Detection of hidden antibiotic resistance through real-time genomics

Supplementary Information

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Supplementray Table 1. Real-time genomic sequencing and assembly metrics before and after treatment with CAZ-AVI. Genomic material of each bacterial isolate was sequenced for 15 hours; the pre-treatment isolate was additionally sequenced for a further 8 hours to simulate adaptive sequencing in the clinical setting.

Genomic sequencing and assembly metrics	Pre-treatment sample		Post-treatment sample
	15h	+8h	15h
Median read quality (Phred)	12.3	13.0	12.3
Median read length [kb]	4.1	5.3	4.5
Total number of bases [Mb]	442	1,551	194
Assembly coverage (median)	68	271	30
Number of contigs (>50 kb)	5	5	5
Contig N50 [Mb]	5.2	5.2	5.3

Supplementary Table 2. MIC measurements with VITEK2 and Liofilchem[®]MIC test strip for CAZ-AVI. The colouring scheme indicates the interpretation of MICs according to EUCAST guidelines (green: susceptible/intermediate; red: resistant).

Antibiotic	Pre-treatment isolate MIC (mg/L)	Post-treatment isolate MIC (mg/L)
Ampicillin	>32	>32
Ampicillin-Sulbactam	>32	>32
Piperacillin	>128	>128
Piperacillin-Tazobactam	>128	>128
Cefuroxime	>64	>64
Cefotaxime	>64	>64
Ceftriaxone	>64	>64
Ceftazidime	>64	>64
Imipenem	>16	1
Meropenem	>16	2
Ertapenem	>8	>8
Gentamicin	>16	>16
Ciprofloxacin	>4	>4
Ceftazidime-Avibactam	2	>256



Supplementary Figure 1. Bioinformatic workflow for species identification and antibiotic resistance prediction from nanopore sequencing data of pre- and post-treatment isolates (HAC: High-accuracy basecalling; ID: Identification; cgMLST: core genome Multi-Locus Sequence Typing). All computational analyses were conducted on a portable laptop with an 8 GB NVIDIA GeForce RTX 4070 GPU, 16 GB 5200 MHz RAM, and an Intel i7-13800H CPU with 14 cores and 20 threads.