

S2 Table: Genotypes of bioreactor-adapted TGC resistant (BTR) clonal strains isolated from the end of adaptation

| Run | Strain ^a | MIC _{TGC} (µg/mL) | Deletions in 5'UTR of <i>tetM</i> | New Tn916 Insertion Sites ^b | Clones ^c |
|-----|---------------------|----------------------------|-------------------------------------|--|---------------------|
| N/A | S613 | 0.125 | None | None | N/A |
| 1 | BTR87a | 2.0 | Δ87-bp, 51-bp upstream <i>tetM</i> | <i>yycI</i> | 2 |
| 1 | BTR87b | 2.0 | Δ87-bp, 51-bp upstream <i>tetM</i> | Intergenic site J | 1 |
| 1 | BTR22 | 2.0 | Δ22-bp, 231-bp upstream <i>tetM</i> | None | 3 |
| 1 | BTR37 | 1.0 | Δ37-bp, 98-bp upstream <i>tetM</i> | <i>yycI</i> , Sulfatase, Intergenic sites C, D, H, and I | 2 |
| 1 | BTR0 | 0.5 | None | None | 1 |
| 2 | BTR125a | 2.0 | Δ125-bp, 36-bp upstream <i>tetM</i> | O-antigen ligase ^d | 1 |
| 2 | BTR125b | 2.0 | Δ125-bp, 36-bp upstream <i>tetM</i> | None | 3 |
| 2 | BTR0 | 0.5 | None | None | 1 |

^aAll BTR strains have the S10^{R53Q-Δ54-58KTHK} allele.

^bAll strains retained the ancestral insertion of Tn916. Strains with multiple copies of Tn916 retained the wildtype sequence at the ancestral site.

^cNumber of clones sequenced with the corresponding genotype.

^dThis copy of Tn916 has an insertion of the IS204 transposon downstream of *tetM*.