

Supporting Information

Design and synthesis of highly oxygenated furo[3,2-c]pyran-4-ones and furo[3,2-c]chromen-4-ones scaffold as potential anticancer and antimicrobial agent

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Experimental and cytotoxicity and antimicrobial assay details, compound characterization and NMR spectra

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Experimental

Synthesis of highly oxygenated furo[3,2-c]pyran-4-one (4, 5, 8 and 9)

Dehydroacetic acid (1, 1 mmol) or its chalcone (4, 1 mmol) was dissolved in acetonitrile (15 mL) and refluxed for 3 hours after addition of α -bromoketone (2, 1 mmol) and potassium carbonate (K_2CO_3) (3 mmol). The reaction was monitored by TLC using hexane from petroleum-ethylacetate (9:1). After completion, the reaction mixture was cooled to room temperature and water was added to precipitate the desired product (4, 5). The precipitates were filtered, washed with water and crystallized from ethanol. Similarly, furo[3,2-c]chromen-4-ones (8, 9) were synthesized using the above protocols.

2-Benzoyl-3,6-dimethyl-4H-furo[3,2-c]pyran-4-one (4a)

Colour: White, m.p. 142°C , yield 98%, I.R. (KBr, cm^{-1}): 1756(-O-C=O), 1622(-C=O); ^1H NMR (400 MHz, CDCl_3) δ : 7.93 (m, 2H, $\text{C}_2'/\text{C}_6\text{-H}$), 7.60 (m, 1H, $\text{C}_4\text{-H}$), 7.50 (m, 2H, $\text{C}_3'/\text{C}_5\text{-H}$), 6.43 (d, 1H, $\text{C}_7\text{-H}$, $J=0.9$ Hz), 2.66 (s, 3H, $\text{C}_3\text{-CH}_3$), 2.37 (s, 3H, $\text{C}_6\text{-CH}_3$); ^{13}C NMR (100 MHz, CDCl_3): 184.00 (-C=O), 163.27 (C-1a), 161.92 (C-6), 159.27 (C-2), 147.98 (C-4), 137.40 (C-1'), 132.77 (C-3), 131.37 (C-4'), 129.34 (C-2'/C-6'), 128.44 (C-3'/C-5'), 109.98 (C-3a), 95.63 (C-7), 20.58 (C-6- CH_3), 10.88 (C-3- CH_3); HRMS: $m/z(\text{M}^+)$ calcd. for $\text{C}_{16}\text{H}_{12}\text{O}_4$: 268.0736, found: 269.0889($\text{M}^+\text{+H}$).

6-Dimethyl-2-(4-methylbenzoyl)-4H-furo[3,2-c]pyran-4-one (4b)

Colour: White, m.p. 205 °C, yield 98%, I.R. (KBr, cm⁻¹): 1736 (-O-C=O), 1624 (-C=O); ¹H NMR (400 MHz, CDCl₃) δ : 7.85 (d, 2H, C₂/C₆-H, J=8.4 Hz), 7.29 (d, 2H, C₃/C₅-H, J=8.0 Hz), 6.42 (d, 1H, C₇-H, J=0.8 Hz), 2.66 (s, 3H, C₃-CH₃), 2.45 (s, 3H, C₆-CH₃), 2.37 (s, 3H, C₄-CH₃); ¹³C NMR (100 MHz, CDCl₃): 183.69 (-C=O), 163.09 (C-1a), 161.80 (C-6), 159.34 (C-2), 148.14 (C-4), 143.71 (C-4'), 134.75 (C-1'), 130.88 (C-3), 129.55 (C-2'/C-6'), 129.15 (C-3'/C-5'), 109.95 (C-3a), 95.65 (C-7), 21.73 (C-4'-CH₃), 20.57 (C-6-CH₃), 10.88 (C-3-CH₃); HRMS: m/z(M⁺) calcd. for C₁₇H₁₄O₄: 282.0892, found: 283.1008 (M⁺+H).

2-(4-Methoxybenzoyl)-3,6-dimethyl-4H-furo[3,2-c]pyran-4-one (4c)

Colour: white, m.p. 209 °C, yield 98%, I.R. (KBr, cm⁻¹): 1746 (-O-C=O), 1629 (-C=O); ¹H NMR (400 MHz, CDCl₃) δ : 7.99 (d, 2H, C₂/C₆-H, J= 6.88 Hz), 6.99 (d, 2H, C₃/C₅-H, J=6.88 Hz), 6.43 (d, 1H, C₇-H, J=0.8 Hz), 3.90 (s, 3H, C₄-OCH₃), 2.66 (s, 3H, C₃-CH₃), 2.37 (s, 3H, C₆-CH₃), ¹³C NMR (100 MHz, CDCl₃): 182.41 (-C=O), 163.50 (C-1a), 162.95 (C-4'), 161.63 (C-6), 159.32 (C-2), 148.30 (C-4), 131.89 (C-2'/C-6'), 130.49 (C-1'), 130.11 (C-3), 113.78 (C-3'/C-5'), 109.96 (C-3a), 95.61 (C-7), 55.53 (C-4'-OCH₃), 20.53 (C-6-CH₃), 10.82 (C-3-CH₃); HRMS: m/z(M⁺) calcd. for C₁₇H₁₄O₅: 298.0841, found: 299.0958 (M⁺+H).

(E)-2-Benzoyl-3-(2,5-dimethoxystyryl)-6-methyl-4H-furo[3,2-c]pyran-4-one(5a)

Colour: light yellow, m.p. 162 °C, yield 95%, I.R. (KBr, cm⁻¹): 1741 (-O-C=O), 1627 (-C=O); ¹H NMR (400 MHz, CDCl₃) δ : 8.47 (d, 1H, -CH=CH-Ar, J=16.3 Hz), 7.91 (t, 2H, C₂/C₆-H, J=7.2, 1.6 Hz), 7.77 (d, 1H, -CH=CH-Ar, J=16.3 Hz), 7.62 (t, 1H, C₄-H, J=7.6 Hz), 7.51 (t, 2H, C₃/C₅-H, J=7.6, 7.6 Hz), 7.19 (dd, 1H, C₄', J=8.0, 1.6 Hz), 7.10 (d, 1H, C₆', J=1.6 Hz), 6.87 (d, 1H, C₃', J=8.4 Hz), 6.45 (s, 1H, C₇-H), 3.92 (s, 6H, C₃-Ar-OCH₃), 2.41 (s, 3H, C₆-CH₃); ¹³C NMR (100 MHz, CDCl₃): 184.33 (-C=O), 163.76 (C-1a), 163.29 (C-6), 159.23 (C-5''), 149.99 (C-2''), 149.09 (C-2), 146.85 (C-4), 140.76 (C-3), 138.04 (C-1'), 132.59 (C-4'), 132.10 (-CH=CH-Ar), 130.12 (C-2'/C-6'), 129.39 (C-3'/C-5'), 128.41 (-CH=CH-Ar), 121.49 (C-3a), 114.44 (C-1''), 111.05 (C-3''), 109.41 (C-4''), 107.41 (C-6''), 95.58 (C-7), 55.95 (C-5'-OCH₃), 55.93 (C-2''-OCH₃), 20.47 (C-6-CH₃); HRMS: m/z (M⁺) calcd. for C₂₅H₂₀O₆: 416.1260, found: 417.1258 (M⁺+H).

(E)-3-(2,5-Dimethoxystyryl)-6-methyl-2-(4-methylbenzoyl)-4H-furo[3,2-c]pyran-4-one (**5b**)

Colour: light yellow, m.p. 140 °C, yield 95%, I.R. (KBr, cm⁻¹): 1815 (-O-C=O), 1658 (-C=O); ¹H NMR (400 MHz, CDCl₃) δ : 8.73 (d, 1H, -CH=CH-Ar, J=16.6 Hz), 7.84 (d, 1H, -CH=CH-Ar, J= 16.5 Hz), 7.83 (d, 2H, C₂'/C₆'-H, J=8.2 Hz), 7.31 (d, 2H, C₃'/C₅'-H, J=8.0 Hz), 7.12 (d, 1H, C₄'-H, J=1.5 Hz), 6.83 (s, 1H, C₆'', C₃''-H, J=1.6 Hz), 6.44 (d, 1H, C₇-H, J=0.7 Hz), 3.84 (s, 6H, C₂'/C₅'-OCH₃), 2.45 (s, 3H, C₆-CH₃), 2.39 (s, 3H, C₄-CH₃); ¹³C NMR (100 MHz, CDCl₃): 183.95 (-C=O), 163.53 (C-1a), 163.09 (C-6), 159.18 (C-5''), 153.74 (C-2''), 152.54 (C-2), 147.31 (C-4), 143.55 (C-4'), 135.58 (C-3a), 135.29 (C-1'), 131.70 (-CH=CH-Ar), 129.64 (C-2'/C-6'), 129.11 (C-3'/C-5'), 126.95 (-CH=CH-Ar), 117.08 (C-3a), 115.70 (C-1''), 112.74 (C-3''), 111.99 (C-4''), 107.40 (C-6''), 95.50 (C-7), 56.59 (C-5''-OCH₃), 55.81 (C-2''-OCH₃), 21.72 (C-4'-CH₃), 20.43 (C-6-CH₃); HRMS: m/z (M⁺) calcd. for C₂₆H₂₂O₆: 430.1416, found: 431.1203 (M⁺+H).

(E)-3-(2,5-Dimethoxystyryl)-2-(4-methoxybenzoyl)-6-methyl-4H-furo[3,2-c]pyran-4-one (**5c**)

Colour: light yellow, m.p. 138 °C, yield 95%, I.R. (KBr, cm⁻¹): 1743 (-O-C=O), 1625 (-C=O); ¹H NMR (400 MHz, CDCl₃) δ : 8.70 (d, 1H, -CH=CH-Ar, J=16.7 Hz), 7.96 (dd, 2H, C₂'/C₆'-H, J= 6.9, 2.0 Hz), 7.82 (d, 1H, -CH=CH-Ar, J=16.5 Hz), 7.13 (t, 1H, C₄'-H, J=1.3, 1.5 Hz), 6.99 (dd, 2H, C₃'/C₅'-H, J=6.9, 2.0 Hz), 6.83 (d, 1H, C₃'/C₆'-H, J=1.6 Hz), 6.45 (d, 1H, C₇-H, J=0.8 Hz), 3.86 (s, 9H, C₃'/C₂'/C₅'-OCH₃), 2.40 (s, 3H, C₆-CH₃); ¹³C NMR (100 MHz, CDCl₃): 182.74 (-C=O), 163.41 (C-1a), 163.38 (C-4'), 162.97 (C-6), 159.21 (C-5''), 153.74 (C-2''), 152.50 (C-2), 147.44 (C-4), 135.28 (C-3), 131.99 (-CH=CH-Ar), 131.30 (C-2'/C-6'), 130.51 (C-1'), 126.99 (-CH=CH-Ar), 117.14 (C-3a), 115.67 (C-1''), 113.71 (C-3''), 112.74 (C-4''), 111.91 (C-3'/C-5'), 107.37 (C-6''), 95.50 (C-7), 56.59 (C-5''-OCH₃), 55.82 (C-2''-OCH₃), 55.53 (C-4'-OCH₃), 20.42 (C-6-CH₃); HRMS: m/z (M⁺) calcd. for C₂₆H₂₂O₇: 446.1366, found: 447.1204 (M⁺+H).

(E)-2-Benzoyl-3-(2,4-dimethoxystyryl)-6-methyl-4H-furo[3,2-c]pyran-4-one (**5d**)

Colour: dark yellow, m.p. 140 °C, yield 95%, I.R. (KBr, cm⁻¹): 1739 (-O-C=O), 1624 (-C=O); ¹H NMR (400 MHz, CDCl₃) δ : 8.73 (d, 1H, -CH=CH-Ar, J= 16.6 Hz), 7.90 (m, 2H, C₂'/C₆'-H), 7.86 (d, 1H, -CH=CH-Ar, J=16.5 Hz), 7.58 (m, 1H, C₄-H), 7.54 (m, 1H, C₆'-H), 7.51 (m, 2H, C₃'/C₅'-H, J=8.9 Hz), 6.50 (dd, 1H, C₅'-H, J=2.0, 8.5 Hz), 6.45 (d, 1H, C₃'-H, J=2.3 Hz),

6.42(d,1H,C₇-H,J=0.8Hz), 3.82(s,6H, C₂"/C₄"-OCH₃), 2.39(s,3H,C₆-CH₃);
¹³CNMR(100MHz,CDCl₃): 184.22(-C=O),163.58(C-1a), 163.20(C-6),161.66(C-4"),159.31(C-2),159.25(C-4), 146.68(C-2"),138.16(C-3), 136.23(C-1'),133.01(C-4'), 132.53(-CH=CH-Ar), 129.46(C-6"), 128.89(C-2'/C-6'),128.42(C-3'/C-5'),119.29(-CH=CH-Ar), 114.61(C-3a), 107.44(C-1"), 105.14(C-5"), 98.4(C-3"),95.53(C-7), 55.72(C-2"-OCH₃),55.43(C-4"-OCH₃), 20.43(C-6-CH₃); HRMS:m/z(M⁺) calcd.for C₂₅H₂₀O₆: 416.1260, found: 417.1308(M⁺+H).

(E)-3-(2,4-Dimethoxystyryl)-6-methyl-2-(4-methylbenzoyl)-4H-furo[3,2-c]pyran-4-one(**5e**)
Colour: dark yellow, m.p.186 °C, yield 95%, I.R.(KBr, cm⁻¹): 1741 (-O-C=O), 1627 (-C=O);
¹HNMR (400 MHz, CDCl₃) δ : 8.71(d, 1H,-CH=CH-Ar, J =16.6 Hz), 7.86 (d, 1H,-CH=CH-Ar, J =16.6 Hz), 7.84 (d, 2H, C₂'/C₆'-H, J= 8.2 Hz), 7.56 (d, 1H, C₆"-H, J= 8.5 Hz), 7.28 (d, 2H, C₃'/C₅'-H, J =7.9 Hz), 6.50 (dd, 1H, C₅"-H, J=2.0, 8.5 Hz), 6.45 (d, 1H, C₃"-H, J =2.4 Hz), 6.42 (d, 1H, C₇-H, J= 0.8 Hz), 3.89 (s, 6H, C₂"/C₄"-OCH₃), 2.45 (s, 3H, C₆-CH₃), 2.39 (s, 3H,C₄-CH₃); ¹³CNMR (100 MHz, CDCl₃) : 183.92 (-C=O), 163.41 (C-1a), 163.05 (C-6), 161.58 (C-4"), 159.27 (C-2), 146.90 (C-4), 143.31 (C-2"), 141.16 (C-4"), 135.90 (C-3), 135.46 (C-1'),132.59 (-CH=CH-Ar), 129.59 (C-6"), 129.05 (C-2'/C-6'), 128.85 (C-3'/C-5'), 119.37 (-CH=CH-Ar), 114.71 (C-3a), 107.42 (C-1"), 105.13 (C-5"), 98.46 (C-3"), 95.52 (C-7), 55.70 (C-2"-OCH₃), 55.42 (C-4"-OCH₃), 21.71 (C-4'-CH₃), 20.41 (C-6-CH₃); HRMS: m/z(M⁺) calcd. for C₂₆H₂₂O₆: 430.1416, found: 431.1361(M⁺+H).

(E)-3-(2,4-Dimethoxystyryl)-2-(4-methoxybenzoyl)-6-methyl-4H-furo[3,2-c]pyran-4-one(**5f**)
Colour: dark yellow, m.p.180 °C, yield 95%, I.R.(KBr, cm⁻¹): 1741 (-O-C=O),1624 (-C=O);
¹HNMR (400 MHz, CDCl₃) δ : 8.68 (d, 1H, -CH=CH-Ar, J =16.6 Hz), 7.97 (dd, 2H, C₂'/C₆'-H, J= 2.0, 6.8 Hz), 7.84 (d, 1H, -CH=CH-Ar, J = 16.5 Hz), 7.58 (d, 1H, C₆"-H, J = 8.6 Hz), 6.98 (dd, 2H, C₃'/C₅'-H, J = 2.0, 6.9 Hz), 6.51 (dd, 1H, C₅"-H, J =2.3, 8.4 Hz), 6.45 (d, 1H, C₇-H, J = 0.8 Hz), 3.82 (s, 9H, C₄'/C₂"/C₄"-OCH₃), 2.40 (s, 3H, C₆-CH₃); ¹³CNMR (100 MHz, CDCl₃) :182.74 (-C=O), 163.30 (C-1a), 163.25 (C-4'), 162.93 (C-6), 161.53 (C-4"), 159.33 (C-2), 159.21 (C-4), 147.02 (C-2"), 135.60 (C-3), 132.21 (-CH=CH-Ar), 131.93 (C-2'/C-6'), 130.67 (C-1'), 128.77 (C-6"), 119.39 (-CH=CH-Ar), 114.75(C-3a),113.66(C-3'/C-5'), 107.39(C-1"), 105.11(C-5"), 98.45(C-3"), 95.53(C-7), 55.71(C-2"-OCH₃),55.53(C-4"-OCH₃),55.43(C-4'-OCH₃),20.42(C-6-CH₃); HRMS:m/z(M⁺) calcd.for C₂₆H₂₂O₇: 446.1366, found:

447.1320(M⁺+H).

(E)-2-Benzoyl-3-(3,4-dimethoxystyryl)-6-methyl-4H-furo[3,2-c]pyran-4-one (5g)

Colour: yellow, m.p. 160° C, yield 95%, I.R.(KBr,cm⁻¹):1739(-O-C=O),1624(-C=O); ¹HNMR (400 MHz, CDCl₃) δ: 8.46(d,1H,-CH=CH-Ar,J=16.4Hz), 7.91(t,2H,C₂/C₆-H,J=7.2,1.4Hz), 7.76(d,1H,-CH=CH-Ar,J=16.4Hz),7.60(t,1H,C₄-H,J=7.4Hz), 7.52(t,2H,C₃/C₅-H,J=1.3,7.4Hz), 7.19(dd,1H,C₆-H,J=1.8,8.3Hz), 7.09(d,1H,C₂-H,J=1.8Hz), 6.86(d,1H,C₅-H,J=8.3Hz), 6.44(d,1H,C₇-H,J=0.8Hz), 3.92(s,6H,C₃/C₄-OCH₃), 2.41(s,3H,C₆-CH₃); ¹³CNMR(100 MHz,CDCl₃):184.32(-C=O),163.75(C-1a),163.28(C-6), 159.20(C-3"), 150.01(C-4"),149.11(C-2), 146.86(C-4), 140.76(C-3), 138.05(C-1'), 132.57(-CH=CH-Ar),132.09(C-4'),130.14(C-1"),129.38(C-2'/C-6'),128.40(C-3'/C-5'), 121.48(-CH=CH-Ar), 114.44(C-3a),111.08(C-6"),109.46(C-5"),107.40(C-2"), 95.56(C-7), 55.94(C-3"/C-4"-OCH₃), 20.45(C-3-CH₃); HRMS:m/z(M⁺) calcd.for C₂₅H₂₀O₆: 416.1260, found: 417.1265(M⁺+H).

(E)-3-(3,4-Dimethoxystyryl)-6-methyl-2-(4-methylbenzoyl)-4H-furo[3,2-c]pyran-4-one(5h)

Colour: yellow, m.p. 134° C, yield 96%, I.R. (KBr, cm⁻¹): 1741 (-O-C=O), 1627 (-C=O); ¹HNMR (400 MHz, CDCl₃) δ : 8.46 (d, 1H,-CH=CH-Ar, J = 16.4 Hz), 7.85 (d, 2H, C₂/C₆-H, J= 8.2 Hz), 7.79 (d, 1H, -CH=CH-Ar, J=16.4 Hz), 7.33 (d, 2H, C₃/C₅-H, J= 8.0 Hz), 7.23 (dd, 1H, C₆-H, J = 2.0, 8.4 Hz), 7.12 (d, 1H, C₂-H, J = 2.0 Hz), 6.88 (d, 1H, C₅-H, J = 8.4 Hz), 6.45 (d, 1H, C₇-H, J = 0.8 Hz), 3.94 (s, 6H, C₃/C₄-OCH₃), 2.46 (s, 3H, C₆-CH₃), 2.41 (s, 3H, C₄-CH₃); ¹³CNMR (100 MHz, CDCl₃) : 184.03 (-C=O), 163.61 (C-1a), 163.14 (C-6), 159.28 (C-3"), 149.93 (C-4"), 149.08 (C-2), 147.08 (C-4), 143.54 (C-4'), 140.46 (C-3), 135.34 (C-1'), 131.71 (-CH=CH-Ar), 130.20 (C-1"), 129.61 (C-2'/C-6'), 129.12 (C-3'/C-5'), 121.44 (-CH=CH-Ar), 114.55 (C-3a), 111.05 (C-6"), 109.40 (C-5"), 107.39 (C-2"), 95.58 (C-7), 55.95 (C-3"/C-4"-OCH₃), 21.74 (C-4'-CH₃), 20.46 (C-6-CH₃); HRMS :m/z (M⁺) calcd.for C₂₆H₂₂O₆: 430.1416, found: 431.1454 (M⁺+H).

(E)-3-(3,4-dimethoxystyryl)-2-(4-methoxybenzoyl)-6-methyl-4H-furo[3,2-c]pyran-4-one(5i)

Colour: yellow, m.p. 139° C, yield 96%, I.R. (KBr, cm⁻¹): 1739 (-O-C=O), 1629 (-C=O); ¹HNMR (400 MHz, CDCl₃) δ : 8.43 (d, 1H,-CH=CH-Ar, J=16.4 Hz), 7.98 (d, 2H, C₂/C₆-H, J =

8 Hz), 7.76 (d, 1H, -CH=CH-Ar, J = 16.4 Hz), 7.21 (d, 1H, C_{2''}-H, J = 8.1 Hz), 7.12 (s, 1H, C_{5''}-H), 7.01 (d, 2H, C_{3'}/C_{5'}-H, J = 8 Hz), 6.88 (dd, 1H, C_{6''}-H, J = 3.6, 8.3 Hz), 6.46 (d, 1H, C₇-H, J = 2.8 Hz), 3.91 (s, 9H, C_{3''}/C_{4''}-OCH₃, C_{4'}-OCH₃), 2.41 (s, 3H, C₆-CH₃); ¹³CNMR (100 MHz, CDCl₃): 182.82 (-C=O), 163.49 (C-1a), 163.37 (C-4'), 163.02 (C-6), 159.30 (C-3''), 149.87 (C-4''), 149.07 (C-2), 147.21 (C-4), 140.16 (C-3), 131.94 (-CH=CH-Ar), 131.33 (C-2'/C-6'), 130.57 (C-1'), 130.24 (C-1''), 121.38 (-CH=CH-Ar), 114.61 (C-3'/C-5'), 113.72 (C-3a), 111.05 (C-6''), 109.37 (C-5''), 107.35 (C-2''), 95.57 (C-7), 55.95 (C-3''/C-4''-OCH₃), 55.54 (C-4'-OCH₃), 20.45 (C-6-CH₃); HRMS : m/z(M⁺) calcd. for C₂₆H₂₂O₇: 446.1366, found: 447.1381 (M⁺+H).

(E)-2-Benzoyl-6-methyl-3-(3,4,5-trimethoxystyryl)-4H-furo[3,2-c]pyran-4-one (5j)

Colour: light yellow, m.p. 167° C, yield 95%, I.R. (KBr, cm⁻¹): 1743 (-O-C=O), 1625 (-C=O); ¹HNMR (400 MHz, CDCl₃) δ: 8.44 (d, 1H, -CH=CH-Ar, J = 16.3 Hz), 7.92 (dd, 2H, C_{2'}/C_{6''}-H, J = 1.4, 8.2 Hz), 7.76 (d, 1H, -CH=CH-Ar, J = 16.3 Hz), 7.62 (t, 1H, C_{4'}-H), 7.53 (t, 2H, C_{3'}/C_{5''}-H, J = 7.6 Hz), 6.80 (s, 2H, C_{2''}/C_{6''}-H), 6.46 (d, 1H, C₇-H, J = 0.8 Hz), 3.93 (s, 9H, C_{3''}/C_{4''}/C_{5''}-OCH₃), 2.43 (s, 3H, C₆-CH₃); ¹³CNMR (100 MHz, CDCl₃) : 184.41 (-C=O), 163.85 (C-1a), 163.35 (C-6), 159.23 (C-3''/C-5''), 153.36 (C-2), 147.08 (C-4), 140.70 (C-3), 138.96 (C-4''), 138.96 (C-1'), 137.96 (C-1''), 132.70 (-CH=CH-Ar), 131.62 (C-4'), 129.41 (C-2'/C-6'), 128.45 (C-3'/C-5'), 115.77 (-CH=CH-Ar), 107.39 (C-3a), 104.49 (C-2''/C-6''), 95.59 (C-7), 60.99 (C-4''-OCH₃), 56.19 (C-3''/C-5''-OCH₃), 20.49 (C-6-CH₃); HRMS: m/z(M⁺) calcd. for C₂₆H₂₂O₇: 446.1366, found: 447.1381 (M⁺+H).

(E)-6-Methyl-2-(4-methylbenzoyl)-3-(3,4,5-trimethoxystyryl)-4H-furo[3,2-c]pyran-4-one (5k)

Colour: light yellow, m.p. 140° C, yield 95%, I.R. (KBr, cm⁻¹): 1737 (-O-C=O), 1634 (-C=O); ¹HNMR (400 MHz, CDCl₃) δ : 8.42 (d, 1H, -CH=CH-Ar, J = 16.3 Hz), 7.84 (d, 2H, C_{2'}/C_{6''}-H, J = 8.1 Hz), 7.77 (d, 1H, -CH=CH-Ar, J = 16.3 Hz), 7.32 (d, 2H, C_{3'}/C_{5''}-H, J = 8.1 Hz), 6.82 (s, 2H, C_{2''}/C_{6''}-H), 6.45 (d, 1H, C₇-H, J = 0.8 Hz), 3.83 (s, 12H, C_{3''}/C_{4''}/C_{5''}-OCH₃), 2.45 (s, 3H, C₆-CH₃), 2.41 (s, 3H, C₄-CH₃); ¹³CNMR (100 MHz, CDCl₃): 184.06 (-C=O), 163.669 (C-1a), 159.25 (C-6), 153.35 (C-3''/C-5''), 152.42 (C-2), 147.30 (C-4), 143.66 (C-4'), 140.39 (C-3), 138.94 (C-4''), 135.27 (C-1''), 132.77 (-CH=CH-Ar), 131.19 (C-1'), 129.63 (C-2'/C-6'), 128.99 (C-3'/C-5'), 115.87 (-CH=CH-Ar), 107.36 (C-3a), 104.51 (C-2''/C-6''), 95.57 (C-7), 60.97 (C-4''-OCH₃), 56.16 (C-3''/C-5''-OCH₃),

21.73(C-4'-CH₃), 20.45(C-6-CH₃); HRMS:m/z(M⁺) calcd.for C₂₇H₂₄O₇: 460.1522, found: 461.1659(M⁺+H).

(E)-2-(4-Methoxybenzoyl)-6-methyl-3-(3,4,5-trimethoxystyryl)-4H-furo[3,2-c]pyran-4-one(5l)

Colour: light yellow, m.p.143 °C,yield 95%, I.R.(KBr,cm⁻¹): 1734(-O-C=O),1616(-C=O); ¹HNMR (400 MHz, CDCl₃)δ: 8.40(d,1H,-CH=CH-Ar,J=16.3Hz), 7.96(d,2H,77,J=8.9 Hz), 7.73(d,1H,-CH=CH-Ar,J=16.3Hz), 7.00(d,2H,C₃'/C₅'-H,J=8.9 Hz), 6.80(s,2H,C₂''/C₆''-H), 6.47(d,1H,C₇-H,J=0.8Hz), 3.84(s,12H,C₃''/C₄''/C₅''/OCH₃,C₄'-OCH₃), 2.41(s,3H,C₆-CH₃);¹³CNMR(100 MHz,CDCl₃):182.88(-C=O),163.56(C-1a), 163.45(C-4'),163.08(C-6),159.30(C-3''/C-5''),153.34(C-2),147.45(C-4), 140.08(C-3),138.84(C-4''),132.82(C-1''),131.97(-CH=CH-Ar),130.81(C-2'/C-6'),130.50(C-1'), 115.95(-CH=CH-Ar), 113.75(C-3a), 107.33(C-3'/C-5'), 104.42(C-2''/C-6''),95.58(C-7),60.97(C-4''-OCH₃),56.16(C-3''/C-5''-OCH₃),55.54(C-4'-OCH₃),20.46(C-6-CH₃); HRMS:m/z(M⁺) calcd.for C₂₇H₂₄O₈: 476.1471, found: 477.1550(M⁺+H).

2-Benzoyl-3-methyl-4H-furo[3,2-c]chromen-4-one (8a)

Colour: white, m.p. 141 ° C, yield 92%, IR (KBr, cm⁻¹) : 1753 (-O-C=O), 1672 (-C=O); ¹HNMR: (400 MHz, CDCl₃) δ: 8.03 (d, 2H, C₂'/C₆'-H, J = 7.08 Hz), 7.91 (dd, 1H, C₉-H, J = 7.8, 7.8 Hz), 7.60-7.65 (m, 2H, C₄'-H and C₇-H), 7.56 (t, 2H, C₃'/C₅'-H), 7.47 (d, 1H, C₆-H, J = 8.3 Hz), 7.38 (t, 1H, C₈-H, J = 6.8, 8.3 Hz), 2.76 (s, 3H, C₃-CH₃), ¹³C NMR (100 MHz, CDCl₃) δ: 183.96 (-C=O), 157.84 (C-9b), 157.75 (C-4), 153.75 (C-5a), 149.07 (C-2), 137.33 (C-1'), 132.97 (C-4'), 132.31 (C-7), 132.01 (C-3), 129.47 (C-2'/C-6'), 128.54 (C-3'/C-5'), 124.83 (C-8), 121.75 (C-9), 117.59 (C-6), 112.20 (C-3a), 111.95 (C-9a), 10.82 (C-3-CH₃); HRMS: (m/z) M⁺ calcd. for C₁₉H₁₂O₄: 304.0736, found: 305.0798 (M⁺+H).

3-Methyl-2-(4-methylbenzoyl)-4H-furo[3,2-c] chromen-4-one (8b)

Colour: light brown, m.p. 108 ° C, yield: 90%, IR(KBr, cm⁻¹) : 1726 (-O-C=O), 1656 (-C=O); ¹HNMR: (400 MHz, CDCl₃) δ: 7.95 (d, 2H, C₂'/C₆'-H, J = 8.4 Hz), 7.91 (dd, 1H, C₉-H, J = 7.8 Hz), 7.57-7.62 (m, 1H, C₇-H), 7.47 (d, 1H, C₆-H, J = 7.6 Hz), 7.35-7.39 (m, 3H, C₈-H, C₃'/C₅'-H), 2.74 (s, 3H, C₃-CH₃), 2.38 (s, 3H, C₄'-CH₃); ¹³C NMR (100 MHz, CDCl₃) δ: 183.64

(-C=O), 163.06 (C-9b), 161.77 (C-5a), 159.29 (C-2), 148.15 (C-4), 143.67 (C-4'), 134.77 (C-1'), 132.30 (C-7), 130.86 (C-3), 129.54 (C-2'/C-6'), 129.14 (C-3'/C-5'), 124.8(C-8), 121.7 (C-9), 117.6 (C-6), 112.2 (C-3a), 111.9 (C-9a), 21.71 (C-4'-CH₃), 10.85 (C-3-CH₃); HRMS: (m/z) M⁺ calcd. for C₂₀H₁₄O₄: 318.892, found: 319.0955 (M⁺+H).

2-(4-Methoxybenzoyl)-3-methyl-4H-furo[3,2-c]chromen-4-one (8c)

Colour: white, m.p. 110° C, yield 90%, IR (KBr, cm⁻¹): 1746 (-O-C=O), 1632 (-C=O); ¹HNMR (400 MHz, CDCl₃) δ: 8.08 (d, 1H, C₉-H, J = 8.7 Hz), 7.94 (t, 1H, C₇-H, J = 8.9, 9.3 Hz), 7.75 (d, 1H, C₆-H, J = 8.4 Hz), 7.59 (t, 1H, C₈-H, J = 6.9, 7.2 Hz), 7.36 (m, 2H, C₂/C₆-H), 6.99-7.05 (m, 2H, C₃/C₅-H), 3.88-3.93 (s, 3H, C₄-OCH₃), 2.75 (s, 3H, C₃-CH₃); ¹³CNMR (100 MHz, CDCl₃): 182.41 (-C=O), 163.46 (C-9b), 162.93 (C-4'), 161.62 (C-5a), 159.34 (C-2), 148.24 (C-4), 132.38 (C-7), 131.88 (C-2'/C-6'), 130.48 (C-1'), 130.04 (C-3), 124.8 (C-8), 121.7 (C-9), 117.6 (C-6), 113.75 (C-3'/C-5'), 112.2 (C-3a), 111.9 (C-9a), 55.52 (C-4'-OCH₃), 10.85 (C-3-CH₃); HRMS: m/z M⁺ calcd. for C₂₀H₁₄O₅: 334.0841, found: 335.0901 (M⁺+H).

(E)-2-Benzoyl-3-(2,5-dimethoxystyryl)-4H-furo[3,2-c]chromen-4-one (9a)

Colour: yellow, m.p. 140° C, yield 97%, I.R. (KBr, cm⁻¹): 1753 (-O-C=O), 1600 (-C=O); ¹HNMR: (400 MHz, CDCl₃) δ: 8.75 (d, 1H, -CH=CH-Ar, J = 16.0 Hz), 8.00 (d, 2H, C₂/C₆-H, J = 8.0 Hz), 7.93 (d, 1H, -CH=CH-Ar, J = 16.0 Hz), 7.90 (dd, 1H, C₉-H, J = 1.2, 8.4 Hz), 7.59-7.65 (m, 2H, C₈-H, C₄-H), 7.55 (t, 2H, C₃/C₅-H, J = 8.9 Hz), 7.48 (d, 1H, C₆-H, J = 7.6 Hz), 7.37 (m, 1H, C₇-H), 7.17 (s, 1H, C₄"-H), 6.85 (s, 2H, C₃"/C₆"-H), 3.81, 3.88 (s, 6H, C₂"-OCH₃ and C₅"-OCH₃); ¹³C NMR (100 MHz, CDCl₃) δ: 184.17(-C=O), 158.93 (C-9b), 157.58 (C-5"), 153.84 (C-2"), 152.67 (C-4), 148.16 (C-5a), 137.94 (C-1'), 136.28 (-CH=CH-Ar), 132.79 (C-4'), 132.70 (C-7), 132.54 (C-3), 129.56 (C-2'/C-6'), 128.49 (C-3'/C-5'), 126.88 (-CH=CH-Ar), 124.75 (C-8), 121.94 (C-9), 117.32 (C-6), 116.70 (C-3a), 115.97 (C-1"), 112.84 (C-6"), 112.17 (C-3"), 111.89 (C-4"), 109.66 (C-9a), 56.64, 55.89 (C-2", C-5"-OCH₃); HRMS: (m/z) M⁺ calcd. for C₂₈H₂₀O₆: 452.1260, found: 453.1354 (M⁺+H).

(E)-3-(2,5-Dimethoxystyryl)-2-(4-methylbenzoyl)-4H-furo[3,2-c]chromen-4-one (9b)

Colour: yellow, m.p. 146° C, yield 95%, I.R. (KBr, cm⁻¹): 1744 (-O-C=O), 1620 (-C=O); ¹HNMR: (400 MHz, CDCl₃) δ: 8.74(d, 1H, -CH=CH-Ar, J = 16.6 Hz), 7.89-7.93(m, 4H, -

CH=CH-Ar, C₉-H, C₂'/C₆'-H), 7.59-7.63 (m, 1H, C₇-H), 7.48 (d, 1H, C₆-H, J = 8.0 Hz), 7.34-7.39 (m, 1H, C₈-H, C₃'/C₅'-H), 7.17 (s, 1H, C₄'-H), 6.85 (s, 2H, C₃"/C₆"-H), 3.81, 3.88 (s, 6H, C₂"/C₅"-OCH₃), 2.47 (s, 1H, C₄'-CH₃); ¹³C NMR (100 MHz, CDCl₃) δ: 183.95 (-C=O), 163.53 (C-9b), 163.09 (C-5a), 159.18 (C-5"), 153.74 (C-2"), 152.54 (C-2), 147.31 (C-4), 143.55 (C-4'), 135.58 (C-3), 135.29 (C-1'), 131.70 (-CH=CH-Ar), 129.64 (C-2'/C-6'), 129.11 (C-3'/C-5'), 126.95 (-CH=CH-Ar), 124.45 (C-8), 121.13 (C-9), 117.46 (C-6), 117.08 (C-3a), 115.70 (C-1"), 112.74 (C-6"), 111.99 (C-3"), 107.40 (C-4"), 95.50 (C-7), 56.59 (C-5"-OCH₃), 55.81 (C-2"-OCH₃), 20.43 (C-4'-CH₃); HRMS: (m/z) M⁺ calcd. for C₂₉H₂₂O₆: 466.1416, found: 467.1518 (M⁺+H).

(E)-3-(2,5-Dimethoxystyryl)-2-(4-methoxybenzoyl)-4H-furo[3,2-c]chromen-4-one (**9c**)

Colour: yellow, m.p. 180° C, yield 95%, I.R. (KBr, cm⁻¹): 1754 (-O-C=O), 1627 (-C=O); ¹H NMR: (400 MHz, CDCl₃) δ: 8.71 (d, 1H, -CH=CH-Ar, J = 16.6 Hz), 8.05 (d, 2H, C₂'/C₆'-H, J = 8.9 Hz), 7.93 (dd, 1H, C₉-H, J = 1.2, 8.4 Hz), 7.89 (d, 1H, -CH=CH-Ar, J = 16.5 Hz), 7.60-7.64 (m, 1H, C₇-H), 7.48 (d, 1H, C₆-H, J = 7.6 Hz), 7.36-7.41 (m, 1H, C₈-H), 7.17 (s, 1H, C₄'-H), 7.03 (d, 2H, C₃'/C₅'-H, J = 8.9 Hz), 6.84 (s, 2H, C₃"/C₆"-H), 3.81, 3.88, 3.92 (s, 9H, C₂"/C₅", C₄'-OCH₃); ¹³C NMR (100 MHz, CDCl₃) δ: 182.69 (-C=O), 163.57 (C-9b), 153.79 (C-4'), 153.74 (C-5"), 152.55 (C-2"), 135.60 (C-5a), 132.39 (C-2), 132.10 (C-2'/C-6'), 130.48 (C-7), 126.96 (C-1'), 124.72 (C-3), 121.85 (C-8), 117.30 (C-9), 116.82 (C-6), 115.77 (C-1"), 113.84 (C-3',C-5'), 112.76 (C-3"), 112.05 (C-4"), 111.97 (C-6") 56.61, 55.85, 55.56 (C-2", C-5", C-4'-OCH₃); HRMS: (m/z) M⁺ calcd. for C₂₉H₂₂O₇: 482.1366, found: 483.1472 (M⁺+H).

(E)-2-Benzoyl-3-(3,4,5-trimethoxystyryl)-4H-furo[3,2-c]chromen-4-one (**9d**)

Colour: yellow, m.p. 191° C, yield 94%, I.R. (KBr, cm⁻¹): 1734 (-O-C=O), 1627 (-C=O); ¹H NMR: (400 MHz, CDCl₃) δ: 8.47 (d, 1H, -CH=CH-Ar, J = 16.3 Hz), 8.01 (d, 2H, C₂'/C₆'-H, J = 7.6 Hz), 7.91 (dd, 1H, C₉-H, J = 7.8, 7.6 Hz), 7.83 (1H, d, CH=CH-Ar, J = 16.3 Hz), 7.83 (d, 1H, CH=CH-Ar, J = 16.3 Hz), 7.62-7.67 (m, 2H, C₈-H, C₄'-H), 7.55-7.59 (m, 2H, C₃'/C₅'-H), 7.50 (d, 1H, C₆-H, J = 8.0 Hz), 7.40 (t, 1H, C₇-H, J = 5.1, 7.6 Hz), 6.84 (s, 2H, C₂", C₆"-H) 3.77-3.92 (s, 9H, C₃", C₄", C₅"-OCH₃); ¹³C NMR (100 MHz, CDCl₃): 184.41 (-C=O), 163.85 (C-9b), 163.35 (C-5a), 159.23 (C-3"/C-5"), 153.36 (C-2), 147.08 (C-4), 140.70 (C-3), 138.96 (C-4"), 138.96 (C-1'), 137.96 (C-1"), 132.79 (-CH=CH-Ar), 132.70 (C-9a), 131.62 (C-

-4'), 129.41 (C-2'/C-6'), 128.45 (C-3'/C-5'), 125.52 (C-8), 120.05 (C-9), 117.34 (C-6) 115.77 (-CH=CH-Ar), 107.39 (C-3a), 104.49 (C-2"/C-6"), 95.59 (C-7), 60.99 (C-4"-OCH₃), 56.19 (C-3"/C-5"-OCH₃); HRMS: (m/z) M⁺ calcd. for C₂₉H₂₂O₇: 482.1366, found: 483.1494 (M⁺+H).

(E)-3-(3,4,5-Trimethoxystyryl)-2-(4-methylbenzoyl)-4H-furo[3,2-c]chromen-4-one (9e)

Colour: yellow, m.p. 181° C, yield 94%, I.R. (KBr, cm⁻¹): 1724 (-O-C=O), 1629 (-C=O); ¹HNMR: (400 MHz, CDCl₃) δ: 8.45 (d, 1H, -CH=CH-Ar, J = 16.4Hz), 7.91-7.94 (m, 3H, -CH=CH-Ar, C₉-H, C₂/C₆-H), 7.83 (d, 1H, -CH=CH-Ar, J = 16.3 Hz), 7.63 (t, 1H, C₇-H, J = 8.4 Hz), 7.50 (d, 1H, C₆-H, J = 8.2 Hz), 7.35-7.41 (m, 3H, C₈-H, C₃/C₅-H), 6.84 (s, 2H, C₂"-H and C₆"-H), 3.88, 3.92 (s, 9H, C₃"-OCH₃, C₄"-OCH₃ and C₅"-OCH₃), 2.48 (s, 1H, C₄'-CH₃); ¹³CNMR (100 MHz, CDCl₃): 183.55 (-C=O), 157.72 (C-9b), 153.77 (C-5", C-3"), 153.41 (C-4"), 148.36 (C-5a), 143.87 (C-4'), 140.67 (C-1"), 135.64 (C-1'), 131.77 (-CH=CH-Ar), 129.75 (C-2'/C-6'), 129.24 (C-3'/C-5'), 128.99 (C-3), 124.82 (C-8), 121.92 (C-9), 118.92 (-CH=CH-Ar), 117.32 (C-6), 115.57 (C-3a), 111.89 (C-9a), 104.46 (C-2", C-6") 60.96 (C-4"-OCH₃), 56.21 (C-3"/C-5"-OCH₃), 21.74 (C-4'-CH₃); HRMS: (m/z) M⁺ calcd. for C₃₀H₂₄O₇: 496.1522, found: 497.1768 (M⁺+H).

(E)-3-(3,4,5-Trimethoxystyryl)-2-(4-methoxybenzoyl)-4H-furo[3,2-c]chromen-4-one (9f)

Colour: yellow, m.p. 183° C, yield 94%, I.R. (KBr, cm⁻¹): 1746 (-O-C=O), 1637 (-C=O); ¹HNMR: (400 MHz, CDCl₃) δ: 8.45 (d, 1H, -CH=CH-Ar, J = 16.4 Hz), 8.07 (d, 2H, C₂' / C₆'-H, J = 8.8 Hz), 7.96 (d, 1H, C₉-H, J = 7.7 Hz), 7.82 (d, 1H, -CH=CH-Ar, J = 16.4 Hz), 7.66 (t, 1H, C₇-H, J = 7.1 Hz), 7.53 (d, 1H, C₆-H, J = 8.4 Hz), 7.42 (t, 1H, C₈-H, J = 7.6 Hz), 7.07 (d, 2H, C₃'/C₅'-H, J = 8.8 Hz), 6.86 (s, 2H, C₂" and C₆"-H), 3.95, 3.94, 3.92, 3.90 (s, 12H, C₄'-OCH₃, C₃", C₄"-OCH₃ and C₅"-OCH₃); ¹³CNMR (100 MHz, CDCl₃): 183.52 (-C=O), 156.51 (C-3", C-5"), 153.19 (C-4"), 141.43 (C-1"), 139.66 (C-4'), 133.25 (-CH=CH-Ar), 131.90 (C-1'), 127.57 (C-2', C-6'), 126.23 (C-3', C-5'), 123.07 (-CH=CH-Ar), 121.23 (C-9), 116.23 (C-6), 105.51 (C-2", C-6"), 60.79 (C-4"-OCH₃), 56.11 (C-3", C-4', C-5"-OCH₃); HRMS: (m/z) M⁺ calcd. for C₃₀H₂₄O₈: 512.1471, found: 513.1692 (M⁺+H).

Pharmacological/biological assays

Anticancer activity culture

To relate the degree of propagation of a tumor cell line in the absence and presence of the test substances furo[3,2-c]pyran-4-ones (**4**, **5**) and furo[3,2-c]chromen-4-ones (**8**, **9**), generally later a listed time the sulphorhodamine B (SRB) [1] evaluation is carried out. This trusts on the uptake of the negatively charged pink aminoxanthine dye, sulphorhodamine B (SRB) using amino acids (basic) in the cells. Larger the number of cells, the more amount of dye is taken up then after fixing, the cells are lysed and the released dye will give a more intense colour and better absorbance. Cells were cultured in a 96 well plate. Inoculation densities per well varies from cell line to cell line under investigation. 100 µL of cell suspension was plated. The cells were then treated with 50 µM concentration of test compound complete growth medium (RPMI-1640) for 48 hours. After 48 hours of incubation at 37°C, cells were washed for 1h with ice cold TCA at 4°C. After fixing the cells in fixative, the plates were rinsed with water for three times and it was further allowed to air dry. 100µL of 0.4% SRB dye was added for half an hour at room temperature after drying. 1% v/v acetic acid is then used for washing plates 3 times to remove any unbound SRB. The plates are then kept for drying at room temperature, after drying the bound dye was solubilized by adding 100 µL of 10mM TRIS buffer having pH- 10.4 to each well. The plates were shaking for 5 minutes to solubilize the dye. Finally, OD was taken at 540 nm in a microplate reader. IC₅₀ was determined by plotting OD against concentration.

$$\text{The \% of cell viability} = \frac{\text{Absorbance of treated cells} - \text{Absorbance of blank}}{\text{Absorbance of control cells} - \text{Absorbance of blank}} \times 100$$

$$\% \text{ Growth inhibition} = 100 - \% \text{ cell viability}$$

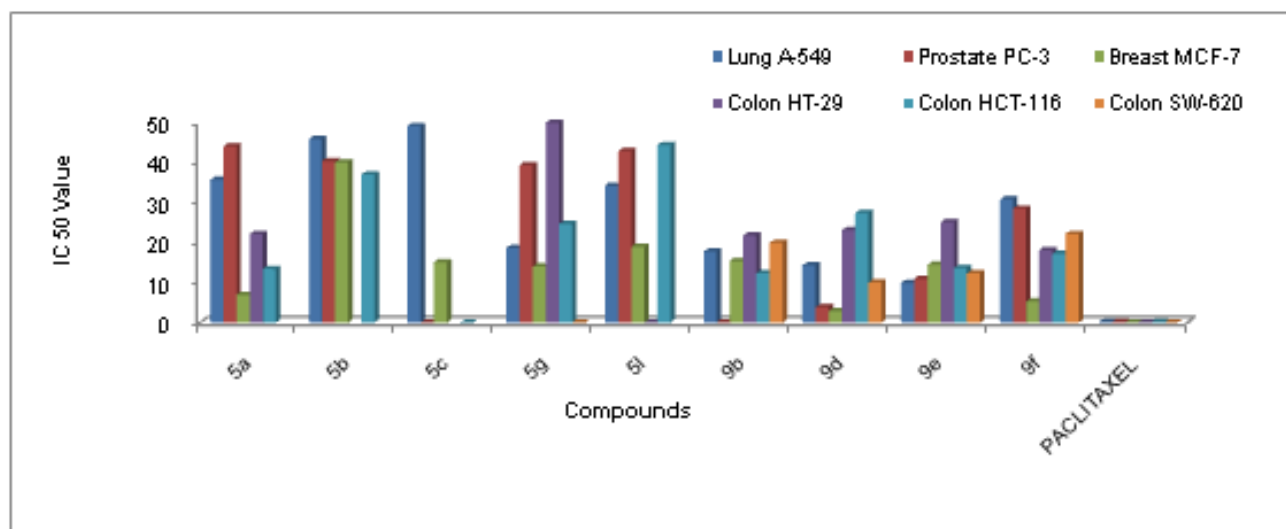


Fig.1. Graphical representation of IC₅₀ value of different compound for different cancer cell lines.

4.2.2 Antimicrobial activity cultures

The antimicrobial screening of the synthesized furo[3,2-c]pyran-4-ones (**4,5**) and furo[3,2-c]chromen-4-ones (**8, 9**) were carried out against three bacterial strains i.e. *Bacillus subtilis* (MTCC-441), *Staphylococcus aureus* (MTCC 3160) and *Escherichia coli* (MTCC-443) and two fungal strains i.e. *Candida albicans* (MTCC-227) and *Aspergillus niger* (MTCC-281) employing tube dilution method [2]. Dilutions of standard and test compounds were done in double strength nutrient broth I.P. and Sabouraud dextrose broth I.P. for bacteria and fungi. The testing compounds were incubated for 24 h at 37°C±1°C for all bacteria and 25°C for seven days for fungi *A. niger* or for 48 h at 37°C±1°C for fungi *C. albicans*. Results of activities were noted in terms of MIC. Graphical representation of antibacterial study is represented in Fig. 2.

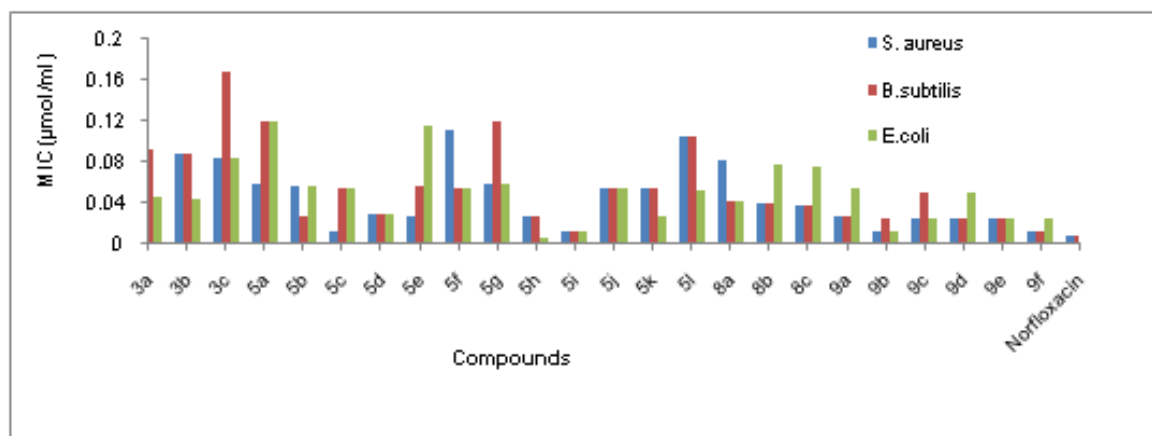


Fig. 2. Show the graphical representation of antibacterial study of synthesized compounds (4, 5, 8 and 9).

Graphical representation of antifungal study is represented in **Fig.3.**

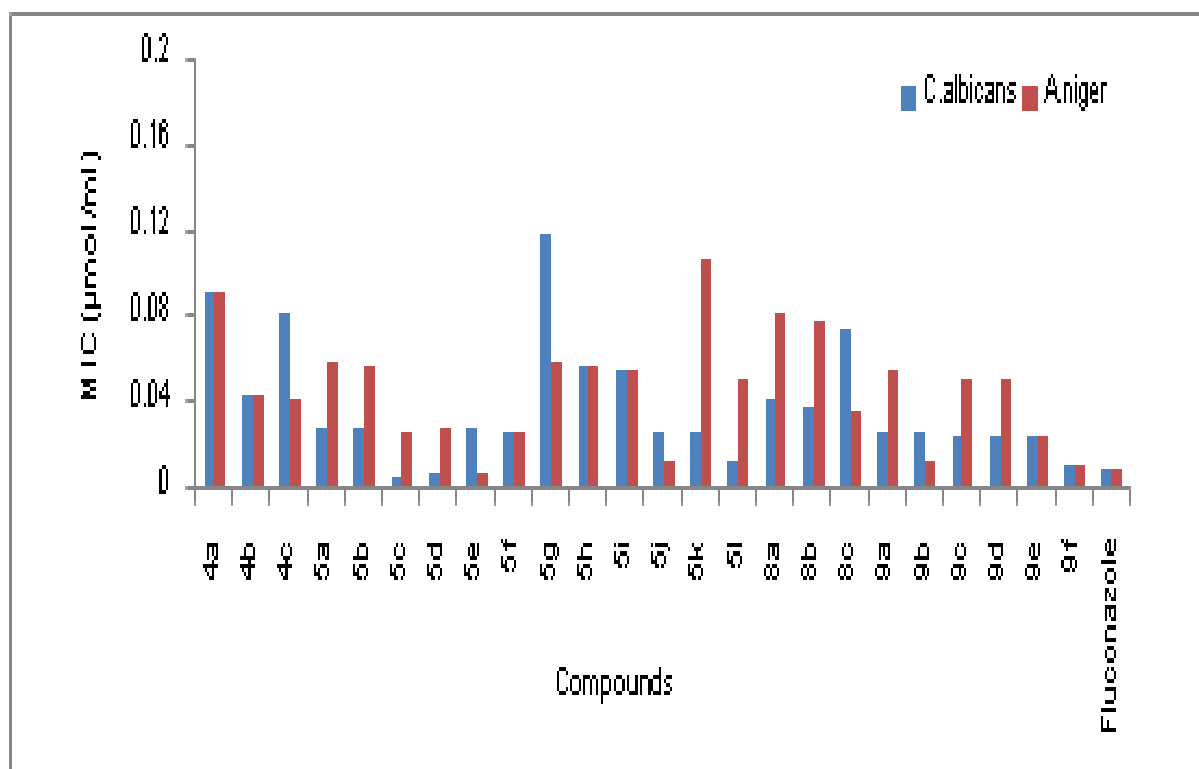


Fig. 3 The graphical representation of antifungal study of synthesized compounds (4, 5, 8, 9)

^1H , ^{13}C NMR, 2D and HRMS spectra of synthesized compounds

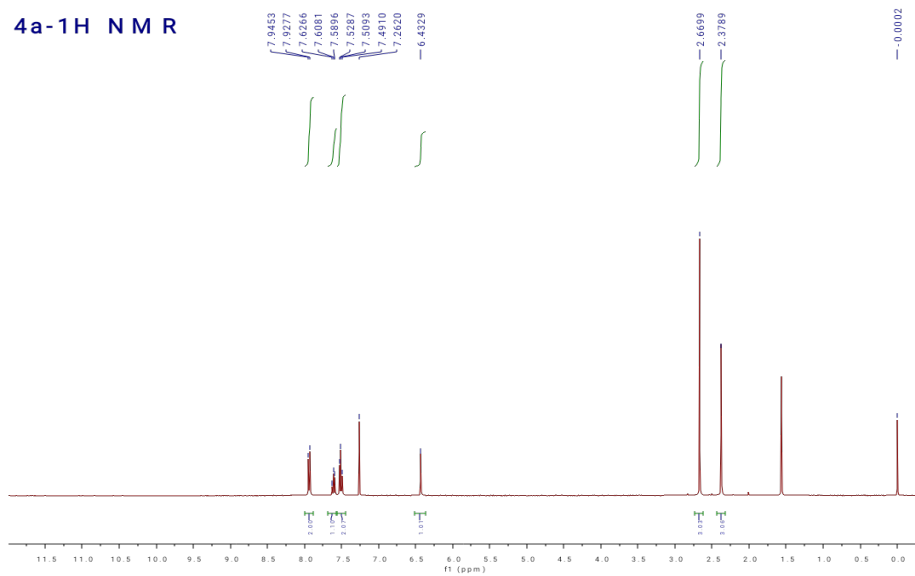


Figure S1. ^1H NMR of compound 4a

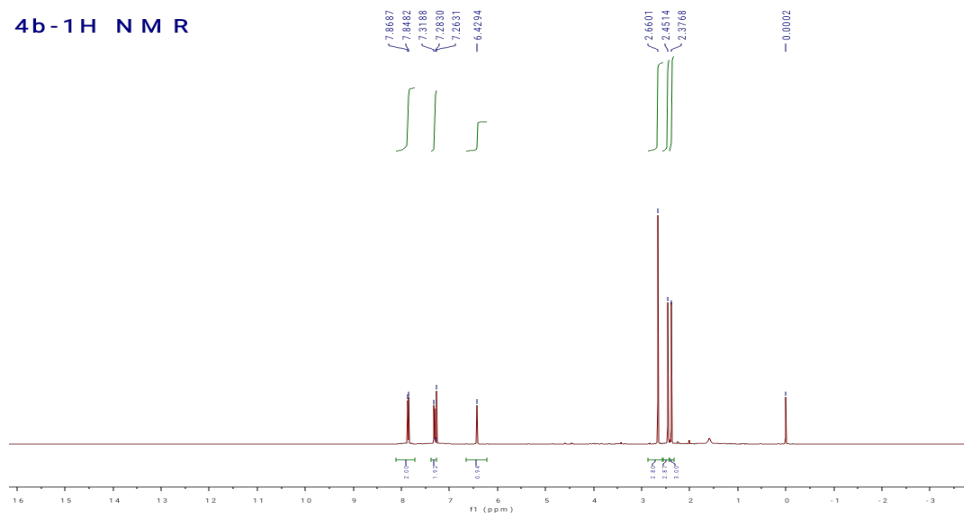


Figure S2. ^1H NMR of compound 4b

4c-1H NMR

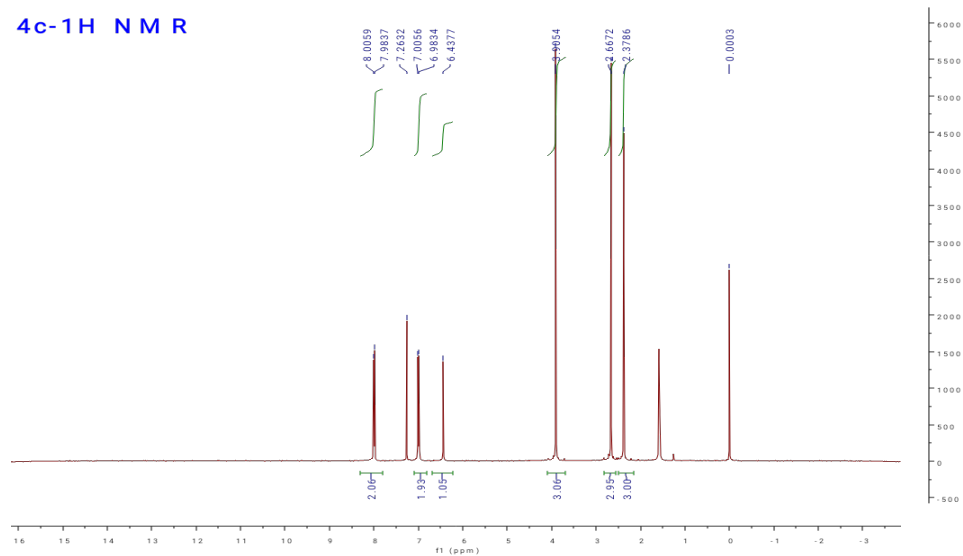


Figure S3. ¹H NMR of compound 4c

5a- 1H NMR

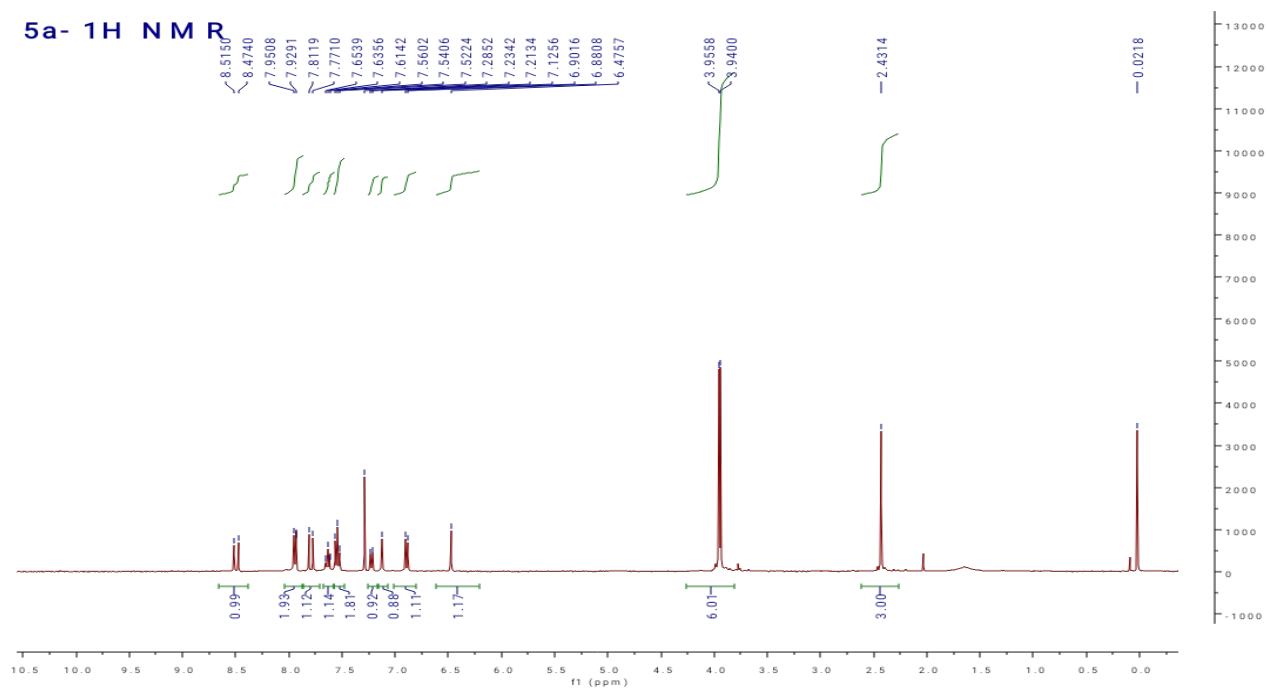


Figure S4. ¹H NMR of compound 5a

5b-1H NMR

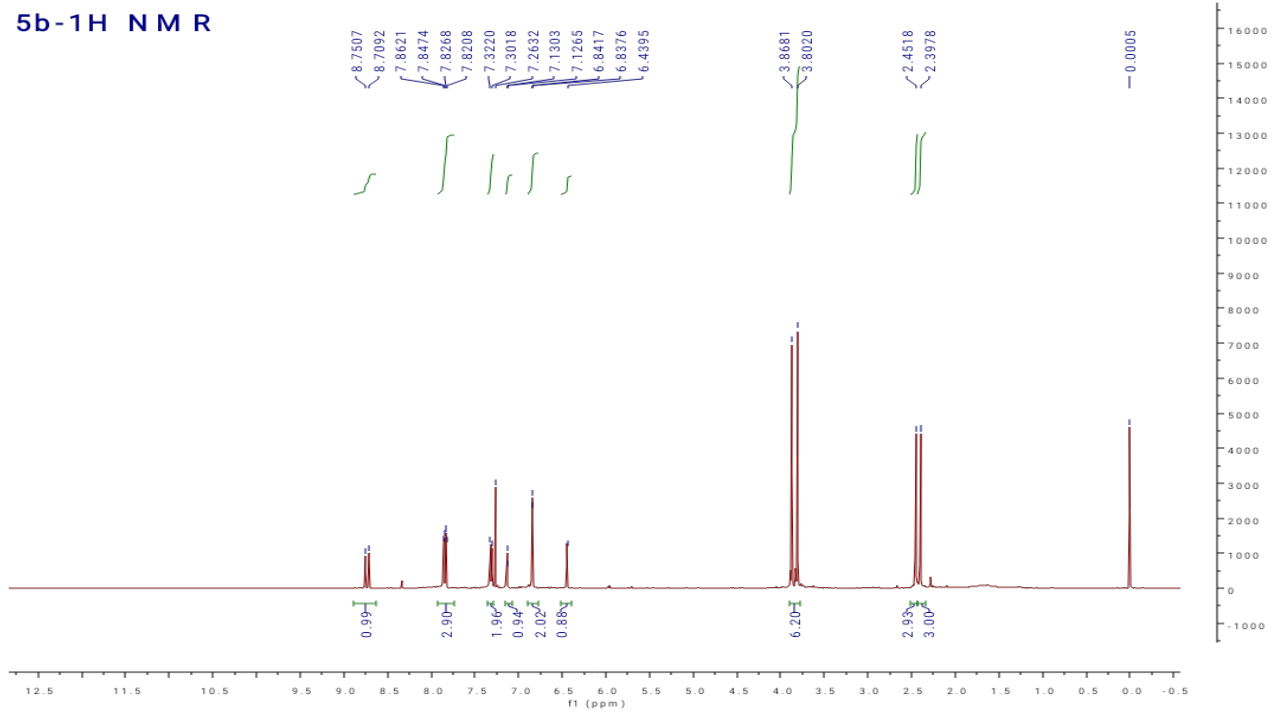


Figure S5. ¹H NMR of compound 5b

5c-1H NMR

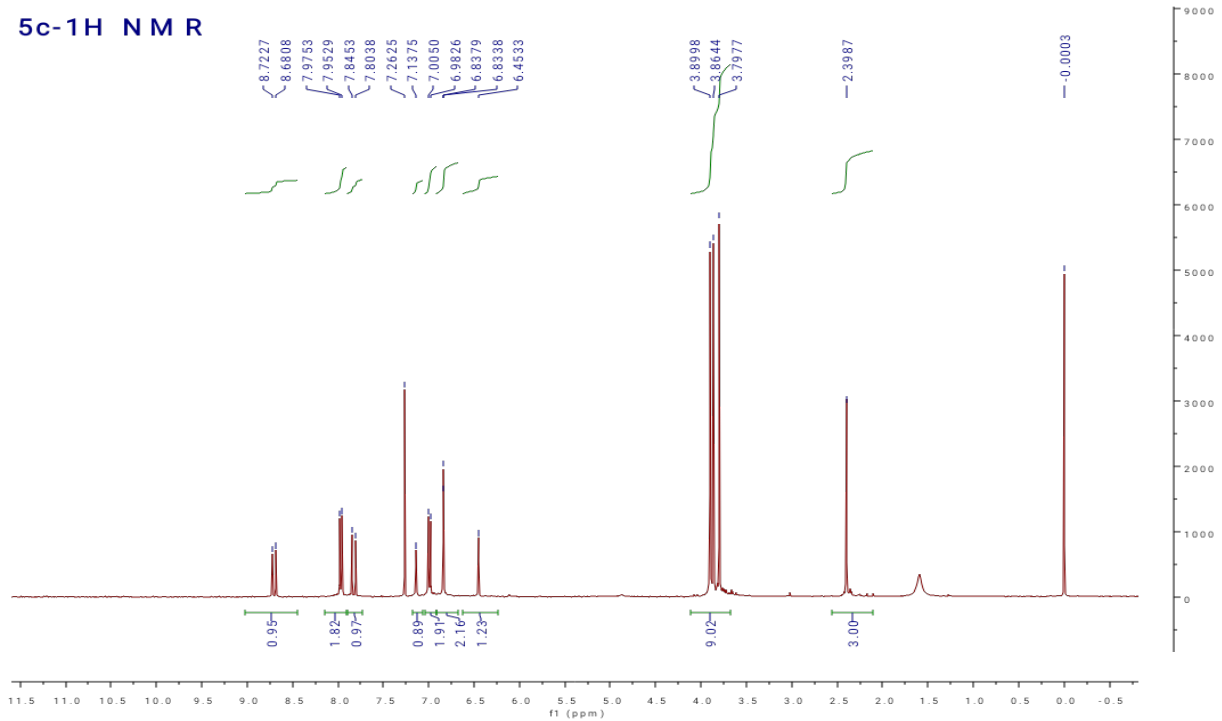


Figure S6. ¹H NMR of compound 5c

5d-1H-NMR

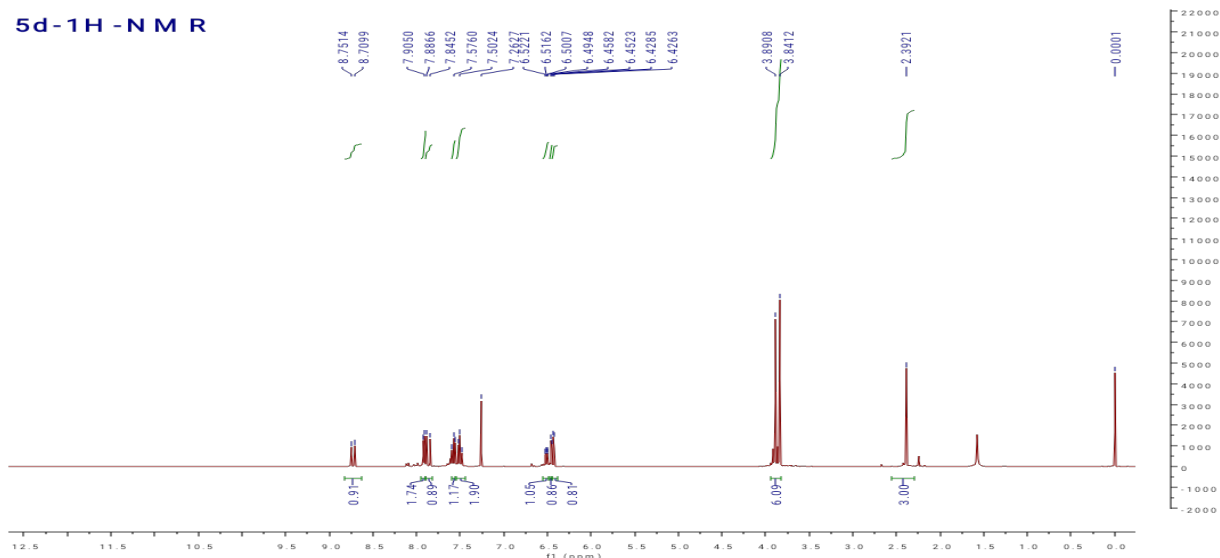


Figure S7. ¹H NMR of compound 5d

5e-1H-NMR

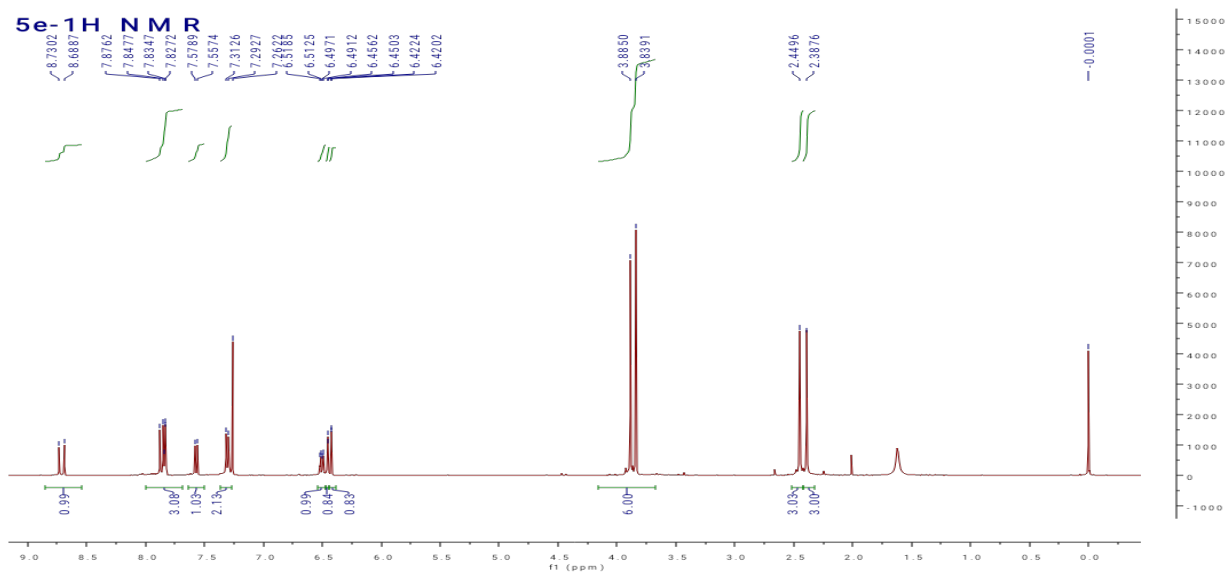


Figure S8. ¹H NMR of compound 5e

5f-1H-NMR

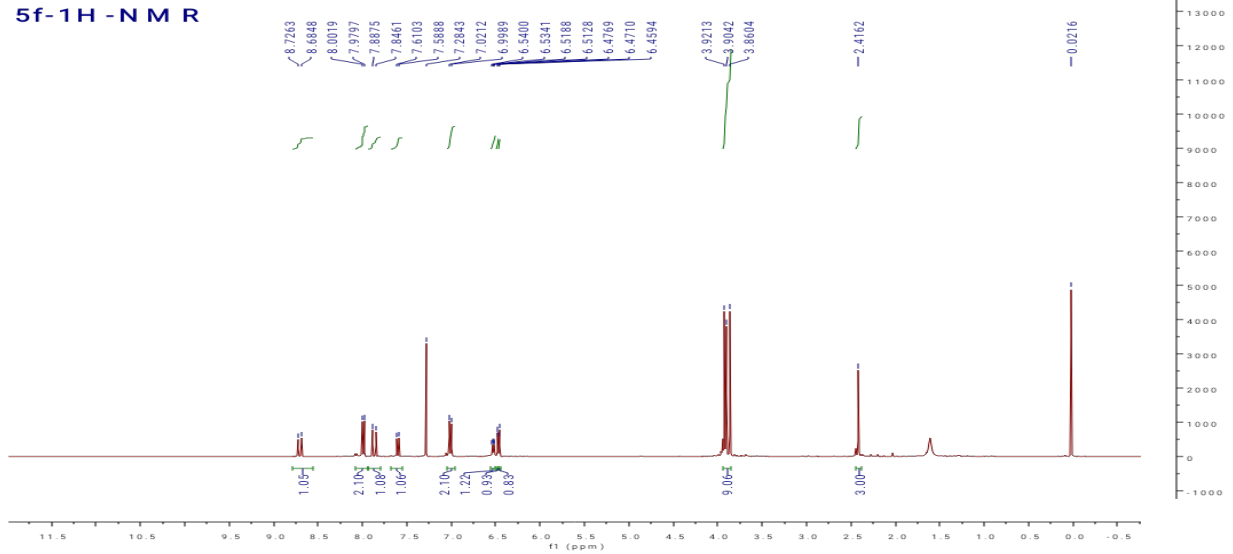


Figure S9. ¹H NMR of compound 5f

5g-1H NMR

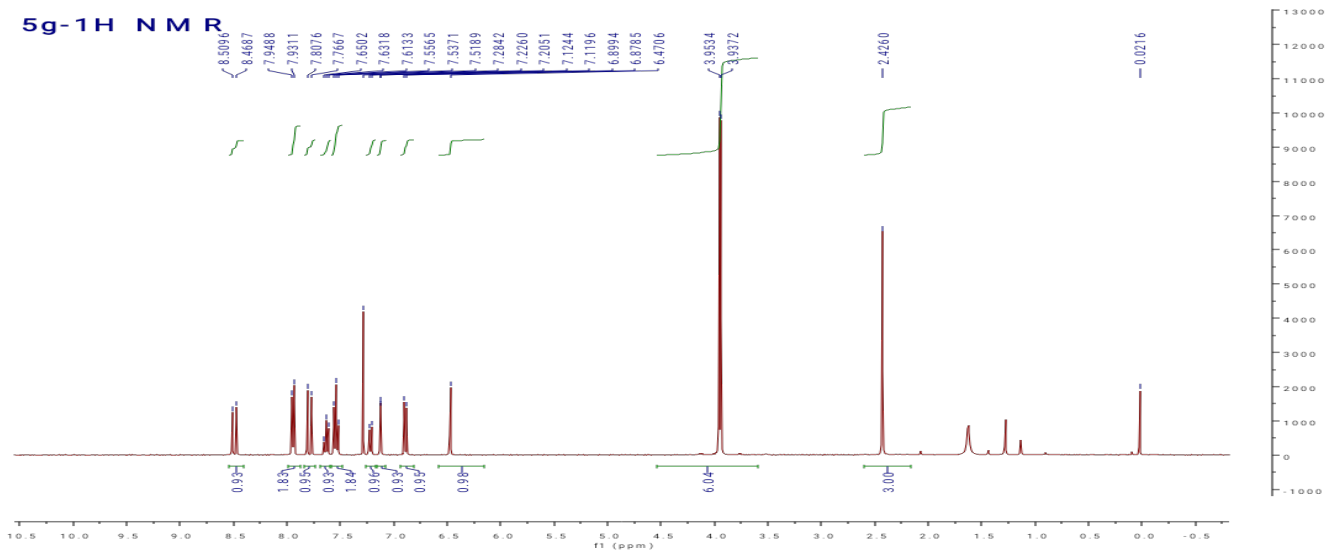


Figure S10. ¹H NMR of compound 5g

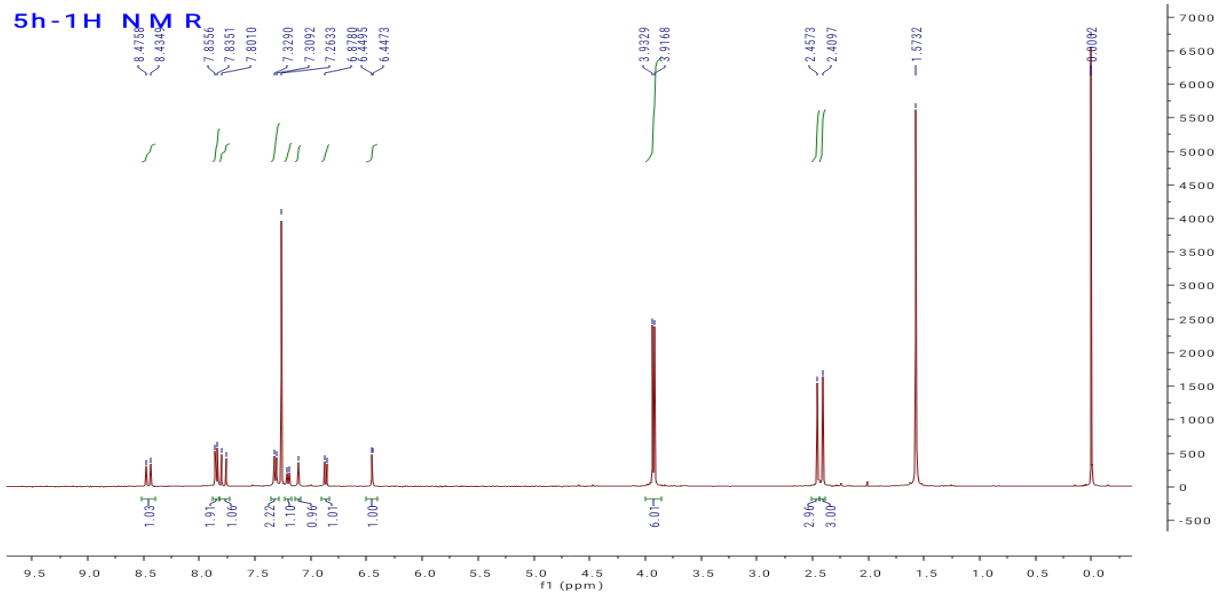


Figure S11. ^1H NMR of compound 5h

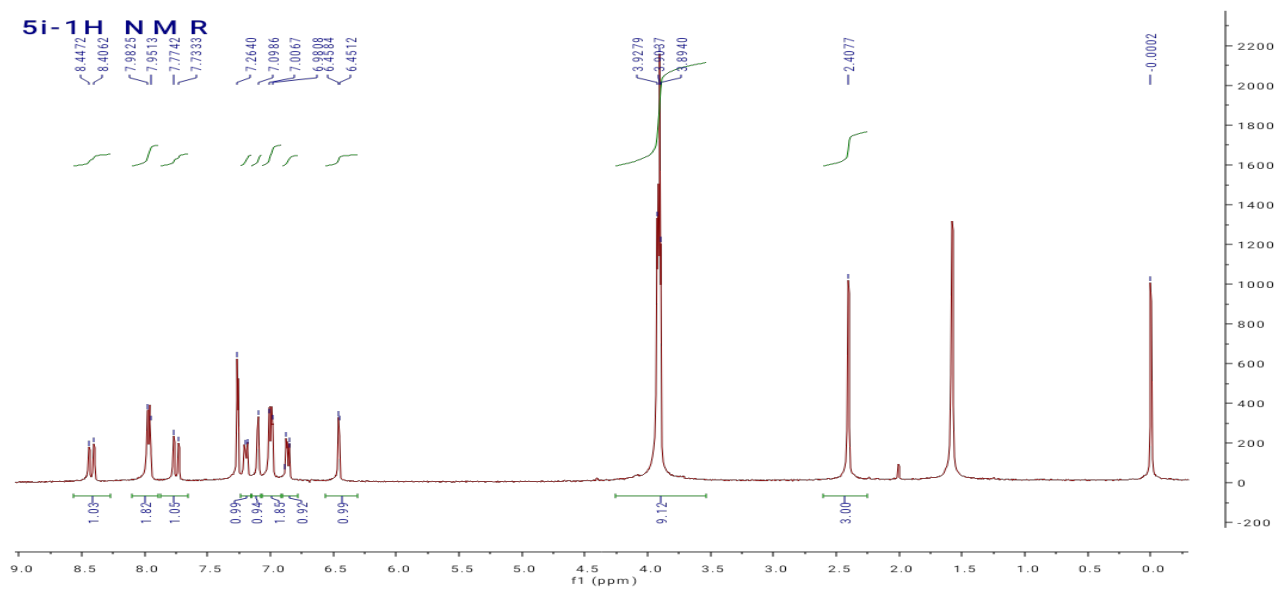


Figure S12. ^1H NMR of compound 5i

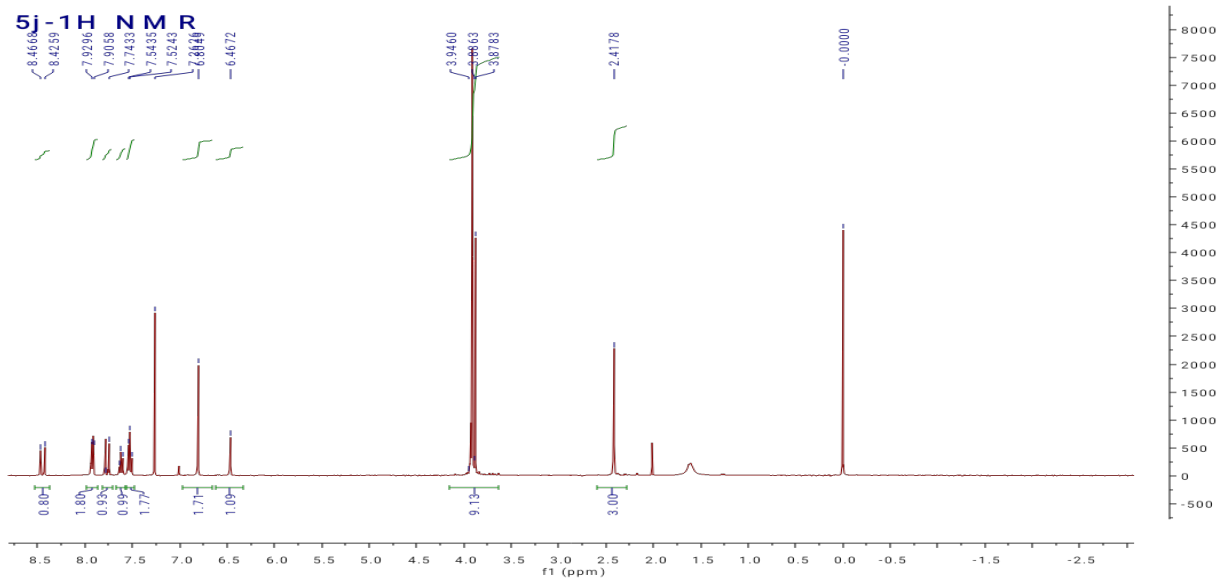


Figure S13. ¹H NMR of compound 5j

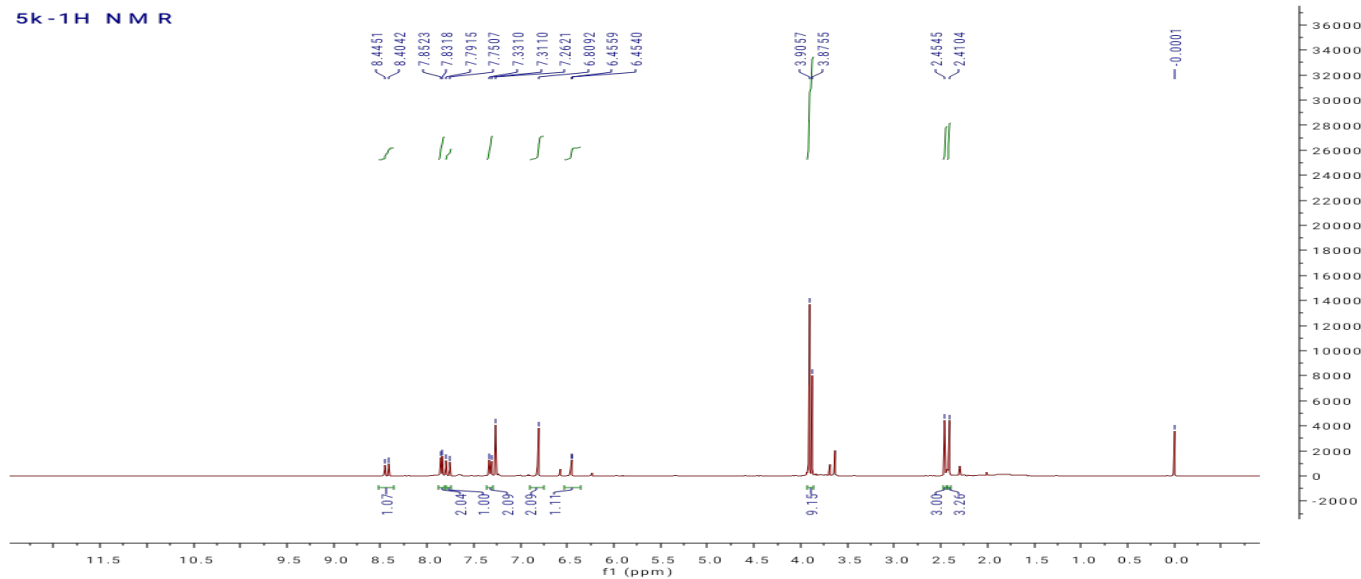


Figure S14. ¹H NMR of compound 5k

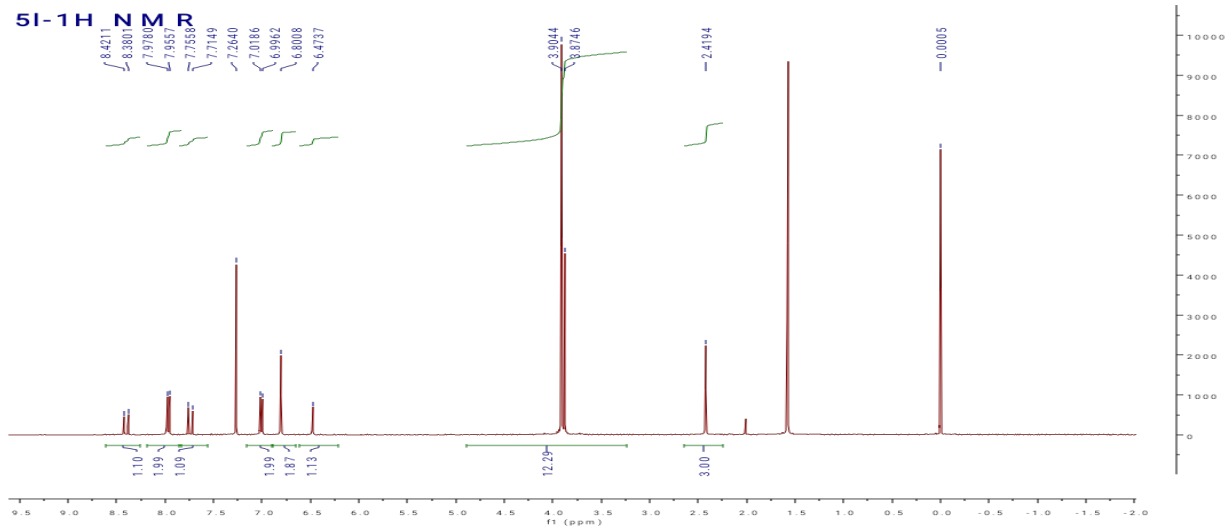


Figure S15. ¹H NMR of compound 5I

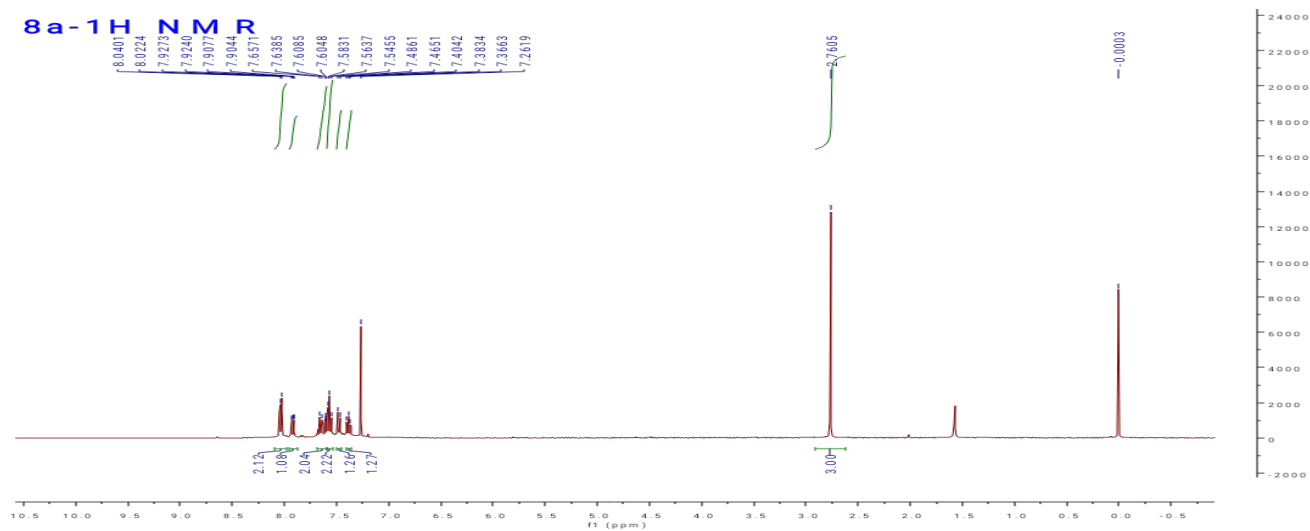


Figure S16. ¹H NMR of compound 8a

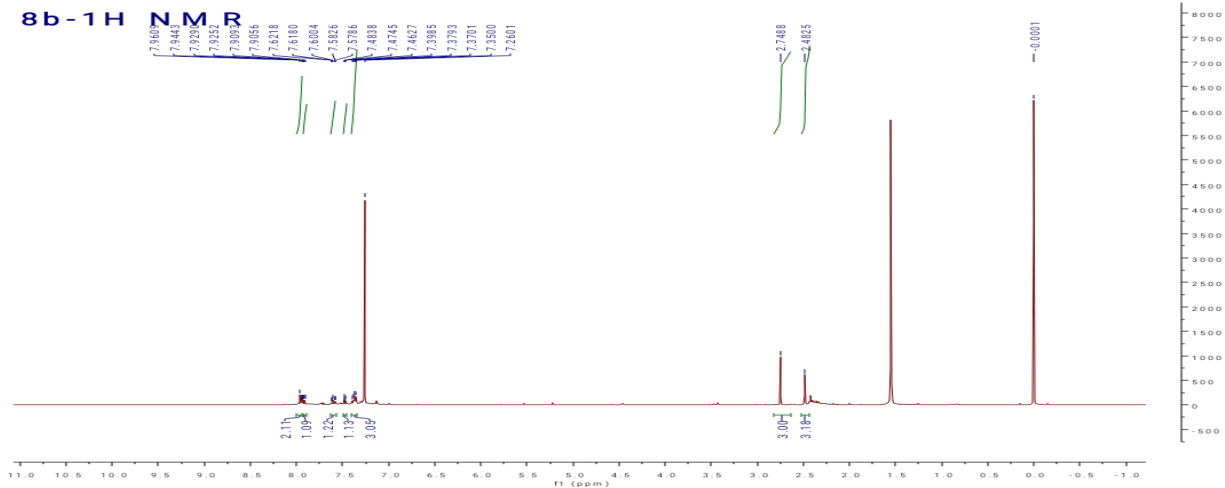


Figure S17. ^1H NMR of compound 8b

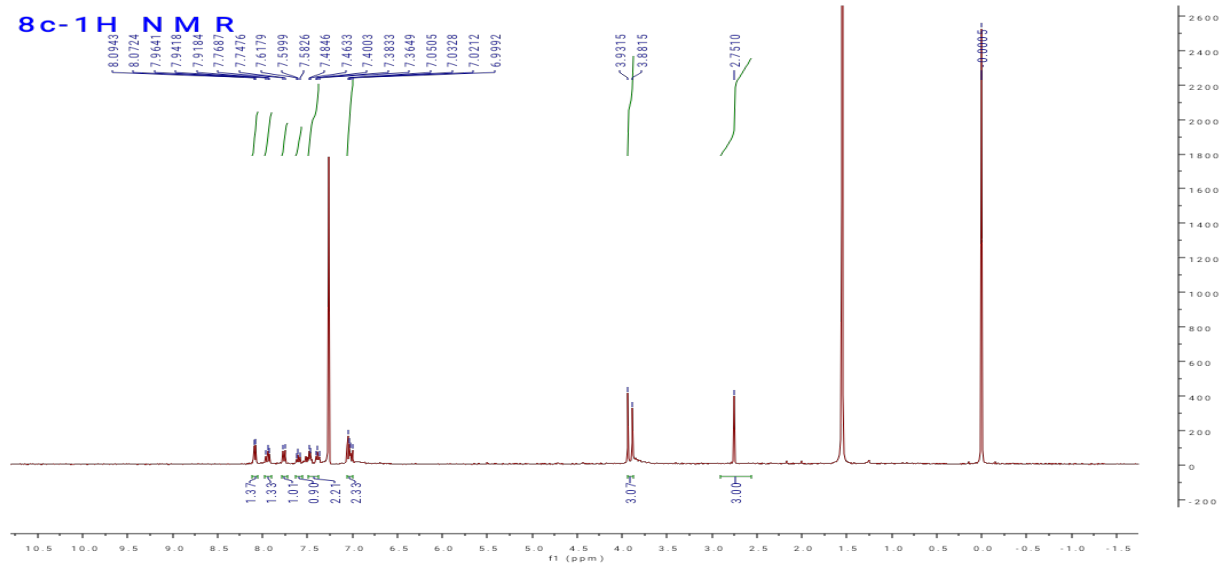


Figure S18. ^1H NMR of compound 8c

9a-1H NMR

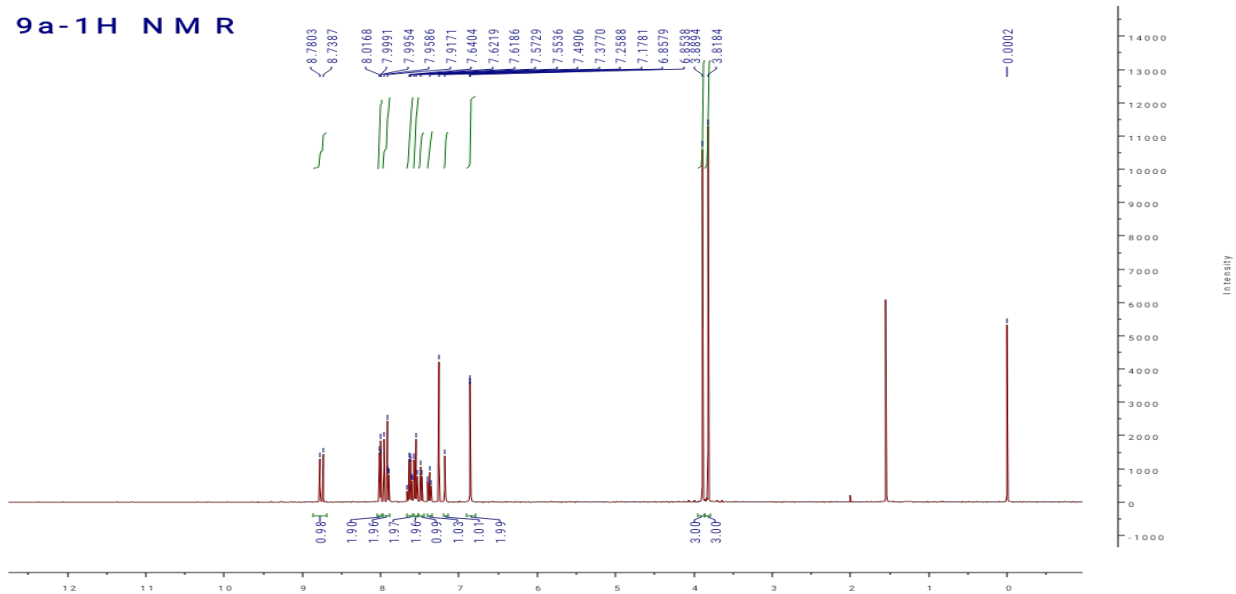


Figure S19. ¹H NMR of compound 9a

9b-1H NMR

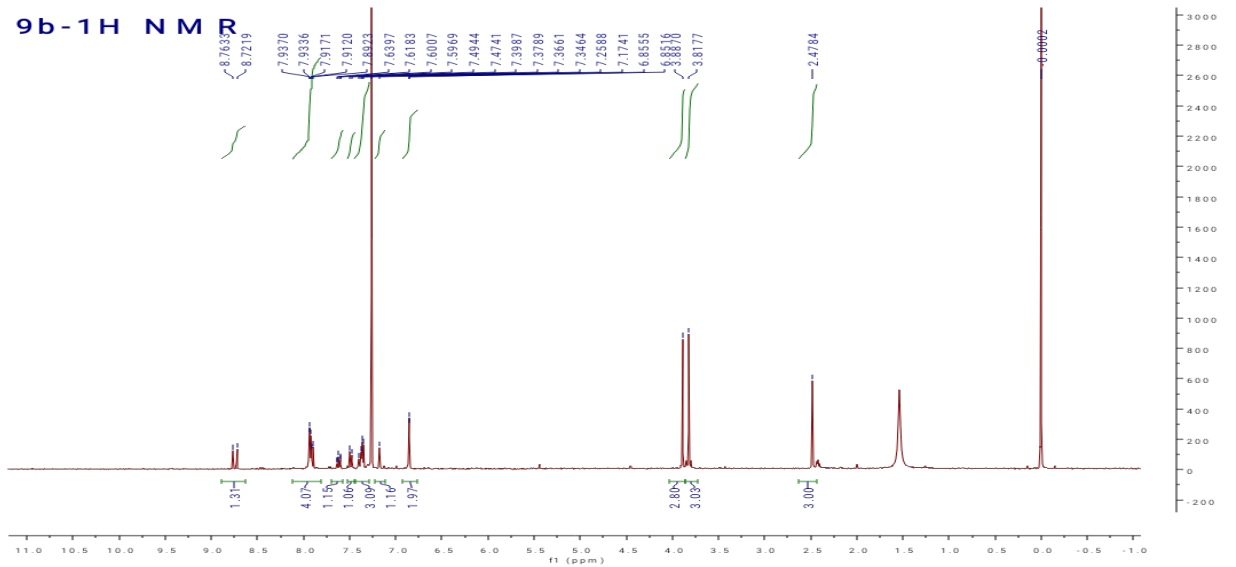


Figure S20. ¹H NMR of compound 9b

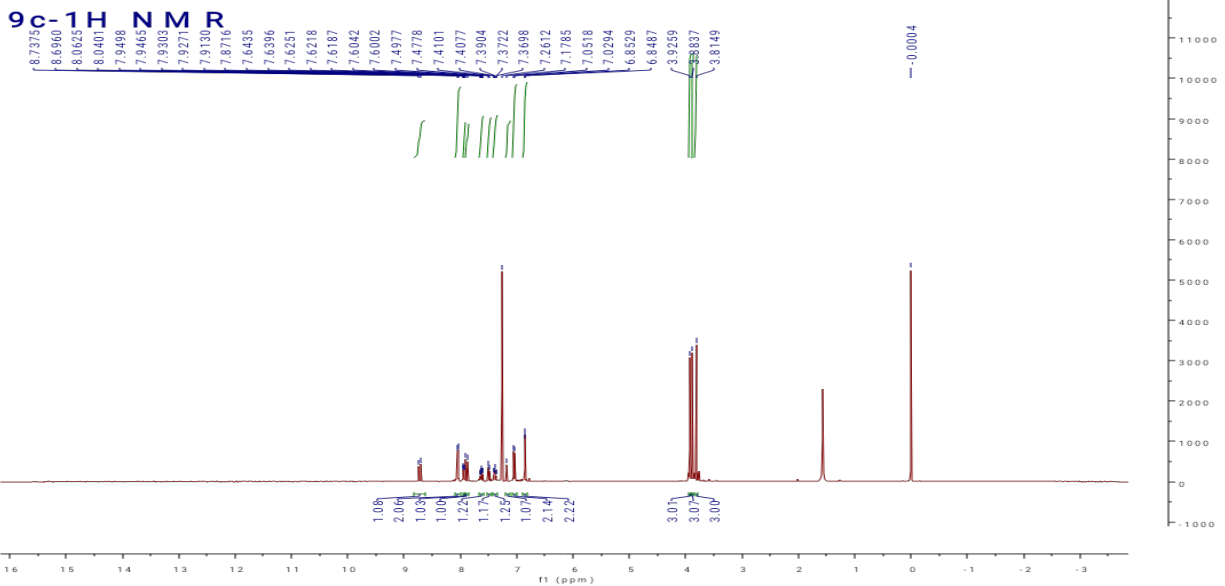


Figure S21. ¹H NMR of compound 9c

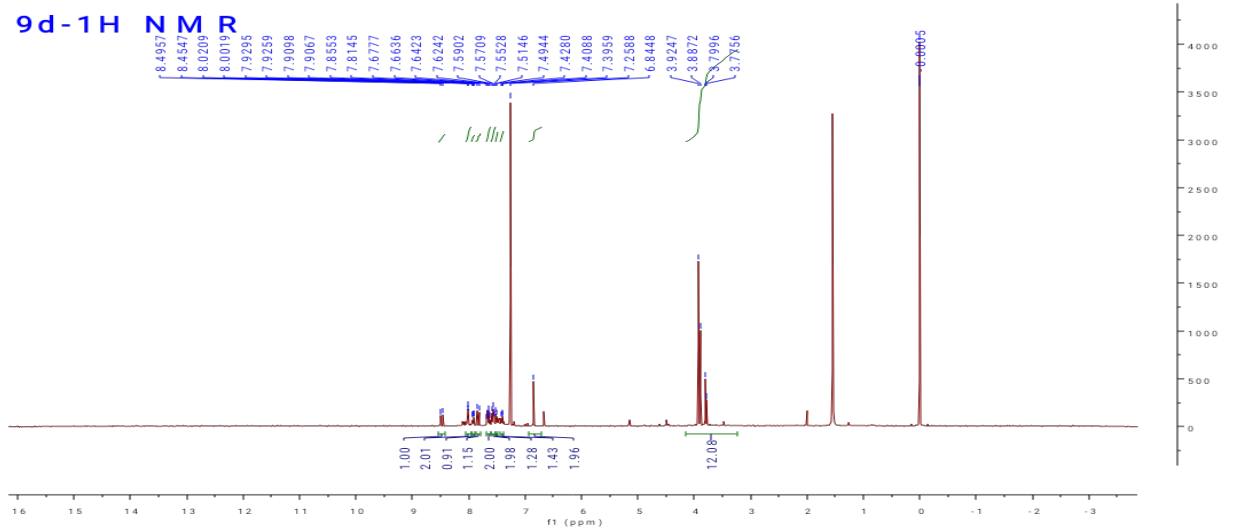


Figure S22. ¹H NMR of compound 9d

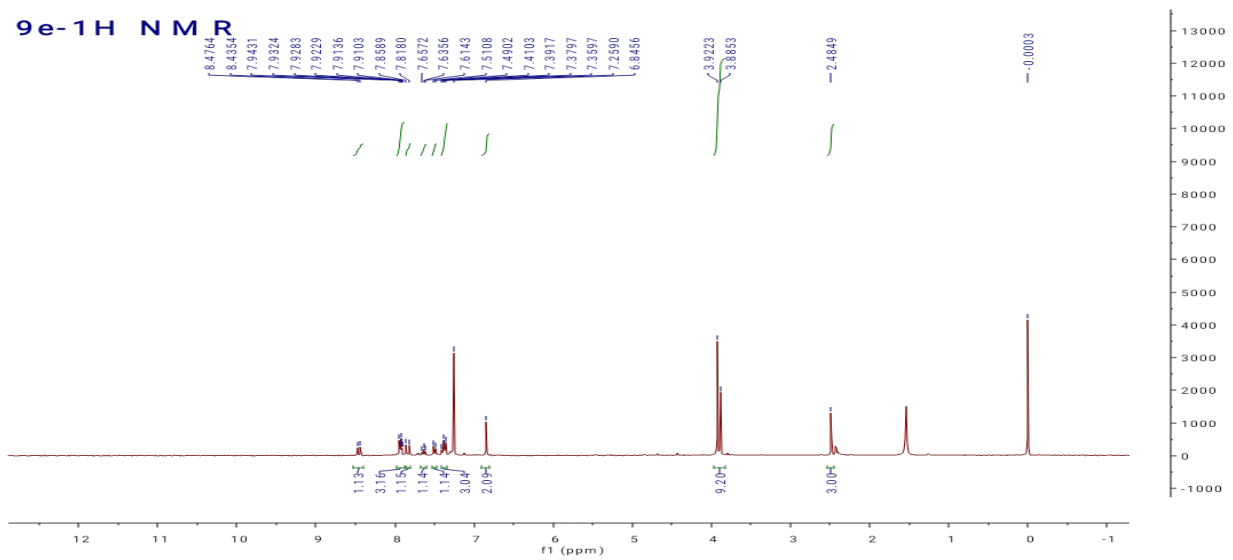


Figure S23. ^1H NMR of compound 9e

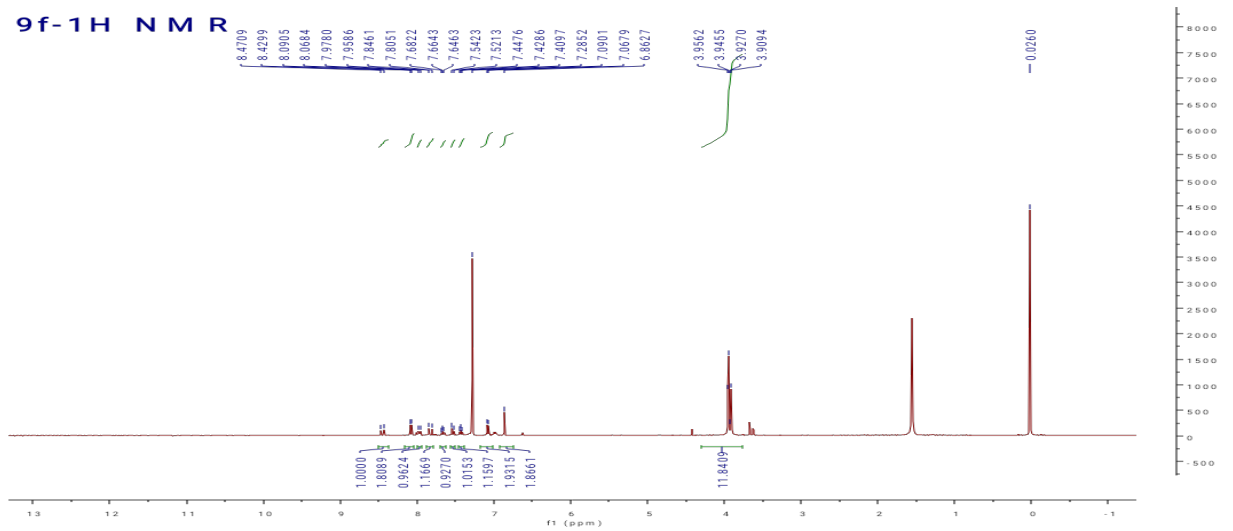


Figure S24. ^1H NMR of compound 9f

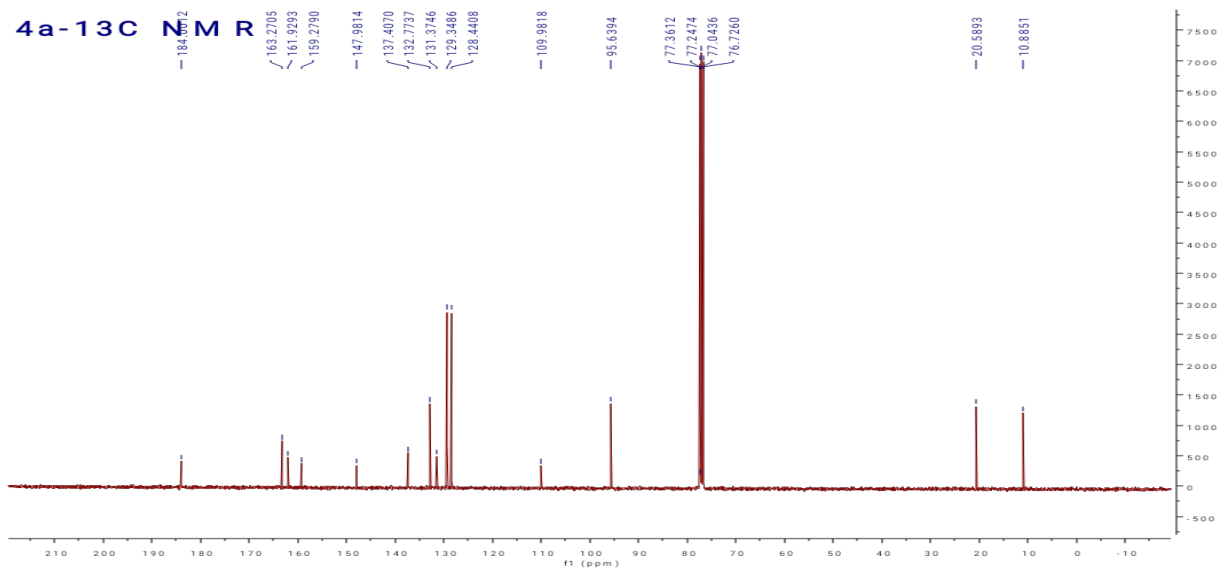


Figure S 25. ¹³C NMR of compound 4a

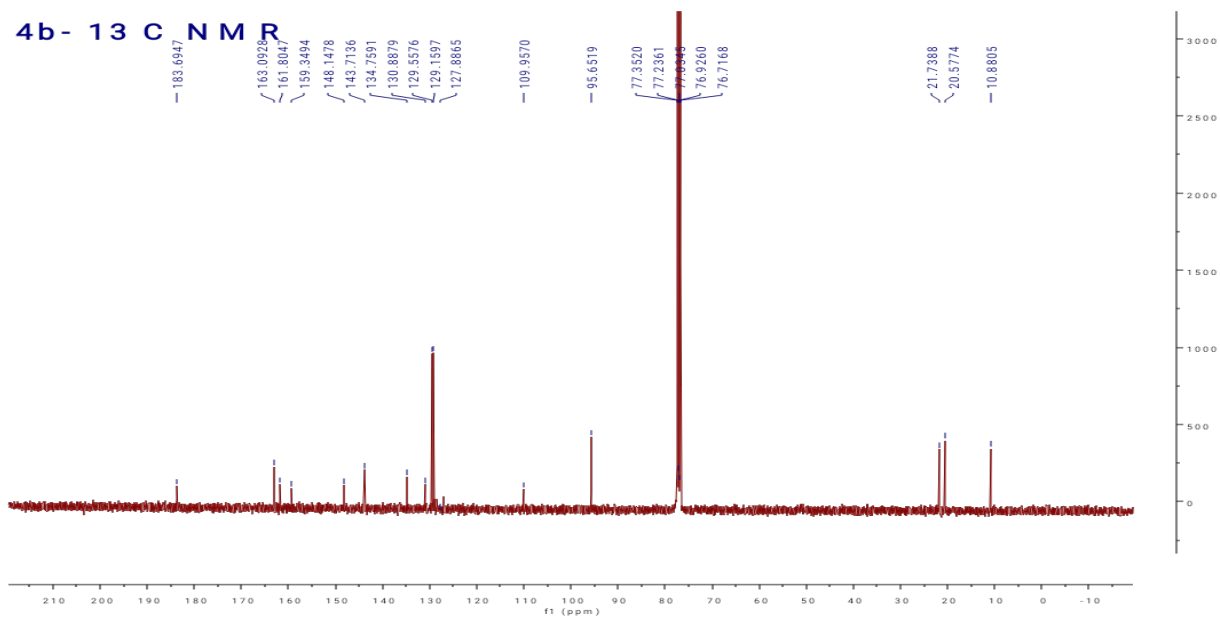


Figure S 26. ¹³C NMR of compound 4b

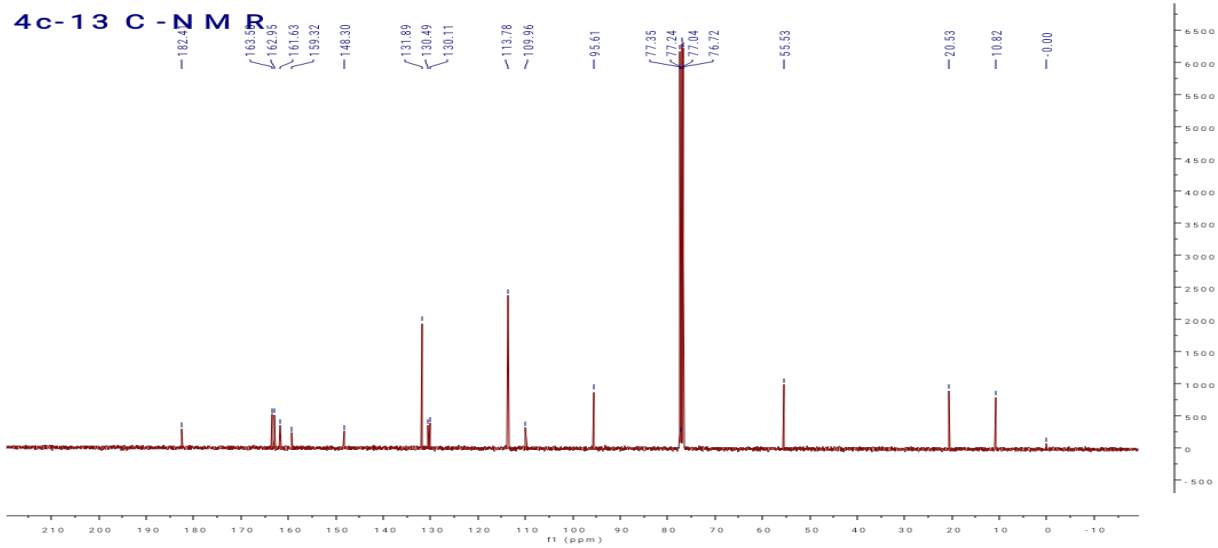


Figure S 27. ¹³C NMR of compound 4c

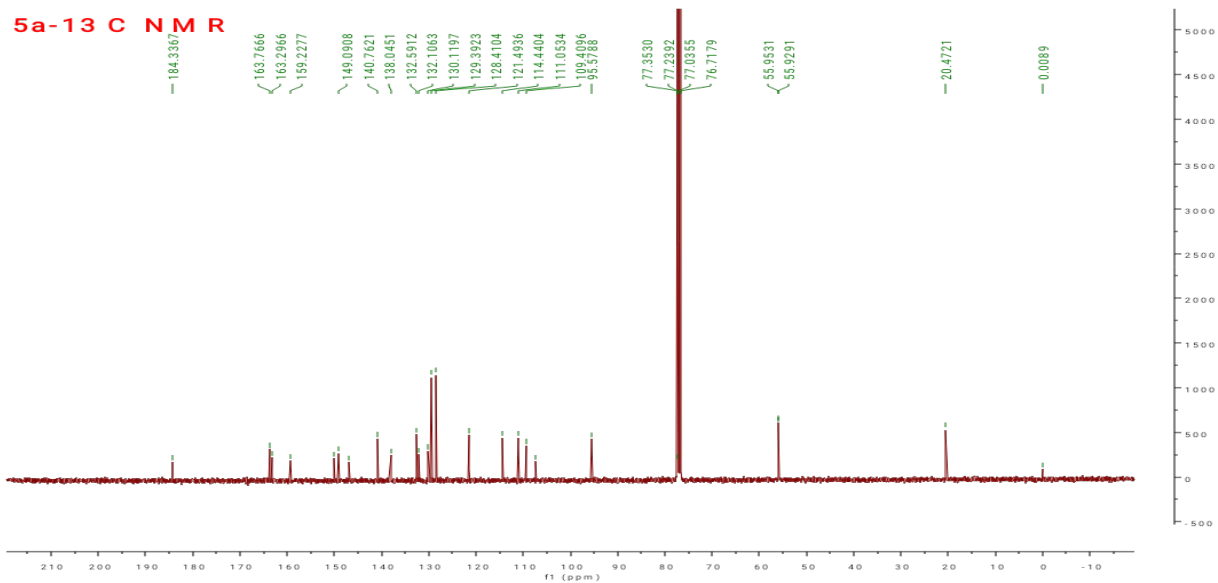


Figure S 28. ¹³C NMR of compound 5a

5b-¹³C NMR

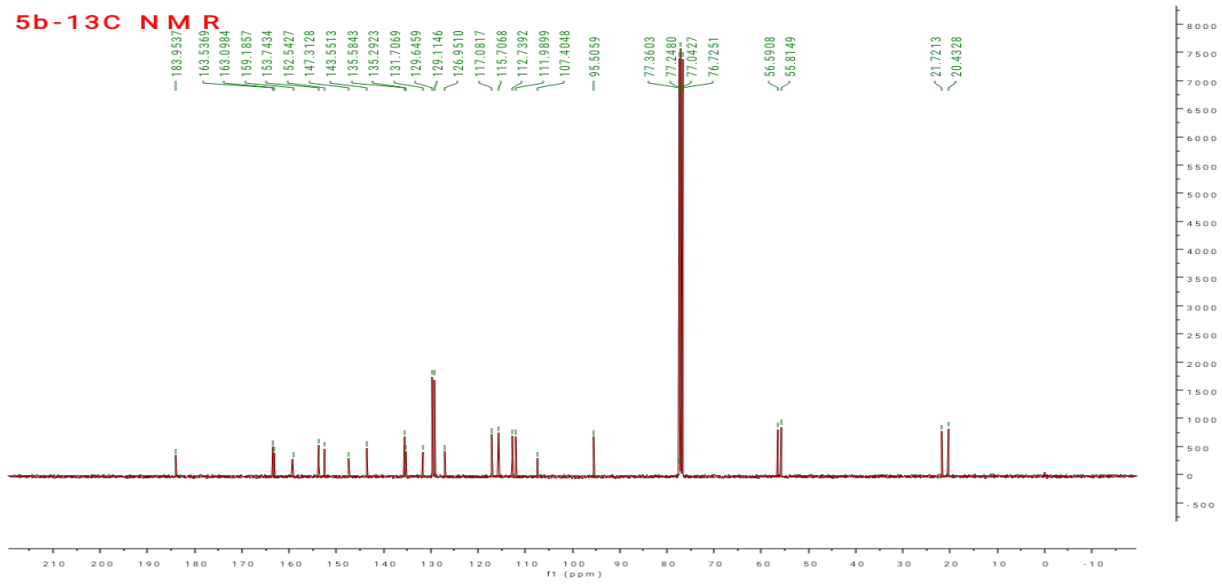


Figure S 29. ¹³C NMR of compound 5b

5c-¹³C NMR

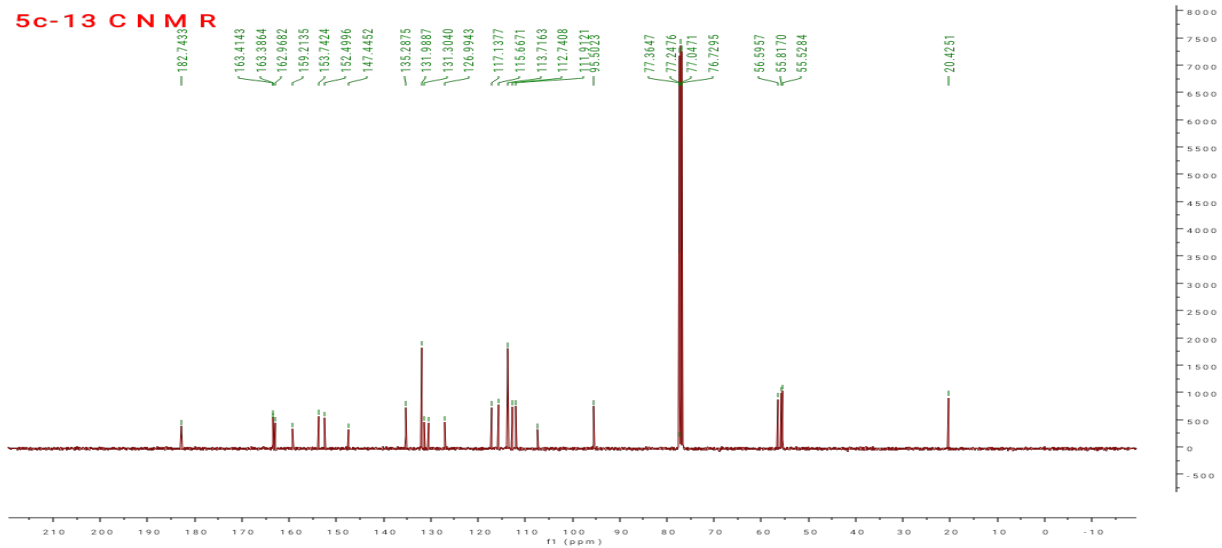


Figure S 30. ¹³C NMR of compound 5c

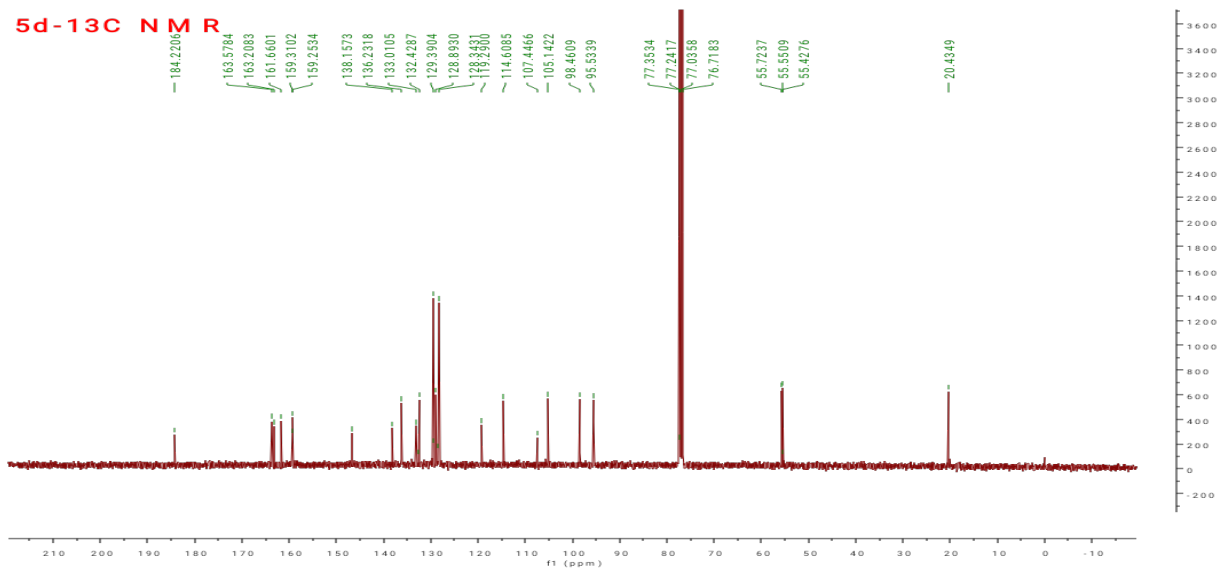


Figure S 31. ¹³C NMR of compound 5d

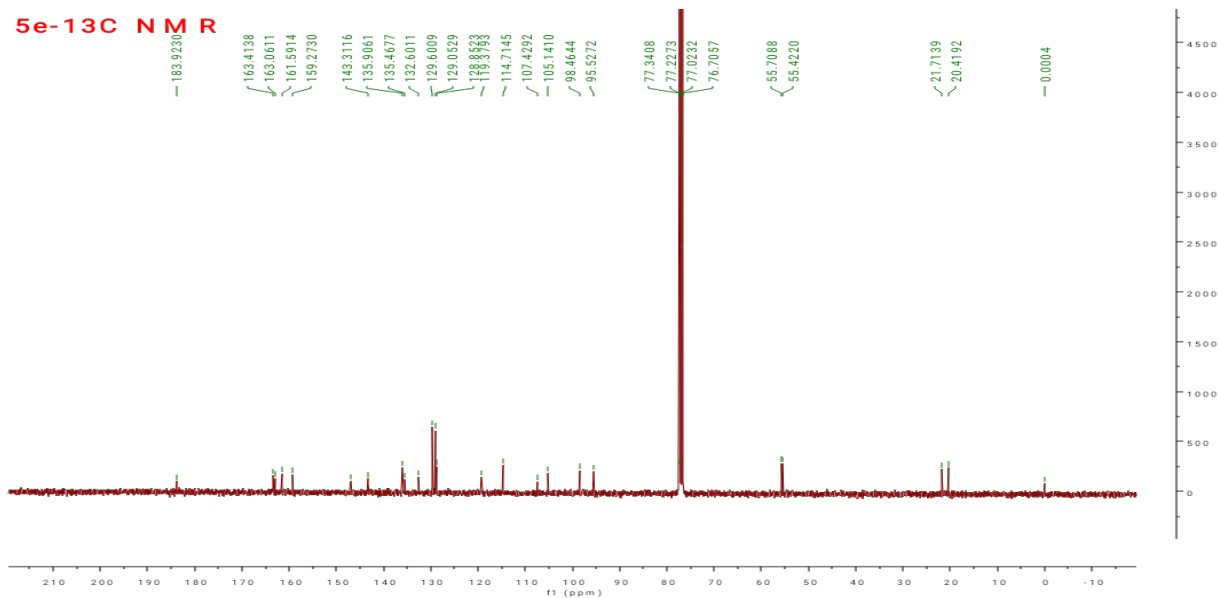


Figure S 32. ¹³C NMR of compound 5e

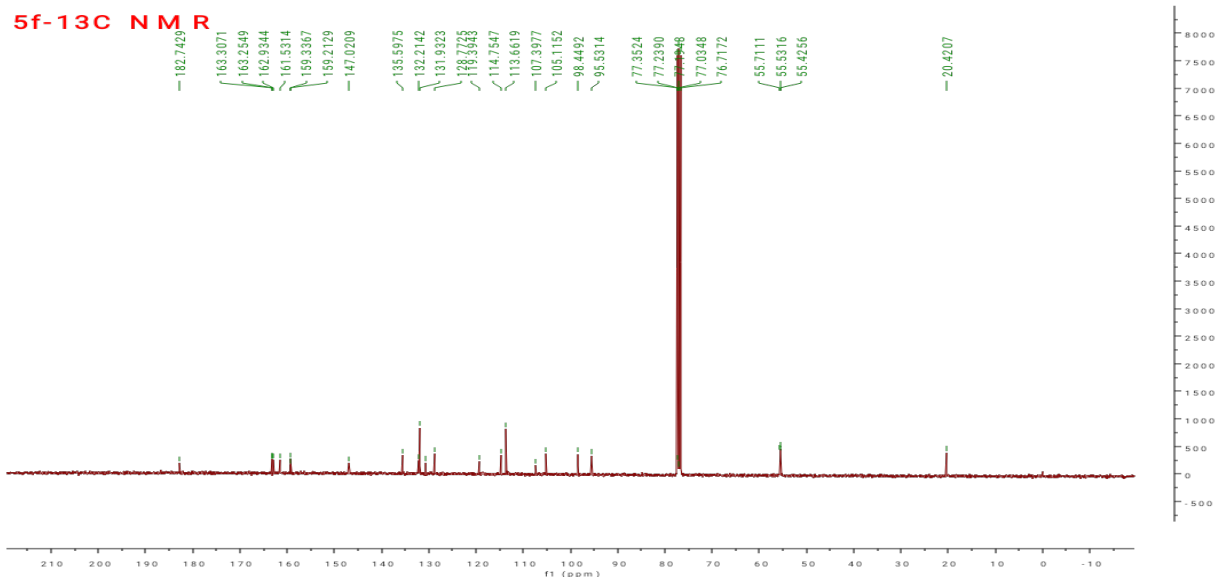


Figure S 33. ¹³C NMR of compound 5f

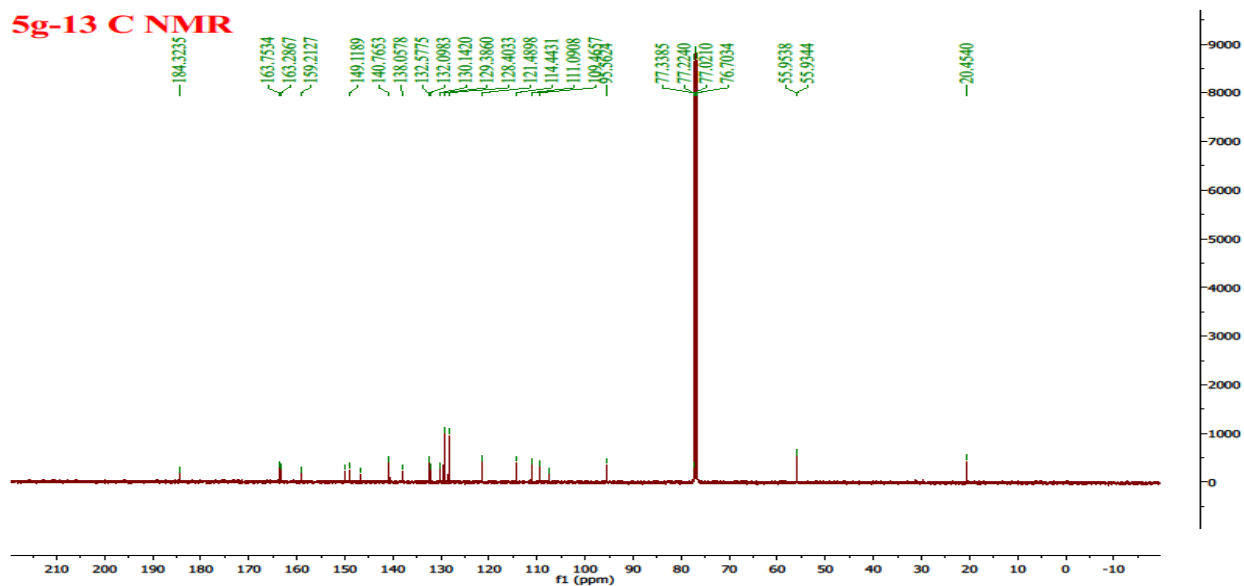


Figure S 34. ¹³C NMR of compound 5g

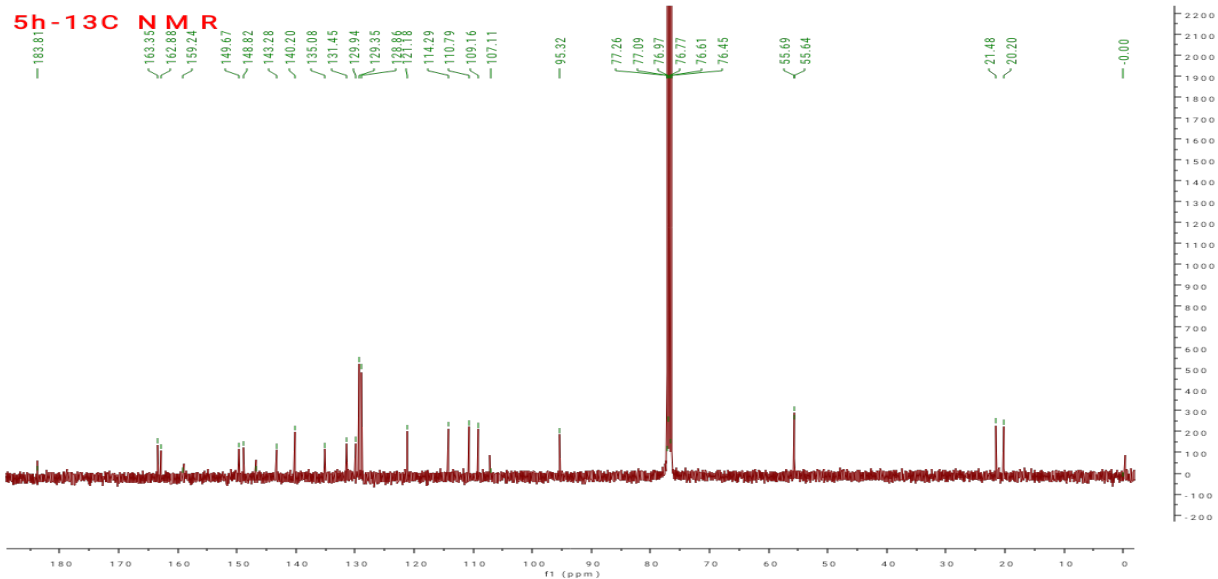


Figure S 35. ¹³C NMR of compound 5h

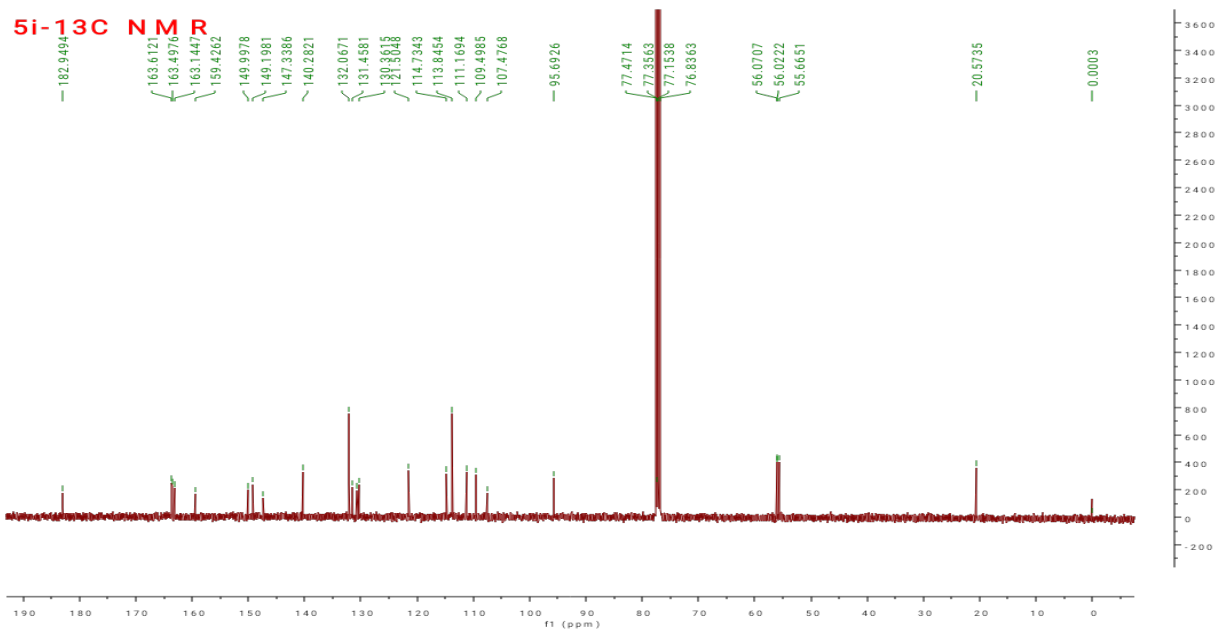


Figure S 36. ¹³C NMR of compound 5i

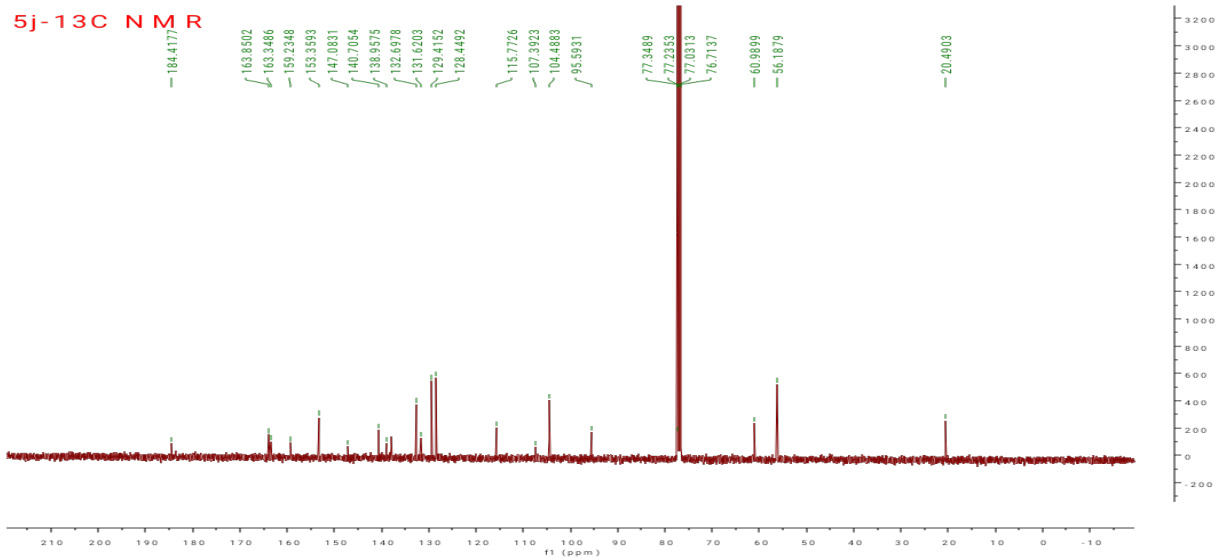


Figure S 37. ¹³C NMR of compound 5j

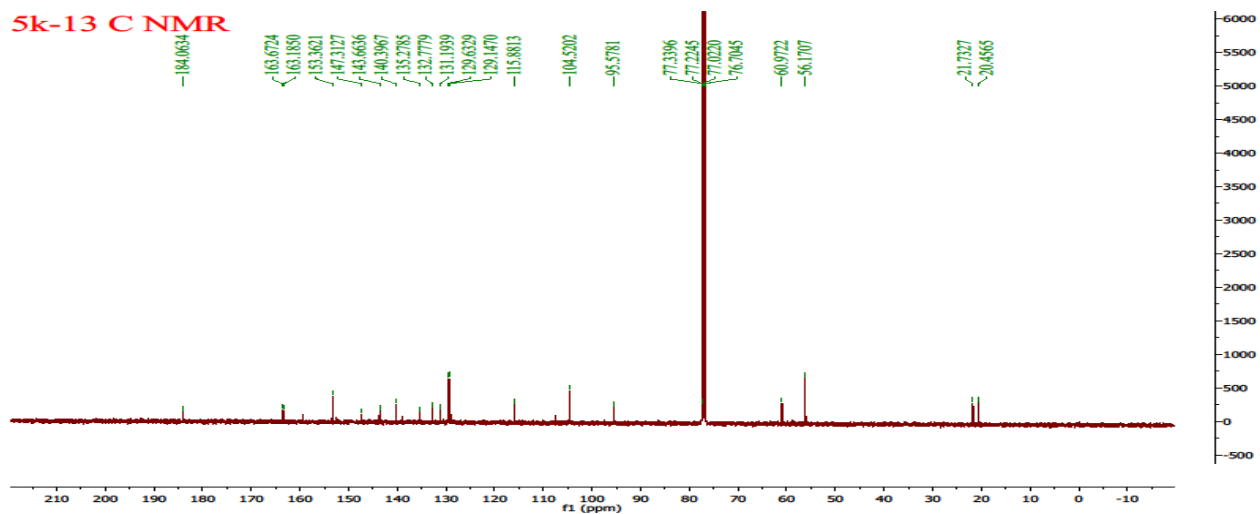


Figure S 38. ¹³C NMR of compound 5k

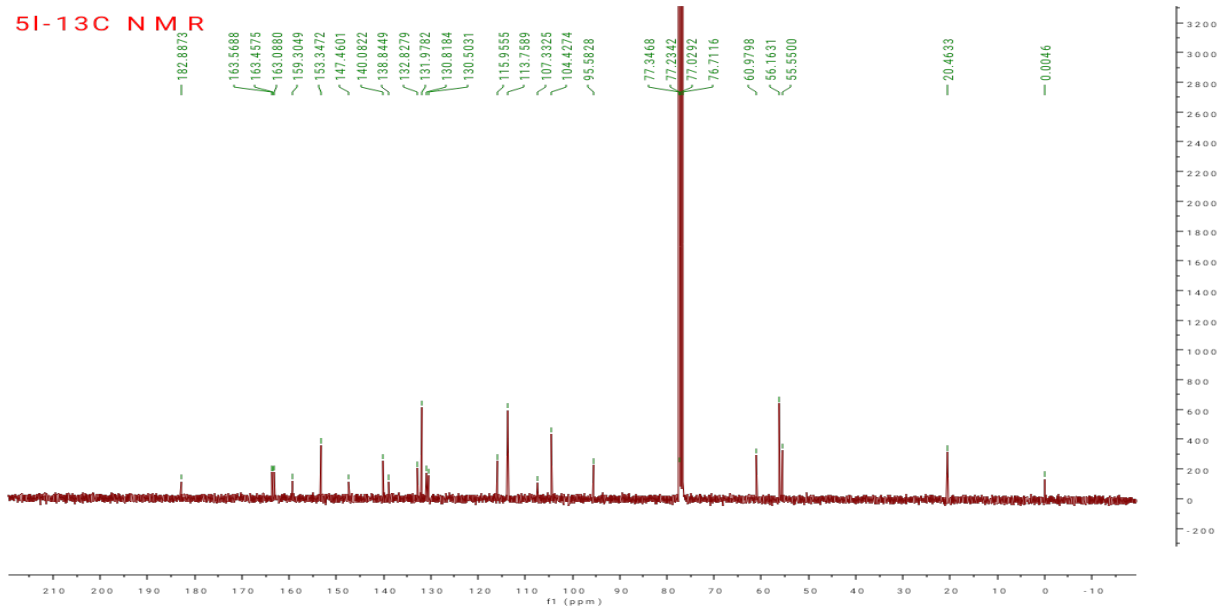


Figure S 39. ¹³C NMR of compound 5I

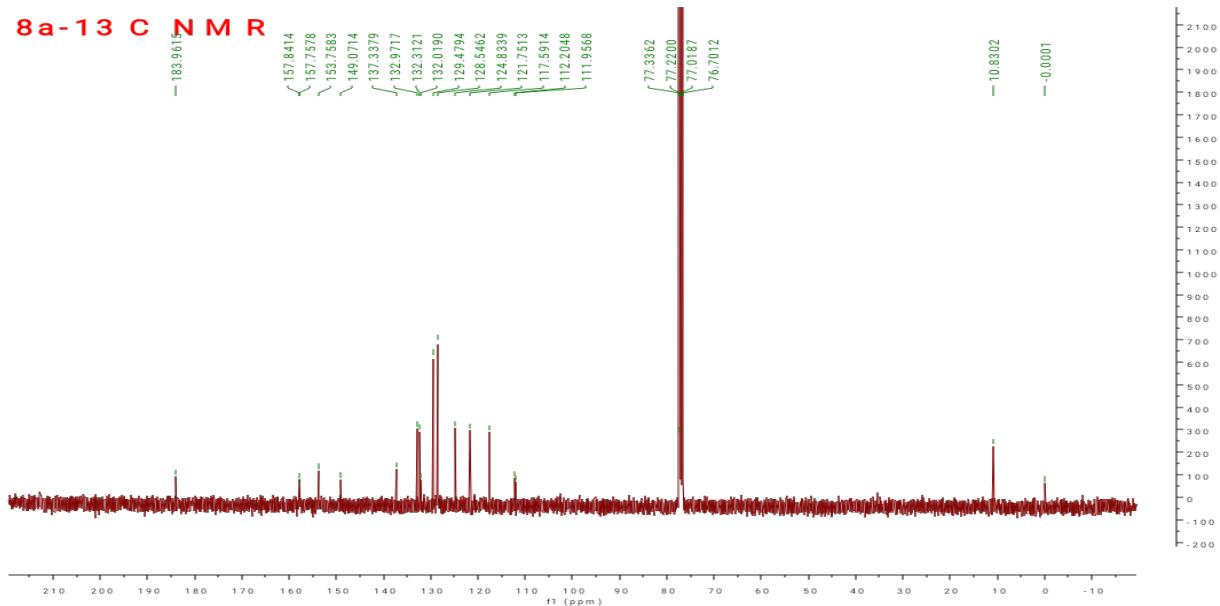


Figure S 40. ¹³C NMR of compound 8a

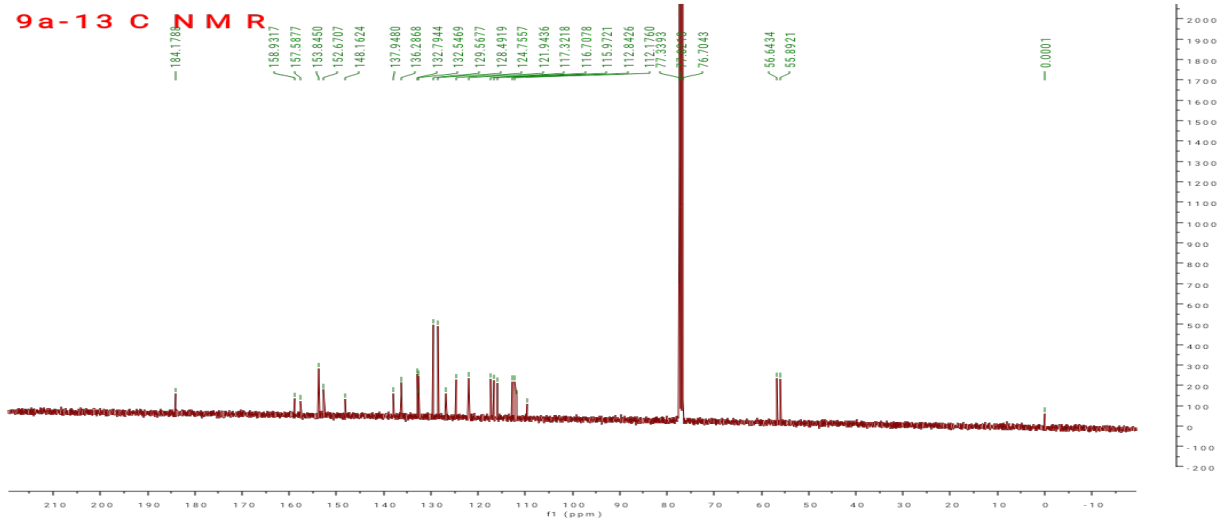


Figure S 41. ¹³C NMR of compound 9a

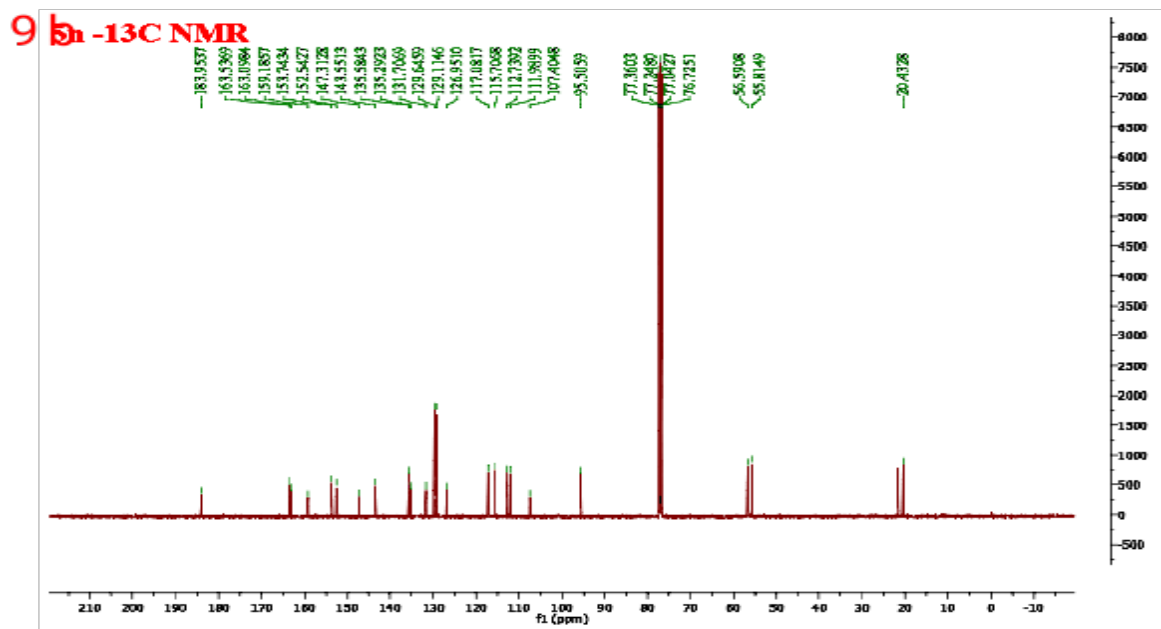


Figure S 42. ¹³C NMR of compound 9b

9c-¹³C NMR

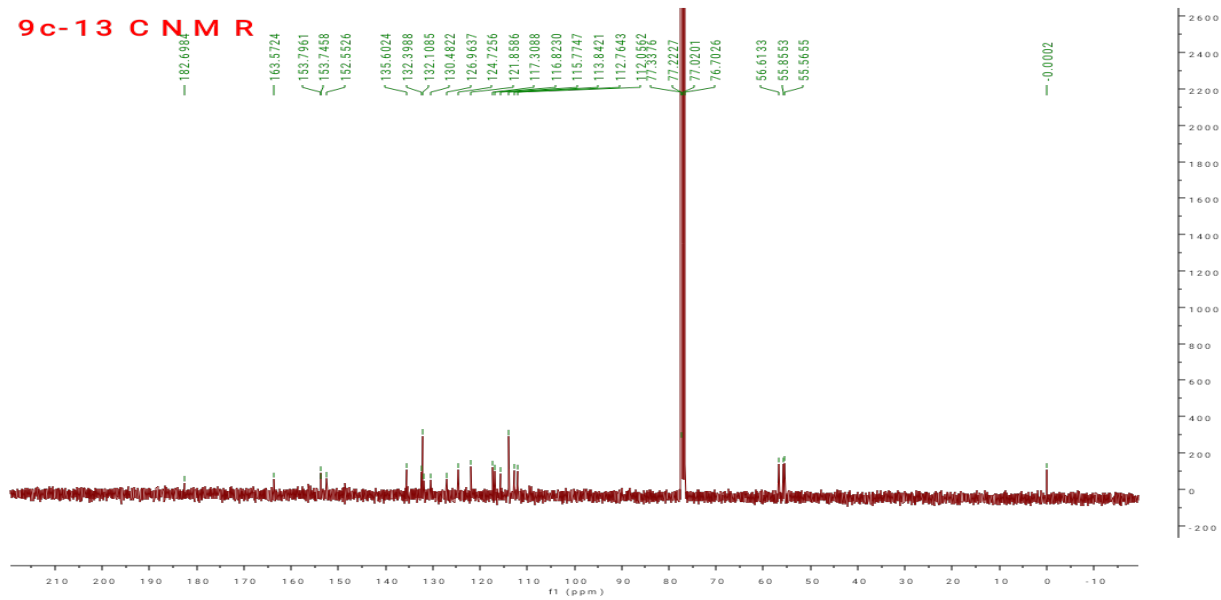


Figure S 43. ¹³C NMR of compound 9c

9d-¹³C NMR

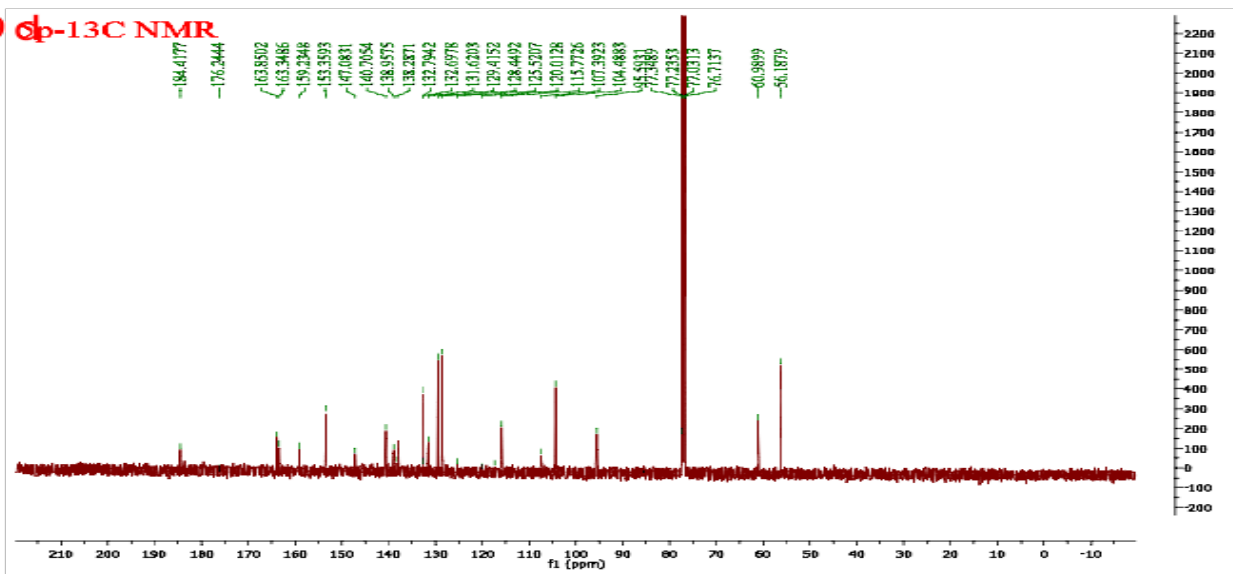


Figure S 44. ¹³C NMR of compound 9d

9e-13C NMR

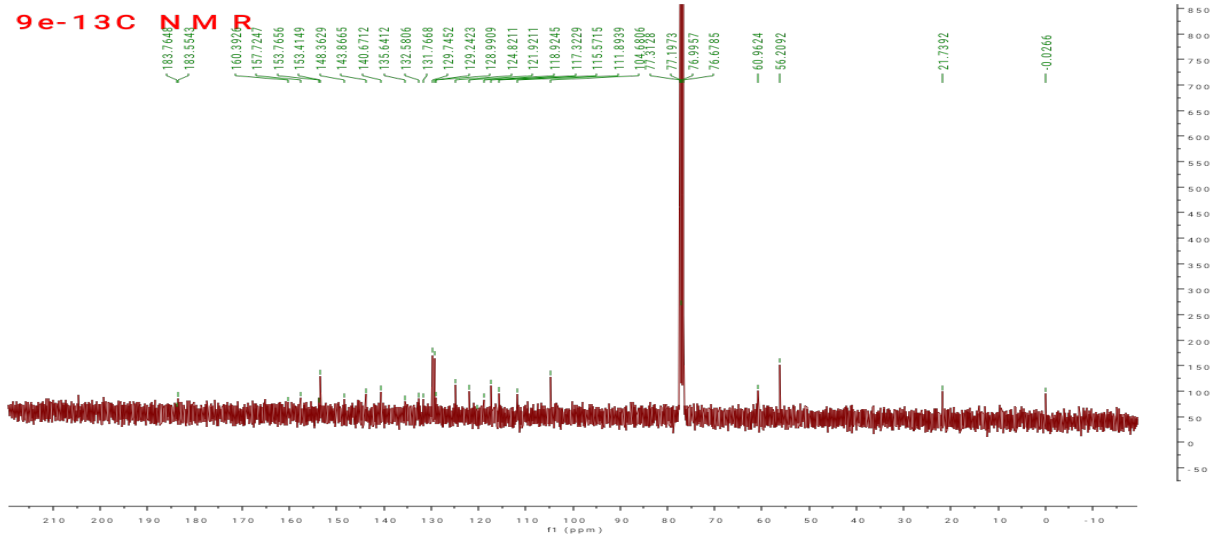


Figure S 45. ¹³C NMR of compound 9c

9f-13C NMR

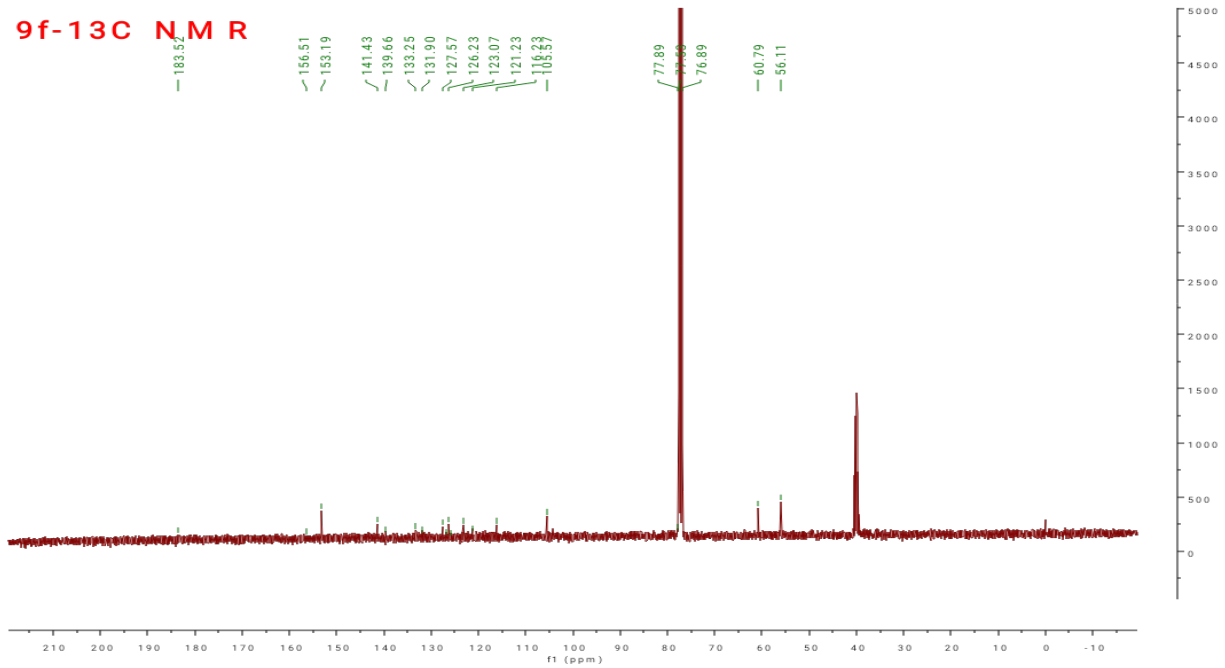


Figure S 46. ¹³C NMR of compound 9f

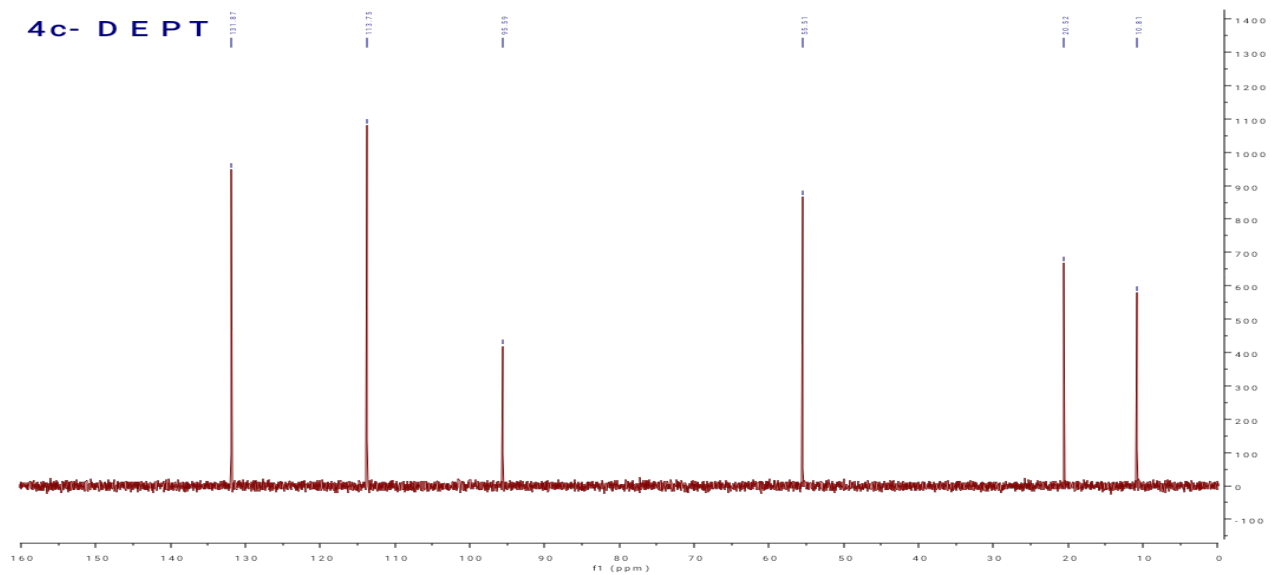


Figure S 47. DEPT of compound 4c

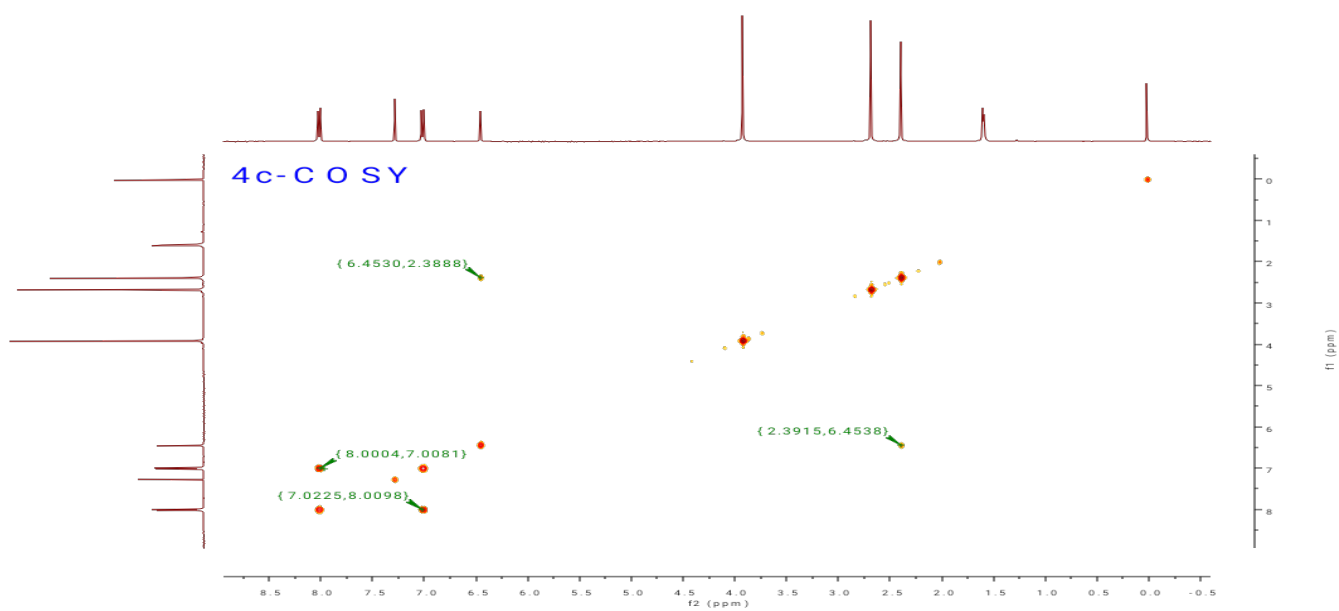


Figure S 48. 2D NMR (COSY) of compound 4c

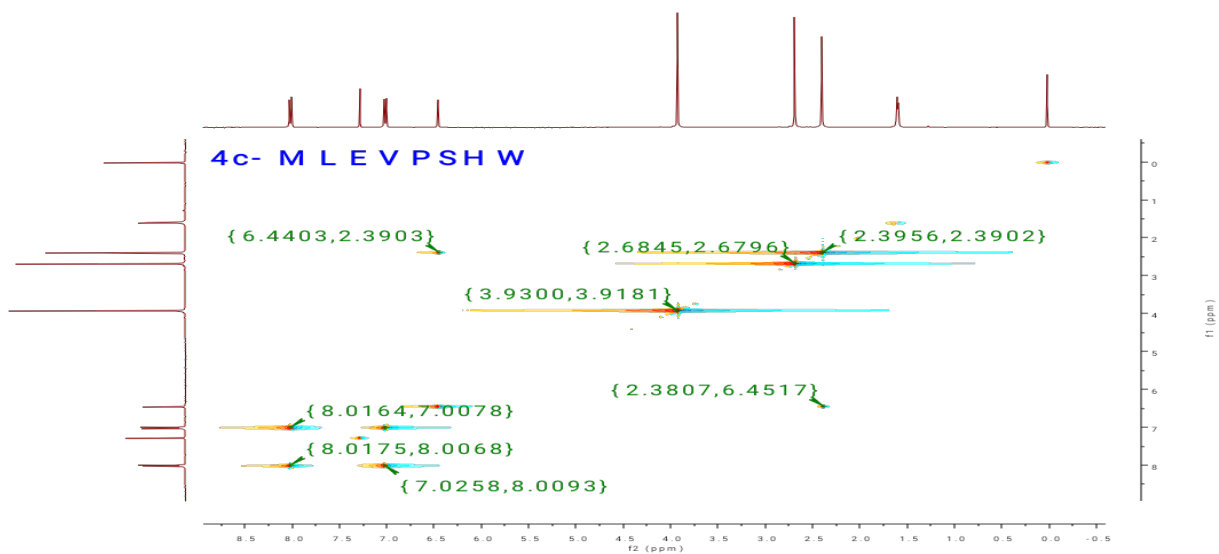


Figure S 49.2D NMR (MLEPSHW) of compound 4c

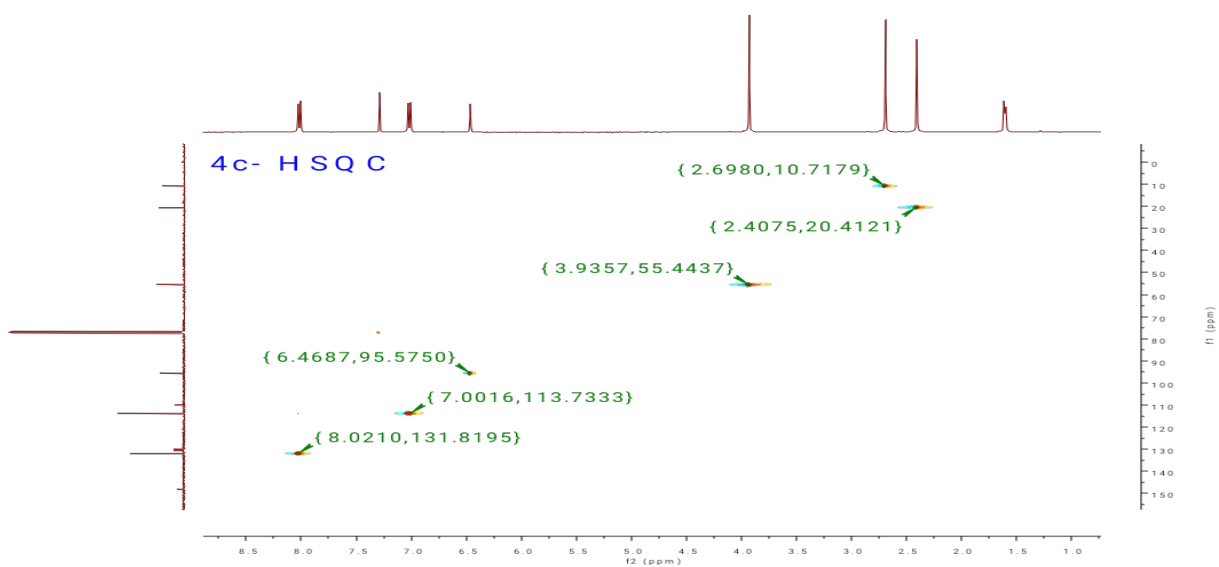


Figure S 50.2D NMR (HSQC) of compound 4c

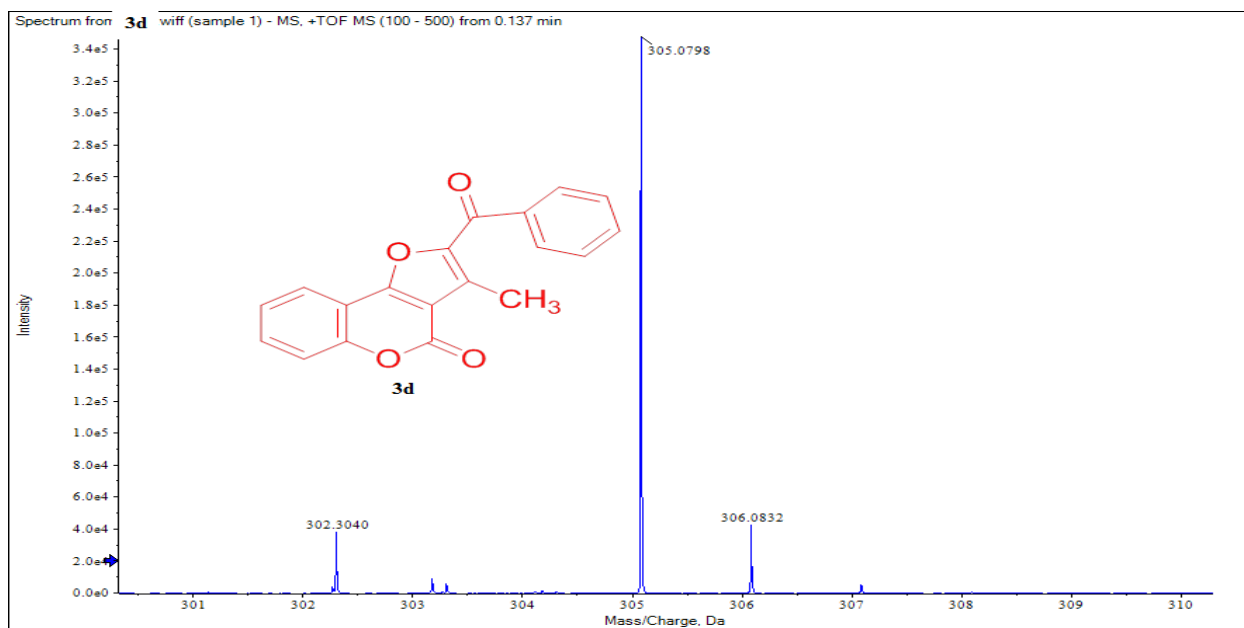


Figure S 51.HRMS of compound 8a

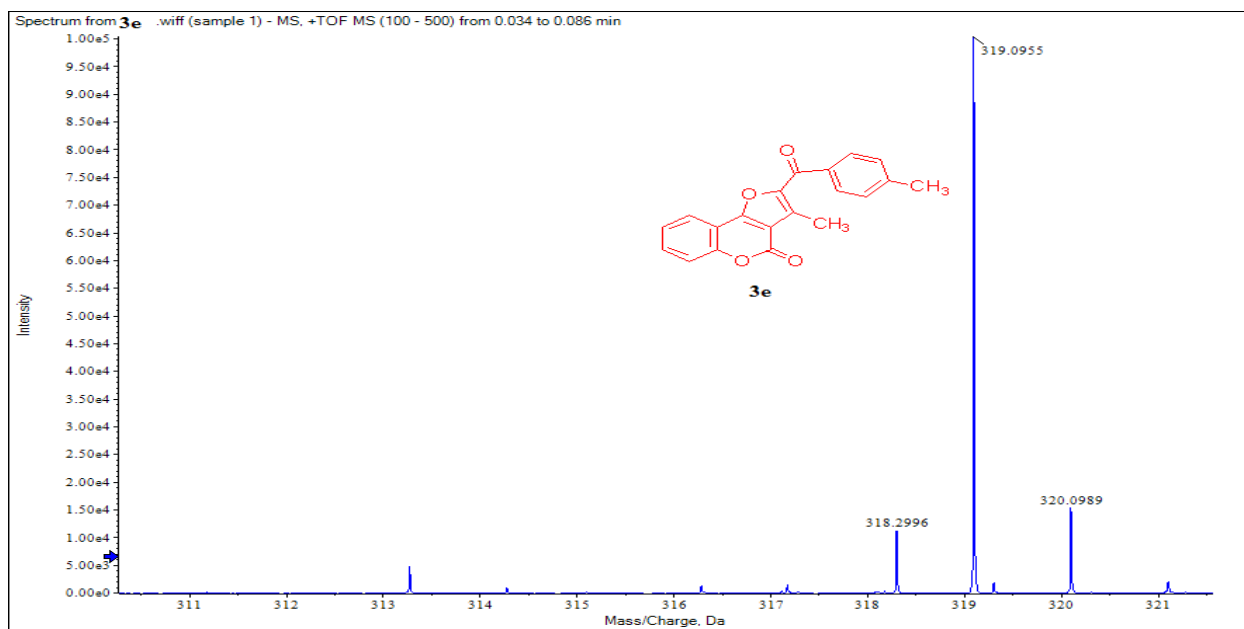


Figure S 52.HRMS of compound 8b

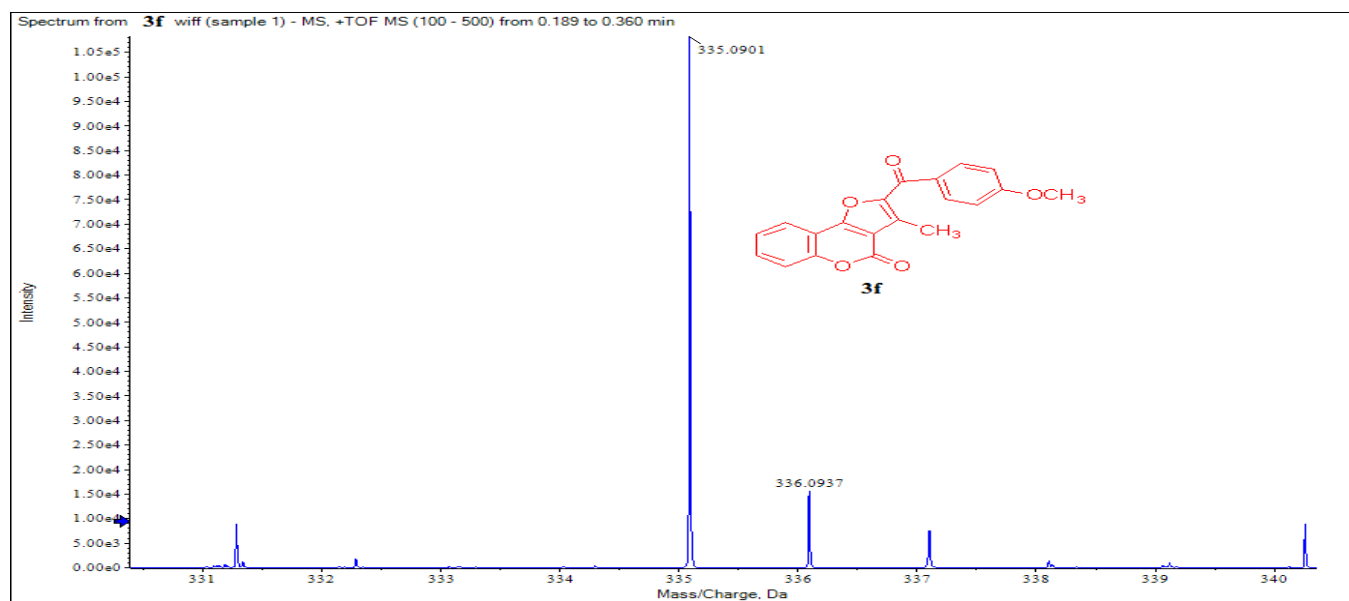


Figure S 53.HRMS of compound 8c

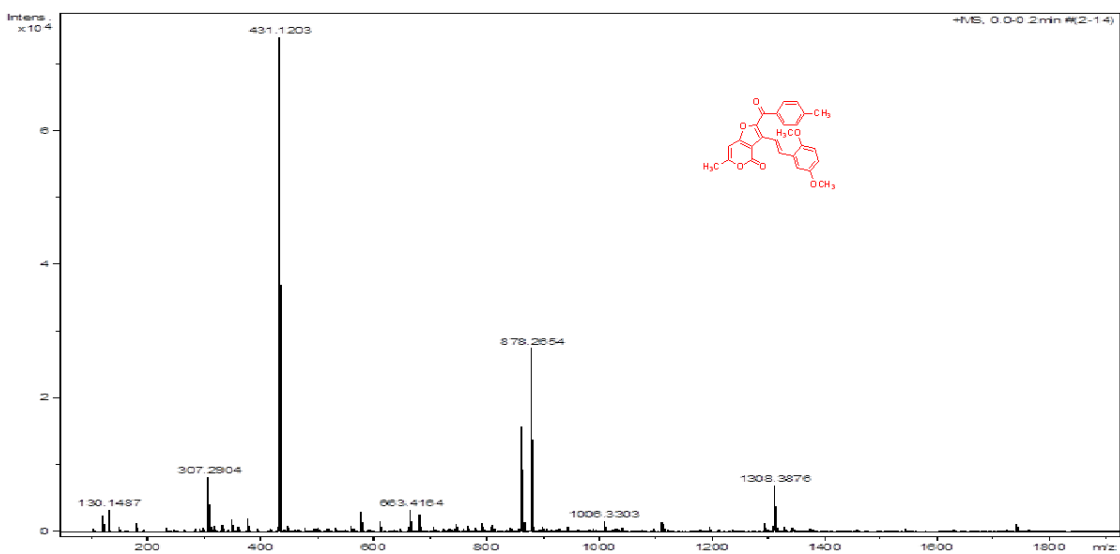


Figure S 54.HRMS of compound 5a

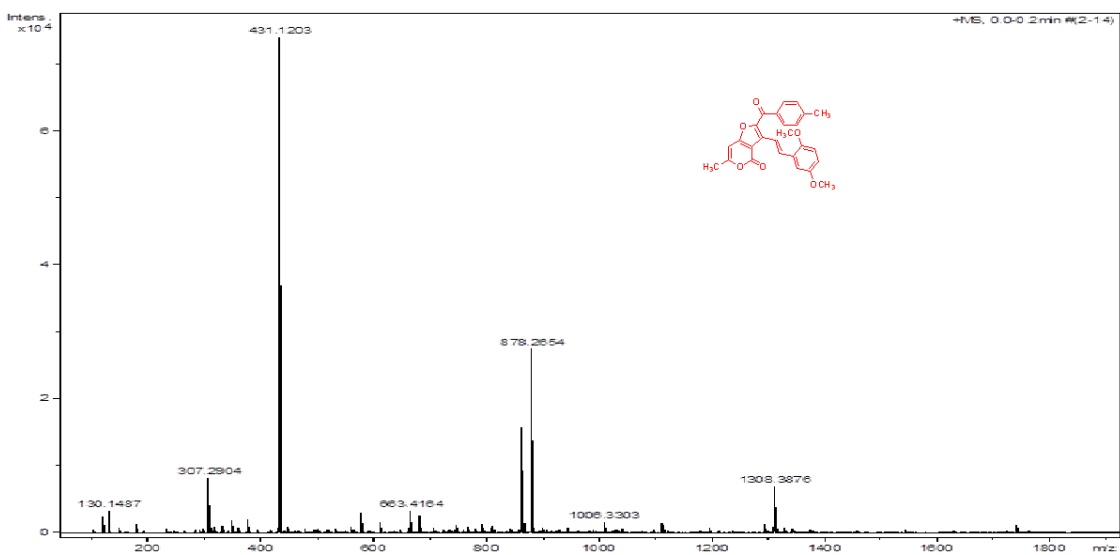


Figure S 55.HRMS of compound 5b

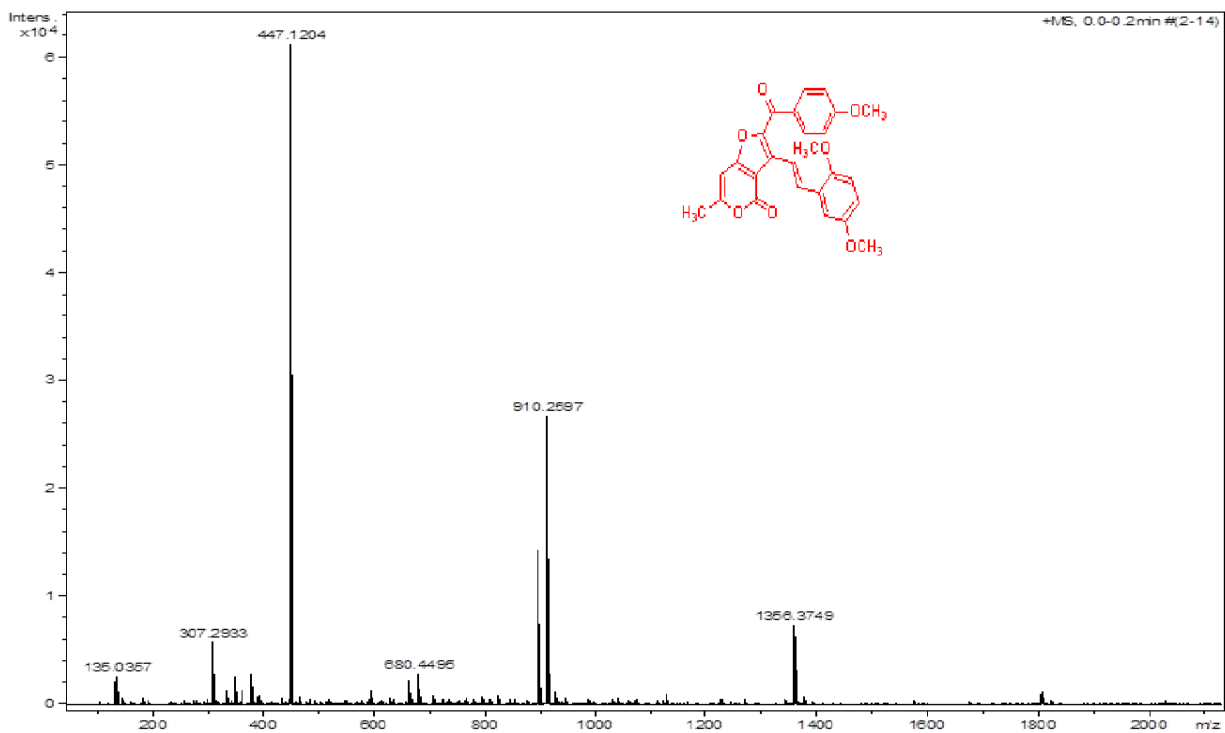


Figure S 56.HRMS of compound 5c

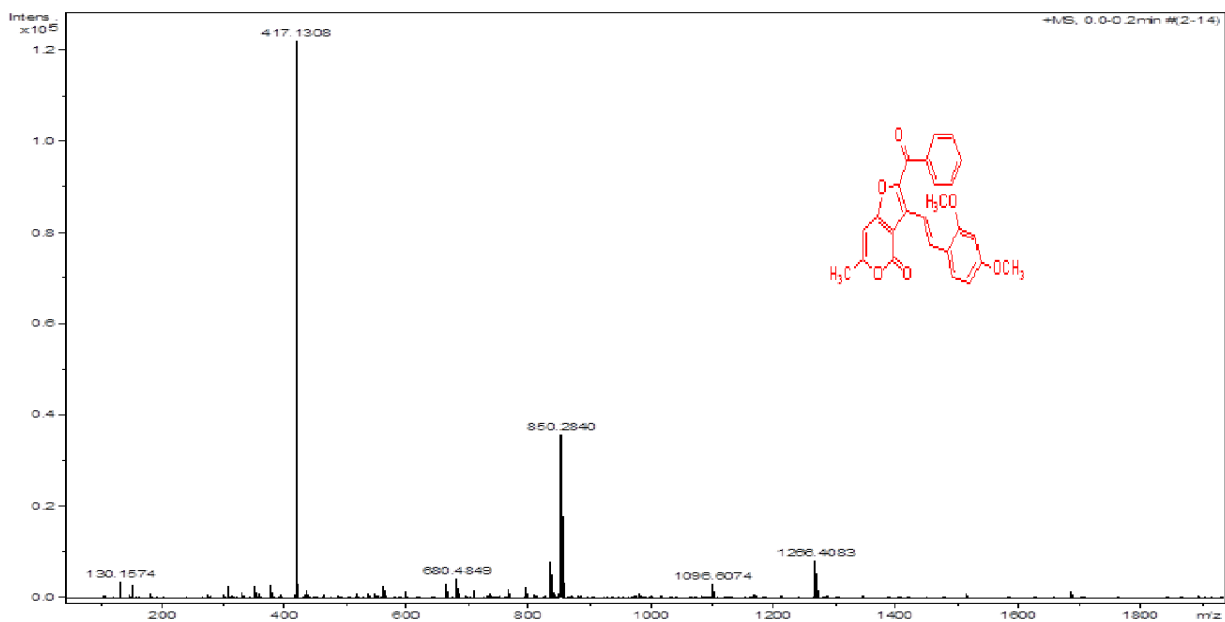


Figure S 57.HRMS of compound 5d

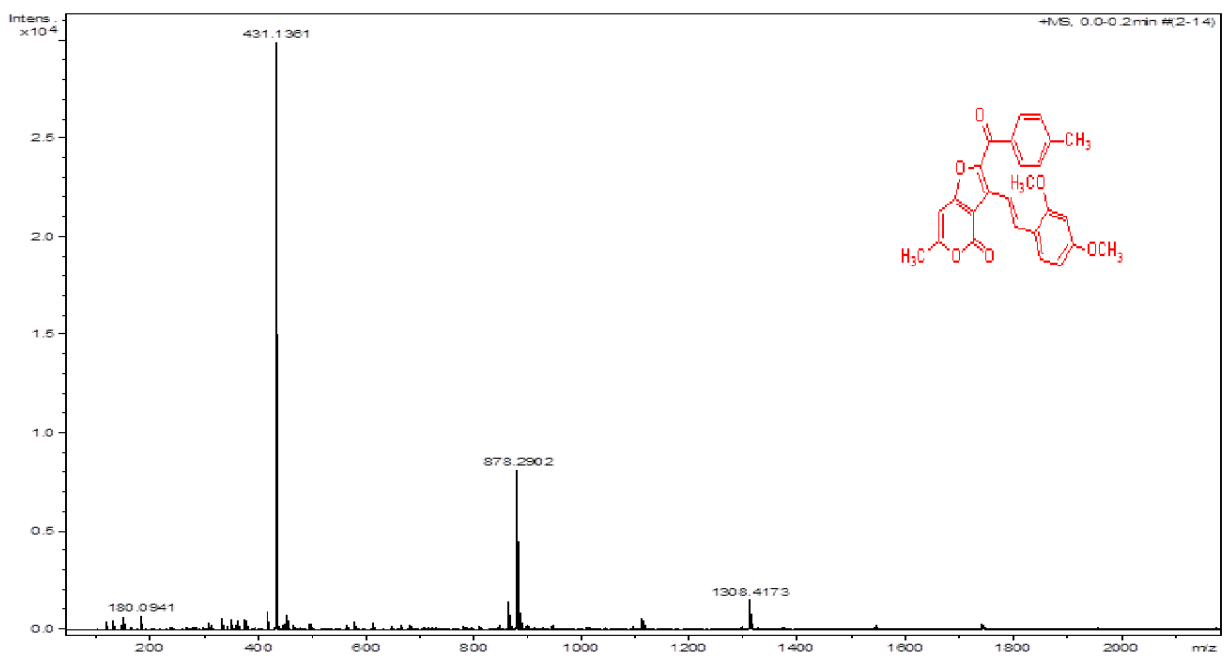


Figure S 58.HRMS of compound 5e

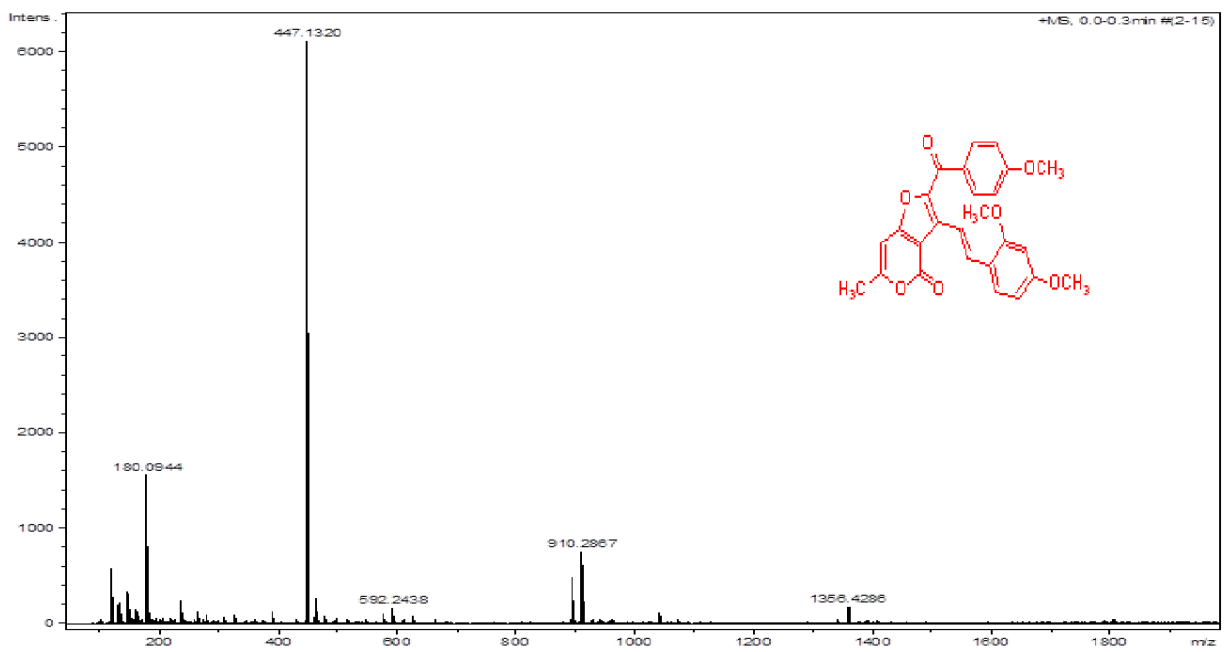


Figure S 59.HRMS of compound 5f

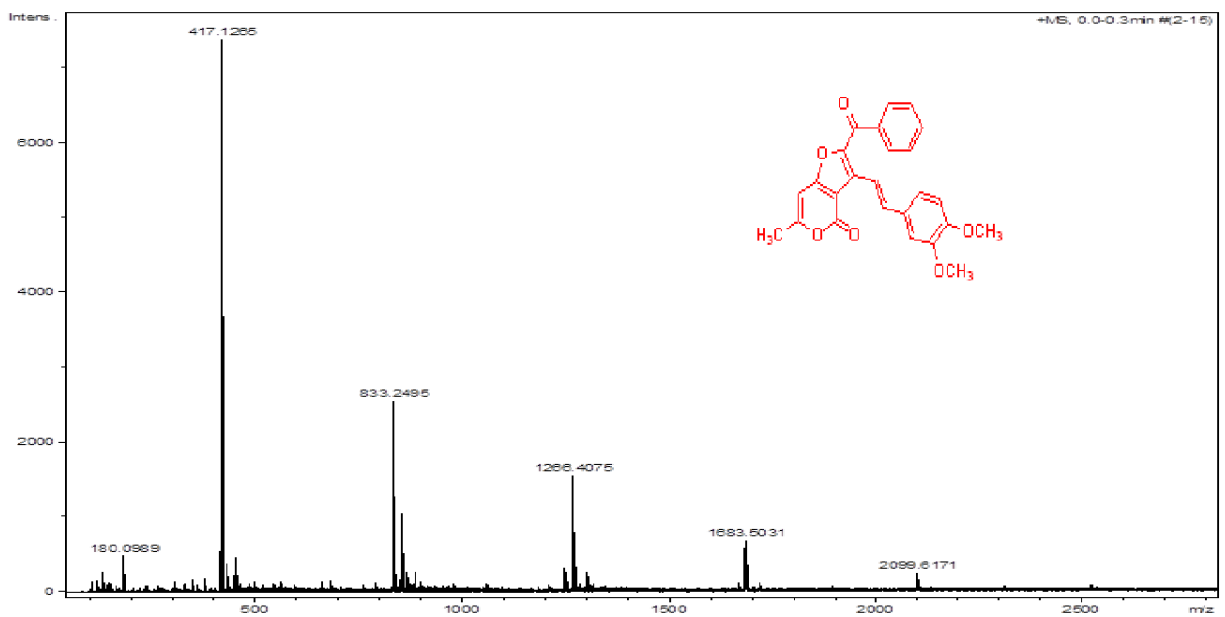


Figure S 60.HRMS of compound 5g

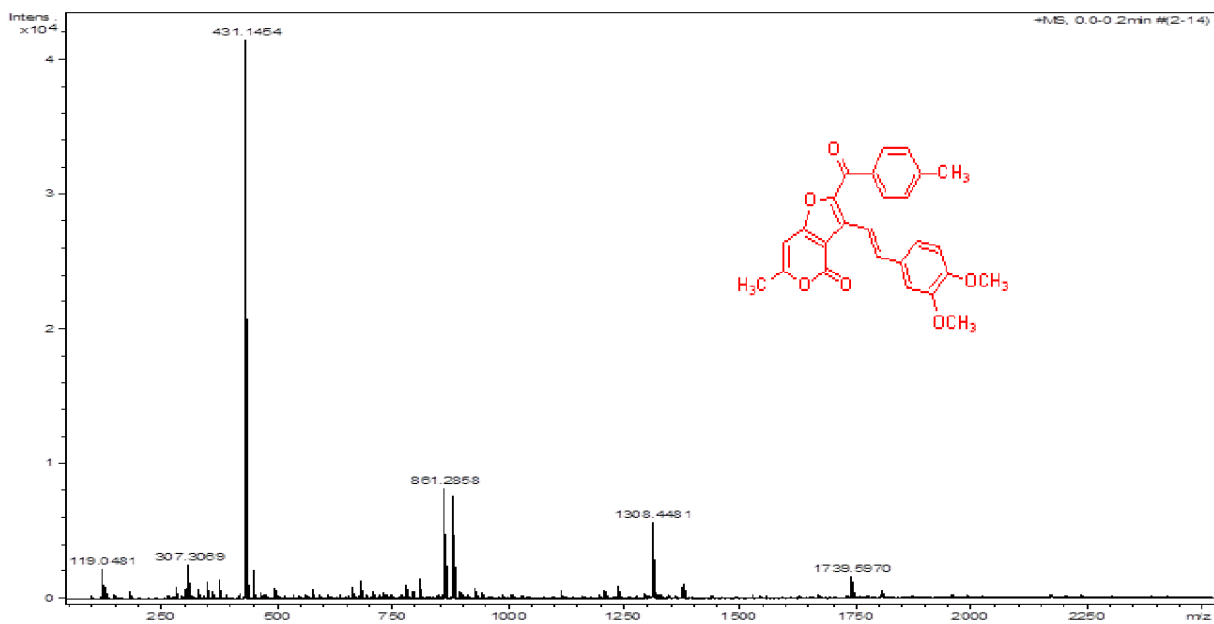


Figure S 61.HRMS of compound 5h

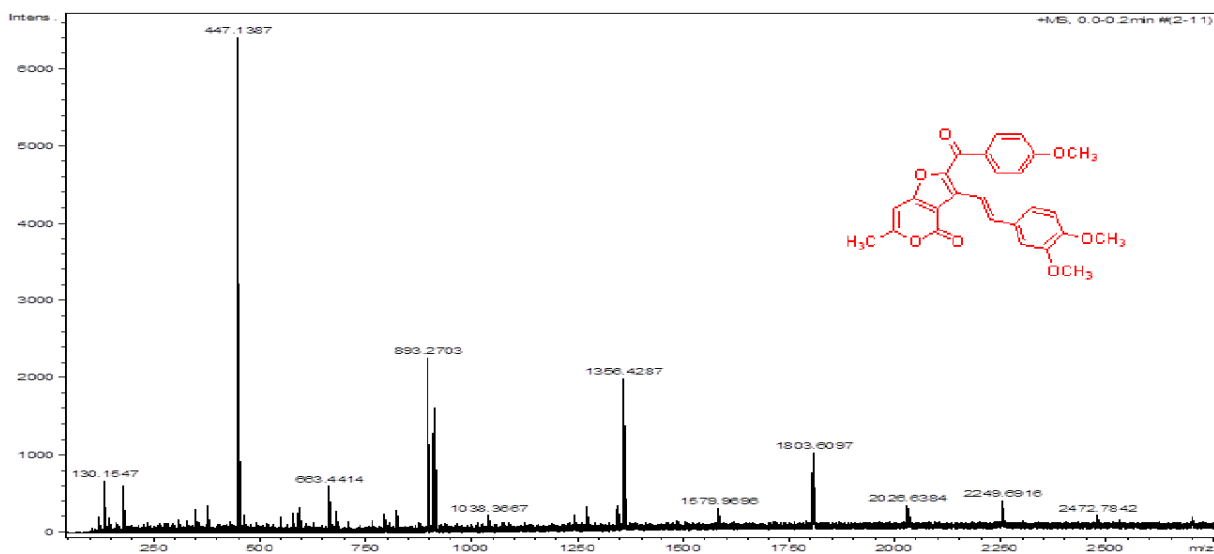


Figure S 62.HRMS of compound 5i

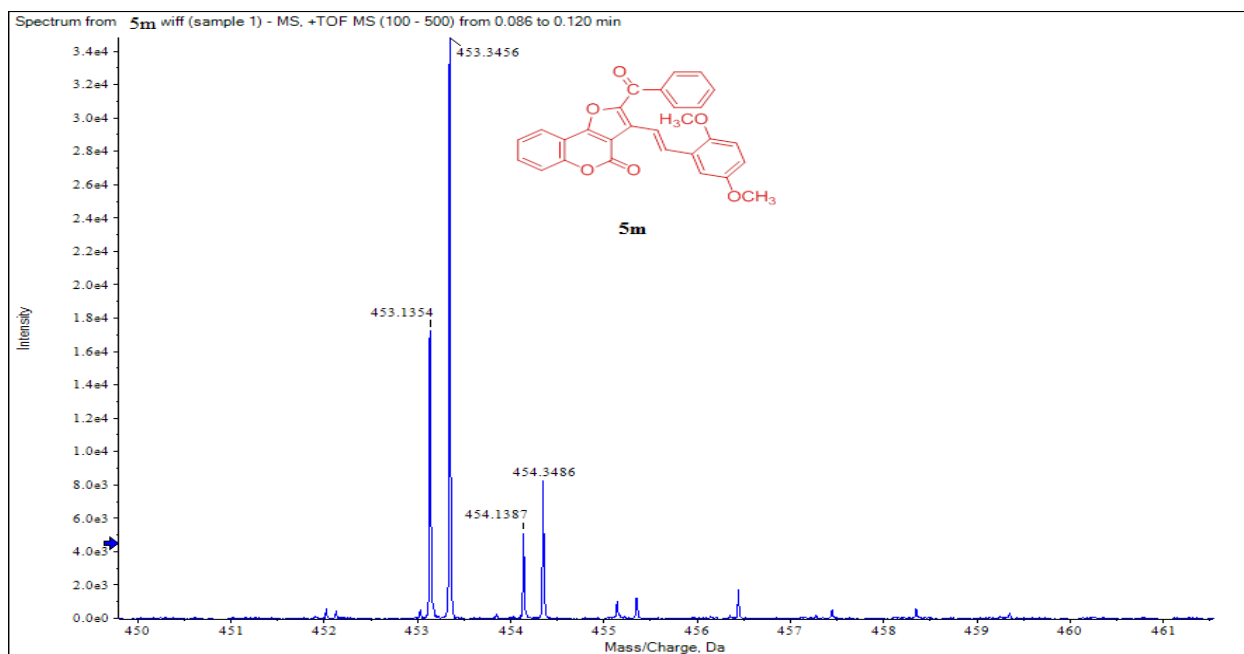


Figure S 63.HRMS of compound 9a

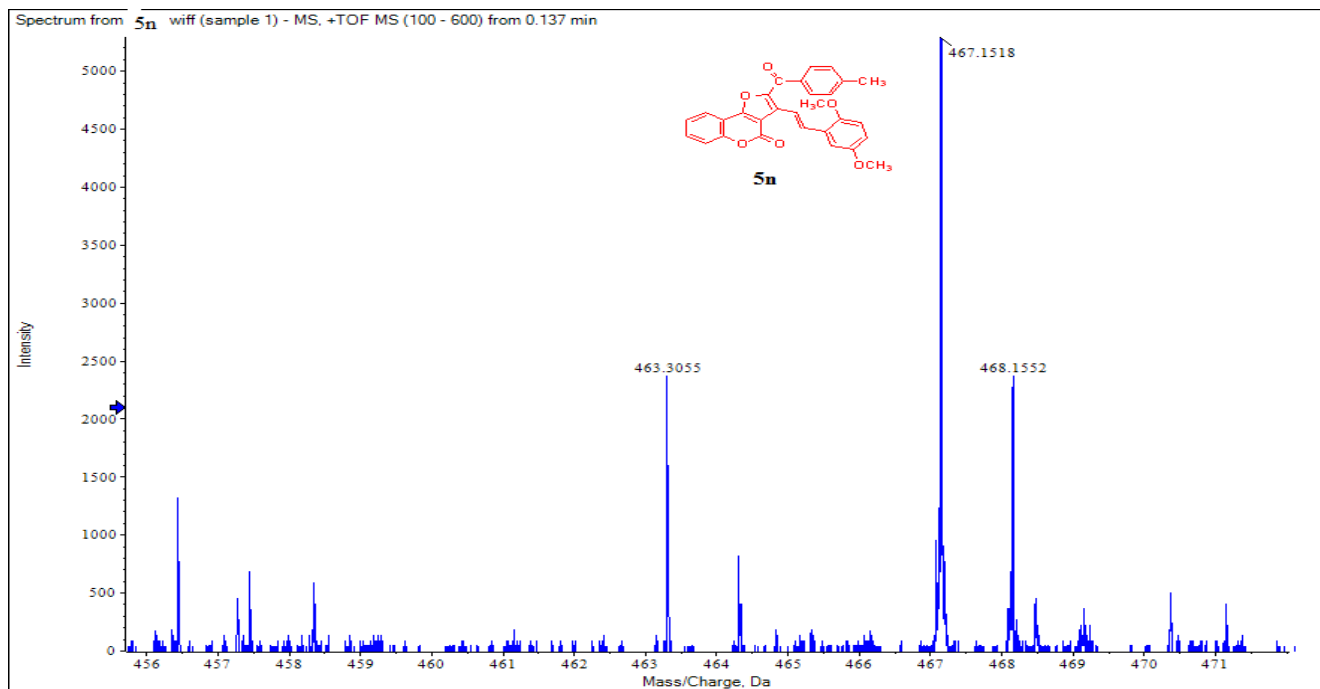


Figure S 64.HRMS of compound 9b

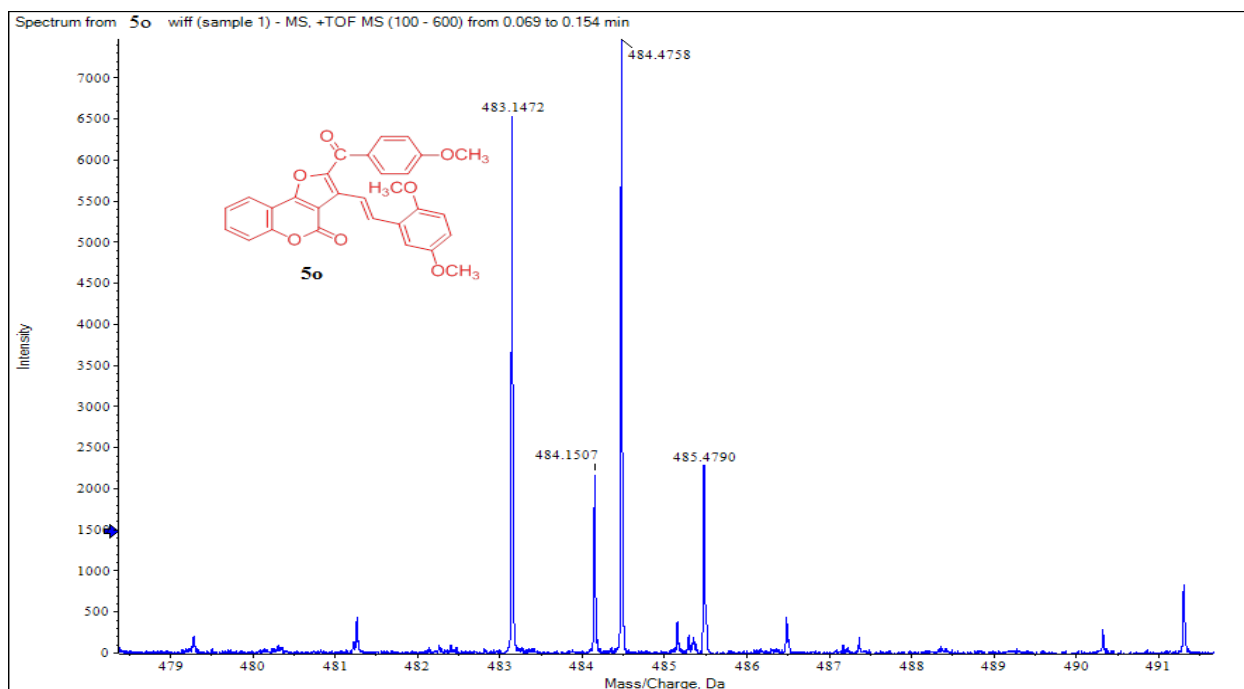


Figure S 65.HRMS of compound 9c

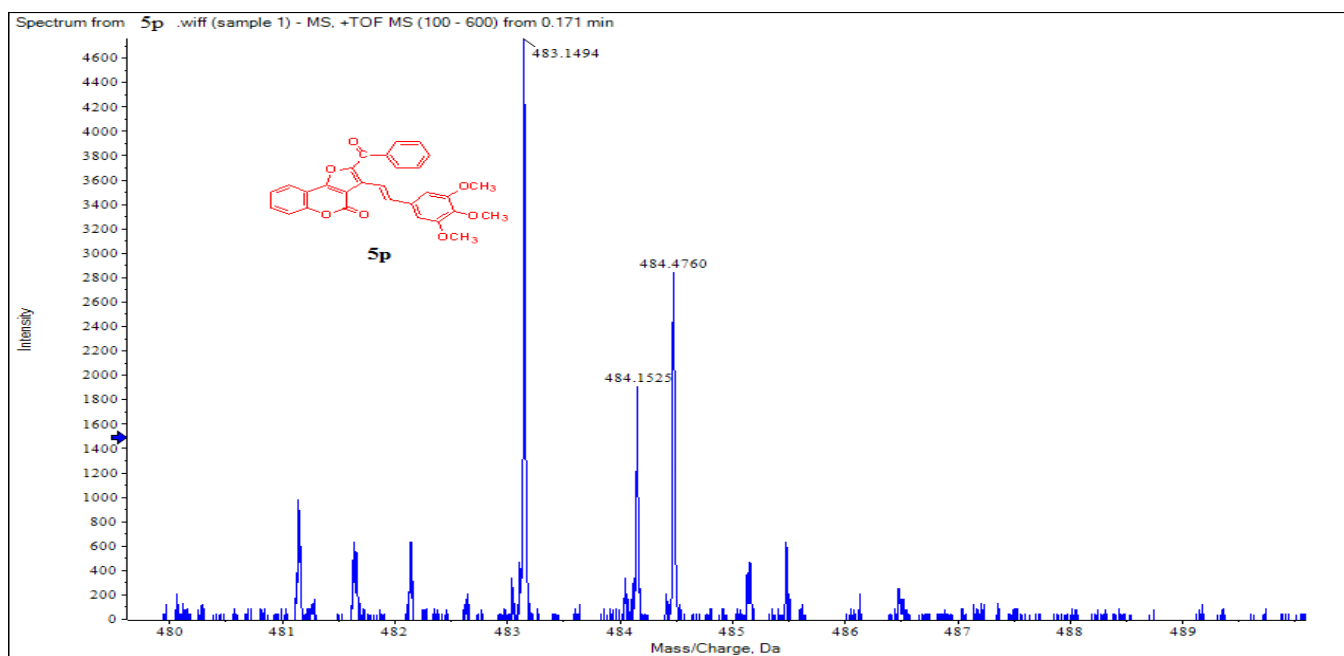


Figure S 66.HRMS of compound 9d

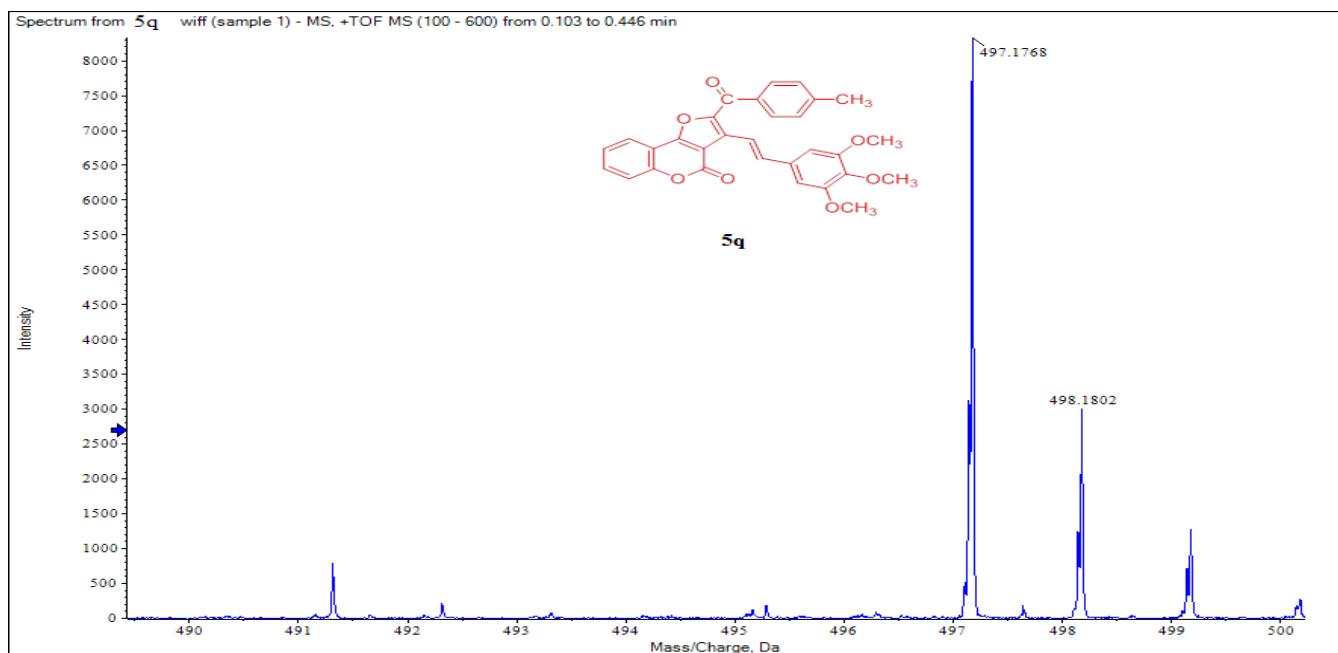


Figure S 67.HRMS of compound 9e

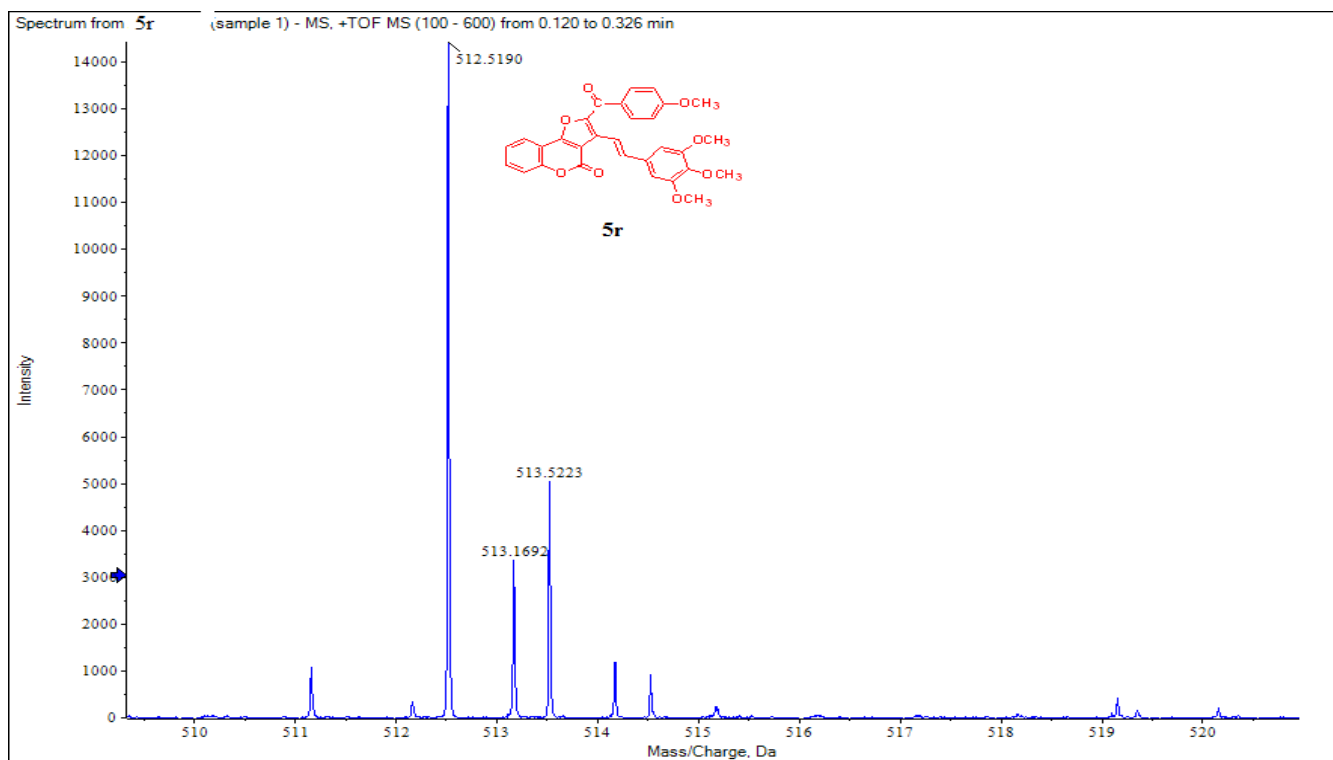


Figure S 68.HRMS of compound 9f

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1. Houghton, P.; Fang, R.; Techatanawat, I.; Steventon, G.; Hylands, P. J.; Lee, C. C.. *Methods*. **2007**, 42, 377-387. doi: [10.1016/j.ymeth.2007.01.003](https://doi.org/10.1016/j.ymeth.2007.01.003).
2. Cappuccino, C. J.; Sherman, N. *Microbiology- a laboratory manual*. Addison Wesley, Californiaa, **1999**, 263.