

(四) *N*-Substituted benzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]-quinolin-3,4-dione (53-67) 之合成

N-Benzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (53) 之合成

取化合物28 (2.15g , 0.01mole) 懸著於DMF 30 ml中 , 加入無水K₂CO₃ (1.38 g , 0.01 mole)加熱(約70-80)使之溶解 , 加入benzyl chloride(12.6ml , 0.1mole), 反應1小時後加冰水 , 以CHCl₃萃取 , 取CHCl₃層 , 以無水MgSO₄乾燥 , 減壓濃縮後 , 收集沉殿物以短程矽膠管柱層析 (CHCl₃/EtOH) 沖提 , 再以MeOH及CHCl₃做再結晶 , 得白色棉絮狀結晶 , 為化合物53 (2.42g , 79.3 %) , mp : 264-266 。光譜數據如下: MS m/z: 305; IR (KBr) cm⁻¹: 1716.8 (C₃=O), 1612.6 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 247 (4.58); ¹H-NMR (DMSO-*d*₆) δ: 2.35 (3H, s, C₇-CH₃), 4.90 (2H, s, H-2), 5.55 (2H, s, H-10), 7.21-7.36 (6H, m, H-6, Ar-H), 7.49 (1H, s, H-8), 8.07 (1H, d, J=8.0 Hz, H-5); ¹³C-NMR (DMSO-*d*₆) δ: 2.49 (C₇-CH₃), 46.95 (C-10), 75.49 (C-2), 100.30 (C-3a), 116.09 (C-8), 126.27 (C-6), 127.82 (C-14), 128.28 (C-5), 129.28 (C-13, C-15), 133.71 (C-11), 138.38 (C-7), 144.40 (C-8a), 172.25 (C-9a), 174.69 (C-4), 190.41 (C-3) .

N-o-Methylbenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (54) 之合成

取化合物 28 (2.15g , 0.01mole) 和 2-methylbenzyl chloride (14ml , 0.1mole) 為原料 , 比照化合物 53 的合成法及處理步驟 , 即可得化合物 54 (2.53g , 19.3 %) , mp : 225 。光譜數據如下 : MS m/z: 319; IR (KBr) cm⁻¹: 1714.5 (C₃=O), 1619.7 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 248 (5.96); ¹H-NMR (DMSO-*d*₆) δ: 2.33 (3H, s, C₇-CH₃), 2.45 (2H, s, C₁₂-CH₃), 4.84 (2H, s, H-2), 5.48 (2H, s, H-10), 6.64 (1H, d, J=7.5Hz, H-6), 7.05-7.29 (5H, m, H-8, Ar-H), 8.12 (1H, d, J=8.5Hz, H-5); ¹³C-NMR (DMSO-*d*₆) δ: 18.90 (C₁₂-CH₃), 21.67 (C₇-CH₃), 44.83 (C-10), 76.12 (C-2), 100.32 (C-3a), 116.84 (C-8), 124.10 (C-6), 124.49 (C-4a), 126.28 (C-15), 126.58 (C-14), 126.87 (C-13), 127.58 (C-16), 130.68 (C-5), 132.89 (C-11), 135.3 (C-12), 136.68 (C-7), 144.26 (C-8a), 171.51 (C-9a), 174.98 (C-4), 191.10 (C-3) .

N-m-Methylbenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione(55) 之合成

取化合物 28 (2.15g , 0.01mole) 和 3-methylbenzyl chloride (14ml , 0.1mole) 為原料 , 比照化合物 53 的合成法及處理步驟 , 即可得化合物 55 (1.71 g , 53.6 %) , mp : 265 。光譜數據如下 : MS m/z: 319; IR (KBr) cm⁻¹: 1706.6 (C₃=O), 1607.9 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 249 (5.79); ¹H-NMR (DMSO-*d*₆) δ: 2.25 (3H, s, C₇-CH₃), 2.35 (3H, s, C₁₃-CH₃), 4.89 (2H, s, H-2), 5.50 (2H, s, H-10), 7.06~7.23 (5H, m, H-6, Ar-H), 7.45 (1H, s, H-8), 8.07 (1H, d, J=8.03Hz, H-5); ¹³C-NMR (DMSO-*d*₆) δ: 21.17 (C₇-CH₃), 21.69 (C₁₃-CH₃), 46.33 (C-10), 76.13 (C-2), 100.27 (C-3a), 117.03 (C-8), 123.88 (C-6), 124.56 (C-4a), 126.11

(C-16), 126.82 (C-14), 127.27 (C-15), 128.68 (C-5), 135.15 (C-11), 138.44 (C-13), 138.49 (C-7), 144.04 (C-8a), 171.43 (C-9a), 171.82 (C-4), 191.16 (C-3).

***N-p*-Methylbenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (56) 之合成**

取化合物 **28** (2.15g , 0.01mole) 和 4-methylbenzyl chloride (14ml , 0.1mole) 為原料 , 比照化合物 **53** 的合成法及處理步驟 , 即可得

化合物 **56** (20.05 g , 64.26 %) , mp : 240~241 。光譜數據如下 : MS *m/z*: 319 ; IR (KBr) cm⁻¹ : 1706.6(C₃=O), 1611.8(C₄=O); UV λ_{max} nm (MeOH) (log ε): 249 (5.79); ¹H-NMR (DMSO-*d*₆) δ: 2.23 (3H, s, C₇-CH₃), 2.35 (3H, s, C₁₄-CH₃), 4.89 (2H, s, H-2), 5.49 (2H, s, H-10), 7.12~7.26 (5H, m, 6-H, Ar-H), 7.47 (1H, s, H-8), 8.06 (1H, d, J=8.06 Hz, H-5); ¹³C-NMR (DMSO-*d*₆) δ: 20.85(C₁₄-CH₃), 21.70(C₇-CH₃), 46.11 (C-10), 76.12 (C-2), 100.24 (C-3a), 117.10 (C-8), 124.53 (C-4a), 126.13 (C-6), 126.90 (C-13, C-15, C-15), 129.67 (C-12, C-16), 1132.14 (C-11), 137.30 (C-14), 138.43 (C-7), 144.08 (C-8a), 171.45 (C-9a), 174.76 (C-4), 191.21 (C-3) .

***N-m*-Methoxybenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (57) 之合成**

取化合物 **28** (2.15g , 0.01mole) 和 3-methoxybenzyl chloride (15.6 ml , 0.1 mole) 為原料 , 比照化合物 **53** 的合成法及處理步驟 , 即可得化合物 **57** (2.58 g , 77.01 %), mp : 215~217 。光譜數據如下 : MS *m/z*: 335 ; IR (KBr) cm⁻¹ : 1718.4(C₃=O), 1611.8 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 248 (5.32); ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₇-CH₃), 3.71 (3H, s, O-CH₃), 4.90 (2H, s, H-2), 5.51 (2H, s, H-10), 6.84~7.25 (5H, m, H-6, H-8, Ar-H), 7.48 (1H, s, H-12), 8.07 (1H, d, J=8.06 Hz, H-5); ¹³C-NMR (DMSO-*d*₆) δ: 21.69 (C₇-CH₃), 48.82 (C-10), 55.30 (OCH₃), 76.15 (C-2), 100.25 (C-3a), 112.93 (C-14), 113.10 (C-8), 117.03 (C-12), 118.72 (C-6), 124.49 (C-4a), 126.15 (C-16), 126.82 (C-15), 130.33 (C-5), 136.78 (C-11), 138.48 (C-7), 144.09 (C-8a), 159.79 (C-13), 171.44 (C-9a), 174.79 (C-4), 191.19 (C-3) .

***N-p*-Methoxybenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (58) 之合成**

取化合物 **28** (2.15g , 0.01mole) 和 4-methoxybenzyl chloride (15.6 ml , 0.1 mole) 為原料 , 比照化合物 **53** 的合成法及處理步驟 , 即可得化合物 **58** (2.54 g , 75.82 %), mp : 275 。光譜數據如下 : MS *m/z*: 335; IR (KBr) cm⁻¹ : 1702.6(C₃=O), 1607.9 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 248 (5.65); ¹H-NMR (DMSO-*d*₆) δ: 2.37 (3H, s, C₇-CH₃), 3.70 (3H, s, O-CH₃), 4.90 (2H, s, H-2), 5.47 (2H, s, H-10), 6.89~7.52 (6H, m, H-6, H-8, Ar-H), 8.06 (1H, d, J=8.4 Hz, H-5); ¹³C-NMR (CF₃COOD) δ: 21.15 (C₇-CH₃), 48.61 (C-10), 55.17 (OCH₃), 78.15 (C-2), 99.75 (C-3a), 155.19 (C-13, C-15), 161.31 (C-8), 125.06 (C-4a), 127.19 (C-6), 128.10 (C-12,

C-16) , 129.92 (C-5) , 140.29 (C-7) , 153.30 (C-8a) , 158.64 (C-14) , 168.34 (C-9a) , 171.27 (C-4) , 194.36 (C-3) .

N-o-Fluorobenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-b]quinolin-3,4-dione (59) 之合成

取化合物 **28**(2.15g , 0.01mole)和 2-fluorobenzyl chloride(14.4 ml , 0.1 mole)為原料 , 比照化合物 **53** 的合成法及處理步驟 , 即可得化合物 **59** (2.66 g , 82.35 %), mp : 250 。光譜數據如下 : MS *m/z*: 323 ; IR (KBr) cm⁻¹ : 1710.5 (C₃=O) , 1619.7 (C₄=O) ; UV λ_{max} nm (MeOH) (log ε): 248 (5.81); ¹H-NMR (DMSO-*d*₆) δ: 2.35 (3H, s, C₇-CH₃) , 4.88 (2H, s, H-2) , 5.57 (2H, s, H-10) , 7.12-7.42 (6H,m,6H, 8-H,Ar-H) , 8.08 (1H, d, J=8.04 Hz, H-5) ; ¹³C-NMR (DMSO-*d*₆) δ: 21.81 (C₇-CH₃) , 41.20 (C-10) , 76.34 (C-2) , 100.52 (C-3a) , 116.28 (C-8) , 116.72 (C-13) , 122.03 (C-4a) , 122.30 (C-6) , 124.56 (C-11) , 125.37 (C-15) , 126.44 (C-14) , 127.07 (C-5) , 128.57 (C-16) , 130.36 (C-7) , 138.52 (C-8a) , 144.49 (C-12) , 171.69 (C-9a) , 175.11 (C-4) , 191.34 (C-3) .

N-m-Fluorobenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-b]quinolin-3,4-dione (60) 之合成

取化合物 **28**(2.15g , 0.01mole)和 3-fluorobenzyl chloride(14.4 ml , 0.1 mole)為原料 , 比照化合物 **53** 的合成法及處理步驟 , 即可得化合物 **60** (1.65 g , 51.08 %), mp : 235~237 。光譜數據如下 : MS *m/z*: 323; IR (KBr) cm⁻¹ : 1726.3(C₃=O), 1635.5 (C₄=O) ; UV λ_{max} nm (MeOH) (log ε): 248 (5.69) ; ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₇-CH₃) , 4.89 (2H, s, H-2) , 5.56 (2H, s, H-10) , 7.12-7.45 (6H,m,6H, H-8,Ar-H) , 8.07 (1H, d, J=8.04 Hz, H-5) ; ¹³C-NMR (DMSO-*d*₆) δ: 21.69 (C₇-CH₃) , 45.85 (C-10) , 76.20 (C-2) , 100.40 (C-3a) , 115.14 (C-8) , 116.90 (C-14) , 122.88 (C-12) , 124.55 (C-4a) , 126.21 (C-6) , 126.90 (C-16) , 131.10 (C-5) , 131.27 (C-15) , 138.07 (C-11) , 138.38 (C-7) , 144.19 (C-8a) , 165.03 (C-13) , 171.48 (C-9a) , 174.88 (C-4) , 191.26 (C-3) .

N-p-Fluorobenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-b]quinolin-3,4-dione (61) 之合成

取化合物 **28**(2.15g , 0.01mole)和 4-fluorobenzyl chloride(14.4 ml , 0.1 mole)為原料 , 比照化合物 **53** 的合成法及處理步驟 , 即可得化合物 **61**(2.09 g , 64.7 %), mp : 250 。光譜數據如下 : MS *m/z*: 323.2; IR (KBr) cm⁻¹ : 1710.5(C₃=O) , 1611.8 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 248 (5.66); ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₇-CH₃) , 4.90 (2H, s, H-2) , 5.53 (2H, s, H-10) , 7.13-7.48 (6H,m,H-6,H-8,Ar-H) , 8.07 (1H, d, J=8.04Hz, H-5) ; ¹³C-NMR (DMSO-*d*₆) δ: 21.69 (C₇-CH₃) , 45.65 (C-10) , 76.17 (C-2) , 100.34 (C-3a) , 115.74 (C-13) , 116.17 (C-15) , 117.00 (C-8) , 124.55 (C-4a) , 126.20 (C-6) , 126.90 (C-5) , 129.13 (C-12) , 129.29 (C-16) , 131.38 (C-11) , 138.34 (C-7) , 144.20 (C-8a) , 159.42 (C-14) , 171.48 (C-9a) , 174.81 (C-4) , 191.24 (C-3) .

*N-o-Chlorobenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (62)* 之合成

取化合物 **28** (2.15g , 0.01mole) 和 2-chlorobenzyl chloride (16 ml , 0.1 mole) 為原料 , 比照化合物 **53** 的合成法及處理步驟 , 即可得化合物 **62** (1.56 g , 46.01 %), mp : 250~251 。光譜數據如下 : MS *m/z*: 339; IR (KBr) cm⁻¹: 1718.4(C₃=O), 1619.7 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 248 (5.78); ¹H-NMR (DMSO-*d*₆) δ: 2.34(3H, s, C₇-CH₃), 4.85(2H, s, H-2), 5.55 (2H, s, H-10), 6.98(1H, d, J=7.8Hz, H-6), 7.24-7.35 (4H, m, H-8, Ar-H), 7.57 (1H, d, J=7.8 Hz, H-13), 8.10 (1H, d, J=8.0 Hz, H-5); ¹³C-NMR (DMSO-*d*₆) δ: 21.64 (C₇-CH₃), 44.68 (C-10), 76.25 (C-2), 100.47 (C-3a), 116.53 (C-8), 124.48 (C-4a), 126.35 (C-6), 126.96 (C-15), 127.26 (C-5), 128.14 (C-14), 129.76 (C-16), 130.00 (C-13), 131.69 (C-12), 132.28 (C-11), 138.54 (C-7), 144.41 (C-8a), 171.51 (C-9a), 175.03 (C-4) 191.10 (C-3) .

*N-m-Chlorobenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione(63)* 之合成

取化合物 **28** (2.31 g , 0.01mole) 和 3-chlorobenzyl chloride (16 ml , 0.1 mole) 為原料 , 比照化合物 **53** 的合成法及處理步驟 , 即可得化合物 **63** (1.51 g , 44.54 %), mp : 245-246 。光譜數據如下 : MS *m/z*: 339; IR (KBr) cm⁻¹: 1710.5(C₃=O), 1623.7 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 248 (5.78); ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₇-CH₃), 4.89 (2H, s, H-2), 5.56 (2H, s, H-10), 7.13-7.49 (6H, m, H-6, H-8, Ar-H) 8.08(1H, d, J=8.0 Hz, H-5) ¹³C-NMR (DMSO-*d*₆) δ: 21.68 (C₇-CH₃), 44.78 (C-10), 76.20 (C-2), 100.42 (C-3a), 116.88 (C-8), 124.54 (C-4a), 125.43 (C-6), 126.24 (C-16), 126.81 (C-5, C-12), 128.07 (C-14), 131.02 (C-15), 133.77 (C-13), 137.80 (C-11), 132.38 (C-7), 144.23 (C-8a), 171.50 (C-9a), 174.92 (C-4), 191.25 (C-3) .

*N-p-Chlorobenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (64)* 之合成

取化合物 **28** (2.31 g , 0.01mole) 和 4-chlorobenzyl chloride (16 ml , 0.1 mole) 為原料 , 比照化合物 **53** 的合成法及處理步驟 , 即可得化合物 **64** (2.09 g , 61.65 %), mp : 260-261 。光譜數據如下 : MS *m/z*: 339; IR (KBr) cm⁻¹: 1710.5(C₃=O), 1615.8 (C₄=O); UV λ_{max} nm (MeOH) (log ε): 249 (5.63); ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₇-CH₃), 4.89 (2H, s, H-2), 5.55 (2H, s, H-10), 7.22 (1H, d, J=8.08Hz, H-6), 7.40 (4H, s, Ar-H), 7.46 (1H, s, H-8), 8.07 (1H, d, J=8.02Hz, H-5); ¹³C-NMR (DMSO-*d*₆) δ: 21.68 (C₇-CH₃), 45.69 (C-10), 76.18 (C-2), 100.34 (C-3a), 116.94 (C-8), 124.55 (C-4a), 126.20 (C-6), 126.89 (C-5), 128.90 (C-13, C-15), 129.07 (C-12, C-16), 132.64 (C-14), 134.31 (C-11), 138.34 (C-7), 144.20 (C-8a), 171.44 (C-9a), 174.83 (C-4), 191.17 (C-3) .

*N-o-Nitrobenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (65)* 之合成

取化合物 **28**(2.15 g , 0.01mole)和 2-nitrobenzyl chloride (17.2 ml , 0.1 mole)為原料, 比照化合物 **53** 的合成法及處理步驟, 即可得化合物 **65**(1.95 g , 55.7 %), mp:293~295 。光譜數據如下: MS *m/z*: 350; IR (KBr) cm⁻¹: 1710.5(C₃=O), 1611.8 (C₄=O) ; UV λ_{max} nm (MeOH) (log ε): 248 (5.78) ; ¹H-NMR (DMSO-*d*₆) δ: 2.29 (3H, s, C₇-CH₃), 4.81 (2H, s, H-2), 5.89 (2H, s, H-10), 7.04 (1H, d, J=8.0 Hz, H-14), 7.24(1H, d, J=8.1Hz, H-6), 7.41(1H, s, H-8), 7.58-7.62(2H, m, H-15, H-16), 8.08(1H, d, J=8.02 Hz, H-5), 8.27(1H, d, J=8.06 Hz, H-13); ¹³C-NMR (DMSO-*d*₆) δ: 21.48 (C₇-CH₃), 45.16 (C-10), 76.23 (C-2), 100.52 (C-3a), 116.74 (C-8), 124.37 (C-4a), 126.02 (C-6), 126.44 (C-13), 126.85 (C-14), 127.02 (C-5), 129.31 (C-16), 130.57 (C-11), 135.07 (C-15), 138.50 (C-7), 144.66 (C-8a), 147.25 (C-12), 171.77 (C-9a), 175.19 (C-4), 191.24 (C-3) .

*N-m-Nitrobenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (66)* 之合成

取化合物 **28**(2.15 g , 0.01mole)和 3-nitrobenzyl chloride(17.2 ml , 0.1 mole)為原料, 比照化合物 **53** 的合成法及處理步驟, 即可得化合物 **66**(2.45 g , 70 %), mp : > 300 。光譜數據如下: MS *m/z*: 350; IR (KBr) cm⁻¹: 1718.4(C₃=O), 1611.8 (C₄=O) ; UV λ_{max} nm (MeOH) (log ε): 248 (5.50) ; ¹H-NMR (DMSO-*d*₆) δ: 2.36 (3H, s, C₇-CH₃), 4.89 (2H, s, H-2), 5.71 (2H, s, H-10), 7.25-7.72 (4H,m, H-6, H-8, H-15, H-16), 8.09 (1H, d, J=8.2 Hz, H-5) 8.14-8.31 (2H, m, H-12,H-14); ¹³C-NMR (CF₃COOD) δ : 21.32 (C₇-CH₃), 48.11 (C-10), 78.56 (C-2), 100.29 (C-3a), 117.31 (C-8), 121.33 (C-6), 121.45 (C-4a), 124.37 (C-14), 127.79 (C-12), 130.27 (C-15), 131.04 (C-15), 134.31 (C-16), 134.17 (C-11), 144.11 (C-7), 148.63 (C-8a), 153.60 (C-13), 169.61 (C-9a), 172.21 (C-4), 194.26 (C-3) .

*N-p-Nitrobenzyl-7-methyl-2,3,4,9-tetrahydrofuro[2,3-*b*]quinolin-3,4-dione (67)* 之合成

取化合物 **28**(2.15 g , 0.01mole)和 4-nitrobenzyl chloride(17.2 ml , 0.1 mole)為原料, 比照化合物 **53** 的合成法及處理步驟, 即可得化合物 **67**(1.66 g , 47.42 %) mp : > 300 .光譜數據如下: MS *m/z*: 350; IR (KBr) cm⁻¹: 1722.4(C₃=O), 1619.7 (C₄=O) ; UV λ_{max} nm (MeOH) (log ε): 248 (5.79) ; ¹H-NMR (DMSO-*d*₆) δ: 2.35 (3H, s, C₇-CH₃), 4.89 (2H, s, H-2), 5.72(2H, s, H-10), 7.25(1H, d, J=8.08 Hz, H-6), 7.44(1H, s, H-8), 7.62(1H, d, J=8.38 Hz, H-5), 8.10(1H,d,J=8.02Hz,H-12,H-16), 8.19(1H, d, J=8.0 Hz, H-5); ¹³C-NMR (DMSO-*d*₆) δ: 21.66(C₇-CH₃), 45.93(C-10), 76.25 (C-2), 100.44 (C-3a), 116.77 (C-8), 124.17 (C-13, C-15), 124.58 (C-4a), 126.30 (C-6), 126.98 (C-5), 128.11 (C-12, C-16), 138.35 (C-11), 143.03 (C-7), 144.33 (C-8a), 147.29 (C-14), 171.47 (C-9a), 174.97 (C-4), 191.11 (C-3) .