**TITLE:** DIVERSITY OF  $bla_{OXA}$  GENES, GR-PLASMIDS AND ISABA1 IN CARBAPENEMASE-PRODUCING ACINETOBACTER BAUMANNII FROM BRAZIL

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## **ABSTRACT**

Acinetobacter baumannii is one of the most important multi drug-resistant (MDR) pathogens causing worrisome nosocomial infections. Plasmid-mediated carbapenemases genes have been also associated to A. baumannii, mainly OXA-carbapenemases. We studied 50 carbapenem resistant A. baumannii isolated from inpatients of a tertiary hospital in Brazil. We investigated the presence of extended-spectrum beta-lactamases (ESBL) ( $bla_{\text{CTX-M-groups-1,-2,-8,-9}}$  and 25,  $bla_{\text{GES}}$ ,  $bla_{\text{BEL}}$ ,  $bla_{\text{VEB}}$ , and  $bla_{\text{PER}}$ ) carbapenemase genes ( $bla_{KPC}$ ,  $bla_{SPM}$ ,  $bla_{NDM}$ ,  $bla_{IMP}$ ,  $bla_{VIM}$ , and  $bla_{OXA}$ ). The presence of the  $bla_{OXA-51}$  was used to confirm the identification of A. baumannii (Ab) species. Zone 1 of the rpoB gene was sequenced for the isolates carrying epidemiologically important carbapenemases. All 50 Ab isolates studied presented bla<sub>OXA-51-like</sub> as expected; 37 of the 50 Ab also presented the bla<sub>OXA-23-like</sub> gene whereas 6/50 presented  $bla_{OXA-143-like}$  and 1/50 presented  $bla_{OXA-58-like}$ . 11 isolates presented only  $bla_{OXA-51-like}$ , however, most isolates carried more than one  $bla_{OXA}$  resistance gene. Besides, we investigated the presence of the insertion sequence ISAba1 in all isolates. Only 5 Ab did not present ISAba1. In all 37 isolates carrying blaOXA-23-like we found ISAba1 located upstream of bla<sub>OXA-23-like</sub>. No ESBL and other carbapenemase genes investigated were identified in any of the 50 isolates studied. For the 7 Ab carrying bla<sub>OXA-143-like</sub> and/or bla<sub>OXA-58-like</sub> we also investigated the presence of 19 AB-GR plasmid groups using AB-PBRT. More than one AB-GR was found per isolate, with exception of the Ab carrying bla<sub>OXA-58-like</sub> that presented only the GR3, being GR-3 followed by GR-8 the most frequently AB-GR detected. Interestingly, none of the isolates showed GR6 which is the most widespread GR group across Europe, China and Taiwan, and most often associated with the  $bla_{OXA-23-like}$  and  $bla_{OXA-58-like}$  genes. The presence of ISAba1, a recognized strong promoter for  $bla_{OXA}$  genes, could increase the  $bla_{OXA}$  carbapenemase activity in the isolates studied. Besides, the wide variety of AB-GR plasmids certainly contributes to dissemination of plasmid-mediated bla<sub>OXA</sub> genes and could also facilitate the spread of other resistance genes beyond A. baumannii. Thereby, plasmids surveillance and characterization in bacteria other than Enterobacteriaceae could improve knowledge and control of antimicrobial resistance.

**Keywords:** Acinetobacter baumannii, bla<sub>OXA</sub>, GR-plasmids, ISAba1

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