

A simple and efficient protocol for the synthesis of quinolines catalyzed by chloramine-T

Y. Venkateswarlu, S. Ramesh Kumar and P. Leelavathi*

Department of Chemistry, University College for Women, Koti, Osmania University, Hyderabad-500095, India

(Received October 20, 2011; Revised March 7, 2012; Accepted April 26, 2012)

Abstract: Chloramine-T has been proved as an efficient catalyst for the synthesis of substituted quinolines. In this method, 2-amino aryl ketones were smoothly reacted with ketones to afford the corresponding quinoline derivatives in very good yields. All the reactions were carried out at acetonitrile reflux.



Keywords: 2-Amino aryl ketones; ketones; chloramine-T; quinolines.

1. Introduction

Quinoline scaffold are found in many natural products and they exhibit remarkable biological activities like Antimalarial, antibacterial, antiasthmatic, antihypertensive, anti-inflammatory agents and antiobesity.¹⁻⁸ Aryl-substituted quinolines act as ligands for 5-lipoxygenase⁹, tyrosine kinase¹⁰, leukotriene¹¹ and other receptors. Furthermore, polyquinolines were shown to undergo hierarchical self-assembly into nanostructures with promising electronic and photonic properties.^{12,13} A variety of methods such as Doebner-von Miller, Skraup, Combes, Friedlander and Knorr synthesis have been used for preparation of quinolines and their derivatives.¹⁴⁻¹⁶

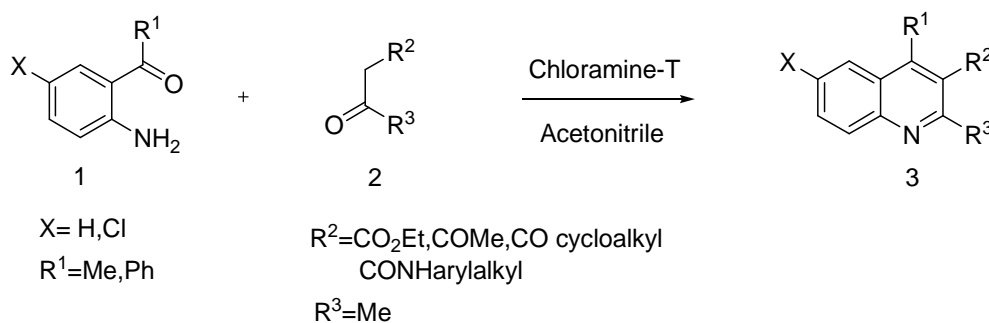
Therefore, the synthesis of quinoline derivatives attracted many researchers and various methods have been developed using a variety of catalysts and conditions.¹⁷⁻²⁶ Among them, Friedlander annulations are one of the simplest and most straightforward protocols. This method involves a condensation and cyclization between a ketone possessing a methylene group and an aromatic 2-aminoaldehyde or ketone. However, many of these methods have some drawbacks such as high reaction temperature (150-200^oC), use of expensive catalysts and extended reaction times. As part of our research program in developing synthetic methodologies²⁷⁻²⁹, herein we report, the synthesis of

* Corresponding author: E-Mail: yekkiralavenkat@gmail.com

quinolines using chloramines-T as a catalyst. The catalyst chloramines-T is known in the literature for various organic transformations.³⁰

2. Results and Discussion

In a typical experiment, 2-amino acetophenone and ethyl acetoacetate were reacted in presence of chloramines-T at acetonitrile reflux to afford the corresponding product, ethyl-2, 4-dimethyl quinoline-3-carboxylate (**3a**) in excellent yield. The reaction was completed within 4 hours.



Scheme 1. synthesis of quinolines

We have examined the effect of temperature on reaction rate and the amount of catalyst used in the reaction and the results were summarized in the table-1. There was no product formation in acetonitrile at room temperature and at reflux conditions even after 24 hours. The product formation was observed in presence of catalyst at room temperature after 24 hours. It was found that the ideal reaction conditions were at acetonitrile reflux and using the catalyst in 10% mole.

Table 1. Optimization of Reaction Conditions for the synthesis of quinolines using Chloramine-T:

SNo	Solvent	Catalyst	Temperature (°C)	Time (h)	Yield (%)
1	CH ₃ CN	0	RT	24	0
2	CH ₃ CN	0	85	24	0
3	CH ₃ CN	0.5	RT	24	60
4	CH ₃ CN	0.5	85	3.0	95
5	CH ₃ CN	1.0	85	3.0	95
6	CH ₃ CN	0.1	85	4.0	95

Encouraged by the result obtained with 2-amino acetophenone and ethyl acetoacetate at established reaction conditions, we have applied this methodology to various substrates. As shown in the table 2, the 2-aminoacetophenone could be replaced by 2-amino-5-chlorobenzophenone and *ortho*-methylene ketones could be extended from ethyl acetoacetate to methylacetoacetate, cyclic-1,3-diketones, such as 1,3-cyclohexanedione and 5,5-dimethyl-1,3-cyclohexanedione and simple cyclic ketones such as cyclohexanone 3-*tert*-cyclohexanone and cyclopentanone. In the optimized reaction conditions, the synthesis of quinoline **1a-o** was successfully obtained in very good yields by Friedlander condensation between 2-aminoacetophenone as well as 2-aminobenzophenone with a variety of carbonyl compounds in the presence of catalyst chloramine-T.

3. Conclusion

In conclusion, the application of various catalysts for the preparation of quinolines via Friedlander annulations such as been studied. chloramines-T been demonstrated here as the most

Synthesis of quinolines catalyzed by chloramine-T

effective catalyst for this synthesis. The simple experimental procedure and impressive yields by applying this inexpensive catalyst have made this protocol practically useful for the synthesis of quinolines.

4. Experimental:

All commercial reagents were used without purification and all solvents were reagent grade. All the reaction mixtures were stirred magnetically and were monitored by TLC using 0.25 mm E-Merck silica gel 60F₂₅₄ precoated glass plates, which were visualized with UV light. Melting points were recorded on Buchi R-535 apparatus. IR spectra were recorded on a Perkin-Elmer FT/IR-240 C spectrophotometer with KBr optics. ¹H NMR spectra were recorded on Varian Gemini 200 MHz spectrometer recorded in CDCl₃ using TMS as an internal standard. ¹³C NMR spectra (75.5 MHz) Mass spectra were recorded on a Bruker Avance-300 finnigan spectrometer with complete proton decoupling, chemical shifts are reported in ppm relative to the solvent resonance as the internal standard (CDCl₃, δ=77.16). MAT 1020 Mass spectrum operating at 70 eV. Mass spectra were recorded on a VG 7070 H Micromass spectrometer. CHN analyses were recorded on a Vario EL Analyser.

4.1. General procedure for the synthesis of quinoilne compounds (4a-o):

A mixture of 2-aminoaryl ketone (1 mmol), ethyl acetoacetate (1.2 mmol), and catalyst chloramines-T (1 mmol) in acetonitrile (5 mL), at reflux for a specified time in Table 1. The progress of the reaction was monitored by thinlayer chromatography (TLC). After completion of the reaction, the organic solvent was removed under reduced pressure. The obtained crude product was purified by column chromatography on silica gel by hexane: ethyl acetate as an eluent.

4.2. Spectral data for all the compounds:

Ethyl-2, 4-dimethylquinoline-3-carboxylate (3a) : Yellow solid. Melting range. 271-272 °C. IR (KBr): 3070, 2930, 2873, 1725, 1614, 1589, 1214, 1082, 578 cm⁻¹. ¹H NMR (300 MHz CDCl₃) (δ/ppm): 1.45 (t, 3H, J = 7.0 Hz), 2.63 (s, 3H), 2.70 (s, 3H), 4.50 (q, 2H, J = 7.0 Hz), 7.50 (t, 1H, J = 7.0 Hz), 7.70 (t, 1H, J = 7.0 Hz), 7.95 (d, 1H, J = 8.3 Hz), 8.0 (d, 1H, J = 8.4 Hz). ¹³C NMR (75 MHz, CDCl₃) (δ/ppm): 14.10, 15.45, 23.62, 61.43, 123.82, 125.73, 126.11, 127.95, 129.22, 129.86, 141.27, 147.0, 154.24, 168.93. EIMS m/z (%): 229 (91), 214 (8), 200 (10), 183 (100), 156 (50), 128 (20), 115 (37), 89 (18), 77 (10); Anal. Calcd for C₁₄H₁₅NO₂ (230): C, 73.34 %; H, 6.59 %; N, 6.11 %. Found: C, 73.12 %; H, 6.48 %; N, 6.05 %.

9-Methyl-3, 4-dihydroacridine-1(2H)-one (3b) : White solid. Melting range. 78-79 °C. IR (KBr): 3068, 2935, 1478, 1614, 1581, 1350, 755 706, 539 cm⁻¹. ¹H NMR (500 MHz CDCl₃) (δ/ppm): 2.15-2.30 (m, 2H), 2.75 (t, 2H, J = 6.0 Hz), 3.05 (s, 3H), 3.40 (d, 2H), 7.55 (t, 1H, J = 7.0 Hz), 7.75 (t, 1H, J = 7.5 Hz), 8.00 (d, 1H, J = 7.5 Hz), 8.20 (d, 1H, J = 7.5 Hz). ¹³C NMR (CDCl₃, 75 MHz) (δ/ppm): 12.89, 24.57, 32.40, 41.28, 122.79, 124.68, 126.45 (2C), 127.16, 128.45, 140.71, 145.41, 157.92, 198.23, Anal Calcd (%) for C₁₄H₁₃NO: C, 79.59 %; H, 6.20 %; N, 6.63 %; O, 7.57 %. Found: C, 79.34 %; H, 6.19 %; N, 6.78 %.

9-Methyl-1, 2, 3, 4-tetrahydroacridine (3c): Solid. Melting range. 75-77 °C. IR (KBr): 2928, 1569, 1478, 1348, 1164, 1076, 939, 839, 819, 775, 752, 708, cm⁻¹. ¹H NMR (300 MHz. CDCl₃) (δ/ppm): 1.55-1.65 (m, 4H), 2.55 (s, 3H), 2.90 (t, 2H, J = 7.6 Hz), 3.10 (t, 2H, J = 7.6 Hz), 7.40-7.95 (m, 4H). ¹³C NMR (75 MHz, CDCl₃) (δ/ppm): 12.91, 22.36, 22.74, 26.53, 33.92, 122.75, 124.54, 126.61 (2C), 127.65, 128.43, 140.77, 145.44, 161.95. EIMS m/z (%): 198 (M⁺, 87), 125 (100). Anal. Calcd for C₁₄H₁₅N (197): C, 85.24 %; H, 7.66 %; N, 7.10 %. Found: C, 85.12 %; H, 6.58 %; N, 7.05 %.

7-Methyl-5, 6-dihydrobenzo[*c*]acridine (3d) : Solid. Melting range. 112 °C. IR (KBr): 3070, 3018, 2946, 2842, 1680, 1582, 1499, 1215, 758 cm⁻¹. ¹H NMR (300 MHz, CDCl₃) (δ/ppm): 2.65 (s, 3H), 3.00 (t, 2H, *J*=7.0 Hz), 3.15 (t, 2H, *J*=6.8 Hz), 7.15-8.55 (m, 8H). ¹³C NMR (75 MHz, CDCl₃) (δ/ppm): 13.60, 25.03, 27.81, 123.35, 125.27, 126.14, 126.95, 127.43, 128.06, 128.90, 129.17, 129.92, 133.17, 134.92, 138.81, 139.49, 146.61, 152.35. MS *m/z* (%): 246 (M⁺, 100), 212 (30), 125 (31). Anal. Calcd for C₁₈H₁₅N (246): C, 88.13 %; H, 6.16 %; N, 5.71 %. Found; C, 88.12%; H, 6.09 %; N, 5.63 %.

Allyl 2, 4-dimethylquinoline-3-carboxylate (3e) : Viscose oil. Melting range 265-267 °C. IR (KBr); 3032, 2988, 2961, 1624, 1577, 1498, 1267, 1221, 755 cm⁻¹. ¹H NMR (300 MHz CDCl₃) (δ/ppm): 2.65 (s, 3H), 2.70 (s, 3H), 4.82 (dt, 2H, *J*=2.40 & 5.25 Hz), 5.50 (dd, 2H, *J*= 1.56 & *J* = 15.0 Hz), 5.95-6.20 (m, 1H), 7.45-8.05 (m, 4H). MS *m/z* (%); 241 (M⁺). Anal. Calcd for C₁₅H₁₅NO₂ (241): C, 74.67 %; H, 7.627 %; N, 5.80 %. Found: C, 7.66 %; H, 6.38 %; N, 5.82 %.

Ethyl 2-methyl-4-phenylquinoline-3-carboxylate (3f) : Solid. Melting range 98-99 °C. IR (KBr): 3030, 2960, 1700, 1605, 1568, 1482, 905 cm⁻¹. ¹H NMR (300 MHz CDCl₃) (δ/ppm): 1.30 (t, 3H, *J* =7.0 Hz), 2.80 (s, 3H), 4.20 (q, 2H, *J* = 7.0 Hz), 7.30-7.60 (m, 6H), 7.75 (d, 1H, *J*=8.1 Hz), 7.80 (t, 1H, *J*=7.9 Hz), 8.10 (d, 1H, *J* = 8.1 Hz). ¹³C NMR (75 MHz) (δ/ppm): 13.65, 23.32, 68.81, 96.14, 125.1, 126.1, 126.4, 127.8, 128.2, 129.1, 129.5, 135.7 (2C), 135.7, 145.7, 147.8, 153.6, 167.7. MS *m/z* (%): 291 (M⁺) (70), 281 (20), 264 (30), 246 (20), 221 (20), 207 (20), 191 (15), 147 (40), 133 (15), 73 (100) Anal. Calcd for C₁₉H₁₇NO₂ (291): C, 78.33 %; H, 5.88 %; N, 4.81 %. Found: C, 78.42 %; H, 5.91 %; N, 4.88 %.

1-(6-Chloro-2-methyl-4-phenylquinolin-3-yl) ethanone (3g) : Solid. Melting range. 149-150 °C. IR (KBr): 3029, 2960, 1701, 1606, 1567, 1481, 909, 602 cm⁻¹. ¹H NMR (300 MHz CDCl₃) (δ/ppm): 1.98 (s, 3H), 2.66 (s, 3H), 7.30-7.20 (m, 7H), 8.00 (d, 1H, *J* =8.9 Hz). ¹³C NMR (δ/ppm): 23.58, 31.60, 124.71, 125.82, 128.83, 129.11, 129.84, 130.85, 132.27, 134.49, 135.46, 142.95, 145.86, 153.87 (2C), 204.91; MS *m/z* (%): 295 (M⁺) (100), 280 (25), 254 (10), 154 (15), 136 (20), 91 (35), 81 (60), 69 (40), 55 (50). Anal. Calcd for C₁₈H₁₄NO₂Cl (295): C, 73.10 %; H, 4.77 %; N, 4.74 %. Found: C, 73.00 %; H, 4.60 %; N, 4.66 %.

7-Chloro-9-phenyl-2, 3-dihydro-1H-cyclopenta[*b*] quinoline (3h) : Solid. Melting range. 104-106 °C. IR (KBr): 3437, 3042, 2950, 1972, 1920, 1751, 1600, 1584, 1482, 1440, 1381, 1339, 1305, 1201, 1160, 1124, 1073, 1027, 949, 878, 828, 755, 705, 658 cm⁻¹. ¹H NMR (δ/ppm): 2.10-2.20 (m, 2H), 2.90 (t, 2H, *J* =7.2 Hz), 3.20 (t, 2H, *J* =7.0 Hz), 7.30-7.55 (m, 7H), 7.95 (d, 1H, *J* = 8.5 Hz). ¹³C NMR (δ/ppm): 23.28, 30.99, 35.01, 124.42, 125.22, 126.91, (2C), 128.25, 129.01 (3C), 129.84, 130.35, 131.20, 138.11, 141.82, 144.90, 162.77. MS *m/z* (%): 279 (M⁺), 280 (100), 276, 246, 230, 203, 190, 179, 158, 150, 128. Anal Calcd for C₁₈H₁₄ClN (279): C, 73.10 %; H, 4.775%; N, 4.74 %. Found: C, 73.0 %; H, 4.60 %; N, 4.66 %.

Isopropyl-6-chloro-2-methyl-4-phenylquinoline-3-carboxylate (3i) : Solid Melting range. 124-125 °C. IR (KBr): 3426, 3055, 2982, 2928, 1900, 1722, 1606, 1579, 1560, 1477, 1448, 1390, 1309, 1278, 1225, 1187, 1107, 1085, 954, 908, 878, 829, 804, 760, 703 cm⁻¹. ¹H NMR (300 MHz CDCl₃) (δ/ppm): 0.90 (d, 6H, *J*=6.1 Hz), 2.70 (s, 3H), 4.80 (heptet, 1H, *J*=6.1 Hz), 7.30 (d, 2H, *J*=7.65 Hz), 7.40 (t, 4H, *J*=7.65 Hz), 7.55 (dd, 1H, *J*=9.0 & 2.2 Hz), 7.90 (d, 1H, *J*=8.4 Hz). MS *m/z* (%): 340 (M⁺, 40), 292 (100), 264 (16). Anal. Calcd for C₂₀H₁₈ClNO₂ (340): C, 70.69 %; H, 5.34 %; N, 4.12 %; O, 9.42 %; Cl, 10.43 %. Found: C, 70.60 %; H, 5.30 %; N, 4.02 %; O, 9.32 %; Cl, 10.33 %.

2-Tert-butyl-7-chloro-9-phenyl-1, 2, 3, 4-tetrahydroacridine (3j) : Solid Melting range. 154-155 °C. IR (KBr): 3040, 2980, 2890, 1600, 1560, 1475, 1370, 1360, 1160, 1070, 950, 825, 760, 700 cm⁻¹. ¹H

Synthesis of quinolines catalyzed by chloramine-T

NMR (300 MHz CDCl₃) (δ /ppm): 0.87 (s, 9H), 1.41-1.62 (m, 2H), 2.20-2.35 (m, 2H), 2.60-2.70 (m, 1H), 3.00-3.15 (m, 1H), 3.22-3.33 (m, 1H), 7.18-7.23 (m, 3H), 7.50-7.60 (m, 4H), 7.90 (d, 1H, J = 8.87 Hz). ¹³C NMR (125 MHz, CDCl₃) (δ /ppm): 23.99, 27.1 (3C), 29.34, 32.49, 34.75, 44.56, 124.50, 127.4 (2C), 128.75, 129.02 (3C), 129.61, 129.85, 130.04, 131.11, 136.32, 144.63, 145.90, 159.72. MS m/z (%): 350 (M⁺ 31.4), 349 (85), 293 (48), 292 (100), 57 (60). Anal. Calcd for C₂₃H₂₄Cl N (350.167); C, 78.965 %; H, 6.91 %; N, 4.00 %, Cl, 10.13 %. Found: C, 78.79 %; H, 6.84 %; N, 3.78 %, Cl, 9.98 %.

7-Chloro-3, 3-dimethyl-9-phenyl-3, 4-dihydroacridine-1(2H)-one (3k) : Yellow solid. Melting range. 219-220 °C. IR (KBr): 3074, 2952, 2866, 1696, 1554, 1477, 1384, 1297, 1198, 1079, 837, 699 cm⁻¹. ¹H NMR (CDCl₃, 300 MHz) (δ /ppm): 1.15 (s, 6H), 2.52 (s, 2H), 3.23 (s, 2H), 7.15 (t, 2H), 7.36-7.37 (s, 1H), 7.48-7.53 (t, 3H) 7.64-7.69 (dd, 1H, J_1 = 9.06, J_2 = 2.26 Hz), 7.95-7.99 (d, 1H, J = 9.06 Hz). ¹³C NMR (75 MHz, CDCl₃) (δ /ppm): 24.14, 27.14 (2C), 45.23, 55.10, 127.23, 127.92 (2C), 128.20, 129.10 (3C), 129.95, 131.15, 132.62, 133.55, 139.26, 146.24, 148.36, 158.77, 197.64; MS m/z (%): 336 (500), 280 (60), 250 (40), 245 (140), 217 (80), 249 (100), 131 (120), 113 (60). Anal. Calcd for C₂₁H₁₈ClNO: C, 75.10 %; H, 5.40 %; N, 4.17 %. Found: C, 75.07 %; H, 5.42 %; N, 4.18 %.


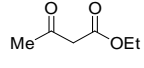
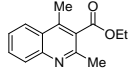


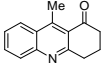


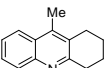


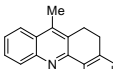

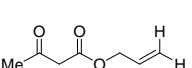
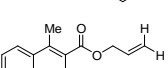

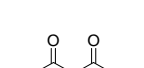
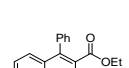

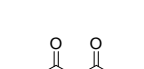
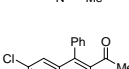


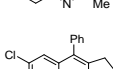

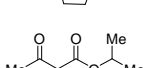
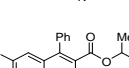


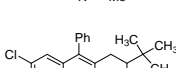


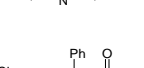

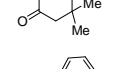
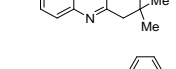

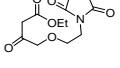
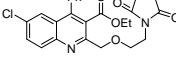

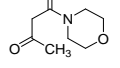
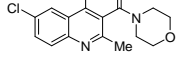

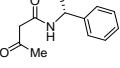
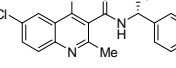
Ethyl-6-chloro-2-(2-phthalimidoethoxy methyl)-4-phenyl quinoline-3-carboxylate (3l) : Yellow solid: Melting range. 165-166 °C. ¹H NMR (CDCl₃, 300 MHz) (δ /ppm): 0.95 (t, 3H J = 7.5 Hz), 3.70 (t, 2H, J = 6.0 Hz), 3.81-3.87 (t, 2H, J = 6.0 Hz), 4.05 (q, 2H J = 7.5 Hz), 4.90 (s, 2H), 7.29-7.34 (m, 3H) 7.45-7.51 (m, 4H) 7.61-7.73 (m, 2H) 7.77-7.83 (m, 2H), 7.80-8.02 (d, 1H, J = 9.0 Hz): ¹³C NMR (75 MHz, CDCl₃) (δ /ppm): 13.50, 37.34, 60.37, 67.55, 73.48, 123.32 (2C), 126.93 (2C), 127.10, 127.50 (2C), 129.22 (3C), 130.06, 131.75, 132.01 (2C), 132.04, 132.20 (2C), 138.15, 148.44, 149.20, 159.55, 167.25 (2C), 168.10; MS m/z (%): 515 (1050), 370 (200), 342 (100), 245 (250), 108 (100), 217 (100), 149 (100), 131 (150). Anal. Calcd for C₂₉H₂₃ClN₂O₅: 515.1373, Found; 515.1359.

(6-Chloro-2-methyl-4-phenyl quinolin-3-yl)(morpholino) methanone (3m) : Yellow solid; Melting range. 187-189 °C. ¹H NMR (CDCl₃, 300 MHz) (δ /ppm): 2.68 (s, 3H), 2.75-2.91 (m, 2H), 2.97-3.22 (m, 2H), 3.27-3.40 (m, 1H), 3.45-3.63 (m, 3H), 7.23-7.33 (m, 1H), 7.46-7.68 (m, 6H), 8.03 (d, 1H, J = 9.14 Hz); ¹³C NMR (75 MHz, CDCl₃) (δ /ppm): 14.43, 41.35 (2C), 66.27 (2C), 124.14, 125.67 (2C), 127.01, 129.14 (3C), 129.27, 130.06, 131.00, 132.42, 138.12, 143.3, 146.11, 155.12, 166.85. MS m/z (%): 367 (M⁺100).

(R)-6-Chloro-2-methyl-4-phenyl-N-(1-phenylethyl) quinoline-3-carboxamide (3n) : Yellow solid; Melting range. 225-227 °C. ¹H NMR (CDCl₃, 300 MHz) (δ /ppm): 1.15 (d, 3H, J = 6.80 Hz), 2.80 (s, 3H), 5.05 (d, 1H, J_1 = 6.80 Hz, J_2 = 7.55 Hz), 5.50 (broad doublet, NH, J = 7.55 Hz), 6.91-7.00 (m, 2H), 7.19-7.29 (m, 4H), 7.34-7.43 (m, 2H), 7.47-7.56 (m, 3H), 7.61-7.66 (dd, 1H, J_1 = 9.06 Hz, J_2 = 2.26 Hz), 7.997-8.02 (d, 1H, J = 9.06 Hz); ¹³C NMR (δ /ppm): 14.43, 23.50, 48.63, 125.10, 126.97, 126.10, 127.31, 127.42, 127.53, 128.71, 128.82, 129.32, 129.43, 129.43, 130.51, 130.72, 130.84, 132.21, 134.66, 138.72, 141.84, 144.01, 145.88, 155.80, 166.67. MS (ESI); m/z (%) 401 (M⁺100).

6-Chloro-2-methyl-4-phenyl-N-P-tolylquinoline-3-carboxamide (3o) : Melting range 227-229 °C. ¹H NMR (CDCl₃, 300 MHz) (δ /ppm): 2.29 (s, 3H), 2.83 (s, 3H), 6.80 (s, 1H, NH), 6.93-7.01 (m, 4H), 7.40-7.55 (m, 7H), 8.00 (d, 1H, J = 9.06, Hz). MS (ESI); m/z (%) 387.20 (M⁺ 100).

Table 2. Chloramine-T catalyzed Friedlander synthesis of quinolines

Entry	2-Amino ketone	Ketone/diketones	Product	Time (h)	Yield (%)
a				4.0	95
b				3.5	89
c				4.0	87
d				3.5	88
e				3.0	92
f				3.0	93
g				4.0	91
h				4.5	85
i				3.0	91
j				4.0	90
k				3.5	94
l				4.5	89
m				5.0	90
n				4.5	86
o				4.0	87

Synthesis of quinolines catalyzed by chloramine-T

References

- [1] Welsch, M. E.; Snyder, S. A.; Stockwell, B. R. Privileged scaffolds for library design & drug discovery. *Curr. Opin. Chem. Biol.* **2010**, *14*, 347-361.
- [2] Michael, J. P. Quinoline, quinazoline & acridone alkaloids. *Nat. Prod. Rep.* **1995**, *12*, 77-89.
- [3] Warshakoon, C. N.; Sheville, J.; Bhatt, T. R.; Ji, Mendez Andino, L. J.; Meyers, M. K.; Kim, N., Wos, A. J.; Mitchell, C.; Jeennifer L. Paris, L. P.; Benth B. Pinney, B. B.; Ofer Reizes, O.; Hu, X. E. Design & synthesis of substituted quinolines as novel & selective MCHA as antiobesity agents. *Bioorg. Med. Chem. Lett.* **2006**, *16*, 5207-5211.
- [4] Larsen, R. D.; Corley, E. G.; King, A. O.; Carroll, J. D.; Davis, P.; Verhoeven, T. R.; Reider, P. J.; Labelle, M.; Gauthier, J. Y.; Xiang, Y. B.; Zamboni, R. J. Practical route to a new class of LTD4 receptor antagonists. *J. Org. Chem.* **1996**, *61*, 3398- 3405.
- [5] Chen, Y. L.; Fang, K. C.; Sheu, J. Y.; Hsu, S. L.; Tzeng, C. C. Synthesis & antibacterialevaluation of certain quinolone derivatives. *J. Med. Chem.* **2001**, *44*, 2374-2377.
- [6] Roma, G.; Di Braccio, M.; Grossi, G., Mattioli, F.; Ghia, M. 1, 8-Naphthyridines IV. 9-Substituted N, N-dialkyl-5(alkylamino or cycloalkylamino) [1, 2, 4] triazolo [4, 3- a][1,8] naphthyridine-6-carboxamides, new compounds with anti-aggressive & potent anti-inflammatory activities. *Eur. J. Med. Chem.* **2000**, *35*, 1021-1035.
- [7] Kalluraya, B.; Sreenivas, S. Synthesis & pharmacological properties of some quinoline derivatives. *Farmaco.* **1998**, *53*, 399-404.
- [8] Doube, D.; Blouin, M.; Brideau, C.; Chan, C.; Desmarais, S.; Eitheir, D.; Falgueyret, J. P.; Friesen, R. W.; Girard, M.; Girard, Y.; Guay, J.; Tagari, P.; Young, R. N. Quinolines as potent 5-lipoxygenase inhibitors; synthesis & biological Profile of L- 746, 530. *Bioorg. Med. Chem. Lett.* **1998**, *8*, 1255-1260.
- [9] Maguire, M. P.; Sheets, K. R.; McVert, K.; Spada, A. P.; Zilberstein, A. A new series of PDGF receptor tyrosine kinase inhibitors: 3-substituted quinoline derivatives. *J. Med. Chem.* **1994**, *37*, 2129-2137.
- [10] Musser, J. H.; Chakraborty, U. R.; Sciortino, S.; Gordon, R. J.; Khandwala, A.; Neiss, E. S.; Pruss, T. P.; Van Inwegen, R.; Weinryb, I.; Coutts, S. M. Substituted arylmethyl phenyl Ethers. A novel series of 5-lipoxygenase inhibitors & leukotriene antagonists. *J. Med. Chem.* **1987**, *30*, 96-104.
- [11] Van Inwegen, R. J.; Khandwala, A.; Gordon, R.; Sonnio, P.; Coutts, S.; Joly, S. Rev 5901: An orally effective peptidoleukotriene antagonist, detailed biochemical /pharmacological Profile. *J. Pharmacol. Exp. Ther.* **1987**, *24*, 117-124.
- [12] Gauthier, J. Y.; Jones, T.; Champion, E.; Charette, L.; Dehaven, R.; Hatchinson, A. W. F.; Hoogsteen, K.; Lord, A.; Masson, P.; Piechuta, H.; Pong, S. S.; Springer, J. P.; Therein, M.; Zamboni, R.; Young, R. N. Stereospecific synthesis, assignment of absolute configuration & biological activity of the enantiomers of 3-[[[3-[2-(7-chloroquinolin-2-yl)-(E)-ethenyl] phenyl-3-(dimethylamino)-3-0-0-propyl] thio] methyl] thio] propionic acid, a potent & specific leukotriene D₄ receptor antagonist. *J. Med. Chem.* **1990**, *33*, 2841-2845.
- [13] Jenekhe, S. A.; Lu, L.; Alan, M. M. New conjugated polymers with donor-acceptor architectures: Synthesis and photophysics of carbazole-quinoline and phenothiazine-quinoline copolymers and oligomers exhibiting large intramolecular charge transfer. *Macromolecules.* **2001**, *34*, 7315-7324.
- [14] Abass, M. Fused quinolines: recent synthetic approaches to azoloquinolines: A Review. *Heterocycles.* **2005**, *65*, 901-965.
- [15] Kouznetsov, V. V.; Mendez, L. Y. V.; Gomez, C. M. M. Recent progress in the synthesis of quinolines. *Curr. Org. Chem.* **2005**, *9*, 141-161.
- [16] Denmark, S. E.; Venkatraman, S. On the mechanism of the Skraup-Doebner Von Miller quinoline synthesis. *J. Org. Chem.* **2006**, *71*, 1668-1676.
- [17] Yadav, J. S.; J. S.; Rao, P. P.; Sreenu, D.; Rao, R S.; Kumar, V. N.; Nagaiah, K.; Prasad, A. R. Sulfamic acid: An efficient, cost-effective & recyclable solid acid catalyst for the Friedlander quinoline synthesis. *Tetrahedron Lett.* **2005**, *46*, 7249-7253.
- [18] Narasimhulu, M.; Reddy, T. S.; Mahesh, K. C.; Prabhakar, P.; Rao, C. B.; Lu, Y. V. Silica supported perchloric acid: A mild & highly efficient heterogeneous catalyst for the synthesis of poly-substituted quinolines via Friedlander hetero annulations. *J. Mol. Catal. A: Chem.* **2007**, *266*, 114-117.
- [19] Wu, J.; Zhang, L.; Diao, T. N. An expeditious approach to quinolines via Friedländer synthesis catalyzed by FeCl₃ or Mg (ClO₄)₂. *Synlett.* **2005**, *17*, 2653-2657.
- [20] Varala, R.; Enugala, R.; Adapa, S. R. Efficient & rapid Friedlander synthesis of functionalized quinolines catalyzed by Nd(NO₃)₃.6H₂O. *Synthesis.* **2006**, *22*, 3825-3830.
- [21] Palimkar, S. S.; Siddiqui, S. A.; Daniel, T.; Lahoti, R. J.; Srinivasan, J V. Ionic liquid promoted regioselective Friedlander annulation: novel synthesis of quinolines & fused polycyclic quinolines. *J. Org. Chem.* **2003**, *68*, 9371-9376.

- [22] Muscia, G. C.; Bollini, M.; Carnevale, J. P.; Bruno, A. M.; Asi's, S. E. Microwave-assisted Friedlander synthesis of quinolines derivatives as potential antiparasitic agents. *Tetrahedron Lett.* **2006**, *47*, 8811-8815.
- [23] Cho, C. S.; Ren, W. X. A recyclable Pd-catalyzed modified Friedlander quinoline synthesis. *J. Organometal. Chem.* **2007**, *692*, 4182-4185.
- [24] Niknam, K.; Zolfigol, M. A.; Dehghani, A. Friedlander quinoline synthesis Catalyzed by M (HSO₄)_n Al, Mg, Ca) under solvent-free conditions. *Heterocycles.* **2008**, *75*, 2513-2521.
- [25] Zhang, X. L.; Wang, Q. Y.; Sheng, S. R.; Wang, Q.; Liu, X. L. Efficient Friedlander synthesis of quinoline derivatives from 2-aminoarylketones & carbonyl compounds mediated by recyclable PEG-supported sulfonic acid. *Synth. Commun.* **2009**, *39*, 3293-3304.
- [26] Ghassamipour, S.; Sardarian, A. R. Friedlander synthesis of poly-substituted quinolines. In the presence of dodecylphosphonic acid as a highly efficient, recyclable & novel catalyst in aqueous media & solvent-free conditions. *Tetrahedron Lett.* **2009**, *50*, 514-519.
- [27] Venkateswarlu, Y.; Leelavathi, P. NbCl₅: An efficient catalyst for the synthesis of Quinoxalines. *Lett. Org. Chem.* **2010**, *7*, 208-211.
- [28] Venkateswarlu, Y.; Kumar, S. R.; Leelavathi, P. Cadmium chloride : A simple and efficient catalyst for the synthesis of 1,4-dihydropyridine (Hantzsch pyridines). *Int J Ind Chem* **2011**, *4*, ASAP;
- [29] Kumar, S. R.; Venkateswarlu, Y.; Leelavathi, P. Lanthanum (III) chloride : An efficient catalyst for the synthesis of substituted thiazoles *OSMC.* 2012, 1.
- [30] Ramesh, N.; Vipin Kumar, V. Iodine catalyzed one-pot diamination of glycols with chloramine-T: A new approach to 2-amino-β-glycosylamines for applications in N-glycopeptide synthesis. *Chem. Commun.* **2006**, 4952-4954.

A C G
publications

© 2012 Reproduction is free for scientific studies