

Unusual Detection of an *Acinetobacter* Class D Carbapenemase Gene, *bla*_{OXA-23}, in a Clinical *Escherichia coli* Isolate

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Carbapenem resistance in *Acinetobacter baumannii* is typically mediated by the acquisition of class D β-lactamases belonging to gene clusters of *bla*_{OXA-23}-like, *bla*_{OXA-40}-like, and *bla*_{OXA-58}-like (1). In the *Enterobacteriaceae*, carbapenem-hydrolyzing class D β-lactamases are encoded by a dissimilar set of genes, the *bla*_{OXA-48}-like genes (2). The presence of *bla*_{OXA-23} in *Enterobacteriaceae* is exceptional and has been described once in *Proteus mirabilis*, where it was chromosomally encoded (3).

As a result of national surveillance efforts to characterize non-carbapenem-susceptible *Enterobacteriaceae* isolates, we detected a case of *bla*_{OXA-23} in *Escherichia coli*, which to the best of our knowledge is the first description of plasmid-borne *bla*_{OXA-23} in *E. coli*.

The isolate was recovered from the urine sample of an elderly woman. However, the isolate was not likely to be a cause of infection, as urine microscopy did not show an increased number of white blood cells. The isolate exhibited resistance to β-lactams, fluoroquinolones, and chloramphenicol. Sensitivity to tigecycline, colistin, and aminoglycosides was observed (Table 1).

Comprehensive PCR screening and sequencing for β-lactamase genes was performed (4, 5). The isolate carried *bla*_{OXA-23}, which had 100% identity with *A. baumannii bla*_{OXA-23}. It was also positive for TEM-1, OXA-1, and CMY-2 but was negative for other carbapenemases and extended spectrum β-lactamases. The isolate had a multilocus sequence type of ST471. (<http://www.pasteur.fr/recherche/genopole/PF8/mlst/EColi.html>).

Two large plasmids of approximately 50 and 100 kb were detected (6), with Southern blot hybridization localizing *bla*_{OXA-23} to the 50-kb plasmid.

Solid-medium conjugation assays were performed to assess the transferability of *bla*_{OXA-23} from the clinical isolate to the azide-resistant recipient *E. coli* J53 and to a spontaneous rifampin-resistant mutant of *A. baumannii*, ATCC 17978. However, we did not manage to isolate any transconjugants, suggesting that *bla*_{OXA-23} was harbored on a non-self-conjugative plasmid.

The immediate nucleotide sequences flanking *bla*_{OXA-23} were determined by inverse PCR and the primer walking approach (7). *E. coli bla*_{OXA-23} is flanked by two copies of insertion sequence IS1 (Fig. 1). In contrast, in *A. baumannii*, *bla*_{OXA-23} is found as part of a transposon (Tn2006, Tn2007, Tn2008) associated with insertion sequence ISAb1 or ISAb4 (8, 9) (Fig. 1).

One hundred eighty-three bases upstream, the *E. coli bla*_{OXA-23} start codon was completely identical to ISAb1 (GenBank accession number GQ861438.1). Within this region, −35 and −10 ISAb1-associated promoters were identified and likely to be

TABLE 1 MICs of antibiotics for the *Escherichia coli* isolate harboring *bla*_{OXA-23}

| Antibiotic | MIC (mg/liter) |
|-------------------------|----------------|
| Aztreonam | 32 |
| Piperacillin-tazobactam | >256 |
| Cefoxitin | 128 |
| Cefotaxime | 32 |
| Ceftazidime | 16 |
| Imipenem | 3 |
| Meropenem | 4 |
| Ertapenem | 6 |
| Tigecycline | 0.125 |
| Tetracycline | 86 |
| Levofloxacin | >32 |
| Ciprofloxacin | >32 |
| Amikacin | 0.125 |
| Gentamicin | 2 |
| Chloramphenicol | 16 |
| Colistin | 0.125 |

functional (10) (Fig. 1). The 764-bp stretch after the *bla*_{OXA-23} stop codon comprised a partially truncated putative AAA ATPase identical to that in *A. baumannii* Tn2006, Tn2007, and Tn2008 (Fig. 1).

We hypothesized that *E. coli bla*_{OXA-23} may be carried within a transposon, not unlike those identified in *A. baumannii* but interrupted by IS1. However, attempts to PCR map the region using both ISAb1 and ISAb4 as reference sequences did not yield amplicons, suggesting that it was in a genetic configuration different from that found in *A. baumannii*.

In summary, we describe the novel detection of an *E. coli* isolate carrying an *Acinetobacter bla*_{OXA-23} gene associated with *Enterobacteriaceae* IS1 on a 50-kb non-self-conjugative plasmid. The discovery highlights the potential for the tremendous spread of carbapenemases.

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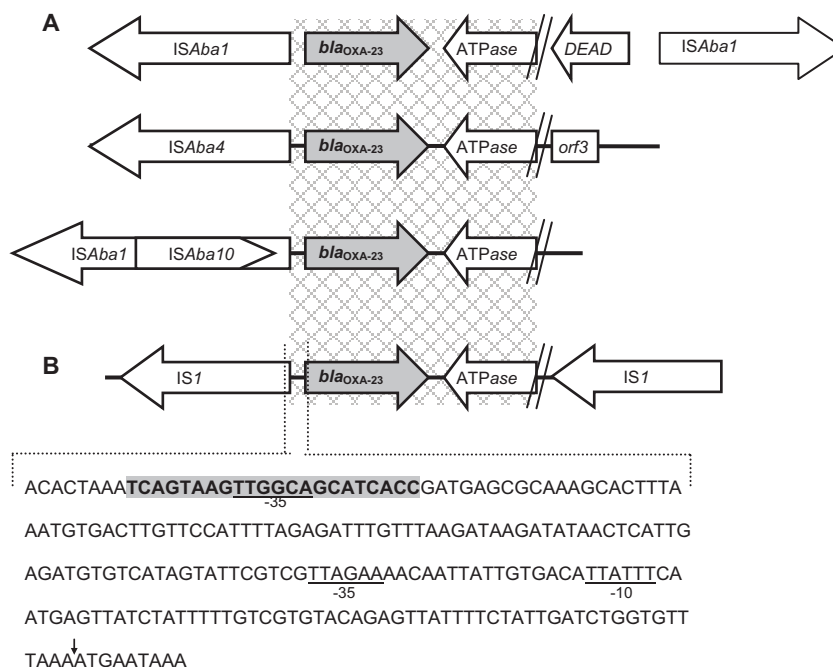


FIG 1 Schematic representation of the genetic organization of *bla*_{OXA-23}. (A) In *Acinetobacter baumannii*, ISAbal insertion sequence elements are typically found immediately upstream of *bla*_{OXA-23} (8, 9). (B) In the clinical *Escherichia coli* isolate, *bla*_{OXA-23} is flanked by two copies of *Enterobacteriaceae* insertion sequence IS1. The terminal left inverted repeat (IRL) of IS1 is indicated by the boldface nucleotides with gray shading. The putative -35 promoter of the IRL is underlined and may form hybrid promoters with existing -10 promoters, enabling transcription (11). The ISAbal-associated -35 and -10 promoters are underlined (10). The ATG start codon of OXA-23 is indicated by the vertical arrow (\downarrow). Gray checked regions represent 100% identity between *E. coli* and *A. baumannii*. ATPase, gene encoding putative AAA ATPase; DEAD, gene encoding the putative DEAD (Asp-Glu-Ala-Asp) helicase.

Nucleotide sequence accession number. The *bla*_{OXA-23} sequence from the clinical *E. coli* isolate has been deposited into GenBank under accession number [KJ716226](https://www.ncbi.nlm.nih.gov/nuclseq/KJ716226).

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We have no conflicts of interest to declare.

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