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## **New Journal of Chemistry**

### **Supplementary Information**

# Mn-mediated Oxidative Radical Cyclization of 2-(azidomethyl)phenyl isocyanides with Carbazate: Access to Quinazoline-2-Carboxylate

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#### 1. Experimental Information.

## A. Genaral Experimental Procedure to Prepare N-formyl Derivatives <sup>16</sup>

To the stirred solution of (2-aminophenyl)methanol (1.0 g, 6.53 mmol) in dry tetrahydrofurone (10 mL) was added ethyl format (4 mL, 52.28 mmol, 8.0 eq) at room temperature, and the mixture was stirred for 5 min. NaH (0.65 g, 16.33 mmol, 2.5 eq) was added in portions at same temperature. After 1 h, the resulting mixture was slowly poured in to ice water. The aqueous phase was extracted with ethyl acetate (3 x 20 mL), the combined organic layers were washed with brine, and dried over anhydrous Na<sub>2</sub>SO<sub>4</sub>, and evaporated under reduced pressure. The crude product could not be isolated as pure product due to the same Rf of starting material and products in most of the common solvent combinations. Hence, the impure products was used for next step without further purification.

## **B.** Genaral Experimental Procedure to Prepare O-methanesulfonate Derivatives <sup>17</sup>

To the stirred solution of N-(2-(hydroxymethyl)phenyl)formamide (1.1 g, 6.508 mmol) in dichloromethane (30 mL) at 0 °C was added triethylamine (1.87 mL, 13.01 mmol, 2 eq), and methane sulphonyl chloride (0.75 mL, 9.761 mmol, 1.5 eq) was added drop wise at same temperature. The resulting solution was stirred for 2 h at room temperature. After the completion of reaction (monitored by TLC). The reaction mixture was quenched with cold water. The aqueous phase was extracted with dichloromethane (3 x 20 mL), the combined organic layers were washed with brine, and dried over anhydrous Na<sub>2</sub>SO<sub>4</sub>, and evaporated under reduced pressure. The crude products could not be isolated as pure product due to the same Rf of starting material and products in most of the common solvent combinations. Hence, the impure products was used for next step without further purification and <sup>13</sup>C NMR were not recorded.

## C. Genaral Experimental Procedure to Prepare Azide Derivatives <sup>17</sup>

To the stirred solution of 2-formamido-5-methoxybenzyl methanesulfonate (1.0 g, 3.862 mmol) in DMF (10 mL) at 0 °C was added sodium azide (0.49 g, 7.722 mmol, 2 eq). The resulting solution was stirred for 2 h at room temperature. After the completion of reaction (monitored by TLC).

The reaction mixture was quenched with cold water. The aqueous phase was extracted with ethyl acetate (3 x 20 mL), the combined organic layers were washed with brine, and dried over anhydrous  $Na_2SO_4$ , and evaporated under reduced pressure. The crude was purified by flash column chromatography on silica gel using ethyl acetate: n-hexane (40: 60) as an eluent. The products were not very pure, the impure products was taken for next step.

#### **D.** Genaral Experimental Procedure to Prepare Isocyanide Derivatives <sup>11, 16</sup>

To the stirred solution of N-(2-(azidomethyl)phenyl)formamide (0.7 g, 3.97 mmol) in tetrhydrofuron (10 mL) at 0 °C was added triethyl amine (1.72 mL, 11.93 mmol, 3 eq), the mixture was stirred for 5 min. phosphorus oxychloride (POCl<sub>3</sub>) (0.58 mL, 5.96 mmol, 1.5 eq) was added drop wise at same temperature. The resulting solution was stirred for 2 h at room temperature. After the completion of reaction (monitored by TLC). The mixture was quenched with saturated NaHCO<sub>3</sub> solution. The aqueous phase was extracted with ethyl acetate (3 x 20 mL), the combined organic layers were washed with brine, and dried over anhydrous Na<sub>2</sub>SO<sub>4</sub>, and evaporated under reduced pressure. The crude was purified by flash column chromatography on silica gel using ethyl acetate: n-hexane (10: 90) as an eluent

#### 2. Crystal structural data for compound 3g (CCDC 1911289)

Bond precision: C-C = 0.0030 A Wavelength: 0.71073 Cell: a = 10.8098 (10) b = 3.9798 (4) c = 24.25 (2)alpha= 90 beta= 92.803 (9) gamma= 90 Temperature: 296

	Calculated
Volume	1041.77(17)
Space group	P 21/n
Hall group	-P 2yn
Moiety formula	C11 H9 F N2 O3

Sum formula	C11 H9 F N2 O3
Mr	236.20
Dx, g cm-3	1.506
Ζ	4
Mu (mm-1)	0.123
F000	488.0
F000'	488.30
h, k, 1max	14, 5, 31
Nref	2380

Data completeness= 0.997Theta (max)= 27.482R (reflections) = 0.0597 (1575)wR2 (reflections) = 0.1449 (2374)S = 1.070 Npar = 154

ORTEP diagram for compound 3g (50% ellipsoid probability)



#### 3. Reaction Schemes for Preparation of Substrates and Products

Scheme 1. General Synthetic route for the Preparation of Starting Materials



#### 4. Characterization data for all isolated products

**N-(2-(hydroxymethyl)-4-methoxyphenyl) formamide (b2):** Following the general procedure A. Pale-yellow solid (1.1 g crude)  $R_f$  (70% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.32 (s, 1H), 8.25 - 8.20 (m, 1H), 7.50 (d, *J* = 8.8 Hz, 1H), 6.99 - 6.96 (m, 1H), 6.78 - 6.76 (m, 1H), 5.23 (s, 1H), 4.46 - 4.42 (m, 2H), 3.71 (s, 3H). LCMS (ESI) m/z calcd for C<sub>9</sub>H<sub>11</sub>NO<sub>3</sub> [M + H]<sup>+</sup> 181.07 found 182.1

**N-(2-(hydroxymethyl)-3-methoxyphenyl) formamide (c2):** Following the general procedure A. Pale-yellow solid (1.02 g crude).  $R_f$  (70% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.32 (s, 1H), 8.25 - 8.20 (m, 1H), 7.50 (d, J = 8.8 Hz, 1H), 6.99 - 6.96 (m, 1H), 6.78 - 6.76 (m, 1H), 5.23 (s, 1H), 4.46 - 4.42 (m, 2H), 3.71 (s, 3H). LCMS (ESI) m/z calcd for C<sub>9</sub>H<sub>11</sub>NO<sub>3</sub> [M + H]<sup>+</sup> 181.07 found 182.1

**N-(2-(hydroxymethyl)-6-methylphenyl) formamide (d2):** Following the general procedure A. Yellow solid (1.1 g crude).  $R_f$  (70% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.35 (s, 1H), 8.21 (s, 1H), 7.28 (d, *J* = 7.2 Hz, 1H), 7.18 – 7.12 (m, 2H), 5.05 - 5.04 (m, 1H), 4.40 - 4.38 (m, 2H), 2.13 (s, 3H). LCMS (ESI) m/z calcd for  $C_9H_{11}NO_2 [M + H]^+$  165.07 found 166.1.

**N-(2-(hydroxymethyl)-3-methylphenyl) formamide (e2):** Following the general procedure A. Yellow solid (1.0 g, crude). R<sub>f</sub> (70% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>**H NMR** (CDCl<sub>3</sub>, 400 MHz): 8.41 (s, 1H), 7.96 (d, J = 8.0 Hz, 1H), 7.15 – 7.02 (m, 3H), 4.69 (s, 2H), 2.32 (s, 3H). **LCMS** (ESI) m/z calcd for C<sub>9</sub>H<sub>11</sub>NO<sub>2</sub> [M + H]<sup>+</sup> 165.07 found 166.1.

**N-(3-fluoro-2-(hydroxymethyl)phenyl)formamide (f2):** Following the general procedure A. Pale-yellow solid (0.98 g crude).  $R_f$  (70% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.58 (s, 1H), 8.25 (s, 1H), 7.30 (2, 1H), 7.14 – 7.12 (m, 1H), 5.24 (s, 1H), 4.42 – 4.41 (m, 2H). LCMS (ESI) m/z calcd for  $C_8H_8FNO_2$  [M + H]<sup>+</sup> 169.05 found 170.0.

**N-(4-fluoro-2-(hydroxymethyl) phenyl) formamide (g2):** Following the general procedure A. Pale-yellow solid (1.0 g crude).  $R_f$  (70% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): 8.46 (s, 1H), 7.96 (bs, 1H), 7.17 - 7.11 (m, 3H), 5.20 (s, 1H), 4.74 (s, 2H). LCMS (ESI) m/z calcd for  $C_8H_8FNO_2$  [M + H]<sup>+</sup> 169.05 found 170.0.

**N-(3-chloro-2-(hydroxymethyl)phenyl)formamide (h2):** Following the general procedure A. Pale-yellow solid (1.0 g crude).  $R_f$  (70% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (DMSO-d6, 400 MHz):  $\delta$  9.70 (s, 1H), 8.24 (s, 1H), 7.46 (d, J = 7.6 Hz, 1H), 7.39 (d, J = 7.6 Hz, 1H), 7.31 (t, J = 7.6 Hz, 2H). 5.26 (t, J = 5.6 Hz, 1H), 4.40 (d, J = 5.6 Hz, 2H). **LCMS** (ESI) m/z calcd for  $C_8H_8CINO_2$  [M + H]<sup>+</sup> 185.0 found 185.9.

**N-(5-chloro-2-(hydroxymethyl)phenyl)formamide (i2):** Following the general procedure A. Yellow solid (1.1 g crude).  $R_f$  (70% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (DMSO-d6, 400 MHz):  $\delta$  9.61 (s, 1H), 8.31 (s, 1H), 7.96 (s, 1H), 7.38 (d, *J* = 8.4 Hz, 1H), 7.20 - 7.24 (m, 1H), 5.36 (brs, 1H), 4.47 (s, 2H). **LCMS** (ESI) m/z calcd for  $C_8H_8CINO_2$  [M + H]<sup>+</sup> 185.0 found 185.9.

**N-(3-bromo-2-(hydroxymethyl)phenyl)formamide (j2)**: Following the general procedure A. Yellow solid (0.9 g crude), 78%).  $R_f$ (70% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): 8.46 (s, 1H), 8.09 (d, *J* = 8.0 Hz, 1H), 7.41 - 7.18 (m, 4H), 4.99 (s, 2H). **LCMS** (ESI) m/z calcd for  $C_8H_8BrNO_2 [M + H]^+$  228.97 found 230.0.

**N-(2-bromo-6-(hydroxymethyl)phenyl)formamide (k2):** Following the general procedure A. Yellow solid (0.95 g crude).  $R_f$  (70% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.38 (s, 1H), 8.23 (s, 1H), 7.59 (d, *J* = 8.0 Hz, 1H), 7.19 - 6.99 (m, 3H), 5.22 (s, 1H), 4.43 - 4.42 (m, 2H). LCMS (ESI) m/z calcd for  $C_8H_8BrNO_2$  [M + H]<sup>+</sup> 228.97 found 231.8.

**N-(4-fluoro-2-(hydroxymethyl)-6-methoxyphenyl)formamide (l2):** Following the general procedure A. White solid (0.9 g crude).  $R_f(70\%$  ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.23 (s, 1H), 8.13 (s, 1H), 6.90 - 6.80 (m, 2H), 5.21 (s, 1H), 4.31 (s, 2H), 3.75 (d, J = 7.2 Hz, 3H). **LCMS** (ESI) m/z calcd for  $C_9H_{10}FNO_3 [M + H]^+$  199.06 found 200.0.

**2-Formamido-5-methoxybenzyl methanesulfonate (b3):** Following the general procedure B. Yellow solid (0.98 g crude).  $R_f$  (30% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.60 (s, 1H), 8.26 - 8.24 (m, 1H), 7.54 (d, *J* = 8.45 Hz, 1H), 7.06 - 7.03 (m, 1H), 6.91 - 6.89 (m, 1H), 4.76 - 4.71 (m, 2H), 3.73 (s, 4H). LCMS (ESI) m/z calcd for  $C_{10}H_{13}NO_5S$  [M + H]<sup>+</sup> 259.05 found 200.1(demesylated product).

**2-Formamido-6-methoxybenzyl methanesulfonate (c3):** Following the general procedure B. Yellow solid (0.75 g crude).  $R_f$  (30% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.50 (s, 1H), 8.22 (s, 1H), 7.29 – 7.20 (m, 3H), 4.61 (s, 2H), 3.78 (s, 6H). LCMS (ESI) m/z calcd for  $C_{10}H_{13}NO_5S [M + H]^+$  259.05 found 199.9 (demesylated product).

**2-Formamido-3-methylbenzyl methanesulfonate (d3):** Following the general procedure B. Yellow solid (0.85 g crude, 57%). R<sub>f</sub> (30% ethyl acetate/ n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.69 (s, 1H), 8.27 (s, 1H), 7.63 (d, J = 8.0 Hz, 1H), 7.27 - 7.25 (m, 1H), 7.16 - 7.12 (m, 2H), 4.79 - 4.74 (m, 3H), 2.25 (s, 3H). LCMS (ESI) m/z calcd for C<sub>10</sub>H<sub>13</sub>NO<sub>4</sub>S [M + H]<sup>+</sup> 243.05 found 184.0 (demesylated product).

**2-Formamido-6-methylbenzyl methanesulfonate (e3):** Following the general procedure B. Yellow solid (0.75 g crude).  $R_f$  (30% ethyl acetate/ n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.69 (s, 1H), 8.27 (s, 1H), 7.63 (d, J = 8.0 Hz, 1H), 7.27 - 7.25 (m, 1H), 7.16 - 7.12 (m, 2H), 4.79 - 4.74 (m, 3H), 2.25 (s, 3H). LCMS (ESI) m/z calcd for  $C_{10}H_{13}NO_4S$  [M + H]<sup>+</sup> 243.05 found 184.0 (demesylated product).

**2-Fluoro-6-formamidobenzyl methanesulfonate (f3):** Following the general procedure B. Yellow solid (0.75 g crude).  $R_f$  (30% ethyl acetate/ n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.84 (s, 1H), 8.30 (s, 2H), 7.34 – 7.26 (m, 4H), 4.80 (s, 2H). LCMS (ESI) m/z calcd for C<sub>9</sub>H<sub>10</sub>FNO<sub>4</sub>S [M + H]<sup>+</sup> 247.03 found 188.0, 152.0 (demesylated product).

**5-Fluoro-2-formamidobenzyl methanesulfonate (g3):** Following the general procedure B. Yellow solid (0.8 g crude).  $R_f$  (30% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): 8.47 (s, 1H), 7.85 - 7.82 (m, 2H), 7.43 (bs, 1H), 7.21 - 7.06 (m, 2H), 4.56 (s, 2H). LCMS (ESI) m/z calcd for C<sub>9</sub>H<sub>10</sub>FNO<sub>4</sub>S [M + H]<sup>+</sup> 247.03 found 188.0, 246.0 (demesylated product).

**2-Chloro-6-formamidobenzyl methanesulfonate (h3):** Following the general procedure B. Yellow solid (0.7 g crude).  $R_f$  (30% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.78 (s, 1H), 8.32 (s, 1H), 8.01 (s, 1H), 7.80 (d, J = 8.0 Hz, 1H), 7.45

(d, J = 6.8 Hz, 1H), 4.80 (s, 2H), 3.33 (s, 3H). LCMS (ESI) m/z calcd for C<sub>9</sub>H<sub>10</sub>ClNO<sub>4</sub>S [M + H]<sup>+</sup> 263.0 found 170.0, 204.0 (demesylated product).

**4-Chloro-2-formamidobenzyl methanesulfonate (i3):** Following the general procedure B. Yellow solid (0.8 g crude).  $R_f$  (30% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.93 (s, 1H), 8.34 (s, 1H), 8.01 (s, 1H), 7.47 (d, *J* = 8.4 Hz, 1H), 7.25 - 7.20 (m, 1H), 4.84 - 4.80 (m, 3H). LCMS (ESI) m/z calcd for C<sub>9</sub>H<sub>10</sub>ClNO<sub>4</sub>S [M + H]<sup>+</sup> 263.0 found 204.0, 206.0 (demesylated product).

**2-Bromo-6-formamidobenzyl methanesulfonate (j3):** Following the general procedure B. Yellow solid (0.65 g crude).  $R_f$  (30% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.99 (s, 1H), 8.34 - 8.32 (m, 1H), 7.84 - 7.82 (m, 1H), 7.54 - 7.48 (m, 1H), 7.44 - 7.27 (m, 1H), 7.52 (d, *J* = 7.6 Hz, 1H), 7.34 - 7.30 (m, 1H), 5.14 (s, 2H), 3.20 (s, 3H). LCMS (ESI) m/z calcd for  $C_9H_{10}BrNO_4S [M + H]^+$  306.95 found 229.0 (demesylated product).

**3-Bromo-2-formamidobenzyl methanesulfonate (k3):** Following the general procedure B. Yellow solid (0.7 g crude).  $R_f$  (30% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.99 (s, 1H), 8.34 – 8.32 (m, 1H), 7.84 – 7.79 (m, 1H), 7.54 – 7.44 (m, 1H), 7.36 - 7.30 (m, 1H), 4.94 (s, 2H), 4.89 – 4.80 (m, 3H). LCMS (ESI) m/z calcd for  $C_9H_{10}BrNO_4S$  [M + H]<sup>+</sup> 306.95 found 229.1 (demesylated product).

**N-(2-(azidomethyl)-4-methoxyphenyl) formamide (b4):** Following the general procedure C. Pale-yellow solid (0.75 g crude).  $R_{f}(40\% \text{ ethyl acetate/n-hexane}) = 0.3.$  <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.49 (s, 1H), 8.20 (s, 1H), 7.30 - 7.26 (m, 1H), 7.13 - 6.98 (m, 2H), 4.31 (s, 1H), 3.78 (s, 3H). LCMS (ESI) m/z calcd for C<sub>9</sub>H<sub>10</sub>N<sub>4</sub>O<sub>2</sub> [M + H]<sup>+</sup> 206.08 found 179.0 (-N<sub>2</sub>).

**N-(2-(azidomethyl)-3-methoxyphenyl) formamide (c4):** Following the general procedure C. Pale-yellow solid (0.7 g crude).  $R_{f}(40\% \text{ ethyl acetate/n-hexane}) = 0.3.$  <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.49 (s, 1H), 8.20 (s, 1H), 7.30 - 7.26 (m, 1H), 7.13 - 6.98 (m, 2H), 4.31 (s, 1H), 3.78 (s, 3H). LCMS (ESI) m/z calcd for C<sub>9</sub>H<sub>10</sub>N<sub>4</sub>O<sub>2</sub> [M + H]<sup>+</sup> 206.08 found 179.0 (-N<sub>2</sub>).

**N-(2-(azidomethyl)-6-methylphenyl)formamide (d4)**: Following the general procedure C. Pale-yellow solid (0.7 g crude). R<sub>f</sub> (40% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>**H NMR** (DMSO-d<sub>6</sub>, 400 MHz): 9.59 (s, 1H), 8.25 (s, 1H), 7.29 - 7.20 (m, 3H), 4.35 (s, 2H), 2.16 (s, 3H). **LCMS** (ESI) m/z calcd for C<sub>9</sub>H<sub>10</sub>N<sub>4</sub>O [M + H]<sup>+</sup> 190.08 found 163.1 (-N<sub>2</sub>).

**N-(2-(azidomethyl)-3-methylphenyl) formamide (e4):** Following the general procedure C. Pale-yellow solid. Isolated yield: (0.7 g crude).  $R_f$  (40% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.80 (s, 1H), 8.27 (s, 1H), 7.52 (d, J = 8.0 Hz, 1H), 7.27 - 7.20 (m, 1H), 7.14 - 7.06 (m, 1H), 4.48 (s, 2H), 2.34 (s, 3H). LCMS (ESI) m/z calcd for  $C_9H_{10}N_4O$  [M + H]<sup>+</sup> 190.08 found 163.1 (-N<sub>2</sub>).

N-(2-(azidomethyl)-3-fluorophenyl) formamide (f4): Following the general procedure C. Pellow solid (0.6 g crude).  $R_f$  (40% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.83 (s, 1H), 8.28 (s, 1H), 7.36 - 7.26 (m, 3H), 4.42 (s, 2H). LCMS (ESI) m/z calcd for  $C_8H_7FN_4O$  [M + H]<sup>+</sup> 194.06 found 167.1 (-N<sub>2</sub>).

**N-(2-(azidomethyl)-4-fluorophenyl)formamide (g4).** Following the general procedure C. Yellow solid (0.62 g crude).  $R_f$  (40% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.82 – 9.76 (m, 1H), 8.26 (s, 1H), 7.71 - 7.70 (m, 1H), 7.34 – 7.16 (m, 2H), 4.47 (s, 2H). LCMS (ESI) m/z calcd for  $C_8H_7FN_4O$  [M + H]<sup>+</sup> 194.06 found 167.1 (-N<sub>2</sub>).

N-(2-(azidomethyl)-3-chlorophenyl) formamide (h4): Following the general procedure C. Pale-yellow solid (0.7 g crude).  $R_f$  (40% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.83 (s, 1H), 8.28 (s, 1H), 7.36 – 7.26 (m, 3H), 4.42 (s, 2H). LCMS (ESI) m/z calcd for  $C_8H_7ClN_4O$  [M + H]<sup>+</sup> 210.03 found 183.1 (-N<sub>2</sub>).

**N-(2-(azidomethyl)-5-chlorophenyl) formamide (i4):** Following the general procedure C. Pale-yellow solid (0.75 g crude). R<sub>f</sub> (40% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): 8.42 (s, 1H), 7.50 (d, J = 8.0 Hz, 1H), 7.37 - 7.33 (m, 1H), 7.31 - 7.25 (m, 2H), 4.40 (s, 2H). LCMS (ESI) m/z calcd for C<sub>8</sub>H<sub>7</sub>ClN<sub>4</sub>O [M + H]<sup>+</sup> 210.03 found 183.1 (-N<sub>2</sub>).

**N-(2-(azidomethyl)-3-bromophenyl) formamide (j4):** Following the general procedure C. Yellow solid (0.60 g crude).  $R_f$  (40% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.99 (s, 1H), 8.28 (s, 1H), 7.76 (d, *J* = 8.0 Hz, 1H), 7.61 - 7.28 (m, 2H), 4.62 (s, 2H). LCMS (ESI) m/z calcd for  $C_8H_7BrN_4O [M + H]^+ 253.98$  found 229.0 (-N<sub>2</sub>).

**N-(2-(azidomethyl)-6-bromophenyl) formamide (k4):** Following the general procedure C. Yellow solid (0.63 g crude).  $R_f$  (40% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d<sub>6</sub>, 400 MHz): 9.75 (s, 1H), 8.29 (s, 1H), 7.77 (d, *J* = 7.6 Hz, 1H), 7.40 - 7.17 (m, 2H), 4.48 (s, 2H). LCMS (ESI) m/z calcd for  $C_8H_7BrN_4O$  [M + H]<sup>+</sup> 253.98 found 229.0 (-N<sub>2</sub>).

**N-(2-(azidomethyl)-4-fluoro-6-methoxyphenyl)formamide (l4):** Following the general procedure C. White solid (0.6 g crude).  $R_f$  (40% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (DMSO-d\_6, 400 MHz): 9.44 (s, 1H), 8.19 (s, 1H), 7.07 - 6.83 (m, 2H), 4.32 (s, 2H), 3.79 (s, 3H). LCMS (ESI) m/z calcd for C<sub>9</sub>H<sub>9</sub>FN<sub>4</sub>O<sub>2</sub> [M + H]<sup>+</sup> 224.07 found 197.1, 225.1 (-N<sub>2</sub>).

**1-(Azidomethyl)-2-isocyanobenzene (1a).**<sup>11</sup> Following the general procedure D. Yellow liquid (0.55 g, 87%).  $R_f$  (10% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  7.46 - 7.38 (m, 4H), 4.55 (s, 2H). <sup>13</sup>C {<sup>1</sup>H} NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  168.19, 132.19, 129.82, 129.27, 129.25, 50.89. LCMS (ESI) m/z calcd for  $C_8H_6N_4$  [M + H]<sup>+</sup> 158.05 found 131.1 (-N<sub>2</sub>).

**2-(Azidomethyl)-1-isocyano-4-methoxybenzene (1b):** Following the general procedure D. Yellow liquid (0.56 g, 87%).  $R_f$  (10% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  7.37 - 7.34 (m, 1H), 7.02 - 6.94 (m, 2H), 4.50 (s, 2H), 3.92 (s, 3H). <sup>13</sup>C {<sup>1</sup>H} NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  171.77, 154.44, 133.62, 130.18, 120.32, 112.45, 56.24, 50.90. LCMS (ESI) m/z calcd for  $C_9H_8N_4O$  [M + H]<sup>+</sup> 188.06 found 161.1 (-N<sub>2</sub>).

**2-(Azidomethyl)-1-isocyano-3-methoxybenzene (1c):** Following the general procedure D. Yellow liquid (0.52 g, 81%).  $R_f(10\%)$  ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  7.37 (t, *J* = 8.0 Hz, 1H), 7.02 - 6.94 (m, 2H), 4.51 (s, 2H), 3.94 (s, 3H). <sup>13</sup>C {<sup>1</sup>H} NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  171.77, 155.44, 133.62, 130.18, 120.32, 111.45, 56.24, 50.90. LCMS (ESI) m/z calcd for  $C_9H_8N_4O$  [M + H]<sup>+</sup> 188.06 found 161.1 (-N<sub>2</sub>).

1-(Azidomethyl)-2-isocyano-3-methylbenzene (1d): Following the general procedure D. Yellow liquid (0.5g, 71%).  $R_f$  (10% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  7.28 - 7.25 (m, 3H), 4.55 (s, 2H), 2.43 (s, 3H). <sup>13</sup>C {<sup>1</sup>H} NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  7.28 - 7.25 (m, 3H), 4.55 (s, 2H), 2.43 (s, 3H).

100 MHz):  $\delta$  167.69, 139.39, 131.74, 130.00, 129.35, 124.99, 47.65, 19.41. **LCMS** (ESI) m/z calcd for C<sub>9</sub>H<sub>8</sub>N<sub>4</sub> [M + H]<sup>+</sup> 172.07 found 145.1 (-N<sub>2</sub>).

**2-(Azidomethyl)-1-isocyano-3-methylbenzene (1e):** Following the general procedure D. Yellow liquid (0.5g, 78%).  $R_f$  (10% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  7.28 - 7.25 (m, 3H), 4.55 (s, 2H), 2.43 (s, 3H). <sup>13</sup>C {<sup>1</sup>H} NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  167.55, 139.39, 131.76, 129.99, 129.37, 125.01, 47.63, 19.45. LCMS (ESI) m/z calcd for  $C_9H_8N_4$  [M + H]<sup>+</sup> 172.07 found 145.1 (-N<sub>2</sub>).

**2-(Azidomethyl)-1-fluoro-3-isocyanobenzene (1f):** Following the general procedure D. Yellow liquid (0.48 g, 75%). R<sub>f</sub> (10% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): δ 7.45 - 7.39 (m, 1H), 7.26 - 7.17 (m, 2H), 4.55 (s, 2H). <sup>13</sup>C {<sup>1</sup>H} NMR (CDCl<sub>3</sub>, 100 MHz): δ 174.31, 159.05, 156.48, 134.50, 130.76, 124.29, 116.42, 50.53. <sup>19</sup>F NMR (376 MHz, CDCl<sub>3</sub>): -110.38 - -110.39 (d, 1F). LCMS (ESI) m/z calcd for C<sub>8</sub>H<sub>5</sub>FN<sub>4</sub> [M + H]<sup>+</sup> 176.04 found 149.1 (-N<sub>2</sub>).

**2-(Azidomethyl)-4-fluoro-1-isocyanobenzene (1g):** Following the general procedure D. Yellow liquid (0.5 g, 78%). R<sub>f</sub> (10% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz): δ 7.44 – 7.41 (m, 1H), 7.26 – 7.20 (s, 1H), 7.09 – 7.05 (m, 1H), 4.56 (s, 2H). <sup>13</sup>C{<sup>1</sup>H} NMR (CDCl<sub>3</sub>, 100 MHz): δ 168.54, 163.66, 161.14, 135.10, 129.03, 116.09, 50.57. <sup>19</sup>F NMR (376 MHz, DMS0-d<sub>6</sub>): - 121.12 (t, 1F). LCMS (ESI) m/z calcd for C<sub>8</sub>H<sub>5</sub>FN<sub>4</sub> [M + H]<sup>+</sup> 176.04 found 149.1 (-N<sub>2</sub>).

2-(Azidomethyl)-1-chloro-3-isocyanobenzene (1h): Following the general procedure D. Yellow liquid. Isolated yield: (0.49 g, 76%).  $R_f(10\%$  ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  7.48 (s, 1H), 7.38 (s, 2H), 4.57 (s, 2H). <sup>13</sup>C{<sup>1</sup>H} NMR

(CDCl<sub>3</sub>, 100 MHz): δ 173.37, 134.40, 131.66, 130.07, 129.75, 127.11, 51.15. **LCMS** (ESI) m/z calcd for C<sub>8</sub>H<sub>5</sub>ClN<sub>4</sub> [M + H]<sup>+</sup> 192.02 found 165.0 (-N<sub>2</sub>).

**1-(Azidomethyl)-4-chloro-2-isocyanobenzene (1i):** Following the general procedure D. Yellow liquid (0.51 g, 80%).  $R_f$  (10% ethyl acetate/n-hexane) = 0.3. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  7.45 – 7.40 (m, 1H), 7.27 – 7.17 (m, 2H), 4.56 (s, 2H). <sup>13</sup>C{<sup>1</sup>H} **NMR** (CDCl<sub>3</sub>, 100 MHz):  $\delta$  174.41, 159.06, 156.49, 134.50, 130.67, 124.28, 116, 40, 50.52. **LCMS** (ESI) m/z calcd for  $C_8H_5ClN_4$  [M + H]<sup>+</sup> 192.02 found 165.0 (-N<sub>2</sub>).

**2-(Azidomethyl)-1-bromo-3-isocyanobenzene (1j):** Following the general procedure D. Yellow liquid (0.48 g, 73%).  $R_f$  (10% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  7.69 (d, J = 8.0 Hz, 1H), 7.42 (d, J = 9.2 Hz, 1H), 7.28 (d, J = 8.0 Hz, 1H), 4.64 (s, 2H). <sup>13</sup>C{<sup>1</sup>H} NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  169.66, 134.39, 132.08, 130.81, 129.81, 126.63, 125.97, 50.77. LCMS (ESI) m/z calcd for  $C_8H_5BrN_4$  [M + H]<sup>+</sup> 235.96 found 209.0 (-N<sub>2</sub>).

**1-(azidomethyl)-3-bromo-2-isocyanobenzene (1k):** Following the general procedure D. Yellow liquid (0.46 g, 70%).  $R_f$  (10% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  7.69 (d, *J* = 8.0 Hz, 1H), 7.42 (d, *J* = 9.6 Hz, 1H), 7.28 – 7.26 (m, 1H), 4.70 (s, 2H). <sup>13</sup>C{<sup>1</sup>H} NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  169.67, 134.39, 132.08, 130.59, 126.63, 125.97, 50.48. LCMS (ESI) m/z calcd for  $C_8H_5BrN_4$  [M + H]<sup>+</sup> 235.96 found 210.9, 208.9 (-N<sub>2</sub>).

1-(Azidomethyl)-5-fluoro-2-isocyano-3-methoxybenzene (11): Following the general procedure D. Yellow liquid (0.46, 70%).  $R_f(10\% \text{ ethyl acetate/n-hexane}) = 0.2$ . <sup>1</sup>H NMR (CDCl<sub>3</sub>, 400 MHz):  $\delta$  6.76 (d, J = 8.0 Hz, 1H), 6.66 (d, J = 9.2 Hz, 1H), 4.52 (s, 2H), 3.93 (s, 3H). <sup>13</sup>C {<sup>1</sup>H} NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  171.84, 164.22, 161.70, 156.94, 135.69, 107.11, 99.88, 56.58, 50.62. LCMS (ESI) m/z calcd for C<sub>9</sub>H<sub>7</sub>FN<sub>4</sub>O [M + H]<sup>+</sup> 206.06 found 179.1 (-N<sub>2</sub>).

**1-(1-Azidoethyl)-2-isocyanobenzene (1m).**<sup>11</sup> following the general procedure D. Pale-green oil (0.42, 65%. R<sub>f</sub> (10% ethyl acetate/n-hexane) = 0.2. <sup>1</sup>H NMR (CDCl<sub>3</sub>, 500MHz):  $\delta$  7.52 (d, J = 7.6 Hz, 1H), 7.46 (d, J = 6.15 Hz, 1H), 7.41 (d, J = 7.6 Hz, 1H), 7.35 (d, J = 7.6 Hz, 1H), 5.09 (q, J = 6.9 Hz, 1H), 1.57 (s, 3H). <sup>13</sup>C{<sup>1</sup>H} NMR (CDCl<sub>3</sub>, 100 MHz):  $\delta$  169.66, 134.39, 132.08, 130.81, 129.81, 126.63, 125.97, 50.77. HRMS (ESI-TOF) (m/z): [M + Na]<sup>+</sup> Calcd for C<sub>9</sub>H<sub>8</sub>N<sub>4</sub> 173.0827; Found 173.0806.

# 5. Copies of <sup>1</sup>H and <sup>13</sup>C NMR for Final products





























NUCLEUS :	F19			
FRQ (MHz):	376.00			
AT (sec):	0.73			
SW (ppm):	237.46			
RD (sec):	1.00		·	
PW (usec):	4.40			
NT :	16	E		
Solvent :	cdcl3	ſ <u>∕</u> i∕∼Ŋ		
NP :	16			
LB (Hz):	1.0			
TEMP (K):	303.15	_0		
EXP :	s2pul	3g		
NMR REPORTS:				
PROJECT NAME :	-R-02			
PROBE: ATB				 

3.0




























## 7. Copies of <sup>1</sup>H and <sup>13</sup>C NMR for Intermediates












































































	7.287	7.258	4.550	2.437	
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NUCLEUS : H1	1				
FRQ (MHz): 399.63					
AT (sec): 2.28					
SW (ppm): 17.98					
RD (sec): 1.00					
PW (usec): 5.35		1			
NT : 64					
Solvent : cdcl3					
NP : 64					
LB (Hz): 1.0		<sup>^</sup> N₃			
TEMP (K): 303.15		NC			
EXP : s2pul	Ĭ	NC			
DATE :Nov 13 2018	(1.4)				
TIME :09:04:31	(10)				
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NUCLEUS : F19		
FRQ (MHz): 376.00		
AT (sec): 0.73		
SW (ppm): 237.46	F	
RD (sec): 1.00	$\downarrow$ $\land$ .	
PW (usec): 4.40	N <sub>3</sub>	
NT : 16	NC	
Solvent : cdcl3	1f	
NP : 16	"	
LB (Hz): 1.0		
TEMP (K): 303.15		
EXP : s2pul		
NMR REPORTS:		
PROJECT NAME : R-02		
PROBE: ATB		



























