Supporting Information

Cobalt catalyzed electrochemical [4+2] annulation for the synthesis of sultams

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General Information

The instrument for electrolysis was dual display potentiostat (DJS-292B) (made in China). The electrodes were purchased from Gaoss Union. Thin layer chromatography (TLC) employed glass 0.25 mm silica gel plates. Flash chromatography columns were packed with 200-300 mesh silica gel in petroleum (boiling point is between 60-90 °C). Gradient flash chromatography was conducted eluting with a continuous gradient from petroleum to the indicated solvent, and they are listed as volume/volume ratios. EPR spectra were recorded on a Bruker X-band A-200 spectrometer. NMR spectra were recorded on a Bruker spectrometer at 400 MHz (¹H NMR), 100 MHz (¹³C NMR), 376 MHz (¹⁹F NMR). Tetramethylsilane was used as an internal standard. All ¹H, ¹³C and ¹⁹F NMR data spectra were reported in delta (δ) units, parts per million (ppm) downfield from the internal standard. Coupling constants are reported in Hertz (Hz). High resolution mass spectra (HRMS) were measured with a Waters Micromass GCT instrument, accurate masses are reported for the molecular ion + sodium ([M+Na]⁺).

Experimental procedure

General procedure for the preparation of the sulfonamide derivatives:¹



Unless stated the standard synthesis is as follows: To a solution of 8-aminoquinoline (3.5 mmol) in pyridine (7.0 mL) was added sulfonyl chloride (3.5 mmol) and the reaction was heated to 130 °C for 30 minutes. After this time the mixture was left to cool to 70 °C and poured over 50 mL of cold water. The resulting suspension was then filtered and washed with cold water (ca.100 mL) until the filtrate became clear. The solid was then dried under vacuum and recrystallized from toluene to afford a pale white/yellow crystalline solid.

General procedure for electrochemical oxidative annulation of sulfonamides with alkynes:

In an oven-dried undivided three-necked bottle equipped with a stir bar, sulfonamides (0.25 mmol), $Co(OAc)_2$ '4H₂O (20 mol%) and NaOAc (0.5 mmol) were combined and added. The bottle was equipped with carbon cloth (15 mm × 20 mm × 0.33 mm) as the anode and stainless steel plate (15 mm × 20 mm × 1 mm) as the cathode. Under air conditions (open to air by four thick syringe needles), alkynes (0.5 mmol), HOAc (1.0 mL), and EtOH (10 mL) were injected respectively into the tubes via syringes. The reaction mixture was **strong** stirred and electrolyzed at a constant current of 4 mA at 75 °C for 8 h. When the reaction was finished, the pure product was obtained by flash column chromatography on silica gel.

Experimental setup:



Figure S1. The experimental setup for electrolysis.

Procedure for gram-scale reaction:

undivided In an oven-dried three-necked bottle equipped with stir а bar. 4-methyl-N-(quinolin-8-yl)benzenesulfonamide (1.942 g, 5.0 mmol), phenylacetylene (1.12 mL, 10 mmol), Co(OAc)₂ 4H₂O (254 mg, 1.0 mmol), NaOAc (820 mg, 10 mmol), were combined and added. The bottle was equipped with two carbon cloths (20 mm \times 30 mm \times 0.33 mm) as the anodes and two stainless steel plates (20 mm \times 30 mm \times 1 mm) as the cathodes. Under air conditions (air bubbling), EtOH (115 mL) and HOAc (11.5 mL) were injected respectively into the tubes via syringes. The reaction mixture was strong stirred and electrolyzed under an air atmosphere (air bubbling) at a constant current of 8 mA at 75 °C for 40 h. When the reaction was finished, the pure product was obtained by flash column chromatography on silica gel.



Figure S2. The experimental setup for gram-scale reaction.

Procedure for cyclic voltammetry (CV):

Cyclic voltammetry was performed in a three-electrode cell connected to a schlenk line under nitrogen at room temperature. The working electrode was a steady glassy carbon disk electrode while the counter electrode was a platinum wire. The reference was an Ag/AgCl electrode submerged in saturated aqueous KCl solution. 11 mL of ethanol containing 0.2 mmol of $^{n}Bu_{4}NBF_{4}$ were poured into the electrochemical cell in all experiments. The scan rate is 0.05 V/s, ranging from 0 V to 2.2 V.



Figure S3. Cyclic voltammograms: 1, 0.1 mmol, Co(OAc) 4H₂O, 0.1 mmol, NaOAc, 0.2 mmol.

Procedure for electron paramagnetic resonance (EPR) experiments:

In an oven-dried undivided three-necked bottle equipped with a stir bar, **1** (0.25 mmol), $Co(OAc)_2 H_2O$ (20 mol%) and NaOAc (0.5 mmol) were combined and added. The bottle was equipped with carbon cloth (15 mm × 20 mm × 0.33 mm) as the anode and stainless steel plate (15 mm × 20 mm × 1 mm) as the cathode. Under air conditions (open to air by four thick syringe needles), **2** (0.5 mmol), HOAc (0.5 mL), and EtOH (10.5 mL) were injected respectively into the tubes via syringes. The reaction mixture was **strong** stirred and electrolyzed at a constant current of 4 mA at 75 °C for 15min. When the reaction was finished, the solution sample was taken out into a small tube and analyzed by EPR. After fitting, we proposed that this radical signal belongs to the alkoxy radical (g = 2.0068, $A_N = 13.58$ G, $A_H = 7.48$ G).

In an oven-dried undivided three-necked bottle equipped with a stir bar, **1** (0.25 mmol), $Co(OAc)_2$ 4H₂O (20 mol%) and NaOAc (0.5 mmol) were combined and added. The bottle was equipped with carbon cloth (15 mm × 20 mm × 0.33 mm) as the anode and stainless steel plate (15 mm × 20 mm × 1 mm) as the cathode. Under air conditions (open to air by four thick syringe needles), **2** (0.5 mmol), HOAc (0.5 mL), and MeCN (10.5 mL) were injected respectively into the tubes via syringes. The reaction mixture was **strong** stirred and electrolyzed at a constant current of 4 mA at 75 °C for 15min. When the reaction was finished, the solution sample was taken out into a small tube and analyzed by EPR. After fitting, we proposed that this radical signal belongs to the superoxide radical anion (g = 2.0068, A_N = 12.9 G, A_H = 10.7 G).

Detailed descriptions for products



6-Methyl-3-phenyl-2-(quinolin-8-yl)-2H-benzo[*e*][1,2]thiazine 1,1-dioxide (3).¹

White solid was obtained in 80% isolated yield (80 mg). ¹H NMR (400 MHz, DMSO-*d*₆) δ 9.06 (d, *J* = 2.68 Hz, 1H), 8.34 (d, *J* = 8.16 Hz, 1H), 7.86 (d, *J* = 8.18 Hz, 1H), 7.82 – 7.73 (m, 2H), 7.67 (s, 1H), 7.64 – 7.53 (m, 2H), 7.46 – 7.34 (m, 2H), 7.31 (s, 1H), 7.26 – 7.12 (m, 4H), 2.48 (s, 3H). ¹³C NMR (100 MHz, DMSO-*d*₆) δ 151.17, 145.01, 143.40, 142.80, 136.18, 135.15, 134.74, 132.83, 129.42, 129.37, 129.09, 128.95, 128.72, 128.40, 128.32, 127.23, 125.99, 122.06, 121.98, 112.87, 112.85, 21.25.



6-Methyl-2-(quinolin-8-yl)-3-(p-tolyl)-2H-benzo[e][1,2]thiazine 1,1-dioxide (4).¹

White solid was obtained in 73% isolated yield (75.3 mg). ¹H NMR (400 MHz, CDCl₃) δ 9.07 (dd, *J* = 4.06, 1.40 Hz, 1H), 8.02 (dd, *J* = 8.28, 1.30 Hz, 1H), 7.69 – 7.57 (m, 4H), 7.41 – 7.35 (m, 2H), 7.27 – 7.20 (m, 2H), 7.20 – 7.16 (m, 1H), 6.91 (d, *J* = 8.03 Hz, 2H), 6.88 (s, 1H), 2.47 (s, 3H), 2.15 (s, 3H). ¹³C NMR (100 MHz, CDCl₃) δ 150.98, 145.72, 144.35, 142.53, 139.16, 135.60, 135.42, 133.26, 132.78, 129.21, 128.92, 128.71, 128.56, 128.53, 127.64, 127.59, 125.56, 122.56, 121.59, 111.85, 111.83, 21.64, 21.12.



3-(4-Fluorophenyl)-6-methyl-2-(quinolin-8-yl)-2H-benzo[*e*][**1,2**]**thiazine 1,1-dioxide**(**5**).² White solid was obtained in 73% isolated yield (75.6 mg). ¹H NMR (400 MHz, DMSO-*d*₆) δ 9.06 (d, *J* = 2.64 Hz, 1H), 8.36 (d, *J* = 6.58 Hz, 1H), 7.89 (d, *J* = 7.88 Hz, 1H), 7.86 – 7.77 (m, 2H), 7.66 (s, 1H), 7.64 – 7.56 (m, 2H), 7.47 – 7.35 (m, 2H), 7.31 (s, 1H), 7.21 (d, J = 6.96 Hz, 1H), 7.05 (t, J = 8.76 Hz, 2H), 2.49 (s, 3H). ¹³C NMR (100 MHz, DMSO- d_6) δ 163.00 (d, ¹ $J_{C-F} = 247.4$ Hz), 151.71, 145.39, 143.31, 142.76, 136.69, 135.00, 133.20, 132.08 (d, ⁴ $J_{C-F} = 3.0$ Hz), 129.94, 129.86 (d, ³ $J_{C-F} = 8.6$ Hz), 129.65, 129.31, 129.19, 128.84, 128.78, 126.48, 122.57, 122.46, 115.91 (d, ² $J_{C-F} = 21.9$ Hz).113.32, 21.68. ¹⁹F NMR (376 MHz, DMSO- d_6) δ -111.51.



3-(3-Fluorophenyl)-6-methyl-2-(quinolin-8-yl)-2H-benzo[*e*][1,2]thiazine 1,1-dioxide(6).

White solid was obtained in 77% isolated yield (79.6 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 9.10 (dd, J = 4.20, 1.72 Hz, 1H), 8.37 (dd, J = 8.34, 1.74 Hz, 1H), 7.89 (d, J = 8.12 Hz, 1H), 7.70 – 7.57 (m, 5H), 7.49 – 7.42 (m, 2H), 7.38 (t, J = 7.84 Hz, 1H), 7.29 – 7.20 (m, 1H), 7.16 (d, J = 7.36 Hz, 1H), 7.08 – 6.99 (m, 1H), 2.48 (s, 3H). ¹³C NMR (100 MHz, DMSO- d_6) δ 161.95 (d, ¹ $J_{C-F} = 243.9$ Hz), 151.31, 145.01, 142.98, 141.92 (d, ⁴ $J_{C-F} = 2.6$ Hz) , 137.61, 137.53, 136.31, 134.59, 132.51, 130.51 (d, ³ $J_{C-F} = 8.4$ Hz), 129.88, 129.32, 129.13, 128.80, 128.60, 128.15, 126.10, 123.19 (d, J = 2.6 Hz), 122.16 (d, ³ $J_{C-F} = 7.0$ Hz), 116.22 (d, ² $J_{C-F} = 20.9$ Hz), 114.10, 113.86 (d, ² $J_{C-F} = 23.1$ Hz), 21.24. ¹⁹F NMR (376 MHz, DMSO- d_6) δ -112.90. HRMS (ESI): calculated for C₂₄H₁₇FN₂NaO₂S [M+Na]⁺: 439.0887; Found: 439.0894.



3-(4-Bromophenyl)-6-methyl-2-(quinolin-8-yl)-2H-benzo[*e*][1,2]thiazine 1,1-dioxide(7).³

White solid was obtained in 81% isolated yield (97.1 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 9.09 (d, J = 4.10 Hz, 1H), 8.41 (d, J = 7.92 Hz, 1H), 7.93 (d, J = 8.22 Hz, 1H), 7.82 – 7.69 (m, 3H), 7.65 (d, J = 8.44 Hz, 2H), 7.53 – 7.35 (m, 5H), 7.24 (d, J = 6.96 Hz, 1H), 2.53 (s, 3H). ¹³C NMR (100 MHz, DMSO- d_6) δ 151.32, 144.95, 142.97, 142.31, 136.30, 134.51, 134.46, 132.69, 131.48, 129.73, 129.32, 129.23, 129.05, 128.80, 128.49, 126.09, 122.87, 122.20, 122.08, 113.40, 21.28.



6-Methyl-2-(quinolin-8-yl)-3-(4-(trifluoromethyl)phenyl)-2H-benzo[*e*][1,2]thiazine 1,1-dioxide(8).¹

White solid was obtained in 80% isolated yield (93.6 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 9.06 (d, J = 2.42 Hz, 1H), 8.36 (d, J = 6.58 Hz, 1H), 7.89 (d, J = 7.88 Hz, 1H), 7.86 – 7.77 (m, 2H), 7.66 (s, 1H), 7.64 – 7.56 (m, 2H), 7.47 – 7.36 (m, 2H), 7.31 (s, 1H), 7.21 (d, J = 6.96 Hz, 1H), 7.05 (t, J = 8.76 Hz, 2H), 2.49 (s, 3H). ¹³C NMR (100 MHz, CD₂Cl₂) δ 151.74, 146.08, 143.82, 143.38, 139.71, 136.52, 135.50, 133.20, 131.13 (q, ² $J_{C-F} = 32.4$ Hz), 130.23, 129.68, 129.13, 128.83, 128.50, 126.21, 125.80 (q, ³ $J_{C-F} = 3.8$ Hz), 124.49 (q, ¹ $J_{C-F} = 272.3$ Hz), 122.98, 122.50, 114.70, 114.67, 21.98. ¹⁹F NMR (376 MHz, DMSO- d_6) δ -61.25.



3-(4-Eethoxyphenyl)-6-methyl-2-(quinolin-8-yl)-2H-benzo[*e*][**1,2**]**thiazine 1,1-dioxide (9).**¹ White solid was obtained in 73% isolated yield (77.9 mg). ¹H NMR (400 MHz, DMSO-*d*₆) δ 9.09 (d, *J* = 2.92 Hz, 1H), 8.37 (d, *J* = 8.14 Hz, 1H), 7.88 (d, *J* = 8.08 Hz, 1H), 7.74 (d, *J* = 8.54 Hz, 2H), 7.65 (s, 1H), 7.64 – 7.57 (m, 2H), 7.39 (t, *J* = 7.36 Hz, 2H), 7.23 (s, 1H), 7.17 (d, *J* = 7.28 Hz, 1H), 6.76 (d, *J* = 8.56 Hz, 2H), 3.64 (s, 3H), 2.49 (s, 3H). ¹³C NMR (100 MHz, DMSO-*d*₆) δ 160.13, 151.16, 145.08, 143.37, 142.73, 136.21, 136.15, 134.96, 133.11, 129.01, 128.75, 128.72, 128.07, 127.55, 126.01, 122.05, 122.00, 113.90, 113.78, 111.48, 111.38, 55.18, 21.29.



3-(Cyclohex-1-en-1-yl)-6-methyl-2-(quinolin-8-yl)-2H-benzo[*e*][**1,2**]**thiazine 1,1-dioxide**(**10**).¹ Yellow solid was obtained in 69% isolated yield (69.5 mg). ¹H NMR (400 MHz, DMSO-*d*₆) δ 9.10 (d, *J* = 3.52 Hz, 1H), 8.44 (d, *J* = 8.42 Hz, 1H), 7.97 (d, *J* = 8.18 Hz, 1H), 7.69 – 7.59 (m, 2H), 7.55 (d, J = 7.92 Hz, 1H), 7.47 (t, J = 7.60 Hz, 1H), 7.39 (d, J = 8.12 Hz, 1H), 7.09 – 6.97 (m, 2H), 6.59 (s, 1H), 2.48 (s, 3H), 2.37 (d, J = 17.76 Hz, 1H), 2.15 (d, J = 17.08 Hz, 1H), 1.95 (d, J = 16.46 Hz, 1H), 1.72 – 1.47 (m, 2H), 1.46 – 1.28 (m, 3H). ¹³C NMR (100 MHz, DMSO- d_6) δ 150.97, 145.23, 145.04, 142.72, 136.17, 135.74, 132.85, 132.12, 131.23, 129.22, 128.88, 128.81, 128.44, 126.56, 126.03, 122.30, 121.94, 111.54, 25.43, 25.04, 21.88, 21.28, 21.18.



3-Butyl-6-methyl-2-(quinolin-8-yl)-2H-benzo[e][1,2]thiazine 1,1-dioxide(11).

Yellow solid was obtained in 85% isolated yield (94.1 mg). ¹H NMR (400 MHz, CDCl₃) δ 8.81 (dd, J = 4.18, 1.76 Hz, 1H), 8.14 (dd, J = 8.30, 1.72 Hz, 1H), 7.85 (dd, J = 8.26, 1.46 Hz, 1H), 7.70 (dd, J = 7.38, 1.42 Hz, 1H), 7.66 (d, J = 8.98 Hz, 1H), 7.61 – 7.49 (m, 1H), 7.36 (dd, J = 8.28, 4.18 Hz, 1H), 7.25 – 7.18 (m, 2H), 6.38 (s, 1H), 2.45 (s, 3H), 2.20 – 2.09 (m, 1H), 2.09 – 1.99 (m, 1H), 1.58 – 1.48 (m, 2H), 1.27 – 1.15 (m, 2H), 0.75 (t, J = 7.32 Hz, 3H). ¹³C NMR (100 MHz, CDCl₃) δ 151.12, 145.68, 144.96, 142.15, 135.79, 133.35, 133.28, 130.75, 129.27, 129.08, 128.77, 127.72, 126.83, 125.73, 121.69, 121.66, 108.53, 33.81, 29.54, 21.77, 21.59, 13.62. HRMS (ESI): calculated for C₂₂H₂₂N₂NaO₂S [M+Na]⁺: 401.1294; Found: 401.1304.



6-Methyl-3-pentyl-2-(quinolin-8-yl)-2H-benzo[e][1,2]thiazine 1,1-dioxide(12).

Yellow solid was obtained in 79% isolated yield (77.1 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 8.78 (d, J = 3.98 Hz, 1H), 8.43 (d, J = 8.32 Hz, 1H), 8.09 (d, J = 7.90 Hz, 1H), 7.72 (d, J = 7.96 Hz, 1H), 7.67 (t, J = 7.78 Hz, 1H), 7.58 (d, J = 8.06 Hz, 1H), 7.54 (dd, J = 8.26, 4.14 Hz, 1H), 7.42 (s, 1H), 7.30 (d, J = 8.02 Hz, 1H), 6.52 (s, 1H), 2.42 (d, J = 2.16 Hz, 3H), 2.11 – 1.99 (m, 1H), 1.98 – 1.87 (m, 1H), 1.39 (t, J = 7.18 Hz, 2H), 1.11 – 1.00 (m, 4H), 0.69 (t, J = 6.12 Hz, 3H). ¹³C NMR (100 MHz, DMSO- d_6) δ 151.36, 144.94, 144.50, 142.35, 136.42, 133.15, 132.42, 130.99, 129.76,

128.89, 128.46, 127.98, 127.00, 126.16, 122.18, 121.13, 108.10, 33.56, 30.18, 26.64, 21.57, 21.19, 13.69. HRMS (ESI): calculated for C₂₃H₂₄N₂NaO₂S [M+Na]⁺: 415.1451; Found: 415.1462.



6-Methyl-3-phenethyl-2-(quinolin-8-yl)-2H-benzo[*e*][1,2]thiazine 1,1-dioxide(13).

Yellow solid was obtained in 89% isolated yield (95.1 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 8.82 (d, J = 2.56 Hz, 1H), 8.45 (d, J = 7.94 Hz, 1H), 8.12 (d, J = 7.38 Hz, 1H), 7.78 (d, J = 7.22 Hz, 1H), 7.69 (t, J = 7.76 Hz, 1H), 7.61 (d, J = 8.02 Hz, 1H), 7.56 (dd, J = 8.34, 4.20 Hz, 1H), 7.40 (s, 1H), 7.32 (d, J = 7.88 Hz, 1H), 7.13 (t, J = 7.06 Hz, 2H), 7.08 (d, J = 6.98 Hz, 1H), 6.91 (d, J = 7.14 Hz, 2H), 6.50 (s, 1H), 2.81 – 2.72 (m, 2H), 2.42 (s, 3H), 2.36 – 2.25 (m, 1H), 2.25 – 2.12 (m, 1H). ¹³C NMR (100 MHz, DMSO- d_6) δ 151.51, 144.98, 143.53, 142.43, 140.54, 136.50, 132.97, 132.43, 131.18, 129.92, 128.93, 128.53, 128.22, 128.16, 127.10, 126.26, 125.95, 122.27, 121.18, 108.42, 36.04, 33.43, 21.18. HRMS (ESI): calculated for C₂₆H₂₂N₂NaO₂S [M+Na]⁺: 449.1294; Found: 449.1305.



3-Cyclopropyl-6-methyl-2-(quinolin-8-yl)-2H-benzo[e][1,2]thiazine 1,1-dioxide(14).²

White solid was obtained in 88% isolated yield (80.1 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 8.77 (dd, J = 4.28, 1.76 Hz, 1H), 8.45 (dd, J = 8.28, 1.76 Hz, 1H), 8.11 (d, J = 7.76 Hz, 1H), 7.81 (d, J = 7.00 Hz, 1H), 7.69 (t, J = 7.78 Hz, 1H), 7.60 (d, J = 8.02 Hz, 1H), 7.55 (dd, J = 8.28, 4.18 Hz, 1H), 7.40 (s, 1H), 7.31 (d, J = 7.88 Hz, 1H), 6.39 (s, 1H), 2.43 (s, 3H), 1.24 – 1.21 (m, 1H), 0.71 – 0.63 (m, 1H), 0.54 – 0.45 (m, 1H), 0.35 – 0.27 (m, 1H), 0.26 – 0.18 (m, 1H). ¹³C NMR (100 MHz, DMSO- d_6) δ 151.27, 146.06, 145.17, 142.25, 136.38, 133.30, 132.94, 131.68, 129.86, 128.88, 128.45, 127.79, 127.03, 126.21, 122.15, 121.06, 104.10, 21.21, 14.79, 7.22, 6.76.



Methyl 6-methyl-2-(quinolin-8-yl)-2H-benzo[*e*][1,2]thiazine-3-carboxylate 1,1-dioxide(15).³ Yellow solid was obtained in 65% isolated yield (61.7 mg). ¹H NMR (400 MHz, DMSO-*d*₆) δ 8.73 (dd, *J* = 4.22, 1.76 Hz, 1H), 8.44 (dd, *J* = 8.32, 1.78 Hz, 1H), 8.06 (dd, *J* = 6.16, 3.54 Hz, 1H), 7.88 (s, 2H), 7.72 (d, *J* = 8.02 Hz, 1H), 7.67 – 7.62 (m, 2H), 7.60 (d, *J* = 8.16 Hz, 1H), 7.55 (dd, *J* = 8.32, 4.18 Hz, 1H), 3.56 (s, 3H), 2.54 (s, 3H). ¹³C NMR (100 MHz, DMSO-*d*₆) δ 162.18, 150.78, 144.52, 143.03, 136.22, 134.32, 132.71, 131.71, 130.98, 130.91, 129.82, 129.53, 129.05, 128.60, 125.96, 121.97, 121.64, 120.53, 52.57, 21.16.



6-Methyl-2-(quinolin-8-yl)-3-(trimethylsilyl)-2H-benzo[e][1,2]thiazine 1,1-dioxide(16).²

White solid was obtained in 51% isolated yield (50.6 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 9.09 (dd, J = 4.18, 1.72 Hz, 1H), 8.70 (dd, J = 8.38, 1.72 Hz, 1H), 8.36 (dd, J = 7.80, 1.92 Hz, 1H), 7.95 – 7.85 (m, 3H), 7.84 – 7.79 (m, 2H), 7.66 (d, J = 8.04 Hz, 1H), 7.17 (s, 1H), 2.72 (s, 3H), 0.00 (s, 9H). ¹³C NMR (100 MHz, DMSO- d_6) δ 151.20, 147.74, 145.64, 142.19, 136.24, 134.78, 132.81, 131.39, 129.88, 129.43, 129.25, 128.88, 127.47, 125.93, 122.14, 121.04, 119.62, 21.23, -1.32.



3-(1-Hydroxyethyl)-6-methyl-2-(quinolin-8-yl)-2H-benzo[*e*][**1,2**]**thiazine 1,1-dioxide(17).** Yellow solid was obtained in 70% isolated yield (64 mg) (a/b = 1.3:1). ¹H NMR (400 MHz, DMSO-*d*₆) δ 9.06 – 8.97 (m, 1H), 8.48 – 8.42 (m, 1H), 8.03 (d, *J* = 8.10 Hz, 1H), 7.62 – 7.58 (m, 2H), 7.55 (d, *J* = 7.34 Hz, 2H), 7.35 (d, *J* = 8.02 Hz, 1H), 7.24 (d, *J* = 7.22 Hz, 1H), 6.97 (s, 1H), 5.43 (d, *J* = 4.96 Hz, 1H), 3.90 – 3.80 (m, 1H), 2.44 (s, 3H), 1.30 – 1.27 (m, 3H). ¹³C NMR (100 MHz, DMSO- d_6) δ 151.37, 149.44, 145.48, 142.51, 136.24, 133.98, 132.42, 132.03, 129.40, 128.88, 128.82, 127.84, 127.52, 126.25, 122.14, 121.57, 108.27, 65.19, 23.20, 21.13. HRMS (ESI): calculated for C₂₀H₁₈N₂NaO₃S [M+Na]⁺: 389.0930; Found: 389.0941.



3-(Hex-5-yn-1-yl)-6-methyl-2-(quinolin-8-yl)-2H-benzo[*e*][**1**,**2**]thiazine **1**,**1**-dioxide(**18**). White solid was obtained in 70% isolated yield (70.3 mg). ¹H NMR (400 MHz, DMSO-*d*₆) δ 8.77 (dd, *J* = 4.20, 1.68 Hz, 1H), 8.43 (dd, *J* = 8.34, 1.68 Hz, 1H), 8.09 (d, *J* = 7.76 Hz, 1H), 7.74 (d, *J* = 6.54 Hz, 1H), 7.67 (t, *J* = 7.76 Hz, 1H), 7.59 (d, *J* = 8.06 Hz, 1H), 7.54 (dd, *J* = 8.30, 4.20 Hz, 1H), 7.42 (s, 1H), 7.31 (d, *J* = 7.96 Hz, 1H), 6.55 (s, 1H), 2.69 (t, *J* = 2.64 Hz, 1H), 2.43 (s, 3H), 2.15 – 2.05 (m, 1H), 2.04 – 1.97 (m, 2H), 1.97 – 1.90 (m, 1H), 1.55 – 1.45 (m, 2H), 1.36 – 1.27 (m, 2H). ¹³C NMR (100 MHz, DMSO-*d*₆) δ 151.38, 144.90, 144.14, 142.38, 136.42, 133.09, 132.31, 131.06, 129.79, 128.88, 128.48, 128.05, 127.02, 126.17, 122.19, 121.13, 108.31, 84.21, 71.24, 33.05, 26.87, 26.10, 21.19, 17.34. HRMS (ESI): calculated for C₂₄H₂₂N₂NaO₂S [M+Na]⁺: 425.1294; Found: 425.1299.



3-(Hydroxymethyl)-4,6-dimethyl-2-(quinolin-8-yl)-2H-benzo[*e*][**1,2**]thiazine **1,1-dioxide(19a).** White solid was obtained in 47% isolated yield (43 mg). ¹H NMR (400 MHz, DMSO-*d*₆) δ 8.86 (dd, *J* = 4.22, 1.68 Hz, 1H), 8.47 (dd, *J* = 8.36, 1.68 Hz, 1H), 8.11 (d, *J* = 7.94 Hz, 1H), 7.77 (s, 1H), 7.66 (t, *J* = 7.78 Hz, 1H), 7.62 – 7.54 (m, 3H), 7.35 (d, *J* = 8.04 Hz, 1H), 5.15 (t, *J* = 5.26 Hz, 1H), 4.59 (d, *J* = 5.22 Hz, 2H), 2.49 (s, 3H), 1.89 (s, 3H). ¹³C NMR (100 MHz, DMSO-*d*₆) δ 151.45, 145.14, 142.05, 139.41, 136.39, 134.17, 133.21, 130.60, 129.75, 129.20, 128.92, 127.68, 126.27, 125.46, 122.22, 121.04, 117.95, 57.45, 21.55, 17.46. HRMS (ESI): calculated for C₂₀H₁₈N₂NaO₃S [M+Na]⁺: 389.0930; Found: 389.0923.



4-(**Hydroxymethyl**)-**3**,**6**-dimethyl-2-(quinolin-8-yl)-2H-benzo[*e*][**1**,**2**]thiazine **1**,**1**-dioxide(**19b**). White solid was obtained in 21% isolated yield (23 mg). ¹H NMR (400 MHz, DMSO-*d*₆) δ 8.79 (d, J = 3.42 Hz, 1H), 8.44 (d, J = 7.82 Hz, 1H), 8.05 (d, J = 7.94 Hz, 1H), 7.68 (s, 1H), 7.65 – 7.53 (m, 3H), 7.51 (d, J = 7.28 Hz, 1H), 7.40 (d, J = 7.82 Hz, 1H), 4.97 (s, 1H), 4.21 (d, J = 11.48 Hz, 1H), 3.68 (d, J = 13.52 Hz, 1H), 2.50 (s, 3H), 2.36 (s, 3H). ¹³C NMR (100 MHz, DMSO-*d*₆) δ 151.24, 145.29, 142.31, 139.28, 136.41, 134.75, 133.30, 130.88, 130.15, 129.41, 128.77, 128.66, 126.21, 125.72, 122.08, 121.22, 116.39, 58.38, 21.46, 14.35. HRMS (ESI): calculated for C₂₀H₁₈N₂NaO₃S [M+Na]⁺: 389.0930; Found: 389.0931.



6-(Tert-butyl)-3-phenyl-2-(quinolin-8-yl)-2H-benzo[*e*][1,2]thiazine 1,1-dioxide(20).²

White solid was obtained in 85% isolated yield (93.9 mg). ¹H NMR (400 MHz, CDCl₃) δ 9.10 (d, *J* = 3.26 Hz, 1H), 8.06 (d, *J* = 8.18 Hz, 1H), 7.80 – 7.70 (m, 3H), 7.63 (d, *J* = 7.74 Hz, 1H), 7.60 (s, 1H), 7.52 (d, *J* = 8.28 Hz, 1H), 7.42 (dd, *J* = 8.26, 4.04 Hz, 1H), 7.28 – 7.20 (m, 2H), 7.15 – 7.08 (m, 3H), 6.97 (s, 1H), 1.41 (s, 9H). ¹³C NMR (100 MHz, CDCl₃) δ 155.71, 151.02, 145.68, 144.09, 135.75, 135.67, 135.39, 132.87, 129.29, 129.05, 128.97, 128.71, 128.61, 128.18, 127.75, 125.63, 124.27, 122.41, 121.67, 112.98, 35.23, 31.19.



8-Methyl-3-phenyl-2-(quinolin-8-yl)-2H-benzo[e][1,2]thiazine 1,1-dioxide(21).¹

White solid was obtained in 56% isolated yield (55.6 mg). ¹H NMR (400 MHz, CDCl₃) δ 9.10 (dd, J = 4.19, 1.69 Hz, 1H), 8.03 (dd, J = 8.32, 1.66 Hz, 1H), 7.77 – 7.71 (m, 2H), 7.62 (dd, J = 6.44, 3.14 Hz, 1H), 7.50 – 7.43 (m, 1H), 7.43 – 7.37 (m, 2H), 7.27 – 7.20 (m, 3H), 7.14 – 7.07 (m, 3H), 6.89 (s, 1H), 2.61 (s, 3H). ¹³C NMR (100 MHz, CDCl₃) δ 151.05, 145.90, 143.24, 135.69, 135.64,

135.51, 135.34, 133.88, 131.45, 131.44, 131.25, 129.03, 128.91, 128.78, 128.77, 128.12, 127.51, 126.06, 125.55, 121.60, 113.03, 20.47.



6-Methoxy-3-phenyl-2-(quinolin-8-yl)-2H-benzo[e][1,2]thiazine 1,1-dioxide(22).¹

White solid was obtained in 71% isolated yield (73.6 mg). ¹H NMR (400 MHz, CDCl₃) δ 9.03 (dd, *J* = 4.18, 1.69 Hz, 1H), 7.97 (dd, *J* = 8.32, 1.64 Hz, 1H), 7.75 – 7.71 (m, 2H), 7.69 (d, *J* = 8.68 Hz, 1H), 7.56 (dd, *J* = 6.70, 2.89 Hz, 1H), 7.33 (dd, *J* = 8.29, 4.20 Hz, 1H), 7.25 – 7.16 (m, 2H), 7.11 – 7.06 (m, 3H), 7.02 (d, *J* = 2.39 Hz, 1H), 6.95 (dd, *J* = 8.68, 2.44 Hz, 1H), 6.87 (s, 1H), 3.86 (s, 3H). ¹³C NMR (100 MHz, CDCl₃) δ 162.27, 150.90, 145.56, 144.69, 135.58, 135.41, 135.12, 135.06, 129.09, 128.86, 128.62, 128.56, 128.08, 127.67, 125.46, 124.49, 124.42, 121.53, 114.58, 112.33, 111.07, 55.56



6,7-Dimethoxy-3-phenyl-2-(quinolin-8-yl)-2H-benzo[*e*][1,2]thiazine 1,1-dioxide(23).²

White solid was obtained in 83% isolated yield (92.2 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 9.13 (d, J = 2.46 Hz, 1H), 8.38 (d, J = 7.40 Hz, 1H), 7.89 (d, J = 8.16 Hz, 1H), 7.85 – 7.78 (m, 2H), 7.62 (dd, J = 8.32, 4.18 Hz, 1H), 7.51 (s, 1H), 7.42 (t, J = 7.80 Hz, 1H), 7.35 (s, 1H), 7.25 – 7.16 (m, 5H), 3.99 (s, 3H), 3.85 (s, 3H). ¹³C NMR (100 MHz, DMSO- d_6) δ 152.02, 151.10, 149.17, 145.09, 141.75, 136.11, 135.24, 134.82, 129.03, 128.96, 128.70, 128.35, 128.15, 126.92, 126.73, 125.95, 123.75, 121.98, 112.86, 110.34, 104.18, 55.97, 55.90.



6-Fluoro-3-phenyl-2-(quinolin-8-yl)-2H-benzo[*e*][1,2]thiazine 1,1-dioxide(24).³

White solid was obtained in 68% isolated yield (68.6 mg). ¹H NMR (400 MHz, DMSO-*d*₆) δ 9.01 (dd, *J* = 4.18, 1.70 Hz, 1H), 8.32 (dd, *J* = 8.34, 1.72 Hz, 1H), 7.87 (dd, *J* = 8.14, 1.48 Hz, 1H), 7.80 (dd, *J* = 8.72, 5.36 Hz, 1H), 7.78 – 7.72 (m, 3H), 7.56 (dd, *J* = 8.34, 4.18 Hz, 1H), 7.47 – 7.41 (m, 1H), 7.39 (d, *J* = 7.92 Hz, 1H), 7.36 – 7.31 (m, 2H), 7.23 – 7.15 (m, 3H). ¹³C NMR (100 MHz, CDCl₃) δ 164.51 (d, ¹*J*_{C-F} = 252.2 Hz), 150.94, 145.62, 145.44, 135.94 (d, ³*J*_{C-F} = 9.7 Hz), 135.66, 135.14, 134.81, 129.42, 129.16, 128.91, 128.90, 128.18, 127.96 (d, ⁴*J*_{C-F} = 2.7 Hz), 127.88, 125.25 (d, ³*J*_{C-F} = 9.8 Hz), 125.20, 121.67, 115.36 (d, ²*J*_{C-F} = 23.5 Hz), 113.55 (d, ²*J*_{C-F} = 23.0 Hz), 111.42 (d, ⁴*J*_{C-F} = 2.4 Hz). ¹⁹F NMR (376 MHz, CDCl₃) δ -106.40.



6-Chloro-3-phenyl-2-(quinolin-8-yl)-2H-benzo[e][1,2]thiazine 1,1-dioxide(25).¹

White solid was obtained in 56% isolated yield (58.3 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 9.01 (dd, J = 4.16, 1.61 Hz, 1H), 8.33 (dd, J = 8.34, 1.52 Hz, 1H), 8.00 (d, J = 1.93 Hz, 1H), 7.87 (dd, J = 8.07, 1.18 Hz, 1H), 7.80 – 7.72 (m, 3H), 7.64 (dd, J = 8.39, 2.01 Hz, 1H), 7.56 (dd, J = 8.30, 4.19 Hz, 1H), 7.43 – 7.37 (m, 1H), 7.37 – 7.31 (m, 2H), 7.23 – 7.16 (m, 3H). ¹³C NMR (100 MHz, DMSO- d_6) δ 151.26, 144.88, 144.69, 137.20, 136.26, 134.92, 134.83, 134.37, 129.88, 129.76, 129.40, 129.19, 128.71, 128.48, 127.62, 127.47, 126.04, 124.12, 122.17, 111.77.



3-Phenyl-2-(quinolin-8-yl)-6-(trifluoromethyl)-2H-benzo[*e*][1,2]thiazine 1,1-dioxide(26).²

White solid was obtained in 63% isolated yield (71.5 mg). ¹H NMR (400 MHz, CDCl₃) δ 8.93 (d, J = 3.96 Hz, 1H), 8.04 (d, J = 8.14 Hz, 1H), 7.94 – 7.87 (m, 2H), 7.74 – 7.68 (m, 3H), 7.66 (d, J = 8.02 Hz, 1H), 7.40 – 7.35 (m, 2H), 7.31 (t, J = 7.72 Hz, 1H), 7.19 – 7.11 (m, 3H), 6.96 (s, 1H). ¹³C NMR (100 MHz, CDCl₃) δ 151.00, 145.98, 145.37, 135.77, 135.07, 134.69, 134.24, 134.05, 133.72, 129.66, 129.61, 129.12, 128.94, 128.28, 128.04, 125.61, 124.52 (q, ${}^{3}J_{C-F} = 3.8$ Hz), 124.32

(q, ${}^{3}J_{C-F} = 3.4$ Hz), 123.40 (q, ${}^{1}J_{C-F} = 273.2$ Hz), 123.39, 121.81, 111.36. ${}^{19}F$ NMR (376 MHz, CDCl₃) δ -62.88.



N-(1,1-Dioxido-3-phenyl-2-(quinolin-8-yl)-2H-benzo[*e*][1,2]thiazin-6-yl)acetamide(27).²

White solid was obtained in 75% isolated yield (82.6 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 10.44 (s, 1H), 9.06 (d, J = 3.88 Hz, 1H), 8.34 (d, J = 8.78 Hz, 1H), 8.18 (s, 1H), 7.86 (d, J = 8.16 Hz, 1H), 7.80 (dd, J = 6.76, 2.90 Hz, 2H), 7.65 (s, 2H), 7.58 (dd, J = 8.32, 4.20 Hz, 1H), 7.38 (t, J = 7.82 Hz, 1H), 7.33 (s, 1H), 7.23 (d, J = 7.48 Hz, 1H), 7.21 – 7.14 (m, 3H), 2.15 (s, 3H). ¹³C NMR (100 MHz, CDCl₃) δ 169.10, 151.20, 145.00, 143.63, 142.84, 136.13, 135.09, 134.74, 133.81, 129.33, 129.02, 128.69, 128.41, 128.29, 127.33, 125.95, 125.83, 123.12, 122.01, 118.80, 117.02, 113.09, 24.18.



3,6-Diphenyl-2-(quinolin-8-yl)-2H-benzo[*e*][1,2]thiazine 1,1-dioxide(28).¹

White solid was obtained in 64% isolated yield (73 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 9.06 (dd, J = 4.18, 1.74 Hz, 1H), 8.35 (d, J = 8.20 Hz, 1H), 8.20 (d, J = 1.78 Hz, 1H), 7.92 – 7.84 (m, 2H), 7.86 – 7.75 (m, 5H), 7.63 – 7.53 (m, 3H), 7.50 (d, J = 7.34 Hz, 1H), 7.46 (s, 1H), 7.39 (t, J = 7.80 Hz, 1H), 7.28 (d, J = 6.02 Hz, 1H), 7.23 – 7.16 (m, 3H). ¹³C NMR (100 MHz, DMSO- d_6) δ 151.14, 144.97, 144.13, 143.64, 138.61, 136.15, 135.08, 134.66, 133.45, 130.16, 129.40, 129.18, 129.12, 128.68, 128.63, 128.59, 128.38, 127.24, 127.14, 126.86, 126.35, 125.96, 122.62, 122.04, 112.95.



3-Phenyl-2-(quinolin-8-yl)-2H-benzo[e][1,2]thiazine 1,1-dioxide(29).¹

White solid was obtained in 76% isolated yield (73.2 mg). ¹H NMR (400 MHz, DMSO-*d*₆) δ 9.05 (d, *J* = 2.80 Hz, 1H), 8.33 (d, *J* = 8.19 Hz, 1H), 7.87 (dd, *J* = 14.02, 7.98 Hz, 2H), 7.83 – 7.76 (m, 3H), 7.73 (d, *J* = 7.69 Hz, 1H), 7.64 – 7.52 (m, 2H), 7.43 – 7.32 (m, 2H), 7.22 (d, *J* = 7.32 Hz, 1H), 7.21 – 7.14 (m, 3H). ¹³C NMR (100 MHz, DMSO-*d*₆) δ 151.16, 144.98, 143.31, 136.17, 135.07, 134.67, 132.80, 132.60, 131.38, 129.39, 129.13, 128.70, 128.68, 128.59, 128.38, 128.27, 127.27, 125.97, 122.06, 121.90, 112.88.



3-Phenyl-2-(quinolin-8-yl)-2H-naphtho[2,3-e][1,2]thiazine 1,1-dioxide(30).¹

White solid was obtained in 65% isolated yield (70.6 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 9.04 (dd, J = 4.24, 1.74 Hz, 1H), 8.42 (d, J = 14.38 Hz, 2H), 8.34 (dd, J = 8.34, 1.74 Hz, 1H), 8.21 – 8.12 (m, 2H), 7.90 – 7.83 (m, 3H), 7.78 – 7.71 (m, 1H), 7.68 – 7.61 (m, 1H), 7.60 – 7.54 (m, 2H), 7.35 (t, J = 7.82 Hz, 1H), 7.28 – 7.16 (m, 4H). ¹³C NMR (100 MHz, CDCl₃) δ 150.93, 145.72, 143.83, 135.63, 135.47, 134.76, 132.10, 131.35, 129.50, 129.12, 129.08, 129.03, 128.69, 128.54, 128.23, 128.10, 127.56, 126.86, 126.70, 125.59, 123.44, 121.64, 113.68.



3-Phenyl-2-(quinolin-8-yl)-2H-thieno[3,2-e][1,2]thiazine 1,1-dioxide(31).¹

White solid was obtained in 75% isolated yield (73.3 mg). ¹H NMR (400 MHz, DMSO- d_6) δ 9.09 (dd, J = 4.18, 1.66 Hz, 1H), 8.36 (dd, J = 8.36, 1.61 Hz, 1H), 8.06 (d, J = 5.04 Hz, 1H), 7.89 (dd, J = 8.22, 1.14 Hz, 1H), 7.75 – 7.68 (m, 2H), 7.60 (dd, J = 8.30, 4.20 Hz, 1H), 7.48 (d, J = 5.05 Hz, 1H), 7.44 – 7.35 (m, 2H), 7.24 (dd, J = 7.41, 1.25 Hz, 1H), 7.18 – 7.12 (m, 3H). ¹³C NMR (100 MHz, DMSO- d_6) δ 151.30, 144.99, 143.62, 141.35, 136.24, 134.96, 134.39, 131.18, 129.35, 129.33, 128.69, 128.61, 128.40, 127.28, 127.13, 126.55, 126.06, 122.13, 109.28.

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Copies of ¹H NMR, ¹³C NMR and ¹⁹F NMR Spectra









10 0 -10 -20 -30 -40 -50 -60 -70 -80 -90 -100 -110 -120 -130 -140 -150 -160 -170 -180 -190 -200 -210 fl (ppm)



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Compound 11



Compound 12



Compound 13 2.024 3.10 1.104 1.054 1.10 1.12 ₹ 1.14 1.00^T 1.07 2.10 1.04 2.04 € 0.99 € 1.04 1.06 1.03 5.0 4.5 fl (ppm) 0.0 9.5 9.0 8.5 . 8.0 . 7.5 7.0 6.5 . 6.0 . 5.5 4.0 3.5 3.0 2.5 2.0 1.5 1.0 0.5 0.0 -($\begin{array}{c} [51.51\\ 143.53\\ 142.43\\ 142.43\\ 136.50\\ 132.97\\ 132.97\\ 132.43\\ 131.8\\ 132.43\\ 131.8\\ 132.43\\ 132.43\\ 132.43\\ 132.43\\ 132.43\\ 132.43\\ 132.5.92\\ 122.27\\$ 40.15 39.99 39.73 39.73 39.73 39.31 39.10 38.89 36.04 33.43 -21.18 ۱[،] 90 80 fl (ppm) 80 170 . 160 150 140 130 120 110 100 70 . 60 50 40 30 20 10 0









Compound 18









S41









Compound 24 9.01 9.01 9.00 9.00 9.00 9.00 8.8.33 8.8.33 8.8.33 8.8.33 7.7.88 7.7.88 7.7.88 7.7.78 7.7.78 7.7.78 7.7.78 7.7.78 7.7.78 7.7.78 7.7.78 7.7.78 7.7.75 7.77 7.75 7.77 7.75 7 1.04 1.03 3.06 1.1.03 3.06 1.09 1.09 1.20 1.20 1.3.18 3.18 1.00-≖ 5.0 4.5 fl (ppm)).0 9.5 . 9.0 8.5 8.0 7.5 7.0 2.5 1.5 6.5 6.0 5.5 4.0 3.5 3.0 2.0 1.0 0.5 0.0 -((150.94)
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- -106.40



Compound 26





50	30	10	-10	-30	-50	-70	-90	-110	-130	-150	-170	-190	-210	-230	-
fl (ppm)															





Compound 29





