

**Table S3.** High- and moderate-impact SNP mutations found in CF28A-L, CF63A-L and CF63A-S isolates showing changes in antibiotic susceptibility over time.

Isolate	Patient 3 visit	Antibiotic resistance <sup>a</sup>	Gene altered	Mutation found
CF28A-L	V2	None	NWMN_0456 <i>ksgA</i> ; dimethyladenosine transferase	Frameshift removing the initiation codon
CF63A-S	V3	TOB (R), ERY (R)	NWMN_0456 <i>ksgA</i> ;	Frameshift removing
CF63A-L	V3	TOB (R), GEN (I), ERY (R), CIP (R)	dimethyladenosine transferase	the initiation codon
			NWMN_2096 <i>lacD</i> ; tagatose 1,6-diphosphate aldolase	730C>T; Pro244Ser

<sup>a</sup> TOB, tobramycin; GEN, gentamicin; ERY, erythromycin; CIP, ciprofloxacin; R, resistant; I, intermediate. See precise minimal inhibitory concentrations in Table S1.