

(八) *N*-Substituted benzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]-quinolin-3,4-dione (109-123) 之合成

N-Benzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (109) 之合成

取化合物 35(2.15g , 0.01mole)懸著於 DMF 30 ml 中 , 加入無水 K_2CO_3 (1.38 g , 0.01 mole)加熱(約 70-80)使之溶解 , 加入 benzyl chloride(11.3g , 0.1mole), 反應 1 小時後加入冰水中 , 以 $CHCl_3$ 萃取 , 取 $CHCl_3$ 層 , 以無水 $MgSO_4$ 乾燥 , 減壓濃縮後 , 收集沉殿物以短程矽膠管柱層析($CHCl_3/EtOH$)沖提 , 再以 MeOH 及 $CHCl_3$ 做再結晶 , 得白色棉絮狀結晶 , 為化合物 109 (1.95 g, 63.93 %) , mp : 205~208 。光譜數據如下 : MS m/z : 305; IR (KBr) cm^{-1} : 1728.8 ($C_3=O$), 1605.3 ($C_4=O$); UV λ_{max} nm (MeOH) (log ε): 243(4.87); 1H -NMR (DMSO- d_6) δ: 2.36 (3H, s, C_6-CH_3), 4.91 (2H, s, H-2), 5.55 (2H, s, H-10), 7.29-7.35 (5H, m, Ar-H), 7.50-7.52 (2H, m, H-7, H-8), 7.99 (1H, s, H-5); ^{13}C -NMR (DMSO- d_6) δ: 20.43 (C_6-CH_3), 46.37 (C-10), 76.14 (C-2), 100.34 (C-3a), 117.16 (C-8), 126.60 (C-13, C-15), 126.85 (C-12, C-16), 127.98 (C-4a), 129.09 (C-6, C-14), 134.38 (C-5), 134.53 (C-7), 135.17 (C-11), 136.20 (C-8a), 171.49 (C-9a), 174.41 (C-4), 191.20 (C-3) .

N-o-Methylbenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (110) 之合成

取化合物 35(2.15g , 0.01mole)和 *o*-methylbenzyl chloride (12.7g , 0.1mole)為原料 , 比照化合物 109 的合成法及處理步驟 , 得化合物 110(1.85g , 57.99 %) , mp:263-265 。光譜數據如下 : MS m/z : 319; IR (KBr) cm^{-1} : 1721.0($C_3=O$), 1605.3 ($C_4=O$); UV λ_{max} nm (MeOH) (log ε): 244 (4.78); 1H -NMR (DMSO- d_6) δ: 2.38 (3H, s, $C_{12}-CH_3$), 2.45 (3H, s, C_6-CH_3), 4.86 (2H, s, H-2), 5.48 (2H, s, H-10), 6.63 (1H, d, $J=7.5$ Hz, H-13), 7.07 (1H, t, $J=7.6$ Hz, H-15), 7.17 (1H, t, $J=6.8$ Hz, H-14), 7.26-7.33 (2H, m, H-8, H-16), 7.46-7.51 (1H, m, H-7), 8.02 (1H, d, $J=1.8$ Hz, H-5); ^{13}C -NMR (DMSO- d_6) δ: 18.90 ($C_{12}-CH_3$), 20.46 (C_6-CH_3), 44.87 (C-10), 76.12 (C-2), 100.35 (C-3a), 117.13 (C-8, C-4a), 124.23 (C-15), 126.58 (C-6, C-14, C-16), 127.57 (C-13), 130.68 (C-5), 132.82 (C-7), 134.49 (C-11), 135.32 (C-12), 136.40 (C-8a), 171.56 (C-9a), 174.60 (C-4), 191.09 (C-3) .

N-m-Methylbenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (111) 之合成

取化合物 35(2.15g , 0.01mole)和 *m*-methylbenzyl chloride(12.7g , 0.1mole)為原料 , 比照化合物 109 的合成法及處理步驟 , 得化合物 111(1.74g , 54.54 %) , mp:128-130 。光譜數據如下 : MS m/z : 319; IR (KBr) cm^{-1} : 1728.8($C_3=O$), 1605.3 ($C_4=O$); UV λ_{max} nm (MeOH) (log ε): 245 (4.59); 1H -NMR (DMSO- d_6) δ: 2.25 (3H, s, $C_{13}-CH_3$), 2.36 (3H, s, C_6-CH_3), 4.91 (2H, s, H-2), 5.50 (2H, s, H-10), 7.11-7.22 (4H, m, Ar-H), 7.50 (2H, d, $J=1.5$ Hz, H-7, H-8), 7.98 (1H, s, H-5); ^{13}C -NMR (DMSO- d_6) δ: 20.43 (C_6-CH_3), 21.18 ($C_{13}-CH_3$), 46.39 (C-10), 76.14 (C-2), 100.32(C-3a), 117.17(C-8), 123.87(C-4a), 126.59(C-6, C-16), 127.25(C-14),

128.68 (C-15), 129.01 (C-12), 134.39 (C-5), 134.52 (C-7), 135.12 (C-11), 136.27 (C-13), 138.43 (C-8a), 171.52 (C-9a), 174.42 (C-4), 191.25 (C-3).

***N-p*-Methylbenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (112) 之合成**

取化合物 35(2.15g , 0.01mole)和 *p*-methylbenzyl chloride(12.7g , 0.1mole)為原料 , 比照化合物 109 的合成法及處理步驟 , 得化合物 112(2.01 , 63.00 %) , mp : 204-206 。光譜數據如下 : MS *m/z*: 318.7 ; IR (KBr) cm⁻¹: 1721.0 (C₃=O), 1613.0 (C₄=O) ; UV λ_{max} nm (MeOH) (log ε): 244 (4.74); ¹H-NMR (DMSO-*d*₆) δ: 2.23 (3H, s, C₁₄-CH₃), 2.35 (3H, s, C₆-CH₃), 4.90 (2H, s, H-2), 5.49 (2H, s, H-10), 7.13 (2H, d, J=8.0 Hz, H-13, H-15), 7.23 (2H, d, J=7.9 Hz, H-12, H-16), 7.50 (1H, m, H-7, H-8), 7.97 (1H, s, H-5), ¹³C-NMR (DMSO-*d*₆) δ: 20.42 (C₆-CH₃), 20.84 (C₁₄-CH₃), 46.19 (C-10), 76.12 (C-2), 100.28 (C-3a), 117.20 (C-8), 126.57 (C-4a, C-6), 126.89 (C-12, C-16), 129.64 (C-13, C-15), 132.12 (C-5), 134.34 (C-11), 134.50 (C-7), 136.19 (C-14), 137.29 (C-8a), 171.48 (C-9a), 174.36 (C-4), 191.26 (C-3).

***N-m*-Methoxybenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (113) 之合成**

取化合物 35(2.15g , 0.01mole)和 *m*-methoxybenzyl chloride(14.3g , 0.1mole)為原料 , 比照化合物 109 的合成法及處理步驟 , 得化合物 113(2.07g , 61.79 %) , mp : 127-129 。光譜數據如下 : MS *m/z*: 335; IR (KBr) cm⁻¹: 1721.0 (C₃=O), 1605.3 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 244 (4.76); ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, d, J=1.7 Hz, C₆-CH₃), 3.75 (3H, d, J=1.1 Hz, C₁₃-OCH₃), 4.67 (2H, d, J=2.8 Hz, H-2) , 5.42 (2H, s, H-10), 6.73-6.85 (3H, m, H-12, H-14, H-16), 7.19-7.32 (3H, m, H-7, H-8, H-15), 8.16 (1H, s, H-5) ; ¹³C-NMR (DMSO-*d*₆) δ: 20.39 (C₆-CH₃), 46.77 (C-10), 55.07 (OCH₃), 75.29 (C-2), 100.11 (C-3a), 112.14 (C-14), 112.77 (C-8), 115.77 (C-12), 117.93 (C-16), 126.55 (C-4a), 127.42 (C-6), 130.21 (C-15), 134.06 (C-5), 134.84 (C-7), 135.30 (C-11), 135.94 (C-8a), 160.01 (C-13), 172.19 (C-9a), 174.08 (C-4), 190.29 (C-3).

***N-p*-Methoxybenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (114) 之合成**

取化合物 35(2.15g , 0.01mole)和 *p*-methoxybenzyl chloride(14.3g , 0.1mole)為原料 , 比照化合物 109 的合成法及處理步驟 , 得化合物 114(2.24g , 66.86 %) , mp : 228-229 。光譜數據如下 : MS *m/z*: 335; IR (KBr) cm⁻¹: 1713.3 (C₃=O), 1605.3 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 243 (4.89); ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₆-CH₃), 3.69 (3H, d, J=6.6 Hz, C₁₄-OCH₃), 4.91 (2H, s, H-2), 5.47 (2H, s, H-10), 6.88 (2H, d, J=8.7 Hz, H-12, H-16), 7.29 (2H, d, J=8.6 Hz, H-13, H-15), 7.56 (2H, m, H-7, H-8), 7.98 (1H, m, H-5) ; ¹³C-NMR (DMSO-*d*₆) δ: 20.43 (C₆-CH₃) ; 45.88 (C-10), 55.32 (OCH₃), 76.11 (C-2), 100.29 (C-3a), 114.47 (C-13, C-15), 117.26 (C-8), 126.57 (C-4a), 126.66 (C-6), 126.95 (C-11), 128.45 (C-12, C-16), 134.34 (C-5), 134.50 (C-7), 136.16 (C-8a), 159.03 (C-14), 171.48 (C-9a),

174.32 (C-4) , 191.22 (C-3) .

N-o-Fluorobenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione(115)之合成

取化合物 35 (2.15g , 0.01mole) 和 *o*-fluorobenzyl chloride (13.1g , 0.1mole) 為原料 , 比照化合物 109 的合成法及處理得化合物 115 (1.96g , 60.68 %), mp : 242-243 。光譜數據如下 : MS *m/z*: 323; IR (KBr) cm⁻¹ : 1721.0 (C₃=O) , 1605.3 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 244(4.62); ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₆-CH₃), 4.89 (2H, s, H-2), 5.58 (2H, s, H-10), 7.11-7.31 (4H, m, Ar-H), 7.49 (2H, d, J=1.9 Hz, H-7, H-8), 7.99 (1H, s, H-5) ; ¹³C-NMR (DMSO-*d*₆) δ: 20.42 (C₆-CH₃), 40.98 (C-10), 76.20 (C-2), 100.43 (C-3a), 115.73 (C-8), 116.15 (C-13), 116.67 (C-4a), 121.91 (C-11), 122.18 (C-15), 125.19 (C-6), 126.63 (C-14), 128.53 (C-5) , 130.33 (C-16) , 134.52 (C-7) , 134.60 (C-8a) , 136.15 (C-12) , 171.15 (C-9a) , 174.61 (C-4) , 191.13 (C-3) .

N-m-Fluorobenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (116) 之合成

取化合物 35 (2.15g , 0.01mole) 和 *m*-fluorobenzyl chloride (13.1g , 0.1mole) 為原料 , 比照化合物 109 的合成法及處理步得化合物 116 (2.05g , 63.47 %), mp : 243-245 。光譜數據如下 : MS *m/z*: M⁺(323) ; IR (KBr) cm⁻¹ : 1721.0 (C₃=O) , 1605.3 (C₄=O) ; UV λ_{max} nm (MeOH) (log ε): 244 (4.77); ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₆-CH₃), 4.90 (2H, s, H-2), 5.56 (2H, s, H-10) , 7.11-7.40 (4H, m, Ar-H), 7.50 (2H, d, J=1.0 Hz, H-7, H-8), 7.98 (1H, s, H-5) ; ¹³C-NMR (DMSO-*d*₆) δ: 20.40 (C₆-CH₃) , 45.89 (C-10) , 76.17 (C-2) , 100.42 (C-3a) , 114.19 (C-8) , 115.10 (C-14) , 116.99 (C-12) , 122.87 (C-4a) , 126.62 (C-6, C-16) , 131.07 (C-5) , 131.24 (C-5) , 134.41 (C-7) , 134.58 (C-11) , 136.10 (C-8a) , 138.17 (C-13) , 171.52 (C-9a) , 174.46 (C-4) , 191.26 (C-3) .

N-p-Fluorobenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione(117)之合成

取化合物 35 (2.15g , 0.01mole) 和 4-fluorobenzyl chloride (13.1g , 0.1mole) 為原料 , 比照化合物 109 的合成法及處理步驟 , 得化合物 117 (2.58g , 79.88 %), mp: 277-279 。光譜數據如下 : MS *m/z*: 323; IR (KBr) cm⁻¹ : 1721.0 (C₃=O) , 1605.3 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 244(4.83); ¹H-NMR (DMSO-*d*₆) δ: 2.35 (3H, s, C₆-CH₃), 4.90 (2H, s, H-2), 5.53 (2H, s, H-10), 7.18 (2H, m, H-13, H-15), 7.38~7.52 (4H, m, H-7, H-8, H-12, H-16), 7.97 (1H, s, H-5) ; ¹³C-NMR (DMSO-*d*₆) δ: 20.41 (C₆-CH₃), 45.71 (C-10), 76.15 (C-2), 100.37 (C-3a), 115.70 (C-8), 116.12 (C-4a), 117.08 (C-6) , 126.64 (C-13, C-15) , 129.09 (C-5) , 129.25 (C-11) , 131.34 (C-12, C-16), 134.40 (C-7), 134.58 (C-8a), 136.09 (C-14), 171.50 (C-9a), 174.40 (C-4) , 191.23 (C-3) .

N-o-Chlorobenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (118) 之合成

取化合物 35 (2.15g , 0.01mole) 和 *o*-chlorobenzyl chloride (14.7g , 0.1mole) 為原料 , 比照化合物 109 的合成法及處理步驟 , 得化合物 118 (1.88g , 55.46 %) , mp: 265-267 。光譜數據如下 : MS *m/z*: 339; IR (KBr) cm⁻¹: 1713.3(C₃=O), 1613.0 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 244(4.67); ¹H-NMR (DMSO-*d*₆) δ: 2.37 (3H, s, C₆-CH₃), 4.87 (2H, s, H-2), 5.55 (2H, s, H-10), 6.69 (1H, d, J=7.5 Hz, H-16), 7.19-7.38 (3H, m, Ar-H), 7.48-7.59 (2H, m, H-7, H-8), 8.02 (1H, m, H-5); ¹³C-NMR (DMSO-*d*₆) δ: 20.44 (C₆-CH₃), 44.73(C-10), 76.25(C-2), 100.51(C-3a), 116.68 (C-8), 126.56 (C-4a), 126.72 (C-15), 127.30 (C-6), 128.09 (C-14), 129.77 (C-13), 130.01 (C-5), 131.67 (C-16), 132.17 (C-12), 134.63 (C-7), 134.74 (C-11), 136.23 (C-8a), 171.57 (C-9a), 174.65 (C-4), 191.13 (C-3) .

N-m-Chlorobenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (119) 之合成

取化合物 35 (2.15g , 0.01mole) 和 *m*-chlorobenzyl chloride (14.7g , 0.1mole) 為原料 , 比照化合物 109 的合成法及處理步驟 , 得化合物 119 (1.98g , 58.41 %) , mp: 226-228 。光譜數據如下 : MS *m/z*: 339; IR (KBr) cm⁻¹: 1713.3(C₃=O), 1605.3 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 245(4.62); ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₆-CH₃), 4.90 (2H, s, H-2), 5.56 (2H, s, H-10), 7.34-7.50 (6H, m, Ar-H, H-7, H-8), 7.99 (1H, s, H-5); ¹H-NMR (DMSO-*d*₆) δ: 20.43 (C₆-CH₃), 45.83 (C-10), 76.20 (C-2), 100.47(C-3a), 117.01(C-8), 125.47(C-4a), 126.65(C-6, C-16), 126.84 (C-14), 128.07 (C-12), 131.00 (C-15), 133.77 (C-5), 134.47 (C-13), 134.63 (C-7), 136.11 (C-11), 137.79 (C-8a), 171.54 (C-9a), 174.50 (C-4), 191.28 (C-3) .

N-p-Chlorobenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (120) 之合成

取化合物 35 (2.15g , 0.01mole) 和 *p*-chlorobenzyl chloride (14.7g , 0.1mole) 為原料 , 比照化合物 109 的合成法及處理步驟 , 得化合物 120 (2.12g , 62.54 %) , mp: 262-264 。光譜數據如下 : MS *m/z*: 339; IR (KBr) cm⁻¹: 1721.0(C₃=O), 1605.3 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 245(4.67); ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₆-CH₃), 4.90 (2H, s, H-2), 5.55 (2H, s, H-10), 7.38 (4H, s, Ar-H), 7.50 (2H, s, H-7, H-8), 7.98 (1H, s, H-5); ¹³C-NMR (DMSO-*d*₆) δ: 20.42 (C₆-CH₃), 45.76 (C-10), 76.17 (C-2), 100.38 (C-3a), 117.06 (C-8), 126.64 (C-4a, C-6), 128.89 (C-13, C-15), 129.04(C-12, C-16), 132.64(C-5), 134.26(C-14), 134.44(C-11), 134.63 (C-7), 136.09 (C-8a), 171.51 (C-9a), 174.44 (C-4), 191.22 (C-3) .

N-o-Nitrobenzyl-6-methyl-2,3,4,9-tetrahydrofuro-[2,3-*b*]quinolin-3,4-dione (121) 之合成

取化合物 35 (2.15g , 0.01mole) 和 *o*-nitrobenzyl chloride (15.8g , 0.1mole) 為原料 , 比照化合物 109 的合成法及處理步驟 , 得化合物 121(1.78g , 50.86 %), mp:275-280 。光譜數據如下 :MS *m/z*: 350; IR (KBr) cm⁻¹: 1721.0(C₃=O), 1613.0 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 244 (4.79); ¹H-NMR (DMSO-*d*₆) δ: 2.38 (3H, s, C₆-CH₃), 4.84 (2H, s, H-2), 5.91 (2H, s, H-10), 7.03 (1H, m, H-16), 7.46 (2H, m, H-7, H-8), 7.57-7.62 (2H, m, H-14, H-15), 8.02 (1H, s, H-5), 8.26 (1H, m, H-13) ; ¹³C-NMR (DMSO-*d*₆) δ: 20.45(C₆-CH₃), 45.14(C-10), 76.23(C-2), 100.59(C-3a), 117.18(C-8), 125.99(C-4a), 126.65(C-13), 127.16(C-6), 129.26(C-14, C-16), 130.57 (C-5) , 134.53 (C-11) , 134.77 (C-15) , 134.98 (C-7) , 136.27 (C-8a) , 147.34 (C-12) , 171.56 (C-9a) , 174.86 (C-4) , 191.16 (C-3) .

N-m-Nitrobenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (122) 之合成

取化合物 35 (2.15g , 0.01mole) 和 *m*-nitrobenzyl chloride (15.8g , 0.1mole) 為原料 , 比照化合物 109 的合成法及處理步驟 , 得化合物 122(2.02g , 57.71 %), mp:272-274 。光譜數據如下 :MS *m/z*: 350; IR (KBr) cm⁻¹: 1713.3(C₃=O), 1605.3 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 243(4.94); ¹H-NMR (DMSO-*d*₆) δ: 2.37 (3H, s, C₆-CH₃), 4.90 (2H, s, H-2), 5.71 (2H, s, H-10), 7.51-7.70 (4H, m, H-7, H-8, H-15, H-16), 8.00 (1H, s, H-5), 8.30 (1H, s, H-12) ; ¹³C-NMR (DMSO-*d*₆) δ: 20.42 (C₆-CH₃), 45.71 (C-10), 76.25 (C-2), 100.53 (C-3a), 116.97 (C-8), 122.05 (C-4a, C-14) , 123.00 (C-12) , 126.75 (C-6) , 130.72 (C-15) , 133.23 (C-5) , 134.55 (C-7) , 134.73 (C-16) , 136.07 (C-11) , 137.61 (C-8a) , 148.29 (C-13) , 171.56 (C-9a) , 174.61 (C-4) , 191.25 (C-3) .

N-p-Nitrobenzyl-6-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (123) 之合成

取化合物 35 (2.15g , 0.01mole) 和 *p*-nitrobenzyl chloride (15.8g , 0.1mole) 為原料 , 比照化合物 109 的合成法及處理步驟得化合物 123 (2.25g , 64.28 %), mp:292-294 。光譜數據如下 :MS *m/z*: 350; IR (KBr) cm⁻¹: 1721.0(C₃=O), 1605.3 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 245(4.69); ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₆-CH₃), 4.89 (2H, s, H-2), 5.71 (2H, s, H-10), 7.48 (2H, s, H-7, H-8), 7.60 (2H, d, J=8.6Hz, H-12, H-16), 8.00 (1H, s, H-5), 8.18 (2H, d, J=8.7Hz, H-13, H-15) ; ¹³C-NMR (DMSO-*d*₆) δ: 20.42(C₆-CH₃), 45.97(C-10), 76.24(C-2), 100.49(C-3a), 116.91 (C-8) , 124.15 (C-4a, C-13, C-15) , 126.73 (C₆), 128.12 (C-12, C-16) , 134.51 (C-5) , 134.75 (C-7) , 136.07 (C-8a) , 143.00 (C-11) , 147.29 (C-14) , 171.55 (C-9a) , 174.56 (C-4) , 191.20 (C-3) .