

Exploration of a structure-activity relationship for the 4(3*H*)-quinazolinone antibiotics

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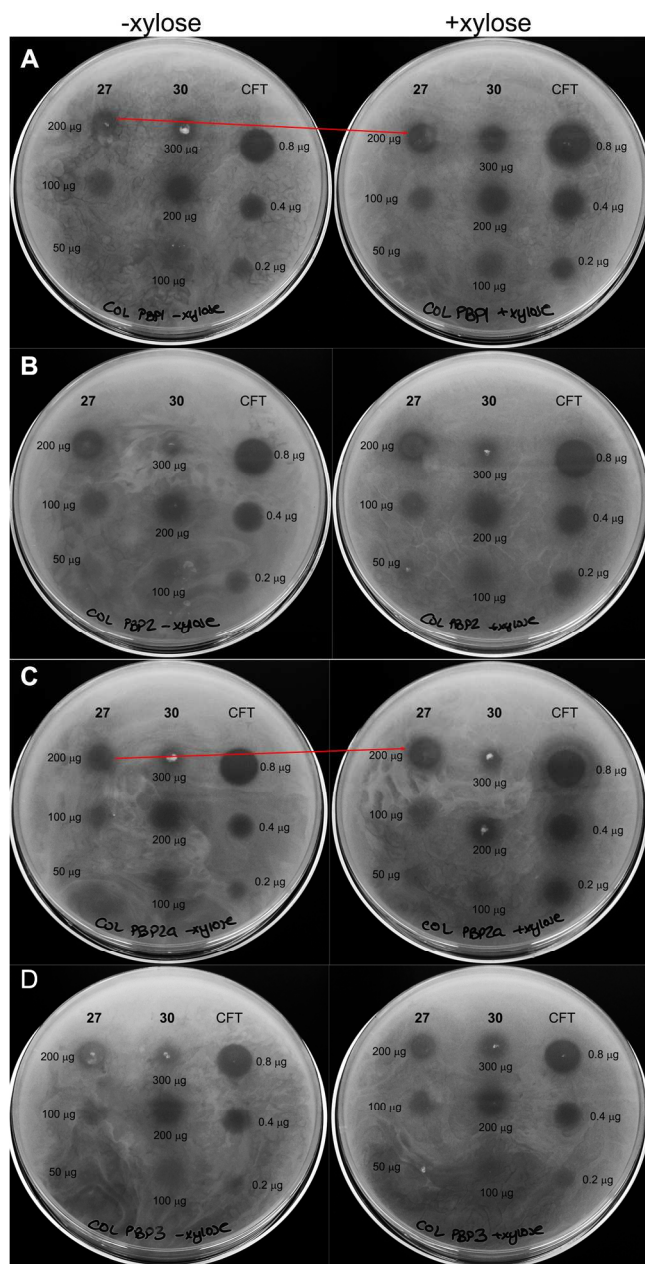
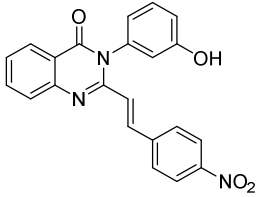
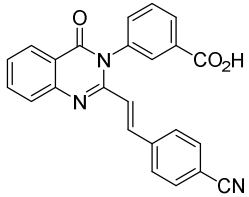
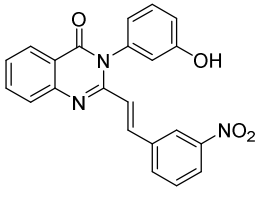
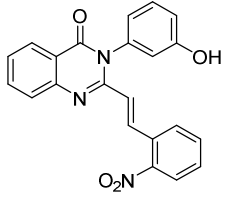
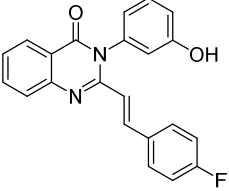
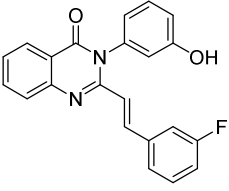
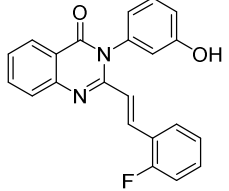
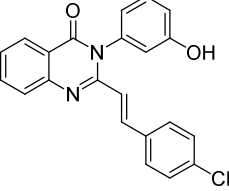
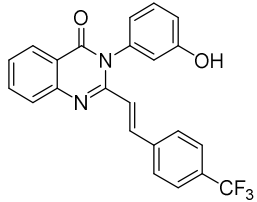
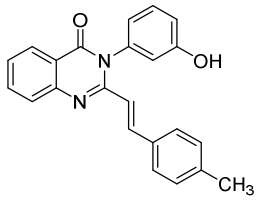
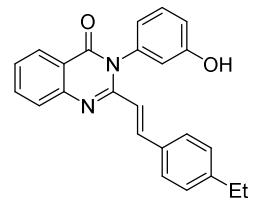
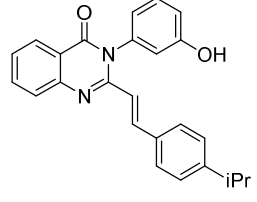


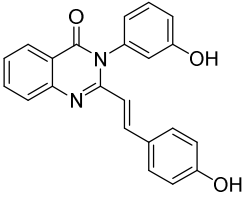
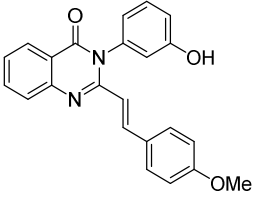
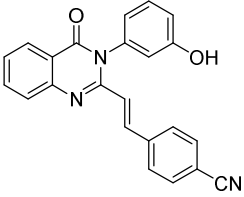
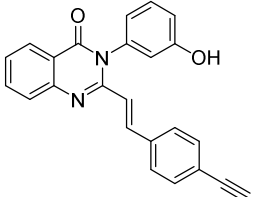
Figure S1. MRSA COL sensitization to antibacterials through antisense PBP knockdown. Left column shows background growth inhibition of the COL strain in the absence of xylose and right column shows activity in the presence of 50 mM xylose. (A) COL PBP1 antisense strain; (B) COL PBP2 antisense strain; (C) COL PBP2a antisense strain; (D) COL PBP3 antisense strain. Plates were spotted with compound **27** (left top to bottom: 200, 100, and 50 µg), compound **30** (middle top to bottom: 300, 200, and 100 µg), and ceftaroline (CFT; right top to bottom: 0.8, 0.4, and 0.2 µg). Precipitation at the highest concentration of compound **30** occurred upon spotting onto the plate. Red arrows highlight large changes in zones of inhibition in the presence of xylose for compound **27**.

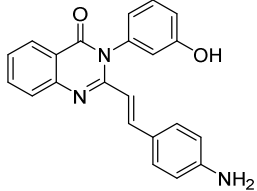
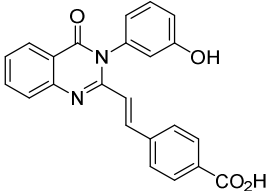
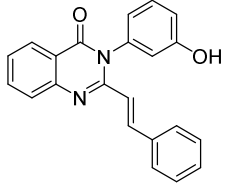
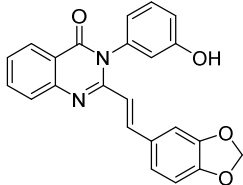
Table S1. Experimental procedures and characterization data of reported compounds

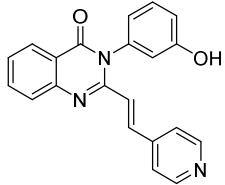
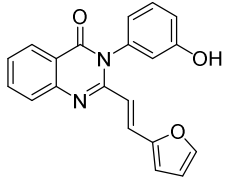
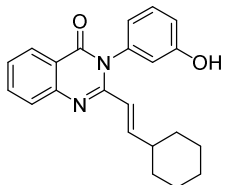
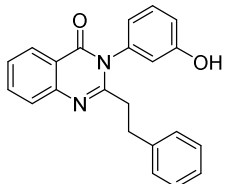
Structure	Experimental procedure and characterization data
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-nitrostyryl)quinazolin-4(3H)-one (1). This compound was prepared according to the procedure for 5 in 65% yield (0.12 g). ¹H NMR (500 MHz, DMSO-<i>d</i>₆) δ 6.56 (d, <i>J</i>=15.55 Hz, 1H), 6.86 (m, 2H), 6.98 (d, <i>J</i> = 7.78 Hz, 1H), 7.38 (t, <i>J</i> = 7.78 Hz, 1H), 7.55 (t, <i>J</i> = 7.38 Hz, 1H), 7.63 (d, <i>J</i> = 8.37 Hz, 2H), 7.77 (d, <i>J</i> = 8.17 Hz, 1H), 7.88 (t, <i>J</i> = 7.58 Hz, 1H), 7.94 (d, <i>J</i> = 15.55 Hz, 1H), 8.13 (d, <i>J</i> = 7.58 Hz, 1H), 8.18 (d, <i>J</i> = 8.57 Hz, 2H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 115.89, 116.50, 118.97, 120.82, 124.20, 126.48, 127.0, 127.31, 128.46, 130.35, 134.77, 136.10, 137.55, 141.28, 147.13, 147.47, 150.78, 158.68, 160.95. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₆N₃O₄, 386.1135; found, 386.1127.</p>
	<p>(E)-3-(3-carboxyphenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (2). The compound was prepared according to the procedure described previously (0.77 g, 75% yield). ¹H NMR (600 MHz, DMSO-<i>d</i>₆) δ 6.47 (d, <i>J</i> = 15.55 Hz, 1H), 7.56 (t, <i>J</i> = 7.34 Hz, 1H), 7.59 (d, <i>J</i> = 8.22 Hz, 2H), 7.73 (m, 2H), 7.79 (m, 3H), 7.90 (m, 2H), 8.05 (s, 1H), 8.12 (m, 2H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 111.56, 118.61, 120.76, 123.42, 126.50, 127.01, 127.35, 128.26, 129.99, 130.06, 130.12, 132.33, 132.83, 133.46, 134.89, 136.95, 137.03, 139.25, 147.21, 150.74, 161.25, 166.52. HRMS (<i>m/z</i>): [M + Na]⁺, calcd for C₂₄H₁₅N₃NaO₃, 416.1006; found, 416.0987. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₄H₁₆N₃O₃, 394.1186; found, 394.1214.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(3-nitrostyryl)quinazolin-4(3H)-one (3). This compound was prepared according to the procedure for 5 in 79% yield (0.18 g). ¹H NMR (500 MHz, DMSO-<i>d</i>₆) δ 5.62 (d, <i>J</i> = 15.55 Hz, 1H), 5.98 (m, 2H), 6.08 (d, <i>J</i> = 6.98 Hz, 1H), 6.51 (t, <i>J</i> = 7.58 Hz, 1H), 6.65 (t, <i>J</i> = 6.58 Hz, 1H), 6.75 (t, <i>J</i> = 7.58 Hz, 1H), 6.67 (d, <i>J</i> = 7.58 Hz, 1H), 6.91 (d, <i>J</i> = 7.18 Hz, 1H), 6.99 (t, <i>J</i> = 6.98 Hz, 1H), 7.06 (d, <i>J</i> = 15.55 Hz, 1H), 7.26 (m, 3H), 9.06 (br. s, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 115.90, 116.39, 119.17, 120.79, 121.94, 122.88, 123.92, 126.50, 126.91, 127.26, 130.36, 130.59, 133.29, 134.80, 136.23, 136.68, 137.64, 147.20, 148.28, 150.91, 158.48, 161.00. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₆N₃O₄, 386.1135; found, 386.1138.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(2-nitrostyryl)quinazolin-4(3H)-one (4). This compound was purchased from ChemDiv.</p>

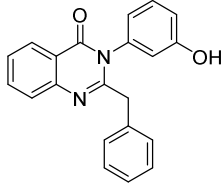
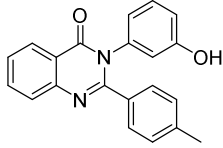
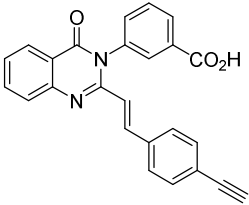
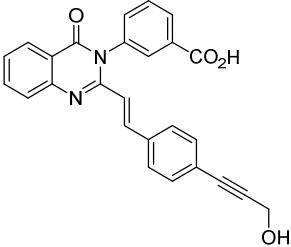
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-fluorostyryl)quinazolin-4(3H)-one (5). 3-(3-hydroxyphenyl)-2-methylquinazolin-4(3H)-one (0.25 g, 1 mmol) was suspended in 5 mL glacial acetic acid and dissolved upon heating, to which 4-fluorobenzaldehyde (0.11 mL, 1 mmol) was added. The reaction was refluxed overnight (18 h) and 10 mL water was added to the cooled reaction mixture. The resulting precipitate was filtered and washed with water, followed by cold ethanol and hexanes to give the product (0.14 g, 40% yield). ¹H NMR (500 MHz, DMSO-<i>d</i>₆) δ 6.32 (d, <i>J</i> = 15.55 Hz, 1H), 6.83 (s, 1H), 6.84 (d, <i>J</i> = 7.78 Hz, 1H), 6.96 (d, <i>J</i> = 8.17 Hz, 1H), 7.22 (m, 2H), 7.40 (t, <i>J</i> = 7.98 Hz, 1H), 7.44 (m, 2H), 7.53 (t, <i>J</i> = 8.17 Hz, 1H), 7.75 (d, <i>J</i> = 8.17 Hz, 1H), 7.87 (m, 2H), 8.12 (d, <i>J</i> = 7.78 Hz, 1H), 9.92 (s, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 116.55, 116.75, 116.92, 117.00, 120.00, 120.57, 121.35, 127.18, 127.18, 127.85, 130.35, 130.41, 131.11, 132.25, 135.45, 138.15, 138.56, 148.05, 151.98, 159.03, 161.80, 162.52. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₆FN₂O₂, 359.1190; found, 359.1186.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(3-fluorostyryl)quinazolin-4(3H)-one (6). This compound was prepared according to the procedure for 5 in 12% yield (0.086 g). ¹H NMR (500 MHz, DMSO-<i>d</i>₆) δ 6.42 (d, <i>J</i> = 15.55 Hz, 1H), 6.83 (s, 1H), 6.84 (d, <i>J</i> = 7.58 Hz, 1H), 6.97 (d, <i>J</i> = 7.98 Hz, 1H), 7.20 (m, 2H), 7.25 (d, <i>J</i> = 8.57 Hz, 1H), 7.40 (m, 2H), 7.55 (t, <i>J</i> = 7.18 Hz, 1H), 7.76 (d, <i>J</i> = 7.98 Hz, 1H), 7.84 (d, <i>J</i> = 15.75 Hz, 1H), 7.88 (t, <i>J</i> = 8.37 Hz, 1H), 8.13 (d, <i>J</i> = 7.98 Hz, 1H), 9.92 (s, 1H). HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₆FN₂O₂, 359.1190; found, 359.1211.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(2-fluorostyryl)quinazolin-4(3H)-one (7). This compound was purchased from ChemDiv.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(3-fluorostyryl)quinazolin-4(3H)-one (8). This compound was prepared according to the procedure for 5 in 66% yield (0.29 g). ¹H NMR (500 MHz, DMSO-<i>d</i>₆) δ 6.38 (d, <i>J</i> = 15.55 Hz, 1H), 6.83 (m, 2H), 6.96 (d, <i>J</i> = 8.17 Hz, 1H), 7.41 (m, 5H), 7.54 (t, <i>J</i> = 8.17 Hz, 1H), 7.76 (d, <i>J</i> = 7.78 Hz, 1H), 7.85 (m, 2H), 8.12 (d, <i>J</i> = 7.78 Hz, 1H), 9.93 (s, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 116.01, 116.49, 119.46, 120.85, 120.91, 126.64, 126.85, 127.35, 129.31, 130.58, 133.97, 134.38, 134.93, 137.43, 137.97, 147.47, 151.33, 158.49, 161.24. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₆ClN₂O₂, 375.0895; found, 375.0892.</p>

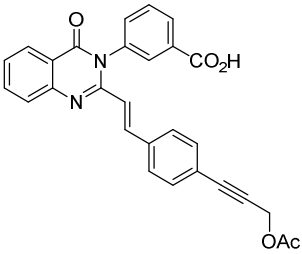
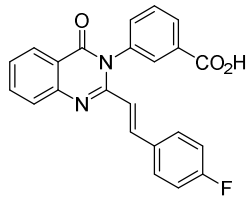
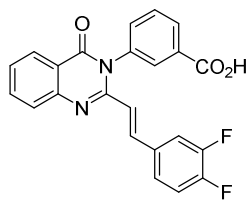
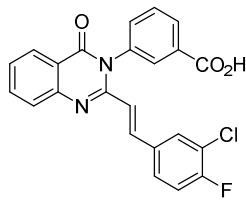
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-(trifluoromethyl)styryl)quinazolin-4(3H)-one (9). This compound was prepared according to the procedure for 5 in 54% yield (0.13 g). ^1H NMR (300 MHz, DMSO-d_6) δ 6.50 (d, J = 15.55 Hz, 1H), 6.81 (m, 2H), 6.93 (d, J = 8.13 Hz, 1H), 7.36 (t, J = 8.13 Hz, 1H), 7.57 (m, 3H), 7.71 (d, J = 8.37 Hz, 2H), 7.77 (d, J = 8.37 Hz, 1H), 7.89 (t, J = 7.18 Hz, 1H), 7.90 (d, J = 15.79 Hz, 1H), 8.12 (d, J = 8.13 Hz, 1H). ^{13}C NMR (126 MHz, DMSO-d_6) δ 115.88, 116.39, 119.20, 120.78, 122.78, 125.91, 126.48, 126.85, 127.26, 128.03, 130.41, 134.76, 136.81, 137.71, 138.84, 147.20, 150.92, 158.47, 161.02. HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{23}\text{H}_{16}\text{F}_3\text{N}_2\text{O}_2$, 409.1158; found, 409.1168.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-methylstyryl)quinazolin-4(3H)-one (10). This compound was prepared according to the procedure for 5 in 47% yield (0.13 g). ^1H NMR (500 MHz, DMSO-d_6) δ 2.29 (s, 3H), 6.31 (d, J = 15.55 Hz, 1H), 6.82 (m, 2H), 6.96 (d, J = 7.98 Hz, 1H), 7.19 (d, J = 7.78 Hz, 2H), 7.26 (d, J = 7.78 Hz, 2H), 7.40 (t, J = 7.98 Hz, 1H), 7.52 (t, J = 7.58 Hz, 1H), 7.75 (d, J = 8.17 Hz, 1H), 7.84 (m, 2H), 8.12 (d, J = 7.78 Hz, 1H), 9.92 (s, 1H). ^{13}C NMR (126 MHz, DMSO-d_6) δ 20.98, 115.84, 116.24, 118.90, 119.30, 120.57, 126.45, 127.12, 127.45, 129.69, 130.39, 132.17, 134.71, 137.93, 138.72, 139.72, 147.41, 151.46, 158.32, 161.13. HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{23}\text{H}_{19}\text{N}_2\text{O}_2$, 355.1441; found, 355.1469.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-ethylstyryl)quinazolin-4(3H)-one (11). This compound was prepared according to the procedure for 5 in 42% yield (0.31 g). ^1H NMR (500 MHz, DMSO-d_6) δ 1.14 (t, J = 7.18 Hz, 3H), 2.57 (d, J = 7.18 Hz, 2H), 6.32 (d, J = 15.55 Hz, 1H), 6.83 (m, 2H), 6.97 (d, J = 7.58 Hz, 1H), 7.20 (d, J = 7.38 Hz, 2H), 7.27 (d, J = 7.38 Hz, 2H), 7.40 (t, J = 7.98 Hz, 1H), 7.52 (t, J = 6.98 Hz, 1H), 7.75 (d, J = 7.58 Hz, 1H), 7.85 (m, 2H), 8.11 (d, J = 7.78 Hz, 1H), 9.92 (s, 1H). ^{13}C NMR (126 MHz, DMSO-d_6) δ 15.35, 28.04, 115.86, 116.25, 119.01, 119.30, 120.58, 126.44, 127.11, 127.52, 128.51, 130.40, 132.45, 134.69, 137.95, 138.71, 145.94, 147.31, 147.41, 158.36, 160.24, 161.13. HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{24}\text{H}_{21}\text{N}_2\text{O}_2$, 369.1598; found, 369.1613.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-isopropylstyryl)quinazolin-4(3H)-one (12). This compound was prepared according to the procedure for 5 in 32% yield (0.15 g). ^1H NMR (500 MHz, DMSO-d_6) δ 1.16 (d, J = 6.78 Hz, 6H), 2.86 (m, J = 6.98 Hz, 1H), 6.32 (d, J = 15.55 Hz, 1H), 6.82 (m, 2H), 6.97 (d, J = 8.17 Hz, 1H), 7.23 (d, J = 7.98 Hz, 2H), 7.28 (d, J = 8.17 Hz, 2H), 7.39 (t, J = 7.98 Hz, 1H), 7.52 (t, J = 7.38 Hz, 1H), 7.75 (d, J = 7.98 Hz, 1H), 7.84 (m, 2H), 8.12 (d, J = 7.78 Hz, 1H), 9.92 (s, 1H). ^{13}C NMR (126 MHz, DMSO-d_6) δ 23.58, 33.30, 115.85,</p>

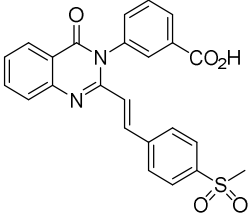
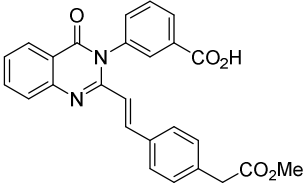
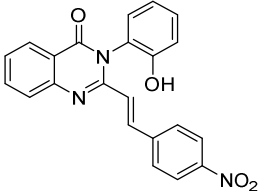
	116.22, 119.09, 119.28, 120.57, 126.44, 127.05, 127.10, 127.52, 130.37, 132.60, 134.69, 137.93, 138.65, 147.40, 160.49, 151.44, 158.32, 161.09. HRMS (m/z): $[M + Na]^+$, calcd for $C_{25}H_{22}N_2NaO_2$, 405.1573; found, 405.1599.
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-hydroxystyryl)quinazolin-4(3H)-one (13). This compound was prepared according to the procedure for 5 in 33% yield (0.096 g). 1H NMR (500 MHz, DMSO-d_6) δ 6.15 (d, $J = 15.55$ Hz, 1H), 6.76 (m, 4H), 6.96 (d, $J = 7.38$ Hz, 1H), 7.21 (d, $J = 7.58$ Hz, 2H), 7.39 (t, $J = 7.58$ Hz, 1H), 7.49 (t, $J = 7.18$ Hz, 1H), 7.72 (d, $J = 7.98$ Hz, 1H), 7.81 (m, 2H), 8.10 (d, $J = 7.78$ Hz, 1H), 9.91 (s, 1H), 9.95 (s, 1H). ^{13}C NMR (126 MHz, DMSO-d_6) δ 116.53, 116.68, 116.89, 120.01, 121.10, 126.65, 127.13, 127.68, 130.07, 131.09, 135.36, 138.75, 139.81, 148.26, 152.51, 159.03, 160.03, 161.89. HRMS (m/z): $[M + H]^+$, calcd for $C_{22}H_{17}N_2O_3$, 357.1234; found, 357.1256.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-methoxystyryl)quinazolin-4(3H)-one (14). This compound was prepared according to the procedure for 5 in 52% yield (0.15 g). 1H NMR (500 MHz, DMSO-d_6) δ 3.76 (s, 3H), 6.21 (d, $J = 15.55$ Hz, 1H), 6.82 (m, 2H), 6.93 (m, 3H), 7.32 (d, $J = 8.77$ Hz, 2H), 7.40 (t, $J = 7.98$ Hz, 1H), 7.51 (t, $J = 7.58$ Hz, 1H), 7.73 (d, $J = 8.17$ Hz, 1H), 7.84 (m, 2H), 8.11 (d, $J = 7.78$ Hz, 1H), 9.92 (s, 1H). ^{13}C NMR (126 MHz, DMSO-d_6) δ 55.29, 114.52, 114.59, 115.83, 116.21, 117.34, 119.30, 120.48, 126.28, 126.44, 127.03, 127.50, 129.14, 130.39, 134.69, 138.00, 138.55, 147.48, 151.63, 158.32, 160.63, 161.15. HRMS (m/z): $[M + H]^+$, calcd for $C_{23}H_{19}N_2O_3$, 371.1390; found, 371.1413.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (15). This compound was prepared according to the procedure for 5 in 84% yield (0.62 g). 1H NMR (500 MHz, DMSO-d_6) δ 6.51 (d, $J = 15.55$ Hz, 1H), 6.85 (m, 2H), 6.97 (d, $J = 7.98$ Hz, 1H), 7.39 (t, $J = 7.78$ Hz, 1H), 7.55 (m, 3H), 7.77 (d, $J = 7.78$ Hz, 1H), 7.81 (d, $J = 8.37$ Hz, 2H), 7.88 (t, $J = 7.18$ Hz, 1H), 7.89 (d, $J = 15.55$ Hz, 1H), 8.12 (d, $J = 7.98$ Hz, 1H), 9.94 (s, 1H). ^{13}C NMR (126 MHz, DMSO-d_6) δ 111.52, 115.87, 116.40, 118.61, 119.30, 120.81, 123.47, 126.50, 126.95, 127.29, 128.08, 130.42, 132.92, 134.80, 136.60, 137.65, 139.38, 147.17, 150.84, 158.32, 160.99. HRMS (m/z): $[M + H]^+$, calcd for $C_{23}H_{16}N_3O_2$, 366.1237; found, 366.1254.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-ethynylstyryl)quinazolin-4(3H)-one (16). This compound was prepared according to the procedure for 5 in 22% yield (0.33 g). 1H NMR (500 MHz, DMSO-d_6) δ 4.32 (s, 1H), 6.40 (d, $J = 15.55$ Hz, 1H), 6.85 (m, 2H), 6.97 (d, $J = 8.18$ Hz, 1H), 7.38 (m, 3H), 7.45 (d, $J = 7.17$ Hz, 2H), 7.53 (t, $J = 7.18$ Hz, 1H), 7.76 (d, $J = 7.98$ Hz, 1H),</p>

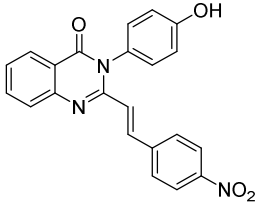
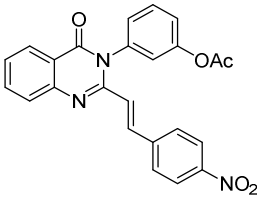
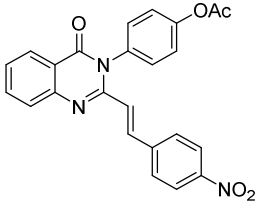
	7.86 (m, 2H), 8.12 (d, $J = 7.98$ Hz, 1H), 9.93 (s, 1H). ^{13}C NMR (126 MHz, DMSO- d_6) δ 82.69, 83.21, 115.86, 116.34, 119.30, 120.68, 121.08, 122.76, 126.46, 126.68, 127.20, 127.62, 130.41, 132.34, 134.72, 135.29, 137.59, 137.79, 147.28, 151.12, 158.32, 161.05. HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{24}\text{H}_{17}\text{N}_2\text{O}_2$, 365.1285; found, 365.1288.
	(E)-3-(3-hydroxyphenyl)-2-(4-aminostyryl)quinazolin-4(3H)-one (17). This compound was prepared according to the procedure for 37 from compound 1 (0.04 g, 0.1 mmol) in 82% yield (0.04 g). ^1H NMR (500 MHz, DMSO- d_6) δ 6.00 (d, $J = 15.35$ Hz, 1H), 6.42 (d, $J = 8.57$ Hz, 1H), 6.50 (d, $J = 8.37$ Hz, 2H), 6.71 (s, 1H), 7.04 (d, $J = 8.37$ Hz, 2H), 7.39 (t, $J = 7.78$ Hz, 1H), 7.45 (t, $J = 6.58$ Hz, 1H), 7.68 (d, $J = 8.17$ Hz, 1H), 7.79 (m, 3H), 8.07 (d, $J = 7.78$ Hz, 1H), 9.91 (s, 1H). HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{22}\text{H}_{18}\text{N}_3\text{O}_2$, 356.1394; found, 356.1392.
	(E)-3-(3-hydroxyphenyl)-2-(4-carboxystyryl)quinazolin-4(3H)-one (18). This compound was prepared according to the procedure for 5 in 83% yield (0.31 g). ^1H NMR (500 MHz, DMSO- d_6) δ 6.49 (d, $J = 15.55$ Hz, 1H), 6.84 (m, 2H), 6.97 (d, $J = 8.37$ Hz, 1H), 7.40 (t, $J = 7.98$ Hz, 1H), 7.49 (d, $J = 8.37$ Hz, 2H), 7.55 (t, $J = 7.98$ Hz, 1H), 7.78 (d, $J = 7.98$ Hz, 1H), 7.90 (m, 4H), 8.14 (d, $J = 7.98$ Hz, 1H), 9.93 (s, 1H). ^{13}C NMR (126 MHz, DMSO- d_6) δ 95.45, 115.86, 116.36, 119.30, 120.76, 122.25, 126.49, 126.83, 127.28, 127.56, 129.57, 129.93, 129.99, 130.44, 131.37, 134.79, 137.44, 137.74, 138.95, 147.26, 151.06, 158.32, 161.06, 166.76. HRMS (m/z): $[\text{M} + \text{Na}]^+$, calcd for $\text{C}_{23}\text{H}_{16}\text{N}_2\text{NaO}_4$, 407.1002; found, 407.1016.
	(E)-3-(3-hydroxyphenyl)-2-styrylquinazolin-4(3H)-one (19). This compound was prepared according to the procedure for 5 in 27% yield (0.64 g). ^1H NMR (500 MHz, DMSO- d_6) δ 5.53 (d, $J = 15.55$ Hz, 1H), 6.00 (m, 2H), 6.12 (d, $J = 8.17$ Hz, 1H), 6.51 (m, 6H), 6.67 (t, $J = 7.58$ Hz, 1H), 6.92 (d, $J = 8.17$ Hz, 1H), 7.02 (m, 2H), 7.27 (d, $J = 7.78$ Hz, 1H), 9.07 (s, 1H). ^{13}C NMR (126 MHz, DMSO- d_6) δ 115.84, 116.27, 119.29, 119.99, 120.63, 126.44, 126.55, 127.15, 127.42, 129.07, 129.77, 130.38, 134.70, 134.87, 137.88, 138.64, 147.34, 151.30, 158.33, 161.08. HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{22}\text{H}_{17}\text{N}_2\text{O}_2$, 341.1285; found, 341.1319.
	(E)-2-(2-(benzo[d][1,3]dioxol-5-yl)vinyl)-3-(3-hydroxyphenyl)quinazolin-4(3H)-one (20). This compound was purchased from ChemDiv.

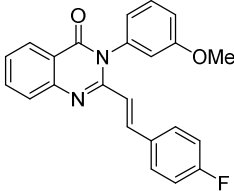
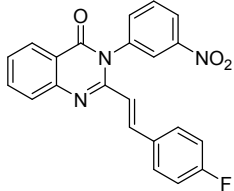
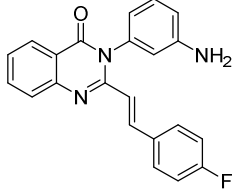
	<p>(E)-3-(3-hydroxyphenyl)-2-(2-(pyridine-4-yl)vinyl)quinazolin-4(3H)-one (21). This compound was prepared according to the procedure for 5 and purified by silica gel column chromatography to give the final product in 43% yield (0.18 g). ¹H NMR (500 MHz, DMSO-<i>d</i>₆) δ 6.60 (d, <i>J</i> = 15.55 Hz, 1H), 6.85 (m, 2H), 6.97 (d, <i>J</i> = 8.37 Hz, 1H), 7.33 (m, 2H), 7.40 (t, <i>J</i> = 7.18 Hz, 1H), 7.57 (t, <i>J</i> = 7.58 Hz, 1H), 7.80 (m, 2H), 7.90 (t, <i>J</i> = 7.38 Hz, 1H), 8.15 (d, <i>J</i> = 7.38 Hz, 1H), 8.56 (m, 2H), 9.94 (s, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 115.84, 115.93, 116.45, 119.33, 120.89, 121.38, 126.53, 127.05, 127.37, 130.44, 134.82, 135.92, 137.62, 141.93, 147.16, 150.50, 150.72, 158.37, 160.99. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₁H₁₆N₃O₂, 342.1237; found, 342.1251.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(2-(furan-2-yl)vinyl)quinazolin-4(3H)-one (22). This compound was prepared according to the procedure for 5 in 61% yield (0.16 g). ¹H NMR (500 MHz, DMSO-<i>d</i>₆) δ 6.17 (d, <i>J</i> = 15.35 Hz, 1H), 6.59 (s, 1H), 6.83 (m, 3H), 6.96 (d, <i>J</i> = 8.17 Hz, 1H), 7.39 (t, <i>J</i> = 7.98 Hz, 1H), 7.51 (t, <i>J</i> = 7.38 Hz, 1H), 7.75 (m, 3H), 7.86 (t, <i>J</i> = 8.17 Hz, 1H), 8.10 (d, <i>J</i> = 7.78 Hz, 1H), 9.93 (s, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 95.45, 112.71, 115.42, 115.82, 116.29, 116.70, 119.30, 120.51, 125.91, 126.40, 126.45, 127.01, 130.48, 134.73, 137.95, 145.49, 147.39, 150.99, 158.34, 161.12. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₀H₁₅N₂O₃, 331.1077; found, 331.1102.</p>
	<p>(E)-2-(2-cyclohexylvinyl)-3-(3-hydroxyphenyl)quinazolin-4(3H)-one (23). 3-(3-hydroxyphenyl)-2-methylquinazolin-4(3H)-one (0.38 g, 1.5 mmol) and cyclohexanecarboxaldehyde (0.37 g, 3.3 mmol) were dissolved in 250 μL pyridine and heated to 165 °C for 1 h. The reaction mixture was purified by silica gel column chromatography to give the product in 9% yield (0.05 g). ¹H (400 MHz, CDCl₃) δ 1.07 (m, 6H), 1.66 (s, br, 6H), 2.04 (s, br, 1H), 5.77 (d, <i>J</i> = 15.2 Hz, 1H), 6.60 (m, 2H), 6.83 (d, <i>J</i> = 7.2 Hz, 1H), 7.11 (d, <i>J</i> = 8.0 Hz, 1H), 7.32 (t, <i>J</i> = 7.2 Hz, 1H), 7.44 (s, br, 1H), 7.75 (s, br, 2H), 8.09 (s, 1H), 8.28 (d, <i>J</i> = 7.6 Hz, 1H). ¹³C NMR (100 MHz, CDCl₃) δ 25.8, 26.1, 32.1, 41.3, 116.1, 117.5, 119.4, 120.1, 120.4, 126.8, 127.3, 130.9, 135.1, 137.3, 150.7, 152.4, 158.8, 163.2. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₂₃N₂O₂, 347.1754; found, 347.1785.</p>
	<p>3-(3-hydroxyphenyl)-2-phenethylquinazolin-4(3H)-one (24). Compound 19 (0.6 g, 1.75 mmol) was dissolved in 250 mL of 50:50 EtOAc:EtOH to which Pd/C (0.12 g) was added. The flask was flushed with H₂ and the reaction stirred at 25 °C under H₂ positive pressure overnight (16 h). The reaction was filtered through celite, concentrated <i>in vacuo</i>, and purified by silica gel column chromatography. Recrystallization from hot ethanol was performed to obtain pure product in 29% yield (0.18 g). ¹H</p>

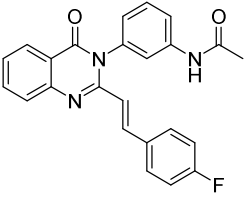
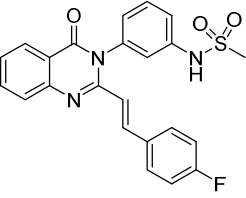
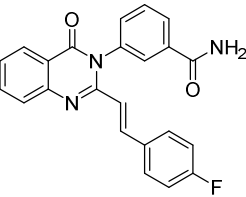
	<p>NMR (500 MHz, DMSO-<i>d</i>₆) δ 2.65 (t, <i>J</i> = 8.77 Hz, 2H), 2.99 (t, <i>J</i> = 8.57 Hz, 2H), 6.79 (m, 2H), 6.90 (d, <i>J</i> = 8.17 Hz, 1H), 7.06 (d, <i>J</i> = 6.98 Hz, 2H), 7.14 (t, <i>J</i> = 7.18 Hz, 1H), 7.22 (t, <i>J</i> = 7.58 Hz, 2H), 7.33 (t, <i>J</i> = 7.78 Hz, 1H), 7.53 (t, <i>J</i> = 7.18 Hz, 1H), 7.71 (d, <i>J</i> = 7.58 Hz, 1H), 7.86 (t, <i>J</i> = 7.18 Hz, 1H), 8.11 (d, <i>J</i> = 7.98 Hz, 1H), 9.85 (s, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 32.17, 37.03, 95.43, 115.61, 116.02, 118.97, 120.55, 126.02, 126.32, 126.54, 126.88, 128.18, 128.35, 130.21, 134.58, 138.13, 147.10, 156.01, 158.26, 161.20. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₉N₂O₂, 343.1441; found, 343.1454.</p>
	<p>2-benzyl-3-(3-hydroxyphenyl)quinazolin-4(3H)-one (25). This compound was purchased from ChemDiv.</p>
	<p>3-(3-hydroxyphenyl)-2-(4-tolyl)quinazolin-4(3H)-one (26). This compound was purchased from ChemDiv.</p>
	<p>(E)-3-(3-carboxyphenyl)-2-(4-ethynylstyryl)quinazolin-4(3H)-one (27). This compound was prepared according to the procedure for 2 and purified by silica gel column chromatography in 24% yield (0.05 g). ¹H NMR (600 MHz, CD₃OD) δ 3.60 (s, 1H), 6.38 (d, <i>J</i> = 15.55 Hz, 1H), 7.29 (d, <i>J</i> = 8.57 Hz, 2H), 7.39 (d, <i>J</i> = 8.17 Hz, 2H), 7.54 (t, <i>J</i> = 6.98 Hz, 1H), 7.64 (d, <i>J</i> = 7.98 Hz, 1H), 7.76 (t, <i>J</i> = 7.98 Hz, 1H), 7.82 (d, <i>J</i> = 8.17 Hz, 1H), 7.88 (m, 2H), 8.07 (s, 1H), 8.20 (d, <i>J</i> = 7.78 Hz, 1H), 8.25 (d, <i>J</i> = 7.78 Hz, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 82.70, 83.24, 120.66, 121.04, 122.84, 126.51, 126.81, 127.31, 127.83, 129.98, 130.12, 132.33, 132.39, 133.49, 134.90, 135.21, 137.21, 137.99, 147.36, 151.04, 161.37, 166.58. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₅H₁₇N₂O₃, 393.1234; found, 393.1250.</p>
	<p>(E)-3-(3-carboxyphenyl)-2-(4-(3-hydroxyprop-1-yn-1-yl)styryl)quinazolin-4(3H)-one (28). 1 M NaOH solution (aq, 0.5 mL) was added to a solution of 29 (100 mg, 0.25 mmol) in 0.5 mL ethanol and stirred for 2 h at room temperature. Water (3 mL) was added and the resulting solution acidified with 3 M HCl (aq). Extraction with dichloromethane followed and the organic layer dried over Na₂SO₄ and concentrated <i>in vacuo</i> to yield pure product in 73% yield (0.068 g). ¹H (400 MHz, DMSO-<i>d</i>₆) δ 4.29 (s, 2H), 6.34 (d, <i>J</i> = 15.6 Hz, 1H), 7.40 (m, 4H), 7.55 (ddd, <i>J</i> = 7.6, 7.2, 1.2 Hz, 1H), 7.74 (m, 2H), 7.78 (d, <i>J</i> = 8 Hz, 1H), 7.88 (d, <i>J</i> = 15.6 Hz, 1H), 7.89 (ddd, <i>J</i> = 8.0, 7.2, 1.6 Hz, 1H), 8.02 (m, 1H), 8.13 (m, 2H). ¹³C NMR (101 MHz,</p>

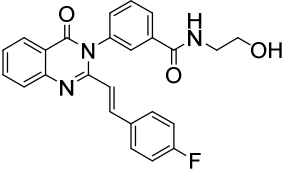
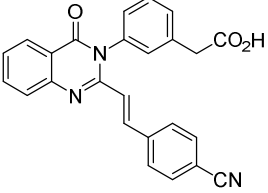
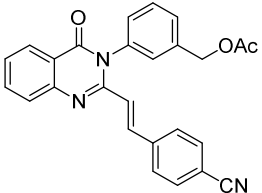
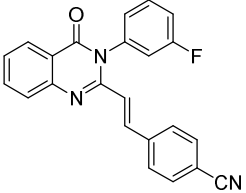
	DMSO- <i>d</i> ₆ δ 50.18, 84.06, 92.61, 121.32, 121.41, 124.33, 127.17, 127.45, 127.95, 128.52 (2C), 130.63, 130.77 (2C), 132.53 (2C), 133.09, 134.15, 135.41, 135.55, 137.88, 138.80, 148.03, 151.77, 162.02, 167.20. HRMS (<i>m/z</i>): [M + H] ⁺ , calcd for C ₂₆ H ₁₉ N ₂ O ₄ , 423.1339; found, 423.1349.
	(E)-3-(3-carboxyphenyl)-2-(4-(3-acetoxyprop-1-yn-1-yl)styryl)quinazolin-4(3H)-one (29). The compound was prepared according to the procedure described previously in 79% yield (0.367 g). ¹ ¹ H (600 MHz, DMSO- <i>d</i> ₆) δ 2.07 (s, 3H), 4.92 (s, 2H), 6.36 (d, <i>J</i> = 15.6 Hz, 1H), 7.40 (m, 4H), 7.54 (t, <i>J</i> = 7.8 Hz, 1H), 7.75 (m, 2H), 7.79 (d, <i>J</i> = 7.8 Hz, 1H), 7.89 (m, 2H), 8.03 (s, 1H), 8.13 (m, 2H), 13.28 (s, 1H). ¹³ C NMR (151 MHz, DMSO- <i>d</i> ₆) δ 20.47, 52.22, 85.34, 85.99, 120.63, 121.05, 122.46, 126.46, 126.75, 127.26, 127.84 (2C), 129.94, 130.05, 130.07, 132.09 (2C), 132.35, 133.44, 134.83, 135.27, 137.15, 137.89, 147.31, 150.98, 161.30, 166.50, 169.73. HRMS (<i>m/z</i>): [M + H] ⁺ , calcd for C ₂₈ H ₂₁ N ₂ O ₅ , 465.1445; found, 465.1421.
	(E)-3-(3-carboxyphenyl)-2-(4-fluorostyryl)quinazolin-4(3H)-one (30). This compound was prepared according to the procedure described previously in 76% yield (0.69 g). ¹ ¹ H NMR (500 MHz, DMSO- <i>d</i> ₆) δ 6.25 (d, <i>J</i> = 15.55 Hz, 1H), 7.19 (m, 2H), 7.46 (m, 2H), 7.55 (d, <i>J</i> = 8.17 Hz, 1H), 7.73 (m, 2H), 7.77 (d, <i>J</i> = 8.17 Hz, 1H), 7.87 (m, 2H), 8.02 (s, 1H), 8.13 (m, 2H). ¹³ C NMR (126 MHz, DMSO- <i>d</i> ₆) δ 115.91, 116.20, 119.77, 120.59, 126.48, 126.66, 127.21, 129.82, 129.93, 130.06, 131.39, 131.44, 132.34, 133.49, 134.84, 137.25, 137.79, 147.39, 151.15, 161.35, 166.55. HRMS (<i>m/z</i>): [M + H] ⁺ , calcd for C ₂₃ H ₁₆ N ₂ O ₃ , 387.1139; found, 387.1140.
	(E)-3-(3-carboxyphenyl)-2-(3,4-difluorostyryl)quinazolin-4(3H)-one (31). This compound was prepared according to the procedure described previously in 37% yield (0.19 g). ¹ ¹ H (500 MHz, DMSO- <i>d</i> ₆) δ 6.34 (d, <i>J</i> = 15.5 Hz, 1H), 7.28 (s, br, 1H), 7.41 (q, <i>J</i> = 8.5 Hz, 1H), 7.53 (m, 2H), 7.71 (m, 2H), 7.78 (d, <i>J</i> = 8.0 Hz, 1H), 7.84 (d, 16.0 Hz, 1H), 7.87 (m, 1H), 8.01 (s, br, 1H), 8.12 (m, 2H). ¹³ C NMR (126 MHz, DMSO- <i>d</i> ₆) δ 117.2, 117.4, 118.7, 118.8, 121.3, 122.1, 125.3, 125.4(3C), 127.2, 127.5, 127.9, 130.6, 130.7 (2C), 133.0, 133.3, 133.4 (4C), 135.5, 137.4, 137.7, 148.0, 149.3, 149.4, 149.7, 149.8, 151.2, 151.3, 151.7, 151.8, 162.0, 167.2. HRMS (<i>m/z</i>): [M + Na] ⁺ , calcd for C ₂₃ H ₁₄ F ₂ N ₂ NaO ₃ , 427.0865; found, 427.0863.
	(E)-3-(3-carboxyphenyl)-2-(3-chloro-4-fluorostyryl)quinazolin-4(3H)-one (32). This compound was prepared according to the procedure described previously in 43% yield (0.23 g). ¹ ¹ H (500 MHz, DMSO- <i>d</i> ₆) δ 6.34 (d, <i>J</i> = 15.5 Hz, 1H), 7.35 (m, 2H), 7.54 (t, <i>J</i> = 7.5 Hz, 1H), 7.71 (m, 4H), 7.82 (m, 2H), 8.02 (s, 1H), 8.12 (d, <i>J</i> = 8.0 Hz, 2H). ¹³ C

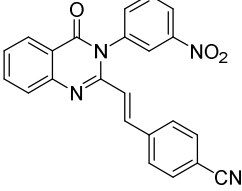
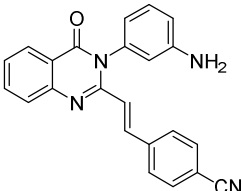
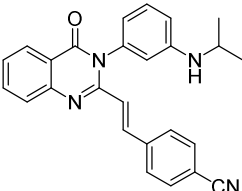
	NMR (126 MHz, DMSO- <i>d</i> ₆) δ 118.1, 118.3, 120.8, 120.9, 121.3, 122.1, 127.1, 127.5, 127.9, 128.6, 128.7, 130.6, 130.7, 133.0, 133.6 (2C), 134.1, 135.5, 137.1, 137.7, 147.9, 151.7, 157.3, 159.3, 161.9, 167.2. HRMS (<i>m/z</i>): [M + H] ⁺ , calcd for C ₂₃ H ₁₅ FCIN ₂ O ₃ , 421.0750; found, 421.0751.
	(E)-3-(3-carboxyphenyl)-2-(4-(methylsulfonyl)styryl)quinazolin-4(3H)-one (33). This compound was prepared according to the procedure described previously in 46% yield (0.26 g). ¹ H NMR (500 MHz, DMSO- <i>d</i> ₆) δ 3.19 (s, 3H), 6.49 (d, <i>J</i> = 15.5 Hz, 1H), 7.57 (t, <i>J</i> = 8.0 Hz, 1H), 7.66 (d, <i>J</i> = 8.0 Hz, 2H), 7.74 (m, 2H), 7.81 (d, <i>J</i> = 8.0 Hz, 1H), 7.86 (m, 4H), 8.05 (s, br, 1H), 8.12 (m, 2H). ¹³ C NMR (100 MHz, DMSO- <i>d</i> ₆) δ 44.0, 121.4, 124.1, 127.2, 127.7, 128.0, 128.3, 128.9, 130.6, 130.7, 130.8, 133.0, 134.1, 135.5, 137.6, 137.7, 140.3, 141.7, 147.9, 151.5, 161.9, 167.2. HRMS (<i>m/z</i>): [M + H] ⁺ , calcd for C ₂₄ H ₁₉ N ₂ O ₅ S, 447.1009; found, 447.1029.
	(E)-3-(3-carboxyphenyl)-2-(4-(2-methoxyethoxy)styryl)quinazolin-4(3H)-one (34). The compound was prepared according to the procedure described previously in 85% yield (0.374 g). ¹ H NMR (600 MHz, DMSO- <i>d</i> ₆) δ 3.59 (s, 3H), 3.67 (s, 2H), 6.29 (d, <i>J</i> = 15.6 Hz, 1H), 7.24 (d, <i>J</i> = 7.8 Hz, 2H), 7.33 (d, <i>J</i> = 7.8 Hz, 2H), 7.54 (ddd, <i>J</i> = 8.4, 7.8, 1.2 Hz, 1H), 7.74 (m, 2H), 7.79 (d, <i>J</i> = 7.8 Hz, 1H), 7.87 (d, <i>J</i> = 15.6 Hz, 1H), 7.88 (ddd, <i>J</i> = 8.4, 7.8, 1.2 Hz, 1H), 8.02 (m, 1H), 8.13 (m, 2H), 13.13 (s, 1H). ¹³ C NMR (151 MHz, DMSO- <i>d</i> ₆) δ 39.86, 51.74, 119.70, 120.57, 126.44, 126.59, 127.20, 127.60 (2C), 129.92, 130.00, 130.02, 130.05 (2C), 132.33, 133.42, 133.44, 134.80, 136.22, 137.25, 138.68, 147.39, 151.21, 161.33, 166.50, 171.22. HRMS (<i>m/z</i>): [M + H] ⁺ , calcd for C ₂₆ H ₂₁ N ₂ O ₅ , 441.1445; found, 441.1475.
	(E)-3-(2-hydroxyphenyl)-2-(4-nitrostyryl)quinazolin-4(3H)-one (35). This compound was prepared according to the procedure for 5 in 22% yield (0.05 g). ¹ H NMR (500 MHz, DMSO- <i>d</i> ₆) δ 6.59 (d, <i>J</i> = 15.55 Hz, 1H), 7.02 (t, <i>J</i> = 6.98 Hz, 1H), 7.41 (t, <i>J</i> = 7.98 Hz, 1H), 7.57 (t, <i>J</i> = 7.58 Hz, 1H), 7.64 (d, <i>J</i> = 7.38 Hz, 2H), 7.79 (d, <i>J</i> = 7.78 Hz, 1H), 7.90 (t, <i>J</i> = 7.18 Hz, 1H), 8.15 (d, <i>J</i> = 7.58 Hz, 1H), 8.20 (d, <i>J</i> = 6.78 Hz, 2H), 10.05 (s, 1H). ¹³ C NMR (126 MHz, DMSO- <i>d</i> ₆) δ 116.79, 119.77, 120.96, 123.52, 123.85, 124.24, 126.49, 127.29, 128.43, 130.07, 130.82, 134.80, 141.30, 147.25, 147.52, 151.36, 153.13, 160.84, 162.38. HRMS (<i>m/z</i>): [M + H] ⁺ , calcd for C ₂₂ H ₁₆ N ₃ O ₄ , 386.1135; found, 386.1159.

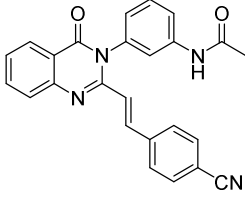
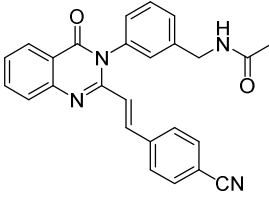
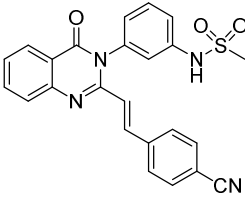
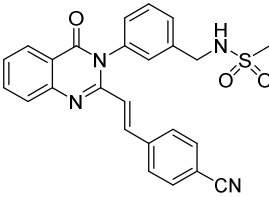
	<p>(E)-3-(4-hydroxyphenyl)-2-(4-nitrostyryl)quinazolin-4(3H)-one (36). This compound was prepared according to the procedure for 5 in 13% yield (0.02 g). ¹H NMR (500 MHz, DMSO-<i>d</i>₆) δ 6.59 (d, <i>J</i> = 15.55 Hz, 1H), 6.94 (d, <i>J</i> = 6.38 Hz, 2H), 7.23 (d, <i>J</i> = 6.58 Hz, 2H), 7.55 (t, <i>J</i> = 6.58 Hz, 1H), 7.65 (d, <i>J</i> = 6.98 Hz, 2H), 7.77 (d, <i>J</i> = 7.98 Hz, 1H), 7.88 (t, <i>J</i> = 6.78 Hz, 1H), 7.92 (d, <i>J</i> = 15.55 Hz, 1H), 8.12 (d, <i>J</i> = 7.18 Hz, 1H), 8.19 (d, <i>J</i> = 7.58 Hz, 2H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 116.15, 120.83, 124.22, 124.45, 126.52, 126.97, 127.31, 127.56, 128.49, 129.90, 134.76, 136.01, 141.35, 147.16, 147.48, 151.44, 157.85, 161.39. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₆N₃O₄, 386.1135; found, 386.1118.</p>
	<p>(E)-3-(3-acetyloxyphenyl)-2-(4-nitrostyryl)quinazolin-4(3H)-one (37). 3-(3-hydroxyphenyl)-2-methylquinazolin-4(3H)-one (0.1 g, 0.4 mmol) and 4-nitrobenzaldehyde (0.08 g, 0.53 mmol) were suspended in 4 mL glacial acetic acid. To the reaction mixture, sodium acetate (0.05 g, 0.65 mmol) and acetic anhydride (400 μL, 3 mmol) were added. The suspension was heated to dissolve the components, refluxed overnight (18 h), and the resulting precipitate was filtered and purified by silica gel column chromatography in 61% yield (0.12 g). ¹H NMR (500 MHz, CDCl₃) δ 2.32 (s, 3H), 6.60 (d, <i>J</i> = 15.35 Hz, 1H), 7.16 (s, 1H), 7.23 (d, <i>J</i> = 7.98 Hz, 1H), 7.32 (d, <i>J</i> = 8.17 Hz, 1H), 7.51 (m, 3H), 7.62 (t, <i>J</i> = 7.98 Hz, 1H), 7.81 (m, 2H), 7.98 (d, <i>J</i> = 15.35 Hz, 1H), 8.16 (d, <i>J</i> = 8.77 Hz, 1H), 8.30 (d, <i>J</i> = 8.18 Hz, 1H). ¹³C NMR (126 MHz, CDCl₃) δ 21.11, 121.00, 122.75, 122.93, 123.76, 124.12, 126.20, 127.25, 127.33, 127.63, 128.44, 130.75, 134.93, 137.28, 137.36, 141.39, 147.41, 147.97, 150.41, 151.54, 161.83, 169.11. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₄H₁₈N₃O₅, 428.1241; found, 428.1238.</p>
	<p>(E)-3-(4-acetyloxyphenyl)-2-(4-nitrostyryl)quinazolin-4(3H)-one (38). This compound was prepared according to the procedure for 37 in 32% yield (0.05 g). ¹H NMR (500 MHz, CDCl₃) δ 2.38 (s, 3H), 6.51 (d, <i>J</i> = 15.35 Hz, 1H), 7.36 (m, 4H), 7.46 (d, <i>J</i> = 8.77 Hz, 2H), 7.52 (t, <i>J</i> = 7.78 Hz, 1H), 7.81 (m, 2H), 7.96 (d, <i>J</i> = 15.35 Hz, 1H), 8.17 (d, <i>J</i> = 8.57 Hz, 2H), 8.30 (d, <i>J</i> = 7.78 Hz, 1H). ¹³C NMR (126 MHz, CDCl₃) δ 21.42, 121.22, 123.49, 124.07, 124.37, 127.45, 127.55, 127.80, 128.54, 129.98, 134.08, 135.11, 137.36, 141.49, 147.61, 148.18, 150.84, 151.48, 162.21, 169.18. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₄H₁₈N₃O₅, 428.1241; found, 428.1224.</p>

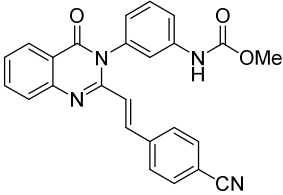
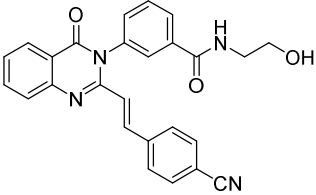
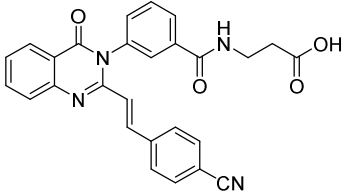
	<p>(E)-3-(3-methoxyphenyl)-2-(4-nitrostyryl)quinazolin-4(3H)-one (39). This compound was prepared according to the procedure for 5 in 52% yield (0.37 g). ¹H NMR (500 MHz, CDCl₃) δ 3.86 (s, 3H), 6.34 (d, <i>J</i> = 15.55 Hz, 1H), 6.87 (s, 1H), 6.90 (d, <i>J</i> = 7.58 Hz, 1H), 7.02 (t, <i>J</i> = 8.57 Hz, 2H), 7.10 (d, <i>J</i> = 8.37 Hz, 1H), 7.33 (m, 2H), 7.49 (m, 2H), 7.80 (m, 2H), 7.95 (d, <i>J</i> = 15.75 Hz, 1H), 8.31 (d, <i>J</i> = 7.98 Hz, 1H). ¹³C NMR (126 MHz, CDCl₃) δ 55.78, 114.37, 115.65, 116.00, 116.30, 119.74, 119.78, 121.03, 121.13, 126.90, 127.40, 127.55, 129.80, 129.91, 130.92, 131.75, 134.90, 138.86, 147.94, 151.74, 160.98, 162.44, 165.35. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₃H₁₈FN₂O₂, 373.1347; found, 373.1335.</p>
	<p>(E)-3-(3-nitrophenyl)-2-(4-nitrostyryl)quinazolin-4(3H)-one (40). This compound was prepared according to the procedure for 5 in 68% yield (1.87 g). ¹H NMR (500 MHz, DMSO-<i>d</i>₆) δ 6.31 (d, <i>J</i> = 15.35 Hz, 1H), 7.18 (t, <i>J</i> = 8.77 Hz, 2H), 7.55 (m, 3H), 7.80 (d, <i>J</i> = 8.17 Hz, 1H), 7.89 (m, 3H), 7.95 (d, <i>J</i> = 7.38 Hz, 1H), 8.13 (d, <i>J</i> = 7.98 Hz, 1H), 8.43 (d, <i>J</i> = 8.37 Hz, 1H), 8.51 (s, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 116.01, 116.18, 119.84, 120.70, 124.42, 124.89, 126.64, 126.86, 127.39, 130.32, 130.39, 131.15, 131.53, 131.56, 135.12, 136.11, 138.19, 138.34, 147.54, 148.73, 151.16, 161.54, 162.06, 164.03. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₅FN₃O₃, 388.1092; found, 388.1117.</p>
	<p>(E)-3-(3-aminophenyl)-2-(4-nitrostyryl)quinazolin-4(3H)-one (41). Compound 40 (1.80 g, 4.6 mmol) and tin (II) chloride dihydrate (5.24 g, 23.2 mmol) were suspended in 50 mL ethanol to which 1 mL concentrated HCl was added. The solution was refluxed for 3 h and then allowed to stir overnight at 25 °C. The solution was concentrated <i>in vacuo</i> and EtOAc was added to completely dissolve the residue. The EtOAc solution was then neutralized with a NaOH solution and washed with saturated potassium fluoride. The mixture was then filtered through a bed of celite to remove the tin precipitate. The resulting filtrate was washed three times with a saturated KF solution and the EtOAc layer was dried with MgSO₄ and concentrated <i>in vacuo</i>. The residue was then purified by silica gel chromatography (EtOAc/hexanes) to give the product in 32% yield (0.54 g). ¹H NMR (500 MHz, DMSO-<i>d</i>₆) δ 5.46 (s, 2H), 6.38 (d, <i>J</i> = 15.55 Hz, 1H), 6.50 (d, <i>J</i> = 7.58 Hz, 1H), 6.54 (s, 1H), 6.73 (d, <i>J</i> = 8.17 Hz, 1H), 7.23 (m, 3H), 7.43 (m, 2H), 7.52 (t, <i>J</i> = 7.18 Hz, 1H), 7.74 (d, <i>J</i> = 7.78 Hz, 1H), 7.85 (d, <i>J</i> = 7.38 Hz, 1H), 7.87 (d, <i>J</i> = 15.75 Hz, 1H), 8.12 (d, <i>J</i> = 7.98 Hz, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 113.50, 114.48, 115.42, 116.05, 116.23, 119.95, 120.62, 126.45, 126.55, 127.12, 129.57, 129.64, 129.99, 131.58, 131.60, 134.69, 137.33, 137.53, 147.32, 150.05, 151.35, 161.03, 163.77. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₇FN₃O,</p>

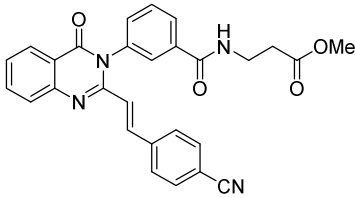
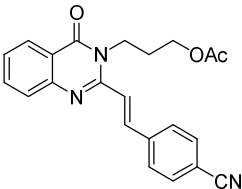
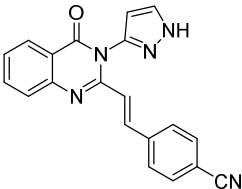
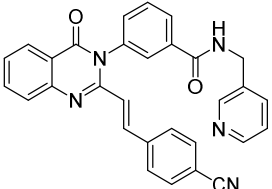
	358.1350; found, 358.1360.
	<p>(E)-3-(3-(acetamido)phenyl)-2-(4-fluorostyryl)quinazolin-4(3H)-one (42). Compound 41 (0.15 g, 0.42 mmol) was dissolved in 4 mL pyridine, to which acetic anhydride was added (0.21 g, 2.1 mmol). The reaction was stirred for 3 h at 25 °C and then refluxed for 1 h. The reaction was then poured over ice in 2M HCl and the resulting precipitate was filtered out and washed with cold water to give the product in 83% yield (0.15 g). ¹H (500 MHz, DMSO-<i>d</i>₆) δ 2.06 (s, 3H), 6.30 (d, <i>J</i> = 15.55 Hz, 1H), 7.12 (d, <i>J</i> = 7.38 Hz, 1H), 7.22 (t, <i>J</i> = 8.77 Hz, 1H), 7.46 (m, 2H), 7.55 (m, 2H), 7.73 (d, <i>J</i> = 6.58 Hz, 2H), 7.82 (d, <i>J</i> = 7.98 Hz, 1H), 7.90 (t, <i>J</i> = 6.78 Hz, 1H), 7.93 (d, <i>J</i> = 15.75 Hz, 1H), 8.13 (d, <i>J</i> = 7.98 Hz, 1H), 10.33 (s, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 24.09, 111.17, 116.12, 116.30, 118.45, 118.87, 119.70, 120.31, 123.21, 126.64, 127.10, 127.41, 130.03, 135.13, 136.70, 140.61, 141.61, 158.28, 160.81, 168.78. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₄H₁₉FN₃O₂, 400.1456; found, 400.1440.</p>
	<p>(E)-3-(3-(methylsulfonamido)phenyl)-2-(4-fluorostyryl)quinazolin-4(3H)-one (43). Compound 41 (0.15 g, 0.42 mmol) was dissolved in 4 mL pyridine and cooled to 0 °C, to which mesyl chloride (0.24 g, 2.1 mmol) was added. The reaction was stirred at 25 °C for 3 h and then poured over ice in 2M HCl. The resulting precipitate was filtered out and washed with cold 2M HCl followed by cold water to give the product in 99% yield (0.19 g). ¹H (500 MHz, DMSO-<i>d</i>₆) δ 3.05 (s, 3H), 6.31 (d, <i>J</i> = 15.75 Hz, 1H), 7.22 (m, 3H), 7.33 (s, 1H), 7.42 (d, <i>J</i> = 8.17 Hz, 2H), 7.48 (m, 2H), 7.58 (t, <i>J</i> = 7.98 Hz, 2H), 7.92 (d, <i>J</i> = 3.59 Hz, 2H), 7.99 (d, <i>J</i> = 15.75 Hz, 1H), 7.8.12 (m, 2H), 10.20 (s, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 54.50, 115.94, 116.11, 118.14, 119.37, 120.08, 120.27, 123.88, 126.52, 127.01, 127.27, 129.92, 129.99, 130.43, 135.05, 139.56, 141.42, 146.46, 160.54, 161.94. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₃H₁₉FN₃O₃S, 436.1126; found, 436.1136.</p>
	<p>(E)-3-(3-carbomylphenyl)-2-(4-fluorostyryl)quinazolin-4(3H)-one (44). This compound was prepared according to the procedure for 5 in 13% yield (0.15 g). ¹H (500 MHz, DMSO-<i>d</i>₆) δ 6.25 (d, <i>J</i> = 15.31 Hz, 1H), 7.20 (t, <i>J</i> = 8.85 Hz, 2H), 7.45 (m, 2H), 7.56 (t, <i>J</i> = 6.70 Hz, 2H), 7.68 (m, 2H), 7.78 (d, <i>J</i> = 8.37 Hz, 1H), 7.93 (m, 3H), 8.09 (m, 2H), 8.13 (d, <i>J</i> = 8.13 Hz, 1H), 11.98 (br. s, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 115.61, 116.01, 116.18, 120.51, 126.49, 127.24, 128.12, 128.31, 129.80, 129.87, 131.86, 134.93, 135.72, 136.98, 146.94, 147.36, 151.21, 161.34, 161.86, 166.76, 172.08. HRMS (<i>m/z</i>): [M + Na]⁺, calcd for C₂₃H₁₆FN₃NaO₂, 408.1119; found, 408.1098.</p>

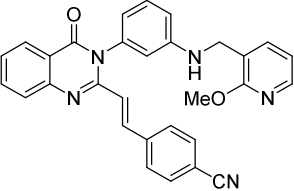
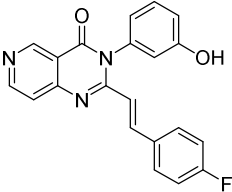
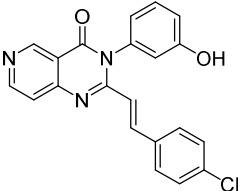
	<p>(E)-3-(3-(2-hydroxyethyl)carbonylphenyl)-2-(4-fluorostyryl)quinazolin-4(3H)-one (45). Compound 30 (0.19 g, 0.5 mmol) was dissolved in 12 mL THF, to which 2-aminoethanol (0.037 g, 0.6 mmol), HOBt (0.074 g, 0.55 mmol), and DIC (0.069 g, 0.55 mmol) were added. The mixture was stirred at room temperature for 8 h and 25 mL EtOAc was added and washed with saturated NH₄Cl. The organic phase was dried over Na₂SO₄, concentrated <i>in vacuo</i>, and purified by silica gel column chromatography to give the product in 69% yield (0.15 g). ¹H (400 MHz, DMSO-<i>d</i>₆) δ 3.28 (m, 2H), 3.49 (m, 2H), 4.75 (d, <i>J</i> = 5.6 Hz, 1H), 6.28 (d, <i>J</i> = 15.6 Hz, 1H), 7.20 (d, <i>J</i> = 8.8 Hz, 2H), 7.43 (m, 2H), 7.53 (m, 1H), 7.64 (d, <i>J</i> = 8.0 Hz, 1H), 7.71 (t, <i>J</i> = 8.0 Hz, 1H), 7.78 (m, 1H), 7.87 (m, 3H), 8.06 (d, <i>J</i> = 7.6 Hz, 1H), 8.14 (dd, <i>J</i> = 1.6, 8.0 Hz, 1H), 8.59 (t, <i>J</i> = 5.2 Hz, 1H). ¹³C NMR (100 MHz, DMSO-<i>d</i>₆) δ 42.3, 59.6, 115.9, 116.2, 119.7, 119.8, 120.5, 126.4, 126.7, 127.2, 127.8, 129.7, 129.8, 129.9, 131.4, 131.5, 131.6, 134.9, 136.0, 136.9, 137.8, 147.3, 151.2, 161.3, 161.6, 164.1, 165.2. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₅H₂₁FN₃O₃, 430.1561; found, 430.1557.</p>
	<p>(E)-3-(3-(carboxymethyl)phenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (46). This compound was prepared according to the procedure for 5 in 47% yield (0.64 g). ¹H (500 MHz, DMSO-<i>d</i>₆) δ 3.32 (s, 2H), 6.50 (d, <i>J</i> = 15.55 Hz, 1H), 7.38 (m, 2H), 7.47 (d, <i>J</i> = 7.58 Hz, 1H), 7.56 (m, 4H), 7.80 (m, 3H), 7.90 (m, 2H), 8.15 (d, <i>J</i> = 7.98 Hz, 1H). HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₅H₁₈FN₃O₃, 408.1343; found, 408.1362.</p>
	<p>(E)-3-(3-(acetoxymethyl)phenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (47). This compound was prepared according to the procedure for 5 in 55% yield (0.39 g). ¹H (500 MHz, DMSO-<i>d</i>₆) δ 2.04 (s, 3H), 5.17 (s, 2H), 6.45 (d, <i>J</i> = 15.55 Hz, 1H), 7.45 (d, <i>J</i> = 7.58 Hz, 1H), 7.49 (s, 1H), 7.57 (m, 4H), 7.62 (t, <i>J</i> = 7.58 Hz, 1H), 7.81 (m, 3H), 7.90 (m, 2H), 8.14 (d, <i>J</i> = 7.78 Hz, 1H). HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₆H₂₀FN₃O₃, 422.1505; found, 422.1558.</p>
	<p>(E)-3-(3-fluorophenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (48). This compound was prepared according to the procedure for 5 in 57% yield (1.36 g). ¹H (400 MHz, DMSO-<i>d</i>₆) δ 6.51 (d, <i>J</i> = 15.6 Hz, 1H), 7.35 (d, <i>J</i> = 7.6 Hz, 1H), 7.43 (m, 6H), 7.82 (d, <i>J</i> = 8.0 Hz, 3H), 7.88 (m, 2H), 8.14 (m, 1H). ¹³C NMR (100 MHz, DMSO-<i>d</i>₆) δ 112.3, 117.1, 117.3, 117.5, 119.3, 121.4, 123.9, 126.1(2C), 127.2, 127.8, 128.0, 128.9, 131.9, 132.0, 133.5, 135.6, 137.7, 138.0, 139.0, 140.0, 147.8, 151.4, 161.7, 161.8, 164.2. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₅H₁₅FN₃O, 368.1194; found, 368.1180.</p>

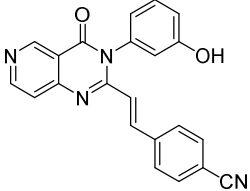
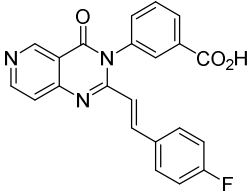
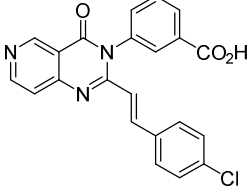
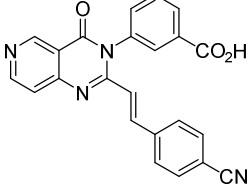
	<p>(E)-3-(3-nitrophenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (49). This compound was prepared according to the procedure for 5 in 99% yield (6.32 g). ¹H (500 MHz, DMSO-<i>d</i>₆) δ 6.54 (d, <i>J</i> = 15.35 Hz, 1H), 7.58 (t, <i>J</i> = 7.78 Hz, 1H), 7.67 (d, <i>J</i> = 8.17 Hz, 2H), 7.80 (m, 3H), 7.91 (m, 4H), 8.14 (d, <i>J</i> = 7.98 Hz, 1H), 8.43 (d, <i>J</i> = 8.18 Hz, 1H), 8.52 (s, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 111.77, 118.80, 120.88, 123.48, 124.50, 124.94, 126.68, 127.24, 127.54, 128.73, 131.16, 132.89, 135.18, 136.10, 137.54, 137.97, 139.35, 147.37, 148.72, 150.76, 161.45. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₃H₁₄N₄NaO₃, 417.0958; found, 417.0941.</p>
	<p>(E)-3-(3-aminophenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (50). This compound was prepared according to the procedure for 41 from compound 49 (6.0 g, 15.2 mmol) in 37 % yield (2.24 g). ¹H (500 MHz, DMSO-<i>d</i>₆) δ 5.47 (s, 2H), 6.51 (d, <i>J</i> = 7.78 Hz, 1H), 6.56 (m, 2H), 6.73 (d, <i>J</i> = 8.37 Hz, 1H), 7.23 (t, <i>J</i> = 7.98 Hz, 1H), 7.55 (m, 3H), 7.76 (d, <i>J</i> = 8.17 Hz, 1H), 7.82 (d, <i>J</i> = 8.37 Hz, 2H), 7.87 (t, <i>J</i> = 8.17 Hz, 1H), 7.90 (d, <i>J</i> = 15.55 Hz, 1H), 8.12 (d, <i>J</i> = 7.98 Hz, 1H). ¹³C NMR (126 MHz, DMSO-<i>d</i>₆) δ 111.50, 113.50, 114.57, 115.42, 118.61, 120.79, 123.52, 126.48, 126.92, 127.27, 128.00, 130.00, 132.92, 134.73, 136.48, 137.34, 139.43, 147.14, 150.04, 150.89, 160.91. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₃H₁₇N₄O, 365.1397; found, 365.1367.</p>
	<p>(E)-3-(3-(isopropylamino)phenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (51). Compound 50 (0.15 g, 0.42 mmol) was dissolved in 10 mL CH₂Cl₂, to which acetone (92 μL), 3 Å activated molecular sieves, and sodium triacetoxyborohydride (0.18 g, 0.84 mmol) were added. The mixture was stirred at room temperature for 6 days, filtered through celite, and concentrated <i>in vacuo</i>. The residue was purified by silica gel column chromatography to give the product in 13% yield (0.022 g). ¹H (500 MHz, CDCl₃) δ 1.19 (d, <i>J</i> = 6.0 Hz, 3H), 1.23 (d, <i>J</i> = 6.5 Hz, 3H), 3.59 (m, 1H), 6.53 (s, 1H), 6.59 (m, 2H), 6.78 (d, <i>J</i> = 8.0 Hz, 1H), 7.34 (t, <i>J</i> = 8.0 Hz, 1H), 7.42 (d, <i>J</i> = 8.5 Hz, 2H), 7.47 (m, 1H), 7.58 (d, <i>J</i> = 8.5 Hz, 2H), 7.77 (m, 2H), 7.93 (d, <i>J</i> = 15.5 Hz, 1H), 8.30 (d, <i>J</i> = 7.5 Hz, 1H). ¹³C NMR (126 MHz, CDCl₃) δ 22.6, 22.9, 112.6, 118.7, 121.4, 123.9, 127.2, 127.4, 127.6, 128.3, 130.8, 132.7, 134.8, 137.3, 137.8, 139.9, 147.7, 151.2, 162.1. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₆H₂₃N₄O, 407.1866; found, 407.1863.</p>

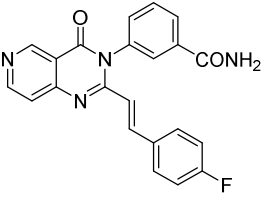
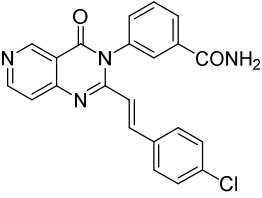
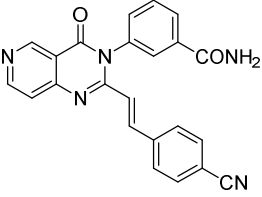
	<p>(E)-3-(3-(acetamido)phenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (52). This compound was prepared according to the procedure for 42 from compound 50 (0.20 g, 0.55 mmol) in 85% yield (0.19 g). ^1H (500 MHz, DMSO-d_6) δ 3.16 (s, 3H), 6.50 (d, J = 15.55 Hz, 1H), 7.13 (d, J = 7.38 Hz, 1H), 7.56 (m, 4H), 7.71 (d, J = 7.98 Hz, 2H), 7.80 (m, 3H), 7.91 (m, 2H), 8.14 (d, J = 7.98 Hz, 1H), 10.22 (s, 1H). ^{13}C NMR (126 MHz, DMSO-d_6) δ 24.23, 111.73, 118.77, 119.17, 119.75, 120.90, 123.50, 126.67, 127.22, 127.52, 128.34, 130.14, 133.06, 135.06, 137.03, 137.07, 139.50, 140.64, 147.33, 150.92, 161.24, 168.86. HRMS (m/z): $[\text{M} + \text{Na}]^+$, calcd for $\text{C}_{25}\text{H}_{18}\text{N}_4\text{NaO}_2$, 429.1322; found, 429.1291.</p>
	<p>(E)-3-(3-(acetamidomethyl)phenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (53). This compound was prepared according to the procedure for 42 in 21% yield (0.17 g). ^1H (500 MHz, DMSO-d_6) δ 1.83 (s, 3H), 4.32 (m, 2H), 6.45 (d, J = 15.5 Hz, 1H), 7.33 (d, J = 5.5 Hz, 2H), 7.44 (d, J = 8.0 Hz, 1H), 7.52 (m, 4H), 7.77 (m, 3H), 7.86 (m, 2H), 8.12 (dd, J = 1.0, 8.0 Hz, 1H), 8.43 (t, J = 11.5 Hz, 1H). ^{13}C NMR (100 MHz, DMSO-d_6) δ 23.3, 42.4, 112.2, 119.2, 121.4, 124.0, 127.1, 127.6, 127.9, 128.0, 128.1, 128.7, 128.8, 130.2, 133.5, 135.5, 137.4, 137.5, 140.0, 142.3, 147.8, 151.4, 161.8, 170.0. HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{26}\text{H}_{21}\text{N}_4\text{O}_2$, 421.1658; found, 421.1675.</p>
	<p>(E)-3-(3-(methylsulfonamido)phenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (54). This compound was prepared according to the procedure for 43 from compound 50 (0.36 g, 1 mmol) in 77% yield (0.33 g). ^1H (500 MHz, DMSO-d_6) δ 3.04 (s, 3H), 6.52 (d, J = 15.75 Hz, 1H), 7.21 (d, J = 7.58 Hz, 1H), 7.27 (s, 1H), 7.37 (d, J = 8.57 Hz, 1H), 7.59 (m, 4H), 7.81 (m, 3H), 7.90 (m, 2H), 8.15 (d, J = 7.98 Hz, 1H), 10.08 (s, 1H). ^{13}C NMR (126 MHz, DMSO-d_6) δ 111.60, 118.65, 119.73, 120.32, 120.76, 123.51, 125.24, 126.56, 127.12, 127.39, 128.22, 130.60, 132.90, 134.98, 136.89, 137.45, 139.39, 139.62, 147.19, 150.78, 161.10. HRMS (m/z): $[\text{M} + \text{Na}]^+$, calcd for $\text{C}_{24}\text{H}_{18}\text{N}_4\text{NaO}_3\text{S}$, 465.0992; found, 465.0973.</p>
	<p>(E)-3-(3-(methylsulfonamidomethyl)phenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (55). The compound was prepared according to the procedure described previously in 93% yield (68 mg, 75% yield). ^1H (400 MHz, DMSO-d_6) δ 2.85 (s, 3H), 4.27 (d, J = 6.4 Hz, 1H), 6.49 (d, J = 15.6 Hz, 1H), 7.38 (d, J = 7.2 Hz, 1H), 7.45 (s, 1H), 7.58 (m, 5H), 7.68 (t, J = 6.4 Hz, 1H), 7.81 (m, 3H), 7.90 (t, J = 7.6 Hz, 1H), 7.92 (d, J = 15.6 Hz, 1H), 8.15 (d, J = 8 Hz, 1H). ^{13}C NMR (100 MHz, DMSO-d_6) δ 40.05, 45.48, 111.54, 118.51, 120.72, 123.37, 126.43, 126.98, 127.29, 127.72, 127.90, 128.14 (2C), 128.51, 129.69,</p>

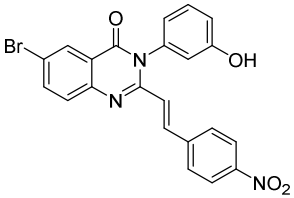
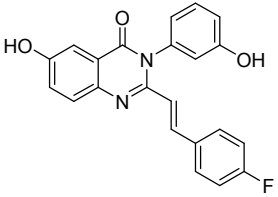
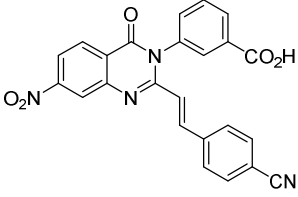
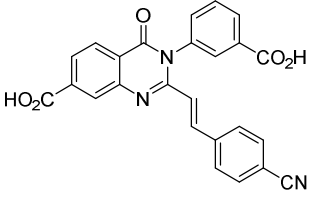
	132.76 (2C), 134.82, 136.74, 136.83, 139.30, 140.30, 147.16, 150.76, 161.12. HRMS (m/z): $[M + H]^+$, calcd for $C_{25}H_{21}N_4O_3S$, 457.1329; found, 457.1328.
	<p>(E)-3-(3-((methyloxycarbonyl)amino)phenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (56). Compound 50 (0.10 g, 0.27 mmol) and methyl chloroformate (100 μL, 1.35 mmol) were dissolved in pyridine (5 mL). The reaction was stirred at 25 $^{\circ}$C overnight (16 h) and water (10 mL) was added. The resulting precipitate was filtered and washed with water, followed by methanol and hexanes to give the product in 68% yield (0.08 g). 1H (500 MHz, DMSO-d_6) δ 3.66 (s, 3H), 6.50 (d, J = 15.35 Hz, 1H), 7.09 (d, J = 6.98 Hz, 1H), 7.57 (m, 6H), 7.81 (m, 3H), 7.91 (m, 2H), 8.14 (d, J = 7.58 Hz, 1H), 9.96 (s, 1H). ^{13}C NMR (126 MHz, DMSO-d_6) δ 51.84, 111.57, 120.72, 122.72, 123.27, 126.51, 127.06, 128.18, 130.09, 132.88, 134.91, 136.90, 137.07, 139.32, 142.52, 147.15, 150.32, 153.97, 154.64, 161.06, 192.50. HRMS (m/z): $[M + Na]^+$, calcd for $C_{25}H_{18}N_4NaO_3$, 445.1271; found, 445.1265.</p>
	<p>(E)-3-(3-(2-hydroxyethyl)carbonylphenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (57). This compound was prepared according to the procedure for 45 from compound 2 (0.19 g, 0.5 mmol) in 63 % yield (0.14 g). 1H (400 MHz, DMSO-d_6) δ 3.28 (m, 2H), 3.51 (q, J = 6.0 Hz, 2H), 4.75 (t, J = 5.6 Hz, 1H), 6.50 (d, J = 15.6 Hz, 1H), 7.57 (m, 3H), 7.63 (m, 2H), 7.81 (d, J = 8.4 Hz, 2H), 7.89 (m, 3H), 8.04 (m, 1H), 8.14 (m, 1H), 8.59 (t, J = 5.6 Hz, 1H). ^{13}C NMR (100 MHz, DMSO-d_6) δ 42.3, 59.6, 111.6, 118.6, 120.6, 126.5, 127.1, 127.4, 127.9, 128.1, 128.2, 129.7, 131.6, 132.8, 135.0, 136.0, 136.7, 137.0, 139.2, 147.2, 150.8, 161.2, 165.2. HRMS (m/z): $[M + H]^+$, calcd for $C_{26}H_{21}N_4O_3$, 437.1608; found, 437.1635.</p>
	<p>(E)-3-(3-(2-carboxyethyl)carbonylphenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (58). The compound was prepared according to the procedure described for 28 in 56% yield (0.130 g). 1H (600 MHz, MeOD-d_4) δ 2.64 (t, J = 6.6 Hz, 2H), 3.64 (td, J = 6.6, 2.4 Hz, 2H), 6.55 (d, J = 15.6 Hz, 1H), 7.51 (m, 2H), 7.58 (ddd, J = 7.8, 7.2, 1.2 Hz, 1H), 7.59 (ddd, J = 7.8, 1.8, 1.2 Hz, 1H), 7.67 (m, 2H), 7.73 (td, J = 7.8, 0.6 Hz, 1H), 7.85 (m, 2H), 7.9 (ddd, J = 7.8, 7.2, 1.2 Hz, 1H), 7.95 (d, J = 15.6 Hz, 1H), 8.04 (ddd, J = 7.8, 1.2, 1 Hz, 1H), 8.24 (ddd, J = 7.8, 1.2, 0.6 Hz, 1H). ^{13}C NMR (100 MHz, DMSO-d_6) δ 34.33, 36.38, 112.30, 119.29, 121.38, 124.10, 127.20, 127.83, 128.09, 128.54, 128.80, 128.95 (2C), 130.47, 132.43, 133.56 (2C), 135.69, 136.47, 137.46, 137.73, 139.98, 147.89, 151.47, 161.92, 165.84, 173.53. HRMS (m/z): $[M + H]^+$, calcd for $C_{27}H_{21}N_4O_4$, 465.1557; found, 465.1539.</p>

	<p>(E)-3-(3-(3-methoxy-3-oxopropyl)carbamoylphenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (59). Compound 26 (0.393 g, 1 mmol) was dissolved in a DMF (15 mL) solution containing HATU (0.418 g, 1.1 mmol). β-Alanine methyl ester hydrochloride (0.139 g, 1 mmol) was added and the mixture was stirred for 20 minutes followed by the addition of NMM (0.22 mL, 2 mmol). The reaction was stirred at 25 °C overnight (16 h) and concentrated <i>in vacuo</i>. Water was added to precipitate the product in 83% yield (0.397 g). ^1H (400 MHz, DMSO-d_6) δ 2.60 (t, $J = 7.2$ Hz, 2H), 3.5 (m, 2H), 3.58 (s, 3H), 6.49 (d, $J = 15.6$ Hz, 1H), 7.58 (m, 3H), 7.65 (m, 1H), 7.72 (m, 1H), 7.81 (m, 3H), 7.91 (m, 3H), 8.03 (d, $J = 7.6$ Hz, 1H), 8.15 (d, $J = 8$ Hz, 1H), 8.68 (t, $J = 5.6$ Hz, 1H). ^{13}C NMR (100 MHz, DMSO-d_6) δ 33.38, 35.56, 51.33, 111.58, 118.53, 120.64, 123.37, 126.46, 127.09, 127.35, 127.79, 128.07, 128.20 (2C), 129.75, 131.74, 132.80 (2C), 134.95, 135.68, 136.73, 137.00, 139.26, 147.16, 150.73, 161.19, 165.18, 171.68. HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{28}\text{H}_{23}\text{N}_4\text{O}_4$, 479.1714; found, 479.1710.</p>
	<p>(E)-3-(3-acetoxypentyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (60). This compound was prepared according to the procedure for 5 in 37% yield (0.14 g). ^1H (500 MHz, DMSO-d_6) δ 1.89 (s, 3H), 2.02 (t, $J = 5.98$ Hz, 2H), 4.09 (quin, $J = 5.98$ Hz, 2H), 4.42 (t, $J = 6.18$ Hz, 2H), 7.52 (t, $J = 7.78$ Hz, 1H), 7.60 (d, $J = 15.35$ Hz, 1H), 7.70 (d, $J = 8.17$ Hz, 1H), 7.83 (t, $J = 8.17$ Hz, 1H), 7.92 (d, $J = 8.37$ Hz, 2H), 7.96 (d, $J = 15.35$ Hz, 1H), 8.04 (d, $J = 8.37$ Hz, 2H), 8.13 (d, $J = 7.98$ Hz, 1H). HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{22}\text{H}_{20}\text{N}_3\text{O}_3$, 374.1505; found, 374.1543.</p>
	<p>(E)-3-(1H-pyrazol-3-yl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (61). This compound was prepared according to the procedure for 5 in 9% yield (0.05 g). ^1H (500 MHz, DMSO-d_6) δ 6.52 (m, 2H), 7.58 (t, $J = 7.98$ Hz, 1H), 7.62 (d, $J = 8.17$ Hz, 2H), 7.79 (d, $J = 8.17$ Hz, 1H), 7.84 (d, $J = 7.78$ Hz, 2H), 7.92 (m, 2H), 8.03 (s, 1H), 8.15 (d, $J = 7.98$ Hz, 1H). HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{20}\text{H}_{14}\text{N}_5\text{O}$, 340.1193; found, 340.1207.</p>
	<p>(E)-3-(pyridine-3-ylmethyl)carbamoylphenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (62). This compound was prepared according to the procedure for 45 from compound 2 (0.16 g, 0.4 mmol) and pyridine-3-ylmethanamine (0.052 g, 0.52 mmol) in 48 % yield (0.093 g). ^1H (500 MHz, CD_3Cl) δ 4.62 (d, $J = 6.0$ Hz, 2H), 6.37 (d, $J = 15.5$ Hz, 1H), 7.28 (m, 1H), 7.37 (d, $J = 8.0$ Hz, 3H), 7.45 (m, 2H), 7.56 (m, 3H), 7.75 (m, 4H), 7.94 (d, $J = 15.5$ Hz, 1H), 8.01 (d, $J = 7.5$ Hz, 1H), 8.17 (d, $J = 8.0$ Hz, 1H), 8.47 (d, $J = 4.5$ Hz, 1H), 8.57 (s, 1H). ^{13}C NMR (100 MHz, DMSO-d_6) δ 41.9, 113.1, 118.6, 120.9, 122.9, 124.1, 127.2, 127.5, 127.8, 128.4, 126.7, 130.6, 131.9, 132.8,</p>

	134.4, 135.3, 136.5, 136.9, 137.2, 138.4, 139.5, 147.7, 148.5, 149.0, 150.4, 162.4, 166.4. HRMS (m/z): $[M + H]^+$, calcd for $C_{30}H_{22}N_5O_2$, 484.1768; found, 484.1788.
	<p>(E)-3-(((2-methoxyphenyl)methyl)aminophenyl)-2-(4-cyanostyryl)quinazolin-4(3H)-one (63). Compound 50 (0.060 g, 0.165 mmol) was dissolved in 5 mL MeOH, to which 2-methoxynicotinaldehyde (0.045 g, 0.33 mmol), 3 Å activated molecular sieves, and sodium cyanoborohydride (0.021 g, 0.033 mmol) were added. The mixture was stirred at room temperature for 5 days, filtered through celite, and concentrated <i>in vacuo</i>. The residue was purified by silica gel column chromatography to give the product in 37% yield (0.030 g). 1H (500 MHz, CD_3Cl) δ 3.96 (s, 3H), 4.29 (s, 2H), 6.51 (m, 2H), 6.61 (d, $J = 7.5$ Hz, 1H), 6.69 (t, $J = 6.0$ Hz, 1H), 6.78 (d, $J = 8.0$ Hz, 1H), 7.32 (m, 3H), 7.47 (m, 2H), 7.60 (d, $J = 8.0$ Hz, 2H), 7.80 (d, $J = 7.5$ Hz, 2H), 7.95 (m, 2H), 8.29 (d, $J = 7.5$ Hz, 1H). ^{13}C NMR (100 MHz, $DMSO-d_6$) δ 43.1, 53.7, 112.8, 112.9, 114.6, 116.9, 117.5, 118.7, 120.9, 121.2, 123.2, 127.3, 127.4, 127.5, 128.4, 130.9, 132.7, 135.0, 136.9, 137.7, 138.0, 139.8, 145.7, 147.2, 149.5, 151.4. HRMS (m/z): $[M + H]^+$, calcd for $C_{30}H_{24}N_5O_2$, 486.1925; found, 486.1941.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-fluorostyryl)pyrido[4,3-d]pyrimidin-4(3H)-one (64). This compound was prepared according to the procedure for 5 in 34% yield (0.098 g) as acetate salt. 1H (400 MHz, $DMSO-d_6$) δ 1.91 (s, 3H), 6.33 (d, $J = 15.6$ Hz, 1H), 6.87 (m, 2H), 6.89 (m, 1H), 7.22 (m, 2H), 7.41 (t, $J = 8.4$ Hz, 1H), 7.46 (m, 2H), 7.62 (m, 1H), 7.99 (d, $J = 15.6$ Hz, 1H), 8.84 (m, 1H), 9.26 (d, $J = 2.4$ Hz, 1H), 9.96 (s, 1H), 11.98 (s, 1H). ^{13}C NMR (100 MHz, $DMSO-d_6$) δ 21.0, 115.8, 116.1, 116.3, 116.4, 116.6, 119.2, 119.3, 119.4, 120.1, 130.0, 130.1, 130.5, 131.1, 131.2, 137.2, 139.7, 149.9, 152.3, 153.4, 155.8, 158.4, 160.6, 161.9, 164.3, 172.1. HRMS (m/z): $[M + H]^+$, calcd for $C_{21}H_{15}FN_3O_2$, 360.1143; found, 360.1163.</p>
	<p>(E)-3-(3-hydroxyphenyl)-2-(4-chlorostyryl)pyrido[4,3-d]pyrimidin-4(3H)-one (65). This compound was prepared according to the procedure for 5 in 29 % yield (0.087g) as acetate salt. 1H (400 MHz, $DMSO-d_6$) δ 1.91 (s, 3H), 6.39 (d, $J = 15.6$ Hz, 1H), 6.88 (m, 2H), 6.97 (m, 1H), 7.39 (m, 5H), 7.63 (m, 1H), 7.98 (d, $J = 15.2$ Hz, 1H), 8.85 (d, $J = 5.6$ Hz, 1H), 9.26 (s, 1H), 9.97 (s, 1H), 11.98 (s, 1H). ^{13}C NMR (100 MHz, $DMSO-d_6$) δ 21.1, 115.7, 116.4, 116.6, 119.1, 120.1, 120.2, 129.2, 129.4, 130.5, 133.4, 134.8, 137.2, 139.5, 149.9, 152.2, 153.4, 155.7, 158.4, 160.5, 172.0. HRMS (m/z): $[M + H]^+$, calcd for $C_{21}H_{15}ClN_3O_2$, 376.0847; found, 376.0863.</p>

	<p>(E)-3-(3-hydroxyphenyl)-2-(4-cyanostyryl)pyrido[4,3-d]pyrimidin-4(3H)-one (66). This compound was prepared according to the procedure for 5 in 45% yield (0.15 g) as acetate salt. ¹H (400 MHz, DMSO-<i>d</i>₆) δ 1.91 (s, 3H), 6.52 (d, <i>J</i> = 15.6 Hz, 1H), 6.88 (m, 2H), 6.97 (m, 1H), 7.41 (t, <i>J</i> = 8.4 Hz, 1H), 7.58 (d, <i>J</i> = 8.4 Hz, 1H), 7.65 (t, <i>J</i> = 5.6 Hz, 1H), 7.84 (d, <i>J</i> = 8.4 Hz, 1H), 8.00 (d, <i>J</i> = 15.6 Hz, 1H), 8.87 (d, <i>J</i> = 5.6 Hz, 1H), 9.26 (s, 1H), 9.98 (s, 1H), 11.98 (s, 1H). ¹³C NMR (100 MHz, DMSO-<i>d</i>₆) δ 21.0, 111.9, 115.7, 116.5, 116.7, 118.5, 119.1, 120.2, 122.9, 128.3, 130.5, 132.9, 137.0, 138.7, 138.9, 149.9, 152.1, 153.5, 155.3, 158.4, 160.5, 172.0. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₅N₄O₂, 367.1190; found, 367.1177.</p>
	<p>(E)-3-(3-carboxyphenyl)-2-(4-fluorostyryl)pyrido[4,3-d]pyrimidin-4(3H)-one (67). This compound was prepared according to the procedure described previously in 27% yield (0.12 g). ¹H (400 MHz, DMSO-<i>d</i>₆) δ 6.26 (d, <i>J</i> = 15.2 Hz, 1H), 7.20 (t, <i>J</i> = 8.8 Hz, 2H), 7.48 (m, 2H), 7.65 (d, <i>J</i> = 6.0 Hz, 1H), 7.75 (m, 2H), 8.00 (d, <i>J</i> = 15.6 Hz, 1H), 8.08 (m, 1H), 8.14 (m, 1H), 8.86 (d, <i>J</i> = 5.6 Hz, 1H), 9.25 (s, 1H), 13.27 (s, br, 1H). ¹³C NMR (100 MHz, DMSO-<i>d</i>₆) δ 116.0, 116.3, 116.4, 119.2, 119.3, 120.2, 129.9, 130.1, 130.2, 130.3, 131.0, 131.1, 132.4, 133.3, 136.6, 140.1, 149.8, 152.3, 153.5, 155.7, 160.8, 161.9, 164.4, 166.5. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₅FN₃O₃, 388.1092; found, 388.1075.</p>
	<p>(E)-3-(3-carboxyphenyl)-2-(4-chlorostyryl)pyrido[4,3-d]pyrimidin-4(3H)-one (68). This compound was prepared according to the procedure described previously in 23 % yield (0.093 g). ¹H (400 MHz, DMSO-<i>d</i>₆) δ 6.33 (d, <i>J</i> = 15.2 Hz, 1H), 7.42 (m, 4H), 7.66 (d, <i>J</i> = 5.6 Hz, 1H), 7.75 (m, 2H), 7.99 (d, <i>J</i> = 15.6 Hz, 1H), 8.08 (d, <i>J</i> = 2.0 Hz, 1H), 8.13 (m, 1H), 8.87 (d, <i>J</i> = 5.6 Hz, 1H), 9.27 (s, 1H). 13.31 (s, br, 1H). ¹³C NMR (100 MHz, DMSO-<i>d</i>₆) δ 116.5, 120.2, 120.3, 129.1, 129.7, 129.9, 130.1, 130.3, 132.4, 133.3, 133.4, 134.8, 136.6, 139.9, 149.8, 152.3, 153.6, 155.6, 160.9, 166.5. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₅ClN₃O₃, 404.0796; found, 404.0769.</p>
	<p>(E)-3-(3-carboxyphenyl)-2-(4-cyanostyryl)pyrido[4,3-d]pyrimidin-4(3H)-one (69). This compound was prepared according to the procedure described previously in 31% yield (0.12 g). ¹H (400 MHz, DMSO-<i>d</i>₆) δ 6.49 (d, <i>J</i> = 15.6 Hz, 1H), 7.63 (d, <i>J</i> = 8.4 Hz, 2H), 7.68 (d, <i>J</i> = 5.6 Hz, 1H), 7.74 (m, 2H), 7.82 (d, <i>J</i> = 8.0 Hz, 2H), 8.02 (d, <i>J</i> = 15.6 Hz, 1H), 8.09 (s, 1H), 8.13 (m, 1H), 8.89 (d, <i>J</i> = 5.6 Hz, 1H), 9.28 (s, 1H), 13.31 (s, br, 1H). ¹³C NMR (100 MHz, DMSO-<i>d</i>₆) δ 2.0, 116.6, 118.5, 120.3, 122.9, 128.6, 129.9, 130.1, 130.4, 132.4, 132.9, 133.3, 136.4, 138.9, 139.1, 149.9, 152.2, 153.7, 155.3, 160.8, 166.5. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₃H₁₅N₄O₃, 395.1139; found,</p>

	395.1143.
	<p>(E)-3-(3-carbomylphenyl)-2-(4-fluorostyryl)pyrido[4,3-d]pyrimidin-4(3H)-one (70). This compound was prepared according to the procedure for 72 from compound 67 (0.12 g, 0.31 mmol) in 69 % yield (0.083 g). ¹H (400 MHz, DMSO-<i>d</i>₆) δ 6.28 (d, <i>J</i> = 15.6 Hz, 1H), 7.21 (t, <i>J</i> = 8.8 Hz, 2H), 7.47 (m, 2H), 7.56 (s, br, 1H), 7.65 (m, 3H), 7.98 (s, br, 1H), 8.01 (d, <i>J</i> = 15.6 Hz, 1H), 8.08 (m, 2H), 8.87 (5.6 Hz, 1H), 9.27 (s, 1H). ¹³C NMR (100 MHz, DMSO-<i>d</i>₆) δ 116.1, 116.2, 116.3, 119.3, 120.2, 128.1, 128.4, 129.8, 130.2, 130.3, 131.0, 131.1, 131.6, 135.8, 136.3, 140.1, 149.9, 152.3, 153.6, 155.8, 160.8, 161.9, 164.4, 166.7. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₆FN₄O₂, 387.1252; found, 387.1262.</p>
	<p>(E)-3-(3-carbomylphenyl)-2-(4-chlorostyryl)pyrido[4,3-d]pyrimidin-4(3H)-one (71). This compound was prepared according to the procedure for 72 from compound 68 (0.125 g, 0.31 mmol) in 68% yield (0.085 g). ¹H (400 MHz, DMSO-<i>d</i>₆) 6.34 (d, <i>J</i> = 15.2 Hz, 1H), 7.41 (m, 4H), 7.55 (s, br, 1H), 7.66 (m, 3H), 7.98 (s, br, 1H), 8.00 (d, <i>J</i> = 15.6 Hz, 2H), 8.08 (m, 2H), 8.88 (d, <i>J</i> = 5.6 Hz, 1H), 9.28 (s, 1H). ¹³C NMR (100 MHz, DMSO-<i>d</i>₆) δ 116.3, 120.1, 120.2, 128.2, 128.4, 129.1, 129.6, 129.8, 131.6, 133.3, 134.8, 135.8, 136.3, 139.9, 149.8, 152.2, 153.6, 155.6, 160.7, 166.7. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₂H₁₆ClN₄O₂, 403.0956; found, 403.0934.</p>
	<p>(E)-3-(3-carbomylphenyl)-2-(4-cyanostyryl)pyrido[4,3-d]pyrimidin-4(3H)-one (72). Compound 69 (0.10 g, 0.26 mmol) was dissolved in 6 mL DMF, to which EDCI (0.082 g, 0.42 mmol), HOAt (0.035 g, 0.26 mmol), and NH₄Cl (0.069 g, 1.29 mmol) were added. The reaction was stirred at room temperature for 14 h and 100 mL EtOAc was added to the mixture and washed three times with saturated NH₄Cl. The organic phase was dried over Na₂SO₄, concentrated <i>in vacuo</i>, and purified by silica gel column chromatography to give the product in 77% yield (0.078 g). ¹H (400 MHz, DMSO-<i>d</i>₆) δ 6.49 (d, <i>J</i> = 15.6 Hz, 1H), 7.56 (s, br, 1H), 7.60 (d, <i>J</i> = 8.0 Hz, 1H), 7.67 (m, 3H), 7.82 (d, <i>J</i> = 8.4 Hz, 2H), 7.99 (s, br, 1H), 8.02 (d, <i>J</i> = 15.6 Hz, 1H), 8.04 (m, 2H), 8.89 (d, <i>J</i> = 5.6 Hz, 1H), 9.28 (s, 1H). ¹³C NMR (100 MHz, DMSO-<i>d</i>₆) δ 112.0, 116.4, 118.5, 120.3, 122.9, 128.1, 128.5, 129.8, 131.6, 132.9, 135.8, 136.1, 138.9, 139.1, 149.9, 152.1, 153.7, 155.3, 160.7, 166.7. HRMS (<i>m/z</i>): [M + H]⁺, calcd for C₂₃H₁₆N₅O₂, 394.1299; found, 394.1318.</p>

	<p>(E)-6-bromo-3-(3-hydroxyphenyl)-2-(4-nitrostyryl)quinazolin-4(3H)-one (73). This compound was prepared according to the procedure for 5 in 43% yield (0.088 g). ^1H (300 MHz, DMSO-d_6) δ 6.49 (d, $J = 15.55$ Hz, 1H), 6.87 (m, 2H), 6.99 (d, $J = 7.65$ Hz, 1H), 7.42 (t, $J = 8.13$ Hz, 1H), 7.58 (d, $J = 8.61$ Hz, 2H), 7.70 (d, $J = 8.61$ Hz, 1H), 7.86 (d, $J = 6.46$ Hz, 1H), 7.93 (d, $J = 15.79$ Hz, 1H), 8.00 (d, $J = 8.61$ Hz, 1H), 8.14 (s, 1H), 8.16 (d, $J = 8.61$ Hz, 2H). HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{22}\text{H}_{15}\text{BrN}_3\text{O}_4$ 464.0240; found, 464.0267.</p>
	<p>(E)-2-(4-fluorostyryl)-6-hydroxy-3-(3-hydroxyphenyl)quinazolin-4(3H)-one (74). This compound was prepared according to the procedure for 5 in 42% yield (0.37 g). ^1H (500 MHz, DMSO-d_6) δ 6.29 (d, $J = 15.55$ Hz, 1H), 6.79 (m, 2H), 6.96 (d, $J = 6.78$ Hz, 1H), 7.21 (t, $J = 8.37$ Hz, 2H), 7.32 (d, $J = 8.57$ Hz, 1H), 7.41 (m, 5H), 7.64 (d, $J = 8.57$ Hz, 1H), 7.73 (d, $J = 15.95$ Hz, 1H), 9.91 (s, 1H), 10.16 (s, 1H). HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{22}\text{H}_{16}\text{FN}_2\text{O}_3$, 375.1139; found, 375.1152.</p>
	<p>(E)-2-(4-cyanostyryl)-7-nitro-3-(3-carboxyphenyl)quinazolin-4(3H)-one (75). The compound was prepared according to the procedure for 2 described previously in 75% yield (0.328 g). ^1H (600 MHz, DMSO-d_6) δ 6.48 (d, $J = 15$ Hz, 1H), 7.60 (d, $J = 8.4$ Hz, 2H), 7.77 (m, 4H), 8.01 (d, $J = 15.6$ Hz, 1H), 8.11 (s, 1H), 8.15 (m, 1H), 8.24 (dd, $J = 9.0, 2.4$ Hz, 1H), 8.33 (d, $J = 9$ Hz, 1H), 8.46 (d, $J = 2.4$ Hz, 1H), 13.3 (s, 1H). ^{13}C NMR (151 MHz, DMSO-d_6) δ 111.88, 118.52, 120.28, 122.11, 122.73, 125.12, 128.46 (2C), 128.78, 129.85, 130.17, 130.39, 132.47, 132.85 (2C), 133.22, 136.53, 138.48, 138.91, 147.61, 151.39, 152.92, 160.49, 166.45. HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{24}\text{H}_{15}\text{N}_4\text{O}_5$, 439.1037; found, 439.1016.</p>
	<p>(E)-2-(4-cyanostyryl)-7-carboxy-3-(3-carboxyphenyl)quinazolin-4(3H)-one (76). The compound was prepared following a modified procedure of the previously reported method (328 mg, 75% yield).¹ Triethylamine (0.81 mL, 5.8 mmol) was added to a suspension of 4-carboxyanthranilic acid (1 g, 5.8 mmol) in toluene (50 mL) and stirred for 20 minutes at 70 °C. Subsequently triethyl orthoacetate (1.8 mL, 9.8 mmol) was added and the resulting mixture was refluxed for 16 h. The product was crystallized by cooling the reaction on ice for 4 h, filtered from the reaction mixture and washed liberally with hexanes to give the product in 63% yield (0.753 g). ^1H (600 MHz, DMSO-d_6) δ 6.48 (d, $J = 15.6$ Hz, 1H), 7.60 (d, $J = 8.4$ Hz, 2H), 7.75 (m, 2H), 7.80 (d, $J = 8.4$ Hz, 2H), 7.92 (d, $J = 15$ Hz, 1H), 8.02 (dd, $J = 8.4, 1.8$ Hz, 1H), 8.07 (m, 1H), 8.14 (m,</p>

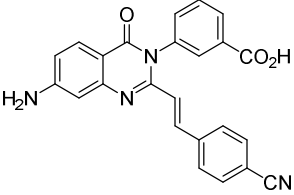
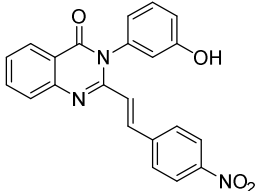
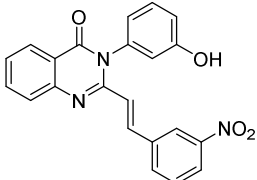
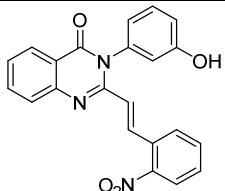
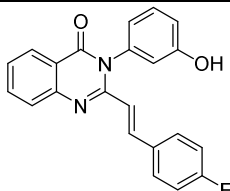
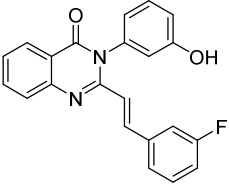
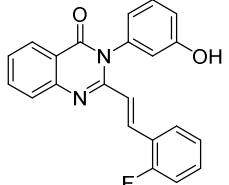
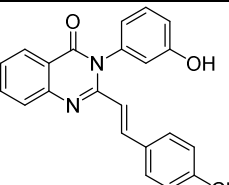
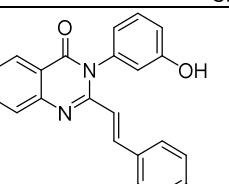
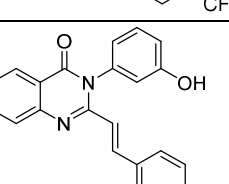
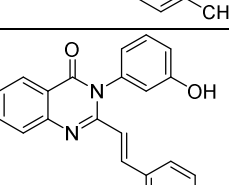
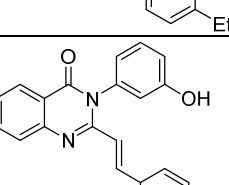
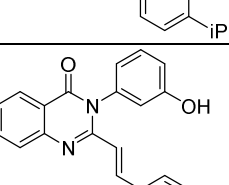
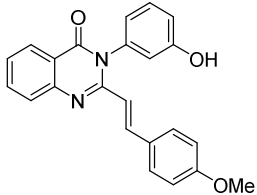
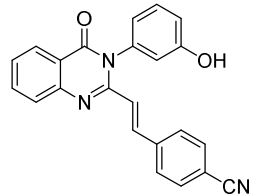
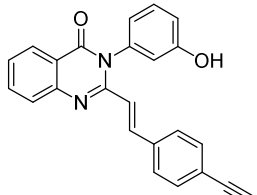
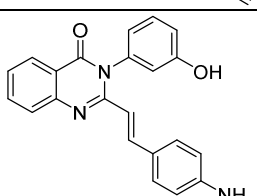
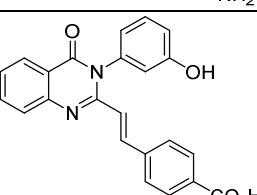
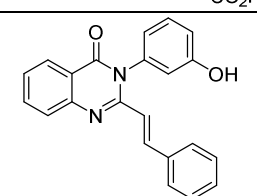
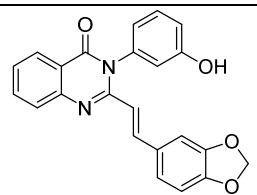
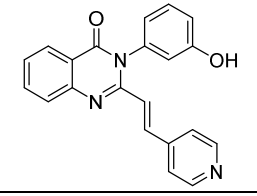
	1H), 8.23 (d, $J = 8.4$ Hz, 1H), 8.27 (d, $J = 1.2$ Hz, 1H), 13.41 (s, 2H). ^{13}C NMR (150 MHz, DMSO- d_6) δ 112.11, 119.02, 123.52, 124.07, 126.98, 127.55, 128.79 (2C), 128.85, 130.37, 130.54, 130.68, 132.83, 133.28 (2C), 133.79, 136.84, 137.27, 138.05, 139.57, 147.54, 152.02, 161.37, 166.89, 166.94. HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{25}\text{H}_{16}\text{N}_3\text{O}_5$, 438.1084; found, 438.1061.
	<p>(E)-2-(4-cyanostyryl)-7-amino-3-(3-carboxyphenyl)quinazolin-4(3H)-one (77). This compound was prepared according to the procedure for 41 from compound 75 in 67 % yield (0.124 g). ^1H (600 MHz, DMSO-d_6) δ 6.42 (d, $J = 15.6$ Hz, 1H), 6.83 (d, $J = 8.4$ Hz, 1H), 6.87 (s, 1H), 7.58 (m, 2H), 7.68 (m, 2H), 7.78 (m, 3H), 7.86 (d, $J = 15.6$ Hz, 1H), 7.91 (s, 1H), 8.08 (d, $J = 7.8$ Hz, 1H). ^{13}C NMR (150 MHz, DMSO-d_6) δ 106.66, 109.17, 111.55, 115.73, 118.69, 123.18, 127.94, 128.35 (2C), 129.77, 129.95, 129.99, 130.09, 132.25, 132.89 (2C), 133.36, 137.03, 137.15, 139.37, 150.99, 154.82, 160.38, 166.52. HRMS (m/z): $[\text{M} + \text{H}]^+$, calcd for $\text{C}_{24}\text{H}_{17}\text{N}_4\text{O}_3$, 409.1295; found, 409.1324.</p>

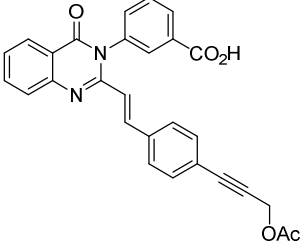
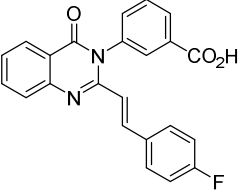
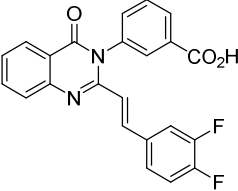
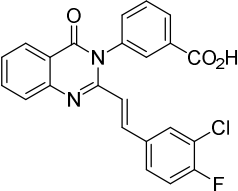
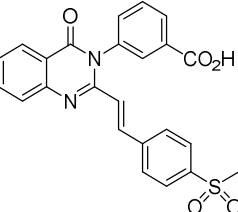
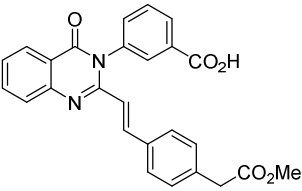
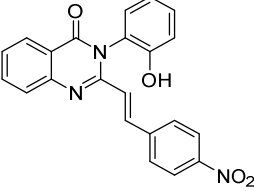
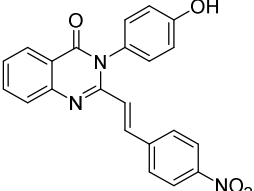
Table S2. MICs of reported compounds against *S. aureus* and *E. faecium*

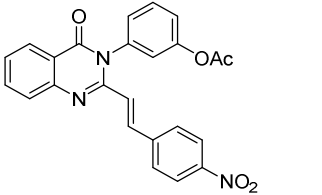
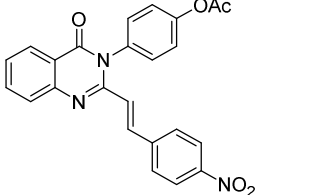
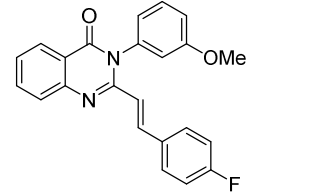
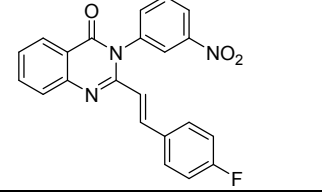
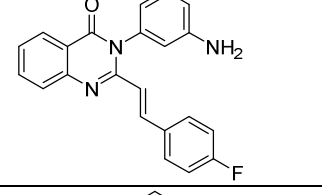
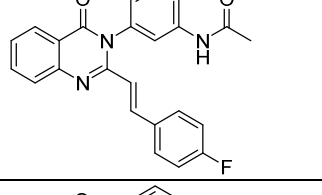
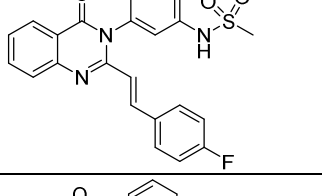
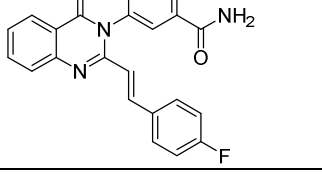
Structure	Number	MIC ($\mu\text{g/mL}$) <i>S. aureus</i> ATCC 29213	MIC ($\mu\text{g/mL}$) <i>E. faecium</i> NCTC 7171
	1	2	32
	2	>128	8
	3	128	64
	4	0.25	>128

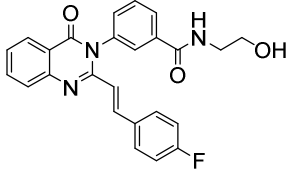
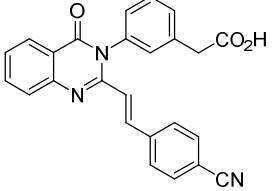
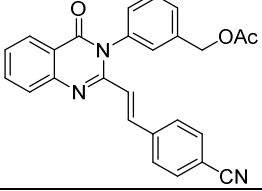
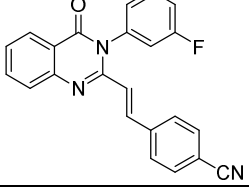
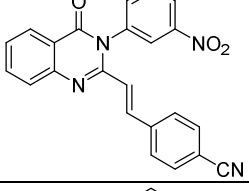
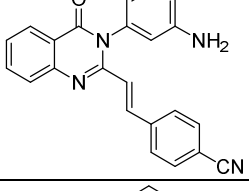
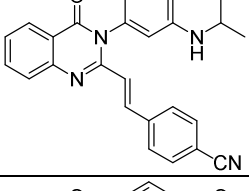
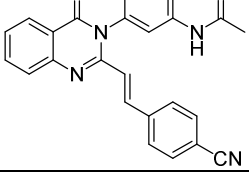
	5	1	128
	6	1	32
	7	0.125	32
	8	128	16
	9	0.25	32
	10	0.5	>128
	11	>128	>128
	12	>128	>128

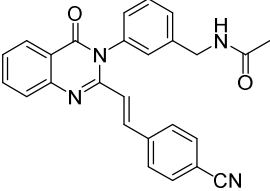
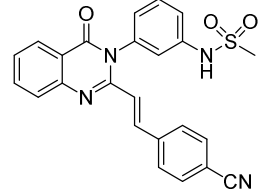
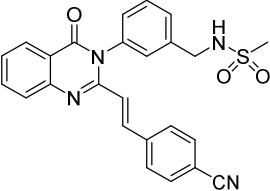
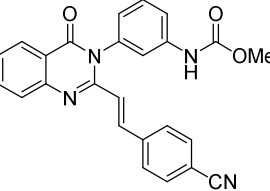
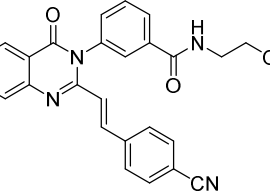
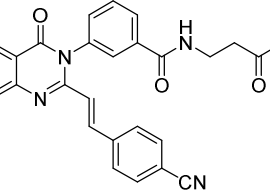
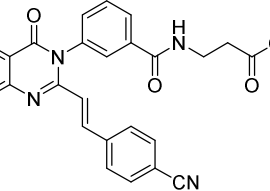
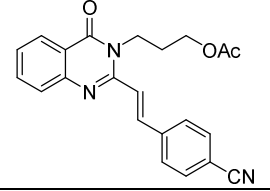
	13	4	>128
	14	0.03	64
	15	0.003	16
	16	16	32
	17	>128	>128
	18	0.5	64
	19	2	32
	20	2	>128

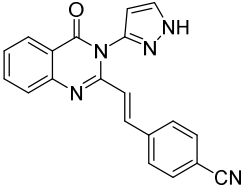
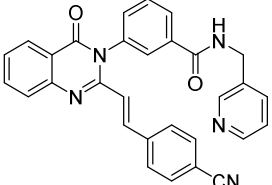
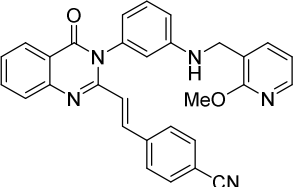
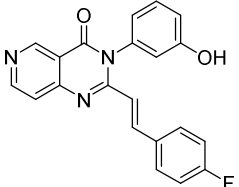
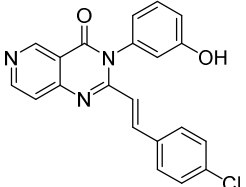
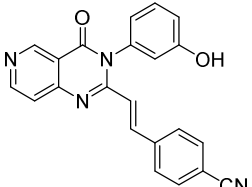
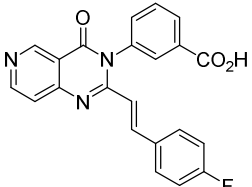
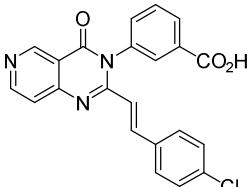
	21	128	128
	22	0.06	>128
	23	0.06	>128
	24	32	>128
	25	128	128
	26	2	>128
	27	0.03	>128
	28	2	>128

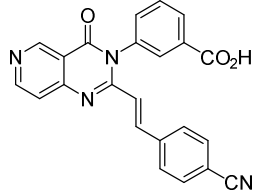
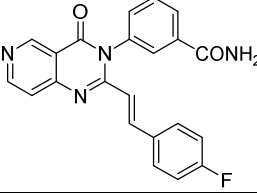
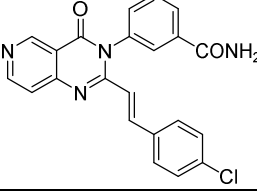
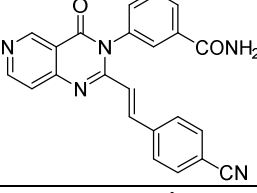
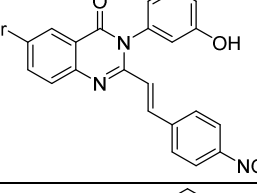
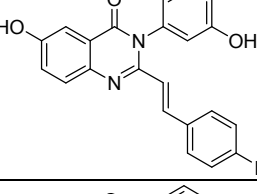
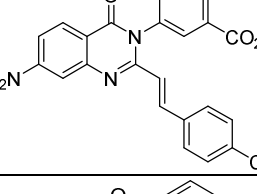
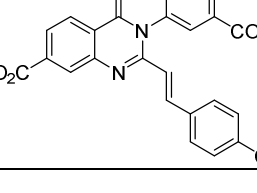
	29	0.25	>128
	30	1	>128
	31	2	>128
	32	2	>128
	33	>128	>128
	34	8	>128
	35	2	>128
	36	>128	64

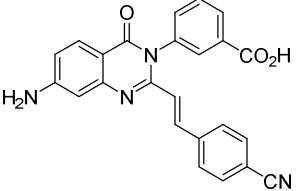
	37	>128	64
	38	>128	32
	39	>128	>128
	40	>128	>128
	41	2	128
	42	1	>128
	43	0.125	>128
	44	4	>128

	45	2	>128
	46	8	>128
	47	0.03	>128
	48	>128	>128
	49	>128	>128
	50	0.5	>128
	51	0.01	>128
	52	0.25	>128

	53	0.015	>128
	54	0.004	>128
	55	4	>128
	56	0.5	>128
	57	1	>128
	58	32	>128
	59	8	>128
	60	>128	>128

	61	0.5	>128
	62	8	>128
	63	2	>128
	64	1	>128
	65	1	>128
	66	0.5	128
	67	64	>128
	68	64	>128

	69	>128	>128
	70	>128	>128
	71	>128	>128
	72	>128	>128
	73	>128	>128
	74	32	32
	75	64	>128
	76	128	>128

	77	32	>128
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