

General Comments

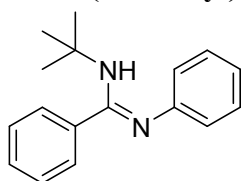
NMR spectra were recorded on Bruker Avance 300 (300 MHz). Chemical shifts (ppm) are given relative to solvent: references for DMSO were 2.50 ppm (^1H -NMR) and 39.50 ppm (^{13}C -NMR). ^{13}C NMR spectra were acquired on a broad band decoupled mode. Multiplets were assigned as s (singlet), d (doublet), t (triplet), dd (doublet of doublet), m (multiplet) and br. s (broad singlet). All measurements were carried out at room temperature unless otherwise stated. Electron impact (EI) mass spectra were recorded on AMD 402 mass spectrometer (70 eV). High resolution mass spectra (HRMS) were recorded on Agilent 6210. The data are given as mass units per charge (m/z). Gas chromatography analysis was performed on an Agilent HP-5890 instrument with a FID detector and HP-5 capillary column (polydimethylsiloxane with 5% phenyl groups, 30 m, 0.32 mm i.d., 0.25 μm film thickness) using argon as carrier gas. The products were isolated from the reaction mixture by column chromatography on silica gel 60, 0.063-0.2 mm, 70-230 mesh (Merck).

All reactions were carried out under air atmosphere. Dimethylformamide (anhydrous, 99.8%) was purchased from ACROS and used as received. All the reagents were purchased from Sigma-Aldrich or Alfa-Aesar chemical company. Various functionalized silanes were prepared based on literature.^[1]

General procedure for the synthesis of *N*-(*tert*-butyl)-*N'*-phenylbenzimidamide

In a 5 mL pressure tube equipped with a stirring bar, trimethoxy(phenyl)silane (0.2 mmol), $\text{Cu}(\text{OAc})_2$ (0.4 mmol), PdCl_2 (5 mol%), KF (0.4 mmol), *tert*-butyl isocyanide (0.3 mmol), aniline (0.3 mmol) and DMF (2 mL) were added. Then close the tube and heat it up to 120 $^\circ\text{C}$ for 24 h. Cool the reaction mixture to room temperature when the reaction completed. The reaction solution was quenched with distilled water and extracted with ethyl acetate three times. The combined organic phases were washed with saturated NaCl solution and dried over Na_2SO_4 . The crude product was purified by column chromatography (ethyl acetate/pentane = 1:10) to give the pure product.

4a: *N*-(*tert*-Butyl)-*N'*-phenylbenzimidamide



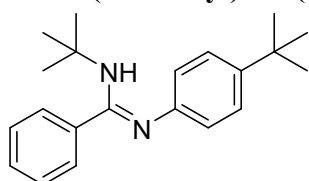
67%, 33.8 mg.

^1H NMR (300 MHz, $\text{DMSO}-d_6$) δ 7.28-7.09 (m, 5H), 7.02-6.89 (m, 2H), 6.71-6.59 (m, 1H), 6.49-6.38 (m, 2H), 6.32 (s, 1H), 1.47 (s, 9H).

^{13}C NMR (75 MHz, DMSO) δ 156.65, 152.24, 136.25, 129.34, 128.83, 128.52, 128.19, 122.96, 120.26, 51.72, 28.98.

GC-MS(EI-70eV): m/z (%)=252 (32), 195.2 (95), 180 (33), 104 (25), 93 (100), 77.1 (71).

4b: *N*-(*tert*-Butyl)-*N'*-(4-(*tert*-butyl)phenyl)benzimidamide



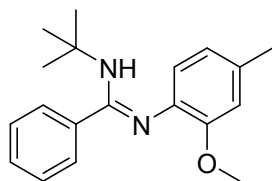
61%, 30.6 mg.

¹H NMR (300 MHz, DMSO-*d*₆) δ 7.29-7.11 (m, 5H), 7.04-6.92 (m, 2H), 6.43-6.31 (m, 2H), 6.22 (s, 1H), 1.46 (s, 9H), 1.16 (s, 9H).

¹³C NMR (75 MHz, DMSO) δ 156.38, 149.36, 142.16, 136.43, 129.33, 128.85, 128.21, 125.18, 122.41, 51.68, 34.07, 31.82, 29.00.

GC-MS(EI-70eV): m/z(%)=308 (61), 251 (80), 237 (24), 134 (100), 104 (28).

4c: *N*-(*tert*-Butyl)-*N'*-(2-methoxy-4-methylphenyl)benzimidamide



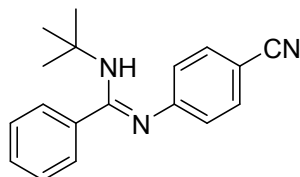
55%, 32.6 mg.

¹H NMR (300 MHz, DMSO-*d*₆) δ 7.23-7.05 (m, 5H), 6.64-6.44 (m, 3H), 6.15 (s, 1H), 3.48 (s, 3H), 2.04 (s, 3H), 1.50 (s, 9H).

¹³C NMR (75 MHz, DMSO) δ 156.75, 150.45, 140.71, 137.47, 129.56, 128.63, 127.71, 127.54, 122.31, 120.84, 109.65, 55.67, 51.52, 29.16, 18.94.

GC-MS(EI-70eV): m/z(%)=296 (84), 225 (33), 209 (100), 137 (24), 122 (25), 104 (20).

4d: *N*-(*tert*-Butyl)-*N'*-(4-cyanophenyl)benzimidamide



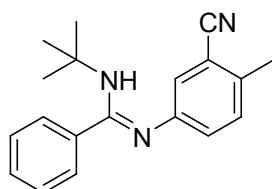
68%, 67.7 mg.

¹H NMR (300 MHz, DMSO-*d*₆) δ 7.37 (d, *J* = 8.4 Hz, 2H), 7.32-7.22 (m, 3H), 7.20-7.13 (m, 2H), 6.90 (s, 1H), 6.54 (d, *J* = 8.5 Hz, 2H), 1.46 (s, 9H).

¹³C NMR (75 MHz, DMSO) δ 157.22, 156.45, 134.66, 132.39, 129.07, 128.93, 127.98, 123.23, 119.83, 101.05, 51.63, 28.26.

GC-MS(EI-70eV): m/z(%)=277 (35), 22 (100), 161(22), 118 (83), 77 (22).

4e: *N*-(*tert*-Butyl)-*N'*-(3-cyano-4-methylphenyl)benzimidamide



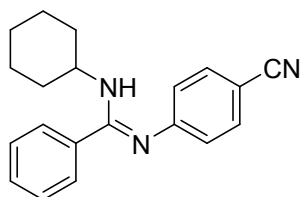
66%, 38,4 mg.

¹H NMR (300 MHz, DMSO-*d*₆) δ 7.36-7.22 (m, 3H), 7.16 (tt, *J* = 4.5, 2.6 Hz, 2H), 7.00 (d, *J* = 8.2 Hz, 1H), 6.74-6.58 (m, 3H), 2.25 (s, 3H), 1.47 (s, 9H).

¹³C NMR (75 MHz, DMSO) δ 157.47, 150.15, 135.01, 131.98, 130.25, 128.85, 128.71, 127.90, 127.72, 125.42, 118.21, 111.12, 51.43, 28.36, 18.88.

GC-MS(EI-70eV): m/z(%)=291 (38), 234 (100), 219 (29), 132 (54), 104 (38).

4f: *N*-(4-Cyanophenyl)-*N'*-cyclohexylbenzimidamide

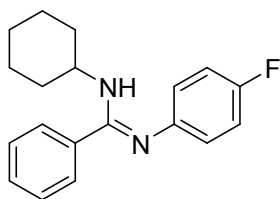


61%, 37 mg.

¹H NMR (300 MHz, DMSO-*d*₆) δ 7.42-7.34 (m, 2H), 7.34-7.23 (m, 4H), 7.22-7.12 (m, 2H), 6.55 (d, *J* = 8.5 Hz, 2H), 3.85 (s, 1H), 2.07-1.94 (m, 2H), 1.80-1.54 (m, 3H), 1.36-1.05 (m, 5H).

¹³C NMR (75 MHz, DMSO) δ 157.83, 157.26, 134.69, 132.87, 129.72, 129.28, 128.58, 123.96, 120.31, 101.73, 50.16, 32.39, 25.89, 25.27.

GC-MS(EI-70eV): m/z(%)=303 (67), 220 (100), 121 (39), 104 (98), 77 (22).

4g: N'-Cyclohexyl-N-(4-fluorophenyl)benzimidamide

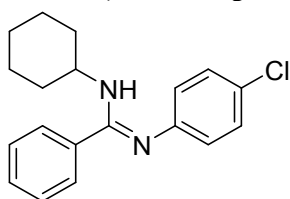
63%, 37.2 mg.

¹H NMR (300 MHz, DMSO-*d*₆) δ 7.25 (dd, *J* = 4.9, 1.6 Hz, 3H), 7.17-7.10 (m, 2H), 6.77 (t, *J* = 8.8 Hz, 3H), 6.42 (dd, *J* = 8.3, 5.2 Hz, 2H), 3.85 (s, 1H), 2.12-1.91 (m, 2H), 1.56-1.77 (m, 3H), 1.36-1.10 (m, 5H).

¹³C NMR (75 MHz, DMSO) δ 160.41 (d, 254.1 Hz), 157.26, 148.72, 135.67, 129.19, 129.08, 128.35, 124.09 (d, *J* = 7.4 Hz), 114.99 (d, *J* = 21.9 Hz), 49.71, 32.54, 25.99, 25.35.

¹⁹F NMR (282 MHz, DMSO) δ -108.5.

GC-MS(EI-70eV): *m/z*(%)=295 (54), 213 (74), 176 (12), 111 (100), 95 (24), 77 (16).

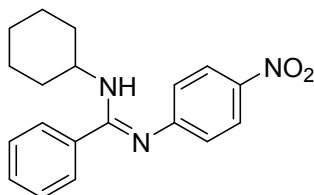
4h: N'-(4-Chlorophenyl)-N-cyclohexylbenzimidamide

64%, 39.9 mg.

¹H NMR (300 MHz, DMSO-*d*₆) δ 7.18-7.10 (m, 3H), 7.07-6.99 (m, 2H), 6.89-6.70 (m, 3H), 6.37-6.26 (m, 2H), 3.71 (s, 1H), 1.93-1.83 (m, 2H), 1.46-1.59 (m, 3H), 1.24-0.99 (m, 5H).

¹³C NMR (75 MHz, DMSO) δ 157.32, 151.35, 135.40, 129.70, 129.20, 128.42, 128.37, 124.74, 124.24, 49.80, 32.49, 25.96, 25.32.

GC-MS(EI-70eV): *m/z*(%)=312 (55), 229 (71), 192 (12), 127 (100), 104 (65), 77 (18).

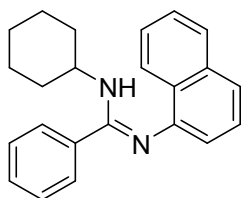
4i: N-Cyclohexyl-N'-(4-nitrophenyl)benzimidamide

47%, 30.4 mg.

¹H NMR (300 MHz, DMSO-*d*₆) δ 7.92-7.80 (m, 2H), 7.53 (d, *J* = 7.5 Hz, 1H), 7.32 (d, *J* = 1.8 Hz, 3H), 7.22 (dd, *J* = 2.6, 1.2 Hz, 2H), 6.57 (d, *J* = 9.0 Hz, 2H), 3.86 (d, *J* = 9.4 Hz, 1H), 2.09-1.89 (m, 2H), 1.83-1.49 (m, 3H), 1.42-1.04 (m, 5H).

¹³C NMR (75 MHz, DMSO) δ 159.97, 158.28, 140.29, 134.42, 130.00, 129.33, 128.69, 124.87, 123.30, 50.38, 32.34, 25.85, 25.23.

GC-MS(EI-70eV): *m/z*(%)=323 (32), 293 (74), 240 (54), 195 (42), 173 (32), 138 (21), 108 (100).

4j: N-Cyclohexyl-N'-(naphthalen-1-yl)benzimidamide

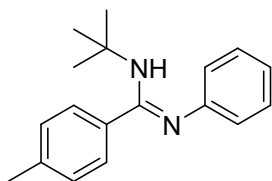
57%, 37.4 mg.

¹H NMR (300 MHz, DMSO-*d*₆) δ 8.15- 8.07 (m, 1H), 7.77-7.64 (m, 1H), 7.49-7.37 (m, 2H), 7.28-6.94 (m, 8H), 6.25 (dd, *J* = 7.2, 1.1 Hz, 1H), 4.00 (s, 1H), 2.28-2.11 (m, 2H), 1.77 (dd, *J* = 8.7, 4.4 Hz, 2H), 1.65-1.60 (m, 1H), 1.45-1.21 (m, 5H).

¹³C NMR (75 MHz, DMSO) δ 156.99, 148.75, 135.75, 134.15, 129.40, 129.16, 128.45, 128.24, 127.99, 126.41, 126.00, 125.09, 124.49, 120.39, 117.04, 50.14, 32.69, 26.03, 25.40.

GC-MS(EI-70eV): *m/z*(%)=328 (59), 245 (68), 230 (50), 208 (21), 143 (100), 127 (44), 104 (28), 77 (10).

4l: *N*-(*tert*-Butyl)-4-methyl-*N'*-phenylbenzimidamide



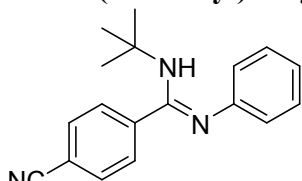
67%, 35.6 mg.

^1H NMR (300 MHz, $\text{DMSO}-d_6$) δ 7.23-7.11 (m, 6H), 6.82 (d, J = 7.3 Hz, 1H), 6.63 (d, J = 8.4 Hz, 2H), 6.39 (s, 1H), 2.40 (s, 3H), 1.65 (s, 9H).

^{13}C NMR (75 MHz, DMSO) δ 156.60, 152.37, 138.22, 133.37, 129.32, 128.73, 128.54, 122.91, 120.18, 51.64, 28.96, 21.20.

GC-MS(EI-70eV): m/z (%)=266 (42), 209 (100), 194 (31), 118 (35), 93 (69), 77 (20).

4m: *N*-(*tert*-Butyl)-4-cyano-*N'*-phenylbenzimidamide



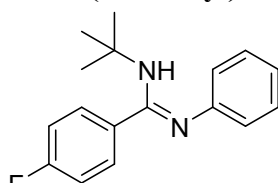
64%, 35.5 mg.

^1H NMR (300 MHz, $\text{DMSO}-d_6$) δ 7.76-7.66 (m, 2H), 7.37-7.26 (m, 2H), 7.05-6.92 (m, 2H), 6.75-6.62 (m, 1H), 6.59 (s, 1H), 6.50-6.39 (m, 2H), 1.47 (s, 9H).

^{13}C NMR (75 MHz, DMSO) δ 155.16, 151.57, 140.81, 132.16, 130.37, 128.70, 122.96, 120.82, 118.85, 111.40, 51.99, 28.87.

GC-MS(EI-70eV): m/z (%)=277 (30), 220 (100), 205 (20), 129 (20), 93 (80), 77 (38).

4n: *N*-(*tert*-Butyl)-4-fluoro-*N'*-phenylbenzimidamide



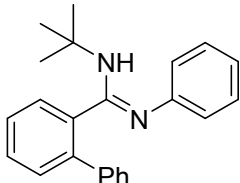
58%, 31.3 mg.

^1H NMR (300 MHz, $\text{DMSO}-d_6$) δ 7.17 (dd, J = 8.8, 5.6 Hz, 2H), 7.07- 6.91 (m, 4H), 6.78- 6.57 (m, 1H), 6.43 (dd, J = 8.3, 1.1 Hz, 2H), 6.37 (s, 1H), 1.46 (s, 9H).

^{13}C NMR (75 MHz, DMSO) δ 162.20 (d, J = 224.7 Hz), 155.69, 152.10, 132.60 (d, J = 3.1 Hz), 131.60(d, J = 8.6 Hz), 128.60, 122.96, 120.40, 114.99(d, J = 21.6 Hz), 51.73, 28.91. ^{19}F NMR (282 MHz, DMSO) δ -108.5.

GC-MS(EI-70eV): m/z (%)=270 (42), 213 (100), 198 (33), 122 (28), 93 (67), 77 (25).

4o: *N*-(*tert*-Butyl)-*N'*-phenyl-[1,1'-biphenyl]-2-carboximidamide



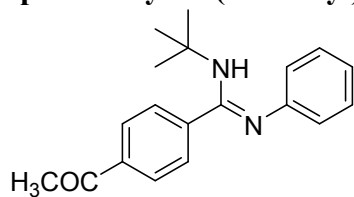
53%, 34.8 mg.

^1H NMR (300 MHz, $\text{DMSO}-d_6$) δ 7.41-7.27 (m, 6H), 7.24-7.18 (m, 2H), 7.16-7.10 (m, 1H), 6.83-6.69 (m, 2H), 6.62-6.53 (m, 1H), 6.50 (s, 1H), 5.95-5.84 (m, 2H), 1.40 (s, 9H).

^{13}C NMR (75 MHz, DMSO) δ 155.64, 152.27, 140.72, 139.95, 130.72, 129.63, 129.19, 129.08, 128.00, 127.77, 127.50, 126.91, 122.72, 120.24, 51.69, 28.71.

GC-MS(EI-70eV): m/z (%)=328 (32), 271 (100), 180 (33), 93 (31).

4p: 4-Acetyl-*N*-(*tert*-butyl)-*N'*-phenylbenzimidamide



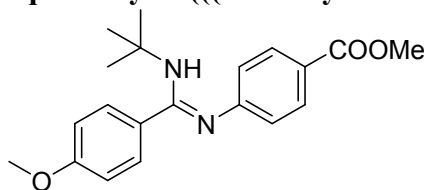
58%, 34.1 mg.

^1H NMR (300 MHz, $\text{DMSO}-d_6$) δ 7.86-7.75 (m, 2H), 7.36-7.23 (m, 2H), 7.04-6.90 (m, 2H), 6.75-6.60 (m, 1H), 6.46 (dtd, J = 8.5, 1.9, 1.2 Hz, 3H), 2.52 (s, 3H), 1.48 (s, 9H).

^{13}C NMR (75 MHz, DMSO) δ 198.04, 155.79, 151.84, 140.67, 136.78, 129.70, 128.65, 128.06, 122.92, 120.62, 51.91, 28.92, 27.20.

GC-MS(EI-70eV): $m/z(\%)$ =294 (41), 237 (100), 222 (20), 146 (26), 93 (81), 77 (43).

4q: Methyl 4-(((*tert*-butylamino)(4-methoxyphenyl)methylene)amino)benzoate



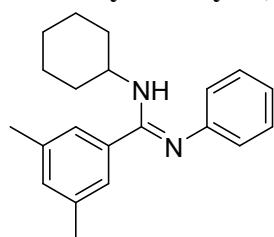
58%, 39.5 mg.

^1H NMR (300 MHz, $\text{DMSO}-d_6$) δ 7.66-7.55 (m, 2H), 7.17-7.05 (m, 2H), 6.85-6.74 (m, 2H), 6.67 (s, 1H), 6.56-6.45 (m, 2H), 3.74 (s, 3H), 3.70 (s, 3H), 1.46 (s, 9H).

^{13}C NMR (75 MHz, DMSO) δ 166.77, 160.05, 157.58, 157.11, 131.04, 130.26, 127.70, 122.80, 120.81, 113.69, 55.57, 51.96, 51.91, 28.86.

GC-MS(EI-70eV): $m/z(\%)$ =340 (35), 283 (90), 191 (22), 151 (100), 120 (51).

4r: *N*-Cyclohexyl-3,5-dimethyl-*N'*-phenylbenzimidamide



57%, 34.9 mg.

^1H NMR (300 MHz, $\text{DMSO}-d_6$) δ 7.06- 6.91 (m, 2H), 6.87 (s, 1H), 6.76 (s, 2H), 6.73-6.56 (m, 2H), 6.45 (d, J = 7.4 Hz, 2H), 3.82 (s, 1H), 2.13 (s, 6H), 2.02-1.99 (m, 2H), 1.76-1.55 (m, 3H), 1.34-1.14 (m, 5H). ^{13}C NMR (75 MHz, DMSO) δ 156.68, 152.32, 137.15, 135.80, 130.37, 128.47, 126.89, 123.01, 120.36, 49.68, 32.59, 26.02, 25.40, 21.24.

GC-MS(EI-70eV): $m/z(\%)$ =306 (50), 223 (100), 149 (25), 132 (45), 93 (58).

HRMS (EI), m/z : $[\text{M}]^+$ calculated for $\text{C}_{21}\text{H}_{26}\text{N}_2$:306.2096, found $[\text{M}+\text{H}]^+$: 307.21715.

[1] A. S. Manoso, C. Ahn, A. Soheli, C. J. Handy, R. Correia, W. M. Segamish, P. Deshong, *J. Org. Chem.* **2004**, *69*, 8305-8314.

