

Electronic Supplementary Information for
The effects of structural changes on anti-microbial and anti-proliferative activity of
diimidazolium salts

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General procedure for the synthesis of the dibromide salts

α,α' -Dibromoxylene (2.5 g, 9.5 mmol) was dissolved in 2-propanol (10 mL). The obtained solution was placed in an oil bath at 85 °C. N-alkylimidazole (20 mmol) was dissolved in 2-propanol (10 mL) and the solution was added, dropwise, to the α,α' -dibromoxylene solution. The reaction mixture was heated at 85 °C for 24 h. After cooling, in the case of *para*-salts the reaction mixture was diluted with acetone (200 mL) and a white solid was collected by filtration. In the case of *meta*- and *ortho*-salts after concentration, yellow viscous oil was obtained, which was washed several times with diethyl ether. The obtained dibromide salts were dissolved in anhydrous methanol and stirred overnight (12 h) at room temperature, in the presence of active charcoal (1% by weight). After filtration on neutral aluminum oxide and concentration in vacuo, the desired salt was obtained.

3,3'-Di-n-hexyl-1,1'-(1,4-phenylenedimethylene)diimidazolium dibromide, [***p*-C₆im**][Br]₂: White solid; m.p. 165-169 °C; yield 83%; ¹H NMR (300 MHz, [D₆]DMSO): δ = 9.47 (s, 2H), 7.89 (m, 4H), 7.46 (s, 4H), 5.51 (s, 4H), 4.23 (t, J=7.2 Hz, 4H), 1.84 (qt, J= 6.5 Hz, 4H), 1.20 (m, 12H), 0.90 (t, J= 6.7 Hz, 6H) ppm; ¹³C NMR (300 MHz, [D₆]DMSO): δ = 136.3, 135.6, 129.0, 123.0, 122.7, 51.6, 49.1, 30.6, 29.33, 25.3, 22.0, 13.9 ppm; elemental analysis calcd (%) for C₂₆H₄₀Br₂N₄ (568): C 54.94, H 7.01, N 9.96, found: C 54.91, H 6.99, N 9.94.

3,3'-Di-n-octyl-1,1'-(1,2-phenylenedimethylene)diimidazolium dibromide, [***o*-C₈im**][Br]₂: White solid; m.p. 182-185 °C; yield 60%; ¹H NMR (300 MHz, [D₆]DMSO): δ = 9.36 (s, 2H), 7.88 (m, 2H), 7.79 (m, 2H), 7.48 (m, 2H), 7.30 (m, 2H), 5.66 (s, 4H), 4.19 (t, J= 7.3 Hz, 4H), 1.80 (t, J= 6.6 Hz, 4H), 1.25 (m, 20H), 0.85 (t, J= 6.9 Hz, 6H) ppm; ¹³C NMR (300 MHz, [D₆]DMSO): δ = 136.9, 133.4, 130.1, 130.0, 123.3, 123.2, 49.5, 31.6, 29.8, 29.0, 28.8, 26.0, 22.5, 14.4 ppm; elemental analysis calcd (%) for C₃₀H₄₈Br₂N₄ (624): C 57.69, H 7.75, N 8.97, found: C 57.72, H 7.73, N 9.00.

3,3'-Di-n-decyl-1,1'-(1,3-phenylenedimethylene)diimidazolium dibromide, [***m*-C₁₀im**][Br]₂: orange wax; yield 77%; ¹H NMR (300 MHz, [D₆]DMSO): δ = 9.40 (s, 2H), 7.84 (m, 4H), 7.57 (s, 1H), 7.48 (m, 1H), 7.40 (m 2H), 5.45 (s, 4H), 4.18 (t, J=7.5 Hz, 4H), 1.79 (qt, J= 7.5 Hz, 4H), 1.24 (m, 28H), 0.86 (t, J= 6.7 Hz, 6H) ppm; ¹³C NMR (300 MHz, [D₆]DMSO): δ = 136.7, 136.0, 130.2, 129.0, 128.9, 123.3, 123.0, 52.1, 49.5, 31.7, 29.7, 29.3, 29.3, 29.1, 28.8, 26.0, 22.5, 14.4 ppm; elemental analysis calcd (%) for C₃₄H₅₆Br₂N₄ (680): C 60.00, H 8.29, N 8.23, found: C 59.98, H 8.31, N 8.22.

3,3'-Di-n-decyl-1,1'-(1,2-phenylenedimethylene)diimidazolium dibromide, [o-C₁₀im][Br]₂: beige waxy-like solid; yield 76%; ¹H NMR (300 MHz, [D₆]DMSO): δ= 9.29 (s, 2H), 7.88 (m, 2H), 7.76 (m, 2H), 7.49 (m, 2H), 7.30 (m, 2H), 5.63 (s, 4H), 4.18 (t, J=6.0 Hz, 4H), 1.80 (qt, J= 7.5 Hz, 4H), 1.24 (m, 28H), 0.86 (t, J= 6.0 Hz, 6H) ppm; ¹³C NMR (300 MHz, [D₆]DMSO): δ= 136.9, 133.4, 130.1, 130.0, 123.3, 123.2, 49.5, 31.7, 29.8, 29.4, 29.3, 29.1, 28.8, 26.0, 22.5, 14.4 ppm; elemental analysis calcd (%) for C₃₄H₅₆Br₂N₄ (680): C 60.00, H 8.29, N 8.23, found: C 60.02, H 8.30, N 8.25.

General procedure for the synthesis of salts different than bromide

A column packed with Amberlite resin IRA-400 (chloride form, 11.50 g) was used to perform the exchange of anions. To convert the chloride form of the resin into the hydroxide form, it was firstly washed with an aqueous solution of NaOH (570 mL, 10% w/v) and subsequently with water until the eluate was neutral. A binary mixture of methanol/water (70:30, v/v) was used as eluent. Bromide salt (1.68 g, 2.69 mmol) was dissolved in the binary mixture (50 mL) and eluted. The eluate was collected in a flask containing a solution of the desired acid in stoichiometric amount. The neutral solution was concentrated in vacuo and the residue was dissolved in ethanol. The obtained solution was heated in the presence of active charcoal, and filtered. After the solvent removal under vacuum, the salt was washed with acetone to eliminate traces of other solvents. When possible the salts were crystallized from the ethanolic solution. The 2,6-naphthalenedisulfonic acid was obtained by elution of a water solution of the corresponding sodium salt through an Amberlite IR 120 plus column. In each case, the silver nitrate test, performed to verify the presence of residual bromide anion, gave a negative result.

3,3'-Di-n-hexyl-1,1'-(1,4-phenylenedimethylene)diimidazolium ditetrafluoroborate, [p-C₆im][BF₄]₂: White solid; m.p. 110-115 °C; yield: 97%; ¹H NMR (400 MHz, [D₆]DMSO): δ= 9.39 (s, 2H), 7.80 (m, 4H), 7.46 (s, 4H), 5.43 (s, 4H), 4.16 (t, J= 8.0 Hz, 4H), 1.77 (qt, J= 8.0 Hz, 4H), 1.24 (m, 12H), 0.83 (t, J= 8.0 Hz, 6H) ppm; ¹³C NMR (300 MHz, [D₆]DMSO): δ= 135.9, 129.4, 123.3, 123.0, 51.9, 49.4, 30.9, 29.6, 25.6, 22.3, 14.2 ppm. elemental analysis calcd (%) for C₂₆H₄₀B₂F₈N₄ (582): C 53.63, H 6.92, N 9.62, found: C 53.60, H 6.91, N 9.60.

3,3'-Di-n-hexyl-1,1'-(1,4-phenylenedimethylene)diimidazolium D-Tartrate, [p-C₆im][D-Tar]: Hygroscopic white solid; yield: 96%; ¹H NMR (400 MHz, [D₆]DMSO): δ= 9.54 (s, 2H), 7.82 (m, 4H), 7.48 (s, 4H), 5.45 (s, 4H), 4.17 (t, J= 6.0 Hz, 4H), 3.73 (s, 2H), 1.77 (qt, J= 7.0 Hz, 4H), 1.24 (m, 12H), 0.84 (t, J= 6 Hz, 6H) ppm; ¹³C NMR (300 MHz, [D₆]DMSO): δ= 175.2, 136.0, 129.4,

123.2, 123.0, 72.1, 51.8, 49.4, 30.9, 29.7, 25.6, 22.3, 14.2 ppm; elemental analysis calcd (%) for $C_{30}H_{44}N_4O_6$ (556): C 64.73, H 7.97, N 10.06, found: C 64.74, H 7.95, N 10.04. $[\alpha]_{24.1}^{589nm} = -3.8^\circ$.

3,3'-Di-n-hexyl-1,1'-(1,4-phenylenedimethylene)diimidazolium 2,6-naphthalendicarboxylate, [**p-C₆im**][**2,6-NDC**]: White solid; m.p. 223-228 °C; yield: 98%; 1H NMR (400 MHz, [D₆]DMSO): δ = 9.45 (s, 2H), 8.35 (m, 2H), 7.96 (d, J= 8.0 Hz, 2H), 7.82 (m, 6H), 7.47 (s, 4H), 5.44 (s, 4H), 4.17 (t, J= 8.0 Hz, 4H), 1.78 (qt, J= 8.0 Hz, 4H), 1.24 (m, 12H), 0.84 (t, J= 8.0 Hz, 6H) ppm; ^{13}C NMR (300 MHz, [D₆]DMSO): δ = 169.1, 137.0, 136.0, 133.3, 129.3, 128.2, 127.3, 124.1, 123.3, 123.0, 122.7, 51.9, 49.4, 31.8, 30.9, 25.6, 22.3, 14.2 ppm. elemental analysis calcd (%) for $C_{38}H_{46}N_4O_4$ (622): C 73.28, H 7.44, N 9.00, found: C 73.27, H 7.41, N 8.98.

3,3'-Di-n-octyl-1,1'-(1,4-phenylenedimethylene)diimidazolium ditetrafluoroborate, [**p-C₈im**][**BF₄**]₂: White solid; m.p. 99.8-104.7 °C; yield: 98%; 1H NMR (300 MHz, [D₆]DMSO): δ = 9.36 (s, 2H), 7.85 (dt, J= 1.6, 7.6 Hz, 4H), 7.51 (s, 4H), 5.48 (s, 4H), 4.22 (t, J= 7.2 Hz, 4H), 1.84 (qt, J= 6.9 Hz, 4H), 1.30 (m, 20H), 0.91 ppm (t, J= 6.8 Hz, 6H); ^{13}C NMR (300 MHz, [D₆]DMSO): δ = 136.3, 135.6, 129.0, 123.0, 122.7, 51.7, 49.2, 31.3, 29.4, 28.6, 28.4, 25.6, 22.2, 14.1 ppm. ^{19}F NMR ([D₆]DMSO): δ : -147.46 ppm. elemental analysis calcd (%) for $C_{30}H_{48}B_2F_8N_4$ (638): C 56.45, H 7.58, N 8.78; found: C 56.40, H 7.60, N 8.75.

3,3'-Di-n-octyl-1,1'-(1,3-phenylenedimethylene)diimidazolium ditetrafluoroborate, [**m-C₈im**][**BF₄**]₂: pale-yellow oil; yield: 98%; 1H NMR (300 MHz, [D₆]DMSO): δ = 9.42 (s; 2H), 7.81 (d; J = 3.6 Hz; 4H), 7.47 (m; 4H), 5.43 (s; 4H), 4.16 (t; J= 7.3 Hz; 4H), 1.78 (qt, J= 7.0 Hz, 4H), 1.24 (m; 20H), 0.85 (t; J = 6.7 Hz; 6H) ppm; ^{13}C NMR (300 MHz, [D₆]DMSO): δ = 138.8, 138.2, 132.3, 131.1, 131.0, 125.4, 125.1, 54.2, 51.5, 33.7, 31.8, 31.0, 30.9, 28.1, 24.6, 16.5 ppm. elemental analysis calcd (%) for $C_{30}H_{48}B_2F_8N_4$ (638): C 56.45, H 7.58, N 8.78; found: C 56.48, H 7.57, N 8.77.

3,3'-Di-n-octyl-1,1'-(1,2-phenylenedimethylene)diimidazolium ditetrafluoroborate, [**o-C₈im**][**BF₄**]₂: pale-yellow oil; yield: 97%; 1H NMR (300 MHz, [D₆]DMSO): δ = 9.28 (s; 2H), 7.85 (s; 2H), 7.74 (s; 2H), 7.48 (m; 2H), 7.29 (m; 2H), 5.60 (s; 4H), 4.16 (t; J= 7.3 Hz; 4H), 1.79 (qt, J= 6.9 Hz, 4H), 1.24 (m; 20H), 0.85 (t; J = 6.6 Hz; 6H) ppm; ^{13}C NMR (300 MHz, [D₆]DMSO): δ = 136.9, 133.4, 130.1, 130.0, 123.3, 123.2, 49.5, 31.6, 29.8, 29.0, 28.8, 26.0, 22.5, 14.4 ppm. elemental analysis calcd (%) for $C_{30}H_{48}B_2F_8N_4$ (638): C 56.45, H 7.58, N 8.78; found: C 56.48, H 7.57, N 8.80.

3,3'-Di-n-octyl-1,1'-(1,4-phenylenedimethylene)diimidazolium L-Tartrate: Hygroscopic white solid; m.p. 202-206 °C; yield: 85%; ¹H NMR (300 MHz, [D₆]DMSO): δ= 9.78 (s; 2H), 7.88 (d; J = 6.4 Hz; 4H), 7.55 (s; 4H), 5.53 (s; 4H), 4.23 (t; J = 4.5 Hz; 4H), 3.77 (s; 2H), 1.83 (m; 4H), 1.24 (m; 20H), 0.90 (t; J = 3.6 Hz; 6H) ppm; ¹³C NMR (300 MHz, [D₆]DMSO): δ= 175.6, 137.0, 135.7, 129.1, 122.9, 122.6, 73.3, 51.5, 49.1, 31.3, 29.4, 28.6, 28.4, 25.7, 22.2, 14.1 ppm. elemental analysis calcd (%) for C₃₄H₅₂N₄O₆ (612): C 66.64, H 8.55, N 9.14; found: C 66.68, H 8.53, N 9.13. $[\alpha]_{23.6}^{589nm} = 10.3^\circ$.

3,3'-Di-n-octyl-1,1'-(1,4-phenylenedimethylene)diimidazolium D-Tartrate: Hygroscopic white solid; m.p. 202-204 °C; yield: 87%; ¹H NMR (300 MHz, [D₆]DMSO): δ= 9.79 (s; 2H), 7.88 (d; J = 7.4 Hz; 4H), 7.55 (s; 4H), 5.52 (s; 4H), 4.23 (t; J = 4.8 Hz; 4H), 3.77 (s; 2H), 1.83 (m; 4H), 1.20 (m; 20H), 0.90 (t; J = 4.4 Hz; 6H) ppm; ¹³C NMR (300 MHz, [D₆]DMSO): δ= 175.6, 136.9, 135.7, 129.2, 122.9, 122.6, 73.1, 51.6, 49.1, 31.3, 29.5, 28.6, 28.5, 25.7, 22.2, 14.1 ppm. elemental analysis calcd (%) for C₃₄H₅₂N₄O₆ (612): C 66.64, H 8.55, N 9.14; found: C 66.62, H 8.54, N 9.16. $[\alpha]_{23.6}^{589nm} = -11.9^\circ$.

3,3'-Di-n-octyl-1,1'-(1,2-phenylenedimethylene)diimidazolium D-Tartrate: Hygroscopic white wax; yield: 88%; ¹H NMR (300 MHz, [D₆]DMSO): δ= 9.68 (s; 2H), 7.82 (s; 2H), 7.78 (s; 2H), 7.46 (m; 2H), 7.33 (m; 2H), 5.71 (s; 4H), 4.17 (t; J = 7.2 Hz; 4H), 3.75 (s; 2H), 1.79 (qt; J = 6.5 Hz; 4H), 1.24 (m; 20H), 0.85 (t; J = 6.6 Hz; 6H) ppm; ¹³C NMR (300 MHz, [D₆]DMSO): δ= 175.7, 137.5, 133.7, 130.2, 130.0, 123.1, 123.1, 72.9, 49.4, 31.6, 29.8, 29.0, 28.8, 26.0, 22.5, 14.4 ppm. elemental analysis calcd (%) for C₃₄H₅₂N₄O₆ (612): C 66.64, H 8.55, N 9.14; found: C 66.62, H 8.54, N 9.16. $[\alpha]_{24.1}^{589nm} = -5.1^\circ$.

3,3'-Di-n-octyl-1,1'-(1,4-phenylenedimethylene)diimidazolium meso-Tartrate: Hygroscopic white solid; m.p. 182-185 °C; yield: 86%; ¹H NMR (300 MHz, [D₆]DMSO): δ= 9.78 (s; 2H), 7.77 (d; J = 6.6 Hz; 4H), 7.48 (s; 4H), 5.49 (s; 4H), 4.19 (tb; 4H), 3.33 (s; 2H), 1.83 (m; 4H), 1.16 (m; 20H), 0.84 (t; J = 4.4 Hz; 6H) ppm; ¹³C NMR (300 MHz, [D₆]DMSO): δ= 174.5, 137.2, 135.6, 129.0, 122.6, 122.3, 79.2, 74.9, 51.3, 48.8, 31.1, 30.6, 29.4, 28.6, 28.5, 28.3 ppm. elemental analysis calcd (%) for C₃₄H₅₂N₄O₆ (612): C 66.64, H 8.55, N 9.14; found: C 66.60, H 8.58, N 9.13.

3,3'-Di-n-octyl-1,1'-(1,4-phenylenedimethylene)diimidazolium ±-Tartrate: Hygroscopic white solid; m.p. 201-204 °C; yield: 90%; ¹H NMR (300 MHz, [D₆]DMSO): δ= 9.81 (s; 2H), 7.88 (dd; J = 1.0, 7.0 Hz; 4H), 7.55 (s; 4H), 5.53 (s; 4H), 4.23 (t; J = 4.8 Hz; 4H), 3.76 (s; 2H), 1.83 (m; 4H), 1.18 (m;

20H), 0.90 (t; J = 4.5 Hz; 6H) ppm; ^{13}C NMR (300 MHz, $[\text{D}_6]\text{DMSO}$): δ = 176.5, 137.8, 136.5, 129.9, 123.6, 123.4, 80.1, 74.2, 52.3, 49.8, 32.0, 31.5, 30.2, 29.5, 29.4, 29.2 ppm. elemental analysis calcd (%) for $\text{C}_{34}\text{H}_{52}\text{N}_4\text{O}_6$ (612): C 66.64, H 8.55, N 9.14; found: C 66.65, H 8.54, N 9.15.

3,3'-Di-n-octyl-1,1'-(1,4-phenylenedimethylene)diimidazolium 2,6-naphthalendicarboxylate: White solid; m.p. 186-190 °C; yield: 96%; ^1H NMR (300 MHz, $[\text{D}_6]\text{DMSO}$): δ = 9.91 (s; 2H), 8.34 (s; 2H), 8.00 (d; J = 8.7 Hz; 2H), 7.89 (s; 4H), 7.76 (d; J = 8.4 Hz; 2H), 7.52 (s; 4H), 5.53 (s; 4H), 4.23 (t; J = 7.2 Hz; 4H), 1.83 (qt; J = 6.3 Hz; 4H), 1.19 (m; 20H), 0.88 (t; J = 6.7 Hz; 6H) ppm; ^{13}C NMR (300 MHz, $[\text{D}_6]\text{DMSO}$): δ = 169.9, 140.0, 137.8, 136.6, 134.0, 130.0, 128.8, 128.1, 127.9, 123.9, 123.6, 52.5, 50.1, 32.2, 30.4, 29.5, 29.4, 26.6, 23.1, 15.0 ppm. elemental analysis calcd (%) for $\text{C}_{42}\text{H}_{54}\text{N}_4\text{O}_4$ (679): C 74.30, H 8.02, N 8.25, found: C 74.31, H 8.00, N 8.23.

3,3'-Di-n-octyl-1,1'-(1,2-phenylenedimethylene)diimidazolium 2,6-naphthalendicarboxylate: White solid; m.p. 196-205 °C; yield: 95%; ^1H NMR (300 MHz, $[\text{D}_6]\text{DMSO}$): δ = 9.76 (s; 2H), 8.32 (s; 2H), 7.96 (d; J = 8.1 Hz; 2H), 7.82 (d; J = 6.6 Hz; 4H), 7.76 (d; J = 7.8 Hz; 2H), 7.45 (m; 2H), 7.35 (m; 2H), 5.74 (s; 4H), 4.16 (t; J = 6.6 Hz; 4H), 1.77 (m; 4H), 1.19 (m; 20H), 0.82 (t; J = 6.3 Hz; 6H) ppm; ^{13}C NMR (300 MHz, $[\text{D}_6]\text{DMSO}$): δ = 169.6, 138.8, 136.0, 133.5, 133.3, 130.1, 128.2, 127.4, 123.2, 49.5, 49.4, 31.6, 29.7, 28.9, 28.7, 26.0, 22.5, 14.4 ppm. elemental analysis calcd (%) for $\text{C}_{42}\text{H}_{54}\text{N}_4\text{O}_4$ (679): C 74.30, H 8.02, N 8.25, found: C 74.27, H 7.99, N 8.24.

3,3'-Di-n-octyl-1,1'-(1,4-phenylenedimethylene)diimidazolium adipate: Hygroscopic white solid; yield: 95%; ^1H NMR (300 MHz, $[\text{D}_6]\text{DMSO}$): δ = 10.41 (s, 2H), 7.84 (s, 2H), 7.68 (s, 2H), 7.45 (s, 4H), 5.36 (s, 4H), 4.09 (t, J = 7.2 Hz, 4H), 1.69 (m, 8H), 1.28 (m, 4H), 1.14 (m, 20H), 0.75 (t, J = 6.3 Hz, 6H) ppm; ^{13}C NMR (300 MHz, $[\text{D}_6]\text{DMSO}$): δ = 176.3, 137.5, 135.8, 129.2, 122.7, 122.6, 51.5, 49.0, 31.3, 29.4, 28.6, 28.4, 27.2, 25.7, 22.2, 14.1. elemental analysis calcd (%) for $\text{C}_{36}\text{H}_{56}\text{N}_4\text{O}_4$ (609): C 71.02, H 9.27, N 9.20; found: C 70.99, H 9.30, N 9.21.

3,3'-Di-n-octyl-1,1'-(1,4-phenylenedimethylene)diimidazolium suberate: Hygroscopic white solid; yield: 92%; ^1H NMR (300 MHz, $[\text{D}_6]\text{DMSO}$): δ = 10.43 (s, 2H), 7.95 (d, J = 1.5 Hz, 2H), 7.85 (s, 2H), 7.57 (s, 4H), 5.55 (s, 4H), 4.23 (t, J = 7.2 Hz, 4H), 1.85 (m, 8H), 1.43 (t, J = 6.9 Hz, 8H), 1.28 (m, 20H), 0.8 (t, J = 7.2 Hz, 6H) ppm; ^{13}C NMR (300 MHz, $[\text{D}_6]\text{DMSO}$): δ = 176.2, 137.7, 135.9, 129.1, 122.8, 122.5, 51.4, 49.0, 31.3, 29.9, 29.5, 28.6, 28.4, 27.0, 25.7, 22.2, 14.1. elemental analysis calcd (%) for $\text{C}_{38}\text{H}_{60}\text{N}_4\text{O}_4$ (637): C 71.66, H 9.50, N 8.80; found: C 71.68, H 9.48, N 8.78.

3,3'-Di-n-decyl-1,1'-(1,4-phenylenedimethylene)diimidazolium ditetrafluoroborate: White solid; m.p. 113-116 °C; yield: 98%; ¹H NMR (400 MHz, [D6]DMSO): δ= 9.29 (s, 2H), 7.78 (dt, J= 1,6 Hz, 9,2 Hz, 4H), 7.45 (s, 4H), 5.41 (s, 4H), 4.15(t, J= 8.0 Hz, 4H), 1.77 (qt, J= 8.0 Hz, 4H), 1.23 (m, 28H), 0.85 (t, J= 8.0 Hz, 6H) ppm; ¹³C NMR (300 MHz, [D6]DMSO): δ= 136.7, 135.9, 129.4, 123.3, 123.0, 51.9, 49.4, 31.7, 29.7, 29.4, 29.3, 29.1, 28.8, 26.0, 22.5, 14.4 ppm. elemental analysis calcd (%) for C₃₄H₅₆B₂F₈N₄ (694): C 58.80, H 8.13, N 8.07, found: C 58.78, H 8.14, N 8.08.

3,3'-Di-n-decyl-1,1'-(1,3-phenylenedimethylene)diimidazolium ditetrafluoroborate: orange wax-like solid; m.p. 55-59 °C; yield: 97%; ¹H NMR (300 MHz, [D6]DMSO): δ= 9.35 (s, 2H), 7.81 (m, 4H), 7.52 (s, 1H), 7.41 (m, 3H), 5.43 (s, 4H), 4.17 (t, J= 7.2 Hz, 4H), 1.79 (qt, J= 6.7 Hz, 4H), 1.24 (m, 28H), 0.85 (t, J= 6.6 Hz, 6H) ppm; ¹³C NMR (300 MHz, [D6]DMSO): δ= 136.7, 136.0, 130.2, 129.0, 128.8, 123.3, 123.0, 52.1, 49.4, 31.7, 29.7, 29.3, 29.3, 29.1, 28.8, 26.0, 22.5, 14.4 ppm. elemental analysis calcd (%) for C₃₄H₅₆B₂F₈N₄ (694): C 58.80, H 8.13, N 8.07, found: C 58.77, H 8.16, N 8.05.

3,3'-Di-n-decyl-1,1'-(1,2-phenylenedimethylene)diimidazolium ditetrafluoroborate: brown oil; yield: 94%; ¹H NMR (300 MHz, [D6]DMSO): δ= 9.20 (s, 2H), 7.85 (s, 2H), 7.71 (s, 2H), 7.48 (m, 2H), 7.29 (m, 2H), 5.58 (s, 4H), 4.16 (t, J= 7.2 Hz, 4H), 1.79 (qt, J= 6.5 Hz, 4H), 1.24 (m, 28H), 0.85 (t, J= 6.7 Hz, 6H) ppm; ¹³C NMR (300 MHz, [D6]DMSO): δ= 136.9, 133.3, 130.1, 130.1, 123.3, 123.2, 49.5, 31.7, 29.8, 29.4, 29.3, 29.1, 28.8, 26.0, 22.5, 14.4 ppm. elemental analysis calcd (%) for C₃₄H₅₆B₂F₈N₄ (694): C 58.80, H 8.13, N 8.07, found: C 58.81, H 8.14, N 8.08.

3,3'-Di-n-decyl-1,1'-(1,4-phenylenedimethylene)diimidazolium D-Tartrate: Hygroscopic white solid; m.p. 102-105 °C; yield: 99%; ¹H NMR (400 MHz, [D6]DMSO): δ= 9.56 (s, 2H), 7.82 (m, 4H), 7.48 (s, 4H), 5.45 (s, 4H), 4.17 (t, J= 8.0 Hz, 4H), 3.72 (s, 2H), 1.77 (qt, J= 8.0 Hz, 4H), 1.23 (m, 28H), 0.85 (t, J= 8.0 Hz, 6H) ppm; ¹³C NMR (300 MHz, [D6]DMSO): δ= 175.4, 136.0, 129.4, 123.2, 122.9, 72.4, 51.9, 49.4, 31.7, 29.8, 29.3, 29.3, 29.1, 28.8, 26.0, 22.5, 14.4 ppm; elemental analysis calcd (%) for C₃₈H₆₀N₄O₆ (668): C 68.23, H 9.04, N 8.38, found: C 68.25, H 9.02, N 8.40. $[\alpha]_{24.1}^{589nm} = -5.0^\circ$.

3,3'-Di-n-decyl-1,1'-(1,4-phenylenedimethylene)diimidazolium 2,6-naphthalendicarboxylate: White solid; m.p. 223-228 °C; yield: 98%; ¹H NMR (400 MHz, [D6]DMSO): δ= 9.61 (s, 2H), 8.42 (s, 2H), 7.98 (d, J= 8.0 Hz, 2H), 7.88 (d, J= 8.0 Hz, 2H), 7.81 (m, 4H), 7.45 (s, 4H), 5.44 (s, 4H), 4.16 (t, J= 8.0 Hz, 4H), 1.76 (qt, J= 8.0 Hz, 4H), 1.24 (m, 28H), 0.83 (t, J= 8.0 Hz, 6H) ppm. ¹³C NMR

(300 MHz, [D6]DMSO): δ = 168.98, 136.94, 135.93, 135.42, 133.34, 129.35, 127.36, 123.31, 122.96, 51.88, 49.41, 48.93, 31.73, 29.72, 29.27, 29.11, 28.78, 25.96, 22.55, 14.42 ppm; elemental analysis calcd (%) for C₄₆H₆₂N₄O₄ (734): C 75.17, H 8.50, N 7.62, found: C 75.19, H 8.51, N 7.65.

Preparation of solution for biological tests

Samples for a typical biological test were prepared by weighing in a screw capped vial the proper amount of desired salt. Each salt was firstly dried under vacuum at 60 °C for at least 2 hours. After drying the sample into the vials for additional two hours. the salts were dissolved in DMSO (1 mL). The concentration of each solution was chosen depending on the salt solubility.

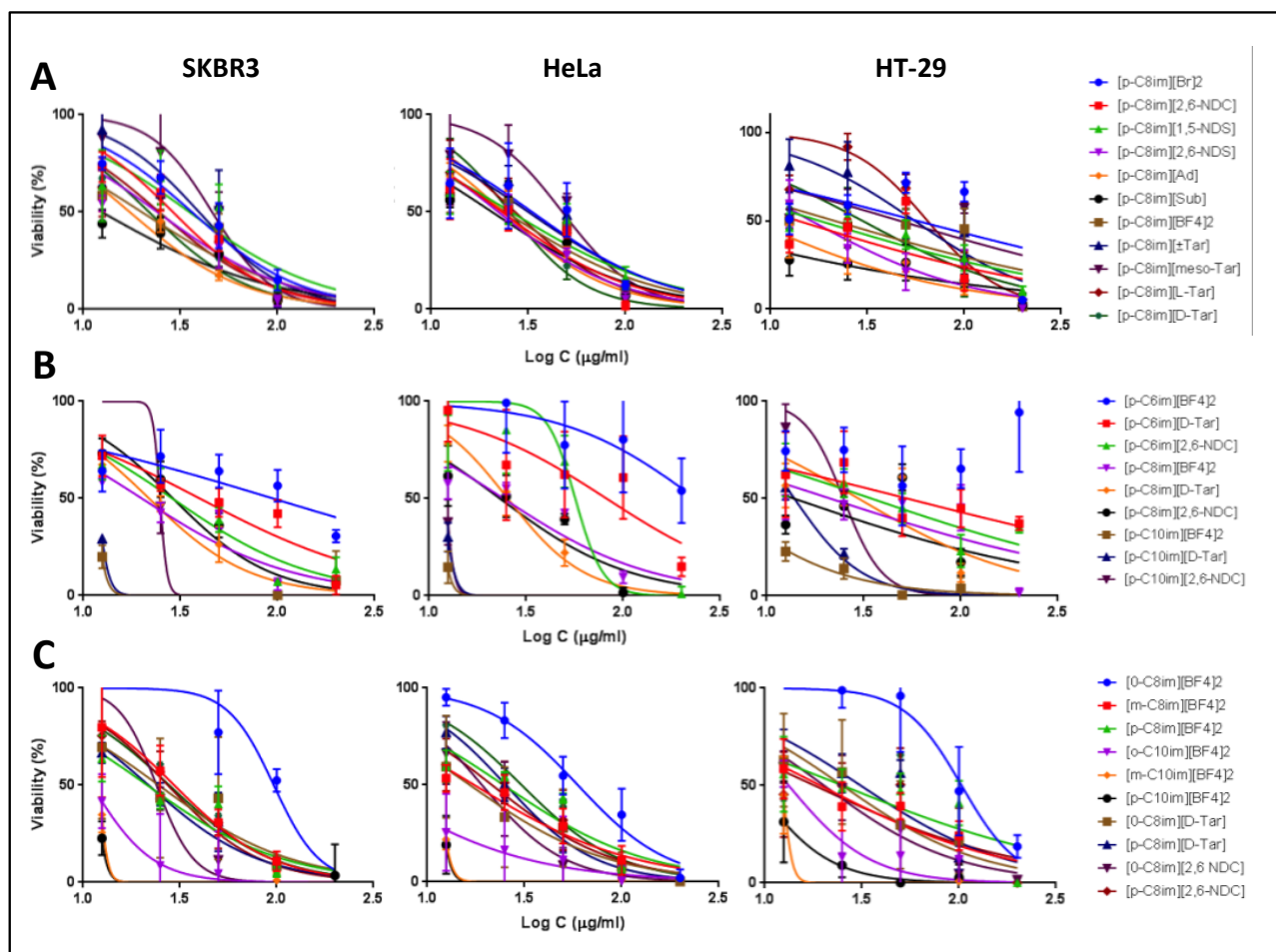


Figure S1. Dose–response curves of DOSs in cancer cell lines SKBR3, HeLa and HT-29 as function of different anion nature (A), alkyl chain length on the imidazolium ion (B) and isomeric substitution (C).

Microbiological assays

Antimicrobial activity was preliminarily evaluated using the diffusion agar technique.

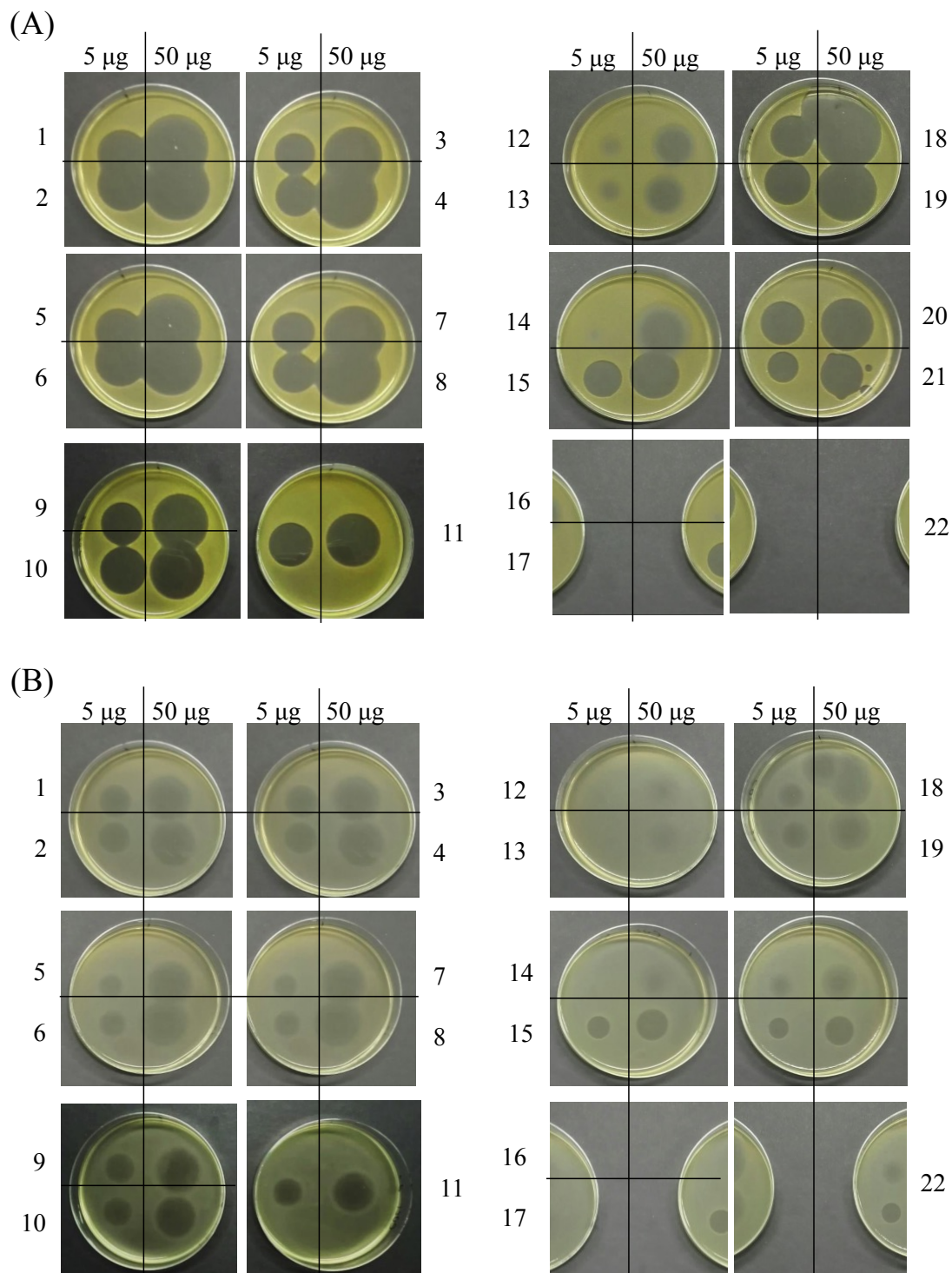


Figure S2. Representative agar diffusion tests performed using *K rhizophila* (A) and *E. coli* (B) as tester strains. DOS amounts were directly spotted on bacterial tester overlay and bacterial growth inhibition halos were observed with the higher DOS amount, the larger halo diameter. Numbers in (A) and (B) refer to : 1) [p-C8im][Br]2; 2) [p-C8im][2,6-NDC]; 3) [p-C8im][1,5-NDS]; 4) [p-C8im][2,6-NDS]; 5)[p-C8im][Ad]; 6) [p-C8im][Sub]; 7) [p-C8im][BF4]2; 8) [p-C8im][$\pm$ Tar]; 9) [p-C8im][meso-Tar]; 10)[p-C8im][L-Tar]; 11)[p-C8im][D-Tar]; 12) [p-C6im][BF4]2; 13) [p-C6im][D-Tar]; 14) [p-C6im][2,6-NDC]; 15) [p-C10im][BF4]2; 16) [p-C10im][D-Tar]; 17)[p-C10im][2,6-NDC]; 18) [o-C8im][D-Tar]; 19) [o-xil-(oim)2][2,6-NDC]; 20) [m-C8im][BF4]2; 21) [m-C10im][BF4]2; 22) [o-C10im][BF4]2.

In order to evaluate MIC₉₀ values, the bacterial tester strains were inoculated at the concentration of 10⁶ cell/ml and incubated for 24 h in presence of different concentration of the different DOSs. Then the bacterial growth was comparatively evaluated using parallel untreated bacterial cultivations as control condition and the growth was reported as percentage values of optical density (OD) in the respect of control. The MIC₉₀ was determined as the minimal DOS concentration among those tested causing more than 90% of bacterial growth reduction in terms OD. Mean values from at least three biological replicates are reported with standard deviation bars. Statistical test (one way ANOVA) was performed to asses significance (P<0.05) of spectrophotometric measurements.

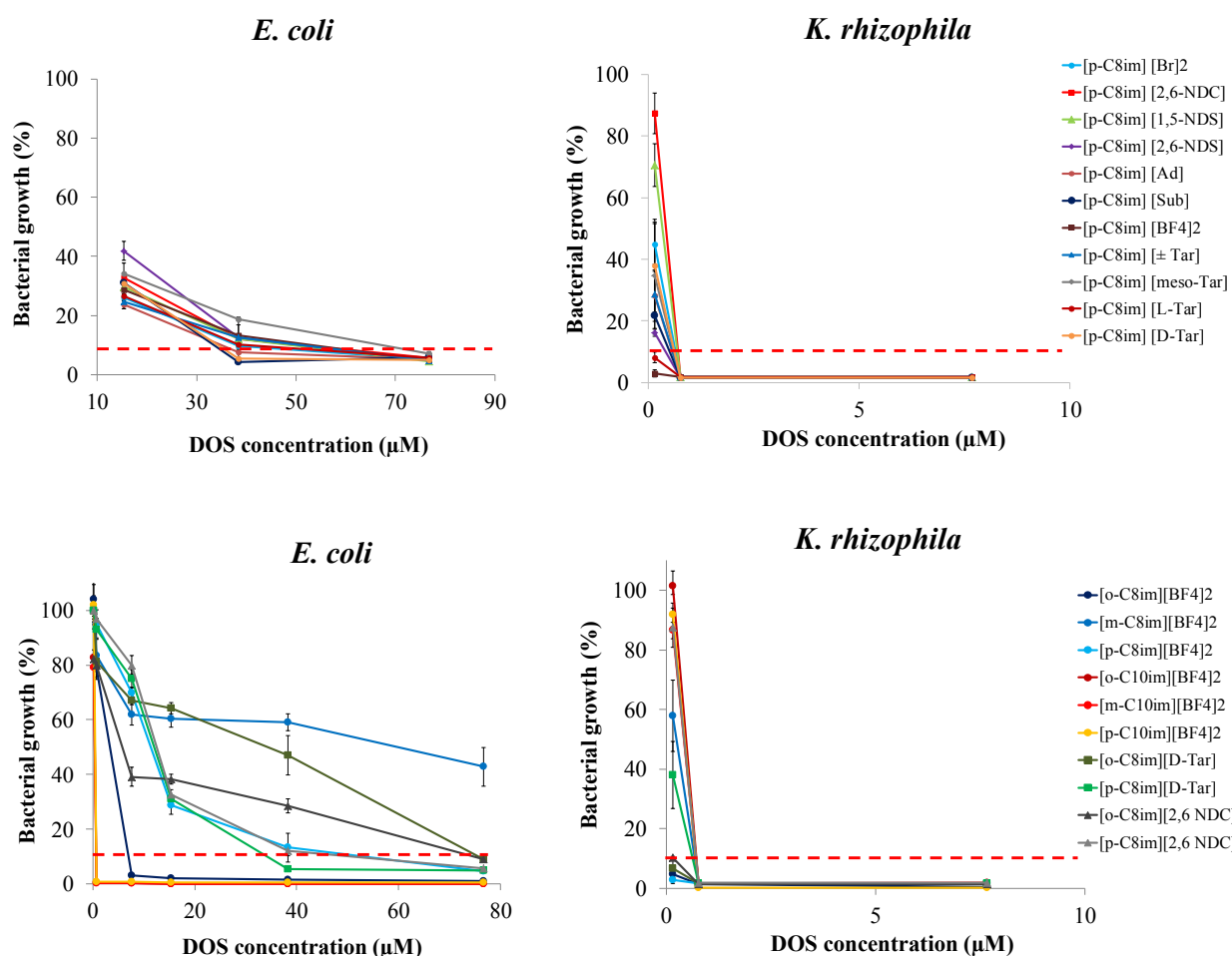


Figure S3. Examples of MIC₉₀ values determination of bacterial cell growth using spectrophotometric measurement at 600 nm. Bacterial growth values are reported as OD percentage in the respect of unexposed cultivations (control condition) and were calculated as mean of three independent biological replicates. Vertical bars represent standard deviation. Dashed red line represent percentage threshold for MIC₉₀ evaluation.

Table S1. IC₅₀ values at 24h of the 23 synthesized DOSs in cancer cell lines SKBR3, HeLa and HT-29.

	IC ₅₀ (μM)±SD		
	SKBR3	HeLa	HT-29
[<i>p</i> -C ₈ im][Br] ₂	72.7±12.0	66.4±19.2	113.9±48.1
[<i>p</i> -C ₈ im][2,6-NDC]	53.4±9.8	42.8±10.3	25.6±14.7
[<i>p</i> -C ₈ im][1,5-NDS]	58.7±22.2	43.0±14.6	29.8±12.0
[<i>p</i> -C ₈ im][2,6-NDS]	40.2±12.0	37.0±4.7	27.1±7.3
[<i>p</i> -C ₈ im][Ad]	35.8±9.8	48.4±6.6	16.3±4.1
[<i>p</i> -C ₈ im][Sub]	24.6±6.3	42.4±7.9	23.6±7.6
[<i>p</i> -C ₈ im][BF ₄] ₂	39.3±8.6	47.6±13.3	41.9±11.8
[<i>p</i> -C ₈ im][±Tar]	80.5±20.4	64.6±13.9	113.1±26.1
[<i>p</i> -C ₈ im][meso-Tar]	94.0±16.3	96.5±19.6	94.6±32.6
[<i>p</i> -C ₈ im][L-Tar]	50.2±10.6	55.3±11.4	131.5±24.5
[<i>p</i> -C ₈ im][D-Tar]	44.6±8.2	52.0±5.7	60.9±12.6
[<i>p</i> -C ₆ im][BF ₄] ₂	>429.5	>429.6	>429.6
[<i>p</i> -C ₆ im][D-Tar]	179.5±44.8	184.5±90.67	119.4±46.7
[<i>p</i> -C ₆ im][2,6-NDC]	64.2±16.9	113.7±24.1	69.7±25.7
[<i>p</i> -C ₁₀ im][BF ₄] ₂	14.4±7.2	14.4±7.2	9.4±3.6
[<i>p</i> -C ₁₀ im][D-Tar]	15.0±7.5	15.0±7.5	14.9±7.5
[<i>p</i> -C ₁₀ im][2,6-NDC]	13.6±6.8	13.6±6.8	13.6±6.8
[<i>o</i> -C ₈ im][BF ₄] ₂	156.7±40.7	116.5±27.4	202.4±45.5
[<i>o</i> -C ₈ im][D-Tar]	73.3±17.1	55.5±10.6	77.2±16.3
[<i>o</i> -C ₈ im][2,6-NDC]	44.3±7.4	33.1±7.4	38.7±8.8
[<i>m</i> -C ₈ im][BF ₄] ₂	60.3±8.6	36.0±7.1	53.9±9.4
[<i>m</i> -C ₁₀ im][BF ₄] ₂	14.4±7.2	14.4±7.2	14.4±7.2
[<i>o</i> -C ₁₀ im][BF ₄] ₂	14.4±7.2	14.4±7.2	14.4±7.2
Doxorubicin	4.1±0.5	1.5±0.3	1.5±0.1

Table S2. IC₅₀ values calculated at 24 and 48h of [*p*-C₈im][BF₄]₂ [*m*-C₈im][BF₄]₂ [*o*-C₈im][BF₄]₂ non tumoral epithelial mammary cell line (HB2) and tumoral mammary cell line (SKBR3)

Compound	IC ₅₀ (μM) against HB2		IC ₅₀ (μM) against SKBR3		(IC ₅₀ ^{HB2} / IC ₅₀ ^{SKBR3})	
	24h	48h	24h	48h	24h	48h
[<i>p</i> -C ₈ im][BF ₄] ₂	82.3±7.5	137.3±11.3	39.3±8.6	15.7±2.5	2.1	8.8
[<i>m</i> -C ₈ im][BF ₄] ₂	117.6±3.7	172.4±12.7	60.3±8.6	29.6±5.0	2.0	5.8
[<i>o</i> -C ₈ im][BF ₄] ₂	>391.9	>391.9	156.7±40.8	109.7±12.6	>2.5	>3.6
Doxorubicin	15.8±2.1	5.3±1.0	4.1±0.5	1.1±0.1	3.8	4.7

Table S3. MIC₉₀ values of DOSs against different bacterial tester strains.

DOS	Gram-negative <i>E. coli</i>	Gram-positive		
	<i>E. coli</i>	<i>K. rhizophila</i>	<i>S. aureus</i>	<i>B. subtilis</i>
DOS	μM (μg/mL)	μM (μg/mL)	μM (μg/mL)	μM (μg/mL)
[<i>p</i> -C ₈ im][Br] ₂	40 (25)	0.8 (0.5)	*	40 (25)
[<i>p</i> -C ₈ im][2,6-NDC]	74 (50)	0.7 (0.5)	*	15 (10)
[<i>p</i> -C ₈ im][1,5-NDS]	66 (50)	0.7 (0.5)	*	33 (25)
[<i>p</i> -C ₈ im][2,6-NDS]	67 (50)	0.7 (0.5)	*	33 (25)
[<i>p</i> -C ₈ im][Ad]	41 (25)	0.8 (0.5)	*	16 (10)
[<i>p</i> -C ₈ im][Sub]	39 (25)	0.8 (0.5)	*	16 (10)
[<i>p</i> -C ₈ im][BF ₄] ₂	78 (50)	0.2 (0.1)	*	39 (25)
[<i>p</i> -C ₈ im][±Tar]	82 (50)	0.8 (0.5)	*	41 (25)
[<i>p</i> -C ₈ im][<i>meso</i> -Tar]	82 (50)	0.8 (0.5)	*	41 (25)
[<i>p</i> -C ₈ im][L-Tar]	41 (25)	0.2 (0.1)	*	16 (10)
[<i>p</i> -C ₈ im][D-Tar]	41 (25)	0.8 (0.5)	*	16 (10)
[<i>p</i> -C ₆ im][BF ₄] ₂	*	86 (50)	*	*
[<i>p</i> -C ₆ im][D-Tar]	*	45 (25)	*	*
[<i>p</i> -C ₆ im][2,6-NDC]	*	161 (100)	*	160 (100)
[<i>p</i> -C ₁₀ im][BF ₄] ₂	0.7 (0.5)	0.7 (0.5)	72 (50)	0.7 (0.5)
[<i>p</i> -C ₁₀ im][D-Tar]	7 (5)	0.7 (0.5)	75 (50)	0.7 (0.5)
[<i>p</i> -C ₁₀ im][2,6-NDC]	7 (5)	0.7 (0.5)	68 (50)	0.1 (0.1)
[<i>o</i> -C ₈ im][BF ₄] ₂	8 (5)	0.2 (0.1)	*	39 (25)
[<i>o</i> -C ₈ im][D-Tar]	81 (50)	0.2 (0.1)	*	8 (5)
[<i>o</i> -C ₈ im][2,6-NDC]	74 (50)	1 (0.1)	*	7 (5)
[<i>m</i> -C ₈ im][BF ₄] ₂	157 (100)	0.8 (0.5)	*	8(5)
[<i>m</i> -C ₁₀ im][BF ₄] ₂	0.7 (0.5)	0.7 (0.5)	36 (25)	7 (5)
[<i>o</i> -C ₁₀ im][BF ₄] ₂	0.7 (0.5)	0.7 (0.5)	14 (10)	7 (5)

* No bacterial growth inhibition at the maximal concentration (100 μg/mL) tested.

Table S4. Selectivity index calculated for [*p*-C₈im][BF₄]₂, [*m*-C₈im][BF₄]₂ and [*o*-C₈im][BF₄]₂ as ratio between 24 h IC₅₀ values against HB2 cells and MIC₉₀ values against *E. coli*, *K. rhizophila* and *B. subtilis*, respectively.

DOS	IC ₅₀ ^{HB2} / MIC ₉₀ ^{<i>E. coli</i>}	IC ₅₀ ^{HB2} / MIC ₉₀ ^{<i>K. rhizophila</i>}	IC ₅₀ ^{HB2} / MIC ₉₀ ^{<i>B. subtilis</i>}
[<i>p</i> -C ₈ im][BF ₄] ₂	1.05	411.5	2.11
[<i>m</i> -C ₈ im][BF ₄] ₂	0.75	147.00	14.7
[<i>o</i> -C ₈ im][BF ₄] ₂	48.99	1959.50	10.05