

Highly regioselectively *meta* arylation of oxaryl amide-protected β-arylethylamine via catellani reaction

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1. Reagents: Unless otherwise noted, all reagents were purchased from Acros, Alfa, Adamas and used without further purification. Column chromatography purifications were performed using 300–400 mesh silica gel.

2. Instruments: NMR spectra were recorded on Varian Inova-400 MHz, Inova-300 MHz, Bruker DRX-400 or Bruker DRX-500 instruments and calibrated using residual solvent peaks as internal reference. Multiplicities are recorded as: s = singlet, d = doublet, t = triplet, dd = doublet of doublets, br s = broad singlet, m = multiplet. HRMS analysis were carried out using TOF-MS instrument with EI source.

3. Optimization of reaction conditions

Table S1. Screening of solvent^a

Entry	Solvent	Yield (%) ^b
1	DCE	36
2	t-AmylOH	10
3	HFIP	7
4	toluene	65
5	m-xylene	78
6	mesitylene	86
7	PhCl	75
8	1,4-dioxane	78

^a1a (0.1 mmol), 2a (0.3 mmol), Pd(OAc)₂ (10 mol%), AgOAc (0.15 mmol), norbornene (0.1 mmol), 1-AdCO₂H (0.05 mmol), solvent (0.5 mL), 100 °C, 24 h. ^bYields were based on LC-MS analysis using biphenyl as an internal standard.

Table S2. Screening of oxidant^a

Entry	Oxidant	Yield (%) ^b
1	Ag ₂ CO ₃	0
2	Ag ₂ O	79
3	AgOPiv	68
4	AgF	24
5	AgCl	0
6	AgNO ₃	0

7	AgOTs	0
8	AgOTf	0
9	AgOAc (1 equiv)	62
10	AgOAc (1.5 equiv)	86
11	AgOAc (2 equiv)	80
12	BQ	0
13	Cu(OAc) ₂	0
14	K ₂ S ₂ O ₈	0

^a1a (0.1 mmol), 2a (0.3 mmol), Pd(OAc)₂ (10 mol%), oxidant (0.15 mmol),

norbornene (0.1 mmol), 1-AdCO₂H (0.05 mmol), mesitylene (0.5 mL), 100

°C, 24 h. ^bYields were based on LC-MS analysis using biphenyl as an internal standard.

Table S3. Screening of additive^a

Entry	Additive	Yield (%) ^b
1	none	75
2	1-AdCO ₂ H	86
3	PivOH	79
4	PhCO ₂ H	65
5	(n-BuO) ₂ PO ₂ H	73
6	HOAc	52
7	Ac-Gly-OH	75
8	K ₂ CO ₃	0
9	PivONa	78

^a1a (0.1 mmol), 2a (0.3 mmol), Pd(OAc)₂ (10 mol%), AgOAc (0.15 mmol),

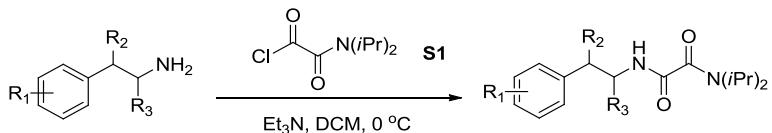
norbornene (0.1 mmol), additive (0.05 mmol), mesitylene (0.5 mL), 100 °C,

24 h. ^bYields were based on LC-MS analysis using biphenyl as an internal standard.

Table S4. Screening of directing group

1a	1 equiv	2a	3 equiv	10 mol% Pd(OAc) ₂ 1.5 equiv AgOAc 1 equiv norbornene 0.5 equiv 1-AdCO ₂ H mesitylene, 100 °C, 24 h	3a
DG:					
Yield	0	0	0	0	86%

4. Preparation of substrates

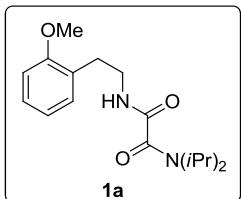


4.1. Preparation of N,N-Diisopropylloxamoyl chloride S1^[1]

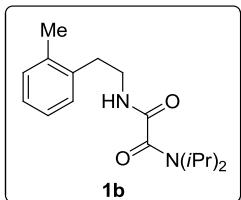
A solution of Diisopropylamine (7.01 mL, 50 mmol, 1.0 eq) in CH₂Cl₂ (50 mL) was added dropwise to a solution of oxalyl chloride (6.44 ml, 75 mmol, 1.5 eq) in CH₂Cl₂ (100 mL) at 0 °C, after stirring for 5 min, triethylamine (7.30 mL, 52.5 mmol, 1.05 eq) was added dropwise. The solution was warmed to room temperature and stirred for 6 hours. The excess of oxalyl chloride and the solvent were removed under reduced pressure and CH₂Cl₂ (30 mL) was added and evaporated. This operation was performed twice to give **S1** as a pale yellow solid. The crude product was used in the next step without any purification.

4.2. General procedures for the preparation of oxalamide substrates

A solution of amine (20 mmol, 1.0 eq) in CH₂Cl₂ (40 mL) was added dropwise to a solution of N,N-Diisopropylloxamoyl chloride **S1** (25 mmol, 1.25 eq) in CH₂Cl₂ (50 mL) at 0 °C, after stirring for 5 min, triethylamine (2.92 mL, 21 mmol, 1.05 eq) was added dropwise and then the mixture was stirred for 6 hours at room temperature before quenched by water (50 mL). The organic layer was separated and the aqueous layer was extracted with CH₂Cl₂ (20 mL × 2). The combined organic phase was washed with brine (30 mL), and then dried over anhydrous Na₂SO₄. Evaporation and column chromatography on silica gel afforded corresponding amide substrates as white solid or colourless oil with >80% yield.

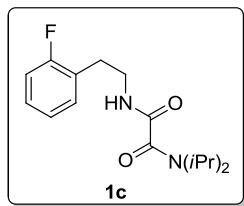


White solid.^[2] ¹H NMR (400 MHz, CDCl₃) δ 7.22–7.18 (m, 1H), 7.14–7.12 (m, 2H), 6.87 (dd, *J* = 17.7, 7.9 Hz, 2H), 4.64–4.57 (m, 1H), 3.82 (s, 3H), 3.53–3.43 (m, 3H), 2.86 (t, *J* = 6.9 Hz, 2H), 1.39 (d, *J* = 6.8 Hz, 6H), 1.18 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 162.40, 156.56, 129.60, 126.95, 126.13, 119.65, 109.35, 54.26, 48.60, 45.39, 38.59, 29.10, 19.88, 19.08; IR ν 3265, 2966, 2935, 2882, 1667, 1616, 1542, 1510, 1486, 1459, 1447, 1437, 1390, 1367, 1350, 1280, 1247, 1236, 1197, 1159, 1137, 1061, 1042, 1020, 874, 823, 777, 759, 742, 693, 681, 632, 613 cm⁻¹.

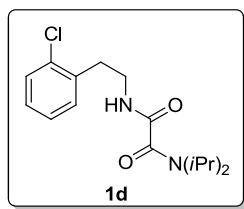


White solid.^[2] ¹H NMR (400 MHz, CDCl₃) δ 7.17–7.12 (m, 5H), 4.69–4.62 (m, 1H), 3.54–3.46 (m,

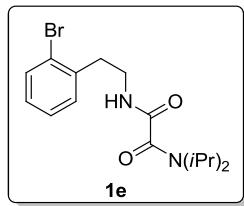
3H), 2.88–2.85 (m, 2H), 2.35 (s, 3H), 1.41 (d, J = 6.8 Hz, 6H), 1.21 (d, J = 6.7 Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 163.41, 163.28, 136.77, 136.46, 130.55, 129.36, 126.78, 126.24, 49.75, 46.58, 39.47, 32.98, 20.96, 20.15, 19.45.



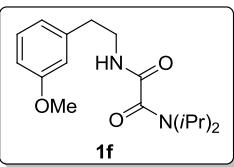
White solid.^[2] ^1H NMR (400 MHz, CDCl_3) δ 7.24–7.19 (m, 2H), 7.10–7.01 (m, 2H), 6.94 (br s, 1H), 4.68–4.61 (m, 1H), 3.58–3.53 (m, 2H), 3.51–3.46 (m, 1H), 2.91 (t, J = 7.1 Hz, 2H), 1.41 (d, J = 6.8 Hz, 6H), 1.20 (d, J = 6.7 Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 163.49, 163.27, 162.40 (d, J_{C-F} = 244.0 Hz), 131.18 (d, J_{C-F} = 5.0 Hz), 128.47 (d, J_{C-F} = 8.0 Hz), 125.63 (d, J_{C-F} = 16.0 Hz), 124.30 (d, J_{C-F} = 4.0 Hz), 115.48 (d, J_{C-F} = 22.0 Hz), 49.76, 46.55, 39.38, 29.03, 29.01, 20.93, 20.15.



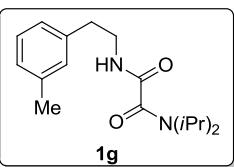
White solid. ^1H NMR (400 MHz, CDCl_3) δ 7.35 (dd, J = 7.5, 1.6 Hz, 1H), 7.28–7.14 (m, 4H), 4.63–4.57 (m, 1H), 3.60–3.55 (m, 2H), 3.52–3.45 (m, 1H), 3.00 (t, J = 7.2 Hz, 2H), 1.40 (d, J = 6.8 Hz, 6H), 1.20 (d, J = 6.7 Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.48, 162.33, 135.36, 133.23, 130.01, 128.68, 127.12, 126.04, 48.73, 45.50, 37.92, 32.25, 19.91, 19.12; HRMS Calcd for $\text{C}_{16}\text{H}_{23}\text{ClN}_2\text{NaO}_2$ [M+Na $^+$]: 333.1346; Found: 333.1352.



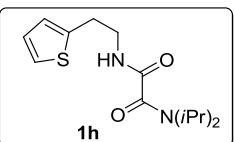
White solid.^[2] ^1H NMR (400 MHz, CDCl_3) δ 7.53 (d, J = 7.9 Hz, 1H), 7.32 (br s, 1H), 7.29–7.23 (m, 2H), 7.10–7.06 (m, 1H), 4.58–4.54 (m, 1H), 3.60–3.55 (m, 2H), 3.52–3.45 (m, 1H), 3.01 (t, J = 7.2 Hz, 2H), 1.40 (d, J = 6.8 Hz, 6H), 1.20 (d, J = 6.7 Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 163.49, 163.40, 138.04, 132.95, 130.97, 128.32, 127.65, 124.63, 49.73, 46.44, 38.96, 35.65, 20.87, 20.09 .



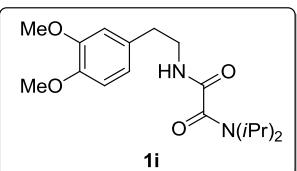
White solid.^[2] ¹H NMR (400 MHz, CDCl₃) δ 7.22–7.18 (m, 1H), 7.10 (br s, 1H), 6.80 (d, *J* = 7.7 Hz, 1H), 6.76–6.75 (m, 2H), 4.58–4.52 (m, 1H), 3.78 (s, 3H), 3.56–3.51 (m, 2H), 3.49–3.44 (m, 1H), 2.82 (t, *J* = 7.2 Hz, 2H), 1.39 (d, *J* = 6.8 Hz, 6H), 1.18 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 163.46, 159.87, 140.22, 129.68, 121.15, 114.36, 112.18, 55.25, 49.80, 46.50, 40.39, 35.57, 20.90, 20.14.



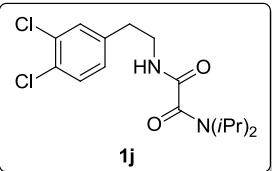
White solid. ¹H NMR (400 MHz, CDCl₃) δ 7.20–7.16 (t, *J* = 7.6 Hz, 2H), 7.03–7.00 (m, 3H), 4.58–4.51 (m, 1H), 3.56–3.51 (m, 2H), 3.49–3.44 (m, 1H), 2.81 (t, *J* = 7.3 Hz, 2H), 2.32 (s, 3H), 1.39 (d, *J* = 6.8 Hz, 6H), 1.19 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 162.52, 162.48, 137.55, 137.22, 128.61, 127.54, 126.32, 124.81, 48.79, 45.45, 39.52, 34.41, 20.43, 19.88, 19.13; HRMS Calcd for C₁₇H₂₇N₂O₂ [M+H⁺]: 291.2073; Found: 291.2076.



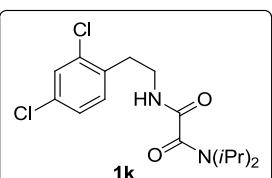
White solid.^[3] ¹H NMR (400 MHz, CDCl₃) δ 7.29 (br s, 1H), 7.13 (d, *J* = 4.9 Hz, 1H), 6.93–6.90 (m, 1H), 6.85 (s, 1H), 4.59–4.52 (m, 1H), 3.58–3.53 (m, 2H), 3.51–3.44 (m, 1H), 3.06 (t, *J* = 6.8 Hz, 2H), 1.39 (d, *J* = 6.8 Hz, 6H), 1.19 (d, *J* = 6.6 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 163.50, 163.42, 140.98, 127.09, 125.47, 123.95, 49.81, 46.47, 40.69, 29.63, 20.88, 20.13.



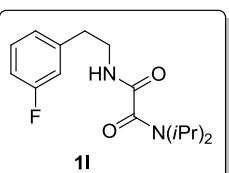
White solid.^[2] ¹H NMR (400 MHz, CDCl₃) δ 7.03 (br s, 1H), 6.80–6.78 (m, 1H), 6.74 (dd, *J* = 5.9, 1.8 Hz, 2H), 4.61–4.54 (m, 1H), 3.86 (s, 3H), 3.84 (s, 3H), 3.54–3.44 (m, 3H), 2.78 (t, *J* = 7.1 Hz, 2H), 1.38 (d, *J* = 6.8 Hz, 6H), 1.18 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 163.41, 163.29, 149.09, 147.80, 131.16, 120.77, 112.02, 111.45, 56.00, 55.94, 49.76, 46.54, 40.60, 35.16, 20.91, 20.14



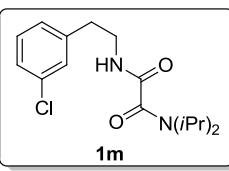
White solid. ^1H NMR (400 MHz, CDCl_3) δ 7.52 (br s, 1H), 7.33 (d, $J = 8.2$ Hz, 1H), 7.29 (d, $J = 1.9$ Hz, 1H), 7.05 (dd, $J = 8.2, 2.0$ Hz, 1H), 4.38–4.31 (m, 1H), 3.53–3.41 (m, 3H), 2.80 (t, $J = 7.1$ Hz, 2H), 1.35 (d, $J = 6.8$ Hz, 6H), 1.15 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.75, 162.68, 138.06, 131.42, 129.84, 129.55, 129.49, 127.34, 76.48, 49.00, 45.42, 38.90, 33.55, 19.81, 19.10; HRMS Calcd for $\text{C}_{16}\text{H}_{22}\text{Cl}_2\text{N}_2\text{NaO}_2$ [$\text{M}+\text{Na}^+$]: 367.0956; Found: 367.0966.



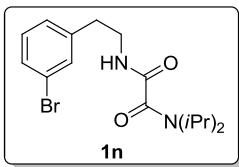
White solid.^[2] ^1H NMR (400 MHz, CDCl_3) δ 7.34 (s, 1H), 7.26 (br s, 1H), 7.19–7.15 (m, 2H), 4.54–4.50 (m, 1H), 3.55–3.50 (m, 2H), 3.48–3.43 (m, 1H), 2.94 (t, $J = 7.1$ Hz, 2H), 1.37 (d, $J = 6.8$ Hz, 6H), 1.18 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 163.52, 163.26, 134.99, 134.92, 133.12, 131.82, 129.45, 127.29, 49.81, 46.55, 38.71, 32.76, 20.90, 20.12.



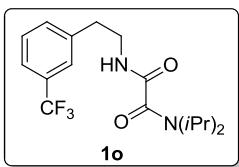
White solid. ^1H NMR (400 MHz, CDCl_3) δ 7.45 (br s, 1H), 7.25–7.19 (m, 1H), 6.98 (d, $J = 7.6$ Hz, 1H), 6.91–6.85 (m, 2H), 4.45–4.38 (m, 1H), 3.55–3.50 (m, 2H), 3.46–3.41 (m, 1H), 2.83 (t, $J = 7.2$ Hz, 2H), 1.36 (d, $J = 6.8$ Hz, 6H), 1.15 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.75, 162.62, 161.94 ($d, J_{C-F} = 244.0$ Hz), 140.32 ($d, J_{C-F} = 8.0$ Hz), 129.04 ($d, J_{C-F} = 8.0$ Hz), 123.49 ($d, J_{C-F} = 3.0$ Hz), 114.70 ($d, J_{C-F} = 21.0$ Hz), 112.40 ($d, J_{C-F} = 21.0$ Hz), 48.89, 45.38, 39.11, 34.13, 34.12, 19.80, 19.09; HRMS Calcd for $\text{C}_{16}\text{H}_{23}\text{FN}_2\text{NaO}_2$ [$\text{M}+\text{Na}^+$]: 317.1641; Found: 317.1650.



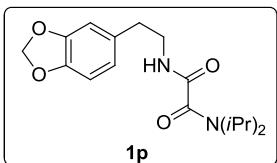
White solid.^[2] ^1H NMR (400 MHz, CDCl_3) δ 7.24–7.17 (m, 4H), 7.10 (d, $J = 7.0$ Hz, 1H), 4.53–4.46 (m, 1H), 3.56–3.50 (m, 2H), 3.49–3.44 (m, 1H), 2.83 (t, $J = 7.2$ Hz, 2H), 1.38 (d, $J = 6.8$ Hz, 6H), 1.18 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 163.55, 163.45, 140.75, 134.40, 129.94, 128.99, 127.07, 126.83, 49.90, 46.54, 40.21, 35.16, 20.91, 20.15.



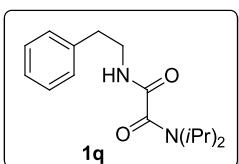
White solid.^[2] ¹H NMR (400 MHz, CDCl₃) δ 7.42 (br s, 1H), 7.35 (s, 1H), 7.32–7.31 (m, 1H), 7.14 (d, *J* = 4.6 Hz, 2H), 4.43–4.38 (m, 1H), 3.54–3.49 (m, 2H), 3.47–3.42 (m, 1H), 2.81 (t, *J* = 7.2 Hz, 2H), 1.36 (d, *J* = 6.8 Hz, 6H), 1.16 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 163.65, 163.61, 141.08, 131.85, 130.15, 129.66, 127.49, 122.58, 49.93, 46.40, 40.11, 35.03, 20.85, 20.11.



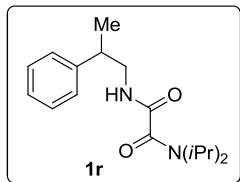
White solid.^[2] ¹H NMR (400 MHz, CDCl₃) δ 7.49–7.47 (m, 2H), 7.45–7.41 (m, 2H), 7.17 (br s, 1H), 4.63–4.58 (m, 1H), 3.59–3.54 (m, 2H), 3.52–3.45 (m, 1H), 2.92 (t, *J* = 7.2 Hz, 2H), 1.39 (d, *J* = 6.8 Hz, 6H), 1.19 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 163.47, 163.16, 139.64, 132.33 (d, *J*_{C-F} = 1.0 Hz), 131.03 (q, *J*_{C-F} = 32.0 Hz), 129.19, 125.62 (q, *J*_{C-F} = 4.0 Hz), 124.23 (q, *J*_{C-F} = 270.0 Hz), 123.60 (q, *J*_{C-F} = 4.0 Hz), 49.82, 46.65, 40.29, 35.39, 20.91, 20.15.



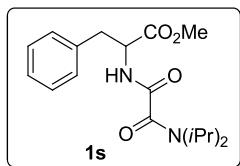
White solid.^[2] ¹H NMR (400 MHz, CDCl₃) δ 7.08 (br s, 1H), 6.73 (d, *J* = 7.9 Hz, 1H), 6.69 (d, *J* = 1.4 Hz, 1H), 6.65 (dd, *J* = 7.9, 1.6 Hz, 1H), 5.91 (s, 2H), 4.62–4.55 (m, 1H), 3.51–3.44(m, 3H), 2.76 (t, *J* = 7.1 Hz, 2H), 1.39 (d, *J* = 6.8 Hz, 6H), 1.19 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 163.41, 163.33, 147.87, 146.30, 132.39, 121.77, 109.18, 108.46, 100.99, 49.79, 46.54, 40.73, 35.24, 20.92, 20.15; HRMS Calcd for C₁₇H₂₃N₂O₄ [M-H⁺]: 319.1658; Found: 319.1673.



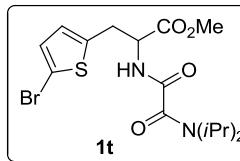
White solid.^[4] ¹H NMR (400 MHz, CDCl₃) δ 7.32–7.28 (m, 2H), 7.23–7.20 (m, 3H), 7.11 (br s, 1H), 4.60–4.53 (m, 1H), 3.58–3.53 (m, 2H), 3.52–3.45 (m, 1H), 2.86 (t, *J* = 7.2 Hz, 2H), 1.40 (d, *J* = 6.8 Hz, 6H), 1.19 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 163.45, 163.41, 138.68, 128.85, 128.70, 126.62, 49.78, 46.52, 40.53, 35.55, 20.92, 20.16.



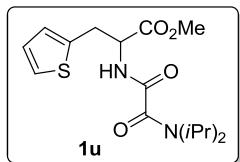
White solid.^[4] ¹H NMR (400 MHz, CDCl₃) δ 7.30 (t, *J* = 7.6 Hz, 2H), 7.23–7.19 (m, 3H), 6.94 (br s, 1H), 4.42–4.37 (m, 1H), 3.54–3.39 (m, 3H), 3.03–2.94 (m, 1H), 1.38–1.35 (m, 6H), 1.28 (d, *J* = 7.0 Hz, 3H), 1.16–1.12 (m, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 163.54, 143.93, 128.72, 127.28, 126.77, 49.83, 46.40, 45.86, 39.78, 20.88, 20.16, 20.14, 19.45.



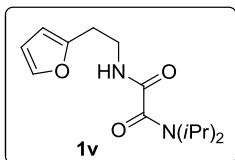
White solid.^[2] ¹H NMR (400 MHz, CDCl₃) δ 7.30–7.24 (m, 4H), 7.17 (d, *J* = 7.1 Hz, 2H), 4.89–4.82 (m, 1H), 4.39–4.33 (m, 1H), 3.72 (s, 3H), 3.51–3.44 (m, 1H), 3.22–3.17 (m, 1H), 3.12–3.07 (m, 1H), 1.42–1.40 (m, 6H), 1.21–1.13 (m, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 184.72, 171.32, 162.95, 135.74, 129.34, 128.69, 127.21, 53.24, 52.52, 49.81, 46.50, 38.03, 20.90, 20.81, 20.10.



Pale yellow solid. ¹H NMR (400 MHz, CDCl₃) δ 7.28 (br s, 1H), 6.88 (d, *J* = 3.7 Hz, 1H), 6.63 (d, *J* = 3.7 Hz, 1H), 4.83–4.78 (m, 1H), 4.53–4.47 (m, 1H), 3.77 (s, 3H), 3.54–3.47 (m, 1H), 3.39–3.26 (m, 2H), 1.42 (dd, *J* = 6.7, 5.3 Hz, 6H), 1.22 (dd, *J* = 6.6, 4.5 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 169.50, 162.09, 161.48, 137.88, 129.10, 126.73, 110.10, 51.97, 51.91, 48.92, 45.70, 31.41, 20.01, 19.94, 19.18; HRMS Calcd for C₁₆H₂₃BrN₂NaO₄S [M+Na⁺]: 441.0460; Found: 441.0468.

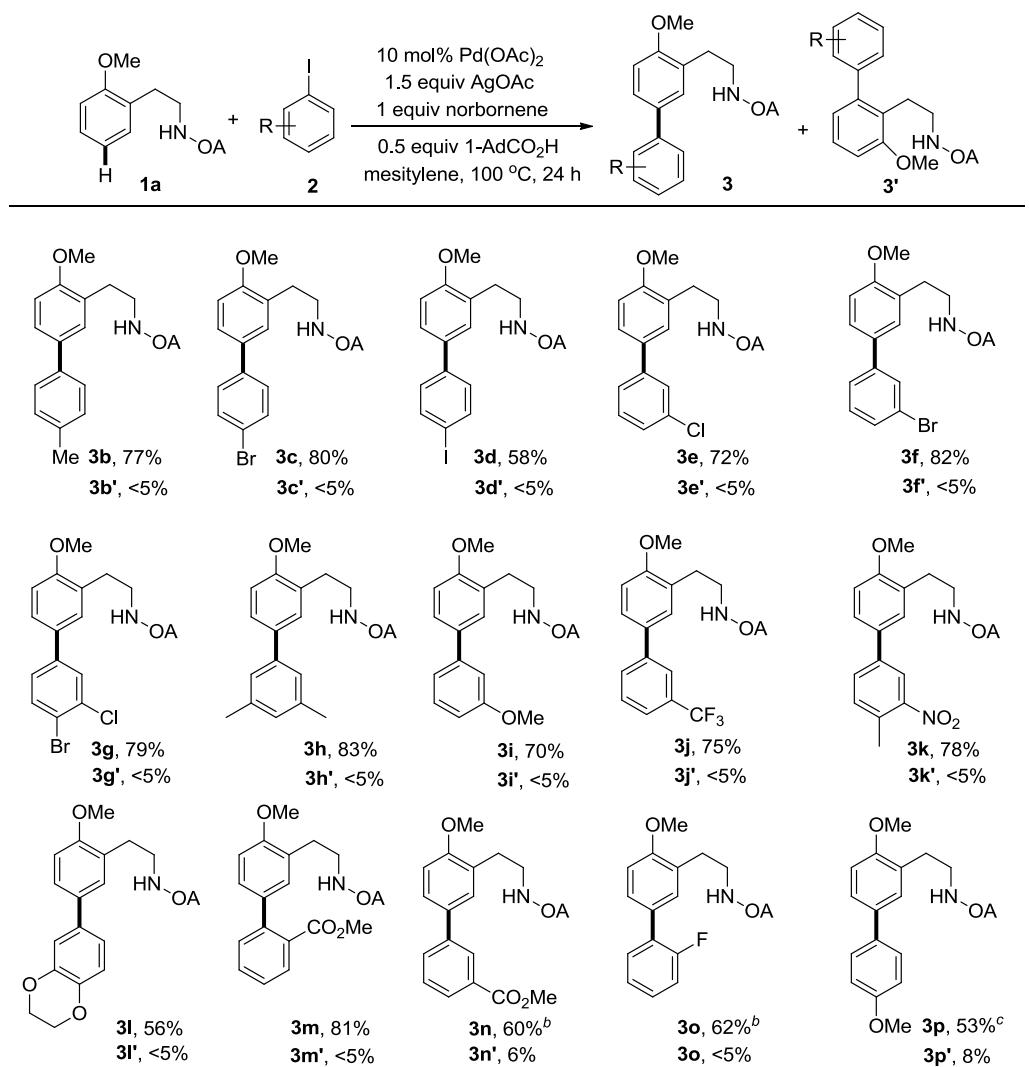


Pale yellow solid. ¹H NMR (400 MHz, CDCl₃) δ 7.28 (br s, 1H), 7.15 (dd, *J* = 5.1, 1.1 Hz, 1H), 6.92 (dd, *J* = 5.1, 3.5 Hz, 1H), 6.84 (d, *J* = 3.1 Hz, 1H), 4.85–4.81 (m, 1H), 4.52–4.46 (m, 1H), 3.74 (s, 3H), 3.52–3.45 (m, 1H), 3.43–3.35 (m, 2H), 1.41 (dd, *J* = 6.7, 4.3 Hz, 6H), 1.20 (dd, *J* = 6.6, 3.3 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 169.69, 162.03, 161.58, 136.04, 126.25, 126.07, 124.01, 52.21, 51.70, 48.82, 45.58, 30.97, 19.95, 19.88, 19.13; HRMS Calcd for C₁₆H₂₅N₂O₄S [M+H⁺]: 341.1535; Found: 341.1536.

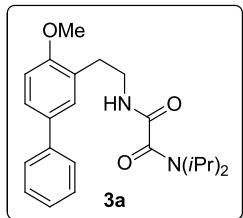


White solid. ^1H NMR (400 MHz, CDCl_3) δ 7.31 (d, $J = 1.1$ Hz, 1H), 7.14 (br s, 1H), 6.27 (dd, $J = 3.0, 1.9$ Hz, 1H), 6.08 (d, $J = 3.1$ Hz, 1H), 4.65–4.58 (m, 1H), 3.58–3.53 (m, 2H), 3.52–3.45 (m, 1H), 2.87 (t, $J = 6.7$ Hz, 2H), 1.39 (d, $J = 6.8$ Hz, 6H), 1.20 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 163.42, 163.23, 152.72, 141.69, 110.39, 106.51, 49.77, 46.57, 37.95, 27.98, 20.94, 20.16; HRMS Calcd for $\text{C}_{14}\text{H}_{22}\text{N}_2\text{NaO}_3$ [M+Na $^+$]: 289.1528; Found: 289.1530.

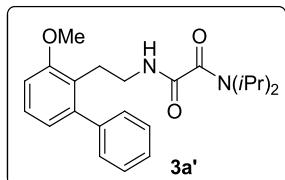
5. Meta arylation with different aryl iodides



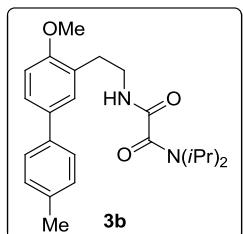
A mixture of **1a** (0.2 mmol, 61.3 mg), **2** (0.6 mmol, 3.0 eq), Pd(OAc)₂ (4.5 mg, 10 mol%), AgOAc (50 mg, 0.3 mmol, 1.5 eq), norbornene (18.8 mg, 0.2 mmol, 1.0 eq), 1-AdCO₂H (18 mg, 0.1 mmol, 0.05 eq) and 1 mL mesitylene in a 15 mL glass vial was heated at 100 °C with vigorous stirring for 24 hours. The reaction mixture was cooled to room temperature, and diluted with ethyl acetate and filtered through celite. The filtrate was concentrated in vacuo and purified by column chromatography on silica gel (Ethyl acetate/Petroleum ether = 1:15 to 1:3) to give product.



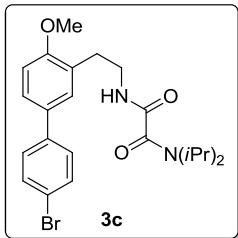
Pale yellow oil; Yield (81%, 61.9 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.55 (d, *J* = 7.5 Hz, 2H), 7.46–7.40 (m, 4H), 7.29 (t, *J* = 7.3 Hz, 1H), 7.12 (br s, 1H), 6.93 (d, *J* = 8.4 Hz, 1H), 4.71–4.65 (m, 1H), 3.88 (s, 3H), 3.59–3.54 (m, 2H), 3.51–3.44 (m, 1H), 2.94 (t, *J* = 6.8 Hz, 2H), 1.40 (d, *J* = 6.7 Hz, 6H), 1.16 (d, *J* = 6.6 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 162.40, 162.19, 156.24, 139.82, 132.87, 128.54, 128.82, 126.49, 125.91, 125.81, 125.60, 109.77, 54.59, 48.64, 45.58, 38.72, 29.30, 19.94, 19.15; HRMS Calcd for C₂₃H₃₀N₂NaO₃ [M+Na⁺]: 405.2154; Found: 405.2152; IR ν 3270, 3089, 2994, 2977, 2937, 2884, 2835, 1674, 1612, 1561, 1495, 1450, 1434, 1381, 1362, 1317, 1306, 1290, 1263, 1244, 1205, 1170, 1158, 1140, 1120, 1093, 1055, 1038, 1022, 988, 956, 925, 876, 805, 774, 752, 740, 728, 702, 675, 612 cm⁻¹.



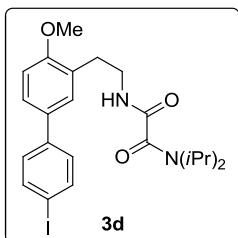
Pale yellow oil; Yield (83%, 63.4 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.41–7.32 (m, 3H), 7.28–7.21 (m, 3H), 7.04 (br s, 1H), 6.89 (d, *J* = 8.1 Hz, 1H), 6.84 (d, *J* = 7.6 Hz, 1H), 4.71–4.64 (m, 1H), 3.89 (s, 3H), 3.51–3.44 (m, 1H), 3.38–3.33 (m, 2H), 2.83 (t, *J* = 7.0 Hz, 2H), 1.40 (d, *J* = 6.8 Hz, 6H), 1.17 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 162.05, 161.99, 156.87, 143.20, 140.49, 128.22, 127.28, 126.20, 126.16, 124.11, 121.77, 108.42, 54.64, 48.44, 45.55, 38.75, 25.41, 19.97, 19.12; HRMS Calcd for C₂₃H₃₁N₂O₃ [M+H⁺]: 383.2335; Found: 383.2347.



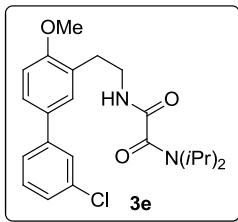
Pale yellow solid; Yield (77%, 61.0 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.45–7.41 (m, 3H), 7.37 (d, *J* = 2.2 Hz, 1H), 7.21 (d, *J* = 7.9 Hz, 2H), 7.10 (br s, 1H), 6.92 (d, *J* = 8.5 Hz, 1H), 4.72–4.65 (m, 1H), 3.88 (s, 3H), 3.58–3.54 (m, 2H), 3.51–3.44 (m, 1H), 2.93 (t, *J* = 6.9 Hz, 2H), 2.38 (s, 3H), 1.41 (d, *J* = 6.8 Hz, 6H), 1.17 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 162.41, 162.20, 156.04, 136.95, 135.52, 132.87, 128.55, 128.38, 126.44, 125.76, 125.41, 109.77, 54.60, 48.66, 45.60, 38.79, 29.32, 20.19, 19.96, 19.17; HRMS Calcd for C₂₄H₃₃N₂O₃ [M+H⁺]: 397.2491; Found: 397.2500.



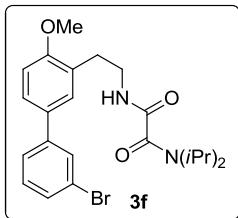
Yellow solid; Yield (80%, 73.6 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.51 (d, $J = 8.5$ Hz, 2H), 7.43–7.39 (m, 3H), 7.35 (d, $J = 2.1$ Hz, 1H), 7.10 (br s, 1H), 6.92 (d, $J = 8.5$ Hz, 1H), 4.72–4.65 (m, 1H), 3.88 (s, 3H), 3.58–3.53 (m, 2H), 3.51–3.45 (m, 1H), 2.93 (t, $J = 6.9$ Hz, 2H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.17 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.38, 162.13, 156.51, 138.76, 131.57, 130.88, 128.37, 127.51, 126.69, 125.47, 119.94, 109.85, 54.63, 48.65, 45.62, 38.61, 29.29, 19.96, 19.16; HRMS Calcd for $\text{C}_{23}\text{H}_{30}\text{BrN}_2\text{O}_3$ [$\text{M}+\text{H}^+$]: 461.1440; Found: 461.1444.



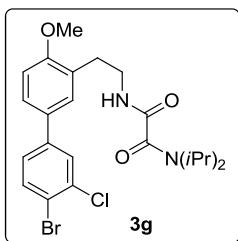
Yellow solid; Yield (58%, 58.9 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.51 (d, $J = 8.5$ Hz, 2H), ^1H NMR (400 MHz, CDCl_3) δ 7.71 (d, $J = 8.5$ Hz, 2H), 7.40 (dd, $J = 8.4, 2.4$ Hz, 1H), 7.35 (d, $J = 2.3$ Hz, 1H), 7.30 (d, $J = 8.5$ Hz, 2H), 7.10 (br s, 1H), 6.92 (d, $J = 8.5$ Hz, 1H), 4.73–4.66 (m, 1H), 3.88 (s, 3H), 3.57–3.52 (m, 2H), 3.51–3.45 (m, 1H), 2.92 (t, $J = 6.9$ Hz, 2H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.17 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 206.11, 162.37, 162.11, 156.56, 139.34, 136.86, 131.59, 128.30, 127.79, 126.70, 125.43, 109.85, 91.31, 54.62, 48.63, 45.61, 38.60, 30.06, 29.29, 19.96, 19.16; HRMS Calcd for $\text{C}_{23}\text{H}_{29}\text{IN}_2\text{NaO}_3$ [$\text{M}+\text{Na}^+$]: 531.1121; Found: 531.1112.



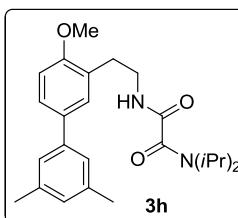
Pale yellow solid; Yield (72%, 59.9 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.54 (t, $J = 1.6$ Hz, 1H), 7.44–7.40 (m, 2H), 7.36 (d, $J = 2.2$ Hz, 1H), 7.32 (t, $J = 7.8$ Hz, 1H), 7.27–7.25 (m, 1H), 7.10 (br s, 1H), 7.10 (s, 1H), 6.93 (d, $J = 8.5$ Hz, 1H), 4.72–4.66 (m, 1H), 3.88 (s, 3H), 3.58–3.53 (m, 2H), 3.51–3.44 (m, 1H), 2.93 (t, $J = 6.9$ Hz, 2H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.17 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.39, 162.14, 156.67, 141.68, 133.71, 131.37, 129.04, 128.53, 126.70, 125.98, 125.78, 125.65, 124.04, 109.83, 54.63, 48.65, 45.62, 38.60, 29.32, 19.96, 19.16; HRMS Calcd for $\text{C}_{23}\text{H}_{30}\text{ClN}_2\text{O}_3$ [$\text{M}+\text{H}^+$]: 417.1945; Found: 417.1938.



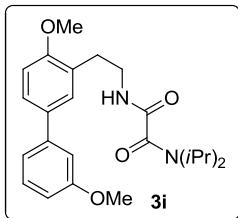
Yellow oil; Yield (82%, 75.4 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.70 (t, $J = 1.6$ Hz, 1H), 7.47 (d, $J = 7.6$ Hz, 1H), 7.41 (dd, $J = 8.5, 2.1$ Hz, 2H), 7.35 (d, $J = 2.2$ Hz, 1H), 7.26 (d, $J = 15.7$ Hz, 1H), 7.09 (br s, 1H), 6.92 (d, $J = 8.5$ Hz, 1H), 4.72–4.66 (m, 1H), 3.58–3.53 (m, 2H), 3.51–3.45 (m, 1H), 2.93 (t, $J = 6.9$ Hz, 2H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.17 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.41, 162.15, 156.68, 141.97, 131.27, 129.32, 128.88, 128.71, 128.53, 126.71, 125.67, 124.51, 121.99, 109.83, 54.63, 48.66, 45.62, 38.60, 29.32, 19.96, 19.17; HRMS Calcd for $\text{C}_{23}\text{H}_{30}\text{BrN}_2\text{O}_3$ [$\text{M}+\text{H}^+$]: 461.1440; Found: 461.1448.



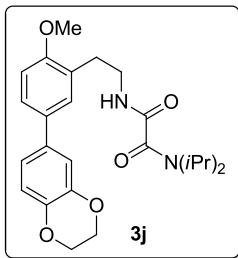
Pale yellow oil; Yield (81%, 80.0 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.65 (d, $J = 2.0$ Hz, 1H), 7.61 (d, $J = 8.3$ Hz, 1H), 7.39 (dd, $J = 8.4, 2.2$ Hz, 1H), 7.34 (d, $J = 2.1$ Hz, 1H), 7.30 (dd, $J = 8.3, 2.0$ Hz, 1H), 7.11 (br s, 1H), 6.92 (d, $J = 8.5$ Hz, 1H), 4.71–7.65 (m, 1H), 3.88 (s, 3H), 3.57–3.45 (m, 3H), 2.92 (t, $J = 6.8$ Hz, 2H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.17 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.41, 162.15, 156.86, 140.58, 133.79, 132.93, 130.24, 128.34, 127.53, 126.84, 125.49, 125.32, 119.53, 109.88, 54.63, 48.65, 45.60, 38.47, 29.28, 19.95, 19.15; HRMS Calcd for $\text{C}_{23}\text{H}_{29}\text{BrClN}_2\text{O}_3$ [$\text{M}+\text{H}^+$]: 495.1050; Found: 495.1040.



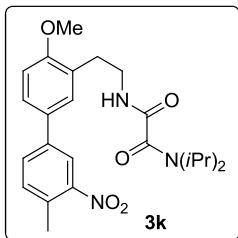
Pale yellow oil; Yield (83%, 68.1 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.43 (dd, $J = 8.4, 2.3$ Hz, 1H), 7.38 (d, $J = 2.2$ Hz, 1H), 7.17 (s, 2H), 7.09 (br s, 1H), 6.95 (s, 1H), 6.91 (d, $J = 8.4$ Hz, 1H), 4.71–4.64 (m, 1H), 3.88 (s, 3H), 3.58–3.53 (m, 2H), 3.51–3.44 (m, 1H), 2.93 (t, $J = 6.9$ Hz, 2H), 2.37 (s, 6H), 1.41 (d, $J = 6.8$ Hz, 6H), 1.17 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.44, 162.22, 156.10, 139.80, 137.32, 133.13, 128.60, 127.49, 126.35, 125.61, 123.87, 109.68, 54.59, 48.66, 45.59, 38.81, 29.33, 20.53, 19.97, 19.17; HRMS Calcd for $\text{C}_{25}\text{H}_{35}\text{N}_2\text{O}_3$ [$\text{M}+\text{H}^+$]: 411.2648; Found: 411.2655.



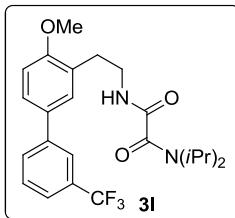
Yellow oil; Yield (70%, 57.7 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.44 (dd, $J = 8.4, 2.4$ Hz, 1H), 7.39 (d, $J = 2.3$ Hz, 1H), 7.31 (t, $J = 7.9$ Hz, 1H), 7.13 (d, $J = 7.7$ Hz, 1H), 7.09–7.06 (m, 2H), 6.93 (d, $J = 8.5$ Hz, 1H), 6.86–6.84 (m, 1H), 4.70–4.63 (m, 1H), 3.87 (d, $J = 6.2$ Hz, 6H), 3.59–3.54 (m, 2H), 3.51–3.44 (m, 1H), 2.93 (t, $J = 6.9$ Hz, 2H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.16 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.43, 162.21, 159.08, 156.36, 141.37, 132.76, 128.81, 128.58, 126.48, 125.69, 118.48, 111.56, 111.39, 109.75, 54.61, 54.47, 48.68, 45.59, 38.72, 29.34, 19.96, 19.16; HRMS Calcd for $\text{C}_{24}\text{H}_{32}\text{N}_2\text{NaO}_4$ [$\text{M}+\text{Na}^+$]: 435.2260; Found: 435.2262.



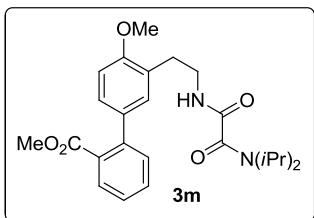
Yellow oil; Yield (56%, 49.3 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.37 (dd, $J = 8.4, 2.0$ Hz, 1H), 7.32 (d, $J = 1.8$ Hz, 1H), 7.10 (br s, 1H), 7.06–7.01 (m, 2H), 6.89 (d, $J = 7.6$ Hz, 2H), 4.72–4.65 (m, 1H), 4.28 (s, 4H), 3.86 (s, 3H), 3.57–3.52 (m, 2H), 3.51–3.44 (m, 1H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.17 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.41, 162.21, 155.92, 142.77, 141.82, 133.51, 132.33, 128.19, 126.42, 125.18, 118.94, 116.58, 114.59, 109.75, 63.58, 63.57, 54.58, 48.65, 45.58, 38.79, 29.29, 19.96, 19.16; HRMS Calcd for $\text{C}_{25}\text{H}_{32}\text{N}_2\text{NaO}_5$ [$\text{M}+\text{Na}^+$]: 463.2209; Found: 463.2208.



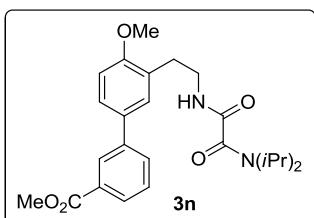
Yellow oil; Yield (79%, 69.7 mg); ^1H NMR (400 MHz, CDCl_3) δ 8.15 (d, $J = 1.8$ Hz, 1H), 7.70 (dd, $J = 7.9, 1.9$ Hz, 1H), 7.46 (dd, $J = 8.4, 2.4$ Hz, 1H), 7.40 (d, $J = 2.3$ Hz, 1H), 7.36 (d, $J = 8.0$ Hz, 1H), 7.10 (br s, 1H), 6.95 (d, $J = 8.5$ Hz, 1H), 4.74–4.67 (m, 1H), 3.89 (s, 3H), 3.58–3.53 (m, 2H), 3.52–3.45 (m, 1H), 2.94 (t, $J = 6.9$ Hz, 2H), 2.61 (s, 3H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.18 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 206.12, 162.39, 162.10, 156.93, 148.72, 138.99, 132.24, 130.61, 130.13, 130.08, 128.35, 126.96, 125.57, 121.55, 109.97, 54.66, 48.63, 45.63, 38.57, 30.06, 29.35, 19.97, 19.22, 19.15; HRMS Calcd for $\text{C}_{24}\text{H}_{32}\text{N}_3\text{O}_5$ [$\text{M}+\text{H}^+$]: 442.2342; Found: 442.2339.



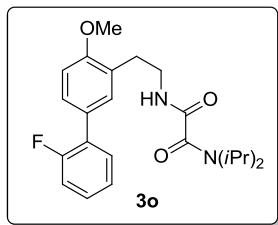
Pale yellow oil; Yield (75%, 67.5 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.79 (s, 1H), 7.73 (d, *J* = 7.2 Hz, 1H), 7.56–7.49 (m, 2H), 7.45 (dd, *J* = 8.4, 2.4 Hz, 1H), 7.39 (d, *J* = 2.3 Hz, 1H), 7.08 (br s, 1H), 6.95 (d, *J* = 8.5 Hz, 1H), 4.74–4.67 (m, 1H), 3.89 (s, 3H), 3.59–3.54 (m, 2H), 3.51–3.45 (m, 1H), 2.95 (t, *J* = 6.9 Hz, 2H), 1.40 (d, *J* = 6.8 Hz, 6H), 1.17 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 162.41, 162.13, 156.80, 140.64, 131.35, 130.22 (q, *J*_{C-F} = 32.0 Hz), 129.22 (d, *J*_{C-F} = 1.0 Hz), 128.63, 128.28, 127.44, 126.85, 126.09 (q, *J*_{C-F} = 271.0 Hz), 125.80, 124.73, 122.62 (q, *J*_{C-F} = 4.0 Hz), 122.45 (q, *J*_{C-F} = 4.0 Hz), 109.93, 54.66, 48.65, 45.63, 38.59, 29.37, 19.95, 19.13; HRMS Calcd for C₂₄H₃₀F₃N₂O₃ [M+H⁺]: 451.2209; Found: 451.2207.



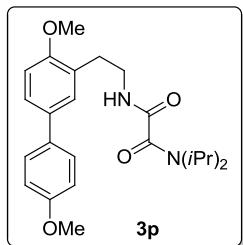
Pale yellow oil; Yield (81%, 71.3 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.77 (d, *J* = 7.4 Hz, 1H), 7.51–7.47 (m, 1H), 7.36 (t, *J* = 7.3 Hz, 2H), 7.17–7.11 (m, 3H), 6.88 (d, *J* = 8.4 Hz, 1H), 4.71–4.64 (m, 1H), 3.87 (s, 3H), 3.67 (s, 3H), 3.56–3.44 (m, 3H), 2.90 (t, *J* = 6.9 Hz, 2H), 1.40 (d, *J* = 6.8 Hz, 6H), 1.17 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 168.40, 162.42, 162.23, 156.04, 141.01, 132.66, 130.32, 129.93, 129.84, 129.82, 128.80, 126.99, 125.91, 125.87, 109.06, 54.48, 51.16, 48.65, 45.53, 38.81, 29.16, 19.93, 19.14; HRMS Calcd for C₂₅H₃₂N₂NaO₅ [M+Na⁺]: 463.2209; Found: 463.2217.



Yellow oil; Yield (60%, 52.8 mg); ¹H NMR (400 MHz, CDCl₃) δ 8.21 (s, 1H), 7.96 (d, *J* = 7.7 Hz, 1H), 7.75 (d, *J* = 7.8 Hz, 1H), 7.48 (dd, *J* = 12.8, 4.9 Hz, 2H), 7.42 (d, *J* = 2.2 Hz, 1H), 7.12 (br s, 1H), 6.94 (d, *J* = 8.5 Hz, 1H), 4.71–4.65 (m, 1H), 3.94 (s, 3H), 3.89 (s, 3H), 3.59–3.54 (m, 2H), 3.51–3.44 (m, 1H), 2.95 (t, *J* = 6.9 Hz, 2H), 1.39 (d, *J* = 6.8 Hz, 6H), 1.16 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 166.29, 162.41, 162.18, 156.61, 140.05, 131.68, 130.33, 129.73, 128.51, 127.93, 126.91, 126.85, 126.71, 125.71, 109.86, 54.62, 51.30, 48.65, 45.58, 38.72, 29.35, 19.94, 19.14; HRMS Calcd for C₂₅H₃₂N₂NaO₅ [M+Na⁺]: 463.2209; Found: 463.2222.

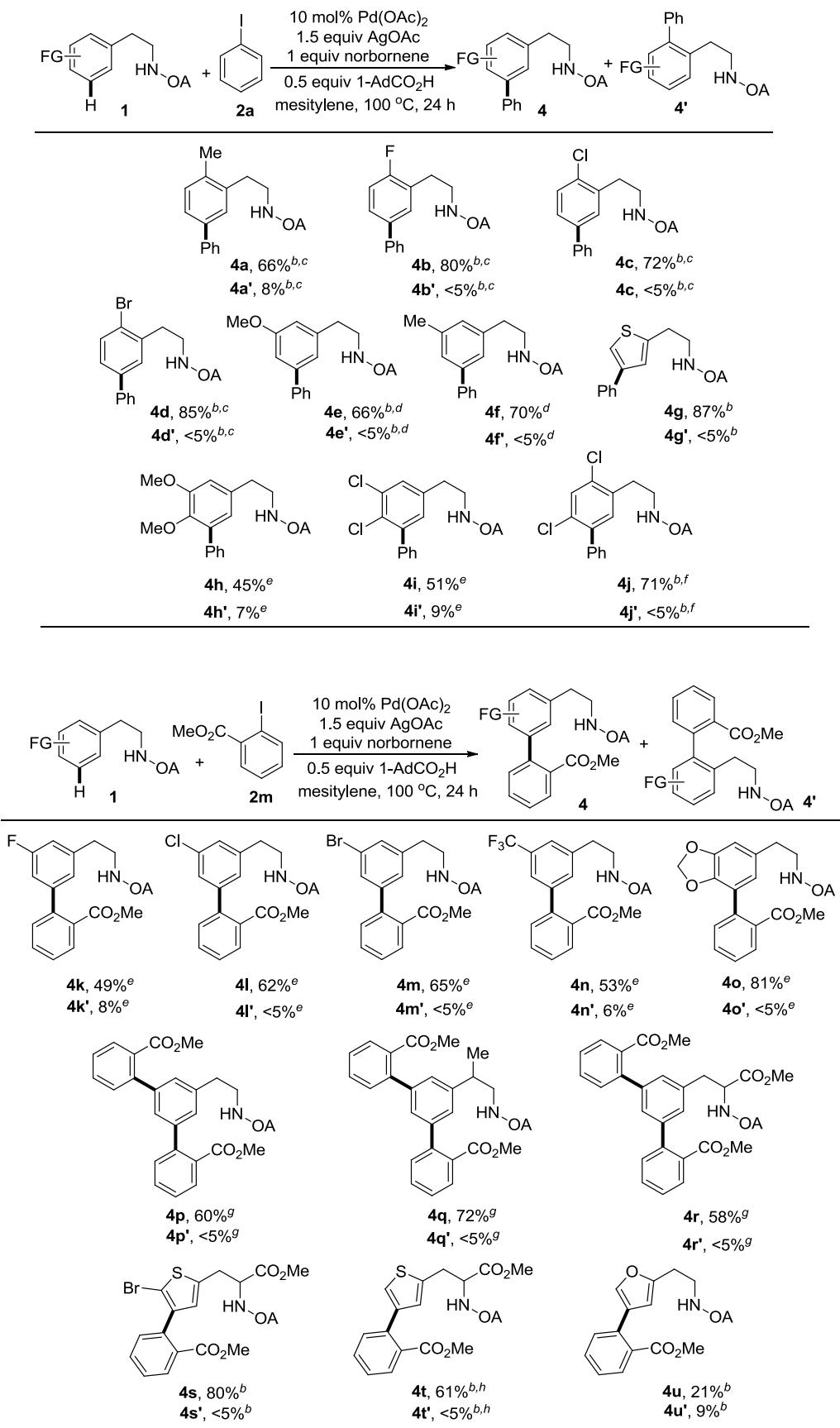


Pale yellow oil; Yield (62%, 49.6 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.44–7.40 (m, 2H), 7.34 (s, 1H), 7.29–7.24 (m, 1H), 7.19–7.08 (m, 3H), 6.94 (d, *J* = 8.5 Hz, 1H), 4.71–4.64 (m, 1H), 3.88 (s, 3H), 3.58–3.53 (m, 2H), 3.51–3.44 (m, 1H), 2.93 (t, *J* = 6.9 Hz, 2H), 1.40 (d, *J* = 6.8 Hz, 6H), 1.17 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 162.41, 162.20, 160.07, 157.62, 156.34, 130.38 (d, *J*_{C-F} = 3.0 Hz), 129.72 (d, *J*_{C-F} = 3.0 Hz), 127.75 (d, *J*_{C-F} = 3.0 Hz), 127.62, 127.60, 127.52, 127.27 (d, *J*_{C-F} = 1.0 Hz), 126.26, 123.44 (d, *J*_{C-F} = 3.0 Hz), 115.14 (d, *J*_{C-F} = 23.0 Hz), 109.46, 54.58, 48.67, 45.59, 38.71, 29.24, 19.94, 19.15; HRMS Calcd for C₂₃H₂₉FN₂NaO₃ [M+Na⁺]: 423.2060.; Found: 423.2068.

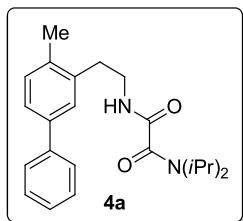


Yellow oil; Yield (53%, 43.7 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.49–7.46 (m, 2H), 7.40–7.38 (dd, *J* = 8.4, 2.4 Hz, 1H), 7.34 (d, *J* = 2.3 Hz, 1H), 7.10 (br s, 1H), 6.95–6.90 (m, 3H), 4.71–4.64 (m, 1H), 3.87 (s, 3H), 3.84 (s, 3H), 3.58–3.53 (m, 2H), 3.51–3.44 (m, 1H), 2.93 (t, *J* = 6.9 Hz, 2H), 1.40 (d, *J* = 6.8 Hz, 6H), 1.16 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 162.41, 162.21, 157.85, 155.80, 132.58, 132.48, 128.19, 126.92, 126.43, 125.16, 113.27, 109.78, 54.59, 54.48, 48.65, 45.58, 38.77, 29.30, 19.96, 19.16; HRMS Calcd for C₂₄H₃₂N₂NaO₄ [M+Na⁺]: 435.2260; Found: 435.2268.

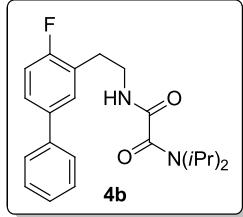
6. Meta arylation with different β -arylethyamides



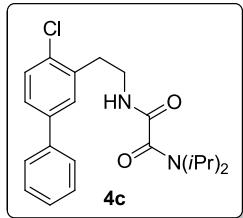
A mixture of oxalamide **1** (0.2 mmol, 1.0 eq), iodobenzene **2a** or methyl 2-iodobenzoate **2m** (0.6 mmol, 3.0 eq), Pd(OAc)₂ (4.5 mg, 10 mol%), AgOAc (50 mg, 0.3 mmol, 1.5 eq), norbornene (18.8 mg, 0.2 mmol, 1.0 eq), 1-AdCO₂H (18 mg, 0.1 mmol, 0.05 eq) and 1 mL mesitylene in a 15 mL glass vial was heated at 100 °C with vigorous stirring for 24 hours. The reaction mixture was cooled to room temperature, and diluted with ethyl acetate and filtered through celite. The filtrate was concentrated in vacuo and purified by column chromatography on silica gel (Ethyl acetate/Petroleum ether = 1:15 to 1:3) to give product.



Pale yellow solid; Yield (66%, 48.3 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.60–7.58 (m, 2H), 7.43–7.37 (m, 4H), 7.32 (t, *J* = 7.3 Hz, 1H), 7.24 (d, *J* = 8.3 Hz, 1H), 7.08 (br s, 1H), 4.76–4.69 (m, 1H), 3.59–3.54 (m, 2H), 3.53–3.46 (m, 1H), 2.93 (t, *J* = 7.4 Hz, 2H), 2.39 (s, 3H), 1.41 (d, *J* = 6.8 Hz, 6H), 1.18 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 162.34, 162.02, 140.06, 138.31, 136.15, 134.60, 130.07, 127.83, 127.20, 126.18, 126.15, 124.52, 48.71, 45.70, 38.53, 32.16, 19.96, 19.16, 18.17; HRMS Calcd for C₂₃H₃₀N₂NaO₂ [M+Na⁺]: 389.2205; Found: 389.2207.

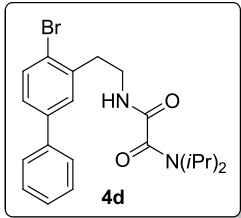


Pale yellow oil; Yield (80%, 59.2 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.54 (d, *J* = 7.5 Hz, 2H), 7.42 (dd, *J* = 9.1, 6.8 Hz, 4H), 7.33 (t, *J* = 7.3 Hz, 1H), 7.12–7.07 (m, 2H), 4.68–4.61 (m, 1H), 3.63–3.58 (m, 2H), 3.51–3.45 (m, 1H), 2.96 (t, *J* = 7.0 Hz, 2H), 1.40 (d, *J* = 6.8 Hz, 6H), 1.16 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 162.43, 162.02, 161.27, 158.82, 139.24, 136.66 (d, *J*_{C-F} = 3.0 Hz), 128.98 (d, *J*_{C-F} = 5.0 Hz), 127.92, 126.41, 126.19, 126.18 (d, *J*_{C-F} = 8.0 Hz), 124.82 (d, *J*_{C-F} = 16.0 Hz), 114.83 (d, *J*_{C-F} = 22.0 Hz), 48.74, 45.66, 38.41, 28.46, 28.18, 19.93, 19.16; HRMS Calcd for C₂₂H₂₇FN₂NaO₂ [M+Na⁺]: 393.1954; Found: 393.1960.

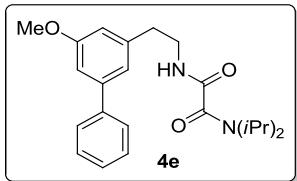


Pale yellow solid; Yield (72%, 55.6 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.58–7.56 (m, 2H), 7.47 (d, *J* = 1.7 Hz, 1H), 7.44–7.38 (m, 4H), 7.36–7.33 (m, 1H), 7.06 (br s, 1H), 4.74–4.67 (m, 1H), 3.65–

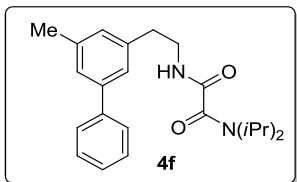
3.59 (m, 2H), 3.52–3.45 (m, 1H), 3.06 (t, J = 7.1 Hz, 2H), 1.40 (d, J = 6.8 Hz, 6H), 1.17 (d, J = 6.7 Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.38, 161.92, 139.34, 138.97, 135.57, 132.38, 129.06, 128.80, 127.98, 126.75, 126.41, 126.21, 125.89, 48.70, 45.70, 38.01, 32.46, 28.46, 19.95, 19.16; HRMS Calcd for $\text{C}_{22}\text{H}_{27}\text{ClN}_2\text{NaO}_2$ [M+Na $^+$]: 409.1659; Found: 409.1672.



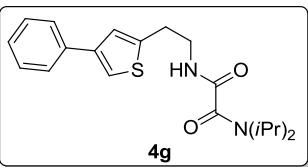
Pale yellow oil; Yield (85%, 73.1 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.59 (dd, J = 12.6, 7.9 Hz, 3H), 7.47–7.41 (m, 3H), 7.37–7.31 (m, 2H), 7.06 (br s, 1H), 4.75–4.69 (m, 1H), 3.64–3.59 (m, 2H), 3.52–3.45 (m, 1H), 3.07 (t, J = 7.1 Hz, 2H), 1.41 (d, J = 6.8 Hz, 6H), 1.17 (d, J = 6.6 Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.38, 161.91, 140.01, 138.95, 137.30, 132.38, 128.79, 127.99, 126.80, 126.18, 126.16, 122.70, 48.70, 45.70, 38.08, 34.88, 19.95, 19.16; HRMS Calcd for $\text{C}_{22}\text{H}_{27}\text{BrN}_2\text{NaO}_2$ [M+Na $^+$]: 453.1154; Found: 453.1136.



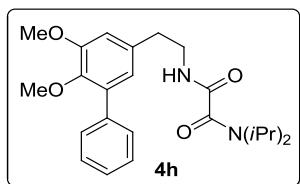
Pale yellow oil; Yield (66%, 50.4 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.58 (d, J = 7.3 Hz, 2H), 7.42 (t, J = 7.5 Hz, 2H), 7.34 (t, J = 7.3 Hz, 1H), 7.04 (s, 1H), 6.99 (s, 2H), 6.76 (s, 1H), 4.68–4.62 (m, 1H), 3.86 (s, 3H), 3.63–3.58 (m, 2H), 3.52–3.45 (m, 1H), 2.90 (t, J = 7.1 Hz, 2H), 1.40 (d, J = 6.8 Hz, 6H), 1.18 (d, J = 6.7 Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.41, 162.13, 159.32, 142.14, 140.08, 139.51, 127.85, 126.61, 126.36, 119.32, 112.25, 110.35, 54.49, 48.78, 45.64, 39.43, 34.77, 19.95, 19.18; HRMS Calcd for $\text{C}_{23}\text{H}_{30}\text{N}_2\text{NaO}_3$ [M+Na $^+$]: 405.2154; Found: 405.2157.



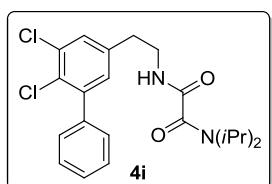
Pale yellow solid; Yield (70%, 51.2 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.58 (d, J = 7.2 Hz, 2H), 7.42 (t, J = 7.5 Hz, 2H), 7.33 (t, J = 7.3 Hz, 1H), 7.27 (s, 1H), 7.24 (s, 1H), 7.03 (s, 1H), 6.98 (br s, 1H), 4.69–4.62 (m, 1H), 3.62–3.57 (m, 2H), 3.52–3.45 (m, 1H), 2.89 (t, J = 7.2 Hz, 2H), 2.40 (s, 3H), 1.40 (d, J = 6.8 Hz, 6H), 1.18 (d, J = 6.7 Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.39, 162.15, 140.75, 140.27, 138.02, 137.81, 127.81, 127.63, 126.36, 126.34, 125.45, 123.90, 48.76, 45.64, 39.57, 34.57, 20.58, 19.96, 19.18; HRMS Calcd for $\text{C}_{23}\text{H}_{30}\text{N}_2\text{NaO}_2$ [M+Na $^+$]: 389.2205; Found: 389.2206.



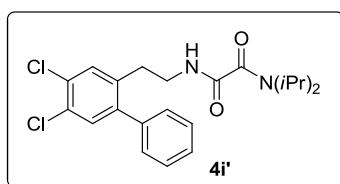
Yellow solid; Yield (87%, 62.3 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.55 (d, $J = 7.3$ Hz, 2H), 7.37 (t, $J = 7.6$ Hz, 2H), 7.29–7.26 (m, 2H), 7.19 (br s, 1H), 7.17 (s, 1H), 4.66–4.59 (m, 1H), 3.65–3.60 (m, 2H), 3.52–3.45 (m, 1H), 3.10 (t, $J = 6.9$ Hz, 2H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.19 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.51, 162.21, 141.31, 140.83, 134.95, 127.88, 126.21, 125.38, 123.95, 117.89, 48.84, 45.61, 39.60, 29.04, 19.93, 19.17; HRMS Calcd for $\text{C}_{20}\text{H}_{26}\text{N}_2\text{NaO}_2\text{S} [\text{M}+\text{Na}^+]$: 381.1613; Found: 381.1626.



Pale yellow solid; Yield (45%, 37.1 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.55–7.53 (m, 2H), 7.42–7.38 (m, 2H), 7.35–7.31 (m, 1H), 7.00 (br s, 1H), 6.78 (s, 2H), 4.72–4.65 (m, 1H), 3.91 (s, 3H), 3.60–3.54 (m, 5H), 3.53–3.46 (m, 1H), 2.84 (t, $J = 7.1$ Hz, 2H), 1.40 (t, $J = 6.2$ Hz, 6H), 1.19 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.35, 162.06, 152.22, 144.28, 137.25, 134.95, 133.43, 128.36, 127.22, 126.27, 121.87, 111.08, 59.73, 55.13, 48.75, 45.67, 39.55, 34.53, 19.96, 19.18; HRMS Calcd for $\text{C}_{24}\text{H}_{32}\text{N}_2\text{NaO}_4 [\text{M}+\text{Na}^+]$: 435.2260; Found: 435.2261.

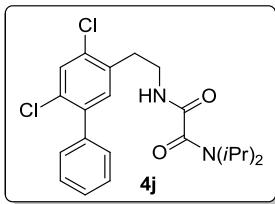


Pale yellow solid; Yield (51%, 42.8 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.45–7.37 (m, 5H), 7.33 (d, $J = 2.0$ Hz, 1H), 7.14 (br s, 1H), 7.10 (d, $J = 2.0$ Hz, 1H), 4.63–4.56 (m, 1H), 3.59–3.53 (m, 2H), 3.52–3.46 (m, 1H), 2.85 (t, $J = 7.2$ Hz, 2H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.18 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 162.41, 161.99, 141.90, 138.26, 137.20, 132.64, 129.23, 128.72, 128.41, 128.36, 127.25, 127.11, 48.84, 45.70, 39.10, 33.74, 19.94, 19.17; HRMS Calcd for $\text{C}_{22}\text{H}_{27}\text{Cl}_2\text{N}_2\text{O}_2 [\text{M}+\text{H}^+]$: 421.1450; Found: 421.1455.

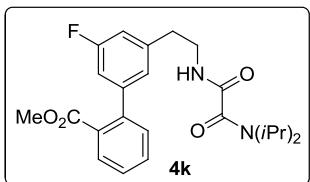


Pale yellow oil; Yield (9%, 7.6 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.43–7.37 (m, 4H), 7.31 (s, 1H), 7.26–7.24 (m, 2H), 7.04 (br s, 1H), 4.60–4.53 (m, 1H), 3.51–3.44 (m, 1H), 3.33–3.28 (m, 2H), 2.77 (t, $J = 7.4$ Hz, 2H), 1.38 (d, $J = 6.8$ Hz, 6H), 1.19 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101

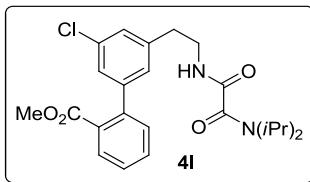
MHz, CDCl₃) δ 163.18, 142.38, 139.15, 136.36, 131.93, 131.43, 131.41, 130.36, 129.00, 128.66, 127.94, 49.80, 46.66, 39.59, 38.82, 36.57, 32.15, 27.99, 20.93, 20.12; HRMS Calcd for C₂₂H₂₇Cl₂N₂O₂ [M+H⁺]: 421.1450; Found: 421.1465.



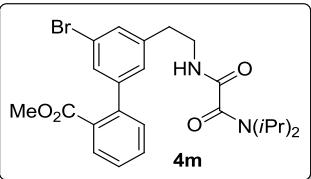
Pale yellow oil; Yield (71%, 59.6 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.50 (s, 1H), 7.44–7.36 (m, 5H), 7.23 (s, 1H), 7.10 (br s, 1H), 4.73–4.67 (m, 1H), 3.57–3.54 (m, 2H), 3.52–3.46 (m, 1H), 3.00 (t, *J* = 7.1 Hz, 2H), 1.40 (d, *J* = 6.8 Hz, 6H), 1.17 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 206.15, 162.33, 161.78, 138.48, 137.22, 134.15, 132.57, 132.51, 130.45, 129.52, 128.51, 127.25, 127.05, 48.69, 45.72, 37.84, 31.77, 30.06, 19.93, 19.14; HRMS Calcd for C₂₂H₂₇Cl₂N₂O₂ [M+H⁺]: 421.1450; Found: 421.1457.



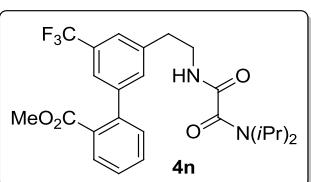
Off-white solid; Yield (49%, 41.9 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.84 (dd, *J* = 7.7, 1.2 Hz, 1H), 7.55–7.51 (m, 1H), 7.45–7.40 (m, 1H), 7.36 (dd, *J* = 7.6, 1.0 Hz, 1H), 7.01 (s, 1H), 6.95 (br s, 1H), 6.93–6.87 (m, 2H), 4.69–4.63 (m, 1H), 3.68 (s, 3H), 3.59–3.54 (m, 2H), 3.52–3.46 (m, 1H), 2.89 (t, *J* = 7.2 Hz, 2H), 1.40 (d, *J* = 6.8 Hz, 6H), 1.19 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 167.68, 162.42, 161.99, 161.65 (d, *J*_{C-F} = 244.0 Hz), 142.81 (d, *J*_{C-F} = 9.0 Hz), 140.35 (d, *J*_{C-F} = 2.0 Hz), 139.70 (d, *J*_{C-F} = 8.0 Hz), 130.58, 129.77, 129.68, 129.11, 126.82, 123.79 (d, *J*_{C-F} = 3.0 Hz), 113.56 (d, *J*_{C-F} = 21.0 Hz), 112.84 (d, *J*_{C-F} = 22.0 Hz), 51.24, 48.78, 45.66, 39.35, 34.36, 19.94, 19.17; HRMS Calcd for C₂₂H₂₇FN₂NaO₂ [M+Na⁺]: 393.1954; Found: 393.1968.



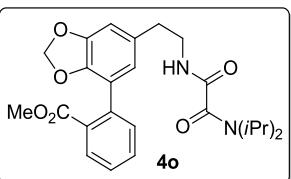
Pale yellow solid; Yield (62%, 55.1 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.85 (dd, *J* = 7.7, 0.9 Hz, 1H), 7.55–7.51 (m, 1H), 7.45–7.41 (m, 1H), 7.35 (d, *J* = 7.6 Hz, 1H), 7.18 (d, *J* = 11.6 Hz, 2H), 7.05 (s, 1H), 7.02 (s, 1H), 4.67–4.60 (m, 1H), 3.68 (s, 3H), 3.59–3.54 (m, 2H), 3.52–3.45 (m, 1H), 2.87 (t, *J* = 7.2 Hz, 2H), 1.40 (d, *J* = 6.8 Hz, 6H), 1.19 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 206.13, 167.59, 162.48, 162.07, 142.42, 140.20, 139.20, 133.04, 130.64, 129.81, 129.56, 129.18, 126.87, 126.73, 126.37, 125.87, 51.24, 48.82, 45.63, 39.32, 34.24, 30.06, 19.93, 19.16; HRMS Calcd for C₂₄H₂₉ClN₂NaO₄ [M+Na⁺]: 467.1714; Found: 467.1716.



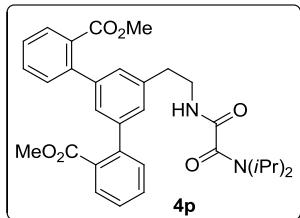
Pale yellow solid; Yield (65%, 63.4 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.86 (d, $J = 7.7$ Hz, 1H), 7.55–7.51 (m, 1H), 7.43 (dd, $J = 7.5, 6.7$ Hz, 1H), 7.34 (dd, $J = 9.4, 4.3$ Hz, 3H), 7.10 (s, 1H), 7.01 (br s, 1H), 4.67–4.61 (m, 1H), 3.68 (s, 3H), 3.59–3.54 (m, 2H), 3.52–3.46 (m, 1H), 2.87 (t, $J = 7.2$ Hz, 2H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.19 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 167.58, 162.45, 161.99, 142.68, 140.09, 139.42, 130.65, 129.84, 129.63, 129.55, 129.21, 128.76, 126.90, 126.86, 121.24, 51.25, 48.82, 45.67, 39.36, 34.22, 19.97, 19.18; HRMS Calcd for $\text{C}_{24}\text{H}_{29}\text{BrN}_2\text{NaO}_4$ [M+Na $^+$]: 511.1208; Found: 511.1206.



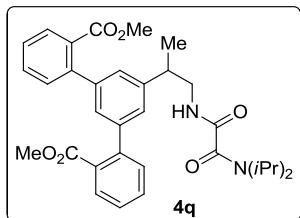
Yellow solid; Yield (53%, 50.7 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.89 (d, $J = 7.6$ Hz, 1H), 7.56 (t, $J = 7.4$ Hz, 1H), 7.47–7.37 (m, 5H), 7.10 (br s, 1H), 4.68–4.62 (m, 1H), 3.66 (s, 3H), 3.62–3.57 (m, 2H), 3.52–3.46 (m, 1H), 2.96 (t, $J = 7.1$ Hz, 2H), 1.40 (d, $J = 6.7$ Hz, 6H), 1.18 (d, $J = 6.6$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 167.45, 162.44, 161.90, 141.61, 140.18, 138.34, 131.41 (d, $J_{C-F} = 1.0$ Hz), 130.79, 129.94, 129.64, 129.54, 129.39, 127.07, 124.53, 123.38 (q, $J_{C-F} = 4.0$ Hz), 123.18 (q, $J_{C-F} = 270.0$ Hz), 122.84 (q, $J_{C-F} = 4.0$ Hz), 51.20, 48.78, 45.70, 39.36, 34.40, 19.92, 19.15; HRMS Calcd for $\text{C}_{25}\text{H}_{29}\text{F}_3\text{N}_2\text{NaO}_4$ [M+Na $^+$]: 501.1977; Found: 501.1983.



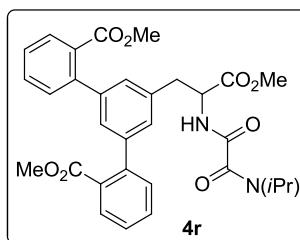
Off-white solid; Yield (81%, 73.5 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.88 (dd, $J = 7.8, 1.1$ Hz, 1H), 7.57–7.53 (m, 1H), 7.46–7.39 (m, 2H), 7.01 (br s, 1H), 6.70 (dd, $J = 5.9, 1.5$ Hz, 2H), 5.90 (s, 2H), 4.71–4.65 (m, 1H), 3.74 (s, 3H), 3.57–3.46 (m, 3H), 1.41 (d, $J = 6.8$ Hz, 6H), 1.19 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 168.51, 163.37, 163.11, 147.43, 143.57, 136.42, 132.37, 131.88, 131.20, 130.86, 130.22, 127.88, 122.85, 122.47, 108.51, 101.13, 52.23, 49.76, 46.64, 40.76, 35.39, 20.95, 20.18; HRMS Calcd for $\text{C}_{25}\text{H}_{30}\text{N}_2\text{NaO}_6$ [M+Na $^+$]: 477.2002; Found: 477.2000.



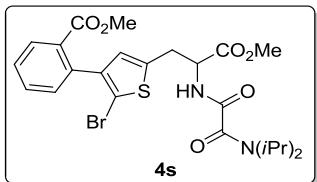
Pale yellow oil; Yield (60%, 65.3 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.83–7.81 (m, 2H), 7.54–7.50 (m, 2H), 7.42–7.39 (m, 4H), 7.16 (d, $J = 1.3$ Hz, 2H), 7.12 (d, $J = 1.4$ Hz, 1H), 7.06 (br s, 1H), 4.70–4.64 (m, 1H), 3.67 (s, 6H), 3.62–3.57 (m, 2H), 3.51–3.44 (m, 1H), 2.93 (t, $J = 7.2$ Hz, 2H), 1.40 (d, $J = 6.8$ Hz, 6H), 1.17 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 168.15, 162.46, 162.10, 141.16, 140.62, 137.14, 130.41, 130.00, 129.91, 128.95, 126.86, 126.43, 125.91, 51.17, 48.76, 45.60, 39.68, 34.51, 19.92, 19.16; HRMS Calcd for $\text{C}_{32}\text{H}_{36}\text{N}_2\text{NaO}_6$ [$\text{M}+\text{Na}^+$]: 567.2471; Found: 567.2478.



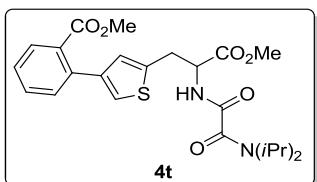
Pale yellow oil; Yield (72%, 80.4 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.82–7.80 (m, 2H), 7.54–7.50 (m, 2H), 7.40 (dd, $J = 12.7, 4.9$ Hz, 4H), 7.15 (d, $J = 2.3$ Hz, 3H), 6.94 (br s, 1H), 4.57–4.51 (m, 1H), 3.70–3.61 (m, 7H), 3.48–3.41 (m, 1H), 3.39–3.33 (m, 1H), 3.09–3.03 (m, 1H), 1.38–1.33 (m, 9H), 1.16 (d, $J = 6.6$ Hz, 3H), 1.11 (d, $J = 6.7$ Hz, 3H); ^{13}C NMR (101 MHz, CDCl_3) δ 168.23, 162.65, 162.31, 142.41, 141.16, 140.58, 130.38, 130.08, 129.83, 128.92, 126.42, 125.93, 125.44, 51.16, 48.81, 45.45, 45.01, 38.69, 19.89, 19.82, 19.15, 19.13, 18.43; HRMS Calcd for $\text{C}_{33}\text{H}_{38}\text{N}_2\text{NaO}_6$ [$\text{M}+\text{Na}^+$]: 581.2628; Found: 581.2644.



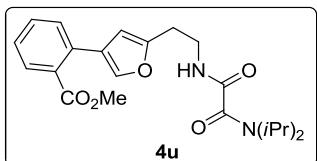
Pale yellow oil; Yield (58%, 69.8 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.81 (d, $J = 7.6$ Hz, 2H), 7.51 (t, $J = 7.1$ Hz, 2H), 7.39 (t, $J = 8.3$ Hz, 5H), 7.09 (s, 3H), 4.90–4.85 (m, 1H), 4.59–4.53 (m, 1H), 3.69 (d, $J = 17.4$ Hz, 9H), 3.49–3.42 (m, 1H), 3.24 (d, $J = 5.9$ Hz, 2H), 1.39 (t, $J = 6.8$ Hz, 6H), 1.18 (d, $J = 6.6$ Hz, 3H), 1.07 (d, $J = 6.6$ Hz, 3H); ^{13}C NMR (101 MHz, CDCl_3) δ 170.27, 168.01, 161.95, 161.38, 141.07, 140.66, 134.30, 130.39, 129.97, 128.94, 127.32, 126.57, 126.46, 52.41, 51.61, 51.19, 48.76, 45.63, 36.93, 19.92, 19.76, 19.14, 19.11; HRMS Calcd for $\text{C}_{34}\text{H}_{38}\text{N}_2\text{NaO}_8$ [$\text{M}+\text{Na}^+$]: 625.2526; Found: 625.2542.



Yellow oil; Yield (80%, 88.3 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.94 (dd, $J = 7.8, 1.1$ Hz, 1H), 7.55–7.51 (m, 1H), 7.45–7.40 (m, 2H), 7.31–7.29 (m, 1H), 6.65 (s, 1H), 4.86–4.82 (m, 1H), 4.61–4.54 (m, 1H), 3.79 (s, 3H), 3.74 (s, 3H), 3.54–3.47 (m, 1H), 3.41–3.31 (m, 2H), 1.42 (dd, $J = 6.8, 2.5$ Hz, 6H), 1.21 (t, $J = 6.3$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 169.52, 166.72, 162.13, 161.39, 140.37, 135.80, 135.04, 130.79, 130.33, 129.82, 129.48, 128.24, 127.16, 107.63, 52.04, 51.92, 51.45, 48.88, 45.70, 31.51, 20.00, 19.94, 19.19, 19.15; HRMS Calcd for $\text{C}_{24}\text{H}_{29}\text{BrN}_2\text{NaO}_6\text{S}$ [$\text{M}+\text{Na}^+$]: 575.0827; Found: 575.0810.

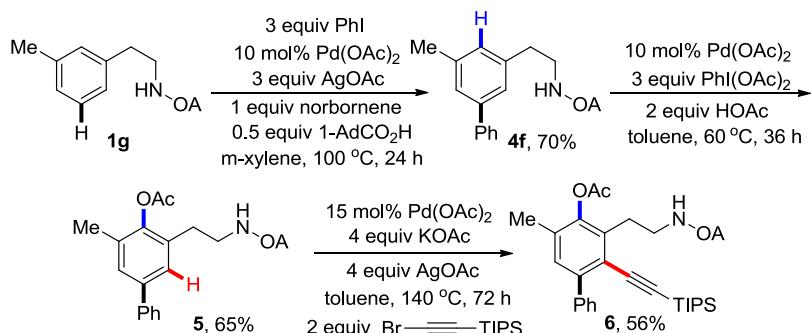


Yellow oil; Yield (61%, 57.8 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.74 (dd, $J = 7.5, 1.9$ Hz, 1H), 7.49–7.45 (m, 1H), 7.38–7.33 (m, 3H), 7.06 (d, $J = 1.4$ Hz, 1H), 6.82 (d, $J = 1.3$ Hz, 1H), 4.88–4.84 (m, 1H), 4.61–4.54 (m, 1H), 3.78 (s, 3H), 3.73 (s, 3H), 3.53–3.46 (m, 1H), 3.42 (d, $J = 5.3$ Hz, 2H), 1.42 (dd, $J = 6.8, 1.7$ Hz, 6H), 1.20 (dd, $J = 8.5, 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 169.70, 168.16, 162.08, 161.44, 140.46, 135.80, 135.72, 130.33, 129.99, 129.63, 128.66, 127.52, 126.40, 121.01, 52.26, 51.83, 51.31, 48.84, 45.67, 31.23, 19.97, 19.95, 19.18; HRMS Calcd for $\text{C}_{24}\text{H}_{31}\text{N}_2\text{O}_6\text{S}$ [$\text{M}+\text{H}^+$]: 475.1903; Found: 475.1918.



Yellow oil; Yield (21%, 16.8 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.72 (dd, $J = 7.7, 1.0$ Hz, 1H), 7.49–7.45 (m, 2H), 7.38–7.32 (m, 2H), 7.08 (br s, 1H), 6.20 (s, 1H), 4.75–4.69 (m, 1H), 3.81 (s, 3H), 3.63–3.58 (m, 2H), 3.53–3.47 (m, 1H), 2.91 (t, $J = 6.7$ Hz, 2H), 1.41 (d, $J = 6.8$ Hz, 6H), 1.21 (d, $J = 6.7$ Hz, 6H); ^{13}C NMR (101 MHz, CDCl_3) δ 168.23, 162.39, 161.97, 151.83, 137.94, 131.69, 130.39, 129.82, 129.41, 128.68, 126.22, 125.03, 107.50, 51.35, 48.78, 45.69, 37.00, 27.14, 19.98, 19.19; HRMS Calcd for $\text{C}_{22}\text{H}_{29}\text{N}_2\text{O}_5$ [$\text{M}+\text{H}^+$]: 401.2076; Found: 401.2085.

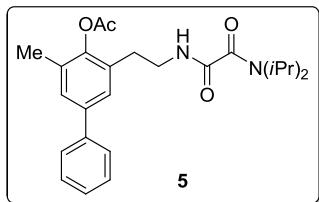
7. Pd-catalyzed trifunctionalization of presubstituted β -arylethamide



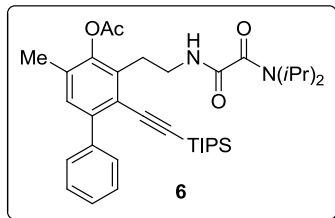
The first step: a mixture of oxalamide **1g** (0.2 mmol, 1.0 eq), iodobenzene (0.6 mmol, 3.0 eq), Pd(OAc)₂ (4.5 mg, 10 mol%), AgOAc (100 mg, 0.6 mmol, 3.0 eq), norbornene (18.8 mg, 0.2 mmol, 1.0 eq), 1-AdOH (18 mg, 0.1 mmol, 0.05 eq) and 1 mL m-xylene in a 15 mL glass vial was heated at 100 °C with vigorous stirring for 24 hours. The reaction mixture was cooled to room temperature, and diluted with ethyl acetate and filtered through celite. The filtrate was concentrated in vacuo and purified by column chromatography on silica gel (Ethyl acetate/Petroleum ether = 1:8) to give product **4f**.

The second step: a mixture of presubstituted β -arylethamide **4f** (0.2 mmol, 1.0 eq), Pd(OAc)₂ (4.5 mg, 10 mol%), PhI(OAc)₂ (0.6 mmol, 3.0 eq), HOAc (0.4 mmol, 2.0 eq) and 1 mL toluene in a 15 mL glass vial was heated at 60 °C with vigorous stirring for 36 hours. The reaction mixture was cooled to room temperature, and diluted with ethyl acetate and filtered through celite. The filtrate was concentrated in vacuo and purified by column chromatography on silica gel (Ethyl acetate/Petroleum ether = 1:4) to give product **5**.

The third step: a mixture of presubstituted β -arylethamide **5** (0.2 mmol, 1.0 eq), bromoalkyne (0.4 mmol, 2.0 eq), Pd(OAc)₂ (6.6 mg, 15 mol%), KOAc (0.8 mmol, 4.0 eq), AgOAc (0.8 mmol, 4.0 eq), and 1 mL toluene in a 15 mL glass vial was heated at 140 °C with vigorous stirring for 72 hours. The reaction mixture was cooled to room temperature, and diluted with ethyl acetate and filtered through celite. The filtrate was concentrated in vacuo and purified by column chromatography on silica gel (Ethyl acetate/Petroleum ether = 1:20) to give product **6**.

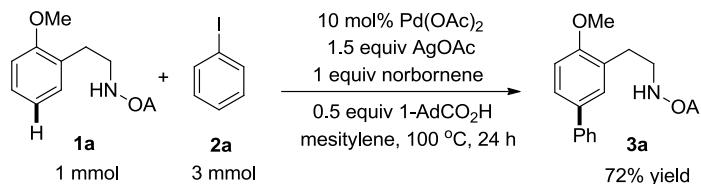


Yellow oil; Yield (65%, 55.1 mg); ¹H NMR (400 MHz, CDCl₃) δ 7.57–7.55 (m, 2H), 7.43–7.39 (m, 2H), 7.34–7.31 (m, 3H), 7.01 (br s, 1H), 4.66–4.59 (m, 1H), 3.57–3.44 (m, 2H), 3.47 (dd, *J* = 13.6, 6.8 Hz, 1H), 2.80 (t, *J* = 7.3 Hz, 2H), 2.41 (s, 3H), 2.22 (s, 3H), 1.41 (d, *J* = 6.8 Hz, 6H), 1.16 (d, *J* = 6.7 Hz, 6H); ¹³C NMR (101 MHz, CDCl₃) δ 168.57, 162.50, 162.15, 146.72, 139.50, 138.47, 130.13, 129.97, 127.83, 127.59, 126.43, 126.31, 125.99, 48.75, 45.61, 38.39, 29.39, 19.92, 19.74, 19.16, 15.77; HRMS Calcd for C₂₅H₃₂N₂NaO₄ [M+Na⁺]: 447.2260; Found: 447.2256.

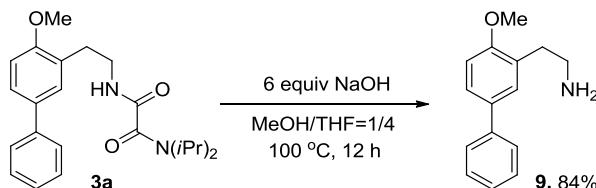


Yellow oil; Yield (56%, 67.6 mg); ^1H NMR (400 MHz, CDCl_3) δ 7.49 (d, $J = 6.8$ Hz, 2H), 7.37–7.28 (m, 3H), 7.10 (s, 1H), 6.98 (br s, 1H), 4.81–4.75 (m, 1H), 3.56–3.48 (m, 3H), 3.07 (s, 2H), 2.43 (s, 3H), 2.18 (s, 3H), 1.43 (d, $J = 6.8$ Hz, 6H), 1.22 (d, $J = 6.7$ Hz, 6H), 0.97 (d, $J = 3.0$ Hz, 21H); ^{13}C NMR (101 MHz, CDCl_3) δ 169.66, 163.35, 162.90, 147.60, 143.42, 140.54, 133.64, 131.38, 130.72, 129.51, 127.96, 127.35, 121.26, 103.74, 98.70, 49.57, 46.64, 39.05, 29.36, 21.03, 20.71, 20.19, 18.68, 16.89, 11.33; HRMS Calcd for $\text{C}_{36}\text{H}_{52}\text{N}_2\text{NaO}_4\text{Si} [\text{M}+\text{Na}^+]$: 627.3594; Found: 627.3597.

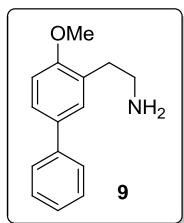
8. Scale up and removal of directing group



A mixture of **1a** (1 mmol, 306.4 mg), **2** (3 mmol, 3.0 eq), $\text{Pd}(\text{OAc})_2$ (22.4 mg, 10 mol%), AgOAc (250 mg, 1.5 mmol, 1.5 eq), norbornene (94 mg, 1 mmol, 1.0 eq), 1-AdCO₂H (90 mg, 0.5 mmol, 0.05 eq) and 5 mL mesitylene in a 15 mL glass vial was heated at 100 °C with vigorous stirring for 24 hours. The reaction mixture was cooled to room temperature, and diluted with ethyl acetate and filtered through celite. The filtrate was concentrated in vacuo and purified by column chromatography on silica gel to give product **3a**.



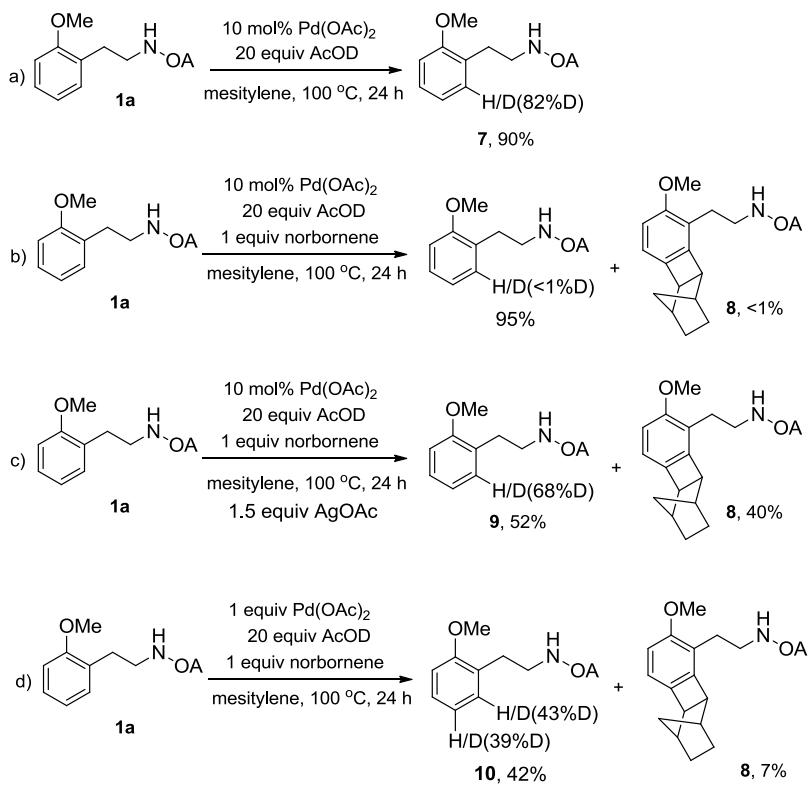
The compound **3a** (76.5 mg, 0.2 mmol) was dissolved in a mixture of MeOH/THF (0.2 mL /0.8 mL), NaOH (48 mg, 0.12 mmol, 6 eq) was then added. The mixture was heated to 100 °C and stirred for 12 hours. Water was added and the mixture was extracted with DCM. The combined organic layers was washed with water and brine, dried over anhydrous Na_2SO_4 , and concentrated in vacuo. The residue was purified by column chromatography on silica gel (Methanol/DCM = 1:20) to give the desired product **9** as white solid in 38.1 mg, 84% yield.



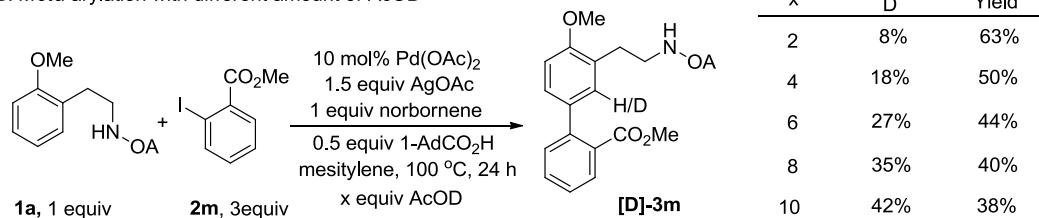
¹H NMR (400 MHz, CDCl₃) δ 7.56–7.54 (m, 2H), 7.45–7.39 (m, 4H), 7.32–7.27 (m, 1H), 6.93 (d, J = 8.4 Hz, 1H), 3.86 (s, 3H), 2.98 (t, J = 6.9 Hz, 2H), 2.84 (t, J = 6.9 Hz, 2H), 2.03 (br s, 2H); ¹³C NMR (101 MHz, CDCl₃) δ 157.39, 140.97, 133.60, 129.50, 128.82, 128.37, 126.87, 126.75, 126.19, 110.78, 55.57, 42.20, 34.83; HRMS Calcd for C₁₅H₁₈NO [M+H⁺]: 228.1388; Found: 228.1381.

9. Deuteration experiment

A: Deuteration study



B: Meta arylation with different amount of AcOD



Procedure for A:

- a) A mixture of compound **1a** (0.2 mmol), Pd(OAc)₂ (4.5 mg, 10 mol%), AcOD (4 mmol, 20.0 eq) and mesitylene (1 mL) was heated at 100 °C for 24 hours. The reaction mixture was cooled to room temperature, and concentrated in vacuo. The resulting residue was purified by column chromatography on silica gel (Ethyl acetate/Petroleum ether = 1:10) to give the deuterated product **7**.
- b) A mixture of compound **1a** (0.2 mmol), Pd(OAc)₂ (4.5 mg, 10 mol%), AcOD (4 mmol, 20.0 eq), norbornene (18.8 mg, 1.0 eq) and mesitylene (1 mL) was heated at 100 °C for 24 hours. The reaction mixture was cooled to room temperature, and concentrated in vacuo. The resulting residue was purified by column chromatography on silica gel (Ethyl acetate/Petroleum ether = 1:10) to give the product.
- c) A mixture of compound **1a** (0.2 mmol), Pd(OAc)₂ (4.5 mg, 10 mol%), AcOD (4 mmol, 20.0 eq), AgOAc (50 mg, 1.5 eq), norbornene (18.8 mg, 1.0 eq) and mesitylene (1 mL) was heated at 100 °C for 24 hours. The reaction mixture was cooled to room temperature, and concentrated in vacuo. The resulting residue was purified by column chromatography on silica gel (Ethyl acetate/Petroleum ether = 1:10) to give the product.

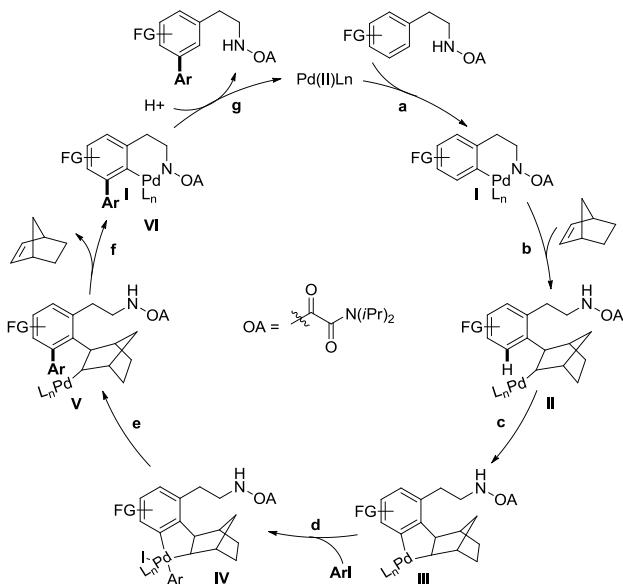
Procedure for B:

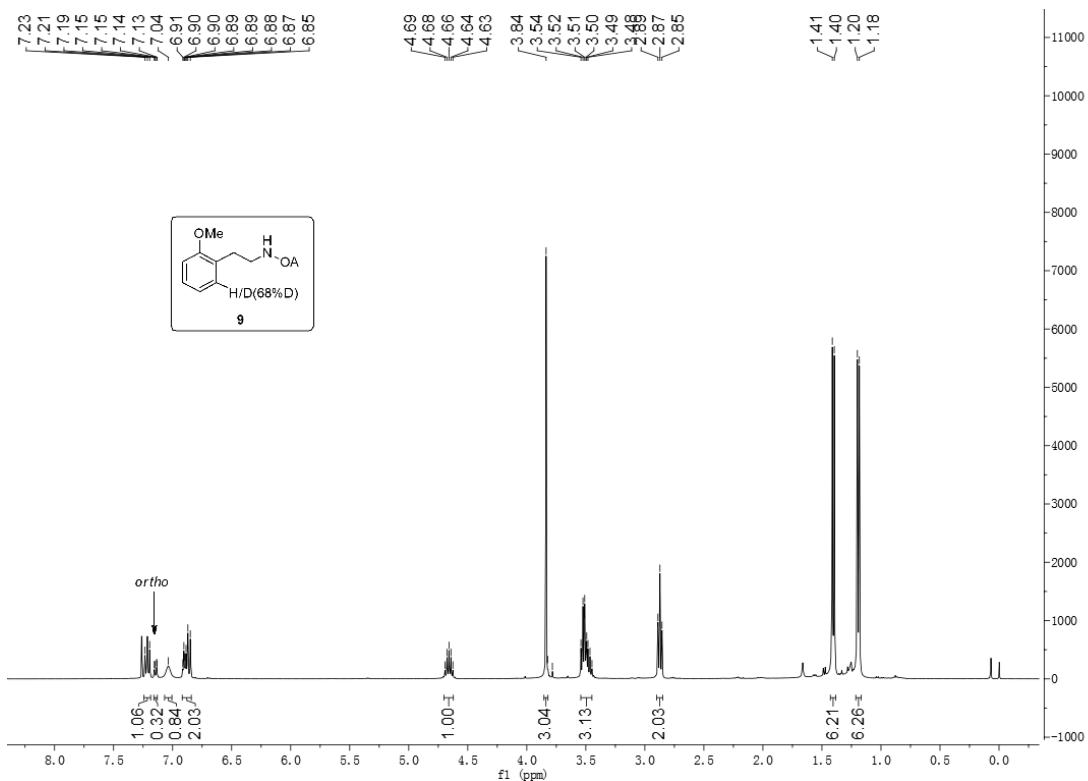
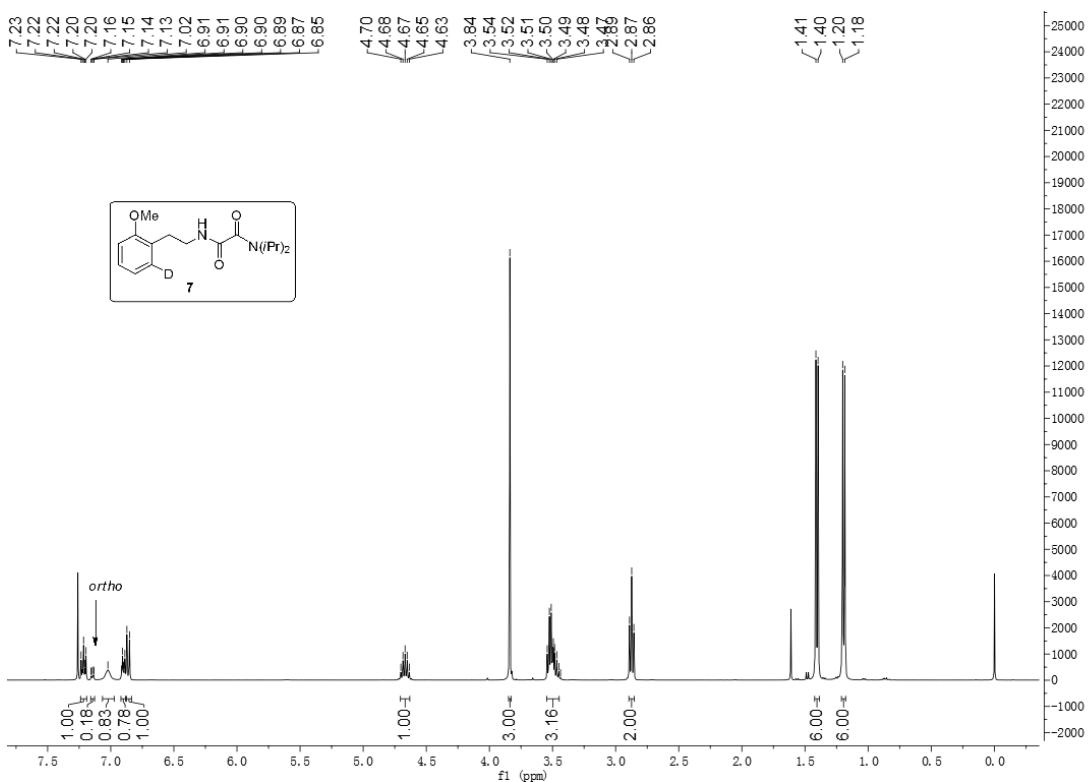
A mixture of compound **7** (0.2 mmol, 1.0 eq), **2m** (0.6 mmol, 3.0 eq), Pd(OAc)₂ (4.5 mg, 10 mol%), AgOAc (50 mg, 1.5 eq), norbornene (18.8 mg, 1.0 eq), 1-AdCO₂H (18 mg, 0.5 eq) and mesitylene (1 mL) was heated at 100 °C for 4 hours. The reaction mixture was cooled to room temperature, and concentrated in vacuo. The resulting residue was purified by column chromatography on silica gel (Ethyl acetate/Petroleum ether = 1:3) to give the product.

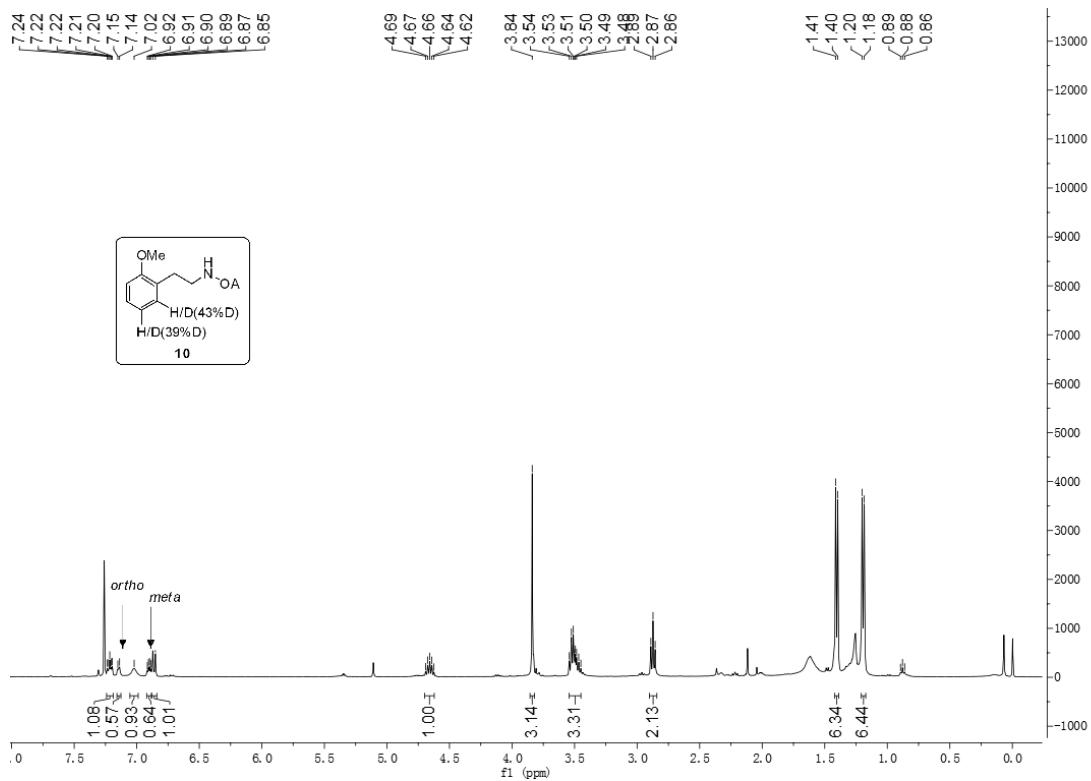
Procedure for C:

A mixture of compound **1a** (0.2 mmol, 1.0 eq), **2m** (0.6 mmol, 3.0 eq), Pd(OAc)₂ (4.5 mg, 10 mol%), AgOAc (50 mg, 1.5 eq), norbornene (18.8 mg, 1.0 eq), 1-AdCO₂H (18 mg, 0.5 eq), AcOD (from 2.0 eq to 10.0 eq) and mesitylene (1 mL) was heated at 100 °C for 24 hours. The reaction mixture was cooled to room temperature, and concentrated in vacuo. The resulting residue was purified by column chromatography on silica gel (Ethyl acetate/Petroleum ether = 1:3) to give the product **[D]-3m**.

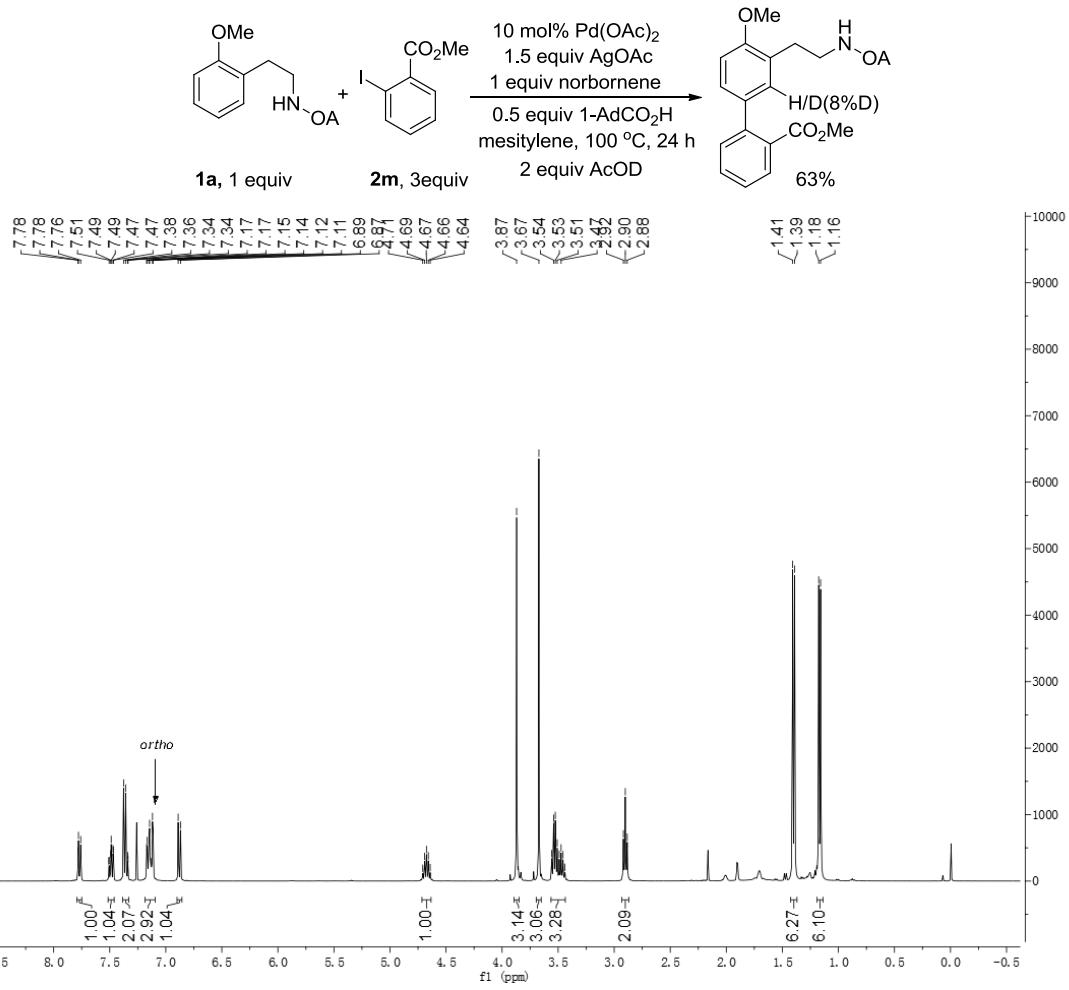
Plausible catalytic cycle for *meta* arylation

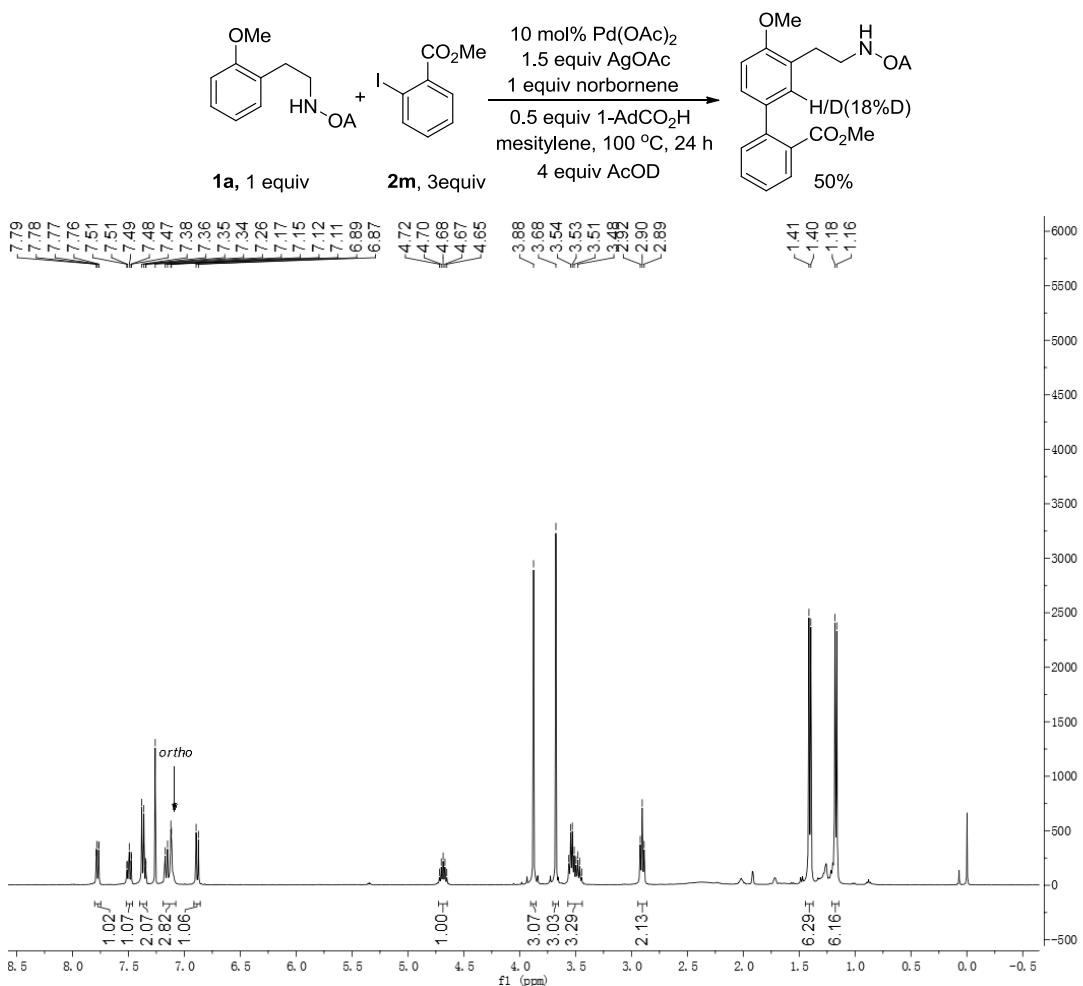


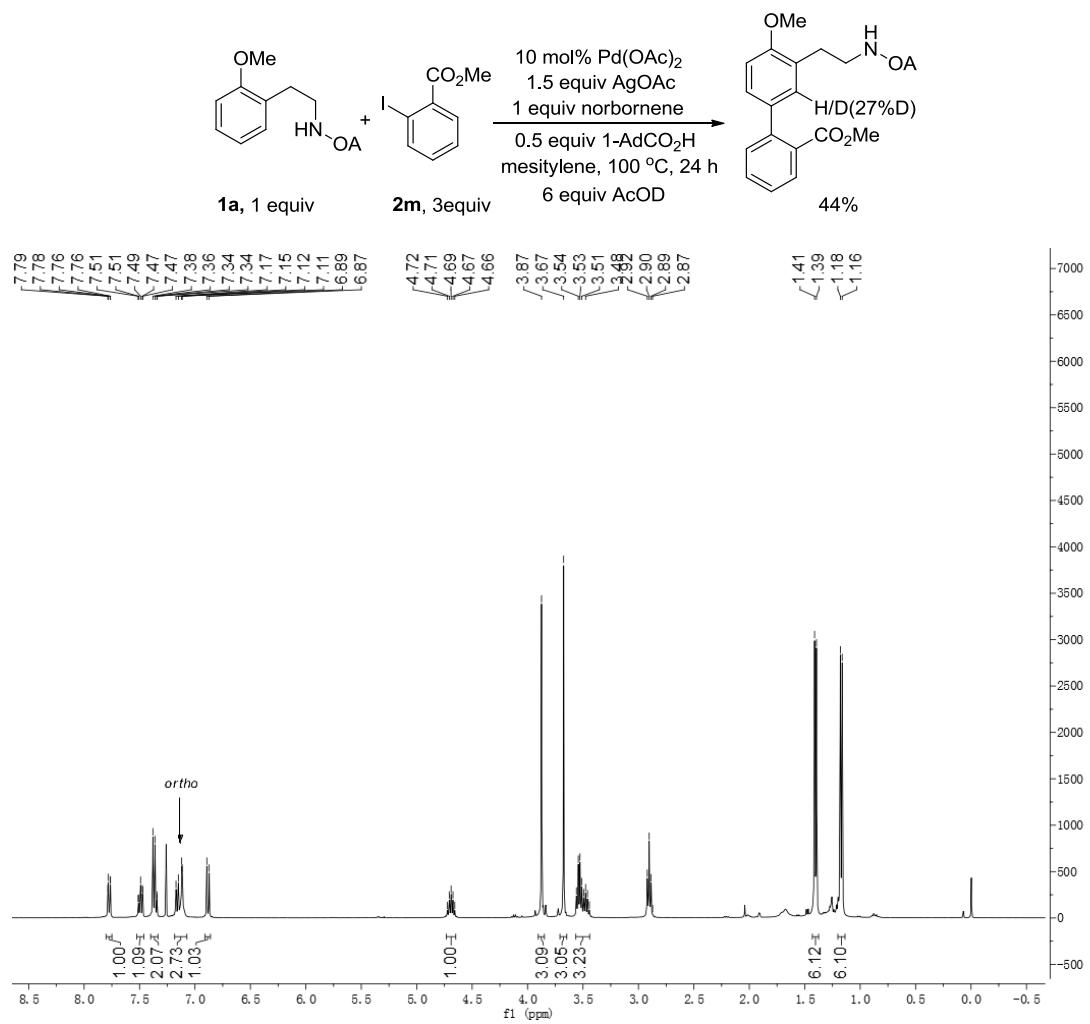


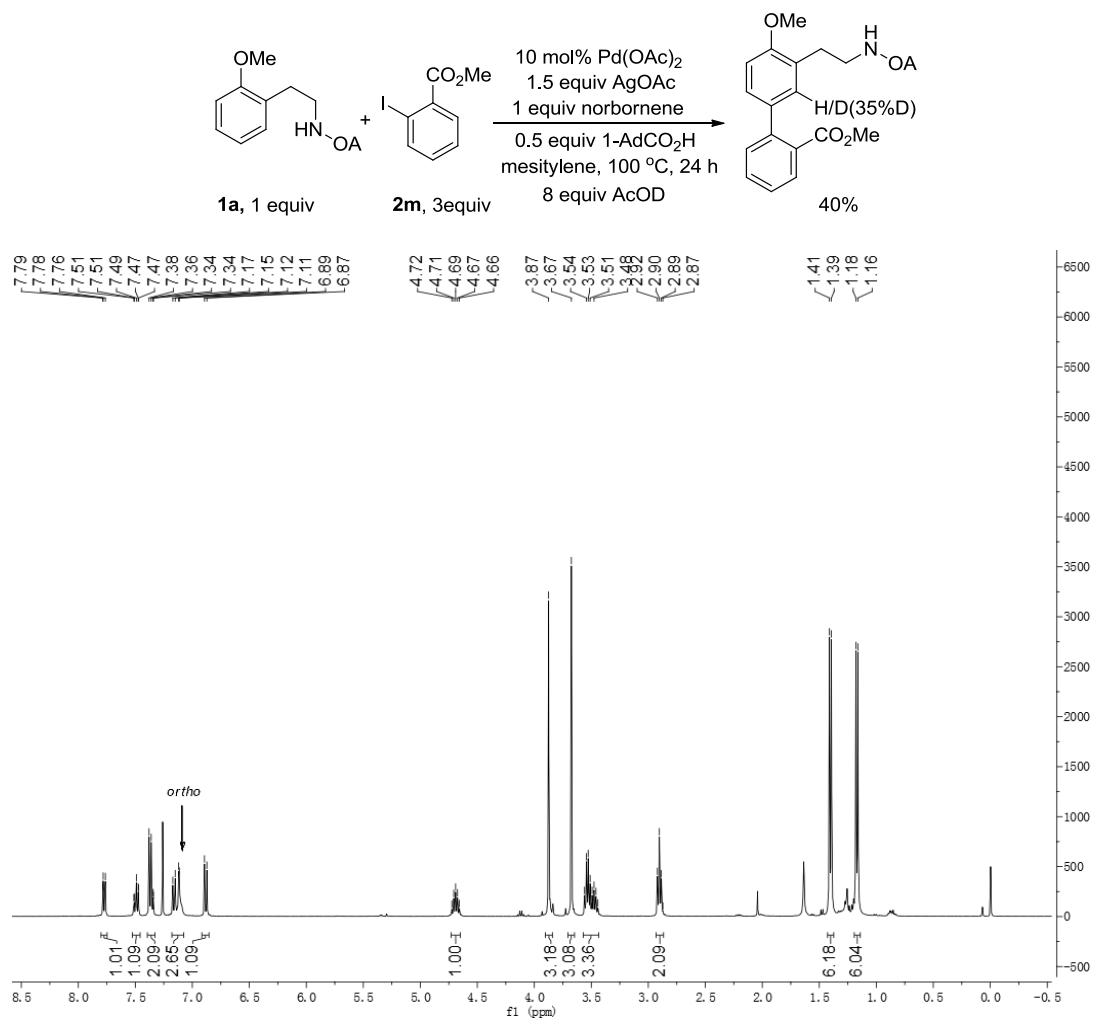


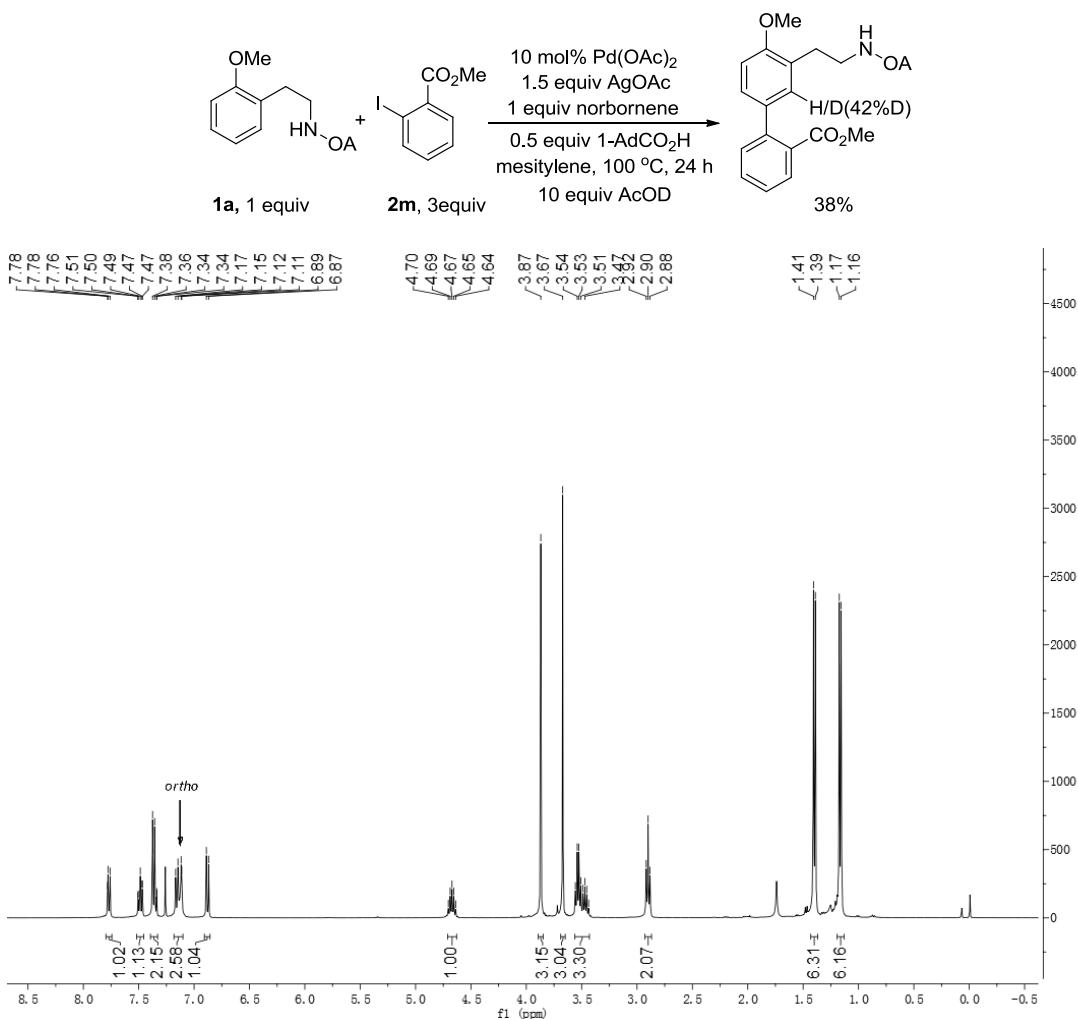
¹H NMR spectra of reaction with different amount fo AcOD







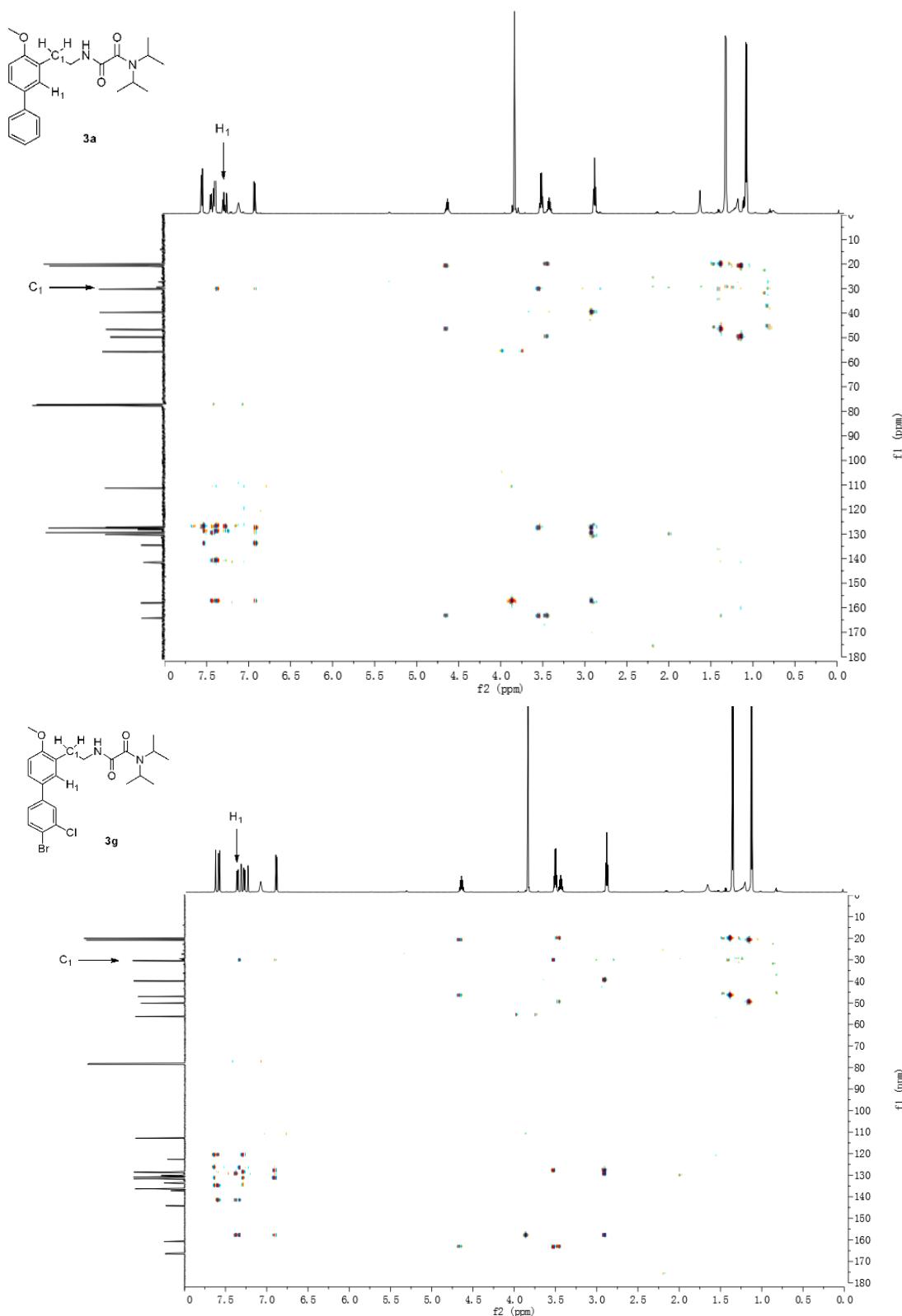


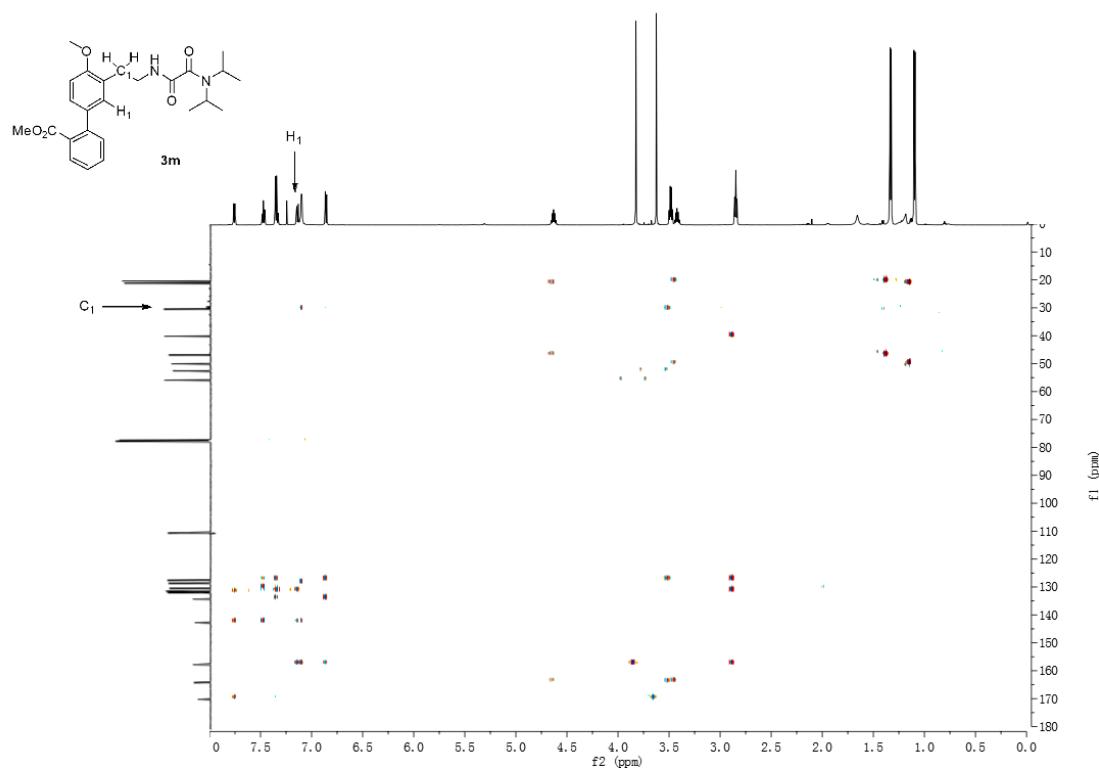
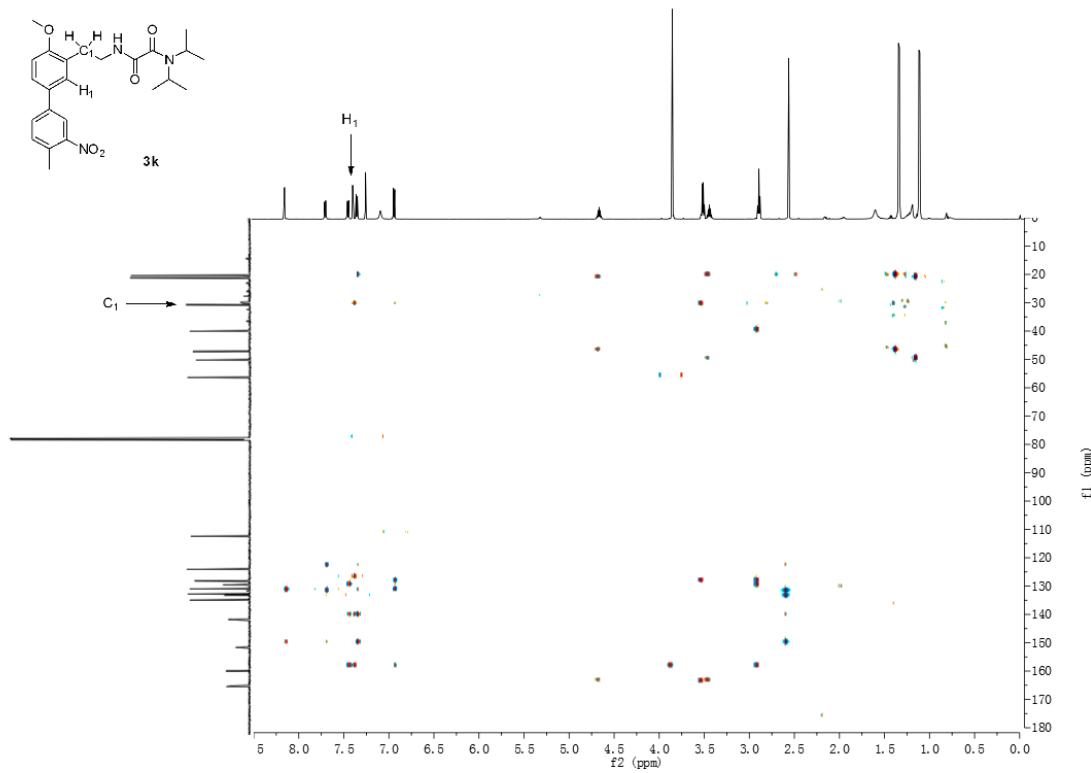


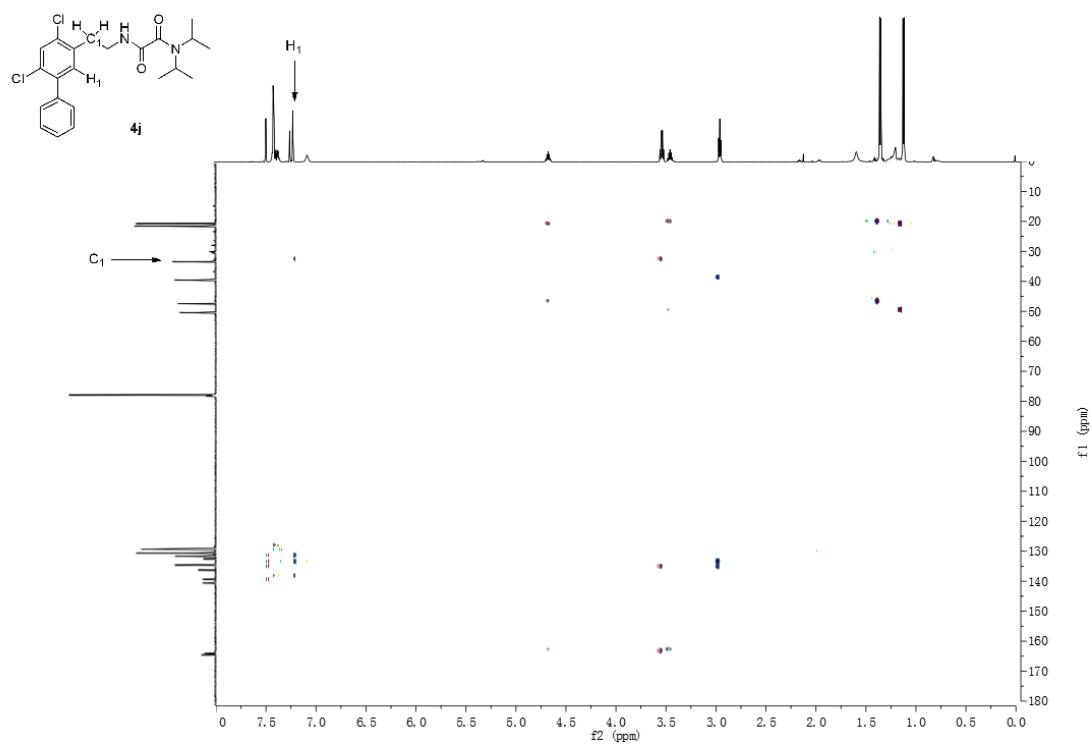
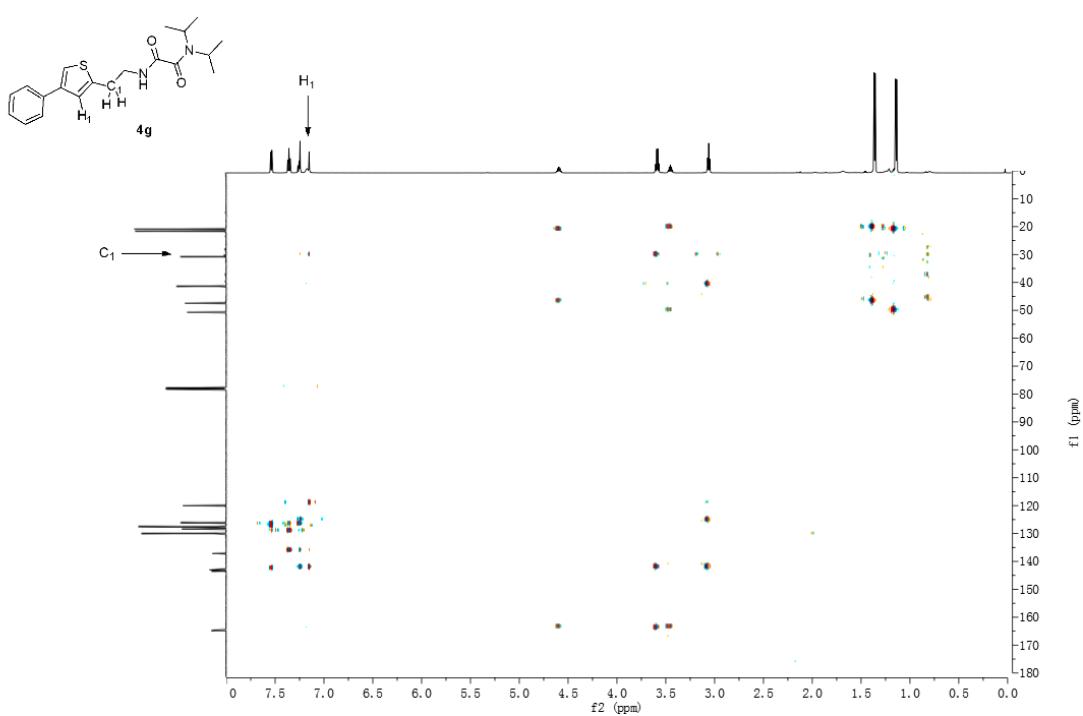
10. References

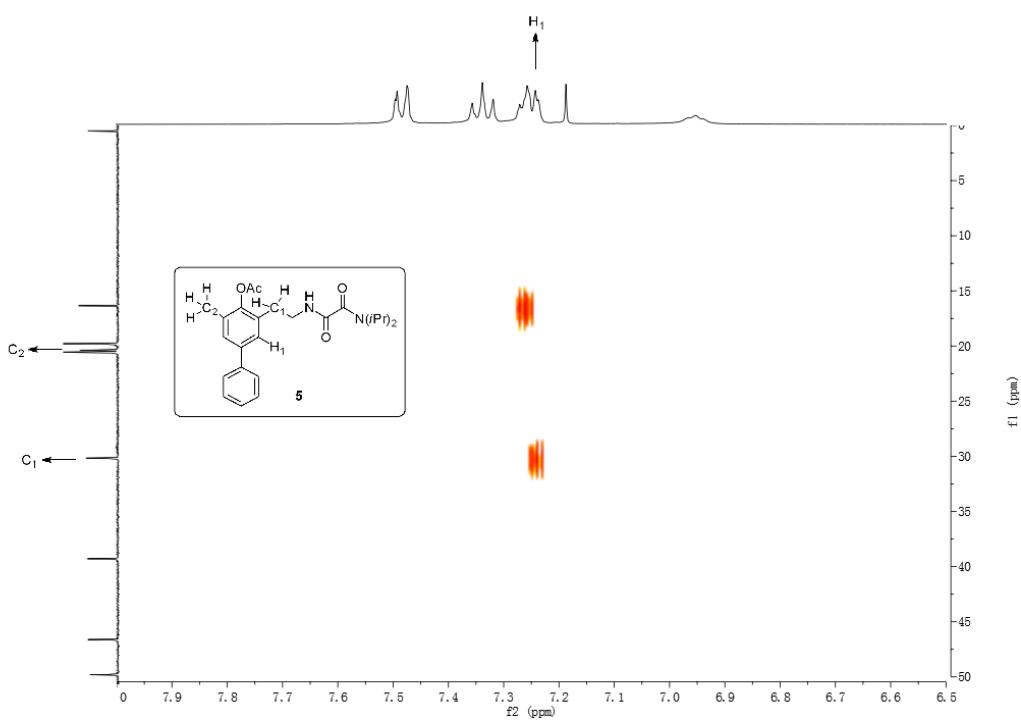
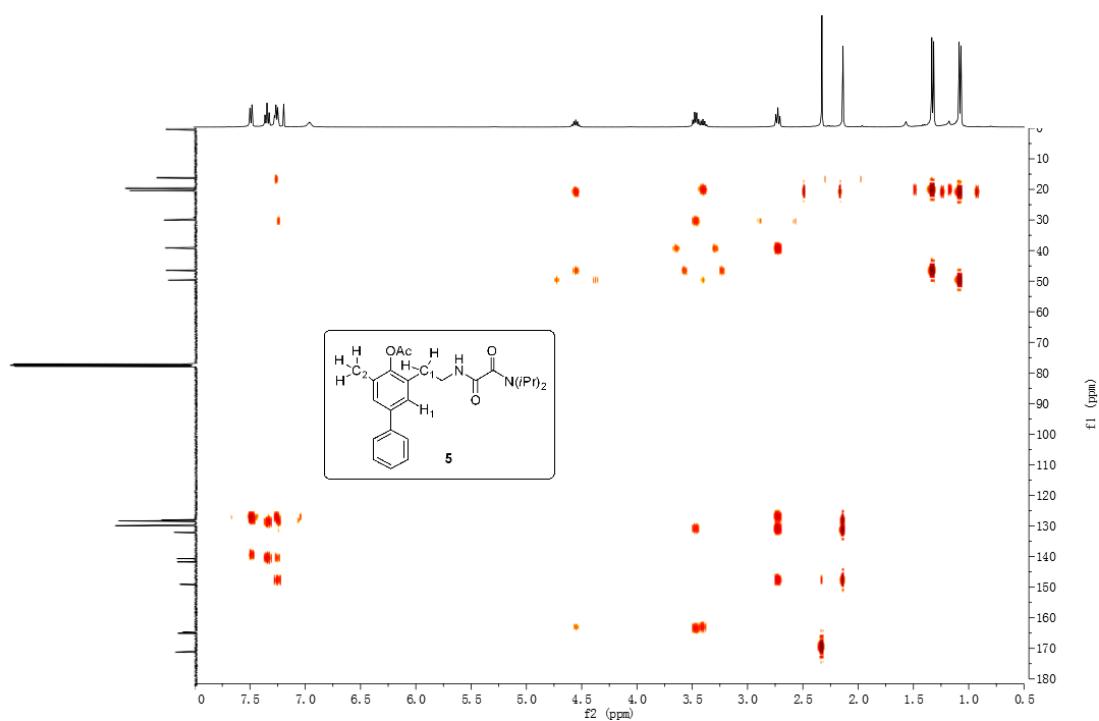
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11. The structure determination of 3a, 3g, 3k, 3m, 4g, 4j and 5 according to HMBC spectrum









12. ^1H and ^{13}C NMR spectra

