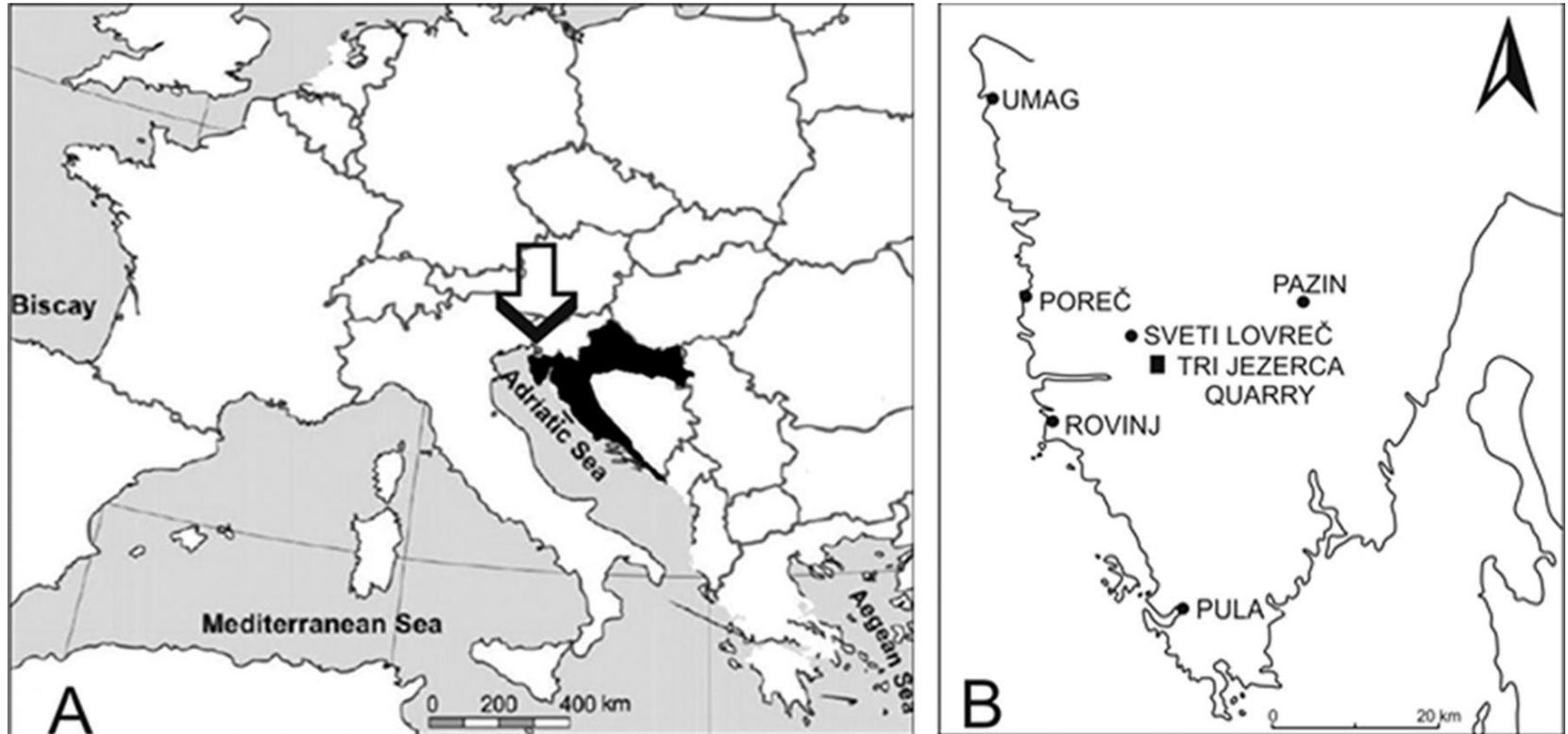


# *Acinetobacter*: an old friend but a new enemy

*A. baumannii* (and its close relatives of clinical importance) are not ubiquitous organisms. While it is certainly true that *A. baumannii* can be isolated from patients and hospital environmental sources during outbreaks, this species has no known natural habitat outside the hospital. This species can be isolated only very rarely from soil, water and other environmental samples; indeed, during non-outbreak periods it is often isolated only rarely inside hospitals.

# Environmental *Acinetobacter baumannii* Strain Similar to a Clinical Isolate in Paleosol

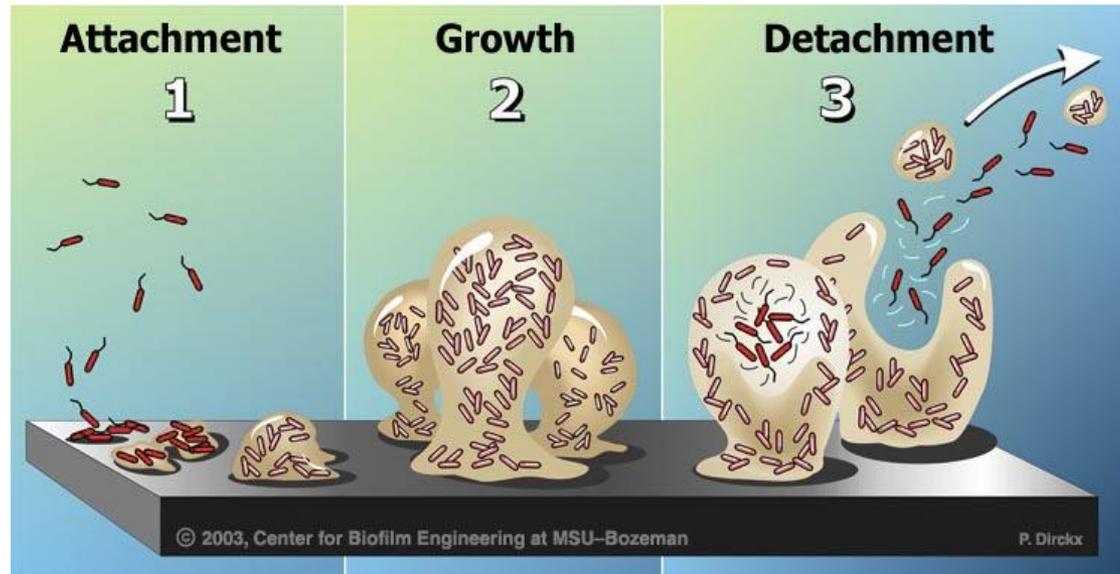
- an accidental discovery in 2013



Jasna Hrenovic et al. *Appl. Environ. Microbiol.*  
2014;80:2860-2866

Applied and Environmental Microbiology

# Multicenter investigation in Croatia



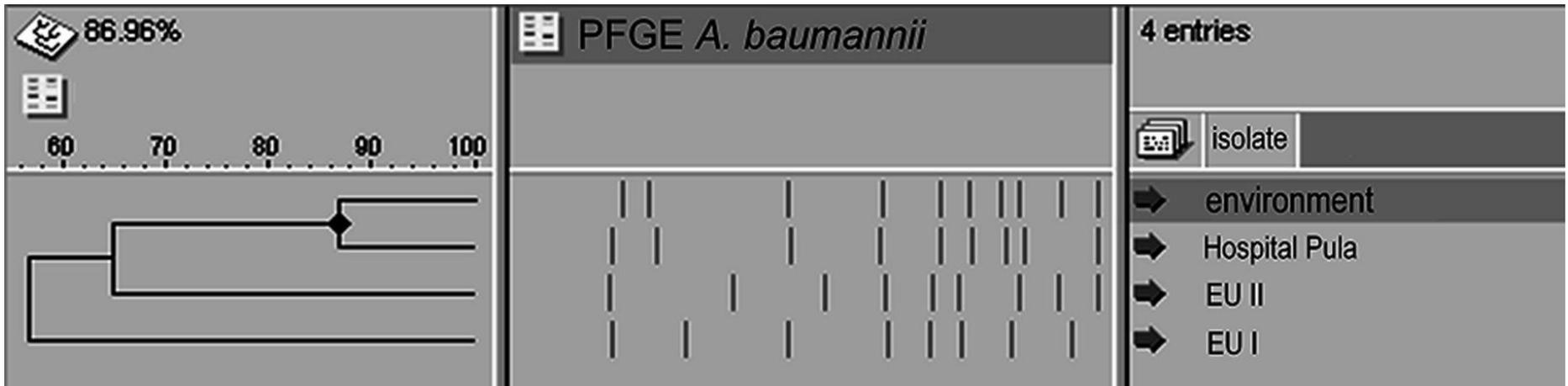
- more than 100 clinical isolates of *A. baumannii* (2009)
- focused on ability to form biofilm in correlation to genotypes (clones), origin of tested isolates and resistance to antibiotics

Kaliterna V, PhD thesis 2014

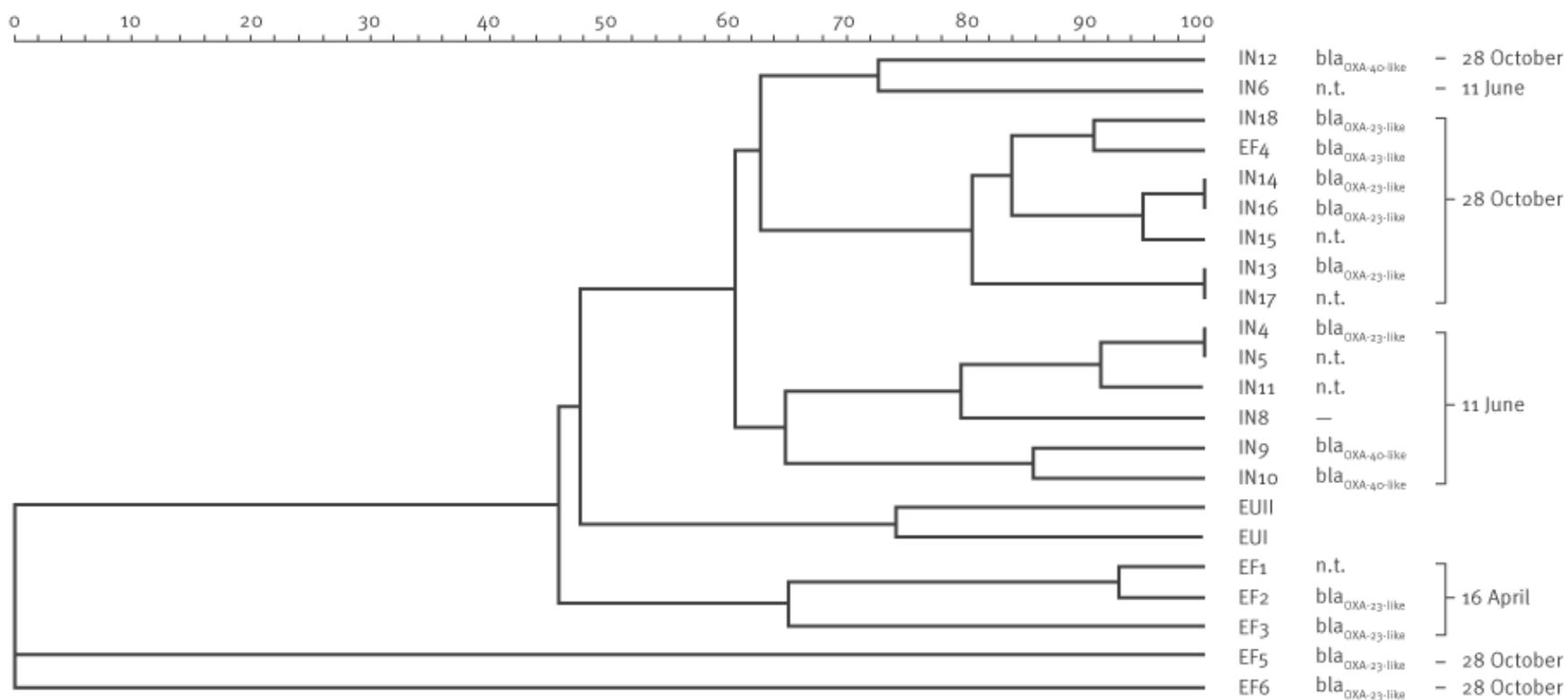
Croatian Committee for Antibiotic Resistance Surveillance



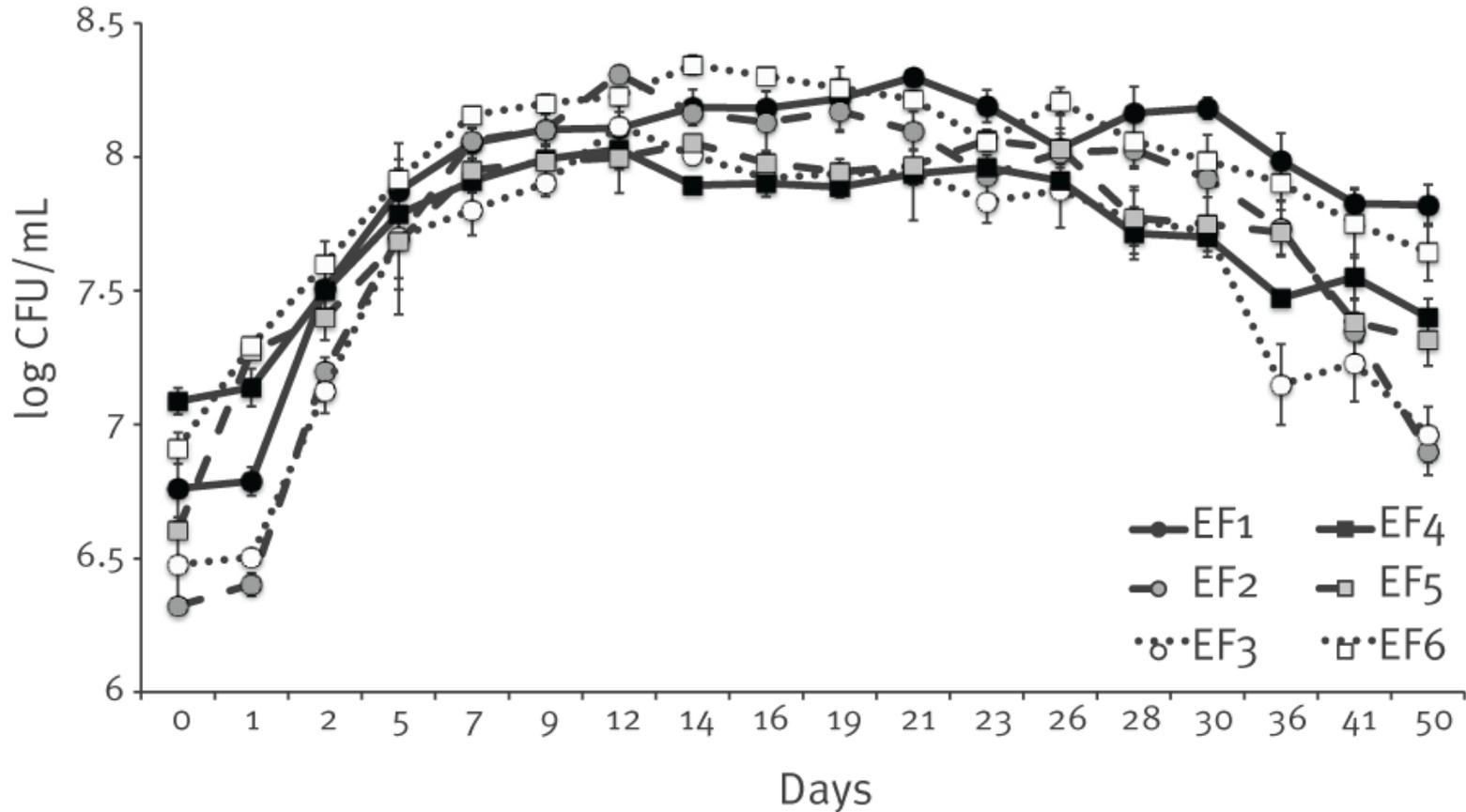
# Dendrogram based on Apal-digested DNA from different isolates of *A. baumannii*



# Carbapenem-resistant isolates of *Acinetobacter baumannii* in a municipal wastewater treatment plant, Croatia, 2014



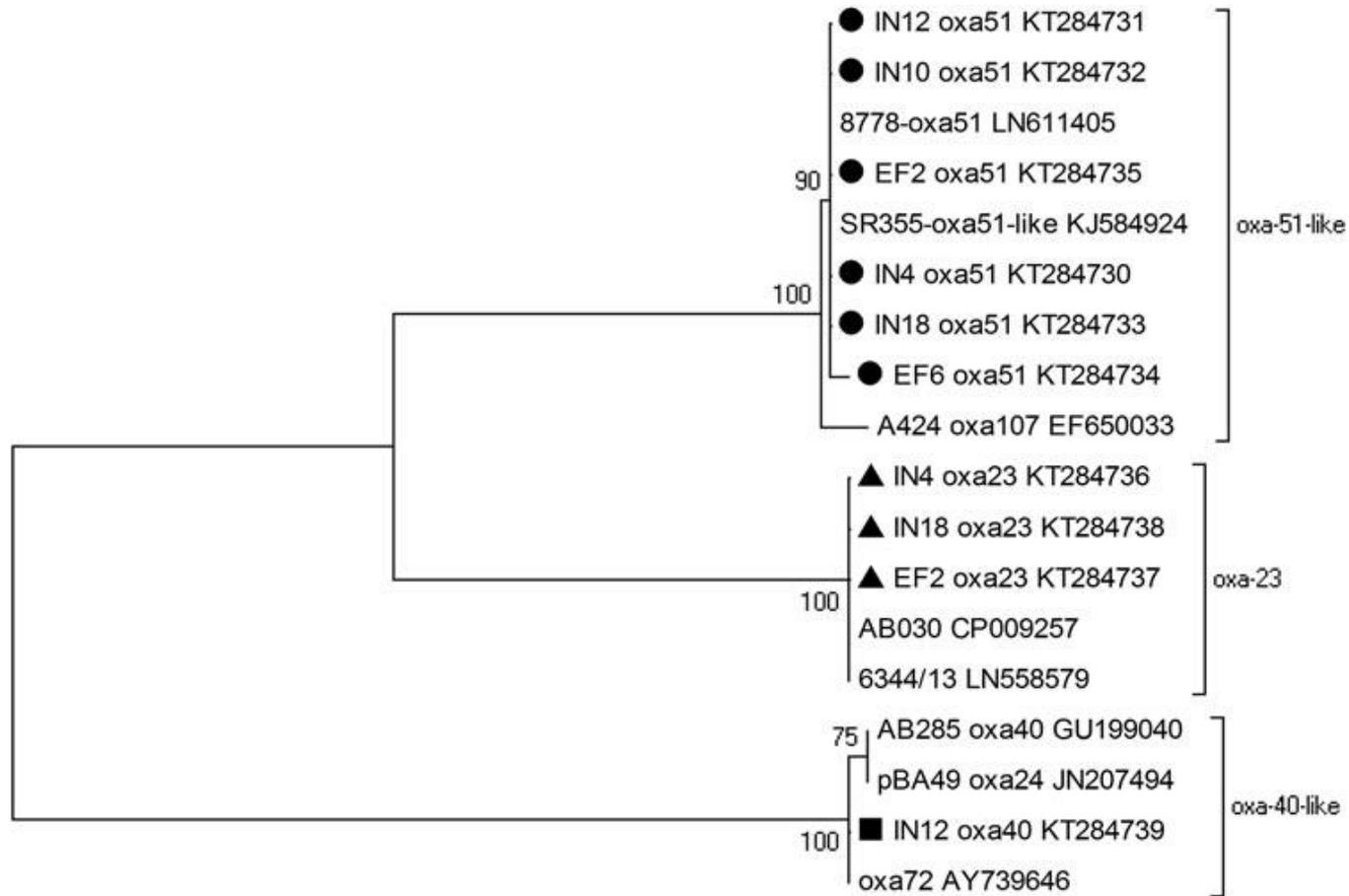
# Survival of six *Acinetobacter baumannii* isolates (EF1–6) recovered from effluent wastewater in the autoclaved effluent wastewater during 50 days, Croatia, 2014



# Point out

- all wastewater *A. baumannii* isolates recovered in this study were MDR and showed comparable levels of antibiotic resistance to clinical *A. baumannii* isolates in Croatia
- the findings of MDR *A. baumannii* after the process of chlorination suggest that conventional disinfection of effluent may not be the best strategy for mitigating the propagation of *A. baumannii* in environment
- currently the standards for discharge of treated municipal wastewater do not prescribe the elimination of MDR bacteria including (untreated) hospital wastewaters

# Phylogenetic tree on the basis of *rpoB* gene confirming molecular identification of *A. baumannii*



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# Strategies to Prevent Transmission

- Hand hygiene
- Isolation precautions
- Active surveillance for MDROs
- Decolonization of patients
- Environmental hygiene including hospital wastewater

The screenshot displays a CDC webpage with the following content:

- Threat Level:** SERIOUS (indicated by 4 yellow circles and 1 red circle).
- Statistics:**
  - 7,300 MULTIDRUG-RESISTANT ACINETOBACTER INFECTIONS
  - 500 DEATHS FROM MULTIDRUG-RESISTANT INFECTIONS
  - 12,000 ACINETOBACTER INFECTIONS PER YEAR
- Warning:** AT LEAST THREE DIFFERENT CLASSES OF ANTIBIOTICS NO LONGER CURE RESISTANT ACINETOBACTER INFECTIONS.
- Public Health Threat:** An estimated 12,000 healthcare-associated Acinetobacter infections occur in the United States each year. Nearly 7,000 (or 63%) of these are multidrug-resistant, and about 500 deaths are attributed to these infections.
- Resistance of Concern:** Some Acinetobacter strains are resistant to nearly all or all antibiotics including carbapenems, often considered antibiotics of last resort.
  - About 63% of Acinetobacter is considered multidrug-resistant, meaning at least three different classes of antibiotics no longer cure Acinetobacter infections.
  - Approximately 2% of healthcare-associated infections reported to CDC's National Healthcare Safety Network are caused by Acinetobacter, but the proportion is higher among critically ill patients on mechanical ventilators (about 7%).
- Table:**

	Percentage of all Acinetobacter healthcare-associated infections that are multidrug-resistant	Estimated number of infections	Estimated number of deaths attributed
Multidrug-resistant Acinetobacter	63%	7,300	500
- Source:** U.S. Department of Health and Human Services, Center for Disease Control and Prevention.



# The role of *ISAbal* in expression of OXA carbapenemase genes in *Acinetobacter baumannii*

Jane F. Turton<sup>1</sup>, M. Elaina Ward<sup>2</sup>, Neil Woodford<sup>2</sup>, Mary E. Kaufmann<sup>1</sup>, Rachel Pike<sup>2</sup>, David M. Livermore<sup>2</sup> & Tyrone L. Pitt<sup>1</sup>



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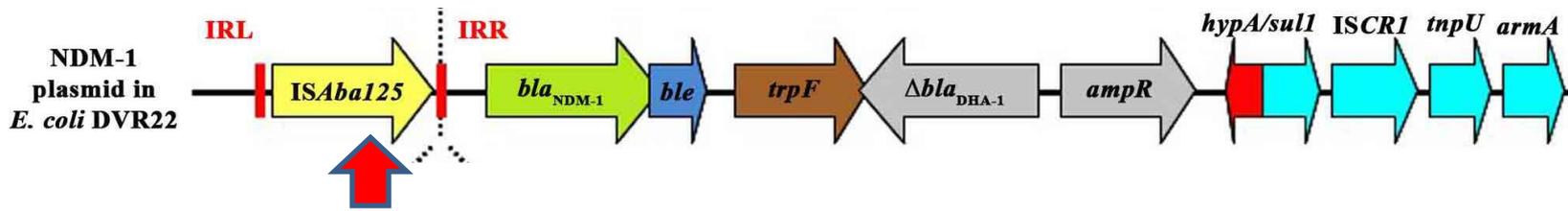
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for Microbiology

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0095-1137/09/\$08.000 doi:10.1128/JCM.02394-08  
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## Occurrence of OXA-107 and *ISAbal* in Carbapenem-Resistant Isolates of *Acinetobacter baumannii* from Croatia

Ivana Goic-Barisic,1\* Branka Bedenic,2 Marija Tonkic,1 Anita Novak,1  
Stjepan Katic,2 Smilja Kalenic,2 Volga Punda-Polic,1 and Kevin J. Towner3

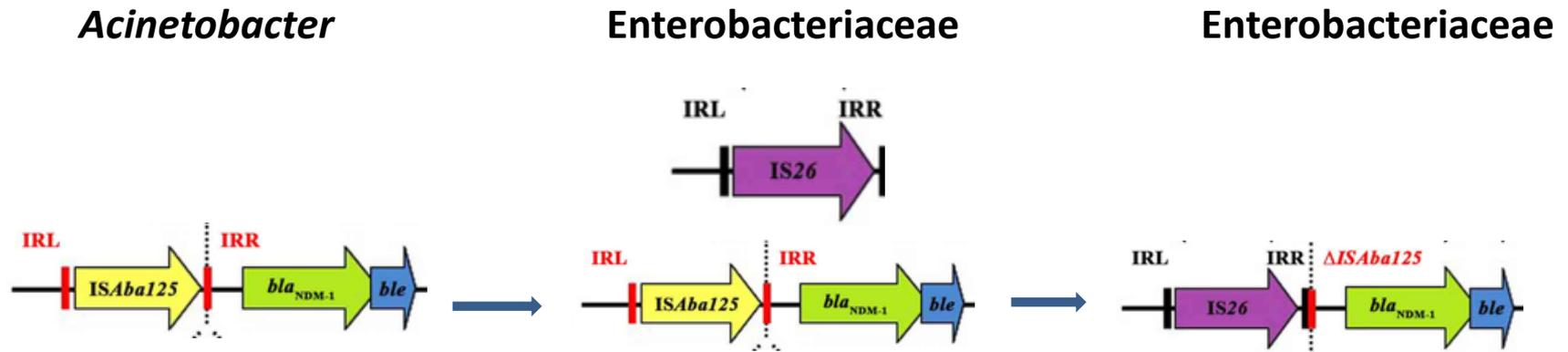
## Genetic environment surrounding *bla*<sub>NDM</sub> in *Acinetobacter baumannii*



**ISAba125 upstream of NDM-1**  
**ISAba125 is an IS specific from *Acinetobacter* spp.**

**The first and unique ISAba125-NDM in Enterobacteriaceae!!**

**ISAba125-NDM was mobilized from *Acinetobacter* to Enterobacteriaceae**



**Multidrug resistant *Acinetobacter baumannii*  
inside and outside hospital setting**



# Thank you

- Croatian science foundation – project 252556  
**Natural habitat of clinically important  
*Acinetobacter baumannii* (NATURACI)**

Jasna Hrenović

Martina Šeruga Musić

Blaženka Hunjak



Ana Kovačić

Marija Tonkić

