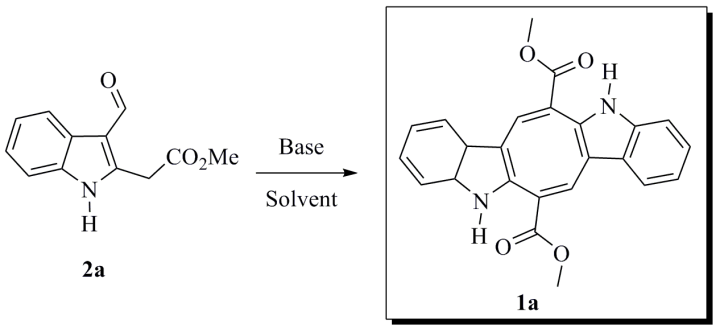
Supplementary Information

Scheme S1. Optimization of the reaction conditions for the synthesis of **1a**.



2a **1a**

Table S1. Optimization of the reaction conditions for the synthesis of **1a**.

Entry	Base	Concentration (M)	Solvent	Temperature (℃)	Time (h)	Yield (%)
1	Diethylamine Piperidine	0.01	Methanol	reflux	48	2[9]
2	Diethylamine Piperidine	0.01	Xylene	reflux	3	12
3	Diethylamine Piperidine	0.023	Xylene	reflux	3	32
4	Diethylamine Piperidine	0.038	Xylene	reflux	3	31
5	Diethylamine Piperidine	0.076	Xylene	reflux	4	32
6	Diethylamine Piperidine	0.230	Xylene	reflux	6	15
7	Diethylamine Piperidine	0.076	Toluene	reflux	4.5	24
8	Diethylamine Piperidine	0.230	Toluene	reflux	7.5	12

Anti-Tuberculosis Activity

Figure S1. IC_{50} values of compounds **1a**, **1b**, **1d** and **1e** against the *M. tuberculosis* strain H37Rv. Values are the means ± S.E.M., *n* = 5.

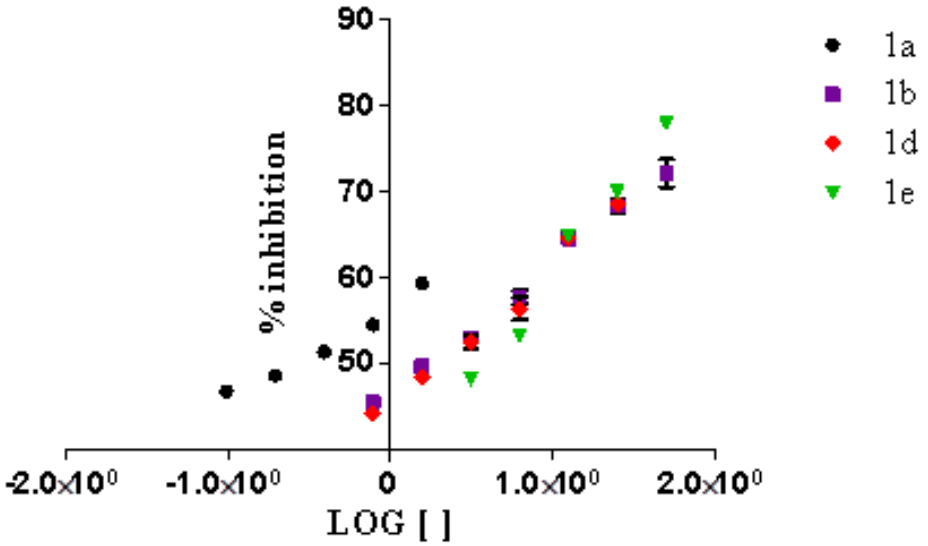


Figure S2. Concentration-response curve of compound **1a** against the *M. tuberculosis* strain H37Rv. This compound was tested at a concentration of 50–1.56 μM. Values are the means ± S.E.M., *n* = 5.

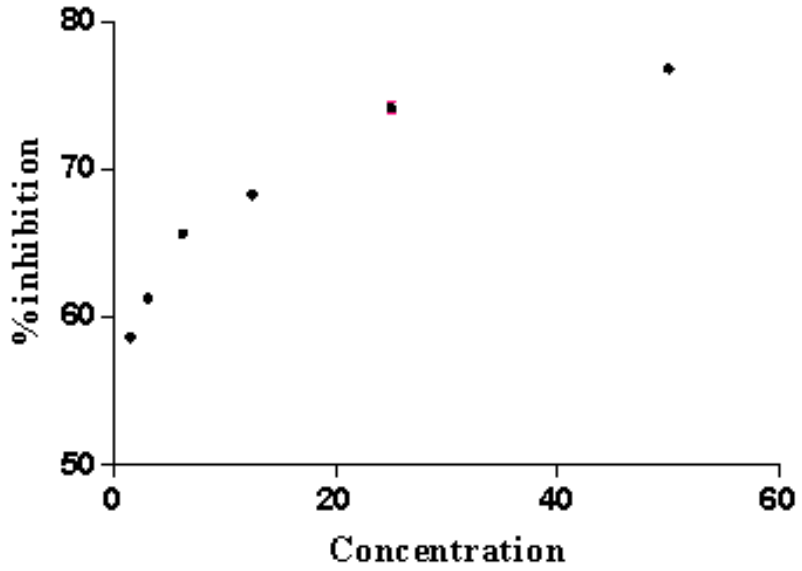


Figure S3. IC_{50} values of rifampin against the *M. tuberculosis* strain H37Rv. Values are the means ± S.E.M., $n = 5$.

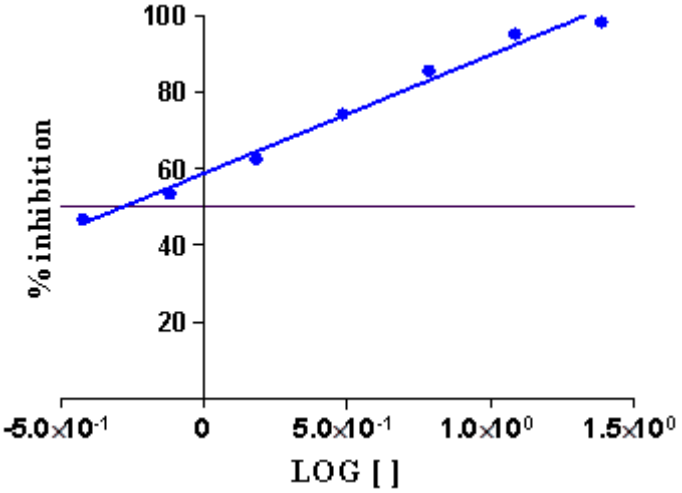


Figure S4. ^{1}H NMR spectrum of the Caulerpin (**1a**).

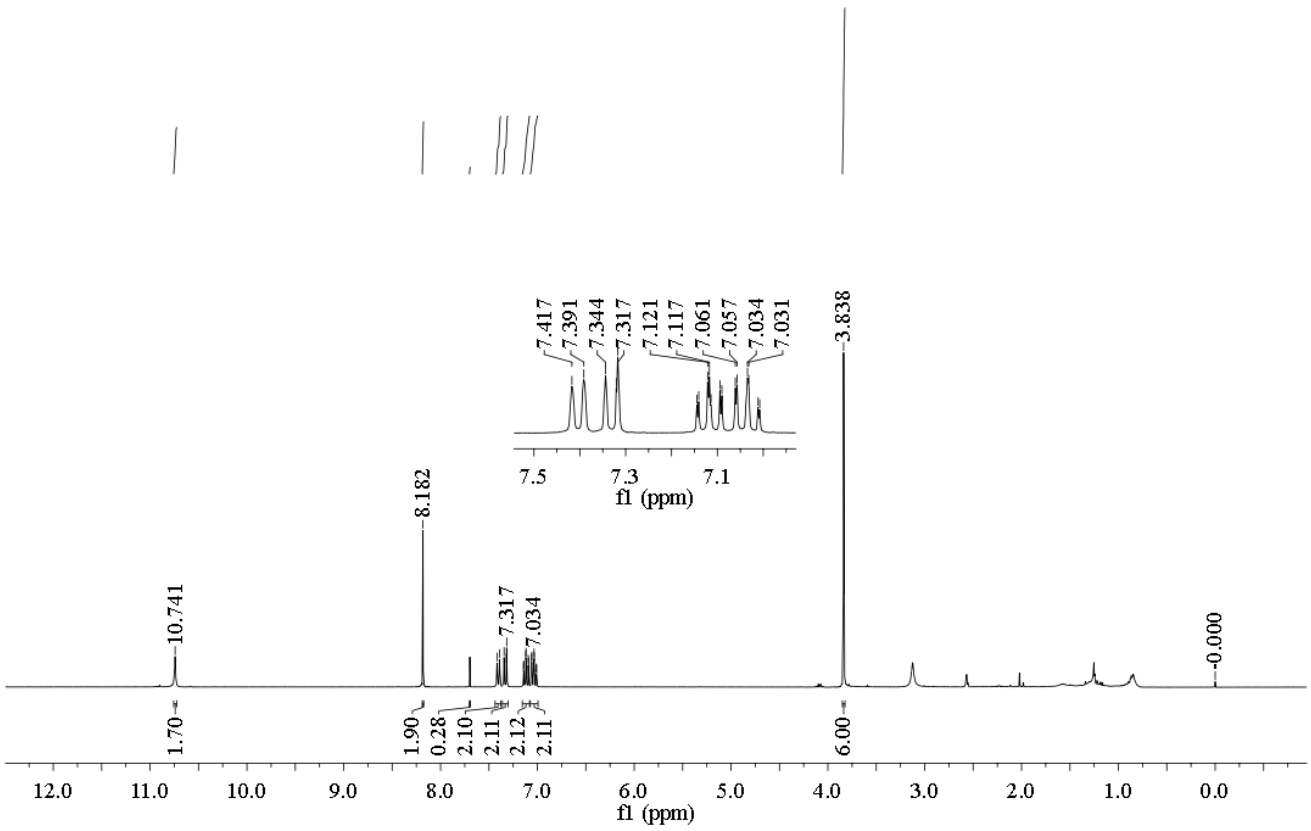


Figure S5. ^{13}C NMR spectrum of the Caulerpin (**1a**).

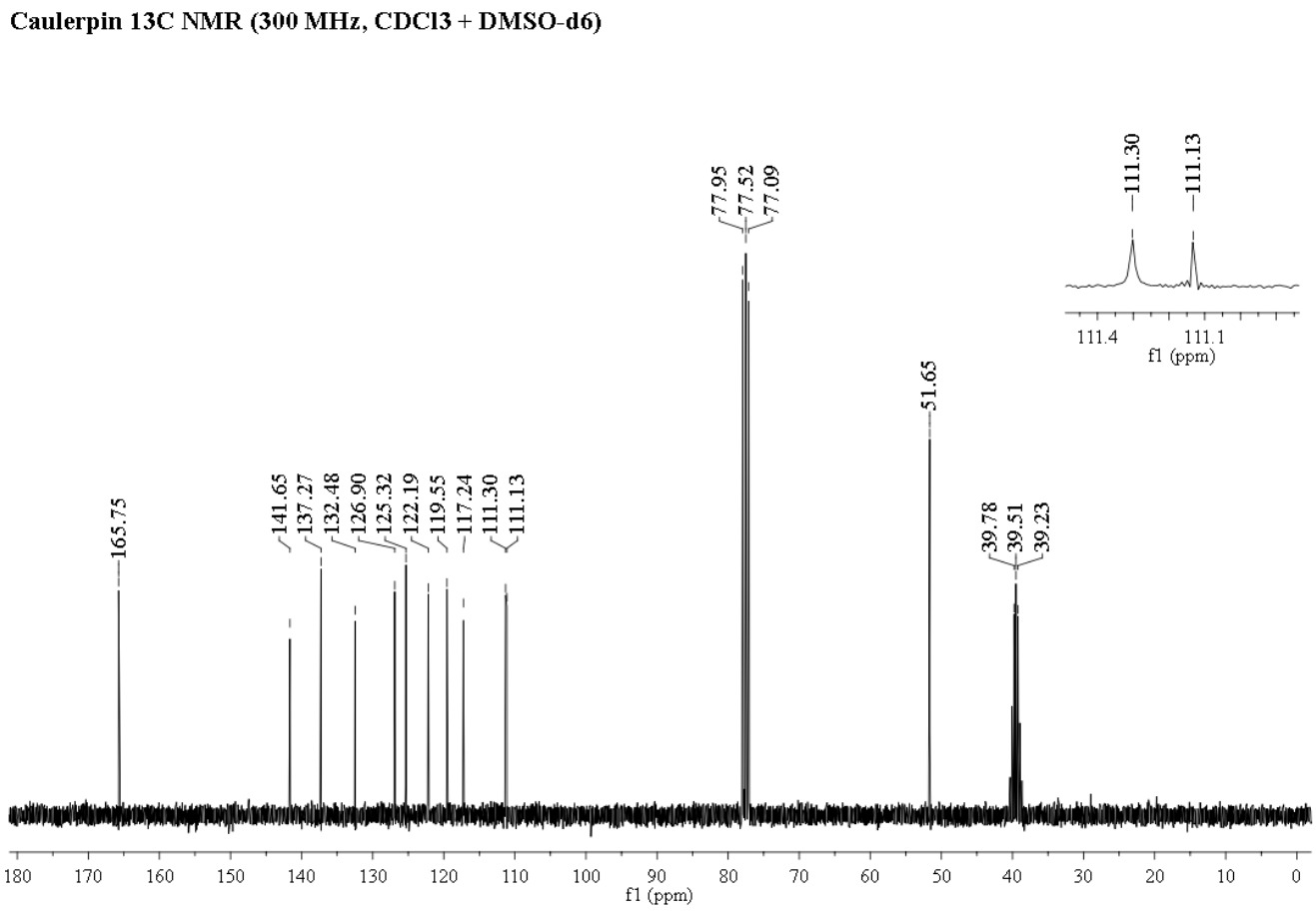


Figure S6. Elemental composition of the Caulerpin (**1a**).

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[ Elemental Composition ]                                    Page: 1
Data  : Dr-Roberto-Martinez003          Date : 02-Dec-2013 13:24
Sample: 2616    la
Note : -luis-velasco
Inlet : Direct                          Ion Mode : FAB+
RT : 1.04 min                           Scan#: (2,7)
Elements : C 40/0, H 49/0, O 5/0, N 3/1
Mass Tolerance      : 1000ppm, 1mmu if m/z > 1
Unsaturation (U.S.) : -0.5 - 20.0

Observed m/z   Int%
  398.1266     100.0
Estimated m/z  Error[ppm]  U.S.     C     H     O     N
  398.1267       -0.3      17.0    24    18     4     2
```

Figure S7. Mass spectrum of the Caulerpin (**1a**).

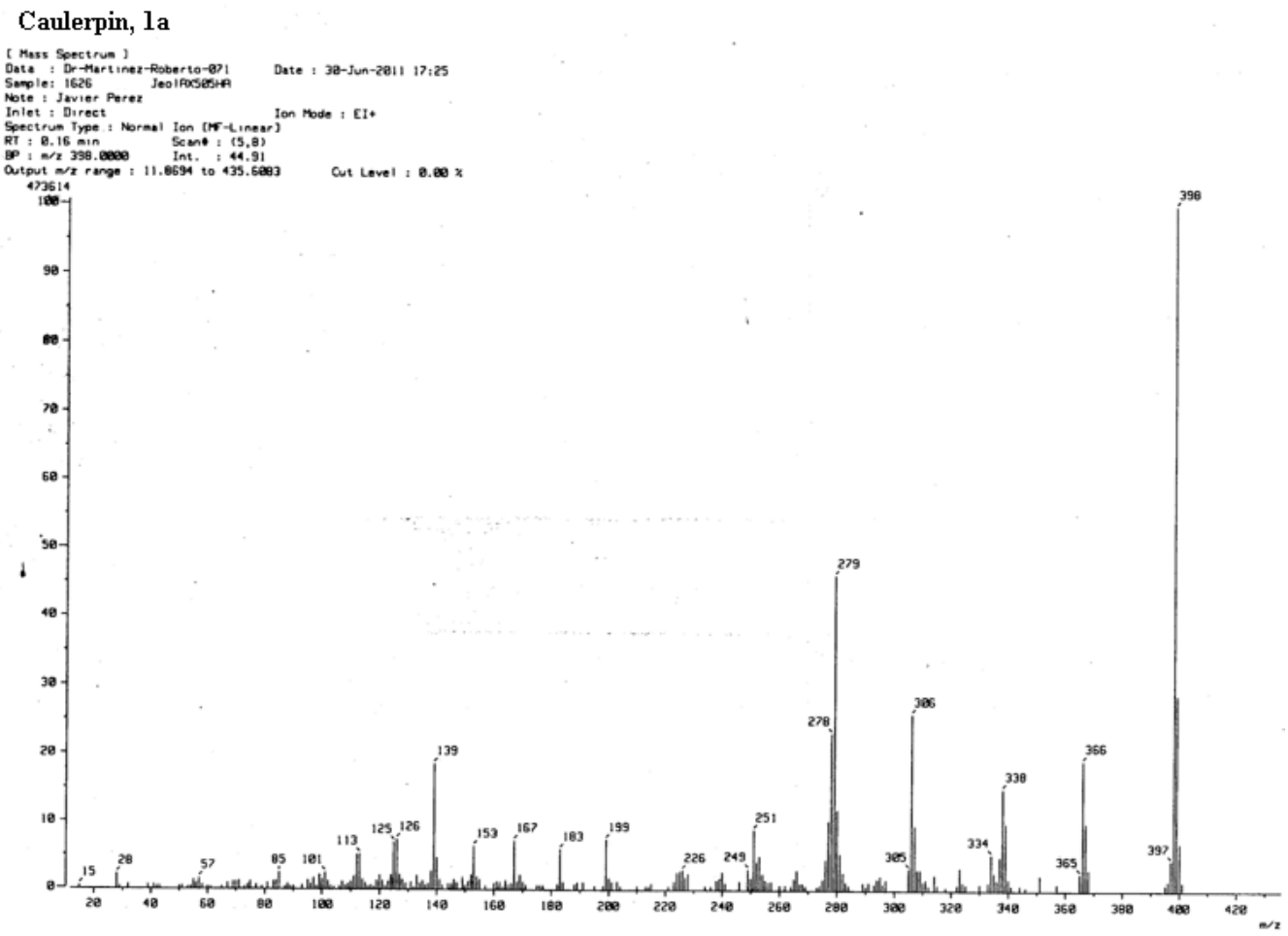

Figure S8. ^{1}H NMR spectrum of diethyl 2-(3-formyl-1*H*-indol-2-yl)malonate (**6a**).

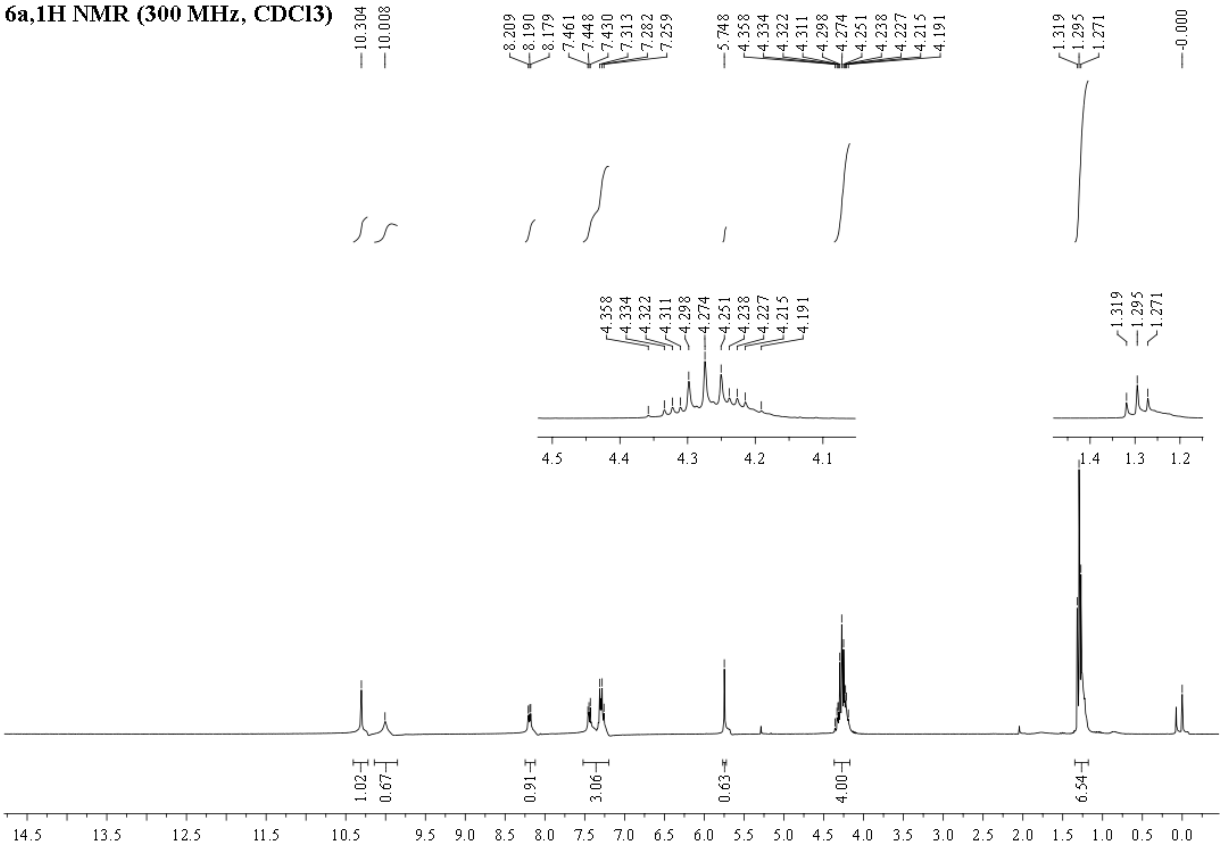

Figure S9. ^{13}C NMR spectrum of diethyl 2-(3-formyl-1*H*-indol-2-yl)malonate (**6a**).

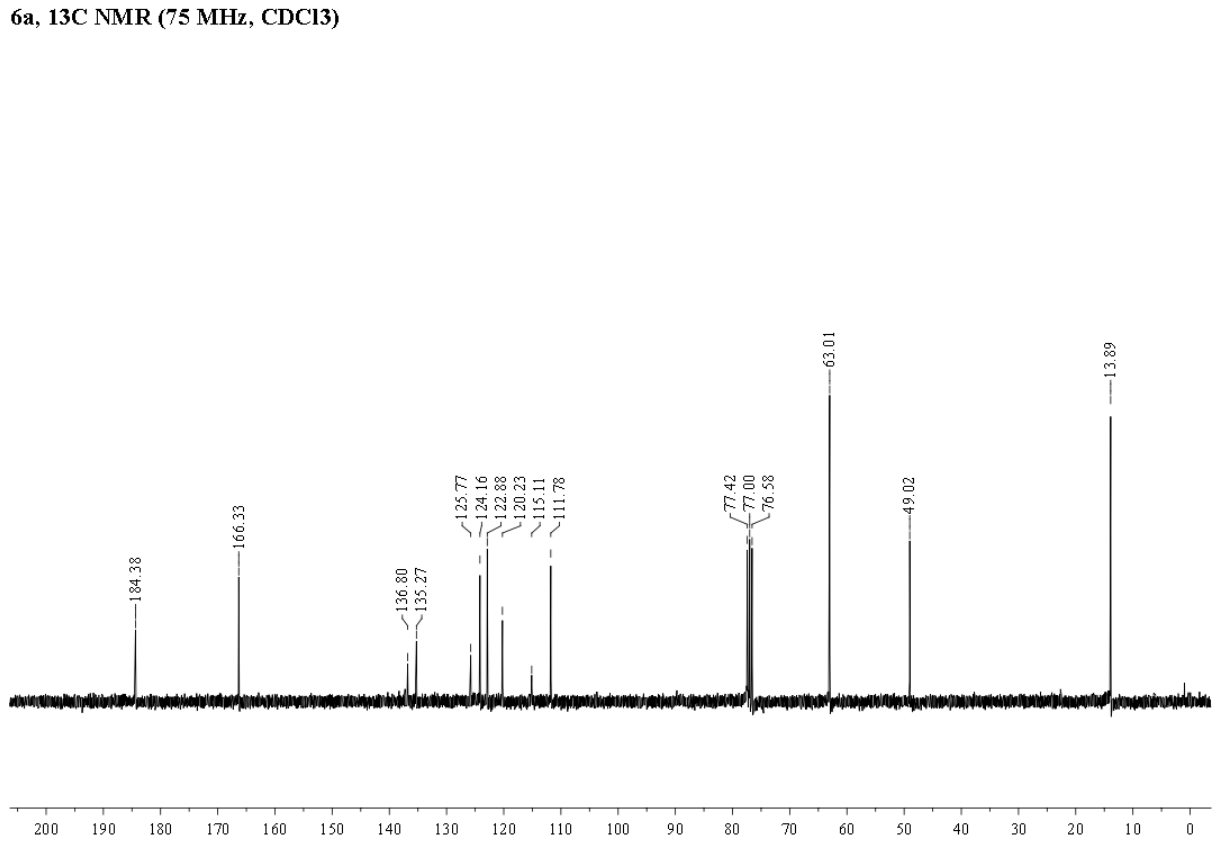

Figure S10. Mass spectrum of diethyl 2-(3-formyl-1*H*-indol-2-yl)malonate (**6a**).

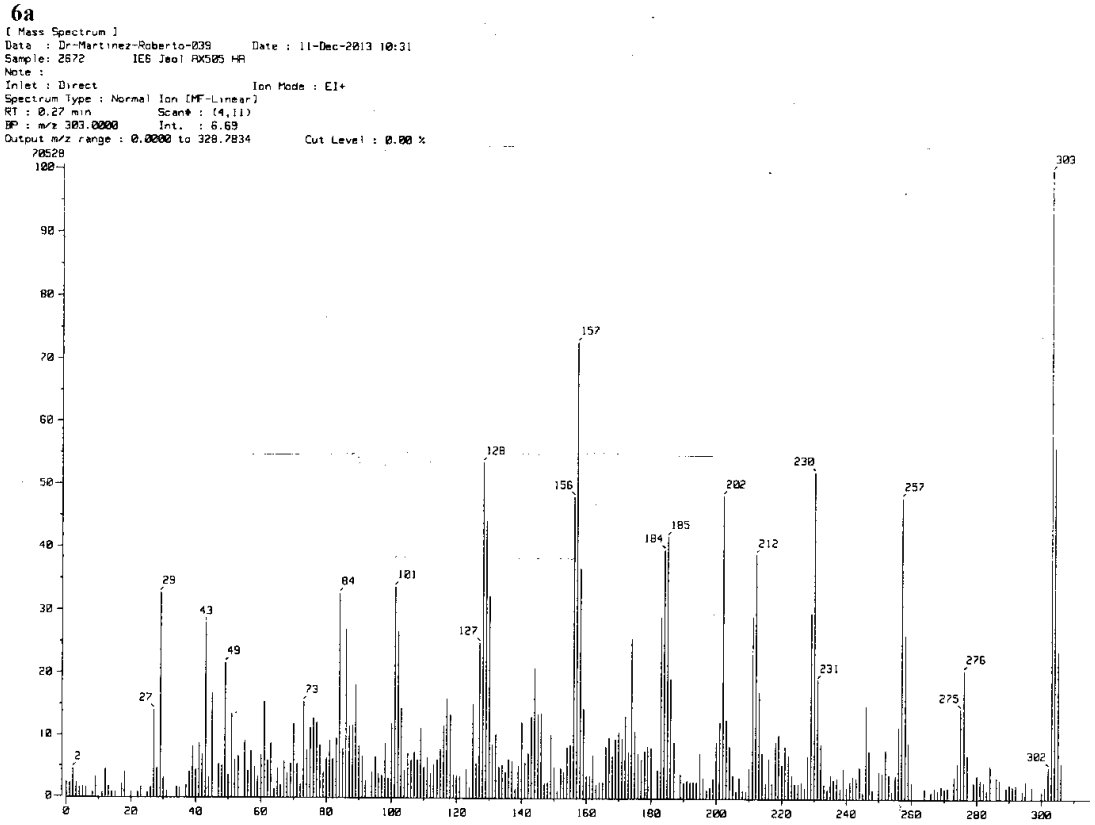

Figure S11. ^{1}H NMR spectrum of dimethyl 2,9-dimethyl-5,12-dihydrocycloocta [1,2-*b*:5,6-*b*′]-diindole-6,13-dicarboxylate (**1f**).

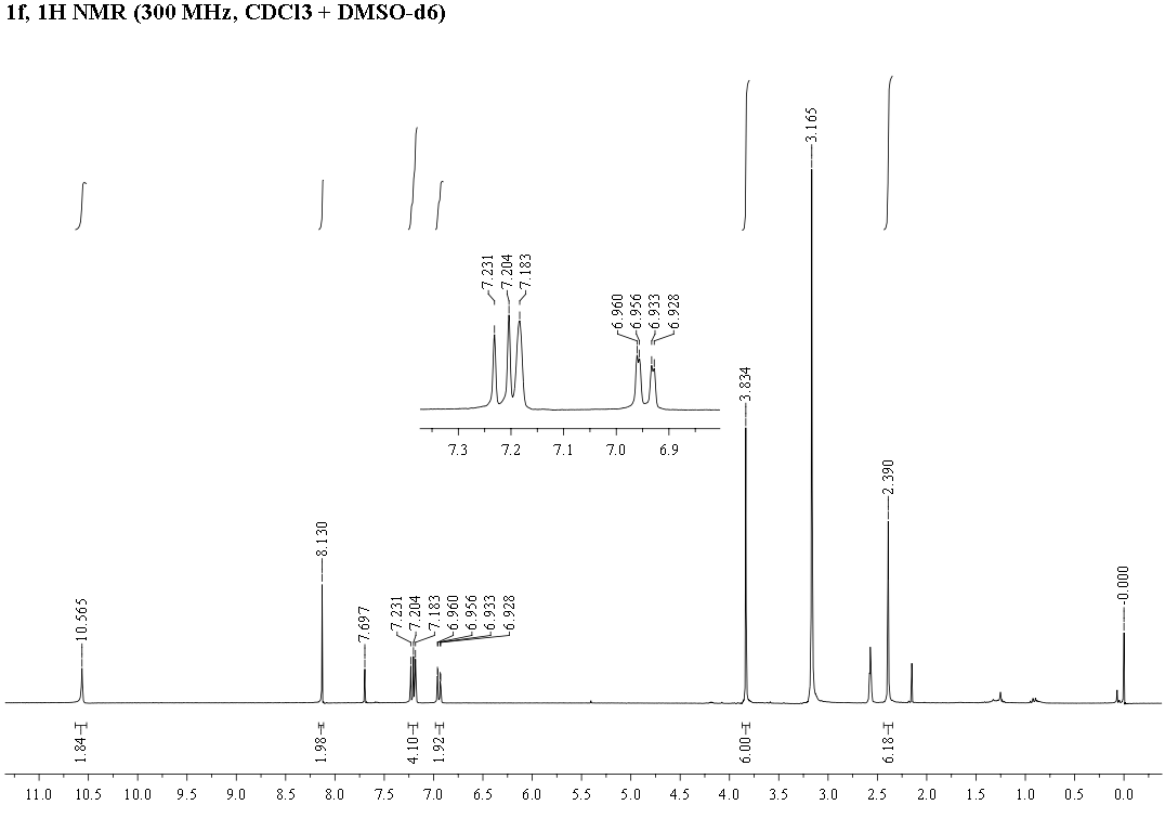

Figure S12. ^{13}C NMR spectrum of dimethyl 2,9-dimethyl-5,12-dihydrocycloocta [1,2-*b*:5,6-*b′*]-diindole-6,13-dicarboxylate (**1f**).

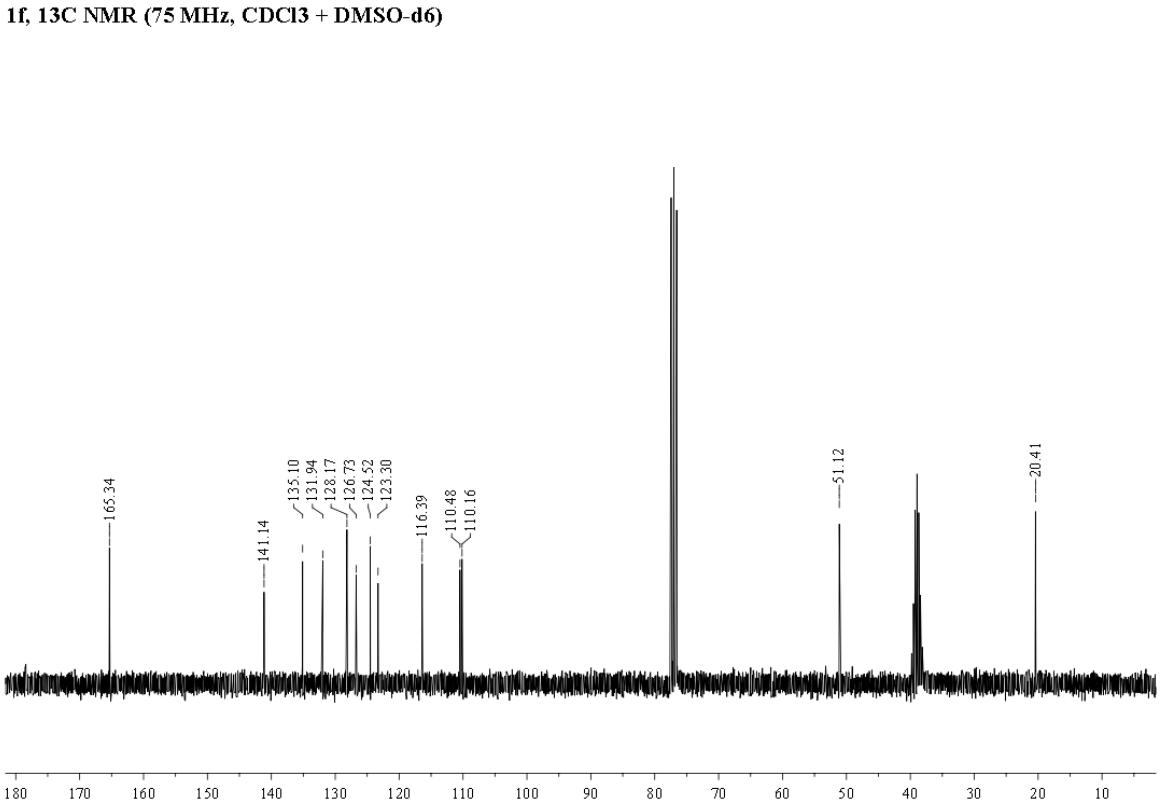

Figure S13. Mass spectrum of dimethyl 2,9-dimethyl-5,12-dihydrocycloocta [1,2-*b*:5,6-*b′*]-diindole-6,13-dicarboxylate (**1f**).

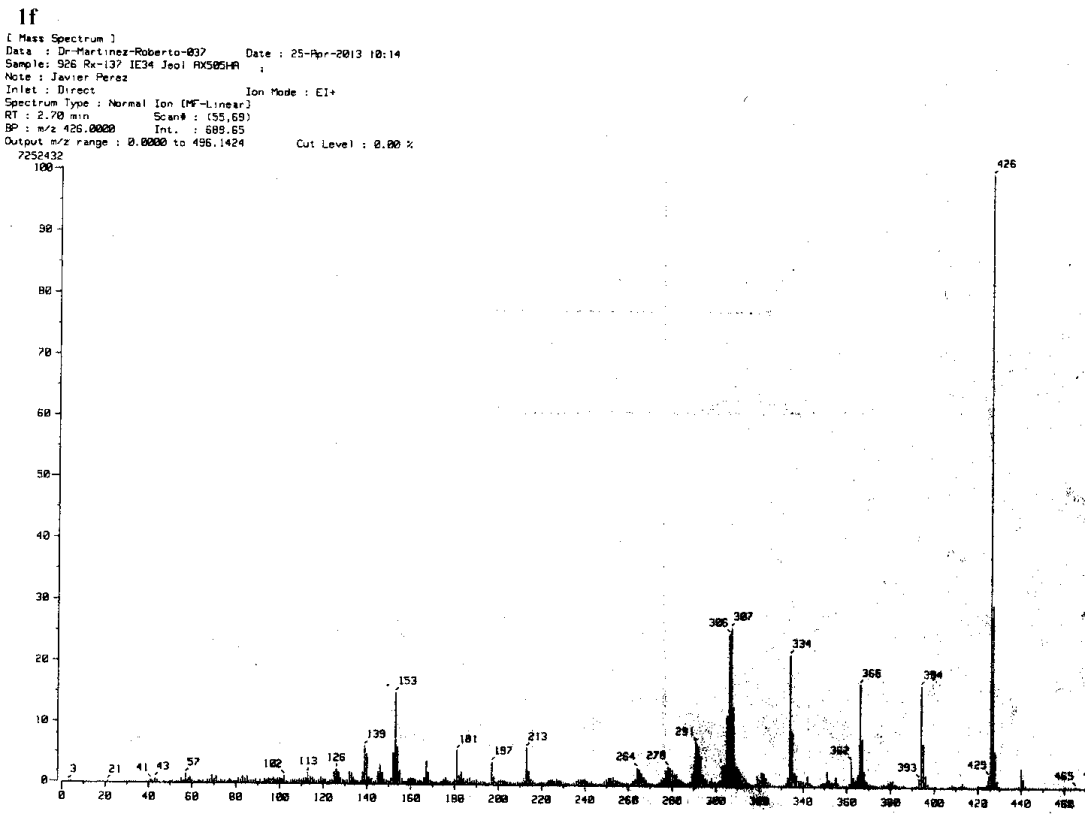

Figure S14. Elemental composition of dimethyl 2,9-dimethyl-5,12-dihydrocycloocta [1,2-*b*:5,6-*b*′]-diindole-6,13-dicarboxylate (**1f**).

1f

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[ Elemental Composition ]                                          Page: 1
Data  : Dr-Roberto-Martinez026        Date : 11-Jun-2013 14:00
Sample: 1351      Rx-137
Note : -luis-velasco
Inlet : Direct                        Ion Mode : FAB+
RT : 2.67 min                         Scan#: (9,13)+(4,6)
Elements : C 40/0, H 49/0, O 7/0, N 3/0
Mass Tolerance       : 1000ppm, 1mmu if m/z > 1
Unsaturation (U.S.) : 0.0 - 34.0

Observed m/z    Int%
  426.1573      100.0
Estimated m/z  Error[ppm]  U.S.      C      H      O      N
  426.1580       -1.6       17.0     26     22      4      2
```

Figure S15. ^{1}H NMR spectrum of diethyl 2-(3-formyl-5-methyl-1*H*-indol-2-yl)malonate (**6f**).

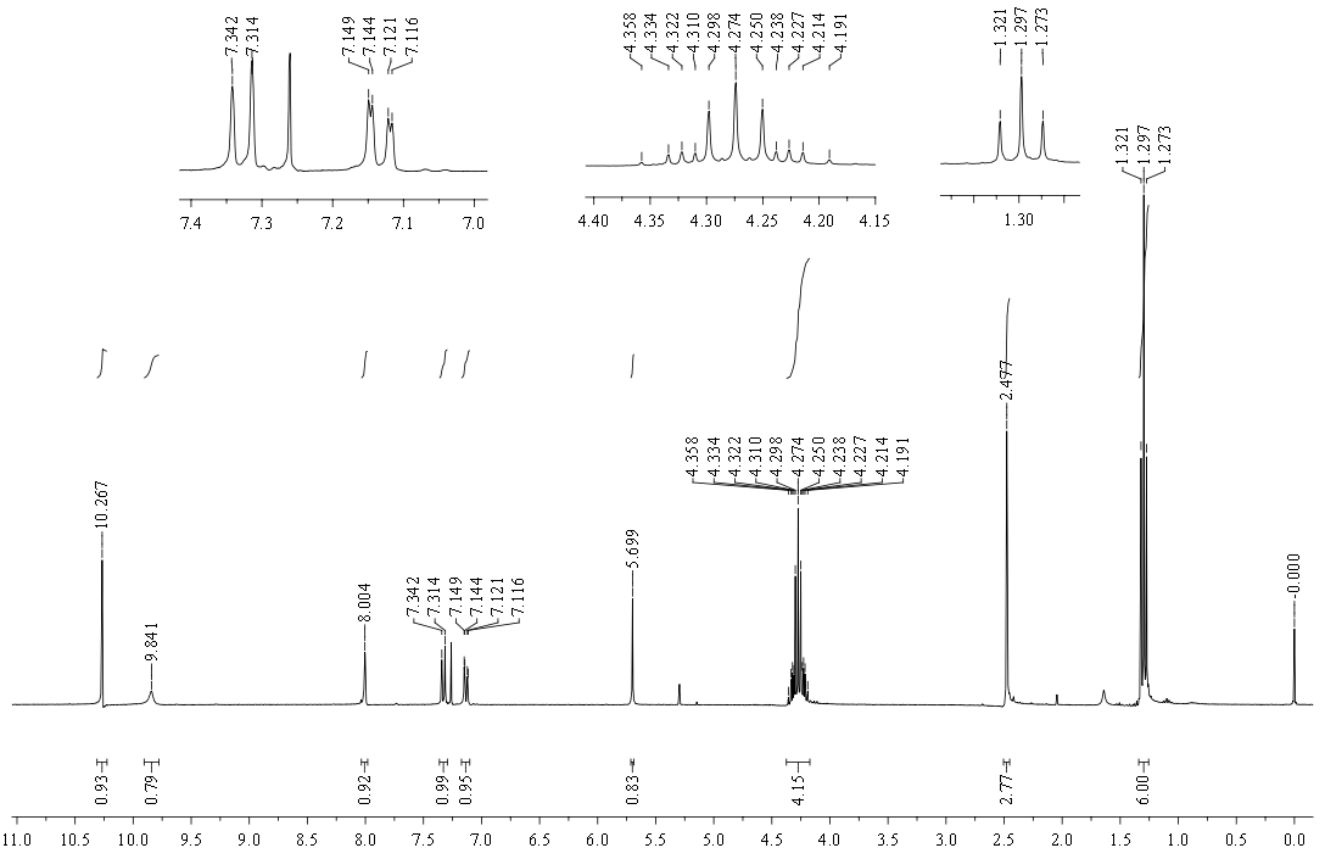

Figure S16. ^{13}C NMR spectrum of diethyl 2-(3-formyl-5-methyl-1*H*-indol-2-yl)malonate (**6f**).

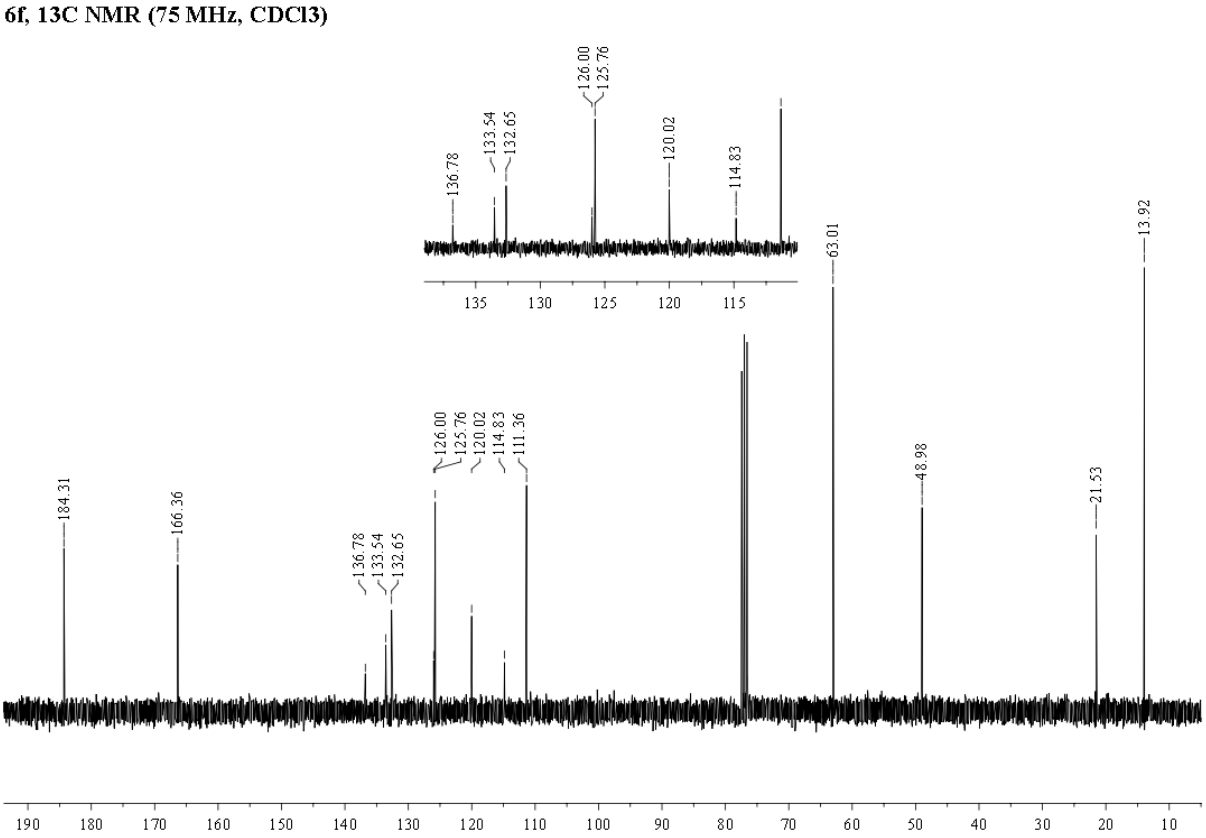

Figure S17. ^{1}H NMR spectrum of methyl 2-(3-formyl-5-methyl-1*H*-indol-2-yl)acetate (**2f**).

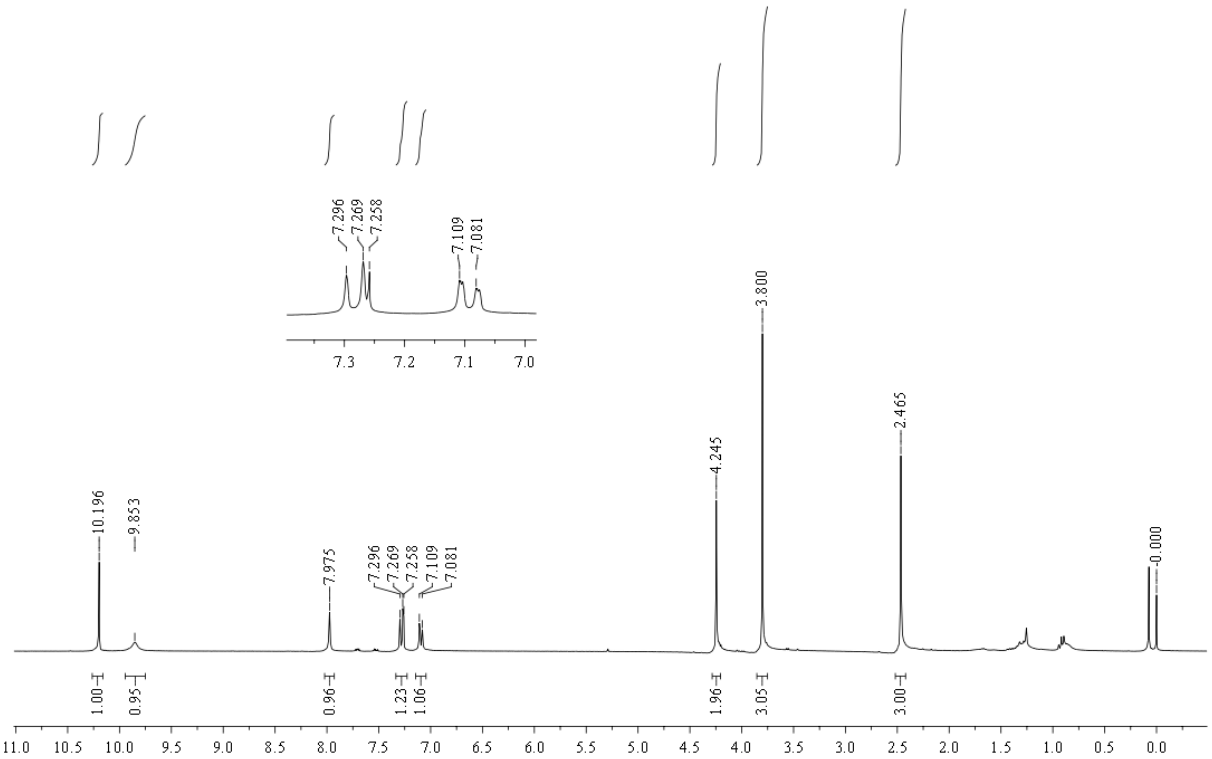

Figure S18. ^{13}C NMR spectrum of methyl 2-(3-formyl-5-methyl-1*H*-indol-2-yl)acetate (**2f**).

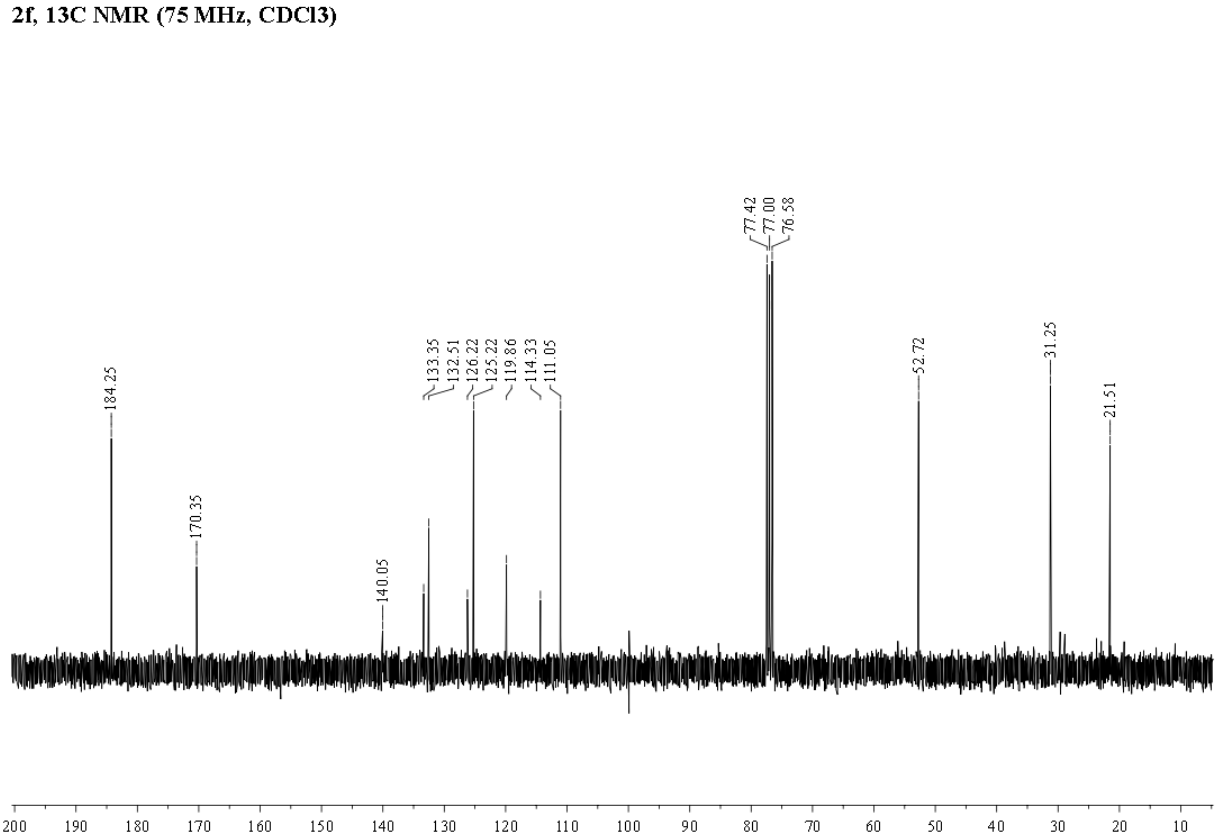

Figure S19. ^{1}H NMR spectrum of 5-methyl-1*H*-indole-3-carbaldehyde (**4f**).

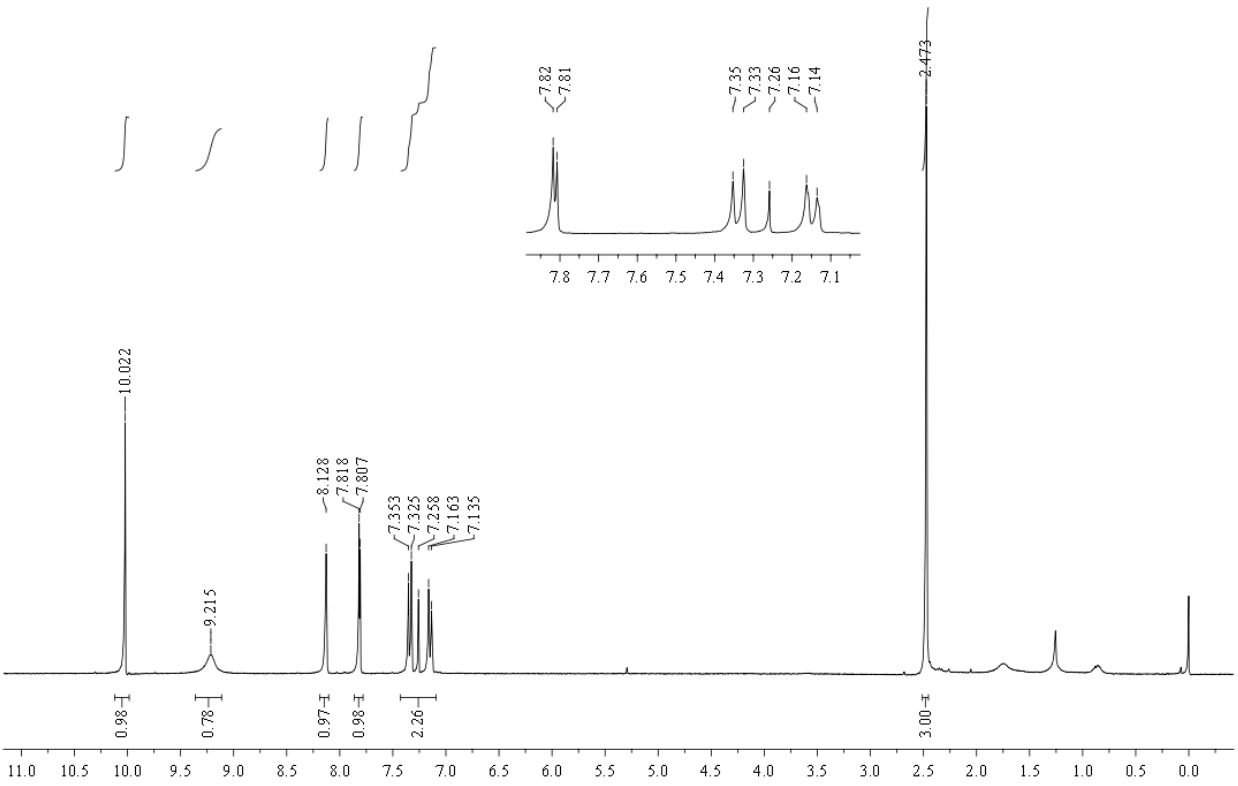

Figure S20. ^{1}H NMR spectrum of diethyl 2-(ethoxycarbonothioylthio)malonate (**3**).

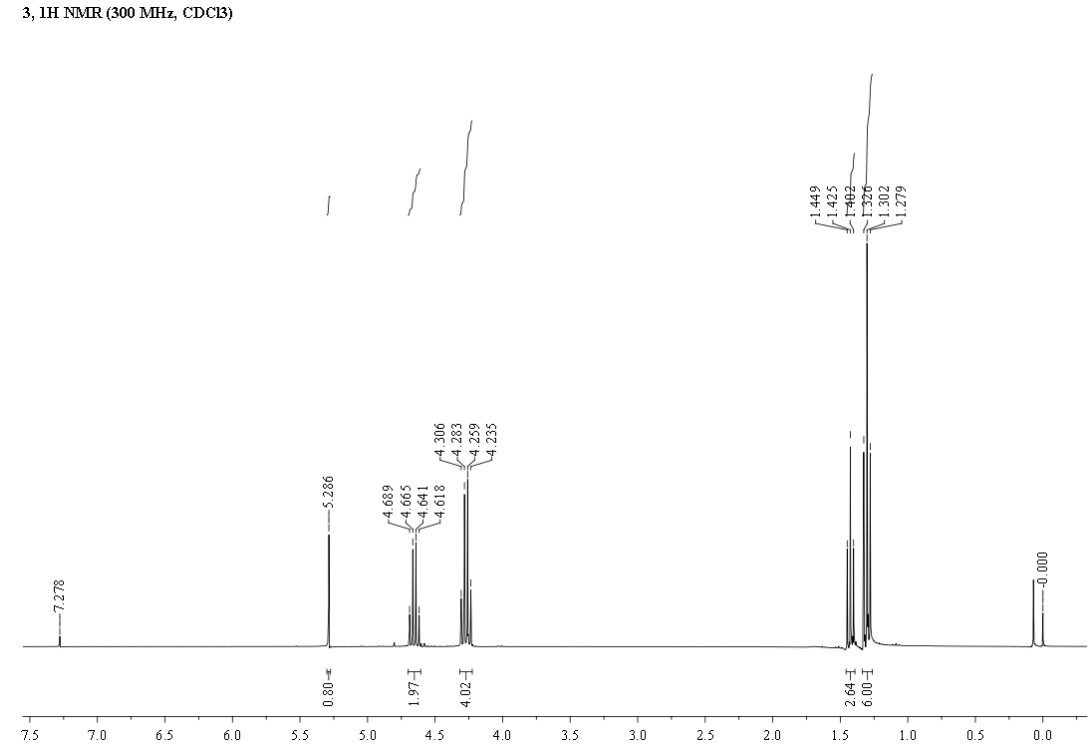

Figure S21. ^{13}C NMR spectrum of diethyl 2-(ethoxycarbonothioylthio)malonate (**3**).