

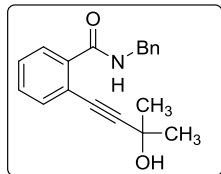
Highly, regioselective electrophile induced cyclizations of 2-(prop-1-ynyl)benzamides

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N-Benzyl-2-(3-hydroxy-3-methylbut-1-yn-1-yl)benzamide (10a)



To a solution of N-benzyl-2-iodobenzamide¹ (1 gm, 2.96 mmol) and 2-methylbut-3-yn-2-ol (298 mg, 3.55 mmol) in dry THF (20 mL) and DIPA (10 mL) were added [(Ph₃P)₂PdCl₂] (20 mg, 0.029 mmol) and CuI (28 mg, 0.28 mmol) under nitrogen. The reaction was stirred at room temperature for

14 h. The reaction mixture was diluted with saturated aq. NH₄Cl (10 mL) and ethyl acetate (20 mL). Aqueous layer was extracted with ethyl acetate (2 x 50 mL). The combined organic layers were washed with brine (10 mL), and dried over Na₂SO₄. Evaporation of the solvent and purification of the crude mixture by column chromatography (4:1, hexane: EtOAc) gave the coupled product **10a** (750 mg, 2.55 mmol, 86 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.08 (1 H, dd, *J* = 1.9 & 6.9 Hz), 7.70 (1 H, br, s), 7.28-7.46 (8 H, m), 4.67 (2 H, d, *J* = 5.5 Hz), 1.39 (6 H, s) ppm.

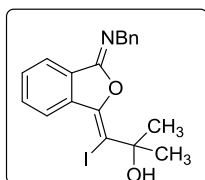
¹³C NMR (100 MHz, CDCl₃): δ = 166.3, 138.2, 135.1, 133.6, 130.6, 130.0, 128.91, 128.88, 128.3, 127.8, 119.3, 100.7, 80.5, 65.3, 44.3 and 30.9 ppm.

IR (neat): 3394, 3057, 2987, 1649, 1531, 1437, 1267, 1163, 903, 735 and 555 cm⁻¹.

HR ESI-MS: [C₁₉H₂₀NO₂]⁺ = [M+H]⁺ requires 294.1494; found 294.1505.

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

1-((1E,3Z)-3-(Benzylimino)isobenzofuran-1(3H)-ylidene)-1-iodo-2-methylpropan-2-ol (11a)



To a solution of coupled product **10a** (30 mg, 0.102 mmol) in dry DCM (3 mL), was added NIS (28 mg, 0.122 mmol) and the reaction mixture was stirred at room temperature for 15 min. The reaction mixture was diluted with water (10 mL) and DCM (20 mL) aqueous layer was extracted with DCM (2 x 20

mL). The combined organic layers were washed with hypo solution (10 mL) and brine (10 mL), and dried over Na₂SO₄. Evaporation of the solvent and purification of the crude mixture by column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **11a** (41 mg, 0.097 mmol, 95 %) as a brown viscous oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.82 (1 H, d, *J* = 8.0 Hz), 7.94 (1 H, d, *J* = 7.6 Hz), 7.60 (1 H, td, *J* = 1.1 & 8.2 Hz), 7.52 (1 H, t, *J* = 7.6 Hz), 7.40 (2 H, d, *J* = 7.2 Hz), 7.33 (2 H, t, *J* = 7.3 Hz), 7.22-7.26 (1 H, m), 4.81 (2 H, s), 3.19 (1 H, br, s), 1.68 (6 H, s) ppm.

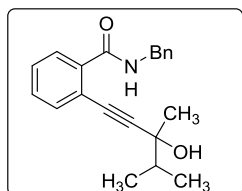
¹³C NMR (100 MHz, CDCl₃): δ = 153.6, 145.9, 139.6, 136.7, 131.5, 131.0, 130.4, 128.6, 127.6, 127.0, 125.4, 123.5, 93.0, 74.6, 52.0 and 29.8 ppm.

IR (neat): 3309, 3059, 2976, 2928, 2856, 1777, 1684, 1645, 1456, 1361, 1262, 1108, 1007, 740, 700, 544 and 411 cm^{-1} .

HR ESI-MS: $[\text{C}_{19}\text{H}_{19}\text{INO}_2]^+ = [\text{M}+\text{H}]^+$ requires 420.0461; found 420.0465.

TLC: $R_f = 0.6$ (4:1; hexane:EtOAc).

N-Benzyl-2-(3-hydroxy-3,4-dimethylpent-1-yn-1-yl)benzamide (10b)



To a solution of N-benzyl-2-iodobenzamide (200 mg, 0.593 mmol) and 3,4-dimethylpent-1-yn-3-ol (80 mg, 0.711 mmol) in dry THF (10 mL) and DIPA (1 mL) were added $[(\text{Ph}_3\text{P})_2\text{PdCl}_2]$ (4 mg, 0.00593 mmol) and CuI (17 mg, 0.088 mmol) under nitrogen. The reaction was stirred at room temperature for 14 h. purification by flash column chromatography (4:1, hexane: EtOAc) gave the coupled product **10b** (150 mg, 0.467 mmol, 79 %) as a yellow oil.

^1H NMR (400 MHz, CDCl_3): δ = 8.06 (1 H, dd, J = 2.2 & 5.9 Hz), 7.74 (1 H, br), 7.29-7.45 (8 H, m), 4.66 (2 H, d, J = 5.6 Hz), 1.91 (1 H, br, s), 1.71-1.76 (1 H, m), 1.31 (3 H, s), 0.98 (3 H, d, J = 6.8 Hz), 0.95 (3 H, d, J = 6.8 Hz) ppm.

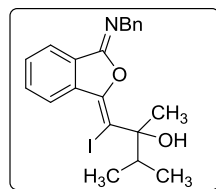
^{13}C NMR (100 MHz, CDCl_3): δ = 166.3, 139.8, 138.1, 135.1, 133.6, 130.5, 129.9, 128.76, 128.71, 128.1, 127.6, 119.4, 82.2, 71.8, 44.1, 38.8, 26.5, 17.7 and 17.6 ppm.

IR (neat): 3408, 3055, 2976, 1657, 1524, 1441, 1271, 1155, 1088, 1034, 908, 731, 473 and 430 cm^{-1} .

HR ESI-MS: $[\text{C}_{21}\text{H}_{23}\text{NaNO}_2]^+ = [\text{M}+\text{Na}]^+$ requires 344.1626; found 344.1645.

TLC: $R_f = 0.4$ (4:1; hexane:EtOAc).

1-((1E,3Z)-3-(Benzylimino)isobenzofuran-1(3H)-ylidene)-1-iodo-2,3-dimethylbutan-2-ol (11b)



To a solution of coupled product **10b** (30 mg, 0.09 mmol) in dry DCM (3 mL), was added NIS (25 mg, 0.112 mmol) and the reaction mixture was stirred at room temperature for 15 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **11b** (29 mg, 0.064 mmol, 72 %) as a yellow oil.

^1H NMR (400 MHz, CDCl_3): δ = 8.82 (1 H, d, J = 8.0 Hz), 7.87 (1 H, d, J = 7.5 Hz), 7.54 (1 H, t, J = 7.5 Hz), 7.45 (1 H, t, J = 7.4 Hz), 7.32 (2 H, d, J = 7.4 Hz), 7.26 (2 H, t, J = 7.3 Hz), 7.17 (1 H, t, J = 6.1 Hz), 4.72 (2 H, s), 2.76 (1 H, br, s), 2.41-2.48 (1 H, m), 1.57 (3 H, s), 0.92 (3 H, d, J = 6.7 Hz), 0.88 (3 H, d, J = 6.8 Hz) ppm.

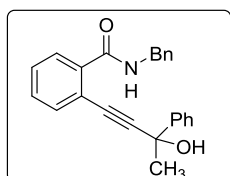
^{13}C NMR (100 MHz, CDCl_3): δ = 153.8, 145.6, 139.7, 136.8, 131.5, 131.4, 130.4, 128.6, 127.6, 127.0, 125.7, 123.5, 95.1, 78.7, 52.1, 36.4, 25.4, 17.9 and 17.4 ppm.

IR (neat): 3055, 2984, 2929, 1786, 1705, 1650, 1423, 1266, 1099, 1027, 896, 744, 708 and 436 cm^{-1} .

HR ESI-MS: $[\text{C}_{21}\text{H}_{23}\text{INO}_2]^+ = [\text{M}+\text{H}]^+$ requires 448.0774; found 448.0803.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

N-Benzyl-2-(3-hydroxy-3-phenylbut-1-yn-1-yl)benzamide (10c)



To a solution of N-benzyl-2-iodobenzamide (200 mg, 0.593 mmol) and 2-phenylbut-3-yn-2-ol (104 mg, 0.711 mmol) in dry THF (10 mL) and DIPA (1 mL) were added $[(\text{Ph}_3\text{P})_2\text{PdCl}_2]$ (4 mg, 0.00593 mmol) and CuI (17 mg, 0.088 mmol) under nitrogen. The reaction was stirred at room temperature for 14 h. purification by flash column chromatography (4:1, hexane: EtOAc) gave the coupled product **10c** (142 mg, 0.423 mmol, 71 %) as a yellow oil.

^1H NMR (400 MHz, CDCl_3): δ = 7.95-7.98 (1 H, m), 7.65 (1 H, br), 7.57 (2 H, dd, J = 1.5 & 8.3 Hz), 7.44-7.47 (1 H, m), 7.28-7.37 (5 H, m), 7.19 (5 H, s), 4.47 (2 H, t, J = 6.2 Hz), 1.68 (3 H, s) ppm.

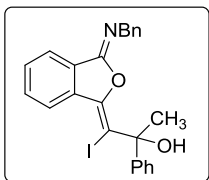
^{13}C NMR (100 MHz, CDCl_3): δ = 166.3, 145.0, 138.1, 135.4, 133.6, 130.6, 130.0, 129.0, 128.7, 128.4, 128.0, 127.9, 127.5, 124.9, 119.1, 99.4, 82.8, 70.1, 44.1 and 32.8 ppm.

IR (neat): 3400, 3059, 2985, 2926, 2846, 1651, 1529, 1437, 1267, 895, 733 and 447 cm^{-1} .

HR ESI-MS: $[\text{C}_{24}\text{H}_{21}\text{NaNO}_2]^+ = [\text{M}+\text{Na}]^+$ requires 378.1470; found 378.1477.

TLC: R_f = 0.4 (4:1; hexane:EtOAc)

1-((1E,3Z)-3-(Benzylimino)isobenzofuran-1(3H)-ylidene)-1-iodo-2-phenylpropan-2-ol (11c)



To a solution of coupled product **10c** (30 mg, 0.084 mmol) in dry DCM (3 mL), was added NIS (23 mg, 0.101 mmol) and the reaction mixture was stirred at room temperature for 15 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodocyclicimidate **11c** (36

mg, 0.076 mmol, 92%) as a yellow oil.

^1H NMR (400 MHz, CDCl_3): δ = 8.85 (1 H, d, J = 8.1 Hz), 7.86 (1 H, d, J = 7.5 Hz), 7.60 (1 H, td, J = 1.0 & 8.2 Hz), 7.50-7.53 (3 H, m), 7.30 (4 H, t, J = 7.6 Hz), 7.19-7.24 (4 H, m), 4.38 (2 H, q, J = 9.3 Hz), 3.43 (1 H, s), 1.96 (3 H, s) ppm.

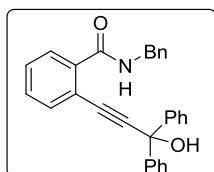
^{13}C NMR (100 MHz, CDCl_3): δ = 153.3, 147.6, 147.1, 139.4, 136.4, 131.5, 131.1, 130.6, 128.5, 128.3, 127.6, 127.3, 126.9, 125.4, 125.1, 123.5, 90.3, 77.5, 51.9 and 32.6 ppm.

IR (neat): 3055, 2986, 2932, 2858, 1785, 1702, 1647, 1608, 1445, 1423, 1265, 1007, 896, 742, 709 and 451 cm^{-1} .

HR ESI-MS: $[\text{C}_{24}\text{H}_{21}\text{INO}_2]^+ = [\text{M}+\text{H}]^+$ requires 482.0617; found 482.0628.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

N-Benzyl-2-(3-hydroxy-3,3-diphenylprop-1-yn-1-yl)benzamide (**10d**)



To a solution of N-benzyl-2-iodobenzamide (200 mg, 0.593 mmol) and 1,1-diphenylprop-2-yn-1-ol (148 mg, 0.711 mmol) in dry THF (10 mL) and DIPA (1 mL) were added $[(\text{Ph}_3\text{P})_2\text{PdCl}_2]$ (4 mg, 0.00593 mmol) and CuI (17 mg, 0.088 mmol). The reaction was stirred at room temperature for 14 h. purification by flash column chromatography (4:1, hexane: EtOAc) gave the coupled product **10d** (152 mg, 0.364 mmol, 61 %) as a yellow oil

^1H NMR (400 MHz, CDCl_3): δ = 7.98-8.00 (1 H, m), 7.50-7.55 (6 H, m), 7.38-7.41 (2 H, m), 7.25-7.33 (6 H, m), 7.13-7.18 (5 H, m), 4.38 (2 H, d, J = 5.8 Hz), 3.05 (1 H, br, s) ppm.

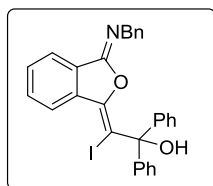
^{13}C NMR (100 MHz, CDCl_3): δ = 166.3, 144.6, 139.8, 138.0, 135.5, 131.1, 130.5, 129.9, 129.0, 128.7, 128.5, 128.3, 128.17, 128.10, 127.87, 127.84, 127.3, 126.0, 119.0, 98.6, 84.9, 74.6 and 43.8 ppm.

IR (neat): 3371, 3300, 3053, 2926, 1639, 1533, 1456, 1282, 1016, 731 and 434 cm^{-1} .

HR ESI-MS: $[\text{C}_{29}\text{H}_{23}\text{NO}_2]^+ = [\text{M}+\text{H}]^+$ requires 440.1626; found 440.1606.

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

2-((1E,3Z)-3-(Benzylimino)isobenzofuran-1(3H)-ylidene)-2-iodo-1,1-diphenylethan-1-ol (**11d**)



To a solution of coupled product **10d** (30 mg, 0.07 mmol) in dry DCM (3 mL), was added NIS (19 mg, 0.086 mmol) and the reaction mixture was stirred at room temperature for 15 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **11d** (37 mg, 0.068 mmol, 98 %) as a brown solid.

^1H NMR (400 MHz, CDCl_3): δ = 8.89 (1 H, d, J = 8.1 Hz), 7.85 (1 H, d, J = 7.6 Hz), 7.60 (1 H, td, J = 1.1 & 8.3 Hz), 7.52 (1 H, td, J = 0.8 & 7.6 Hz), 7.43-7.46 (4 H, m), 7.20-7.34 (9 H, m), 7.11 (2 H, d, J = 6.9 Hz), 3.99 (2 H, s), 3.81 (1 H, br, s) ppm.

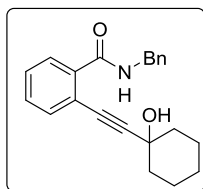
^{13}C NMR (100 MHz, CDCl_3): δ = 152.8, 147.5, 145.4, 139.5, 136.4, 131.5, 131.3, 130.8, 128.4, 128.2, 128.19, 128.15, 127.9, 127.88, 127.84, 127.81, 127.7, 126.8, 125.6, 123.5, 90.9, 83.0 and 51.4 ppm. **IR** (neat): 3438, 3055, 2984, 2926, 2855, 1706, 1652, 1609, 1449, 1266, 1023, 898, 740, 461 and 405 cm^{-1} .

HR ESI-MS: $[\text{C}_{29}\text{H}_{23}\text{INO}_2]^+ = [\text{M}+\text{H}]^+$ requires 544.0774; found 544.0775.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

M.P: 118-120 $^\circ\text{C}$

N-Benzyl-2-((1-hydroxycyclohexyl)ethynyl)benzamide (**10e**)



To a solution of N-benzyl-2-iodobenzamide (300 mg, 0.89 mmol) and 1-ethynylcyclohexan-1-ol (132 mg, 1.068 mmol) in dry THF (10 mL) and DIPA (2 mL) were added $[(\text{Ph}_3\text{P})_2\text{PdCl}_2]$ (6 mg, 0.0089 mmol) and CuI (25 mg, 0.133 mmol) under nitrogen. The reaction was stirred at room temperature for 14 h. purification by flash column chromatography (4:1, hexane: EtOAc) gave the coupled product **10e** (250 mg, 0.75 mmol, 84 %) as a yellow oil.

^1H NMR (400 MHz, CDCl_3): δ = 8.01-8.03 (1 H, m), 7.82 (1 H, br), 7.43-7.45 (1 H, m), 7.26-7.38 (7 H, m), 4.63 (2 H, d, J = 5.6 Hz), 2.60 (1 H, br, s), 1.73-1.75 (2 H, m), 1.62-1.65 (2 H, m), 1.44-1.54 (6 H, m) ppm.

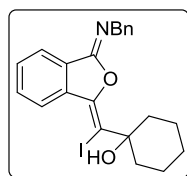
^{13}C NMR (100 MHz, CDCl_3): δ = 166.3, 138.2, 135.1, 133.7, 130.5, 130.0, 128.8, 128.7, 128.2, 127.7, 119.5, 100.1, 82.3, 68.7, 44.2, 39.4, 25.0 and 23.1 ppm.

IR (neat): 3388, 3055, 2935, 1649, 1531, 1441, 1271, 1153, 1068, 966, 901, 739 and 540 cm^{-1} .

HR ESI-MS: $[\text{C}_{22}\text{H}_{24}\text{NO}_2]^+ = [\text{M}+\text{H}]^+$ requires 334.1807; found 344.1808.

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

1-(((1E,3E)-3-(Benzylimino)isobenzofuran-1(3H)-ylidene)iodomethyl)cyclohexan-1-ol (**11e**)



To a solution of coupled product **10e** (30 mg, 0.09 mmol) in dry DCM (3 mL), was added NIS (24 mg, 0.108 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imidate **11e** (35 mg, 0.076 mmol, 85 %) as a yellow oil.

^1H NMR (400 MHz, CDCl_3): δ = 8.85 (1 H, d, J = 8.1 Hz), 7.94 (1 H, d, J = 7.5 Hz), 7.60 (1 H, td, J = 1.2 & 7.4 Hz), 7.52 (1 H, td, J = 0.8 & 7.5 Hz), 7.40 (2 H, d, J = 7.2 Hz), 7.34 (2 H, t, J =

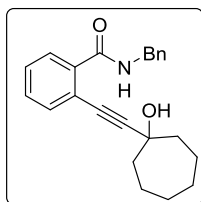
7.3 Hz), 7.23-7.26 (1 H, m), 4.82 (2 H, s), 2.89 (1 H, br, s), 2.12-2.19 (2 H, m), 1.58-1.81 (8 H, m) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 153.6, 145.9, 139.6, 136.8, 131.4, 130.9, 130.3, 128.5, 127.5, 126.9, 125.5, 123.4, 94.9, 75.3, 52.0, 36.7, 25.4 and 22.5 ppm.

IR (neat): 3415, 3059, 2976, 2875, 1780, 1691, 1433, 1265, 1149, 997, 895, 73 and 436 cm⁻¹.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

N-Benzyl-2-((1-hydroxycycloheptyl)ethynyl)benzamide (10f)



To a solution of N-benzyl-2-iodobenzamide (150 mg, 0.445 mmol) and 1-ethynylcycloheptan-1-ol (74 mg, 0.534 mmol) in dry THF (8 mL) and DIPA (1 mL) were added [(Ph₃P)₂PdCl₂] (3 mg, 0.004 mmol) and CuI (13 mg, 0.066 mmol) under nitrogen. The reaction was stirred at room temperature for 14 h. purification by flash column chromatography (8:2, hexane: EtOAc) gave the coupled product **10f** (80 mg, 0.23 mmol, 52 %) as a yellow solid.

¹H NMR (400 MHz, CDCl₃): δ = 8.06-8.08 (1 H, m), 7.81 (1 H, br), 7.44-7.46 (1 H, m), 7.33-7.42 (6 H, m), 7.29 (1 H, dt, *J* = 1.4 & 6.2 Hz), 4.66 (2 H, d, *J* = 4.5 Hz), 2.02 (1 H, br, s), 1.84-1.91 (3 H, m), 1.72-1.77 (2 H, m), 1.44-1.68 (7 H, m) ppm.

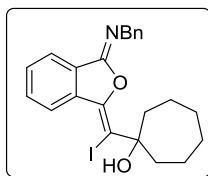
¹³C NMR (100 MHz, CDCl₃): δ = 166.2, 138.3, 135.0, 133.7, 130.6, 130.2, 128.9, 128.8, 128.3, 127.8, 119.4, 101.1, 81.7, 71.9, 44.3, 42.7, 28.0 and 22.1 ppm.

IR (neat): 3391, 3055, 2985, 2931, 2857, 1654, 1595, 1530, 1453, 1424, 1266, 1155, 1027, 897, 743, 707, 529 and 464 cm⁻¹.

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

M.P: 80-82 °C

1-(((1E,3E)-3-(Benzylimino)isobenzofuran-1(3H)-ylidene)iodomethyl)cycloheptan-1-ol (11f)



To a solution of coupled product **10f** (30 mg, 0.086 mmol) in dry DCM (3 mL), was added NIS (23 mg, 0.103 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **11f** (28

mg, 0.059 mmol, 68 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.86 (1 H, d, *J* = 8.1 Hz), 7.94 (1 H, d, *J* = 7.6 Hz), 7.60 (1 H, td, *J* = 1.2 & 7.4 Hz), 7.52 (1 H, td, *J* = 0.8 & 7.6 Hz), 7.40 (2 H, d, *J* = 7.5 Hz), 7.34 (2 H, t, *J* =

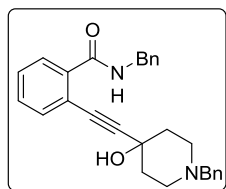
7.3 Hz), 7.25 (1 H, t, $J = 6.5$ Hz), 4.84 (2 H, s), 2.83 (1 H, br, s), 2.36-2.43 (2 H, m), 1.87-1.92 (2 H, m), 1.54-1.80 (8 H, m) ppm.

^{13}C NMR (100 MHz, CDCl_3): $\delta = 153.7, 145.5, 139.7, 137.0, 131.4, 131.1, 130.3, 128.6, 127.6, 127.0, 125.6, 123.5, 97.1, 78.6, 52.1, 40.4, 28.3$ and 22.7 ppm.

IR (neat): 3443, 3055, 2927, 2857, 1781, 1700, 1645, 1458, 1265, 1110, 1032, 997, 897, 739 and 442 cm^{-1} .

TLC: $R_f = 0.6$ (4:1; hexane:EtOAc)

N-Benzyl-2-((1-benzyl-4-hydroxypiperidin-4-yl)ethynyl)benzamide (10g)



To a solution of N-benzyl-2-iodobenzamide (300 mg, 0.89 mmol) and 1-benzyl-4-ethynylpiperidin-4-ol² (229 mg, 1.06 mmol) in dry THF (10 mL) and DIPA (2 mL) were added $[(\text{Ph}_3\text{P})_2\text{PdCl}_2]$ (6 mg, 0.0089 mmol) and CuI (25 mg, 0.133 mmol) under nitrogen. The reaction was stirred at room temperature for 14 h. purification by flash column chromatography (7:3, hexane: EtOAc) gave the coupled product **10g** (210 mg, 0.495 mmol, 56 %) as a yellow oil

^1H NMR (400 MHz, CDCl_3): $\delta = 8.01$ (1 H, d, $J = 6.5$ Hz), 7.60 (1 H, br, s), 7.36-7.43 (5 H, m), 7.23-7.34 (8 H, m), 4.65 (2 H, d, $J = 5.5$ Hz), 3.51 (2 H, s), 2.58-2.63 (2 H, m), 2.34-2.42 (4 H, m), 1.92-1.96 (2 H, m) ppm.

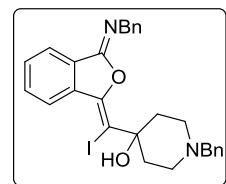
^{13}C NMR (100 MHz, CDCl_3): $\delta = 166.4, 138.3, 138.2, 135.4, 133.8, 130.6, 129.9, 129.27, 129.25, 129.0, 128.9, 128.3, 127.8, 127.2, 119.3, 101.2, 82.7, 70.7, 62.7, 49.9, 44.3$ and 38.9 ppm.

IR (neat): 3473, 2978, 1734, 1450, 1383, 1248, 1053, 920, 733, 629 and 446 cm^{-1} .

HR ESI-MS: $[\text{C}_{28}\text{H}_{29}\text{N}_2\text{O}_2]^+ = [\text{M}+\text{H}]^+$ requires 425.2229; found 425.2234.

TLC: $R_f = 0.4$ (7:3; hexane:EtOAc)

1-Benzyl-4-(((1E,3E)-3-(benzylimino)isobenzofuran-1(3H)-ylidene)iodomethyl)piperidin-4-ol (11g)



To a solution of coupled product **10g** (30 mg, 0.07 mmol) in dry DCM (3 mL), was added NIS (19 mg, 0.084 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (8:2, hexane: EtOAc) gave the iodo cyclic imide **11g** (31

mg, 0.056 mmol, 80 %) as a yellow oil.

¹H NMR (500 MHz, CDCl₃): δ = 8.84 (1 H, d, *J* = 6.4 Hz), 7.94 (1 H, d, *J* = 6.0 Hz), 7.60 (1 H, td, *J* = 0.9 & 6.6 Hz), 7.53 (1 H, td, *J* = 0.6 & 6.1 Hz), 7.42 (2 H, d, *J* = 5.8 Hz), 7.24-7.36 (8 H, m), 4.84 (2 H, s), 3.58 (2 H, s), 2.73-2.79 (2 H, m), 2.45-2.48 (4 H, m), 1.80 (2 H, *J* = 8.5 Hz) ppm.

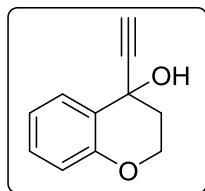
¹³C NMR (125 MHz, CDCl₃): δ = 153.4, 146.8, 139.7, 136.7, 131.5, 131.0, 130.6, 129.3, 128.7, 128.5, 128.4, 127.7, 127.2, 127.0, 125.6, 123.5, 90.1, 73.7, 63.0, 52.2, 49.8 and 37.2 ppm.

IR (neat): 3439, 3055, 2984, 2927, 2855, 1699, 1643, 1455, 1424, 1266, 1025, 896, 740, 538 and 451 cm⁻¹.

HR ESI-MS: [C₂₈H₂₈IN₂O₂]⁺ = [M+H]⁺ requires 551.1196; found 551.1224.

TLC: R_f = 0.4 (4:1; hexane:EtOAc)

4-Ethynylchroman-4-ol (S0)



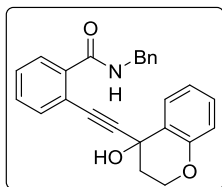
To an ice cold solution of chroman-4-one (300 mg, 2.02 mmol) in dry THF (10 mL) and ethynyl magnesium bromide (0.5 M in THF 4.04 mmol) under nitrogen. The reaction was stirred at same temperature for 2 h. The reaction mixture was diluted with saturated aq. NH₄Cl (10 mL) and ethyl acetate (20 mL). Aqueous layer was extracted with ethyl acetate (2 x 50 mL). The combined organic layers were washed with brine (10 mL), and dried over Na₂SO₄. Evaporation of the solvent and purification of the crude mixture by column chromatography (8:2, hexane: EtOAc) gave the alcohol **S0** (280 mg, 1.64 mmol, 86%) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 7.64 (1 H, dd, *J* = 1.3 & 7.8 Hz), 7.20 (1 H, 1.5 & 8.3 Hz), 6.93 (1 H, t, *J* = 7.7 Hz), 6.80 (1 H, d, *J* = 8.2 Hz), 4.27-4.30 (2 H, m), 2.76 (1 H, t, *J* = 7.4 Hz), 2.61 (1 H, s), 2.24-2.36 (2 H, m) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 153.5, 130.3, 128.2, 124.6, 120.9, 117.3, 86.4, 73.1, 63.6, 62.3 and 37.1 ppm.

TLC: R_f = 0.5 (4:1; hexane:EtOAc).

N-Benzyl-2-((4-hydroxychroman-4-yl)ethynyl)benzamide (10h)



To a solution of N-benzyl-2-iodobenzamide (200 mg, 0.593 mmol) and 4-ethynylchroman-4-ol **S0** (124 mg, 0.71 mmol) in dry THF (10 mL) and DIPA (2 mL) were added [(Ph₃P)₂PdCl₂] (4 mg, 0.005 mmol) and CuI (17 mg, 0.088 mmol) under nitrogen. The reaction was stirred at room temperature

for 14 h. purification by flash column chromatography (4:1, hexane: EtOAc) gave the coupled product **10h** (145 mg, 0.378 mmol, 64 %) as a yellow solid.

¹H NMR (400 MHz, CDCl₃): δ = 7.99-8.02 (1 H, m), 7.55 (1 H, dd, *J* = 1.6 & 7.8 Hz), 7.46-7.48 (2 H, m), 7.38-7.42 (2 H, m), 7.19-7.28 (6 H, m), 6.87-6.91 (1 H, m), 6.83 (1 H, d, *J* = 8.2 Hz), 4.57-4.63 (1 H, m), 4.44-4.49 (1 H, m), 4.21-4.30 (2 H, m), 2.94 (1 H, m), 2.14 (2 H, t, *J* = 3.5 Hz) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 166.5, 153.5, 138.0, 135.6, 133.6, 130.5, 130.2, 129.7, 129.0, 128.7, 128.3, 128.0, 127.5, 124.7, 120.8, 119.1, 117.3, 98.4, 82.8, 64.0, 62.3, 44.1 and 36.8 ppm.

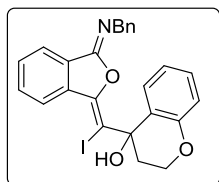
IR (neat): 3059, 2980, 2943, 2858, 1643, 1506, 1439, 1267, 1043, 883, 739 and 430 cm⁻¹.

HR ESI-MS: [C₂₅H₂₁NaNO₃]⁺ = [M+Na]⁺ requires 406.1419; found 406.1434.

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

M.P: 135-137 °C

4-(((1E,3E)-3-(Benzylimino)isobenzofuran-1(3H)-ylidene)iodomethyl)chroman-4-ol (**11h**)



To a solution of coupled product **10h** (30 mg, 0.078 mmol) in dry DCM (3 mL), was added NIS (21 mg, 0.093 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **11h** (35 mg, 0.068 mmol, 88 %) as a brown solid.

¹H NMR (400 MHz, CDCl₃): δ = 8.85 (1 H, d, *J* = 8.1 Hz), 7.83 (1 H, d, *J* = 7.6 Hz), 7.60 (1 H, td, *J* = 1.0 & 8.2 Hz), 7.49 (1 H, t, *J* = 7.6 Hz), 7.20-7.30 (6 H, m), 7.09 (1 H, td, *J* = 1.6 & 7.4 Hz), 6.87 (1 H, dd, *J* = 0.9 & 8.2 Hz), 6.81 (1 H, td, *J* = 1.0 & 7.6 Hz), 4.33 (2 H, dd, *J* = 2.4 & 7.6 Hz), 4.23 (1 H, d, *J* = 15.8 Hz), 4.02 (1 H, d, *J* = 15.8 Hz), 2.60-2.67 (1 H, m), 2.03-2.07 (1 H, m) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 154.0, 153.0, 145.9, 139.8, 136.3, 136.0, 131.3, 130.4, 129.3, 128.2, 127.6, 127.4, 127.1, 126.5, 125.1, 123.4, 121.2, 117.2, 91.2, 70.6, 63.2, 51.2 and 34.5 ppm.

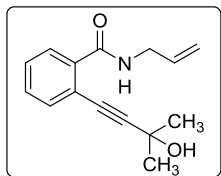
IR (neat): 3367, 3059, 2983, 1784, 1699, 1603, 1437, 1265, 1001, 897, 731 and 455 cm⁻¹.

HR ESI-MS: [C₂₅H₂₁INO₃]⁺ = [M+H]⁺ requires 510.0566; found 510.0594.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

M.P: 125-127 °C

N-Allyl-2-(3-hydroxy-3-methylbut-1-yn-1-yl)benzamide (**14a**)



To a solution of N-allyl-2-iodobenzamide¹ (500 mg, 1.74 mmol) and 2-methylbut-3-yn-2-ol (176 mg, 2.09 mmol) in dry THF (20 mL) and DIPA (4 mL) were added [(Ph₃P)₂PdCl₂] (12 mg, 0.017 mmol) and CuI (50 mg, 0.26 mmol) under nitrogen. The reaction was stirred at room temperature for 14 h.

purification by flash column chromatography (8:2, hexane: EtOAc) gave the coupled product **14a** (380 mg, 1.56 mmol, 90 %) as a white semi solid.

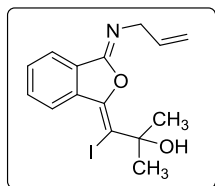
¹H NMR (400 MHz, CDCl₃): δ = 7.97 (1 H, dd, *J* = 3.6 & 6.6 Hz), 7.49 (1 H, br), 7.43 (1 H, dd, *J* = 3.0 & 5.6 Hz), 7.36 (2 H, dd, *J* = 3.8 & 5.5 Hz), 5.89-5.98 (1 H, m), 5.27 (1 H, dd, *J* = 1.3 & 17.1 Hz), 5.16 (1 H, dd, *J* = 1.1 & 10.2 Hz), 4.07-4.10 (2 H, m), 1.60 (6 H, s) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 166.4, 135.4, 134.1, 133.5, 130.5, 129.8, 128.8, 119.3, 116.9, 100.7, 80.4, 65.4, 42.6 and 31.2 ppm.

IR (neat): 3410, 3061, 2985, 1720, 1651, 1535, 1433, 1267, 1169, 906, 755 and 465 cm⁻¹.

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

1-((1E,3Z)-3-(Allylimino)isobenzofuran-1(3H)-ylidene)-1-iodo-2-methylpropan-2-ol (**15a**)



To a solution of coupled product **14a** (30 mg, 0.123 mmol) in dry DCM (3 mL), was added NIS (33 mg, 0.148 mmol) and the reaction mixture was stirred at room temperature for 15 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **15a** (37

mg, 0.1 mmol, 82 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.73 (1 H, d, *J* = 8.0 Hz), 7.84 (1 H, d, *J* = 7.5 Hz), 7.50 (1 H, t, *J* = 7.4 Hz), 7.45 (1 H, t, *J* = 7.4 Hz), 5.93-6.02 (1 H, m), 5.22 (1 H, dd, *J* = 1.5 & 17.1 Hz), 5.09 (1 H, dd, *J* = 1.3 & 10.2 Hz), 4.17 (2 H, d, *J* = 5.4 Hz), 3.25 (1 H, br, s), 1.63 (6 H, s) ppm.

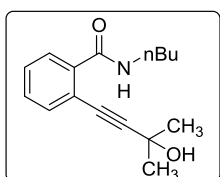
¹³C NMR (100 MHz, CDCl₃): δ = 153.7, 145.9, 136.6, 135.4, 131.5, 130.9, 130.4, 125.4, 123.4, 115.8, 92.7, 74.6, 50.8 and 29.9 ppm.

IR (neat): 3402, 3055, 2985, 2930, 1782, 1684, 1648, 1422, 1266, 1105, 1009, 896, 743, 463 and 417 cm⁻¹.

HR ESI-MS: [C₁₅H₁₇INO₂]⁺ = [M+H]⁺ requires 370.0304; found 370.0339.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

N-Butyl-2-(3-hydroxy-3-methylbut-1-yn-1-yl)benzamide (**14b**)



To a solution of N-butyl-2-iodobenzamide¹ (250 mg, 0.825 mmol) and 2-methylbut-3-yn-2-ol **xx** (83 mg, 0.99 mmol) in dry THF (10 mL) and DIPA

(2 mL) were added $[(\text{Ph}_3\text{P})_2\text{PdCl}_2]$ (6 mg, 0.0082 mmol) and CuI (24 mg, 0.123 mmol) under nitrogen. The reaction was stirred at room temperature for 14 h. purification by flash column chromatography (8:2, hexane: EtOAc) gave the coupled product **14b** (148 mg, 0.57 mmol, 69 %) as a yellow oil.

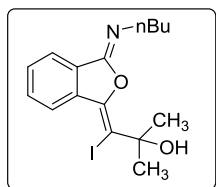
^1H NMR (400 MHz, CDCl_3): δ = 7.92-7.94 (1 H, m), 7.40-7.42 (1 H, m), 7.33-7.35 (3 H, m), 3.61 (1 H, br, s), 3.43 (2 H, q, J = 7.0 Hz), 1.62 (6 H, s), 1.54-1.60 (2 H, m), 1.37 (2 H, q, J = 7.5 Hz), 0.91 (3 H, t, J = 6.8 Hz) ppm.

^{13}C NMR (100 MHz, CDCl_3): δ = 166.6, 135.8, 133.5, 130.3, 129.6, 128.7, 119.3, 100.5, 80.4, 65.4, 40.0, 31.6, 31.3, 20.3 and 13.8 ppm.

IR (neat): 3406, 3059, 2974, 2877, 1651, 1537, 1437, 1267, 1165, 903, 733 and 536 cm^{-1} .

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

1-((1E,3Z)-3-(Butylimino)isobenzofuran-1(3H)-ylidene)-1-iodo-2-methylpropan-2-ol (**15b**)



To a solution of coupled product **14b** (30 mg, 0.115 mmol) in dry DCM (3 mL), was added NIS (31 mg, 0.138 mmol) and the reaction mixture was stirred at room temperature for 15 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **15b** (36 mg, 0.093 mmol, 81 %) as a brown viscous oil.

^1H NMR (400 MHz, CDCl_3): δ = 8.81 (1 H, d, J = 8.0 Hz), 7.89 (1 H, d, J = 7.6 Hz), 7.58 (1 H, t, J = 8.0 Hz), 7.51 (1 H, t, J = 7.5 Hz), 3.59 (2 H, t, J = 7.1 Hz), 3.41 (1 H, br, s), 1.72 (6 H, s), 1.65-1.69 (2 H, m), 1.43 (2 H, q, J = 7.6 Hz), 0.95 (3 H, t, J = 7.3 Hz) ppm.

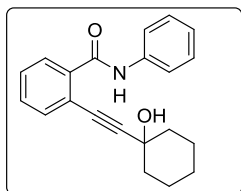
^{13}C NMR (100 MHz, CDCl_3): δ = 152.8, 145.9, 136.5, 131.2, 131.1, 130.3, 125.4, 123.3, 92.0, 74.6, 48.2, 32.9, 29.9, 20.8 and 14.0 ppm.

IR (neat): 3059, 2974, 1782, 1687, 1437, 1265, 1011, 901, 731, 534 and 451 cm^{-1} .

HR ESI-MS: $[\text{C}_{16}\text{H}_{21}\text{INO}_2]^+ = [\text{M}+\text{H}]^+$ requires 386.0617; found 386.0616.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

2-((1-Hydroxycyclohexyl)ethynyl)-N-phenylbenzamide (**14c**)



To a solution of 2-iodo-N-phenylbenzamide³ (500 mg, 1.55 mmol) and 1-ethynylcyclohexan-1-ol (230 mg, 2.09 mmol) in dry THF (10 mL) and DIPA (3 mL) were added $[(\text{Ph}_3\text{P})_2\text{PdCl}_2]$ (11 mg, 0.015 mmol) and CuI (44 mg, 0.23 mmol) under nitrogen. The reaction was stirred at room

temperature for 14 h. purification by flash column chromatography (8:2, hexane: EtOAc) gave the coupled product **14c** (380 mg, 1.19 mmol, 77 %) as a yellow oil

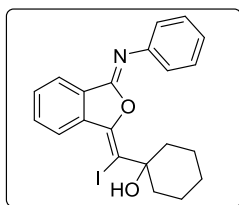
¹H NMR (400 MHz, CDCl₃): δ = 8.95 (1 H, br, s), 8.03 (1 H, dd, J = 4.0 & 8.9 Hz), 7.72 (2 H, d, J = 7.8 Hz), 7.52 (1 H, dd, J = 2.0 & 4.8 Hz), 7.42 (2 H, t, J = 6.6 Hz), 7.34 (2 H, t, J = 7.8 Hz), 7.12 (1 H, t, J = 7.4 Hz), 2.43 (1 H, br, s), 1.94-1.98 (2 H, m), 1.60-1.71 (5 H, m), 1.46-1.57 (3 H, m) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 164.7, 138.0, 136.1, 133.8, 130.8, 130.1, 129.17, 129.10, 124.6, 120.2, 119.3, 100.4, 82.3, 69.3, 39.9, 25.1 and 23.3 ppm.

IR (neat): 3364, 3058, 2939, 2861, 1736, 1666, 1600, 1532, 1443, 1367, 1320, 1236, 1133, 1023, 961, 903, 742 and 414 cm⁻¹.

TLC: R_f = 0.4 (4:1; hexane:EtOAc)

1-(Iodo((1E,3Z)-3-(phenylimino)isobenzofuran-1(3H)-ylidene)methyl)cyclohexan-1-ol (**15c**)



To a solution of coupled product **14c** (30 mg, 0.094 mmol) in dry DCM (3 mL), was added NIS (25 mg, 0.112 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **15c** (37 mg, 0.083 mmol, 87 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.87 (1 H, d, J = 8.0 Hz), 8.01 (1 H, dt, J = 0.9 & 7.5 Hz), 7.65 (1 H, td, J = 1.2 & 7.4 Hz), 7.58 (1 H, td, J = 1.0 & 7.6 Hz), 7.35-7.39 (2 H, m), 7.20 (2 H, dd, J = 0.4 & 8.5 Hz), 7.13-7.18 (1 H, m), 2.70 (1 H, br, s), 1.96-2.04 (2 H, m), 1.68-1.73 (2 H, m), 1.50-1.60 (6 H, m) ppm.

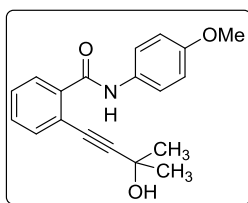
¹³C NMR (100 MHz, CDCl₃): δ = 152.3, 146.2, 146.0, 136.9, 132.0, 130.9, 130.5, 129.0, 125.5, 124.8, 123.9, 122.6, 96.1, 75.3, 36.8, 24.9 and 22.3 ppm.

IR (neat): 3372, 3056, 2980, 2936, 2860, 1731, 1694, 1600, 1479, 1369, 1266, 1142, 1088, 1018, 892 and 742 cm⁻¹.

HR ESI-MS: [C₂₁H₂₀NaINO₂]⁺ = [M+Na]⁺ requires 468.0437; found 468.0435.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

2-(3-Hydroxy-3-methylbut-1-yn-1-yl)-N-(4-methoxyphenyl)benzamide (**14d**)



To a solution of 2-iodo-N-(4-methoxyphenyl)benzamide³ (350 mg, 1.02 mmol) and 2-methylbut-3-yn-2-ol (103 mg, 1.22 mmol) in dry THF (10 mL) and Et₃N (2 mL) were added [(Ph₃P)₂PdCl₂] (7 mg, 0.01 mmol) and

CuI (29 mg, 0.153 mmol) under nitrogen. The reaction was stirred at 60 °C for 14 h. purification by flash column chromatography (8:2, hexane: EtOAc) gave the coupled product **14d** (260 mg, 0.84 mmol, 82 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.95 (1 H, br), 8.00-8.02 (1 H, m), 7.61 (2 H, d, *J* = 8.9 Hz), 7.38-7.45 (3 H, m), 6.84 (2 H, d, *J* = 9.0 Hz), 3.77 (3 H, s), 1.57 (6 H, s) ppm.

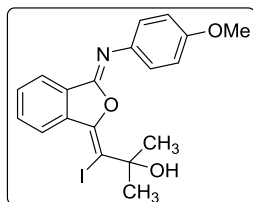
¹³C NMR (100 MHz, CDCl₃): δ = 164.4, 156.6, 135.9, 133.6, 131.0, 130.7, 130.0, 129.0, 122.0, 119.2, 114.2, 101.3, 80.2, 65.5, 55.5 and 31.4 ppm.

IR (neat): 3414, 3056, 2986, 2934, 2848, 1659, 1515, 1420, 1265, 1172, 1122, 1037, 829, 740 and 527 cm⁻¹.

HR ESI-MS: [C₁₉H₁₉NaNO₃]⁺ = [M+Na]⁺ requires 332.1263; found 332.1253.

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

1-Iodo-1-((1E,3Z)-3-((4-methoxyphenyl)imino)isobenzofuran-1(3H)-ylidene)-2-methylpropan-2-ol (15d)



To a solution of coupled product **14d** (30 mg, 0.097 mmol) in dry DCM (3 mL), was added NIS (26 mg, 0.116 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **15d** (37 mg, 0.085 mmol, 88 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.84 (1 H, d, *J* = 8.0 Hz), 8.01 (1 H, d, *J* = 7.5 Hz), 7.64 (1H, td, *J* = 1.2 & 7.6 Hz), 7.58 (1 H, td, *J* = 0.8 & 7.5 Hz), 7.25 (2 H, d, *J* = 6.5 Hz), 6.92 (2 H, d, *J* = 6.8 Hz), 3.82 (3 H, s), 3.27 (1 H, br, s), 1.61 (6 H, s) ppm.

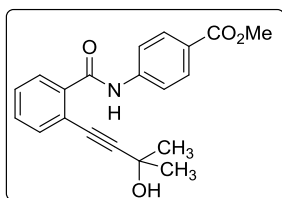
¹³C NMR (100 MHz, CDCl₃): δ = 157.0, 151.1, 146.2, 138.6, 136.3, 131.8, 131.4, 130.5, 125.4, 124.4, 123.8, 114.3, 92.6, 74.9, 55.5 and 30.4 ppm.

IR (neat): 3443, 3055, 2985, 2929, 2844, 1692, 1628, 1506, 1463, 1424, 1265, 1169, 1107, 1027, 895, 840, 738 and 433 cm⁻¹.

HR ESI-MS: [C₁₉H₁₉INO₃]⁺ = [M+H]⁺ requires 436.0410; found 436.0442.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

Methyl 4-(2-(3-hydroxy-3-methylbut-1-yn-1-yl)benzamido)benzoate (14e)



To a solution of methyl 4-(2-iodobenzamido)benzoate³ (300 mg, 0.789 mmol) and 2-methylbut-3-yn-2-ol (80 mg, 0.947 mmol) in dry THF (10 mL) and Et₃N (2 mL) were added [(Ph₃P)₂PdCl₂] (5.5 mg, 0.007 mmol)

and CuI (22 mg, 0.118 mmol) under nitrogen. The reaction was stirred at 60 °C for 14 h. purification by flash column chromatography (8:2, hexane: EtOAc) gave the coupled product **14e** (245 mg, 0.727 mmol, 92 %) as a yellow oil.

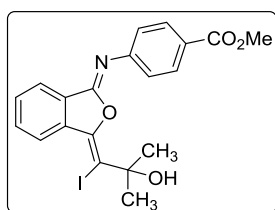
¹H NMR (400 MHz, CDCl₃): δ = 9.23 (1 H, br, s), 8.06 (1 H, dd, *J* = 3.7 & 6.6 Hz), 8.00 (2 H, d, *J* = 8.6 Hz), 7.82 (2 H, d, *J* = 8.7 Hz), 7.50 (1 H, dd, *J* = 3.2 & 5.5 Hz), 7.44 (2 H, dd, *J* = 3.8 & 5.2 Hz), 3.89 (3 H, s), 1.60 (6 H, s) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 166.7, 164.7, 142.1, 135.3, 133.8, 131.2, 130.9, 130.3, 129.2, 125.8, 119.3, 119.2, 101.8, 80.1, 65.8, 52.1 and 31.4 ppm.

IR (neat): 3369, 2962, 1653, 1601, 1417, 1277, 1101, 748, 640, 573 and 430 cm⁻¹.

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

Methyl 4-(((1Z,3E)-3-(2-hydroxy-1-iodo-2-methylpropylidene)isobenzofuran-1(3H)-ylidene)amino)benzoate (15e)



To a solution of coupled product **14e** (30 mg, 0.089 mmol) in dry DCM (3 mL), was added NIS (24 mg, 0.106 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **15e** (39 mg, 0.084 mmol, 95 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.86 (1 H, d, *J* = 8.0 Hz), 8.06 (2 H, d, *J* = 8.3 Hz), 8.02 (1 H, d, *J* = 7.6 Hz), 7.69 (1 H, t, *J* = 7.7 Hz), 7.62 (1 H, t, *J* = 7.5 Hz), 7.21 (2 H, d, *J* = 8.3 Hz), 3.92 (3 H, s), 2.97 (1 H, br, s), 1.52 (6 H, s) ppm.

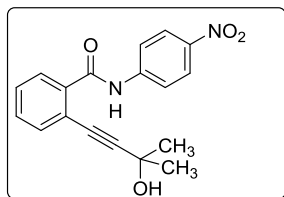
¹³C NMR (100 MHz, CDCl₃): δ = 166.9, 152.8, 150.8, 145.7, 136.8, 132.5, 130.79, 130.73, 130.5, 126.2, 125.5, 124.1, 122.3, 95.0, 74.7, 52.1 and 30.1 ppm.

IR (neat): 3055, 2986, 2925, 1713, 1600, 1423, 1266, 1109, 1015, 896 and 736 cm⁻¹.

HR ESI-MS: [C₂₀H₁₉INO₄]⁺ = [M+H]⁺ requires 464.0359; found 464.0342.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

2-(3-Hydroxy-3-methylbut-1-yn-1-yl)-N-(4-nitrophenyl)benzamide (14f)



To a solution of 2-iodo-N-(4-nitrophenyl)benzamide³ (300 mg, 0.817 mmol) and 2-methylbut-3-yn-2-ol **xx** (82 mg, 0.980 mmol) in dry THF (10 mL) and Et₃N (2 mL) were added [(Ph₃P)₂PdCl₂] (6 mg, 0.008 mmol) and CuI (23 mg, 0.122 mmol) under nitrogen. The reaction was

stirred at 60 °C for 14 h. purification by flash column chromatography (8:2, hexane: EtOAc) gave the coupled product **14f** (230 mg, 0.709 mmol, 87 %) as a yellow solid.

¹H NMR (400 MHz, CDCl₃): δ = 9.61 (1 H, br), 8.15 (2 H, d, *J* = 9.0 Hz), 8.08 (1 H, t, *J* = 5.2 Hz), 7.96 (2 H, d, *J* = 9.0 Hz), 7.45-7.47 (3 H, m), 2.82 (1 H, br), 1.64 (6 H, s) ppm.

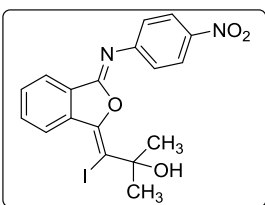
¹³C NMR (100 MHz, CDCl₃): δ = 164.6, 143.77, 143.73, 134.3, 133.9, 131.7, 130.6, 129.4, 125.1, 119.7, 119.1, 102.3, 80.1, 65.9 and 31.5 ppm.

IR (neat): 3408, 3061, 2985, 1672, 1612, 1527, 1429, 1325, 1263, 1167, 1134, 897, 748 and 546 cm⁻¹.

HR ESI-MS: [C₁₈H₁₆NaN₂O₄]⁺ = [M+Na]⁺ requires 347.1008; found 347.1034.

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

1-Iodo-2-methyl-1-((1E,3Z)-3-((4-nitrophenyl)imino)isobenzofuran-1(3H)-ylidene)propan-2-ol (**15f**)



To a solution of coupled product **14f** (30 mg, 0.09 mmol) in dry DCM (3 mL), was added NIS (25 mg, 0.11 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **15f**

(37 mg, 0.082 mmol, 92 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.81 (1 H, d, *J* = 8.0 Hz), 8.19 (2 H, d, *J* = 8.8 Hz), 7.95 (1 H, d, *J* = 7.4 Hz), 7.66 (1 H, t, *J* = 7.6 Hz), 7.59 (1 H, t, *J* = 7.3 Hz), 7.19 (2 H, d, *J* = 8.3 Hz), 2.73 (1 H, br, s), 1.45 (6 H, s) ppm.

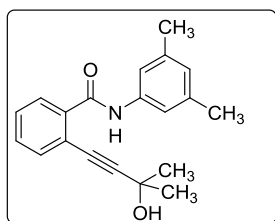
¹³C NMR (100 MHz, CDCl₃): δ = 153.5, 152.8, 145.5, 144.4, 136.9, 132.9, 130.8, 130.2, 125.7, 124.9, 124.3, 123.0, 96.5, 74.6 and 30.0 ppm.

IR (neat): 3055, 2984, 2926, 2854, 1698, 1589, 1515, 1454, 1428, 1341, 1266, 1108, 1024, 891, 854, 742, 463 and 430 cm⁻¹.

HR ESI-MS: [C₁₈H₁₆IN₂O₄]⁺ = [M+H]⁺ requires 451.0155; found 451.0166.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

N-(3,5-Dimethylphenyl)-2-(3-hydroxy-3-methylbut-1-yn-1-yl)benzamide (**14g**)



To a solution of N-(3,5-dimethylphenyl)-2-iodobenzamide⁴ (500 mg, 1.42 mmol) and 2-methylbut-3-yn-2-ol (143 mg, 1.02 mmol) in dry THF (10 mL) and Et₃N (3 mL) were added [(Ph₃P)₂PdCl₂] (10 mg, 0.0142 mmol) and CuI (40 mg, 0.213 mmol) under nitrogen. The reaction was

stirred at 60 °C for 14 h. purification by flash column chromatography (8:2, hexane: EtOAc) gave the coupled product **14g** (360 mg, 1.17 mmol, 82 %) as a yellow oil

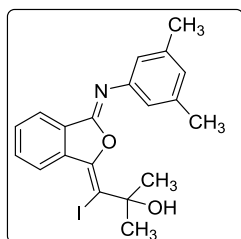
¹H NMR (400 MHz, CDCl₃): δ = 8.88 (1 H, br, s), 8.00 (1 H, dd, *J* = 2.2 & 6.7 Hz), 7.36-7.45 (3 H, m), 7.35 (2 H, s), 6.74 (1 H, s), 3.00 (1 H, br, s), 2.28 (6 H, s), 1.58 (6 H, s) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 164.5, 138.7, 137.7, 136.0, 133.6, 130.7, 129.9, 128.9, 126.4, 119.2, 117.9, 101.3, 80.1, 65.5, 31.3 and 21.4 ppm.

IR (neat): 3377, 3051, 2978, 2933, 2862, 1659, 1549, 1454, 1273, 1161, 957, 897, 737 and 436 cm⁻¹.

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

1-((1E,3Z)-3-((3,5-Dimethylphenyl)imino)isobenzofuran-1(3H)-ylidene)-1-iodo-2-methylpropan-2-ol (**15g**)



To a solution of coupled product **14g** (30 mg, 0.097 mmol) in dry DCM (3 mL), was added NIS (26 mg, 0.117 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **15g** (36 mg, 0.083 mmol, 86 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.75 (1 H, d, *J* = 8.0 Hz), 7.95 (1H, d, *J* = 7.3 Hz), 7.57 (1 H, dd, *J* = 1.2 & 7.4 Hz), 7.50 (1 H, dd, *J* = 0.9 & 7.5 Hz), 6.79 (2 H, s), 6.72 (1 H, s), 3.14 (1 H, br), 2.24 (6 H, s), 1.51 (6 H, s) ppm.

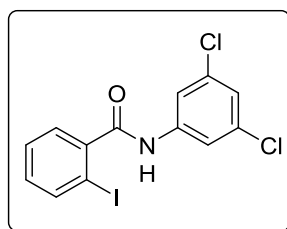
¹³C NMR (100 MHz, CDCl₃): δ = 151.7, 146.2, 145.6, 138.5, 136.5, 131.9, 131.2, 130.5, 126.6, 125.4, 123.9, 120.6, 92.8, 74.8, 30.3 and 21.4 ppm.

IR (neat): 3392, 3052, 2981, 2926, 2862, 1691, 1598, 1466, 1365, 1264, 1193, 1160, 1106, 1030, 935, 895, 849, 734 and 413 cm⁻¹.

HR ESI-MS: [C₂₀H₂₁INO₂]⁺ = [M+H]⁺ requires 434.0617; found 434.0586.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

N-(3,5-Dichlorophenyl)-2-iodobenzamide (**S1**)



To an ice cold solution of 2-iodo benzoic acid (300 mg, 1.209 mmol) in dry DCM (10 mL) and 3,5dichloro aniline (215 mg, 1.33 mmol) were added DCC (298 mg, 1.45 mmol) and DMAP (29 mg, 0.24 mmol) under nitrogen. The reaction was stirred at room temperature for 12 h. The reaction mixture was diluted water and DCM (10 mL). Aqueous

layer was extracted DCM (2 x 10 mL). The combined organic layers were washed with brine (10 mL), and dried over Na₂SO₄. Evaporation of the solvent and purification of the crude mixture by column chromatography (9:1, hexane: EtOAc) gave the acyl protected product **S1** (250 mg, 0.64 mmol, 53 %) as a white solid.

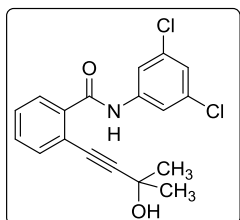
¹H NMR (400 MHz, CDCl₃): δ = 7.89 (1 H, d, J = 7.9 Hz), 7.66 (1 H, br, s), 7.58 (2 H, s), 7.47 (1 H, d, J = 6.4 Hz), 7.42 (1 H, t, J = 7.4 Hz), 7.13-7.17 (2 H, m) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 167.4, 141.3, 140.2, 139.3, 135.5, 132.0, 128.7, 128.5, 125.0, 118.4 and 92.3 ppm.

IR (neat): 3271, 3109, 2868, 1645, 1585, 1531, 1425, 1275, 1014, 945, 851 and 646 cm⁻¹.

TLC: R_f = 0.5 (9:1; hexane:EtOAc).

N-(3,5-Dichlorophenyl)-2-(3-hydroxy-3-methylbut-1-yn-1-yl)benzamide (**14h**)



To a solution of N-(3,5-dichlorophenyl)-2-iodobenzamide **S1** (300 mg, 0.769 mmol) and 2-methylbut-3-yn-2-ol (78 mg, 0.92 mmol) in dry THF (10 mL) and Et₃N (2 mL) were added [(Ph₃P)₂PdCl₂] (6 mg, 0.007 mmol) and CuI (28 mg, 0.28 mmol) under nitrogen. The reaction was stirred at 60 °C for 14 h. purification by flash column chromatography (8:2, hexane: EtOAc) gave the coupled product **14h** (215 mg, 0.619 mmol, 80 %) as a yellow oil.

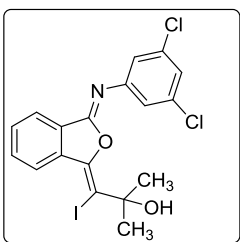
¹H NMR (400 MHz, CDCl₃): δ = 9.22 (1 H, br, s), 8.08 (1 H, dd, J = 4.2 & 7.0 Hz), 7.77 (2 H, s), 7.46-7.53 (3 H, m), 7.10 (1 H, t, J = 1.6 Hz), 2.29 (1 H, br, s), 1.65 (6 H, s) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 164.4, 139.7, 135.3, 133.8, 131.4, 130.5, 129.3, 124.5, 121.6, 119.0, 118.5, 102.1, 80.2, 65.9 and 31.5 ppm.

IR (neat): 3406, 3086, 2968, 2924, 2870, 1660, 1583, 1425, 1257, 1128, 928, 839, 741, 617, 544 and 480 cm⁻¹.

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

1-((1E,3Z)-3-((3,5-Dichlorophenyl)imino)isobenzofuran-1(3H)-ylidene)-1-iodo-2-methylpropan-2-ol (**15h**)



To a solution of coupled product **14h** (30 mg, 0.086 mmol) in dry DCM (3 mL), was added NIS (23 mg, 0.103 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane:EtOAc) gave the iodo cyclic imide **15h** (30 mg, 0.063 mmol, 74 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.86 (1 H, d, *J* = 7.6 Hz), 7.98 (1 H, d, *J* = 7.2 Hz), 7.69 (1 H, t, *J* = 7.4 Hz), 7.61 (1 H, t, *J* = 6.8 Hz), 7.14 (3 H, s), 2.80 (1 H, br, s), 1.60 (6 H, s) ppm.

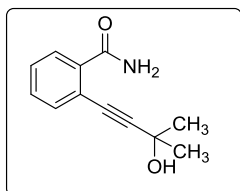
¹³C NMR (100 MHz, CDCl₃): δ = 153.6, 148.1, 145.5, 136.8, 135.0, 132.6, 130.7, 130.6, 125.6, 124.7, 124.2, 121.7, 96.6, 74.5 and 29.9 ppm.

IR (neat): 3059, 2985, 1693, 1579, 1435, 1265, 1109, 1020, 910, 735, 546 and 428 cm⁻¹.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

M.P: 95-97 °C

2-(3-Hydroxy-3-methylbut-1-yn-1-yl)benzamide (14i)



To a solution of 2-iodo benzamide⁴ (200 mg, 0.809mmol) and 2-methylbut-3-yn-2-ol (77 mg, 0.92mmol) in dry THF (10 mL) and DIPA (1 mL) were added [(Ph₃P)₂PdCl₂] (6 mg, 0.0089 mmol) and CuI (23 mg, 0.121 mmol) under nitrogen. The reaction was stirred at 60 °C for 14 h. purification by flash column chromatography (6:4, hexane: EtOAc) gave the coupled product **14i** (130 mg, 0.64mmol, 79 %) as a yellow oil.

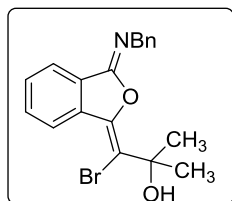
¹H NMR (400 MHz, CDCl₃): δ = 7.99 (1 H, s), 7.45 (1 H, s), 7.39 (2 H, t, *J* = 5.2 Hz), 6.37 (1 H, s), 3.50 (1 H, s), 1.62 (6 H, s) ppm.

¹³C NMR (125 MHz, CDCl₃): δ = 169.2, 134.8, 133.3, 130.8, 129.6, 128.5, 120.0, 100.7, 80.2, 65.2 and 31.0 ppm.

IR (neat): 3454, 2980, 2931, 1642, 1597, 1450, 1385, 1261, 1165, 964, 910, 760, 715, 622, 576, 504 and 467 cm⁻¹.

TLC: R_f = 0.3 (3:2; hexane:EtOAc).

1-((1E,3E)-3-(Benzylimino)isobenzofuran-1(3H)-ylidene)-1-bromo-2-methylpropan-2-ol (13a)



To a solution of coupled product **10a** (30 mg, 0.102 mmol) in dry DCM (3 mL), was added NBS (22 mg, 0.122 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the bromocyclic imidate **13a** (30 mg, 0.08 mmol, 79 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.50 (1 H, dd, *J* = 0.6 & 7.9 Hz), 7.94 (1 H, dd, *J* = 0.8 & 7.6 Hz), 7.59 (1 H, td, *J* = 1.1 & 8.5 Hz), 7.52 (1 H, td, *J* = 1.0 & 7.6 Hz), 7.40 (2 H, d, *J* = 7.6 Hz),

7.34 (2 H, t, $J = 7.3$ Hz), 7.25 (1 H, t, $J = 7.1$ Hz), 4.80 (2 H, s), 3.26 (1 H, br, s), 1.66 (6 H, s) ppm.

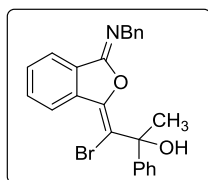
^{13}C NMR (100 MHz, CDCl_3): $\delta = 153.9, 144.6, 139.6, 136.3, 132.0, 130.5, 130.3, 128.6, 127.6, 127.0, 125.4, 123.5, 114.8, 74.7, 52.0$ and 29.6 ppm.

IR (neat): 3443, 3055, 2986, 2912, 1785, 1685, 1647, 1423, 1266, 1018, 896, 743, 706, 481 and 430 cm^{-1} .

HR ESI-MS: $[\text{C}_{19}\text{H}_{19}\text{BrNO}_2]^+ = [\text{M}+\text{H}]^+$ requires 372.0599; found 372.0619.

TLC: $R_f = 0.6$ (4:1; hexane:EtOAc)

1-((1E,3E)-3-(Benzylimino)isobenzofuran-1(3H)-ylidene)-1-bromo-2-phenylpropan-2-ol (13b)



To a solution of coupled product **10b** (30 mg, 0.084 mmol) in dry DCM (3 mL), was added NBS (18 mg, 0.10 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the bromo cyclic imide **13b** (35 mg, 0.08 mmol, 97 %) as a yellow oil.

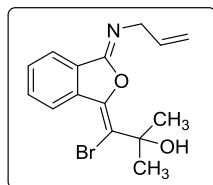
^1H NMR (400 MHz, CDCl_3): $\delta = 8.51$ (1 H, d, $J = 8.0$ Hz), 7.88 (1 H, d, $J = 7.6$ Hz), 7.58 (1 H, t, $J = 7.8$ Hz), 7.49-7.53 (3 H, m), 7.22-7.33 (8 H, m), 4.46 (2 H, d, $J = 3.0$ Hz), 3.55 (1 H, br, s), 1.93 (3 H, s) ppm.

^{13}C NMR (100 MHz, CDCl_3): $\delta = 153.5, 147.4, 145.9, 139.4, 136.0, 131.9, 130.7, 130.5, 128.6, 128.4, 127.6, 127.3, 126.9, 125.3, 124.8, 123.5, 113.4, 78.1, 51.9$ and 31.5 ppm.

IR (neat): 3055, 2986, 2932, 2858, 1785, 1702, 1647, 1608, 1423, 1265, 1007, 896, 742, 451 and 415 cm^{-1} .

TLC: $R_f = 0.6$ (4:1; hexane:EtOAc).

1-((1E,3Z)-3-(Allylimino)isobenzofuran-1(3H)-ylidene)-1-bromo-2-methylpropan-2-ol (13c)



To a solution of coupled product **14a** (30 mg, 0.123 mmol) in dry DCM (3 mL), was added NBS (26 mg, 0.147 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the bromocyclic imide **13c** (25 mg, 0.077 mmol, 63 %) as a yellow oil.

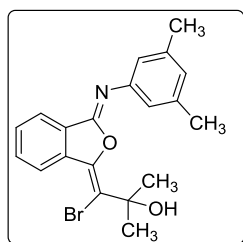
¹H NMR (400 MHz, CDCl₃): δ = 8.50 (1 H, d, J = 8.0 Hz), 7.91 (1 H, d, J = 7.6 Hz), 7.60 (1 H, td, J = 1.1 & 7.6 Hz), 7.53 (1 H, td, J = 7.5 Hz), 6.01-6.11 (1 H, m), 5.28 (1 H, dd, J = 1.7 & 17.1 Hz), 5.17 (1 H, dd, J = 1.6 & 10.2 Hz), 4.24 (2 H, dt, J = 1.6 & 5.5 Hz), 1.68 (6 H, s) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 154.0, 144.6, 136.2, 135.4, 132.0, 130.3, 125.4, 125.3, 123.5, 115.8, 114.7, 74.8, 50.9 and 29.6 ppm.

IR (neat): 3444, 3055, 2986, 2928, 1786, 1701, 1653, 1423, 1266, 1102, 1017, 896, 740 and 466 cm⁻¹

TLC: R_f = 0.6 (4:1; hexane:EtOAc)

1-Bromo-1-((1E,3Z)-3-((3,5-dimethylphenyl)imino)isobenzofuran-1(3H)-ylidene)-2-methylpropan-2-ol (13d)



To a solution of coupled product **14g** (30 mg, 0.097mmol) in dry DCM (5 mL), was added NBS (21 mg, 0.116mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the bromo cyclic imide **13d** (35 mg, 0.09 mmol, 93 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.43 (1 H, d, J = 7.9 Hz), 7.95 (1 H, d, J = 7.3 Hz), 7.56 (1 H, dd, J = 1.2 & 7.5 Hz), 7.50 (1 H, dd, J = 1.0 & 7.5 Hz), 6.80 (2 H, s), 6.72 (1 H, s), 3.19 (1 H, br), 2.24 (6 H, s), 1.50 (6 H, s) ppm.

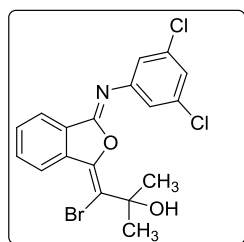
¹³C NMR (100 MHz, CDCl₃): δ = 151.9, 145.4, 144.6, 138.4, 136.0, 132.3, 130.6, 130.3, 126.6, 125.2, 123.8, 120.6, 115.1, 74.7, 29.7 and 21.3 ppm.

IR (neat): 3380, 3054, 2984, 2926, 1694, 1596, 1459, 1423, 1376, 1265, 1198, 1151, 1099, 1017, 896, 740 and 452 cm⁻¹

HR ESI-MS: [C₂₀H₂₁BrNO₂]⁺ = [M+H]⁺ requires 386.0756; found 386.0757

TLC: R_f = 0.6 (4:1; hexane:EtOAc)

1-Bromo-1-((1E,3Z)-3-((3,5-dichlorophenyl)imino)isobenzofuran-1(3H)-ylidene)-2-methylpropan-2-ol (13e)



To a solution of coupled product **14h** (30 mg, 0.086mmol) in dry DCM (3 mL), was added NBS (18 mg, 0.103mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane:EtOAc) gave the bromo cyclic imide **13e** (25 mg, 0.058 mmol, 68 %) as a yellow oil.

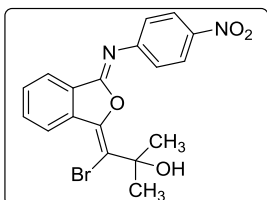
¹H NMR (400 MHz, CDCl₃): δ = 8.47 (1 H, d, J = 7.9 Hz), 7.92 (1 H, d, J = 7.5 Hz), 7.61 (1 H, t, J = 7.3 Hz), 7.53 (1 H, t, J = 7.0 Hz), 7.07 (3 H, s), 2.78 (1 H, br), 1.52 (6 H, s) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 153.5, 147.9, 144.3, 140.2, 136.4, 135.1, 133.1, 130.6, 125.5, 124.8, 124.1, 121.8, 117.6, 74.6 and 29.8 ppm.

IR (neat): 3061, 2978, 2941, 2852, 1695, 1579, 1435, 1265, 1028, 733 and 428 cm⁻¹

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

1-Bromo-2-methyl-1-((1E,3Z)-3-((4-nitrophenyl)imino)isobenzofuran-1(3H)-ylidene)propan-2-ol (13f)



To a solution of coupled product **14f** (30 mg, 0.09mmol) in dry DCM (3 mL), was added NBS (19 mg, 0.108mmol) and the reaction mixture was stirred at room temperature for 30 min.purificationby flash column chromatography (9:1, hexane:EtOAc) gave the bromo cyclic imidate **13f** (36 mg, 0.089 mmol, 98 %) as a yellow solid.

¹H NMR (400 MHz, CDCl₃): δ = 8.47 (1 H, d, J = 8.0 Hz), 8.16 (2 H, d, J = 9.0 Hz), 7.95 (1 H, d, J = 7.5 Hz), 7.63 (1 H, t, J = 7.3 Hz), 7.55 (1 H, t, J = 7.4 Hz), 7.20 (2 H, d, J = 8.8 Hz), 2.85 (1 H, br, s), 1.45 (6 H, s) ppm.

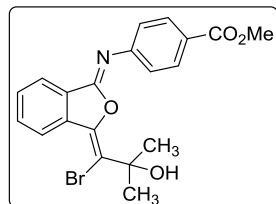
¹³C NMR (100 MHz, CDCl₃): δ = 153.9, 152.5, 144.5, 144.2, 136.5, 133.3, 130.7, 129.7, 125.5, 124.9, 124.3, 123.1, 117.7, 74.8 and 29.8 ppm.

IR (neat): 3421, 3228, 3082, 2929, 2008, 1819, 1666, 1589, 1506, 1335, 1105, 1034, 852, 754 and 619 cm⁻¹

TLC: R_f = 0.6 (4:1; hexane:EtOAc)

M.P: 142-144 °C

Methyl4-(((1Z,3E)-3-(1-bromo-2-hydroxy-2-methylpropylidene)isobenzofuran-1(3H)-ylidene)amino)benzoate (13g)



To a solution of coupled product **14e** (30 mg, 0.089mmol) in dry DCM (3 mL), was added NBS (19 mg, 0.106mmol) and the reaction mixture was stirred at room temperature for 30 min.purificationby flash column chromatography (9:1, hexane:EtOAc) gave the bromo cyclic imidate **13g** (35 mg, 0.084mmol, 94 %) as a yellow oil.

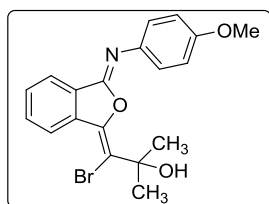
¹H NMR (400 MHz, CDCl₃): δ = 8.55 (1 H, d, *J* = 7.7 Hz), 8.06 (3 H, t, *J* = 8.2 Hz), 7.69 (1 H, t, *J* = 7.4 Hz), 7.62 (1 H, t, *J* = 7.3 Hz), 7.22 (2 H, d, *J* = 7.9 Hz), 3.92 (3 H, s), 2.97 (1 H, br, s), 1.53 (6 H, s) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 166.9, 153.1, 150.5, 144.4, 136.4, 132.9, 130.8, 130.5, 130.1, 126.3, 125.4, 124.1, 122.4, 116.7, 74.8, 52.1 and 29.8 ppm.

IR (neat): 3055, 2985, 2954, 2930, 1786, 1711, 1599, 1436, 1267, 1171, 1110, 1025, 896, 743, 518 and 464 cm⁻¹

TLC: R_f = 0.6 (4:1; hexane:EtOAc)

1-Bromo-1-((1E,3Z)-3-((4-methoxyphenyl)imino)isobenzofuran-1(3H)-ylidene)-2-methylpropan-2-ol (13h)



To a solution of coupled product **14d** (30 mg, 0.097mmol) in dry DCM (3 mL), was added NBS (21 mg, 0.116mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane:EtOAc) gave the bromo cyclic imide **13h** (34 mg, 0.087 mmol, 90 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.44 (1 H, d, *J* = 7.8 Hz), 7.93 (1 H, dd, *J* = 1.0 & 7.1 Hz), 7.54 (1 H, td, *J* = 1.3 & 7.4 Hz), 7.49 (1 H, td, *J* = 1.4 & 7.5 Hz), 7.18 (2 H, d, *J* = 9.0 Hz), 6.83 (2 H, d, *J* = 9.0 Hz), 3.74 (3 H, s), 3.24 (1 H, br, s), 1.52 (6 H, s) ppm.

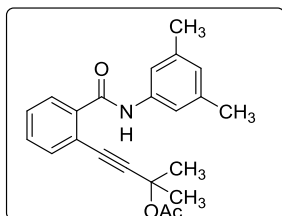
¹³C NMR (100 MHz, CDCl₃): δ = 157.1, 151.3, 144.8, 138.5, 135.9, 132.2, 130.9, 130.4, 125.3, 124.5, 123.8, 115.0, 114.3, 75.0, 55.5 and 29.9 ppm.

IR (neat): 3423, 3056, 2985, 2926, 2842, 1695, 1619, 1503, 1430, 1265, 1162, 1026, 896 and 547 cm⁻¹

TLC: R_f = 0.6 (4:1; hexane:EtOAc)

M.P: 78-80 °C

4-(2-((3,5-Dimethylphenyl)carbamoyl)phenyl)-2-methylbut-3-yn-2-yl acetate (16a)



To an ice cold solution of N-(3,5-dimethylphenyl)-2-(3-hydroxy-3-methylbut-1-yn-1-yl)benzamide **14g** (100 mg, 0.325 mmol) in dry DCM (5 mL) and Et₃N (1 mL) were added Ac₂O (50 mg, 0.488 mmol) and DMAP (8 mg, 0.065mmol) under nitrogen. The reaction was stirred at same temperature for 2h. The reaction mixture was diluted water and DCM (10 mL). Aqueous layer was extracted DCM (2 x 10 mL). The combined organic layers were

washed with brine (10 mL), and dried over Na₂SO₄. Evaporation of the solvent and purification of the crude mixture by column chromatography (9:1, hexane: EtOAc) gave the acyl protected product **16a** (110 mg, 0.315 mmol, 97 %) as a yellow solid.

¹H NMR (400 MHz, CDCl₃): δ = 8.79 (1 H, br, s), 8.03 (1 H, dd, J = 2.2 & 7.6 Hz), 7.52-7.54 (1 H, m), 7.42-7.45 (2 H, m), 7.36 (2 H, s), 6.79 (1 H, s), 2.32 (6 H, s), 1.93 (3 H, s), 1.70 (6 H, s) ppm.

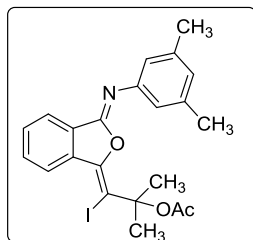
¹³C NMR (100 MHz, CDCl₃): δ = 169.6, 164.7, 138.6, 137.8, 136.5, 133.8, 130.7, 130.3, 129.2, 126.4, 118.9, 118.8, 97.2, 82.4, 72.0, 28.9, 21.8 and 21.5 ppm.

IR (neat): 3055, 2988, 2925, 1739, 1669, 1598, 1553, 1424, 1266, 1191, 1133, 1017, 896, 843 and 737 cm⁻¹

TLC: R_f = 0.5 (9:1; hexane:EtOAc)

M.P: 102-104 °C

1-((1E,3Z)-3-((3,5-Dimethylphenyl)imino)isobenzofuran-1(3H)-ylidene)-1-iodo-2-methylpropan-2-yl acetate (17a**)**



To a solution of coupled product **16a** (30 mg, 0.085mmol) in dry DCM (3 mL), was added NIS (23 mg, 0.102mmol) and the reaction mixture was stirred at room temperature for 1 h. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **17a** (35 mg, 0.08mmol, 94 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.86 (1 H, d, J = 8.0 Hz), 7.99 (1 H, d, J = 7.3 Hz), 7.61 (1 H, td, J = 1.2 & 7.4 Hz), 7.56 (1 H, td, J = 0.9 & 7.5 Hz), 7.04 (2 H, s), 6.80 (1 H, s), 2.32 (6 H, s), 1.89 (3 H, s), 1.77 (6 H, s) ppm.

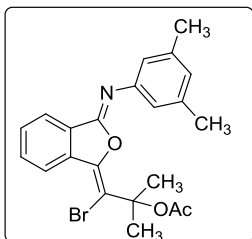
¹³C NMR (100 MHz, CDCl₃): δ = 169.7, 152.0, 146.4, 145.4, 138.1, 136.6, 132.0, 131.6, 130.5, 126.6, 125.7, 123.8, 121.8, 87.3, 82.3, 28.6, 21.6 and 21.4 ppm.

IR (neat): 3050, 2983, 2925, 2858, 1739, 1690, 1598, 1467, 1435, 1368, 1258, 1183, 1136, 1015, 941, 849, 741, 704, 671 and 422 cm⁻¹

HR ESI-MS: [C₂₂H₂₃INO₃]⁺ = [M+H]⁺ requires 476.0717; found 476.0719

TLC: R_f = 0.6 (9:1; hexane:EtOAc)

1-Bromo-1-((1E,3Z)-3-((3,5-dimethylphenyl)imino)isobenzofuran-1(3H)-ylidene)-2-methylpropan-2-yl acetate (17a'**)**



To a solution of coupled product **16a** (30 mg, 0.085mmol) in dry DCM (3 mL), was added NBS (18 mg, 0.102mmol) and the reaction mixture was stirred at room temperature for 1.5 h. purification by flash column chromatography (9:1, hexane:EtOAc) gave the bromo cyclic imide **17a'** (29 mg, 0.07 mmol, 82 %) as a yellow solid.

¹H NMR (400 MHz, CDCl₃): δ = 8.46 (1 H, d, J = 7.8 Hz), 7.91 (1 H, td, J = 1.0 & 6.8 Hz), 7.45-7.56 (2 H, m), 6.98 (2 H, s), 6.72 (1 H, s), 2.24 (6 H, s), 1.79 (3 H, s), 1.68 (6 H, s) ppm.

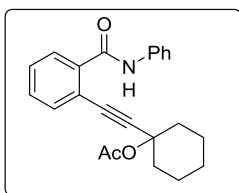
¹³C NMR (100 MHz, CDCl₃): δ = 169.7, 152.3, 145.2, 138.1, 136.1, 132.1, 130.3, 126.7, 125.5, 124.2, 123.7, 122.0, 121.9, 111.1, 81.4, 27.9, 21.4 and 21.3 ppm.

IR (neat): 3054, 2986, 2927, 2858, 1737, 1692, 1627, 1596, 1465, 1423, 1369, 1265, 1140, 1038, 1017, 896, 708 and 421 cm⁻¹

TLC: R_f = 0.6 (9:1; hexane:EtOAc)

M.P: 63-65 °C

1-((2-(Phenylcarbamoyl)phenyl)ethynyl)cyclohexyl acetate (**16b**)



To an ice cold solution of 2-((1-hydroxycyclohexyl)ethynyl)-N-phenylbenzamide **14c** (100 mg, 0.313mmol) in dry DCM (5 mL) and Et₃N (1 mL) were added Ac₂O (51 mg, 0.47mmol) and DMAP (8 mg, 0.06mmol) under nitrogen. The reaction was stirred at same temperature

for 2h. The reaction mixture was diluted with water and DCM (10 mL). Aqueous layer was extracted with DCM (2 x 10 mL). The combined organic layers were washed with brine (10 mL), and dried over Na₂SO₄. Evaporation of the solvent and purification of the crude mixture by column chromatography (9:1, hexane: EtOAc) gave the acyl protected product **16b** (102 mg, 0.282 mmol, 90 %) as a yellow oil.

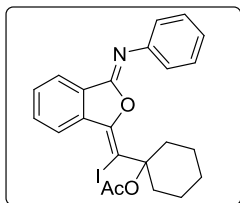
¹H NMR (400 MHz, CDCl₃): δ = 8.94 (1 H, br, s), 8.01 (1 H, d, J = 7.1 Hz), 7.71 (2 H, d, J = 7.8 Hz), 7.52-7.54 (1 H, m), 7.43 (2 H, t, J = 3.7 Hz), 7.35 (2 H, t, J = 7.5 Hz), 7.13 (1 H, t, J = 7.4 Hz), 2.12-2.15 (2 H, m), 1.92 (3 H, s), 1.79-1.82 (2 H, m), 1.55-1.56 (4 H, m), 1.44-1.47 (1 H, m), 1.25-1.28 (1 H, m) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 169.5, 165.0, 137.9, 136.6, 133.6, 130.6, 130.2, 129.1, 128.8, 124.6, 121.3, 119.1, 96.1, 84.4, 75.6, 36.9, 25.0, 22.6 and 21.8 ppm.

IR (neat): 3377, 3142, 3057, 2940, 2861, 1737, 1667, 1599, 1444, 1369, 1321, 1267, 1235, 1137, 1021, 960, 911, 741 and 699 cm⁻¹

TLC: $R_f = 0.5$ (9:1; hexane:EtOAc)

1-(Iodo((1E,3Z)-3-(phenylimino)isobenzofuran-1(3H)-ylidene)methyl)cyclohexyl acetate (18a)



To a solution of coupled product **16b** (30 mg, 0.083mmol) in dry DCM (3 mL), was added NIS (22 mg, 0.099mmol) and the reaction mixture was stirred at room temperature for 1 h. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **18a** (38 mg, 0.073 mmol, 89 %) as a yellow semi solid.

^1H NMR (400 MHz, CDCl_3): $\delta = 8.89$ (1 H, d, $J = 8.1$ Hz), 8.00 (1 H, d, $J = 7.7$ Hz), 7.62 (1 H, t, $J = 7.4$ Hz), 7.56 (1 H, t, $J = 6.0$ Hz), 7.41 (2 H, d, $J = 6.6$ Hz), 7.35 (2 H, t, $J = 7.6$ Hz), 7.15 (1 H, td, $J = 1.0$ & 7.4 Hz), 2.50 - 2.53 (2 H, m), 1.85 (3 H, s), 1.76 - 1.82 (3 H, m), 1.50 - 1.64 (5 H, m) ppm.

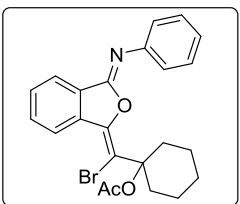
^{13}C NMR (100 MHz, CDCl_3): $\delta = 169.5, 152.5, 146.4, 145.8, 136.8, 131.8, 131.7, 130.5, 128.6, 125.8, 124.9, 124.0, 123.8, 88.7, 83.2, 36.1, 25.0, 22.5$ and 21.2 ppm.

IR (neat): 3055, 2984, 2938, 2860, 1733, 1693, 1593, 1488, 1451, 1423, 1368, 1266, 1234, 1141, 1093, 1017, 966, 894, 740, 707, 521, 447 and 412 cm^{-1}

HR ESI-MS: $[\text{C}_{23}\text{H}_{23}\text{INO}_3]^+ = [\text{M}+\text{H}]^+$ requires 488.0717; found 488.0707

TLC: $R_f = 0.6$ (9:1; hexane:EtOAc)

1-(Bromo((1E,3Z)-3-(phenylimino)isobenzofuran-1(3H)-ylidene)methyl)cyclohexyl acetate (18a')



To a solution of coupled product **16b** (30 mg, 0.083mmol) in dry DCM (3 mL), was added NBS (18 mg, 0.099mmol) and the reaction mixture was stirred at room temperature for 1.5 h. purification by flash column chromatography (9:1, hexane: EtOAc) gave the bromo cyclic imide **18a'**

(36 mg, 0.082 mmol, 98 %) as a yellow solid.

^1H NMR (400 MHz, CDCl_3): $\delta = 8.86$ (1 H, d, $J = 7.8$ Hz), 8.01 (1 H, dd, $J = 1.0$ & 7.0 Hz), 7.61 (1 H, td, $J = 1.3$ & 7.4 Hz), 7.56 (1 H, td, $J = 1.1$ & 7.5 Hz), 7.33 - 7.42 (4 H, m), 7.15 (1 H, tt, $J = 1.3$ & 2.8 Hz), 2.51 - 2.53 (2 H, m), 1.86 (3 H, s), 1.76 - 1.84 (2 H, m), 1.52 - 1.69 (6 H, m) ppm.

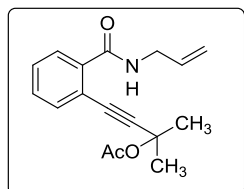
^{13}C NMR (100 MHz, CDCl_3): $\delta = 169.5, 152.9, 145.7, 145.1, 136.4, 132.2, 131.3, 130.3, 128.6, 125.7, 124.9, 124.1, 123.7, 112.1, 82.8, 34.9, 25.1, 22.1$ and 21.1 ppm.

IR (neat): 3055, 2985, 2938, 2860, 1734, 1695, 1623, 1592, 1487, 1449, 1424, 1369, 1266, 1236, 895 and 746 cm^{-1}

TLC: R_f = 0.6 (9:1; hexane:EtOAc)

M.P: 96-98 $^{\circ}\text{C}$

4-(2-(Allylcarbamoyl)phenyl)-2-methylbut-3-yn-2-yl acetate (**16c**)



To an ice cold solution of N-allyl-2-(3-hydroxy-3-methylbut-1-yn-1-yl)benzamide **14a** (100 mg, 0.411 mmol) in dry DCM (5 mL) and Et_3N (1 mL) were added Ac_2O (67 mg, 0.617 mmol) and DMAP (10 mg, 0.08 mmol) under nitrogen. The reaction was stirred at same temperature for 1.5h. The reaction mixture was diluted with water and DCM (10 mL). Aqueous layer was extracted with DCM (2 x 10 mL). The combined organic layers were washed with brine (10 mL), and dried over Na_2SO_4 . Evaporation of the solvent and purification of the crude mixture by column chromatography (9:1, hexane: EtOAc) gave the acyl protected product **16c** (105 mg, 0.368 mmol, 90 %) as a yellow oil.

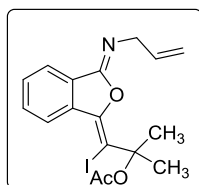
^1H NMR (400 MHz, CDCl_3): δ = 8.15 (1 H, d, J = 7.3 Hz), 8.03 (1 H, br, s), 7.47 (1 H, t, J = 7.1 Hz), 7.41 (2 H, t, J = 8.4 Hz), 5.94-6.04 (1 H, m), 5.27 (1 H, d, J = 17.1 Hz), 5.13 (1 H, d, J = 10.2 Hz), 4.17 (2 H, t, J = 5.8 Hz), 2.05 (3 H, s), 1.73 (6 H, s) ppm.

^{13}C NMR (100 MHz, CDCl_3): δ = 170.0, 165.7, 135.4, 134.7, 133.9, 130.7, 130.4, 129.2, 118.7, 115.7, 96.3, 82.8, 71.9, 42.6, 28.8 and 22.0 ppm.

IR (neat): 3394, 3056, 2989, 2935, 1736, 1653, 1594, 1525, 1475, 1425, 1369, 1266, 1191, 1133, 1017, 969, 928, 738, 557 and 454 cm^{-1}

TLC: R_f = 0.5 (9:1; hexane:EtOAc)

1-((1E,3Z)-3-(Allylimino)isobenzofuran-1(3H)-ylidene)-1-bromo-2-methylpropan-2-ylacetate (**19a**)



To a solution of coupled product **16c** (30 mg, 0.105 mmol) in dry DCM (3 mL), was added N1S (28 mg, 0.126 mmol) and the reaction mixture was stirred at room temperature for 1.5 h. Purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **19a** (25 mg, 0.058 mmol, 55 %) as a yellow oil.

^1H NMR (400 MHz, CDCl_3): δ = 8.84 (1 H, d, J = 8.0 Hz), 7.89 (1 H, d, J = 7.6 Hz), 7.58 (1 H, td, J = 1.2 & 7.4 Hz), 7.51 (1 H, td, J = 1.0 & 7.5 Hz), 6.03-6.10 (1 H, m), 5.28 (1 H, dq, J = 1.7

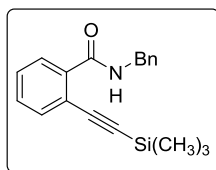
& 17.1 Hz), 5.14 (1 H, dq, $J = 1.6$ & 10.2 Hz), 4.24 (2 H, dt, $J = 1.6$ & 5.6 Hz), 2.08 (3 H, s), 1.87 (6 H, s) ppm.

^{13}C NMR (100 MHz, CDCl_3): $\delta = 169.7, 154.2, 146.3, 136.9, 135.7, 131.5, 131.3, 130.5, 125.8, 123.3, 115.7, 86.8, 82.5, 50.9, 28.4$ and 22.1 ppm.

IR (neat): 2925, 2858, 1780, 1736, 1654, 1459, 1412, 1365, 1254, 1146, 1109, 1012, 932, 838, 762 and 696 cm^{-1}

TLC: $R_f = 0.6$ (9:1; hexane:EtOAc)

N-Benzyl-2-((trimethylsilyl)ethynyl)benzamide (20a)



To a solution of N-benzyl-2-iodobenzamide (200 mg, 0.593 mmol) and trimethylsilylacetylene (70 mg, 0.712 mmol) in dry THF (10 mL) and Et_3N (1 mL) were added $[(\text{Ph}_3\text{P})_2\text{PdCl}_2]$ (4 mg, 0.0059 mmol) and CuI (17 mg, 0.088 mmol) under nitrogen. The reaction was stirred at room temperature for

14 h. purification by flash column chromatography (9:1, hexane: EtOAc) gave the coupled product **20a** (120 mg, 0.390 mmol, 59 %) as a brown oil

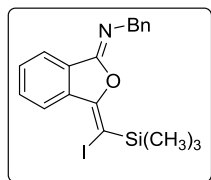
^1H NMR (400 MHz, CDCl_3): $\delta = 8.06\text{--}8.08$ (2 H, m), 8.04 (1 H, br, s), 7.14–7.35 (7 H, m), 4.58 (2 H, d, $J = 5.6$ Hz), -0.0002 (9 H, s) ppm.

^{13}C NMR (100 MHz, CDCl_3): $\delta = 165.5, 138.1, 135.0, 134.1, 130.6, 130.5, 129.2, 128.8, 127.9, 127.6, 119.4, 103.7, 102.2, 44.2$ and -0.3685 ppm.

IR (neat): 3390, 3055, 2985, 1655, 1595, 1534, 1423, 1266, 1146, 1089, 1037, 895, 864, 849, 740, 707, 536 and 469 cm^{-1}

TLC: $R_f = 0.5$ (9:1; hexane:EtOAc)

(1Z,3E)-N-Benzyl-3-(iodo(trimethylsilyl)methylene)isobenzofuran-1(3H)-imine (21a)



To a solution of coupled product **20a** (30 mg, 0.097 mmol) in dry DCM (3 mL), was added NIS (26 mg, 0.117 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **21a** (25

mg, 0.057 mmol, 60 %) as a brown oil.

^1H NMR (400 MHz, CDCl_3): $\delta = 8.94$ (1 H, d, $J = 7.9$ Hz), 7.92 (1 H, d, $J = 7.4$ Hz), 7.60 (1 H, t, $J = 7.4$ Hz), 7.53 (1 H, t, $J = 7.4$ Hz), 7.41 (2 H, d, $J = 7.2$ Hz), 7.34 (2 H, t, $J = 7.3$ Hz), 7.24 (1 H, t, $J = 7.2$ Hz), 4.81 (2 H, s), 0.3952 (9 H, s) ppm.

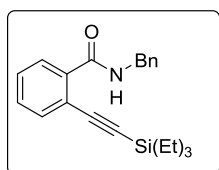
^{13}C NMR (100 MHz, CDCl_3): δ = 155.3, 154.4, 140.0, 136.8, 132.5, 131.4, 130.8, 128.5, 127.8, 126.9, 125.9, 123.4, 79.1, 52.1 and 0.6676 ppm.

IR (neat): 3054, 2982, 2903, 1702, 1589, 1456, 1423, 1350, 1264, 1178, 1104, 1026, 851, 742, 483 and 438 cm^{-1}

HR ESI-MS: $[\text{C}_{19}\text{H}_{21}\text{INOSi}]^+ = [\text{M}+\text{H}]^+$ requires 434.0442; found 434.0442

TLC: R_f = 0.6 (9:1; hexane:EtOAc)

N-Benzyl-2-((triethylsilyl)ethynyl)benzamide (20b)



To a solution of N-benzyl-2-iodobenzamide (200 mg, 0.593mmol) and triethylsilylacetylene (100 mg, 0.712 mmol) in dry THF (10 mL) and Et_3N (1 mL) were added $[(\text{Ph}_3\text{P})_2\text{PdCl}_2]$ (4 mg, 0.0059 mmol) and CuI (17 mg, 0.088mmol) under nitrogen. The reaction was stirred at room temperature for

14 h. purification by flash column chromatography (9:1, hexane: EtOAc) gave the coupled product **20b** (130 mg, 0.372 mmol, 63 %) as a brown oil

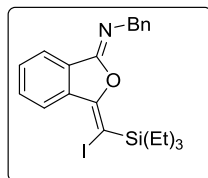
^1H NMR (400 MHz, CDCl_3): δ = 8.16 (1 H, br, s), 8.11 (1 H, dd, J = 1.3 & 7.8 Hz), 7.46 (1 H, d, J = 6.8 Hz), 7.16-7.37 (7 H, m), 4.60 (2 H, d, J = 5.6 Hz), 0.87 (9 H, t, J = 7.9 Hz), 0.46 (6 H, q, J = 7.9 Hz) ppm.

^{13}C NMR (100 MHz, CDCl_3): δ = 165.8, 138.2, 134.8, 134.4, 130.6, 130.5, 129.1, 128.7, 127.8, 127.4, 119.6, 104.9, 100.0, 44.1, 7.53 and 4.14 ppm.

IR (neat): 3385, 3055, 2957, 2916, 2877, 1655, 1593, 1533, 1457, 1420, 1266, 1146, 1093, 1013, 974, 901, 846, 738, 706 and 507 cm^{-1}

TLC: R_f = 0.5 (9:1; hexane:EtOAc)

(1Z,3E)-N-Benzyl-3-(iodo(triethylsilyl)methylene)isobenzofuran-1(3H)-imine (21b)



To a solution of coupled product **20b** (30 mg, 0.085mmol) in dry DCM (3 mL), was added NIS (23 mg, 0.103mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the iodo cyclic imide **21b** (30 mg, 0.063 mmol, 75 %) as a brown oil.

^1H NMR (400 MHz, CDCl_3): δ = 9.00 (1 H, d, J = 7.9 Hz), 7.92 (1 H, d, J = 7.6 Hz), 7.58-7.62 (1 H, m), 7.53 (1 H, t, J = 7.4 Hz), 7.41 (2 H, d, J = 7.5 Hz), 7.34 (2 H, t, J = 7.3 Hz), 7.22-7.26 (1 H, m), 4.81 (2 H, s), 1.01 (9 H, t, J = 6.8 Hz), 0.93 (6 H, t, J = 7.3 Hz) ppm.

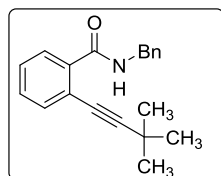
^{13}C NMR (100 MHz, CDCl_3): δ = 155.8, 154.5, 140.0, 136.8, 132.5, 131.3, 130.7, 128.5, 127.8, 126.8, 126.1, 123.4, 79.9, 52.0, 7.66 and 4.97 ppm.

IR (neat): 2953, 2930, 2875, 1781, 1703, 1654, 1586, 1415, 1355, 1264, 1102, 1024, 1001, 821, 730, 696, 669, 504 and 435 cm^{-1}

HR ESI-MS: $[\text{C}_{26}\text{H}_{27}\text{INOSi}]^+ = [\text{M}+\text{H}]^+$ requires 476.0907; found 476.0898

TLC: R_f = 0.6 (9:1; hexane:EtOAc)

N-Benzyl-2-(3,3-dimethylbut-1-yn-1-yl)benzamide (20c)



To a solution of N-benzyl-2-iodobenzamide (200 mg, 0.59 mmol) and 3,3-dimethylbut-1-yne (67 mg, 0.71 mmol) in dry THF (8 mL) and DIPA (2 mL) were added $[(\text{Ph}_3\text{P})_2\text{PdCl}_2]$ (4.0 mg, 0.0059 mmol) and CuI (17 mg, 0.088 mmol) under nitrogen. The reaction was stirred at room temperature for 14 h. The reaction mixture was diluted with saturated aq. NH_4Cl (10 mL) and ethyl acetate (20 mL). Aqueous layer was extracted with ethyl acetate (2 x 50 mL). The combined organic layers were washed with brine (10 mL), and dried over Na_2SO_4 . Evaporation of the solvent and purification of the crude mixture by column chromatography (8:2, hexane: EtOAc) gave the coupled product **20c** (120 mg, 0.412 mmol, 70 %) as a yellow oil.

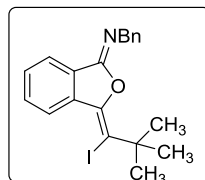
^1H NMR (400 MHz, CDCl_3): δ = 8.14-8.18 (2 H, m), 7.25-7.44 (8 H, m), 4.69 (2 H, s), 1.13 (9 H, s) ppm.

^{13}C NMR (100 MHz, CDCl_3): δ = 166.0, 138.3, 134.3, 133.9, 130.6, 130.5, 128.8, 128.3, 128.2, 127.6, 120.3, 105.7, 78.3, 44.2, 30.5 and 28.3 ppm.

IR (neat): 3385, 3058, 2972, 2873, 1656, 1599, 1533, 1464, 1362, 1271, 1201, 1155, 743, 603, 548, 502 and 484 cm^{-1} .

TLC: R_f = 0.4 (4:1; hexane:EtOAc).

(1E,3E)-N-Benzyl-3-(1-iodo-2,2-dimethylpropylidene)isobenzofuran-1(3H)-imine (21c)



To a solution of coupled product **20c** (30 mg, 0.103 mmol) in dry DCM (3 mL), was added NIS (28 mg, 0.123 mmol) and the reaction mixture was stirred at room temperature for 30 min. The reaction mixture was diluted with water (10 mL) and DCM (20 mL) aqueous layer was extracted with DCM (2 x 20 mL).

The combined organic layers were washed with hypo solution (10 mL) and brine (10 mL), and dried over Na_2SO_4 . Evaporation of the solvent and purification of the crude mixture by column

chromatography (9:1, hexane: EtOAc) gave the five membered iodo cyclic imide **21c** (28 mg, 0.067 mmol, 65%) as a yellow oil.

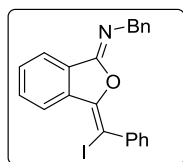
¹H NMR (400 MHz, CDCl₃): δ = 8.91 (1 H, d, *J* = 8.0 Hz), 7.92 (1 H, d, *J* = 7.4 Hz), 7.56 (1 H, t, *J* = 7.3 Hz), 7.48 (1 H, t, *J* = 7.4 Hz), 7.42 (2 H, d, *J* = 7.2 Hz), 7.33 (2 H, t, *J* = 7.2 Hz), 7.24 (1 H, t, *J* = 6.0 Hz), 4.83 (2 H, s), 1.49 (9 H, s) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 154.3, 146.3, 140.0, 137.5, 131.3, 131.0, 129.8, 128.4, 127.7, 126.7, 125.8, 123.2, 95.7, 52.1, 40.1 and 32.1 ppm.

IR (neat): 3404, 3056, 2972, 1775, 1696, 1604, 1464, 1358, 1265, 1109, 1031, 922, 901, 742, 629 and 447 cm⁻¹.

TLC: R_f = 0.6 (4:1; hexane:EtOAc).

(1Z,3E)-N-Benzyl-3-(iodo(phenyl)methylene)isobenzofuran-1(3H)-imine (24a)



To a solution of coupled product⁵ **22** (30 mg, 0.096 mmol) in dry DCM (3 mL), was added NIS (26 mg, 0.115 mmol) and the reaction mixture was stirred at room temperature for 15 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the five membered iodo cyclic imide **24a** (35 mg, 0.08

mmol, 83 %) as a yellow oil. and six membered iodo cyclic imide **24b** (5 mg, 0.011 mmol, 13%) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.0 (1 H, dt, *J* = 0.8 & 8.0 Hz), 7.93 (1 H, d, *J* = 7.6 Hz), 7.63 (1 H, td, *J* = 1.2 & 7.4 Hz), 7.51-7.59 (3 H, m), 7.38 (2 H, t, *J* = 7.3 Hz), 7.20-7.31 (6 H, m), 4.60 (2 H, s) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 154.5, 147.3, 140.7, 139.9, 136.0, 131.8, 131.5, 130.2, 130.1, 128.47, 128.44, 128.18, 128.11, 126.8, 125.0, 123.5, 73.7 and 51.9 ppm.

IR (neat): 3059, 2989, 1772, 1691, 1435, 1267, 1009, 903 and 731 cm⁻¹

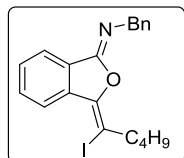
TLC: R_f = 0.5 (9:1; hexane:EtOAc)

(Z)-N-Benzyl-4-iodo-3-phenyl-1H-isochromen-1-imine (24b)

¹H NMR (400 MHz, CDCl₃): δ = 8.32 (1 H, td, *J* = 1.0 & 8.0 Hz), 7.71 (1 H, dd, *J* = 0.7 & 7.9 Hz), 7.60-7.63 (3 H, m), 7.45-7.48 (3 H, m), 7.41-7.43 (3 H, m), 7.32 (2 H, t, *J* = 7.2 Hz), 7.27 (1 H, t, *J* = 4.3 Hz), 4.69 (2 H, s) ppm.

TLC: R_f = 0.6 (9:1; hexane:EtOAc)

(1Z,3E)-N-Benzyl-3-(1-iodopentylidene)isobenzofuran-1(3H)-imine (25a)



To a solution of coupled product⁶ **23** (30 mg, 0.103mmol) in dry DCM (3 mL), was added NIS (28 mg, 0.123 mmol) and the reaction mixture was stirred at room temperature for 30 min. purification by flash column chromatography (9:1, hexane: EtOAc) gave the five membered iodo cyclic imide product **25a** (22 mg, 0.059mmol, 59 %) as a yellow oil. and six membered iodo cyclic imide **25b** (4 mg, 0.007 mmol, 7%) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 8.65 (1 H, d, J = 7.8 Hz), 7.90 (1 H, d, J = 7.4 Hz), 7.57 (1 H, t, J = 7.3 Hz), 7.49 (1 H, t, J = 7.4 Hz), 7.43 (2 H, d, J = 7.0 Hz), 7.33 (2 H, t, J = 7.1 Hz), 7.24 (1 H, t, J = 1.5 Hz), 4.80 (2 H, s), 2.96 (2 H, t, J = 7.1 Hz), 1.61 (2 H, t, J = 7.1 Hz), 1.39 (2 H, q, J = 7.4 Hz), 0.96 (3 H, t, J = 7.2 Hz) ppm.

¹³C NMR (100 MHz, CDCl₃): δ = 154.5, 147.1, 140.2, 135.9, 131.9, 131.3, 129.9, 128.5, 128.0, 126.8, 124.1, 123.3, 81.4, 51.8, 39.2, 31.2, 21.6 and 14.0 ppm.

IR (neat): 3057, 2976, 1655, 129, 1714, 901 and 731 cm⁻¹

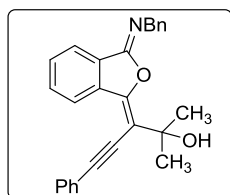
TLC: R_f = 0.5 (9:1; hexane:EtOAc)

(E)-N-Benzyl-3-butyl-4-iodo-1H-isochromen-1-imine (25b)

¹H NMR (400 MHz, CDCl₃): δ = 8.22 (1 H, t, J = 7.4 Hz), 7.22-7.29 (1 H, m), 7.51-7.54 (2 H, m), 7.44 (1 H, d, J = 7.2 Hz), 7.31-7.35 (3 H, m), 7.23 (1 H, t, J = 7.3 Hz), 4.69 (2 H, s), 2.81-2.95 (2 H, m), 1.64-1.75 (2 H, m), 1.41-1.46 (2 H, m), 0.94-0.98 (3 H, m) ppm.

TLC: R_f = 0.6 (9:1; hexane:EtOAc)

3-((1Z,3E)-3-(Benzylimino)isobenzofuran-1(3H)-ylidene)-2-methyl-5-phenylpent-4-yn-2-ol (27)



To a solution of 1-((1E,3Z)-3-(Benzylimino)isobenzofuran-1(3H)-ylidene)-1-iodo-2-methylpropan-2-ol **11a** (150 mg, 0.357 mmol) and phenyl acetylene (44 mg, 0.429mmol) in Et₃N (5 ml) were added [(Ph₃P)₂PdCl₂] (2.4 mg, 0.0035 mmol) and CuI (10 mg, 0.05mmol) under nitrogen. The

reaction was stirred at 60 °C for 14 h. purification by flash column chromatography (9:1, hexane: EtOAc) gave the coupled product **27** (100 mg, 0.254 mmol, 71 %) as a yellow oil.

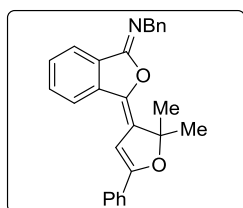
¹H NMR (500 MHz, CDCl₃): δ = 8.45 (1 H, d, J = 7.9 Hz), 7.94 (1 H, d, J = 7.7 Hz), 7.59 (1 H, t, J = 7.3 Hz), 7.53-7.55 (2 H, m), 7.50 (1 H, t, J = 7.5 Hz), 7.33-7.43 (7 H, m), 7.25 (1 H, t, J = 7.4 Hz), 4.87 (2 H, s), 3.24 (1 H, br, s), 1.68 (6 H, s) ppm.

¹³C NMR (125 MHz, CDCl₃): δ = 153.9, 151.5, 139.6, 136.3, 132.2, 131.4, 130.2, 129.7, 128.8, 128.75, 128.70, 127.7, 127.0, 123.8, 123.3, 123.2, 109.1, 99.6, 85.8, 72.8, 52.2 and 29.6 ppm.

IR (neat): 3055, 2986, 2931, 1784, 1732, 1704, 1655, 1611, 1423, 1266, 1078, 896, 741, 707, 606 and 527 cm⁻¹

TLC: R_f = 0.6 (4:1; hexane:EtOAc)

(1E,3Z)-N-Benzyl-3-(2,2-dimethyl-5-phenylfuran-3(2H)-ylidene)isobenzofuran-1(3H)-imine (26)



To a solution of coupled product **27** (30 mg, 0.076mmol) in dry DCM (3 mL), was added AgOTf (5 mg, 0.019mmol) and the reaction mixture was stirred at room temperature for 14 h. purification by flash column chromatography (9:1, hexane: EtOAc) gave the cyclised product **26** (25 mg, 0.063 mmol, 82 %) as a yellow oil.

¹H NMR (400 MHz, CDCl₃): δ = 7.86 (1 H, d, J = 7.7 Hz), 7.65 (2 H, dd, J = 1.8 & 8.2 Hz), 7.59 (1 H, d, J = 7.9 Hz), 7.48 (1 H, td, J = 1.0 & 8.1 Hz), 7.42 (1 H, d, J = 7.5 Hz), 7.35 (2 H, t, J = 7.0 Hz), 7.24-7.33 (4 H, m), 7.17 (1 H, t, J = 6.4 Hz), 7.04-7.10 (1 H, m), 6.33 (1 H, s), 4.77 (2 H, s), 1.70 (6 H, s) ppm.

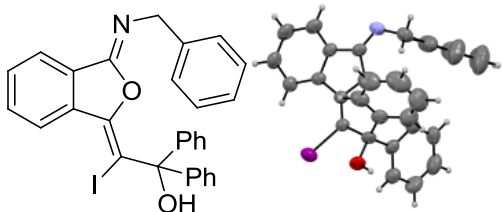
¹³C NMR (100 MHz, CDCl₃): δ = 162.1, 156.7, 140.5, 136.8, 136.3, 131.8, 130.5, 129.6, 128.6, 128.5, 128.2, 127.9, 127.6, 126.8, 125.8, 123.7, 120.7, 95.4, 90.3, 51.8 and 25.7 ppm.

IR (neat): 3433, 3055, 2986, 2929, 1778, 1701, 1654, 1450, 1423, 1266, 1020, 896, 737, 707 and 444 cm⁻¹

TLC: R_f = 0.5 (4:1; hexane:EtOAc)

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Crystallographic Data and Structure Refinements Summary for Compound 11d	
Molecular Structure (ORTEP Diagram)	
CCDC number	CCDC 1589065
Formula (M+ CH ₂ Cl ₂)	C ₃₀ H ₂₄ Cl ₂ I NO ₂
Formula weight	628.30
Colour	colourless
Crystal morphology	orthogonal
Temperature/K	296(2)
Radiation	Mo K α
Wavelength/Å	0.71073
Crystal system	tetragonal
Space group	I4 ₁ /a
a (Å)	18.2568(5)
b (Å)	18.2568(5)
c (Å)	33.7580(19)
α (°)	90
β (°)	90
γ (°)	90
Volume(Å ³)	11252.0(9)
Z	16
Density (g/cm ³)	1.484
μ (1/mm)	1.355
F (000)	5024
θ (min, max)	1.268 to 24.648
No. of unique reflns	4740
No. of parameters	302
R _{obs} , wR _{2_obs}	R ₁ = 0.0472, wR ₂ = 0.1185
$\Delta\rho_{min}$, $\Delta\rho_{max}$ (eÅ ⁻³)	0.787 and -0.722
GooF	1.125

