

Supporting Information

Visible-Light-Photocatalyzed C5-H Nitration of 8-Aminoquinoline Amides

Pugen Liu, Huijie Qiao, Xiaoxue Su, Peirong Bai and Fan Yang *

Henan Key Laboratory of Chemical Biology and Organic Chemistry, Key Laboratory of Applied Chemistry of Henan Universities, Green Catalysis Center, College of Chemistry, Zhengzhou University, Zhengzhou 450052, China; liupugen@163.com (P.L.)

* Correspondence: yangf@zzu.edu.cn

Table of Contents

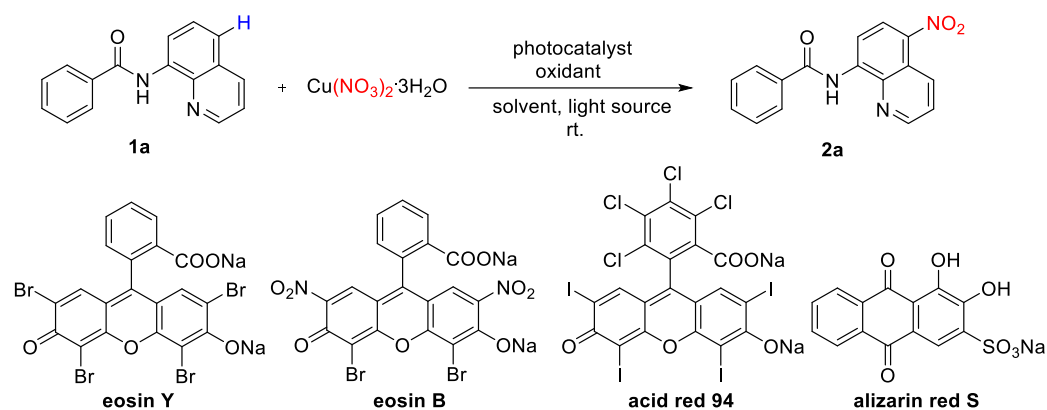
| | |
|--|------------|
| 1. General Information | S2 |
| 2. Optimization of Reaction Conditions | S2 |
| 3. Typical Procedure for Synthesizing the Products | S4 |
| 4. References | S12 |
| 5. Copies of ^1H and ^{13}C NMR Spectra of the Products | S13 |

1. General Information

^1H and ^{13}C NMR spectra were recorded on a Bruker DPX-400 spectrometer with CDCl_3 as the solvent and TMS as an internal standard. Melting points were measured using a WC-1 microscopic apparatus and are uncorrected. High-resolution mass spectra were obtained with MALDI-FTMS. All solvents were used directly without further purification. Dichloromethane, ethyl acetate, and hexane were used for column chromatography. The commercials were obtained from commercial sources and used as-received without further purification unless otherwise noted.

2. Optimization of Reaction Conditions

Table S1. Screening of reaction conditions^a.



| Entry. | Catalyst | Oxidant | Solvent | Yield (%) ^b |
|-----------------|--|--|------------------------|------------------------|
| 1 | EosinB | $\text{K}_2\text{S}_2\text{O}_8$ | CH_3CN | <5 |
| 2 | EosinB | $\text{K}_2\text{S}_2\text{O}_8$ | dioxane | <5 |
| 3 | EosinB | $\text{K}_2\text{S}_2\text{O}_8$ | DMF | <5 |
| 4 | EosinB | $\text{K}_2\text{S}_2\text{O}_8$ | toluene | <5 |
| 5 | EosinB | $\text{K}_2\text{S}_2\text{O}_8$ | acetone | <5 |
| 6 | EosinB | $\text{K}_2\text{S}_2\text{O}_8$ | THF | <5 |
| 7 | EosinB | $\text{K}_2\text{S}_2\text{O}_8$ | DCM | 56 |
| 8 | EosinB | $\text{K}_2\text{S}_2\text{O}_8$ | CHCl_3 | 49 |
| 9 | EosinB | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 68 |
| 10 | EosinB | $\text{K}_2\text{S}_2\text{O}_8$ | DMSO | <5 |
| 11 | EosinB | $\text{Na}_2\text{S}_2\text{O}_8$ | DCE | 61 |
| 12 | EosinB | $(\text{NH}_4)_2\text{S}_2\text{O}_8$ | DCE | 58 |
| 13 | EosinB | $\text{PhI}(\text{OAc})_2$ | DCE | 36 |
| 14 | EosinB | TBHP | DCE | <5 |
| 15 | EosinY | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 65 |
| 16 | $\text{Ru}(\text{bpy})_3\text{Cl}_2 \cdot 6\text{H}_2\text{O}$ | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 76 |
| 17 | $\text{Ru}(\text{bpy})_3(\text{PF}_6)_2$ | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 69 |
| 18 | $\text{Ru}(\text{bpy})_3\text{Cl}_2$ | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 75 |
| 19 | Alizarin Red S | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 72 |
| 20 | Acid Red 94 | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 82 |
| 21 ^c | Acid Red 94 | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 76 |
| 22 ^d | Acid Red 94 | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 81 |
| 23 ^e | Acid Red 94 | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 72 |
| 24 ^f | Acid Red 94 | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 78 |
| 25 ^g | Acid Red 94 | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 69 |
| 26 ^h | Acid Red 94 | $\text{K}_2\text{S}_2\text{O}_8$ | DCE | 76 |

| | | | | |
|-----------------|-------------|--|-----|----|
| 27 ⁱ | Acid Red 94 | K ₂ S ₂ O ₈ | DCE | 68 |
| 28 ^j | Acid Red 94 | K ₂ S ₂ O ₈ | DCE | 73 |
| 29 ^k | Acid Red 94 | K ₂ S ₂ O ₈ | DCE | 66 |
| 30 ^l | Acid Red 94 | K ₂ S ₂ O ₈ | DCE | 62 |
| 31 ^m | Acid Red 94 | K ₂ S ₂ O ₈ | DCE | 43 |
| 32 | — | K ₂ S ₂ O ₈ | DCE | 25 |
| 33 ^o | Acid Red 94 | K ₂ S ₂ O ₈ | DCE | 23 |

^areaction conditions: 1a (0.20 mmol), Cu(NO₃)₂·3H₂O (1.5 equiv), catalyst (5 mol %), oxidant (2.5 equiv), solvent (1.5 mL), and under household light at room temperature in air for 10 h; ^bisolated yield of **2a**; ^cK₂S₂O₈ (2 equiv); ^dK₂S₂O₈ (3 equiv); ^eCu(NO₃)₂·3H₂O (1 equiv); ^fCu(NO₃)₂·3H₂O (2 equiv); ^gcatalyst (4 mol %); ^hunder N₂; ⁱunder O₂; ^jfor 8 h; ^kblue LED; ^lgreen LED; ^mred LED; and ^oin the dark.

3. Typical Procedure for Synthesizing the Products

(2a–2p, 2a'–2m'): To a 10 mL reaction tube, a mixture of amide (0.2 mmol), Cu(NO₃)₂·3H₂O (0.3 mmol, 1.5 equiv), Acid Red 94 (0.01 mmol, 5 mol %), and K₂S₂O₈ (0.5 mmol, 2.5 equiv) was added in DCE (1.5 mL). The resulting mixture was stirred under the irradiation of 26 W household light under air at room temperature for 10 h. Upon completion, the mixture was filtered through a celite pad and washed with CH₂Cl₂, the solvent was removed under reduced pressure and then followed by recrystallization to produce the corresponding product using hexane–CH₂Cl₂ as a solvent.

(2q, 2r, and 2n'): To a 10 mL reaction tube, a mixture of amide (0.2 mmol), Cu(NO₃)₂·3H₂O (0.3 mmol, 1.5 equiv), Acid Red 94 (0.01 mmol, 5 mol %), K₂S₂O₈ (0.5 mmol, 2.5 equiv), and DCE (1.5 mL) was added. Upon completion, the solvent was removed under reduced pressure, and the residue was purified by silica gel column chromatography to produce the corresponding product.

Spectral Data of Product 2 *N*-(5-nitroquinolin-8-yl)benzamide (**2a**) [1]. Yellow solid in 82% yield; mp 215–216 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.07 (s, 1H), 9.30 (d, *J* = 8.6 Hz, 1H), 9.00 (d, *J* = 8.8 Hz, 1H), 8.95 (d, *J* = 3.3 Hz, 1H), 8.59 (d, *J* = 8.8 Hz, 1H), 8.08 (d, *J* = 7.28 Hz, 2H), 7.75 (dd, *J* = 8.8, 4.1 Hz, 1H), 7.66–7.63 (m, 1H), 7.60–7.56 (m, 2H); ¹³C NMR (100 MHz, CDCl₃) δ 165.7, 149.1, 140.9, 138.7, 137.8, 134.1, 133.4, 132.7, 129.0, 127.9, 127.5, 124.7, 121.8, 113.7.

2-methyl-N-(5-nitroquinolin-8-yl)benzamide (**2b**) [1]. Yellow solid in 79% yield; mp 192–193 °C; ¹H NMR (400 MHz, CDCl₃) δ 10.58 (s, 1H), 9.30 (d, *J* = 8.8 Hz, 1H), 9.01 (d, *J* = 8.8 Hz, 1H), 8.88 (d, *J* = 2.8 Hz, 1H), 8.61 (d, *J* = 8.8 Hz, 1H), 7.74–7.69 (m, 2H), 7.74–7.69 (m, 1H), 7.38–7.34 (m, 2H), 2.62 (s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 168.3, 149.1, 141.0, 138.8, 137.6, 137.3, 135.5, 133.4, 131.7, 131.1, 127.9, 127.3, 126.2, 124.7, 121.8, 113.7, 20.3.

3-methyl-N-(5-nitroquinolin-8-yl)benzamide (**2c**). Yellow solid in 82% yield; mp 201–202 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.04 (s, 1H), 9.30 (d, *J* = 8.5 Hz, 1H), 9.01 (d, *J* = 8.9 Hz, 1H), 8.96 (d, *J* = 2.8 Hz, 1H), 8.60 (d, *J* = 8.8 Hz, 1H), 7.88 (d, *J* = 11.9 Hz, 2H), 7.75 (dd, *J* = 9.1, 4.1 Hz, 1H), 7.49–7.45 (m, 2H), 2.49 (s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 165.94, 149.1, 141.0, 139.0, 138.6, 137.8, 134.2, 133.4, 128.9, 128.2, 127.9, 124.7, 124.4, 121.8, 113.7, 21.5; HRMS (ESI): calculated for C₁₇H₁₃N₃O₃: [M+H]⁺ requires 308.1030; found 308.1031.

4-methyl-N-(5-nitroquinolin-8-yl)benzamide (**2d**) [2]. Yellow solid in 80% yield; mp 210–211 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.05 (s, 1H), 9.31 (d, *J* = 8.7 Hz, 1H), 9.02 (d, *J* = 8.8 Hz, 1H), 8.96 (d, *J* = 3.4 Hz, 1H), 8.61 (d, *J* = 8.8 Hz, 1H), 7.99 (d, *J* = 7.8 Hz, 2H), 7.74 (dd, *J* = 8.9, 4.1 Hz, 1H), 7.38 (d, *J* = 7.9 Hz, 2H), 2.48 (s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 165.7, 149.0, 143.4, 141.0, 138.6, 137.8, 133.5, 131.4, 129.7, 128.0, 127.5, 124.7, 121.9, 113.7, 21.6; HRMS (ESI): calculated for C₁₇H₁₃N₃O₃: [M+H]⁺ requires 308.1030; found 308.1033.

2-methoxy-N-(5-nitroquinolin-8-yl)benzamide (**2e**). Yellow solid in 75% yield; mp 208–209 °C; ¹H NMR (400 MHz, CDCl₃) δ 12.68 (s, 1H), 9.26 (d, *J* = 7.9 Hz, 1H), 9.06 (d, *J* = 8.9 Hz, 1H), 8.93 (d, *J* = 2.8 Hz, 1H), 8.56 (d, *J* = 8.8 Hz, 1H), 8.31 (d, *J* = 6.7 Hz, 1H), 7.69 (dd, *J* = 8.8, 4.1 Hz, 1H), 7.57–7.54 (m, 1H), 7.17–7.14 (m, 1H), 7.08 (d, *J* = 8.2 Hz, 1H), 4.20 (s, 3H); ¹³C

NMR (100 MHz, CDCl₃) δ 164.2, 157.9, 148.9, 142.2, 138.4, 138.2, 134.0, 133.1, 132.6, 128.1, 124.4, 121.9, 121.5, 121.4, 114.4, 111.7, 56.2; HRMS (ESI): calculated for C₁₇H₁₃N₃O₄: [M+H]⁺ requires 324.0979; found 324.0977.

4-methoxy-N-(5-nitroquinolin-8-yl)benzamide (2f) [3]. Yellow solid in 78% yield; mp 260-261 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.03 (s, 1H), 9.34 (d, *J* = 8.6 Hz, 1H), 9.01 (d, *J* = 8.9 Hz, 1H), 8.96 (d, *J* = 3.6 Hz, 1H), 8.63 (d, *J* = 8.9 Hz, 1H), 8.07 (d, *J* = 8.7 Hz, 2H), 7.76 (dd, *J* = 8.9, 4.1 Hz, 1H), 7.07 (d, *J* = 8.6 Hz, 2H), 3.92 (s, 3H); ¹³C NMR (100 MHz, CDCl₃) δ 165.2, 163.2, 149.0, 141.2, 138.5, 137.8, 133.5, 129.5, 128.1, 126.4, 124.7, 121.9, 114.3, 113.5, 55.6.

3,5-dimethoxy-N-(5-nitroquinolin-8-yl)benzamide (2g). Yellow solid in 78% yield; mp 200-201 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.00 (s, 1H), 9.30 (d, *J* = 8.8 Hz, 1H), 8.99-8.95 (m, 2H), 8.61 (d, *J* = 8.8 Hz, 1H), 7.76 (dd, *J* = 8.9, 4.1 Hz, 1H), 7.20 (s, 2H), 6.71 (s, 2H), 3.91 (s, 6H); ¹³C NMR (100 MHz, CDCl₃) δ 165.6, 161.2, 149.2, 140.8, 138.8, 137.8, 136.3, 133.4, 127.9, 124.8, 121.8, 113.8, 105.5, 104.4, 55.7; HRMS (ESI): calculated for C₁₈H₁₅N₃O₅: [M+H]⁺ requires 354.1084; found 354.1087.

3-fluoro-N-(5-nitroquinolin-8-yl)benzamide (2h). Yellow solid in 73% yield; mp 236-237 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.06 (s, 1H), 9.31 (d, *J* = 8.7 Hz, 1H), 9.01-8.98 (m, 2H), 8.61 (d, *J* = 8.8 Hz, 1H), 7.88 (d, *J* = 7.6 Hz, 1H), 7.80-7.76 (m, 2H), 7.60-7.55 (m, 1H), 7.36-7.33 (m, 1H); ¹³C NMR (100 MHz, CDCl₃) δ 164.4, 163.0, 149.2, 140.5, 139.0, 137.8, 136.4, 133.5, 130.8, 127.8, 124.8, 122.9, 121.8, 119.7, 114.9, 113.9; HRMS (ESI): calculated for C₁₆H₁₀FN₃O₃: [M+H]⁺ requires 312.0779; found 312.0781.

4-fluoro-N-(5-nitroquinolin-8-yl)benzamide (2i). Yellow solid in 75% yield; mp 218-219 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.03 (s, 1H), 9.31 (d, *J* = 8.6 Hz, 1H), 9.01-8.96 (m, 2H), 8.60 (d, *J* = 8.8 Hz, 1H), 8.13-8.09 (m, 2H), 7.77 (dd, *J* = 9.2, 4.1 Hz, 1H), 7.29-7.24 (m, 2H); ¹³C NMR (100 MHz, CDCl₃) δ 165.5, 164.6, 149.1, 140.7, 138.8, 137.7, 133.5, 130.4, 130.0, 127.9, 124.8, 121.8, 116.2, 113.7; HRMS (ESI): calculated for C₁₆H₁₀FN₃O₃: [M+H]⁺ requires 312.0779; found 312.0778.

3-chloro-N-(5-nitroquinolin-8-yl)benzamide (2j). Yellow solid in 74% yield; mp 234-235 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.04 (s, 1H), 9.31 (d, *J* = 8.7 Hz, 1H), 8.99 (d, *J* = 8.8 Hz, 2H), 8.62 (d, *J* = 8.8 Hz, 1H), 8.07 (s, 1H), 7.96 (d, *J* = 7.5 Hz, 1H), 7.78 (dd, *J* = 9.1, 4.1 Hz, 1H), 7.62 (d, *J* = 7.8 Hz, 1H), 7.52 (t, *J* = 7.8 Hz, 1H); ¹³C NMR (100 MHz, CDCl₃) δ 164.4, 149.2, 140.5, 139.0, 137.8, 135.9, 135.3, 133.5, 132.7, 130.3, 127.9, 127.8, 125.4, 124.8, 121.8, 114.0; HRMS (ESI): calculated for C₁₆H₁₀ClN₃O₃: [M+H]⁺ requires 328.0483; found 328.0483.

4-chloro-N-(5-nitroquinolin-8-yl)benzamide (2k) [2]. Yellow solid in 76% yield; mp 210-212 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.05 (s, 1H), 9.32 (d, *J* = 8.1 Hz, 1H), 9.01-8.97 (m, 2H), 8.62 (d, *J* = 8.8 Hz, 1H), 8.04 (d, *J* = 8.5 Hz, 2H), 7.77 (dd, *J* = 9.1, 4.1 Hz, 1H), 7.56 (d, *J* = 8.4 Hz, 2H); ¹³C NMR (100 MHz, CDCl₃) δ 164.7, 149.2, 140.6, 139.1, 138.9, 137.8, 133.5, 132.5, 129.4, 128.9, 127.8, 124.8, 121.8, 113.9; HRMS (ESI): calculated for C₁₆H₁₀ClN₃O₃: [M+H]⁺ requires 328.0483; found 328.0484.

3-bromo-N-(5-nitroquinolin-8-yl)benzamide (2l). Yellow solid in 78% yield; mp 213-214 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.00 (s, 1H), 9.36 (d, *J* = 8.8 Hz, 1H), 8.96 (d, *J* = 8.9 Hz, 2H), 8.61 (d, *J* = 8.8 Hz, 1H), 8.23 (s, 1H), 8.01 (d, *J* = 7.7 Hz, 1H), 7.78-7.75 (m, 2H), 7.49 (t, *J* = 7.8 Hz, 1H); ¹³C NMR (100 MHz, CDCl₃) δ 164.2, 149.2, 140.4, 139.0, 137.8, 136.1, 135.6, 133.5, 130.8, 130.5, 127.8, 125.9, 124.8, 123.3, 121.8, 114.0; HRMS (ESI): calculated for C₁₆H₁₀BrN₃O₃: [M+H]⁺ requires 371.9978; found 371.9979.

4-bromo-N-(5-nitroquinolin-8-yl)benzamide (2m). Yellow solid in 72% yield; mp 279-280 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.06 (s, 1H), 9.32 (d, *J* = 8.2 Hz, 1H), 9.01-8.97 (m, 2H), 8.62 (d, *J* = 8.9 Hz, 1H), 7.96 (d, *J* = 8.3 Hz, 2H), 7.78 (dd, *J* = 8.9, 4.3 Hz, 1H), 7.72 (d, *J* = 8.5 Hz, 2H); ¹³C NMR (100 MHz, CDCl₃) δ 164.8, 149.2, 140.6, 138.9, 137.8, 133.5, 133.0, 132.3, 129.0, 127.8, 127.6, 124.8, 121.8, 113.9; HRMS (ESI): calculated for C₁₆H₁₀BrN₃O₃: [M+H]⁺ requires 371.9978; found 371.9977.

N-(5-nitroquinolin-8-yl)-3-(trifluoromethyl)benzamide (2n). Yellow solid in 75% yield; mp 181-182 °C; ¹H NMR (400 MHz, CDCl₃) δ 11.08 (s, 1H), 9.30 (d, *J* = 8.6 Hz, 1H), 8.99-8.97 (m, 2H), 8.61 (d, *J* = 8.8 Hz, 1H), 8.36 (s, 1H), 8.25 (d, *J* = 7.6 Hz, 1H), 7.90 (d, *J* = 7.6 Hz, 1H), 7.79-7.72 (m, 2H); ¹³C NMR (100 MHz, CDCl₃) δ 164.2, 149.3, 140.3, 139.1, 137.7, 135.0,

133.7, 133.5, 130.4, 129.7, 129.2, 127.7, 124.9, 124.7, 122.1, 114.0; HRMS (ESI): calculated for $C_{17}H_{10}F_3N_3O_3$: $[M+H]^+$ requires 362.0747; found 362.0748.

2-chloro-4-methoxy-N-(5-nitroquinolin-8-yl)benzamide (2p). Yellow solid in 65% yield; mp 262-263 °C; 1H NMR (400 MHz, $CDCl_3$) δ 10.99 (s, 1H), 9.33 (d, J = 8.8 Hz, 1H), 8.99-8.97 (m, 2H), 8.62 (d, J = 8.8 Hz, 1H), 8.14 (d, J = 1.2 Hz, 1H), 8.02 (d, J = 8.6 Hz, 1H), 7.77 (dd, J = 8.9, 4.1 Hz, 1H), 7.10 (d, J = 8.5 Hz, 1H), 4.03 (s, 3H); ^{13}C NMR (100 MHz, $CDCl_3$) δ 164.1, 158.5, 149.1, 140.8, 138.7, 137.8, 133.5, 129.7, 127.9, 127.7, 127.2, 124.8, 123.3, 121.9, 113.7, 111.8, 56.5; HRMS (ESI): calculated for $C_{17}H_{12}ClN_3O_4$: $[M+H]^+$ requires 358.0589; found 358.0591.

N-(5-nitroquinolin-8-yl)thiophene-2-carboxamide (2q) [3]. Yellow solid in 69% yield; mp 220-221 °C; 1H NMR (400 MHz, $CDCl_3$) δ 10.90 (s, 1H), 9.30 (d, J = 8.8 Hz, 1H), 8.96-8.95 (m, 1H), 8.91 (d, J = 8.8 Hz, 1H), 8.59 (d, J = 8.8 Hz, 1H), 7.88 (d, J = 3.1 Hz, 1H), 7.76 (dd, J = 8.9, 4.1 Hz, 1H), 7.65 (d, J = 4.8 Hz, 1H), 7.24-7.21 (m, 1H); ^{13}C NMR (100 MHz, $CDCl_3$) δ 160.2, 149.1, 140.6, 139.0, 138.7, 137.5, 133.5, 132.2, 129.4, 128.2, 127.9, 124.8, 121.9, 113.7.

*N-(5-nitroquinolin-8-yl)pivalamide (2r)*². Yellow solid in 73% yield; mp 172-173 °C; 1H NMR (400 MHz, $CDCl_3$) δ 10.59 (s, 1H), 9.27-9.25 (m, 1H), 8.92 (d, J = 3.0 Hz, 1H), 8.84 (d, J = 8.8 Hz, 1H), 8.54 (d, J = 8.8 Hz, 1H), 7.71 (dd, J = 8.8, 4.1 Hz, 1H), 1.45 (s, 9H); ^{13}C NMR (100 MHz, $CDCl_3$) δ 177.8, 149.0, 141.0, 138.3, 137.7, 133.3, 127.9, 124.6, 121.7, 113.4, 40.7, 27.6.

N-(2-methyl-5-nitroquinolin-8-yl)benzamide (2a') [2]. Yellow solid in 75% yield; mp 202-203 °C; 1H NMR (400 MHz, $CDCl_3$) δ 11.08 (s, 1H), 9.12 (d, J = 8.9 Hz, 1H), 8.91 (d, J = 8.8, 1H), 8.49 (d, J = 8.8 Hz, 1H), 8.05 (d, J = 7.3 Hz, 2H), 7.66-7.57 (m, 4H), 2.81 (s, 3H); ^{13}C NMR (100 MHz, $CDCl_3$) δ 165.5, 158.5, 140.1, 138.7, 137.3, 134.3, 133.3, 132.6, 129.1, 127.4, 126.7, 125.6, 119.9, 113.7, 25.2; HRMS (ESI): calculated for $C_{17}H_{13}N_3O_3$: $[M+H]^+$ requires 308.1030; found 308.1035.

2-methyl-N-(2-methyl-5-nitroquinolin-8-yl)benzamide (2b'). Yellow solid in 75% yield; mp 209-210 °C; 1H NMR (400 MHz, $CDCl_3$) δ 10.62 (s, 1H), 9.15 (d, J = 8.9 Hz, 1H), 8.96 (d, J = 8.8, 1H), 8.51 (d, J = 8.8 Hz, 1H), 7.72 (d, J = 7.4 Hz, 1H), 7.58 (d, J = 8.9 Hz, 1H), 7.48-7.44 (m, 1H), 7.39-7.35 (m, 2H), 2.75 (s, 3H), 2.64 (s, 3H); ^{13}C NMR (100 MHz, $CDCl_3$) δ 168.2, 158.6, 140.3, 138.9, 137.2, 137.2, 135.6, 133.3, 131.8, 131.1, 127.5, 126.7, 126.3, 125.6, 120.0, 113.7, 25.1, 20.4; HRMS (ESI): calculated for $C_{18}H_{15}N_3O_3$: $[M+H]^+$ requires 322.1186; found 322.1190.

3-methyl-N-(2-methyl-5-nitroquinolin-8-yl)benzamide (2c'). Yellow solid in 79% yield; mp 184-185 °C; 1H NMR (400 MHz, $CDCl_3$) δ 11.04 (s, 1H), 9.15 (d, J = 8.9 Hz, 1H), 8.93 (d, J = 8.8 Hz, 1H), 8.51 (d, J = 8.8 Hz, 1H), 7.91 (s, 1H), 7.81 (d, J = 6.9 Hz, 1H), 7.59 (d, J = 8.9 Hz, 1H), 7.49-7.43 (m, 2H), 2.83 (s, 3H), 2.47 (s, 3H); ^{13}C NMR (100 MHz, $CDCl_3$) δ 165.7, 158.4, 140.2, 139.0, 138.7, 137.3, 134.3, 133.4, 133.3, 128.9, 128.3, 126.8, 125.6, 124.2, 120.0, 113.7, 25.2, 21.5; HRMS (ESI): calculated for $C_{18}H_{15}N_3O_3$: $[M+H]^+$ requires 322.1186; found 322.1190.

4-methyl-N-(2-methyl-5-nitroquinolin-8-yl)benzamide (2d'). Yellow solid in 78% yield; mp 206-207 °C; 1H NMR (400 MHz, $CDCl_3$) δ 10.99 (s, 1H), 9.09 (d, J = 8.9 Hz, 1H), 8.86 (d, J = 8.8 Hz, 1H), 8.44 (d, J = 8.8 Hz, 1H), 7.92 (d, J = 7.9 Hz, 2H), 7.55 (d, J = 8.9 Hz, 1H), 7.37 (d, J = 7.8 Hz, 2H), 2.81 (s, 3H), 2.45 (s, 3H); ^{13}C NMR (100 MHz, $CDCl_3$) δ 165.4, 158.4, 143.3, 140.2, 138.5, 137.2, 133.2, 131.3, 129.7, 127.4, 126.8, 125.6, 120.0, 113.5, 25.2, 21.6; HRMS (ESI): calculated for $C_{18}H_{15}N_3O_3$: $[M+H]^+$ requires 322.1186; found 322.1189.

2-methoxy-N-(2-methyl-5-nitroquinolin-8-yl)benzamide (2e'). Yellow solid in 75% yield; mp 258-259 °C; 1H NMR (400 MHz, $CDCl_3$) δ 12.46 (s, 1H), 9.17 (d, J = 8.9 Hz, 1H), 9.10 (d, J = 8.9 Hz, 1H), 8.51 (d, J = 8.9 Hz, 1H), 8.33 (d, J = 6.6 Hz, 1H), 7.60-7.55 (m, 2H), 7.18 (t, J = 7.6 Hz, 1H), 7.11 (d, J = 8.3 Hz, 1H), 4.23 (s, 3H), 2.86 (s, 3H); ^{13}C NMR (100 MHz, $CDCl_3$) δ 164.3, 157.9, 157.8, 141.6, 137.8, 134.0, 133.2, 132.8, 127.0, 125.3, 121.7, 121.6, 120.1, 114.7, 111.6, 56.2, 25.3; HRMS (ESI): calculated for $C_{18}H_{15}N_3O_4$: $[M+H]^+$ requires 338.1135; found 338.1139.

3-fluoro-N-(2-methyl-5-nitroquinolin-8-yl)benzamide (2f'). Yellow solid in 71% yield; mp 251-252 °C; 1H NMR (400 MHz, $CDCl_3$) δ 11.10 (s, 1H), 9.17 (d, J = 8.9 Hz, 1H), 8.94 (d, J =

8.8 Hz, 1H), 8.53 (d, J = 8.8 Hz, 1H), 7.85 (d, J = 7.7 Hz, 1H), 7.78 (d, J = 9.2 Hz, 1H), 7.63-7.55 (m, 2H), 7.36-7.32 (m, 1H), 2.84 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 164.3, 161.8, 158.7, 139.8, 139.1, 137.4, 136.6, 133.5, 130.8, 126.7, 125.7, 122.8, 120.0, 119.6, 114.9, 113.9, 25.3; HRMS (ESI): calculated for $\text{C}_{17}\text{H}_{12}\text{FN}_3\text{O}_3$: $[\text{M}+\text{H}]^+$ requires 326.0935; found 326.0938.

4-fluoro-N-(2-methyl-5-nitroquinolin-8-yl)benzamide (2g'). Yellow solid in 66% yield; mp 218-219 °C; ^1H NMR (400 MHz, CDCl_3) δ 11.07 (s, 1H), 9.16 (d, J = 8.9 Hz, 1H), 8.93 (d, J = 8.8, 1H), 8.52 (d, J = 8.8 Hz, 1H), 8.11-8.08 (m, 2H), 7.62 (d, J = 8.9 Hz, 1H), 7.29-7.25 (m, 2H), 2.83 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 166.7, 164.3, 158.5, 140.0, 138.9, 137.3, 133.5, 130.5, 129.9, 126.7, 125.7, 120.0, 116.2, 113.8, 25.3; HRMS (ESI): calculated for $\text{C}_{17}\text{H}_{12}\text{FN}_3\text{O}_3$: $[\text{M}+\text{H}]^+$ requires 326.0935; found 326.0938.

3-chloro-N-(2-methyl-5-nitroquinolin-8-yl)benzamide (2h'). Yellow solid in 67% yield; mp 215-216 °C; ^1H NMR (400 MHz, CDCl_3) δ 11.08 (s, 1H), 9.15 (d, J = 8.9 Hz, 1H), 8.91 (d, J = 8.8 Hz, 1H), 8.51 (d, J = 8.8 Hz, 1H), 8.07 (s, 1H), 7.94 (d, J = 7.6 Hz, 1H), 7.62 (d, J = 9.2 Hz, 2H), 7.53 (t, J = 7.8 Hz, 1H), 2.83 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 164.1, 158.7, 139.7, 139.1, 137.3, 136.1, 135.4, 133.4, 132.6, 130.3, 127.9, 126.6, 125.7, 125.2, 120.0, 113.9, 25.3; HRMS (ESI): calculated for $\text{C}_{17}\text{H}_{12}\text{ClN}_3\text{O}_3$: $[\text{M}+\text{H}]^+$ requires 342.0640; found 342.0642.

4-chloro-N-(2-methyl-5-nitroquinolin-8-yl)benzamide (2i') [2]. Yellow solid in 68% yield; mp 221-222 °C; ^1H NMR (400 MHz, CDCl_3) δ 11.08 (s, 1H), 9.16 (d, J = 8.9 Hz, 1H), 8.93 (d, J = 8.8 Hz, 1H), 8.52 (d, J = 8.8 Hz, 1H), 8.02 (d, J = 8.2 Hz, 2H), 7.62 (d, J = 8.9 Hz, 1H), 7.56 (d, J = 8.2 Hz, 2H), 2.84 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 164.5, 158.6, 139.9, 140.0, 139.0, 137.3, 133.5, 132.7, 129.4, 128.8, 126.7, 125.7, 120.0, 113.9, 25.3; HRMS (ESI): calculated for $\text{C}_{17}\text{H}_{12}\text{ClN}_3\text{O}_3$: $[\text{M}+\text{H}]^+$ requires 342.0640; found 342.0641.

3-bromo-N-(2-methyl-5-nitroquinolin-8-yl)benzamide (2j'). Yellow solid in 66% yield; mp 203-204 °C; ^1H NMR (400 MHz, CDCl_3) δ 11.09 (s, 1H), 9.16 (d, J = 8.9 Hz, 1H), 8.91 (d, J = 8.8 Hz, 1H), 8.51 (d, J = 8.8 Hz, 1H), 8.23 (s, 1H), 7.99 (d, J = 7.7 Hz, 1H), 7.76 (d, J = 7.9 Hz, 1H), 7.62 (d, J = 8.9 Hz, 1H), 7.47 (t, J = 7.8 Hz, 1H), 2.88 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 164.0, 158.7, 139.7, 139.1, 137.3, 136.2, 135.5, 133.4, 130.9, 130.5, 126.6, 125.7, 125.7, 123.3, 119.9, 113.9, 25.3; HRMS (ESI): calculated for $\text{C}_{17}\text{H}_{12}\text{BrN}_3\text{O}_3$: $[\text{M}+\text{H}]^+$ requires 386.0135; found 386.0136.

4-bromo-N-(2-methyl-5-nitroquinolin-8-yl)benzamide (2k'). Yellow solid in 71% yield; mp 249-250 °C; ^1H NMR (400 MHz, CDCl_3) δ 11.09 (s, 1H), 9.15 (d, J = 8.9 Hz, 1H), 9.17 (d, J = 9.0, 1H), 8.93 (d, J = 8.8 Hz, 1H), 8.52 (d, J = 8.8 Hz, 1H), 7.94 (d, J = 8.4 Hz, 2H), 7.44 (d, J = 8.4 Hz, 2H), 7.61 (d, J = 8.9 Hz, 1H), 2.83 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 164.7, 158.6, 139.8, 139.0, 137.3, 133.5, 133.2, 132.4, 128.9, 127.5, 126.7, 125.7, 120.0, 113.9, 25.3; HRMS (ESI): calculated for $\text{C}_{17}\text{H}_{12}\text{BrN}_3\text{O}_3$: $[\text{M}+\text{H}]^+$ requires 386.0135; found 386.0136.

N-(2-methyl-5-nitroquinolin-8-yl)-3-(trifluoromethyl)benzamide (2l'). Yellow solid in 56% yield; mp 191-192 °C; ^1H NMR (400 MHz, CDCl_3) δ 11.19 (s, 1H), 9.15 (d, J = 8.9 Hz, 1H), 8.92 (d, J = 8.8 Hz, 1H), 8.53 (d, J = 8.8 Hz, 1H), 8.34 (s, 1H), 8.26 (d, J = 7.7 Hz, 1H), 7.90 (d, J = 7.7 Hz, 1H), 7.74 (t, J = 7.7 Hz, 1H), 7.63 (d, J = 8.9 Hz, 1H), 2.84 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 164.0, 158.7, 139.6, 139.2, 137.3, 135.1, 133.5, 131.7, 130.5, 129.8, 129.1, 126.6, 125.8, 124.5, 123.6, 119.9, 113.9, 25.2; HRMS (ESI): calculated for $\text{C}_{18}\text{H}_{12}\text{F}_3\text{N}_3\text{O}_3$: $[\text{M}+\text{H}]^+$ requires 376.0904; found 376.0907.

N-(2-methyl-5-nitroquinolin-8-yl)-4-(trifluoromethyl)benzamide (2m'). Yellow solid in 59% yield; mp 208-209 °C; ^1H NMR (400 MHz, CDCl_3) δ 11.16 (s, 1H), 9.18 (d, J = 8.9 Hz, 1H), 8.96 (d, J = 8.8 Hz, 1H), 8.54 (d, J = 8.8 Hz, 1H), 8.20 (d, J = 8.0 Hz, 2H), 7.87 (d, J = 8.1 Hz, 2H), 7.64 (d, J = 8.9 Hz, 1H), 2.84 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 164.3, 158.7, 139.6, 139.3, 137.6, 137.4, 134.1, 133.5, 127.9, 126.6, 126.2, 125.8, 123.6, 120.0, 114.1, 25.3; HRMS (ESI): calculated for $\text{C}_{18}\text{H}_{12}\text{F}_3\text{N}_3\text{O}_3$: $[\text{M}+\text{H}]^+$ requires 376.0904; found 376.0905.

N-(2-methyl-5-nitroquinolin-8-yl)thiophene-2-carboxamide (2n'). Yellow solid in 72% yield; mp 257-258 °C; ^1H NMR (400 MHz, CDCl_3) δ 10.98 (s, 1H), 9.16 (d, J = 8.8 Hz, 1H), 8.85 (d, J = 8.8 Hz, 1H), 8.51 (d, J = 8.7 Hz, 1H), 7.85 (d, J = 2.4 Hz, 1H), 7.67 (d, J = 4.5 Hz, 1H), 7.61 (d, J = 8.9 Hz, 1H), 7.23 (d, J = 3.9 Hz, 1H), 2.84 (s, 3H); ^{13}C NMR (100 MHz, CDCl_3) δ 160.1, 158.5, 139.9, 139.2, 138.8, 137.1, 133.4, 132.0, 129.3, 128.2, 126.8, 125.7, 120.0, 113.7, 25.2; HRMS (ESI): calculated for $\text{C}_{15}\text{H}_{11}\text{N}_3\text{O}_3\text{S}$: $[\text{M}+\text{H}]^+$ requires 314.0594; found 314.0596.

References

1. Zhu, X.; Qiao, L.; Ye, P.; Ying, B.; Xu, J.; Shen, C.; Zhang, P. Copper-catalyzed rapid C–H nitration of 8-aminoquinolines by using sodium nitrite as the nitro source under mild conditions. *RSC Adv.* **2016**, *6*, 89979–89983.
2. He, Y.; Zhao, N.-N.; Qiu, L.-Q.; Zhang, X.-Y.; Fan, X.-S. Regio- and chemoselective mono- and bisnitration of 8-aminoquinoline amides with Fe (NO₃)₃·9H₂O as promoter and nitro source. *Organic letters. Org. Lett.* **2016**, *18*, 6054–6057.
3. Whiteoak, C.J.; Planas, O.; Company, A.; Ribas, X. A First Example of Cobalt-Catalyzed Remote C–H Functionalization of 8-Aminoquinolines Operating through a Single Electron Transfer Mechanism. *Adv. Synth. Catal.* **2016**, *358*, 1679–1688.

4. Copies of ^1H and ^{13}C NMR Spectra of the Products

