Table S1 | MIC values (µg/ml) of *P. aeruginosa* PA14 and 12 clinical isolates after 3 and 9 days of ALE assays under sub-MIC concentrations of ciprofloxacin.

| STRAINS          | CONCENTRATION (relative to MIC) | REPLICATE | 0d  | 3d  | 9d  |
|------------------|---------------------------------|-----------|-----|-----|-----|
| Without antibiotic |                                | LX1       | 0.047 | 0.064 | 0.064 |
|                  |                                 | LX2       | 0.047 | 0.064 | 0.064 |
|                  |                                 | LX3       | 0.047 | 0.064 | 0.064 |
|                  |                                 | LX4       | 0.047 | 0.064 | 0.064 |
| PA14             | 1/200                           | X1        | 0.047 | 0.064 | 0.064 |
|                  |                                 | X2        | 0.047 | 0.094 | 0.064 |
|                  |                                 | X3        | 0.047 | 0.064 | 0.064 |
|                  |                                 | X4        | 0.047 | 0.094 | 0.094 |
|                  | 1/100                           | X5        | 0.047 | 0.064/0.1 | 0.125 |
|                  |                                 | X6        | 0.047 | 0.064 | 0.125 |
|                  |                                 | X7        | 0.047 | 0.064/0.5 | 0.094 |
|                  |                                 | X8        | 0.047 | 0.047/0.38 | 0.094 |
|                  |                                 | X9        | 0.047 | 0.75 | 0.75 |
|                  |                                 | X10       | 0.047 | 1 | 0.75 |
|                  |                                 | X11       | 0.047 | 0.75 | 0.75 |
|                  |                                 | X12       | 0.047 | 1 | 0.75 |
|                  | 1/25                            | X13       | 0.047 | 1.5 | 1 |
|                  |                                 | X14       | 0.047 | 1.5 | 1 |
|                  |                                 | X15       | 0.047 | 1 | 0.75 |
|                  |                                 | X16       | 0.047 | 1 | 1.5 |
|                  | 1/10                            | X17       | 0.047 | 1.5 | 1.5 |
|                  |                                 | X18       | 0.047 | 2 | 1.5 |
|                  |                                 | X19       | 0.047 | 1.5 | 1 |
|                  |                                 | X20       | 0.047 | 1 | 1.5 |
|                  | 1/5                             | X21       | 0.047 | 1 | 16 |
|                  |                                 | X22       | 0.047 | 1.5 | 3 |
|                  |                                 | X23       | 0.047 | 2 | 3 |
|                  |                                 | X24       | 0.047 | 2 | 4 |
|                  |                                 | X25       | 0.047 | 1.5 | ≥32 |
|                  |                                 | X26       | 0.047 | 1.5 | 6 |
|                  |                                 | X27       | 0.047 | 2 | 12 |
|                  |                                 | X28       | 0.047 | 1.5 | 6 |
| Without antibiotic |                                | LA1       | 0.094 | 0.125 | 0.125 |
|                  |                                 | LA2       | 0.094 | 0.125 | 0.125 |
|                  |                                 | LA3       | 0.094 | 0.125 | 0.125 |
|                  |                                 | LA4       | 0.094 | 0.125 | 0.125 |
|    | A1  | A2  | A3  | A4  | A5  | A6  | A7  | A8  | A9  | A10 | A11 | A12 | A13 | A14 | A15 | A16 | A17 | A18 | A19 | A20 | A21 | A22 | A23 | A24 | A25 | A26 | A27 | A28 |
|----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| 1/200 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 |
| 1/100 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 |
| 1/50 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 |
| 1/25 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 |
| 1/10 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 |
| 1/5 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 |
| 1/2 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,125 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 | 0,094 |

**Without antibiotic**

|    | LB1 | LB2 | LB3 | LB4 | LB5 | LB6 | LB7 | LB8 | LB9 | LB10 |
|----|-----|-----|-----|-----|-----|-----|-----|-----|-----|------|
| 1/200 | 0,064 | 0,064 | 0,064 | 0,047 | 0,064 | 0,064 | 0,064 | 0,064 | 0,064 | 0,064 |
| 1/100 | 0,064 | 0,064 | 0,064 | 0,047 | 0,064 | 0,064 | 0,064 | 0,064 | 0,064 | 0,064 |
| 1/50 | 0,064 | 0,064 | 0,064 | 0,047 | 0,064 | 0,064 | 0,064 | 0,064 | 0,064 | 0,064 |

**AND04-004A**
|   | B11 | B12 | B13 | B14 | B15 | B16 | B17 | B18 | B19 | B20 | B21 | B22 | B23 | B24 | B25 | B26 | B27 | B28 |
|---|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| 1/25 | 0.064 | 0.094 | 0.047/0.38 | 0.064 | 0.094 | 0.047/0.38 | 0.064 | 0.094 | 0.047/0.38 | 0.064 | 0.094 | 0.047/0.38 | 0.064 | 0.094 | 0.047/0.38 | 0.064 | 0.094 | 0.047/0.38 |
| 1/10 | 0.064 | 0.75 | 0.75 | 0.064 | 0.75 | 0.75 | 0.064 | 0.75 | 0.75 | 0.064 | 0.75 | 0.75 | 0.064 | 0.75 | 0.75 | 0.064 | 0.75 | 0.75 |
| 1/5 | 0.064 | 0.75 | 0.75 | 0.064 | 0.75 | 0.75 | 0.064 | 0.75 | 0.75 | 0.064 | 0.75 | 0.75 | 0.064 | 0.75 | 0.75 | 0.064 | 0.75 | 0.75 |
| 1/2 | 0.064 | 1.5 | 4 | 0.064 | 1 | 3 | 0.064 | 0.5/3 | 4 | 0.064 | 1 | 1.5 |

| Without antibiotic | LC1 | LC2 | LC3 | LC4 |
|---------------------|-----|-----|-----|-----|
| 1/200               | 0.19 | 0.19 | 0.25 | 0.19 |
| 1/100               | 0.19 | 0.19 | 0.25 | 0.19 |
| 1/50                | 0.19 | 0.19 | 0.25 | 0.19 |
| 1/25                | 0.19 | 0.19 | 0.25 | 0.19 |
| 1/10               | 0.19 | 0.19 | 0.25 | 0.19 |

| FQSE06-0403 | C1 | C2 | C3 |
|-------------|-----|-----|-----|
| 1/200       | 0.19 | 0.25 | 0.19 |
| 1/100       | 0.19 | 0.25 | 0.19 |
| 1/50        | 0.19 | 0.25 | 0.19 |
| 1/25        | 0.19 | 0.19 | 0.25 |
| 1/10        | 0.19 | 0.19 | 0.25/1 |

| C4 | C5 | C6 | C7 | C8 | C9 | C10 | C11 | C12 | C13 | C14 | C15 | C16 | C17 | C18 | C19 | C20 |
|----|----|----|----|----|----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| 0.19 | 0.19 | 0.25 | 0.19 | 0.19 | 0.19 | 0.19 | 0.19 | 0.19 | 0.19 | 0.19 | 0.19 | 0.094 | 0.19 | 0.19 | 0.25/0.75 |
|  | C21 | 0.19 | 1 | 1.5 |
|---|-----|------|---|-----|
|  | C22 | 0.19 | 1.5 | 2 |
|  | C23 | 0.19 | 1 | 1 |
|  | C24 | 0.19 | 1 | 1.5 |
|  | C25 | 0.19 | 1.5 | 1.5 |
|  | C26 | 0.19 | 0.75 | 1 |
|  | C27 | 0.19 | 0.75 | 1 |
|  | C28 | 0.19 | 0.5 | 1.5 |
|  | C29 | 0.19 | 1 | 1.5 |
|  | C30 | 0.19 | 1.5 | 1.5 |
|  | C31 | 0.19 | 0.75 | 1 |
|  | C32 | 0.19 | 0.75 | 1.5 |
|  | C33 | 0.19 | 0.5 | 1.5 |
|  | C34 | 0.19 | 1 | 1.5 |
|  | C35 | 0.19 | 1.5 | 1.5 |
|  | C36 | 0.19 | 0.75 | 1 |
|  | C37 | 0.19 | 0.75 | 1.5 |
|  | C38 | 0.19 | 0.5 | 1.5 |

**Without antibiotic**

|  | LD1 | 0.094 | 0.094 | 0.094 |
|---|-----|-------|-------|-------|
|  | LD2 | 0.094 | 0.094 | 0.094 |
|  | LD3 | 0.094 | 0.094 | 0.094 |
|  | LD4 | 0.094 | 0.094 | 0.064 |
|  | D1  | 0.094 | 0.094 | 0.064 |
|  | D2  | 0.094 | 0.094 | 0.094 |
|  | D3  | 0.094 | 0.094 | 0.064 |
|  | D4  | 0.094 | 0.064 | 0.094 |
|  | D5  | 0.094 | 0.094 | 0.064/0.19 |
|  | D6  | 0.094 | 0.064 | 0.064/0.19 |
|  | D7  | 0.094 | 0.064 | 0.064/0.25 |
|  | D8  | 0.094 | 0.125 | 0.125 |
|  | D9  | 0.094 | 0.5 | 0.75 |
|  | D10 | 0.094 | 0.5 | 0.75 |
|  | D11 | 0.094 | 0.5 | 0.75 |
|  | D12 | 0.094 | 0.094/0.38 | 0.5 |
|  | D13 | 0.094 | 0.75 | 0.75 |
|  | D14 | 0.094 | 1 | 0.75 |
|  | D15 | 0.094 | 1 | 0.5 |
|  | D16 | 0.094 | 1 | 0.75 |
|  | D17 | 0.094 | 1 | 1 |
|  | D18 | 0.094 | 1 | 1 |
|  | D19 | 0.094 | 0.75 | 0.75 |
|  | D20 | 0.094 | 1 | 0.75 |
|  | D21 | 0.094 | 0.75 | 1.5 |
|  | D22 | 0.094 | 1.5 | 1 |
|  | D23 | 0.094 | 1/6 | 4 |
|  | D24 | 0.094 | 2 | 2 |
|  | D25 | 0.094 | 6 | 6 |
|  | D26 | 0.094 | 1.5 | 8 |
|  | D27 | 0.094 | 1 | 3 |
|  | D28 | 0.094 | 3 | 2 |
|  | LE1 | 0.25 | 0.25 | 0.19 |
|  | LE2 | 0.25 | 0.25 | 0.19 |

**BAL04-002**
|    | E1   | E2   | E3   | E4   | E5   | E6   | E7   | E8   |
|----|------|------|------|------|------|------|------|------|
| 1/200 | 0.25 | 0.25 | 0.25 | 0.25 | 0.25 | 0.25 | 0.25 | 0.25 |
| 1/100 |      |      |      |      |      |      |      |      |
| 1/50  |      |      |      |      |      |      |      |      |
| 1/25  |      |      |      |      |      |      |      |      |
| 1/10  |      |      |      |      |      |      |      |      |
| 1/5   |      |      |      |      |      |      |      |      |
| 1/2   |      |      |      |      |      |      |      |      |

**Without antibiotic**

|    | LF1  | LF2  | LF3  | LF4  |
|----|------|------|------|------|
| 1/200 | 0.5  | 0.75 | 0.5  | 0.5  |
| 1/100 |      |      |      |      |
| 1/50  |      |      |      |      |
| 1/25  |      |      |      |      |

**CAT02-004**

|    | LE3  | LE4  |
|----|------|------|
| 1/200 | 0.25 | 0.25 |
| 1/100 |      |      |
| 1/50  |      |      |
| 1/25  |      |      |
| 1/10  |      |      |
| 1/5   |      |      |
| 1/2   |      |      |

|    | F1   | F2   | F3   | F4   | F5   | F6   | F7   | F8   |
|----|------|------|------|------|------|------|------|------|
| 1/200 | 0.5  | 0.5  | 0.5  | 0.5  | 0.5  | 0.5  | 0.5  | 0.5  |
| 1/100 |      |      |      |      |      |      |      |      |
|       | 1/50      | 1/100     | 1/250     |
|-------|-----------|-----------|-----------|
|       | F9        | F10       | F11       |
|       | 0,5       | 0,5       | 0,5       |
|       | 0,5       | 0,5       | 0,75      |
|       | 0,5       | 0,5       | 0,75      |
|       | 0,5       | 0,75      | 0,75      |
|       | 0,5       | 0,75      | 0,75      |
|       | 0,5       | 0,75      | 0,75      |
|       | 0,75      | 0,75      | 0,75      |
|       | 0,19      | 0,25      | 0,25      |
|       | 0,19      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,75      | 0,75      | 0,75      |
|       | 0,75      | 0,75      | 0,75      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,38      | 0,38      | 0,38      |

Without antibiotic:

|       | LG1       | LG2       | LG3       |
|-------|-----------|-----------|-----------|
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,19      | 0,38      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |

|       | G1        | G2        | G3        |
|-------|-----------|-----------|-----------|
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |

|       | G5        | G6        | G7        |
|-------|-----------|-----------|-----------|
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |

|       | G8        | G9        | G10       |
|-------|-----------|-----------|-----------|
|       | 0,25      | 0,25      | 0,25      |
|       | 0,38      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,19      | 0,25      | 0,25      |

|       | G11       | G12       | G13       |
|-------|-----------|-----------|-----------|
|       | 0,25      | 0,38      | 0,25      |
|       | 0,19      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,19      | 0,25      | 0,25      |

|       | G14       | G15       | G16       |
|-------|-----------|-----------|-----------|
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,38      | 0,25      |
|       | 0,25      | 0,19      | 0,25      |
|       | 0,25      | 0,25      | 0,25      |
|       | 0,25      | 0,19      | 0,25      |

|       | G17       | G18       | G18       |
|-------|-----------|-----------|-----------|
|       | 0,25      | 0,38      | 0,19      |
|       | 0,25      | 0,38      | 0,25      |
| Dilution | FQSE110603 | Without antibiotic |
|----------|------------|-------------------|
| 1/10     |            |                   |
|          | G19 0.25   | LH1 0.125 0.094   |
|          | G20 0.25   | LH2 0.125 0.094   |
|          | G21 0.25   | LH3 0.125 0.125   |
|          | G22 0.25   | LH4 0.125 0.125   |
|          | G23 0.25   |                   |
|          | G24 0.25   |                   |
|          | G25 0.25   |                   |
|          | G26 0.25   |                   |
|          | G27 0.25   |                   |
|          | G28 0.25   |                   |
| 1/5      |            |                   |
|          | H1 0.125   | H1 0.125 0.125   |
|          | H2 0.125   | H2 0.125 0.19    |
|          | H3 0.125   | H3 0.125 0.19    |
|          | H4 0.125   | H4 0.125 0.125   |
| 1/2      |            |                   |
|          | H5 0.125   | H5 0.125 0.75    |
|          | H6 0.125   | H6 0.125 0.25    |
|          | H7 0.125   | H7 0.125 1.1     |
|          | H8 0.125   | H8 0.125 0.75    |
|          | H9 0.125   | H9 0.125 0.5     |
|          | H10 0.125  | H10 0.125 0.75   |
|          | H11 0.125  | H11 0.125 0.75   |
|          | H12 0.125  | H12 0.125 0.75   |
|          | H13 0.125  | H13 0.125 0.75   |
|          | H14 0.125  | H14 0.125 1.1    |
|          | H15 0.125  | H15 0.125 1.1    |
|          | H16 0.125  | H16 0.125 1.5    |
| 1/25     |            |                   |
|          | H17 0.125  | H17 0.125 2.1    |
|          | H18 0.125  | H18 0.125 1.5    |
|          | H19 0.125  | H19 0.125 1.5    |
|          | H20 0.125  | H20 0.125 1.5    |
| 1/10     |            |                   |
|          | H21 0.125  | H21 0.125 1.5    |
|          | H22 0.125  | H22 0.125 1.5    |
|          | H23 0.125  | H23 0.125 1.5    |
|          | H24 0.125  | H24 0.125 1.5    |
|          | H25 0.125  | H25 0.125 1.5    |
|          | H26 0.125  | H26 0.125 1.5    |
|          | H27 0.125  | H27 0.125 1.5    |
|          | H28 0.125  | H28 0.125 1.5    |
| ICA01-004                                      | LJ1   | 0.064 | 0.064 | 0.064 |
|                                               | LJ2   | 0.064 | 0.064 | 0.094 |
|                                               | LJ3   | 0.064 | 0.064 | 0.094 |
|                                               | LJ4   | 0.064 | 0.064 | 0.064 |
| Without antibiotic                            | I1    | 0.064 | 0.064 | 0.064 |
|                                               | I2    | 0.064 | 0.064 | 0.094 |
|                                               | I3    | 0.064 | 0.064 | 0.064 |
|                                               | I4    | 0.064 | 0.064 | 0.064 |
| 1/200                                          | I5    | 0.064 | 0.064 | 0.064 |
|                                               | I6    | 0.064 | 0.064 | 0.064 |
|                                               | I7    | 0.064 | 0.064 | 0.064 |
|                                               | I8    | 0.064 | 0.064 | 0.064 |
|                                               | I9    | 0.064 | 0.064 | 0.094 |
|                                               | I10   | 0.064 | 0.064 | 0.094 |
|                                               | I11   | 0.064 | 0.064 | 0.064 |
|                                               | I12   | 0.064 | 0.064 | 0.094 |
| 1/100                                          | I13   | 0.064 | 0.5   | 0.38  |
|                                               | I14   | 0.064 | 0.38  | 0.38  |
|                                               | I15   | 0.064 | 0.38  | 0.38  |
|                                               | I16   | 0.064 | 0.38  | 0.38  |
|                                               | I17   | 0.064 | 0.5   | 0.25  |
|                                               | I18   | 0.064 | 0.38  | 0.38  |
|                                               | I19   | 0.064 | 0.5   | 0.38  |
|                                               | I20   | 0.064 | 0.75  | 0.75  |
|                                               | I21   | 0.064 | 1     | 0.5   |
|                                               | I22   | 0.064 | 0.75  | 0.75  |
|                                               | I23   | 0.064 | 1     | 1.5   |
|                                               | I24   | 0.064 | 1     | 0.5   |
|                                               | I25   | 0.064 | 1/6   | 8     |
|                                               | I26   | 0.064 | 2     | 4     |
|                                               | I27   | 0.064 | 1     | 12    |
|                                               | I28   | 0.064 | 2     | 8     |
| 1/5                                            | L1    | 2     | 3     | 2     |
|                                               | L2    | 2     | 2     | 2     |
|                                               | L3    | 2     | 3     | 2     |
|                                               | L4    | 2     | 2     | 3     |
| Without antibiotic                            | J1    | 2     | 2     | 2     |
|                                               | J2    | 2     | 2     | 2     |
|                                               | J3    | 2     | 2     | 2     |
|                                               | J4    | 2     | 3     | 2     |
| 1/200                                          | J5    | 2     | 2     | 3     |
|                                               | J6    | 2     | 2     | 3     |
|   | J7 | J8 | J9 | J10 | J11 | J12 | J13 | J14 | J15 | J16 | J17 | J18 | J19 | J20 | J21 | J22 | J23 | J24 | J25 | J26 | J27 | J28 |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| 1/100 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| 1/50 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| 1/25 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| 1/10 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| 1/5 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| 1/2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 |
| Without antibiotic | | | | | | | | | | | | | | | | | | | | | | | |
| LK1 | 0,094 | 0,125 | 0,094 | | | | | | | | | | | | | | | | | | | |
| LK2 | 0,094 | 0,125 | 0,064 | | | | | | | | | | | | | | | | | | | |
| LK3 | 0,094 | 0,094 | 0,094 | | | | | | | | | | | | | | | | | | | |
| LK4 | 0,094 | 0,094 | 0,094 | | | | | | | | | | | | | | | | | | | |
| 1/200 | | | | | | | | | | | | | | | | | | | | | | | |
| K1 | 0,094 | 0,094 | 0,094 | | | | | | | | | | | | | | | | | | | |
| K2 | 0,094 | 0,125 | 0,064 | | | | | | | | | | | | | | | | | | | |
| K3 | 0,094 | 0,125 | 0,094 | | | | | | | | | | | | | | | | | | | |
| K4 | 0,094 | 0,094 | 0,094 | | | | | | | | | | | | | | | | | | | |
| 1/100 | | | | | | | | | | | | | | | | | | | | | | | |
| K5 | 0,094 | 0,125 | 0,094 | | | | | | | | | | | | | | | | | | | |
| K6 | 0,094 | 0,125 | 0,094 | | | | | | | | | | | | | | | | | | | |
| K7 | 0,094 | 0,125 | 0,125 | | | | | | | | | | | | | | | | | | | |
| K8 | 0,094 | 0,094 | 0,094 | | | | | | | | | | | | | | | | | | | |
| CLE03-004 | | | | | | | | | | | | | | | | | | | | | | | |
| 1/50 | | | | | | | | | | | | | | | | | | | | | | | |
| K9 | 0,094 | 0,094 | 0,25 | | | | | | | | | | | | | | | | | | | |
| K10 | 0,094 | 0,125 | 0,19 | | | | | | | | | | | | | | | | | | | |
| K11 | 0,094 | 0,125 | 0,25 | | | | | | | | | | | | | | | | | | | |
| K12 | 0,094 | 0,094 | 0,19 | | | | | | | | | | | | | | | | | | | |
| K13 | 0,094 | 0,75 | 0,094/0,38 | | | | | | | | | | | | | | | | | | | |
| K14 | 0,094 | 0,75 | 0,75 | | | | | | | | | | | | | | | | | | | |
| K15 | 0,094 | 0,75 | 0,5 | | | | | | | | | | | | | | | | | | | |
| K16 | 0,094 | 0,75 | 0,38 | | | | | | | | | | | | | | | | | | | |
|    |    |    |    |
|----|----|----|----|
|    | K17 | 0.094 | 1  | 0.75 |
|    | K18 | 0.094 | 1  | 1    |
|    | K19 | 0.094 | 1  | 1    |
|    | K20 | 0.094 | 1  | 1    |
| 1/10 | K21 | 0.094 | 1.5 | 2 |
|    | K22 | 0.094 | 1.5 | 1 |
|    | K23 | 0.094 | 1  | 1.5  |
|    | K24 | 0.094 | 1.5 | 1 |
| 1/5 | K25 | 0.094 | 1.5 | 2 |
|    | K26 | 0.094 | 1.5 | 4 |
|    | K27 | 0.094 | 1.5 | 3 |
|    | K28 | 0.094 | 2  | 4    |
| 1/2 |  |  |  |  |
|    |  |  |  |  |
| 1/10 |  |  |  |  |
| 1/5 |  |  |  |  |
| 1/2 |

|    |    |
|----|----|
| LL1 | 0.25 |
| LL2 | 0.25 |
| LL3 | 0.25 |
| LL4 | 0.25 |

| FQSE111010 |
|-----------|
| 1/200     |
| L1        | 0.25 |
| L2        | 0.25 |
| L3        | 0.25 |
| L4        | 0.25 |

| 1/100     |
| L5        | 0.25 |
| L6        | 0.25 |
| L7        | 0.25 |
| L8        | 0.25 |

| 1/50      |
| L9        | 0.25 |
| L10       | 0.25 |
| L11       | 0.25 |
| L12       | 0.25 |

| 1/25      |
| L13       | 0.25 |
| L14       | 0.25 |
| L15       | 0.25 |
| L16       | 0.25 |

| 1/10      |
| L17       | 0.25 |
| L18       | 0.25 |
| L19       | 0.25 |
| L20       | 0.25 |

| 1/5       |
| L21       | 0.25 |
| L22       | 0.25 |
| L23       | 0.25 |
| L24       | 0.25 |

| 1/2       |
| L25       | 0.25 |
| L26       | 0.25 |
The table shows the MIC values, at 3 and 9 days, for each replicate population of *P. aeruginosa* PA14 and 12 clinical isolates, evolved under sub-MIC concentrations of ciprofloxacin. Relative sub-MIC concentrations are determined based on the MIC of each strain in LBB. The concentrations belonging to the 9 days sub-MIC selective window are represented as a black cell. All MICs are obtained by E-test strips. Double inhibition halos are shown as two MIC values separated by a slash (X/X), being the highest value the one that was taken into consideration. Ciprofloxacin MIC values ≥ EUCAST breakpoint (0,5 µg/ml) are represented as a grey cell. L(A-L)1-4 refer to control populations evolved in absence of drug (in LBB). The data on PA14 strain come from [1].
Table S2 | MIC values (µg/ml) to antibiotics of different structural families in *P. aeruginosa* PA14 and 12 clinical isolates’ populations evolved under 9 days sub-MIC selective windows of ciprofloxacin.

| STRAINS | CONCENTRATION (relative to MIC) | REPLICATE | LEV (1) | CAZ (8) | AMK (16) | ATM (16) | IPM (4) | FOF (-) |
|---------|--------------------------------|-----------|---------|---------|----------|----------|--------|--------|
| PA14    | Parental strain                |           | 0,125   | 0,75    | 2        | 1,5      | 1      | 64     |
|         | LX1                             |           | 0,19    | 1       | 3        | 2        | 0,75   | 48     |
|         | LX2                             |           | 0,19    | 0,75    | 3        | 2        | 0,75   | 64     |
|         | LX3                             |           | 0,19    | 0,75    | 3        | 2        | 0,75   | 48     |
|         | LX4                             |           | 0,19    | 1       | 3        | 2        | 1      | 64     |
|         | X5                              |           | 0,38    | 2       | 8        | 12       | 1      | 48     |
|         | X6                              |           | 0,25    | 1       | 12       | 4        | 1      | 24     |
|         | X7                              |           | 0,25    | 0,75/3  | 3/8      | 4        | 1      | 24     |
|         | X8                              |           | 0,38    | 1,5     | 8        | 3        | 1,5    | 32     |
|         | X9                              |           | 2       | 1,5     | 6        | 2        | 1      | 32     |
|         | X10                             |           | 2       | 1,5     | 4        | 2        | 1      | 24     |
|         | X11                             |           | 2       | 1       | 6        | 1/6      | 1      | 32     |
|         | X12                             |           | 0,5/2   | 1       | 3        | 1,5/4    | 1      | 32     |
|         | X13                             |           | 2       | 1,5     | 4        | 6        | 1      | 24     |
|         | X14                             |           | 3       | 1       | 2        | 0,75     | 0,75   | 24     |
|         | X15                             |           | 2       | 1       | 2        | 2        | 1      | 24     |
|         | X16                             |           | 3       | 1       | 3        | 2        | 1,5    | 64     |
|         | X17                             |           | 3       | 1       | 4        | 1        | 1      | 16     |
|         | X18                             |           | 4       | 1       | 6        | 2        | 0,75   | 24     |
|         | X19                             |           | 4       | 2       | 4        | 1,5/4    | 0,75   | 24     |
| PA14    | X20                             |           | 4       | 1,5     | 4        | 2        | 0,75   | 24     |
|         | X21                             |           | 24      | 0,75    | 2        | 1        | 2      | 16     |
|         | X22                             |           | 6       | 1,5     | 2        | 1        | 1,5    | 24     |
|         | X23                             |           | 12      | 1       | 1,5      | 1        | 2      | 24     |
|         | X24                             |           | 12      | 0,75    | 1,5      | 1        | 0,75   | 16     |
|         | X25                             |           | ≥32     | 1       | 3        | 0,75     | 0,75   | 8      |
|         | X26                             |           | ≥32     | 0,75    | 1,5      | 1,5      | 2      | 24     |
|         | X27                             |           | ≥32     | 0,75    | 1,5      | 0,75     | 0,75   | 16     |
|         | X28                             |           | ≥32     | 0,75    | 1        | 1        | 1,5    | 16     |
|         | Parental strain                |           | 0,25    | 0,5     | 4        | 0,19     | 12     | 4      |
|         | LA1                             |           | 0,38    | 0,75    | 1,5      | 0,25     | 16     | 12     |
|         | LA2                             |           | 0,38    | 0,75    | 3        | 0,38     | 16     | 12     |
|         | LA3                             |           | 0,38    | 0,75    | 1,5      | 0,38     | 16     | 8      |
|         | LA4                             |           | 0,38    | 0,5     | 3        | 0,25/2   | 16     | 12     |
|         | A9                              |           | 0,38    | 0,75    | 4        | 2        | 12     | 4      |
| CAN01-002 | 1/50 | A10 | 0,19/0,75 | 1 | 2 | 0,38 | 16 | 6 |
| A11 | 0,75 | 0,38 | 2 | 0,19 | 12 | 4 |
| A12 | 0,75 | 0,75 | 3 | 0,38 | 24 | 6 |
| 1/25 | A13 | 0,5 | 0,5 | 3 | 0,25 | 16 | 8 |
| A14 | 0,75 | 0,25 | 3 | 0,19 | 12 | 4 |
| A15 | 0,5 | 0,75 | 3 | 0,19/2 | 12 | 4 |
| A16 | 0,5 | 0,5 | 3 | 0,19/1 | 12 | 6 |
| 1/10 | A17 | 0,75 | 0,5 | 1,5 | 0,19 | 6 | 4 |
| A18 | 1 | 0,75 | 2 | 0,19 | 16 | 3 |
| A19 | 0,38/4 | 0,5 | 3 | 0,19 | 16 | 4 |
| 1/5 | A20 | 1,5 | 0,5 | 2 | 0,38 | 16 | 3 |
| A21 | 1,5 | 0,5 | 2 | 0,19 | 12 | 4 |
| A22 | 1,5 | 0,5 | 2 | 0,25 | 16 | 3 |
| A23 | 1,5 | 0,38 | 2 | 0,25 | 16 | 4 |
| A24 | 1,5 | 0,5 | 2 | 0,19 | 16 | 4 |
| 1/2 | A25 | 1,5 | 0,38 | 1 | 0,125 | 1,5 | 4 |
| A26 | 1,5 | 0,38 | 1 | 0,125 | 1,5 | 4 |
| A27 | 2 | 0,38 | 1 | 0,125 | 1,5 | 4 |
| A28 | 1,5 | 0,38 | 1,5 | 0,19 | 2 | 4 |
| Parental strain | 0,25 | 1 | 3 | 3 | 2 | 256 |
| Parental strain | LB1 | 0,38 | 1 | 3 | 3 | 384 |
| Parental strain | LB2 | 0,25 | 1 | 3 | 3 | 384 |
| Parental strain | LB3 | 0,38 | 1 | 3 | 4 | 2 | 384 |
| Parental strain | LB4 | 0,38 | 1 | 3 | 4 | 2 | 256 |
| Parental strain | B9 | 0,19/1,5 | 1 | 4 | 4 | 1 | 256 |
| Without antibiotic | 1/50 | B10 | 0,19/1,5 | 1 | 3 | 4 | 3 | 256 |
| Without antibiotic | B11 | 0,19/1,5 | 1 | 3 | 4 | 2 | 256 |
| Without antibiotic | B12 | 0,19/1 | 1 | 3 | 4 | 4 | 256 |
| 1/25 | B13 | 1/3 | 1 | 3 | 4 | 6 | 256 |
| Without antibiotic | B14 | 3 | 1 | 2 | 4 | 4 | 384 |
| Without antibiotic | B15 | 4 | 1 | 3 | 3 | 4 | 256 |
| Without antibiotic | B16 | 3 | 1 | 1,5 | 2/6 | 4 | 256 |
| 1/10 | B17 | 2/4 | 0,75/2 | 3 | 8 | 3 | 256 |
| Without antibiotic | B18 | 16 | 0,75 | 1,5 | 3 | 4 | 256 |
| Without antibiotic | B19 | 3 | 0,75 | 1 | 2 | 4 | 384 |
| Without antibiotic | B20 | 3 | 0,5 | 1 | 1,5 | 6 | 256 |
| 1/5 | B21 | 3 | 0,75 | 1 | 2 | 6 | 384 |
| Without antibiotic | B22 | 3 | 0,75 | 1 | 1,5 | 6 | 384 |
| Without antibiotic | B23 | 3 | 0,5 | 1 | 1,5 | 6 | 256 |
| Without antibiotic | B24 | 3 | 0,5 | 1 | 1 | 6 | 256 |
| 1/2 | Without antibiotic | B25 | ≥32 | 0,5 | 1 | 1 | 4 | 192 |
| Without antibiotic | B26 | 24 | 0,75 | 1 | 1,5 | 6 | 256 |
|        | 1/2 |        |        |        |        |        |
|--------|-----|--------|--------|--------|--------|--------|
|        | B27 | 24     | 0,75   | 1      | 2      | 6      | 256   |
|        | B28 | 6      | 0,75   | 0,75   | 1,5    | 6      | 384   |
|        | Parental strain | 0,75 | 0,38   | 64     | 0,125  | 2      | 64    |
|        | LC1 | 0,75   | 0,25   | 64     | 0,125  | 1,5    | 96    |
|        | LC2 | 0,75   | 0,25   | 64     | 0,094  | 1,5    | 96    |
|        | LC3 | 0,5    | 0,25   | 192    | 0,125  | 2      | 48    |
|        | LC4 | 0,75   | 0,25   | 96     | 0,094  | 1,5    | 64    |
|        | C17 | 1/3    | 0,38   | ≥256   | 0,125  | 1,5    | 48    |
|        | C18 | 4      | 0,25   | ≥256   | 0,19   | 1,5    | 64    |
|        | C19 | 1/3    | 0,38   | ≥256   | 0,125  | 2      | 128   |
|        | C20 | 1      | 0,38   | ≥256   | 0,125  | 3      | 64    |
|        | C21 | 3      | 0,38   | 32     | 0,125  | 1,5    | 64    |
|        | C22 | 4      | 0,5    | 64     | 0,19   | 2      | 96    |
|        | C23 | 3      | 0,38   | 128    | 0,19   | 2      | 48    |
|        | C24 | 4      | 0,38   | 48     | 0,19   | 1,5    | 64    |
|        | C25 | 4      | 0,5    | 48     | 0,19   | 6      | 64    |
|        | C26 | 3      | 0,38   | 96     | 0,125  | 1,5    | 96    |
|        | C27 | 3      | 0,38   | 128    | 0,125  | 1,5    | 192   |
|        | C28 | 3      | 0,5    | 96     | 0,125  | 1,5    | 48    |

|        | 1/10|        |        |        |        |        |
|--------|-----|--------|--------|--------|--------|--------|
|        | Parental strain | 0,25 | 0,75 | 2      | 2      | 1      | ≥1024 |
|        | LD1 | 0,25   | 1      | 2      | 3      | 1      | ≥1024 |
|        | LD2 | 0,25   | 0,75   | 2      | 2      | 0,5    | ≥1024 |
|        | LD3 | 0,25   | 0,75   | 1,5    | 3      | 0,75   | ≥1024 |
|        | LD4 | 0,38   | 1,5    | 1,5    | 2      | 0,75   | ≥1024 |
|        | D5  | 0,19/0,5 | 1     | 1,5    | 2/6    | 1      | ≥1024 |
|        | D6  | 0,25/1  | 1      | 2      | 2/8    | 1      | ≥1024 |
|        | D7  | 0,25/0,75 | 1     | 2      | 2/6    | 0,75   | ≥1024 |
|        | D8  | 0,25/0,5 | 0,75   | 2      | 2/6    | 0,75   | ≥1024 |
|        | D9  | 2      | 2      | 2      | 8      | 1      | ≥1024 |
|        | D10 | 1,5    | 2      | 3      | 12     | 0,75   | ≥1024 |
|        | D11 | 2      | 3      | 3      | 12     | 0,75   | ≥1024 |
|        | D12 | 2      | 2      | 6      | 8      | 1      | ≥1024 |
|        | D13 | 3      | 1,5    | 2      | 2      | 0,75   | ≥1024 |
|        | D14 | 2      | 1,5    | 1,5    | 2/8    | 1      | ≥1024 |
|        | D15 | 2      | 1,5    | 2      | 1,5/8  | 0,75   | ≥1024 |
|        | D16 | 2      | 2      | 1,5    | 1/12   | 0,75   | ≥1024 |
|        | D17 | 2      | 0,75   | 1      | 2      | 0,75   | ≥1024 |
|        | D18 | 2      | 0,5    | 1,5    | 0,75   | 0,75   | ≥1024 |
|        | D19 | 2      | 1      | 1      | 1,5    | 0,75   | ≥1024 |
|        | D20 | 2      | 0,75   | 1,5    | 2      | 0,75   | ≥1024 |
|        | D21 | 3      | 0,75   | 0,75   | 1      | 0,38   | ≥1024 |
|        | D22 | 3      | 0,75   | 0,75   | 0,75   | 0,38   | ≥1024 |
|    |    |    |    |    |    |    |
|----|----|----|----|----|----|----|
| D23 | 16 | 1 | 0,75 | 1 | 0,25 | ≥1024 |
| D24 | 4  | 0,5 | 1 | 1 | 0,38 | ≥1024 |
| D25 | 16 | 0,5 | 1 | 1,5 | 0,5 | ≥1024 |
| D26 | 12 | 0,5 | 1 | 1 | 0,38 | ≥1024 |
| D27 | 4  | 0,5 | 0,75 | 0,5 | 0,25 | ≥1024 |
| D28 | 6  | 0,5 | 0,75 | 0,5 | 0,25 | ≥1024 |

### Without antibiotic

| Parental strain | 1,5 | 2 | 1 | 16 | 2 | 128 |
| LE1 | 2 | 3 | 2 | 16 | 2 | 192 |
| LE2 | 2 | 2 | 1,5 | 16 | 1,5 | 192 |
| LE3 | 1,5 | 3 | 2 | 16 | 2 | 128 |
| LE4 | 1,5 | 3 | 3 | 16 | 1,5 | 96 |

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| E13 | 3 | 3 | 4 | 24 | 3 | 128 |
| E14 | 4 | 4 | 3 | 32 | 2 | 192 |
| E15 | 4 | 6 | 3 | 24 | 2 | 192 |
| E16 | 3 | 6 | 6 | 24 | 3 | 192 |

### 1/10

| E17 | 3/16 | 3 | 1,5 | 16 | 3 | 128 |
| E18 | 6 | 3 | 1,5 | 16 | 1,5 | 192 |
| E19 | 6 | 4 | 4 | 16 | 4 | 192 |
| E20 | 8 | 3 | 2 | 16 | 3 | 128 |

### 1/5

| E21 | ≥32 | 3 | 1,5 | 16 | 3 | 128 |
| E22 | 4 | 3 | 3 | 16 | 2 | 128 |
| E23 | 12 | 3 | 3 | 16 | 3 | 128 |
| E24 | 3/12 | 3 | 4 | 16 | 2 | 96 |

### 1/2

| E25 | ≥32 | 3 | 2 | 16 | 4 | 384 |
| E26 | 12 | 2 | 1 | 8 | 2 | 128 |
| E27 | 24 | 3 | 2 | 16 | 3 | 192 |
| E28 | 12 | 3 | 1 | 16 | 3 | 96 |

### Without antibiotic

| Parental strain | 3 | 1 | 48 | 0,75 | 24 | ≥1024 |
| LF1 | 3 | 1,5 | 48 | 0,75 | 24 | ≥1024 |
| LF2 | 2 | 1 | 48 | 0,5/4 | 24 | ≥1024 |
| LF3 | 2 | 1 | 48 | 0,5 | 24 | ≥1024 |
| LF4 | 2 | 1 | 48 | 0,75 | 16 | ≥1024 |

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| F17 | 8 | 1 | 24 | 0,5 | 24 | ≥1024 |
| F18 | 8 | 1 | 32 | 0,5 | 32 | ≥1024 |
| F19 | 4 | 1 | 32 | 0,38 | 32 | ≥1024 |
| F20 | 6 | 1 | 32 | 0,38 | 24 | ≥1024 |
| F21 | 16 | 0,5 | 32 | 0,19 | 24 | ≥1024 |
| F22 | 6 | 1 | 32 | 0,38 | 24 | ≥1024 |
| F23 | 16 | 0,5 | 32 | 0,19 | 16 | ≥1024 |
| F24 | 16 | 0,75 | 48 | 0,38 | 24 | ≥1024 |
| F25 | 12 | 0,75 | 48 | 0,25 | 24 | ≥1024 |
| F26 | 12 | 1 | 48 | 0,38 | 24 | ≥1024 |
|       | F27 | F28 | Parental strain | LG1  | LG2  | LG3  | LG4  | G23 | G24 | G25 | G26 | G27 | G28 |
|-------|-----|-----|-----------------|------|------|------|------|-----|-----|-----|-----|-----|-----|
|       | 24  | 0,75| 32              | 0,38 | 24   | ≥1024 |      | 2   |     | 3   | 32  | 6   | 8   |
|       | 16  | 0,5 | 32              | 0,19 | 24   | ≥1024 |      | 2   |     | 2   | 32  | 6   | 8   |
| Without antibiotic | Parental strain | 0,75 | 3 | 32 | 6 | 8 | 384 | LG1  | 0,094/0,75 | 4 | 32 | 6 | 8 | 192 |
|       | LG2  | 0,125/0,75 | 2 | 32 | 6 | 8 | 192 | LG3  | 0,125/0,75 | 3 | 24 | 6 | 8 | 192 |
|       | LG4  | 0,125/0,75 | 4 | 32 | 6 | 8 | 192 | 1/5 G23 | 2 | 2 | 32 | 1/8 | 8 | 384 |
|       | G24  | 2 | 2 | 16 | 4 | 8 | 384 |       | G25 | 3 | 2 | 16 | 3 | 12 | 32 |
|       | G26  | 3 | 2 | 12 | 1 | 8 | 48 |       | G27 | 2 | 1,5 | 3/12 | 0,75 | 0,125/8 | 64 |
|       | G28  | 4 | 3 | 24 | 3 | 12 | 48 |       |       |       |       |       |       |       |
| GAL02-002 | 1/2 Parental strain | 0,25 | 1 | 6 | 0,19 | 1,5 | 64 | LH1  | 0,38 | 0,75 | 8 | 0,25 | 2 | 192 |
|       | LH2  | 0,38 | 1 | 6 | 0,25 | 2 | 192 |       | LH3  | 0,38 | 0,75 | 6 | 0,25 | 1,5 | 128 |
|       | LH4  | 0,38 | 0,75 | 6 | 0,25 | 1,5 | 128 |       |       | H5  | 2 | 0,75 | 12 | 0,25 | 2 | 128 |
|       | H6  | 2 | 1 | 8 | 0,25 | 1,5 | 96 |       | H7  | 2 | 0,75 | 8 | 0,25 | 1,5 | 192 |
|       | H8  | 1,5 | 0,75 | 4 | 0,25 | 2 | 192 |       | H9  | 1,5 | 1 | 3 | 0,38 | 1,5 | 128 |
|       | H10 | 1,5 | 0,75 | 2 | 0,38 | 1,5 | 128 |       | H11 | 2 | 1 | 2 | 0,38 | 1,5 | 128 |
|       | H12 | 2 | 1 | 4 | 0,25 | 1,5 | 192 |       | H13 | 1,5 | 0,75 | 3 | 0,25 | 1,5 | 192 |
|       | H14 | 2/4 | 0,75 | 3 | 0,19 | 3 | 128 |       | H15 | 1,5 | 1 | 2 | 0,38 | 3 | 128 |
|       | H16 | 1,5 | 0,75 | 2 | 0,19 | 1,5 | 192 |       | H17 | 8 | 0,75 | 3 | 0,19 | 8 | 128 |
|       | H18 | 8 | 0,75 | 1,5 | 0,25 | 4 | 128 |       | H19 | 6 | 1 | 2 | 0,38 | 6 | 128 |
|       | H20 | 8 | 0,5 | 2 | 0,38 | 6 | 128 |       | H21 | ≥32 | 0,75 | 3 | 0,38 | 3 | 96 |
| FQSE110603 | H22 | 8 | 1 | 2 | 0,19 | 6 | 192 |       | H23 | 8 | 0,5 | 3 | 0,125 | 4 | 128 |
|       | H24 | 12 | 1 | 1 | 0,19 | 8 | 128 |       | H25 | 12 | 0,5 | 2 | 0,19 | 4 | 128 |
|       | H26 | ≥32 | 0,5 | 2 | 0,19 | 6 | 64 |       | H27 | 12 | 0,38 | 3 | 0,19 | 4 | 128 |
|       | H28 | ≥32 | 0,75 | 3 | 0,19 | 4 | 96 |       |       |       |       |       |       |       |
| Parental strain | 0.38 | 0.75 | 8    | 3    | 4    | 32   |
|----------------|------|------|------|------|------|------|
| LI1            | 0.38 | 2    | 4    | 3    | 4    | 32   |
| LI2            | 0.38 | 1    | 4    | 4    | 3    | 32   |
| LI3            | 0.38 | 1    | 4    | 4    | 2    | 32   |
| LI4            | 0.38 | 1    | 4    | 4    | 3    | 32   |
| I13            | 2    | 6    | 12   | 4    | 4    | 48   |
| I14            | 2    | 2    | 6    | 12   | 3    | 64   |
| I15            | 1.5  | 2    | 8    | 16   | 3    | 48   |
| I16            | 2    | 2    | 6    | 16   | 3    | 48   |
| I17            | 2    | 4    | 6    | 16   | 2    | 128  |
| I18            | 2    | 4    | 6    | 16   | 3    | 64   |
| I19            | 2    | 2    | 4    | 16   | 3    | 48   |
| I20            | 2    | 1.5  | 4    | 3    | 2    | 48   |
| I21            | 2    | 4    | 6    | 16   | 2    | 128  |
| I22            | 3    | 6    | 4    | 16   | 3    | 64   |
| I23            | 8    | 3    | 4    | 24   | 3    | 48   |
| I24            | 2    | 3    | 6    | 24   | 3    | 64   |
| I25            | 16   | 0.5  | 3    | 1    | 2    | 48   |
| I26            | 24   | 0.5  | 3    | 1.5  | 8    | 48   |
| I27            | ≥32  | 0.5  | 2    | 1    | 12   | 48   |
| I28            | ≥32  | 0.75 | 2    | 1.5  | 6    | 64   |

ICA01-004

| Parental strain | 0.38 | 0.75 | 8    | 3    | 4    | 32   |
|----------------|------|------|------|------|------|------|
| LJ1            | 8    | 2    | 2    | 8    | 3    | ≥1024|
| LJ2            | 8    | 1.5  | 3    | 6    | 3    | ≥1024|
| LJ3            | 8    | 2    | 3    | 8    | 3    | ≥1024|
| LJ4            | 12   | 3    | 2    | 8    | 3    | ≥1024|
| J21            | 24   | 3    | 4    | 8    | 3    | ≥1024|
| J22            | ≥32  | 2    | 4    | 12   | 4    | ≥1024|
| J23            | 24   | 3    | 6    | 12   | 3    | ≥1024|
| J24            | 24   | 3    | 4    | 8    | 3    | ≥1024|
| J25            | ≥32  | 2    | 3    | 6    | 3    | ≥1024|
| J26            | ≥32  | 3    | 2    | 6    | 3    | ≥1024|
| J27            | ≥32  | 1.5  | 3    | 6    | 3    | ≥1024|
| J28            | 24   | 3    | 3    | 6    | 3    | ≥1024|

AND04-003

| Parental strain | 0.25 | 1    | 3    | 0.75 | 96   |
|----------------|------|------|------|------|------|
| LK1            | 0.25 | 1    | 3    | 1.5  | 0.75 | 64   |
| LK2            | 0.38 | 1.5  | 3    | 3    | 0.5  | 64   |
| LK3            | 0.25 | 1    | 3    | 2    | 0.75 | 96   |
| LK4            | 0.25 | 0.75 | 3    | 2    | 0.75 | 128  |
| K9             | 1    | 2    | 6    | 6    | 0.5  | 128  |
| K10            | 0.75 | 1    | 6    | 2/6  | 0.75 | 128  |
| K11            | 0.5  | 1    | 4    | 2/6  | 0.75 | 192  |
All MICs are obtained by E-test strips. Double inhibition halos are shown as two MIC values separated by a slash (X/X), being the highest value the one that is taken into consideration. Breakpoints of the antibiotics that are encompassed in EUCAST for *P. aeruginosa* are included in brackets. MIC values ≥ breakpoints are represented as a grey cell. L(A-L)1-4 refer to control populations evolved in absence of drug (in LBB). The
data on PA14 strain come from [1]. LEV: levofloxacin, CAZ: ceftazidime, AMK: amikacin, ATM: aztreonam, IPM: imipenem, FOF: fosfomycin.
Table S3 | MIC values (µg/ml) to antibiotics of different structural families in *P. aeruginosa* PA14 and 10 clinical isolates’ populations evolved under 0.04 µg/ml of ciprofloxacin “risk concentration” for 3 days.

| STRAINS | CONDITIONS | REPLICATE | ANTIBIOTIC |
|---------|------------|-----------|------------|
|         |            |           | CIP (0.5)  | LEV (1)  | CAZ (8)  | AMK (16) | ATM (16) | IPM (4)  | FOF (-) |
| PA14    | Parental strain | 0.064 | 0.19 | 0.75 | 1.5 | 1.5 | 1 | 32 |
|         | 3 days LB    | 1       | 0.064 | 0.19 | 0.5 | 2 | 2 | 1 | 16 |
|         |              | 2       | 0.047 | 0.125 | 0.75 | 3 | 1.5 | 1 | 24 |
|         |              | 3       | 0.047 | 0.19 | 0.5 | 2 | 2 | 1 | 24 |
|         | 3 days CIP   | 1*      | 1.5 | 3 | 0.75 | 1.5 | 2 | 0.75 | 24 |
|         |              | 2*      | 1 | 3 | 0.75 | 1 | 1.5 | 0.75 | 16 |
|         |              | 3*      | 1.5 | 3 | 1 | 1.5 | 1.5 | 0.75 | 24 |
| CAN01-002 | Parental strain | 0.047 | 0.125 | 0.5 | 3 | 0.19 | 8 | 2 |
|         | 3 days LB    | 1       | 0.064 | 0.19 | 0.5 | 3 | 0.25 | 12 | 3 |
|         |              | 2       | 0.064 | 0.19 | 0.5 | 4 | 0.25 | 8 | 3 |
|         |              | 3       | 0.064 | 0.19 | 0.75 | 3 | 0.25 | 12 | 3 |
|         | 3 days CIP   | 1*      | 0.5 | 1.5 | 0.75 | 2 | 0.094 | 6 | 3 |
|         |              | 2*      | 0.75 | 1 | 1 | 3 | 0.38 | 8 | 3 |
|         |              | 3*      | 0.25 | 0.75 | 0.5 | 2 | 0.19 | 12 | 2 |
| AND04-004A | Parental strain | 0.064 | 0.25 | 0.75 | 2 | 3 | 3 | 256 |
|         | 3 days LB    | 1       | 0.064 | 0.25 | 0.75 | 3 | 2 | 4 | 192 |
|         |              | 2       | 0.064 | 0.25 | 1 | 2 | 2 | 3 | 256 |
|         |              | 3       | 0.064 | 0.25 | 1 | 2 | 3 | 3 | 384 |
|         | 1*           | 0.75 | 3 | 0.75 | 0.75 | 1.5 | 6 | 192 |
|                | 3 days CIP | 3 days LB | 3 days CIP | 3 days CIP |
|----------------|------------|-----------|------------|------------|
|                | 2*         | 0,5       | 3          | 0,75       |
|                | 3*         | 0,75      | 2          | 0,75       |
| Parental strain| 0,125      | 0,38      | 0,19       | 64         |
|                | 1*         | 0,75      | 2          | 0,38       |
|                | 2*         | 1         | 2          | 0,38       |
|                | 3*         | 1         | 2          | 0,25       |
| FQSE06-0403    | 1          | 0,19      | 0,38       | 0,25       |
|                | 2          | 0,19      | 0,38       | 0,38       |
|                | 3          | 0,125     | 0,38       | 0,38       |
|                | 1*         | 0,75      | 2          | 0,38       |
|                | 2*         | 1         | 2          | 0,38       |
|                | 3*         | 1         | 2          | 0,25       |
|                | 1          | 0,064     | 0,25       | 0,5        |
|                | 2          | 0,047     | 0,25       | 0,5        |
|                | 3          | 0,064     | 0,25       | 0,38       |
| BAL04-002      | 1*         | 0,75      | 2          | 0,25       |
|                | 2*         | 0,5       | 1,5       | 2          |
|                | 3*         | 0,5       | 2          | 1,5       |
|                | 1          | 0,25      | 1          | 2          |
|                | 2          | 0,25      | 1          | 2          |
|                | 3          | 0,19      | 1,5       | 2          |
| CAT02-004      | 1*         | 0,5       | 3          | 1,5       |
|                | 2*         | 0,75      | 2          | 1          |
|                | 3*         | 0,5       | 2          | 1,5       |
|                | 1          | 0,125     | 0,38      | 2          |

**Note:** The values in the table represent concentrations in milligrams per liter (mg/L) and the times in hours (h).
| Strain        | 3 days LB | Parental strain | 3 days CIP | Parental strain | 3 days CIP | Parental strain |
|--------------|-----------|-----------------|-----------|-----------------|-----------|-----------------|
| **GAL02-002** |           |                 |           |                 |           |                 |
| LB           | 2         | 0,125           | 3         | 24              | 4         | 6               | 32            |
|              | 3         | 0,19            | 0,5       | 2               | 24        | 6               | 8             | 48            |
| CIP          | 1*        | 0,5             | 0,5       | 2               | 24        | 6               | 6             | 48            |
|              | 2*        | 0,5             | 0,5       | 2               | 24        | 4               | 8             | 64            |
|              | 3*        | 0,25            | 0,38      | 2               | 16        | 6               | 8             | 64            |
| **FQSE110603** |           |                 |           |                 |           |                 |
| LB           | 1         | 0,094           | 0,25      | 0,75            | 12        | 0,25            | 4             | 48            |
|              | 2         | 0,064           | 0,38      | 0,75            | 6         | 0,25            | 0,5           | 96            |
|              | 3         | 0,064           | 0,38      | 0,5             | 8         | 0,25            | 1,5           | 64            |
| CIP          | 1*        | 1               | 3         | 0,5             | 2         | 0,19            | 2             | 48            |
|              | 2*        | 1,5             | 3         | 0,19            | 2         | 0,25            | 6             | 64            |
|              | 3*        | 0,5             | 2         | 0,5             | 3         | 0,25            | 3             | 48            |
| **ICA01-004** |           |                 |           |                 |           |                 |
| LB           | 1         | 0,094           | 0,25      | 1               | 6         | 3               | 4             | 32            |
|              | 2         | 0,064           | 0,25      | 0,75            | 6         | 3               | 3             | 24            |
|              | 3         | 0,094           | 0,19      | 1               | 6         | 3               | 4             | 32            |
| CIP          | 1*        | 1,5             | 4         | 4               | 8         | 24              | 3             | 48            |
|              | 2*        | 1               | 3         | 4               | 12        | 16              | 8             | 24            |
|              | 3*        | 1               | 3         | 3               | 8         | 16              | 6             | 32            |
| **CLE03-004** |           |                 |           |                 |           |                 |
| LB           | 1         | 0,125           | 0,25      | 1               | 6         | 2               | 1             | 64            |
|              | 2         | 0,125           | 0,25      | 0,75            | 6         | 3               | 0,5           | 128           |
|              | 3         | 0,125           | 0,25      | 1               | 6         | 3               | 0,5           | 192           |
| CIP          | 1*        | 1,5             | 4         | 1,5             | 4/16      | 1/8             | 1             | 128           |
|              | 2*        | 2               | 6         | 1,5             | 16        | 1,5/8           | 1,5           | 128           |
| FQSE111010 | 3*  | 1,5 | 4  | 1,5 | 8  | 1,5/12 | 0,75 | 128 |
|------------|-----|-----|-----|-----|----|--------|------|-----|
| **Parental strain** |     |     |     |     |    |        |      |     |
|             | 0,25| 0,75| 0,75| 16  | 0,38| 12     | 48   |     |
| **3 days LB** | 1   | 0,38| 0,75| 0,75| 12 | 0,25   | 12   | 48  |
|             | 2   | 0,25| 0,75| 0,75| 16 | 0,25   | 12   | 48  |
|             | 3   | 0,38| 0,75| 0,75| 12 | 0,25   | 16   | 48  |
| **3 days CIP** | 1*  | 0,38| 0,75| 0,75| 16 | 0,38   | 8    | 48  |
|             | 2*  | 0,5 | 1   | 1   | 16 | 0,38   | 12   | 48  |
|             | 3*  | 0,5 | 1   | 0,75| 16 | 0,38   | 12   | 48  |

For each strain, 1-3 are the replicate populations evolved in LBB, whereas 1*-3* are the ones evolved in presence of 0,04 µg/ml of ciprofloxacin. All MICs are obtained by E-test strips. Double inhibition halos are shown as two MIC values separated by a slash (X/X), being the highest value the one that is taken into consideration. Breakpoints of the antibiotics that are encompassed in EUCAST for *P. aeruginosa* are included in brackets. MIC values ≥ breakpoints are represented as a grey cell. CIP: ciprofloxacin, LEV: levofloxacin, CAZ: ceftazidime, AMK: amikacin, ATM: aztreonam, IPM: imipenem, FOF: fosfomycin.
Table S4 | MIC values (µg/ml) to antibiotics and mutations in genes involved in quinolones resistance of *P. aeruginosa* single clones from PA14 and 10 clinical isolates’ populations evolved under 0,04 µg/ml of ciprofloxacin “risk concentration” for 3 days.

| STRAINS | SINGLE CLONES | ANTIBIOTIC | QUINOLONE RESISTANCE MUTATIONS |
|---------|---------------|------------|--------------------------------|
|         |               | CIP (0,5)  | LEV (1) | CAZ (8) | AMK (16) | ATM (16) | IPM (4) | MER (8) | CHL | ERY | TET | TGC | Nucleotide change | Amino acid change |
| PA14    | Parental strain | 0.064 | 0.19 | 1 | 1.5 | 2 | 1 | 0.125 | 48 | 48 | 4 | 1 | - | - |
|         | 1* SC | 1.5 | 4 | 1 | 1 | 1 | 1 | 0.125 | ≥256 | 64 | 12 | 4 | - | - |
|         | 3* SC | 1 | 2 | 1 | 1 | 1.5 | 0.75 | 0.38 | ≥256 | 64 | 12 | 4 | mexS T98G | MexS V33G |
| CAN01-002 | Parental strain | 0.094 | 0.38 | 1 | 4 | 0.19 | 12 | 0.38 | ≥256 | 96 | 4 | 1.5 | - | - |
|         | 1* SC | 0.38 | 1.5 | 0.38 | 2 | 0.094 | 2 | 1 | 96 | ≥256 | 3 | 0,75 | nfxB 467DelC | NfxB V156fs |
|         | 2* SC | 0.5 | 1 | 0.25 | 1 | 0.094 | 1.5 | 1 | ≥256 | ≥256 | 12 | 2 | nfxB A564C | NfxB W188fs |
| AND04-004A | Parental strain | 0.064 | 0.25 | 1 | 4 | 3 | 3 | 1 | ≥256 | 96 | 6 | 3 | - | - |
|         | 1* SC | 0.38 | 1.5 | 0.75 | 0.75 | 2 | 4 | 1.5 | ≥256 | 48 | 12 | 3 | - | - |
|         | 3* SC | 0.5 | 2 | 0.75 | 1 | 1.5 | 8 | 2 | ≥256 | 48 | 12 | 3 | gyrA A346C | mexS C961T | GyrA N116H MexS H321L |
| FQSE06-0403 | Parental strain | 0.19 | 0.5 | 0.38 | 96 | 0.125 | 3 | 0.19 | 12 | 192 | 6 | 6 | - | - |
|         | 3* SC | 0.5 | 0.75 | 0.25 | 96 | 0.094 | 4 | 0.25 | 12 | ≥256 | 6 | 6 | - | - |
| BAL04-002 | Parental strain | 0.094 | 0.19 | 1 | 3 | 2 | 1 | 0.25 | ≥256 | 128 | 8 | 3 | - | - |
|         | 1* SC | 0.75 | 1.5 | 0.75 | 1.5 | 0.75 | 0.5 | 0.38 | ≥256 | ≥256 | 12 | 3 | nfxB T562A | NfxB W188fs |
|         | 2* SC | 4 | 3 | 0.75 | 1.5 | 0.75 | 0.38 | 0.38 | ≥256 | ≥256 | 16 | 2 | nfxB A564C | NfxB W188fs |
| CAT02-004 | Parental strain | 0.25 | 1.5 | 4 | 1 | 12 | 2 | 16 | ≥256 | ≥256 | 32 | 6 | - | - |
|         | 2* SC | 0.5 | 2 | 3 | 1 | 8 | 2 | 4 | ≥256 | ≥256 | 24 | 6 | - | - |
| GAL02-002 | Parental strain | 0.19 | 0.5 | 1.5 | 24 | 4 | 8 | 24 | 96 | ≥256 | 4 | 6 | - | - |
|         | 1* SC | 0.5 | 0.5 | 2 | 24 | 6 | 6 | 24 | 96 | ≥256 | 6 | 6 | - | - |
| FQSE110603 | Parental strain | 0.094 | 0.25 | 0.75 | 8 | 0.25 | 3 | 0.047 | 32 | 96 | 3 | 1.5 | - | - |
|         | 1* S | 1.5 | 4 | 0.5 | 1.5 | 0.125 | 4 | 0.064 | ≥256 | 96 | 8 | 6 | mexS 548Del 10 bp | MexS R183fs |
|         | 2* SC | 1.5 | 6 | 0.5 | 2 | 0.19 | 6 | 0.125 | ≥256 | 64 | 8 | 3 | mexS 80D 18 bp | MexS A27fs |
|         | Parental strain | 0.094 | 0.25 | 1 | 4 | 2 | 3 | 0.75 | 96 | 96 | 6 | 2 | - | - |
|          | 1* SC | 1,5 | 3 | 4 | 16 | 6 | 1 | ≥256 | 128 | 24 | 4 |   |   |
|----------|-------|-----|---|---|----|---|---|------|-----|----|---|---|---|
| ICA01-004 | 0,38  |     |   |   |    |   |   |      |     |    |   | - | - |
| 2* SC    | 1     | 3   | 0,75 | 1,5 | 1,5 | 12 | 1 | ≥256 | 128 | 8  | 4 |   | - |
| Parental strain | 0,094 | 0,25 | 1 | 3 | 2 | 0,75 | 0,064 | 48 | 192 | 6 | 4 | - | - |
| CLE03-004 | 1,5   | 3   | 0,75 | 1,5 | 0,75 | 1,5 | 0,094 | ≥256 | 192 | 12 | 6 | mexS G406T | MexS G136C |
| 2* SC    | 0,38  | 1   | 1   | 2 | 1,5 | 1 | 0,094 | ≥256 | 192 | 6 | 4 | parE C1103G | mexS 302InsCGA |
| Parental strain | 0,25  | 0,75 | 1 | 12 | 0,38 | 12 | 32 | ≥256 | ≥256 | 6 | 3 | - | - |
| FQSE111010 | 0,5   | 0,75 | 0,75 | 12 | 0,38 | 8 | 4 | ≥256 | ≥256 | 4 | 4 | - | - |

For each strain, 1*-3* SC are the representative single clones from populations 1*-3* (Table S3) evolved in presence of 0.04 µg/ml of ciprofloxacin for 3 days. All MICs are obtained by E-test strips. Breakpoints of the antibiotics that are encompassed in EUCAST for *P. aeruginosa* are included in brackets. MIC values ≥ breakpoints are represented as a grey cell. The single clones that, based on its MDR phenotype and genotype, were chosen for gene expression analysis, are underlined. Del: deletion, Ins: insertion, Fs: frameshift, bp: base pairs, CIP: ciprofloxacin, LEV: levofloxacin, CAZ: ceftazidime, AMK: amikacin, ATM: aztreonam, IPM: imipenem, MER: meropenem, CHL: chloramphenicol, ERY: erythromycin, TET: tetracycline, TGC: tigecycline.
| Primer | Sequence (5’-3’) | Description |
|--------|-----------------|-------------|
| **PCR amplification** | | |
| **gyrA.fw** | AGTCCTATCTCGACTACGCGAT | To amplify QRDR* in *gyrA* |
| **gyrA.rv** | AGTCGACGTTTTCCCTTTCCAG | | |
| **gyrB.fw** | CGCGGTGGAACAGGAGATGGGCAAGTAC | To amplify QRDR in *gyrB* |
| **gyrB.rv** | CTGGCGGAAGAAGAAGGTCAACACGAGGTT | | |
| **parC.fw** | CGAGCAGGCTATCTGAATTAT | To amplify QRDR in *parC* |
| **parC.rv** | GAAGGACTTGGGATCGTGCCG | | |
| **parE.fw** | CGGCGTTCGTCTCGGCGTGTTAGGAAGGA | To amplify QRDR in *parE* |
| **parE.rv** | TCGAGGCGTAGTAGATTTCCCTGCGGA | | |
| **nfxB.fw** | TTCTGCACAATGCGCACAA | To amplify the open reading frame of *nfxB* |
| **nfxB.rv** | CGTGCCATGCGGCGACGAG | | |
| **mexS.fw** | TGAGGTTATTCAACCCGTGA | To amplify the open reading frame of *mexS* |
| **mexS.rv** | TCCGCTGCGCGGTCCCAAT | | |
| **mexT.fw** | ACGCAAGGCTTGACGGCGA | To amplify the open reading frame of *mexT* |
| **mexT.rv** | GCGGTGCGCGATCGATTT | | |
| **RT-PCR amplification** | | |
| **mexC.fw** | GACCTGCTGTTCAGATCG | To amplify *mexC* by RT-PCR |
| **mexC.rv** | GTGGCGGTATCGAAGTCCT | | |
| **mexE.fw** | ACTTCCTCGACAAACCAGGTC | To amplify *mexE* by RT-PCR |
| **mexE.rv** | GTGCAGTGACCGGTCTTGT | | |
To check DNA contamination from RNA samples and to amplify the housekeeping gene *rplU* by RT-PCR.

| **rplU.fw** | CGCAGTGATTGTACCAGTG |
|-------------|----------------------|
| **rplU.rv** | AGGCCTGAATGCGGTGATC  |

*QRDR: Quinolone Resistance Determining Region.*
Table S6 | MIC values (µg/ml) to antibiotics of different structural families in presence and absence of the EPI PAβN of single clones from populations of *P. aeruginosa* PA14 and of clinical strains of *P. aeruginosa* evolved under ciprofloxacin “risk concentration”.

|        | CIP  | LEV  | IPM  | MER  | CHL  | ERY  | TET  | TGC  |
|--------|------|------|------|------|------|------|------|------|
|        | MH PAβN | MH PAβN | MH PAβN | MH PAβN | MH PAβN | MH PAβN | MH PAβN | MH PAβN |
| PA14 3* SC | 1   | 0.19 | 2   | 0.094 | 0.75 | 0.75 | 0.38 | 0.5 | ≥256 | 128 | 64 | 16 | 12 | 2   | 4   | 0.75 |
| AND04-004A 3* SC | 0.5 | 0.125 | 3   | 0.064 | 4   | 4   | 3   | 2   | ≥256 | ≥256 | 48 | 24 | 12 | 6   | 4   | 0.38 |
| FQSE110603 2* SC | 1.5 | 0.25 | 6   | 1   | 4   | 4   | 0.125 | 0.125 | ≥256 | ≥256 | 48 | 24 | 8  | 4   | 3   | 0.75 |
| ICA01-004 2* SC | 1   | 0.25 | 3   | 0.5  | 8   | 6   | 0.75 | 0.75 | ≥256 | ≥256 | 128 | 32 | 12 | 6   | 4   | 1   |
| CLE03-004 1* SC | 1   | 0.125 | 4   | 0.25  | 0.75 | 0.75 | 0.094 | 0.094 | ≥256 | 64  | 192 | 12 | 16 | 4   | 8   | 0.38 |
| CAN01-002 2* SC | 0.38 | X    | 1   | X   | 1.5  | X    | 0.75 | X    | ≥256 | X   | ≥256 | X | 12 | X   | 3   | X   |
| BAL04-002 2* SC | 2   | 0.25 | 2   | 0.125 | 0.25 | 0.25 | 0.38 | 0.5 | ≥256 | 16  | ≥256 | ≥256 | 12 | 6   | 4   | 0.5  |

For each strain, 1*-3* SC are the representative single clones from populations 1*-3* (Table S3) evolved in presence of 0,04 µg/ml of ciprofloxacin for 3 days. All MICs were obtained by E-test strips, in absence and in presence of 25 µg/ml of the EPI PAβN. “X” indicates the clone that was unable to grow in presence of PAβN.

EPI: efflux pump inhibitor, CIP: ciprofloxacin, LEV: levofloxacin, IPM: imipenem, MER: meropenem, CHL: chloramphenicol, ERY: erythromycin, TET: tetracycline, TGC: tigecycline.
1. Sanz-Garcia, F., S. Hernando-Amado, and J.L. Martinez, *Evolution under low antibiotic concentrations: a risk for the selection of Pseudomonas aeruginosa multidrug-resistant mutants in nature*. Environ Microbiol, 2021.