

SUPPLIMENTARY INFORMATION

UNUSUAL TITANIUM INDUCED McMURRY COUPLING OF 4-OXO-4H-CHROMENE-2-CARBALDEHYDES ENROUTE TO BIS-CHROMONES

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Experimental section

General Information

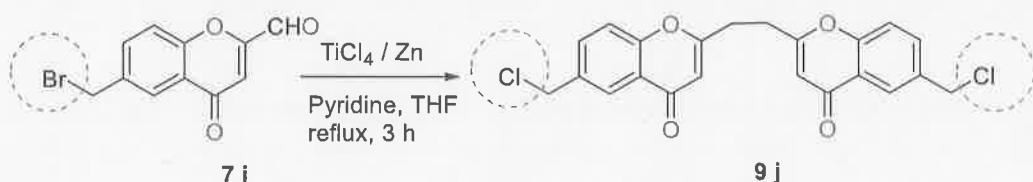
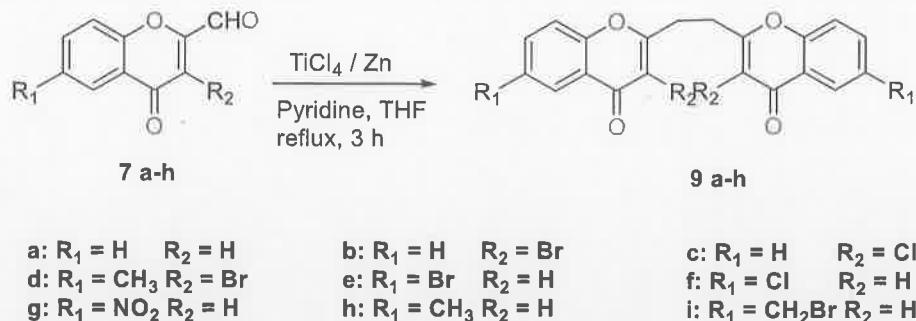
Electrospray ionization and tandem mass spectrometry experiments were performed using a triple quadrupole mass spectrometer (PE Sciex model API 3000). The positive and negative electrospray data were obtained by switching the capillary voltage between +5000 and -4500 V, respectively. For HRMS, UPLC-TOF-MS system consisted of an ACQUITY™ Ultra Performance Liquid Chromatography system and Micromass LCT Premier XE Mass Spectrometer (High sensitivity orthogonal time-of-flight instrument; Waters, Milford, USA) equipped with an ESI lock spray source for accurate mass values. Leucine-enkephalin was used as reference compound and introduced via the lock spray channel.

The NMR experiments were performed on Varian spectrometers operating at 400 and 500 MHz in DMSO (D_6) and $CDCl_3$ at 30 °C. The 1H chemical shift values were reported on the δ scale in ppm, relative to TMS ($\delta = 0.00$) and the ^{13}C chemical shift values were reported relative to DMSO ($\delta = 40$ ppm), $CDCl_3$ ($\delta = 77$ ppm) as internal standards.

- For the compounds **9b**, **9f** and **9g** ^{13}C NMR could not be recorded due to low solubility in selective NMR solvents.

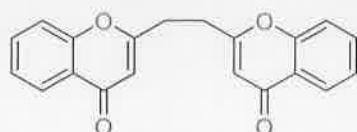
X-ray reflections for compound **9h** were collected at 290 K on an Oxford Xcalibur Mova E diffractometer equipped with an EOS CCD detector and a microfocus sealed tube using Mo K α radiation ($\lambda = 0.7107 \text{ \AA}$). Data collection and reduction was performed using CrysAlisPro (version 1.171.36.32) and OLEX2 (version 1.2) was used to solve and refine the crystal structures. All non-hydrogen atoms were refined anisotropically. All C–H atoms were fixed geometrically. The final CIF files were validated in PLATON and doesn't resulted in any missed symmetry. General procedure for McMurry coupling of 4-oxo-4*H*-chromen-2-carbaldehydes **7a-i** Zn powder (4 mmol) was added to 15 volumes of dry THF under stirring at 0–5°C followed by $TiCl_4$ (2 mmol). Reaction temperature was raised 60–65°C, maintained for about 3 h and then cooled to 0–5°C. Pyridine (2.5 mmol) and corresponding chromone-2-carbaldehyde (**7a-i**) (1.0 mmol) in 15 volumes of dry THF were then added to the mixture. Reaction temperature was raised to 60–65°C and maintained for 3–4 h. Reaction mass was cooled to 10–15°C and quenched with 10% K_2CO_3 solution. After filtering the inorganic salts product was extracted into

dichloromethane, which was concentrated under reduced pressure to get the product in crude form. Pure products (**9a-h**, **9j**) were obtained by passing the crude products through silica column (ethylacetate : petether, 40 : 60).



Characterization data for products **9 a-i**, **11-12**

2,2'-(Ethane-1,2-diyl)bis(4H-chromen-4-one) (9a)



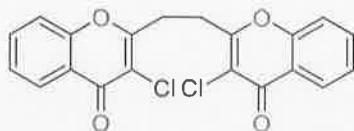
Pale yellow crystallin solid (165 mg, 52%), mp: 190-193 °C: ¹H NMR (400 MHz, DMSO-d₆, ppm): δ = 8.01 (dd, 1H, J = 8.4, 2.0 Hz, H-5), 7.79 (dt, 1H, J = 8.8, 2.0 Hz, H-7), 7.56 (d, 1H, J = 8.4 Hz, H-8), 7.48 (t, 1H, J = 7.6 Hz, H-6), 6.35 (s, 1H, H-3) and 3.15 (s, 2H, H-2a). ¹³C NMR (100 MHz, DMSO-d₆, ppm): δ = 30.3 (CH₃), 109.8 (C-3), 118.0 (C-Ar), 123.0 (C-Ar), 124.8 (C-Ar), 125.3 (C-Ar), 134.1 (C-Ar), 155.8 (C-8a), 167.6 (C-2), 176.7 (C-4, CO). IR (KBr): ν = 1650, 1380 cm⁻¹. HRMS (ESI-TOF/MS): m/z calcd. for C₂₀H₁₅O₄ [M+H]⁺: 319.0970; found: 319.0956.

2,2'-(Ethane-1,2-diyl)bis(3-bromo-4H-chromen-4-one) (9b)



Pale yellow crystalline powder (240 mg, 51%), mp: 266-288 °C. ^1H NMR (400 MHz, DMSO-d₆, ppm): δ = 8.00 (dd, 1H, J = 7.6, 1.2 Hz, H-5), 7.79-7.83 (m, 1H, H-7), 7.56 (d, 1H, J = 8.0 Hz, H-8), 7.52 (t, 1H, J = 6.8 Hz, H-6) and 3.50 (s, 2H, H-2a). FT-IR (KBr): ν = 1646 cm⁻¹ (C=O), HRMS (ESI-TOF/MS): m/z calcd. for C₂₀H₁₃O₄Br₂ [M+H]⁺: 474.9181; found: 474.9202.

2,2'-(Ethane-1,2-diyl)bis(3-chloro-4H-chromen-4-one) (9c)



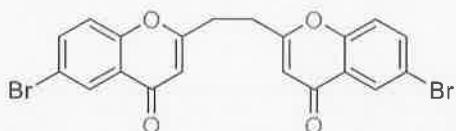
Off white crystalline powder (232 mg, 60%). mp: 254-256 °C. ^1H NMR (400 MHz, DMSO-d₆, ppm): δ = 8.07 (d, 1H, J = 8.0 Hz, H-5), 7.84 (t, 1H, J = 7.6 Hz, H-7), 7.58 (d, 1H, J = 8.8 Hz, H-8), 7.54 (t, 1H, J = 7.6 Hz, H-6) and 3.47 (s, 2H, H-2a). ^{13}C NMR (100 MHz, CDCl₃, ppm): δ = 29.1 (C-2a), 117.6 (C-3), 118.8 (C-Ar), 122.3 (C-Ar), 125.7 (C-Ar), 126.3 (C-Ar), 134.1 (C-Ar), 155.2 (C-8a), 162.6 (C-2), 172.0 (C-4, CO). FT-IR (KBr): ν = 1648 cm⁻¹ (C=O). HRMS (ESI-TOF/MS): m/z calcd. for C₂₀H₁₃O₄Cl₂ [M+H]⁺: 387.0191; found: 387.0178.

2,2'-(Ethane-1,2-diyl)bis(3-bromo-6-methyl-4H-chromen-4-one) (9d)



Off white crystalline powder (300 mg, 60%), mp: 250-253 °C. ^1H NMR (400 MHz, CDCl₃, ppm): δ = 7.99 (s, 1H, H-5), 7.47 (dd, 1H, J = 8.8, 2.0 Hz, H-7), 7.25 (d, 1H, J = 8.8 Hz, H-8), 3.49 (s, 2H, H-2a) and 2.45 (s, 3H, 6-CH₃). ^{13}C NMR (100 MHz, CDCl₃, ppm): δ = 20.8 (6-CH₃), 31.5 (C-2a), 110.0 (C-3), 117.2 (C-Ar), 121.4 (C-Ar), 125.6 (C-Ar), 135.3 (C-Ar), 135.8 (C-Ar), 153.5 (C-8a), 163.5 (C-2), 172.2 (C-4, CO). FT-IR (KBr): ν = 1643 cm⁻¹ (C=O). HRMS (ESI-TOF/MS): m/z calcd. for C₂₂H₁₇O₄Br₂ [M+H]⁺: 502.9494, found: 502.9487.

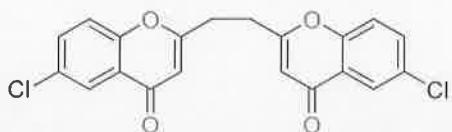
2,2'-(Ethane-1,2-diyl)bis(6-bromo-4H-chromen-4-one) (9e)



Yellow crystalline powder (240 mg, 50%), mp: 245-247 °C. ^1H NMR (400 MHz, CDCl₃, ppm): δ = 8.30 (d, 1H, J = 2.4 Hz, H-5), 7.75 (dd, 1H, J = 2.4 Hz, 8.8 Hz, H-7), 7.28 (d, 1H, J = 10.4 Hz, H-8), 6.23 (s, 1H, H-3) and 3.09 (s, 2H, H-2a). ^{13}C NMR (100 MHz, CDCl₃, ppm): δ = 31.5 (C-2a), 110.5 (C-3), 118.8 (C-Ar), 119.6 (C-Ar), 124.9 (C-Ar), 128.4 (C-Ar), 136.8 (C-Ar), 155.0 (C-

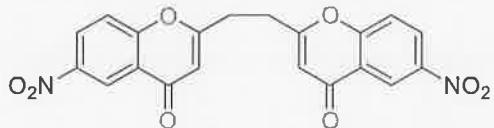
8a), 166.5 (C-2), 176.5 (C-4, CO). FT-IR (KBr): $\nu = 1649 \text{ cm}^{-1}$ (C=O). HRMS (ESI-TOF/MS): m/z calcd. for $\text{C}_{20}\text{H}_{13}\text{O}_4\text{Br}_2 [\text{M}+\text{H}]^+$: 474.9181; found: 474.9182.

2,2'-(Ethane-1,2-diyl)bis(6-chloro-4H-chromen-4-one) (9f)



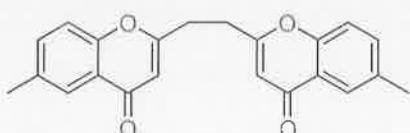
Brown crystalline powder (235 mg, 60%), mp: 233-236 °C; ^1H NMR (400 MHz, $\text{CDCl}_3 + \text{DMSO-d}_6$, ppm): $\delta = 7.93$ (d, 1H, $J = 3.2$ Hz, H-5), 7.82 (dd, 1H, $J = 8.8, 2.8$ Hz, H-7), 7.60 (d, 1H, $J = 8.8$ Hz, H-8), 6.40 (s, 1H, H-3) and 3.15 (s, 1H, H-2a). FT-IR (KBr): $\nu = 1651 \text{ cm}^{-1}$ (C=O). HRMS (ESI-TOF/MS): m/z calcd. for $\text{C}_{20}\text{H}_{13}\text{O}_4\text{Cl}_{12} [\text{M}+\text{H}]^+$: 387.0191; found: 387.0198.

2,2'-(Ethane-1,2-diyl)bis(6-nitro-4H-chromen-4-one) (9g)



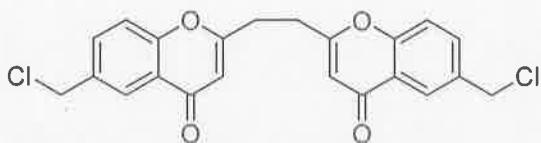
Yellow crystalline powder (210 mg, 50%), mp: 199-201 °C; ^1H NMR (400 MHz, CDCl_3 , ppm): $\delta = 9.06$ (dd, 1H, $J = 6.8, 2.4$ Hz, Ar-H), 8.45-8.52 (m, 2H, Ar-H), 8.33 (t, 1H, $J = 6.4$ Hz, Ar-H), 7.51-7.58 (m, 2H, Ar-H), 6.32 (dd, 1H, $J = 6.4, 1.6$ Hz, H-3) and 3.25 (d, 2H, $J = 15.4$ Hz, H-2a). FT-IR (KBr): $\nu = 1656 \text{ cm}^{-1}$ (C=O). HRMS (ESI-TOF/MS): m/z calcd. for $\text{C}_{20}\text{H}_{13}\text{N}_2\text{O}_8 [\text{M}+\text{H}]^+$: 409.0672; found: 409.0661.

2,2'-(Ethane-1,2-diyl)bis(6-methyl-4H-chromen-4-one) (9h).



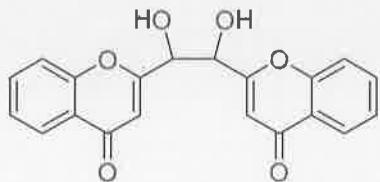
Off white crystalline powder (210 mg, 60%). mp: 195-197 °C; ^1H NMR (400 MHz, DMSO-d₆, ppm): $\delta = 7.78$ (d, 1H, $J = 1.2$ Hz, H-5), 7.59 (dd, 1H, $J = 8.0, 1.6$ Hz, H-7), 7.43 (d, 1H, $J = 8.4$ Hz, H-8), 6.30 (s, 1H, H-3), 3.12 (s, 2H, H-2a) and 2.40 (s, 3H, 6-CH₃). ^{13}C NMR (100 MHz, $\text{CDCl}_3 + \text{DMSO-d}_6$, ppm): $\delta = 20.4$ (6-CH₃), 30.4 (C-2a), 109.6 (C-3), 117.8 (C-Ar), 122.7 (C-Ar), 124.1 (C-Ar), 134.8 (C-Ar), 135.1 (C-Ar), 154.1 (C-8a), 167.4 (C-2), 176.7 (C-4, CO). FT-IR (KBr): $\nu = 1643 \text{ cm}^{-1}$ (C=O). HRMS (ESI-TOF/MS): m/z calcd. for $\text{C}_{22}\text{H}_{19}\text{O}_4 [\text{M}+\text{H}]^+$: 347.1283; found: 347.1278.

2,2'-(Ethane-1,2-diyl)bis(6-(chloromethyl)-4H-chromen-4-one) (9i)



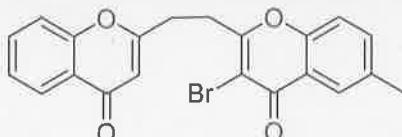
Pale yellow crystalline solid (215 mg, 52%). mp: 119-122 °C. ^1H NMR (400 MHz, CDCl_3 , ppm): δ = 8.16 (s, 1H, H-5), 7.70 (d, 1H, J = 8.8 Hz, H-7), 7.40 (d, 1H, J = 8.8 Hz, H-8), 6.23 (s, 1H, H-3), 4.66 (s, 2H, 6- CH_2Cl) and 3.10 (s, 2H, H-2a). ^{13}C NMR (100 MHz, CDCl_3 , ppm): δ = 31.4 (C-2a), 45.0 (C-6a), 110.6 (C-3), 118.5 (C-Ar), 123.5 (C-Ar), 125.6 (C-Ar), 134.1 (C-Ar), 134.9 (C-Ar), 155.1 (C-2), 166.4 (C-8a), 177.8 (C-4, CO). FT-IR (KBr): ν = 1644 cm^{-1} (C=O). HRMS (ESI-TOF/MS): m/z calcd. for $\text{C}_{22}\text{H}_{17}\text{Cl}_2\text{O}_4$ [M+H] $^+$: 415.0504; found: 415.0512.

2,2'-(1,2-Dihydroxyethane-1,2-diyl)bis(4H-chromen-4-one) (11).



White crystalline solid (217 mg, 62%). mp: 219-221 °C. ^1H NMR (400 MHz, DMSO-d_6 , ppm): δ = 8.04 (dd, 1H, J = 8.4, 1.2 Hz, H-5), 7.80 (dt, 1H, J = 8.8, 1.6 Hz, H-7), 7.57 (d, 1H, J = 8.4 Hz, H-8), 7.50 (t, 1H, J = 8.0 Hz, H-6), 6.46 (s, 1H, H-3), 6.22 (d, 1H, J = 6.0 Hz, 2a-OH) and 4.97 (d, 1H, J = 6.4 Hz, H-2a). ^{13}C NMR (100 MHz, DMSO-d_6 , ppm): δ = 71.2 (C-2a), 109.0 (C-3), 118.2 (C-8), 123.9 (C-6), 124.8 (C-4a), 125.3 (C-5), 134.1 (C-7), 155.6 (C-8a), 168.6 (C-2a), 176.7 (C-4, CO). FT-IR (KBR): ν = 3345 cm^{-1} (OH), 1650 cm^{-1} (C=O). ESI-MS: m/z 351.2 [M+H] $^+$.

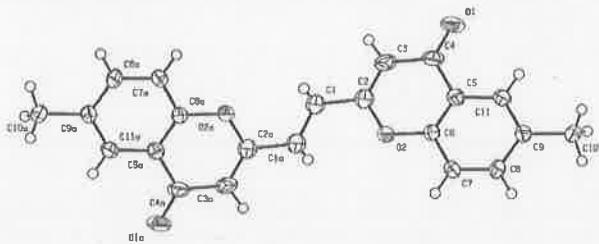
3-Bromo-6-methyl-2-(2-(4-oxo-4H-chromen-2-yl)ethyl)-4H-chromen-4-one (12).



Pale yellow crystalline solid (290 mg, 70%). mp: 141-143 °C. ^1H NMR (400 MHz, CDCl_3 , ppm): δ = 8.18 (dd, 1H, J = 7.6, 1.6 Hz, H-5), 8.01 (s, 1H, H-7), 7.66 (dt, 1H, J = 8.0, 1.2 Hz, Ar-H), 7.46 (dd, 1H, J = 8.0, 1.2 Hz, Ar-H), 7.35-7.40 (m, 3H, Ar-H), 7.23 (s, 1H, Ar-H), 6.24 (s, 1H, H-3), 3.44 (t, 2H, J = 7.6 Hz, H-2a), 3.16 (t, 2H, J = 8.0 Hz, H-2a') and 2.46 (s, 3H, 6- CH_3). ^{13}C NMR (100 MHz, CDCl_3 , ppm): δ = 20.9 (6'- CH_3), 32.1-32.0 (C-2a, C-2'a), 110.0 (C-3'), 110.5 (C-3), 117.2 (C-Ar), 117.7 (C-Ar), 121.5 (C-Ar), 123.6 (C-Ar), 125.2 (C-Ar), 125.7 (C-Ar), 133.7 (C-Ar), 135.4 (C-Ar), 135.9 (C-Ar), 153.5 (C-Ar), 156.3 (C-Ar), 163.6 (C-8a), 166.3 (C-2a), 178.0 (C-4, CO), 172.2 (C-4', CO). FT-IR (KBr): ν = 1644 cm^{-1} (C=O). HRMS (ESI-TOF/MS): m/z calcd. for $\text{C}_{21}\text{H}_{16}\text{O}_4\text{Br}$ [M+H] $^+$: 411.0232; found: 411.0238.

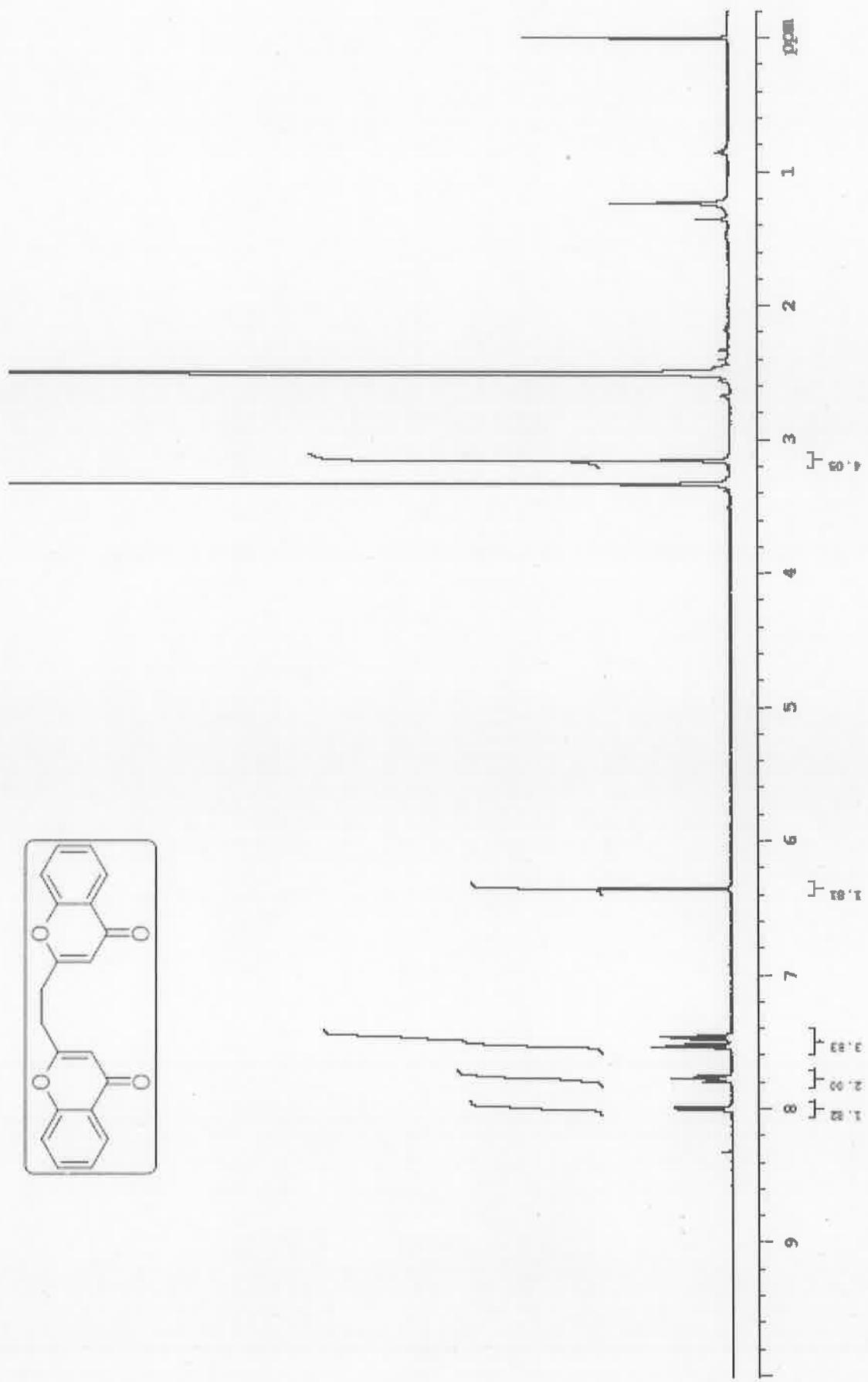
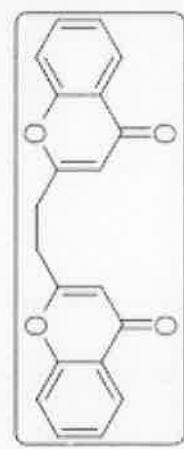
Single Crystal Data (Crystallographic parameters of compound 9h).

Temperature / K	290K
Compound	9h
Formula weight	346.36
Crystal system	Orthorhombic
Space group	<i>C</i> 2cb
<i>a</i> / Å	7.4743(5)
<i>b</i> / Å	15.2098(8)
<i>c</i> / Å	14.2562(8)
α / °	90
β / °	90
γ / °	90
Volume / Å ³	1620.68(2)
<i>Z</i>	4
Density / (g cm ⁻³)	1.42
μ / mm ⁻¹	0.097
<i>F</i> ₀₀₀	728
<i>h</i> _{min} , <i>h</i> _{max}	-8, 8
<i>k</i> _{min} , <i>k</i> _{max}	-18, 18
<i>l</i> _{min} , <i>l</i> _{max}	-16, 16
No. of measured reflections	11806
No. of unique reflections	1427
No. of reflections used	1133
<i>R</i> _{all} , <i>R</i> _{obs}	0.095, 0.075
<i>wR</i> _{2,all} , <i>wR</i> _{2,obs}	0.202, 0.186
$\Delta\rho_{\text{min,max}}$ / (e Å ⁻³)	-0.518, 0.408
<i>GooF</i>	1.082
CCDC No.	1818348

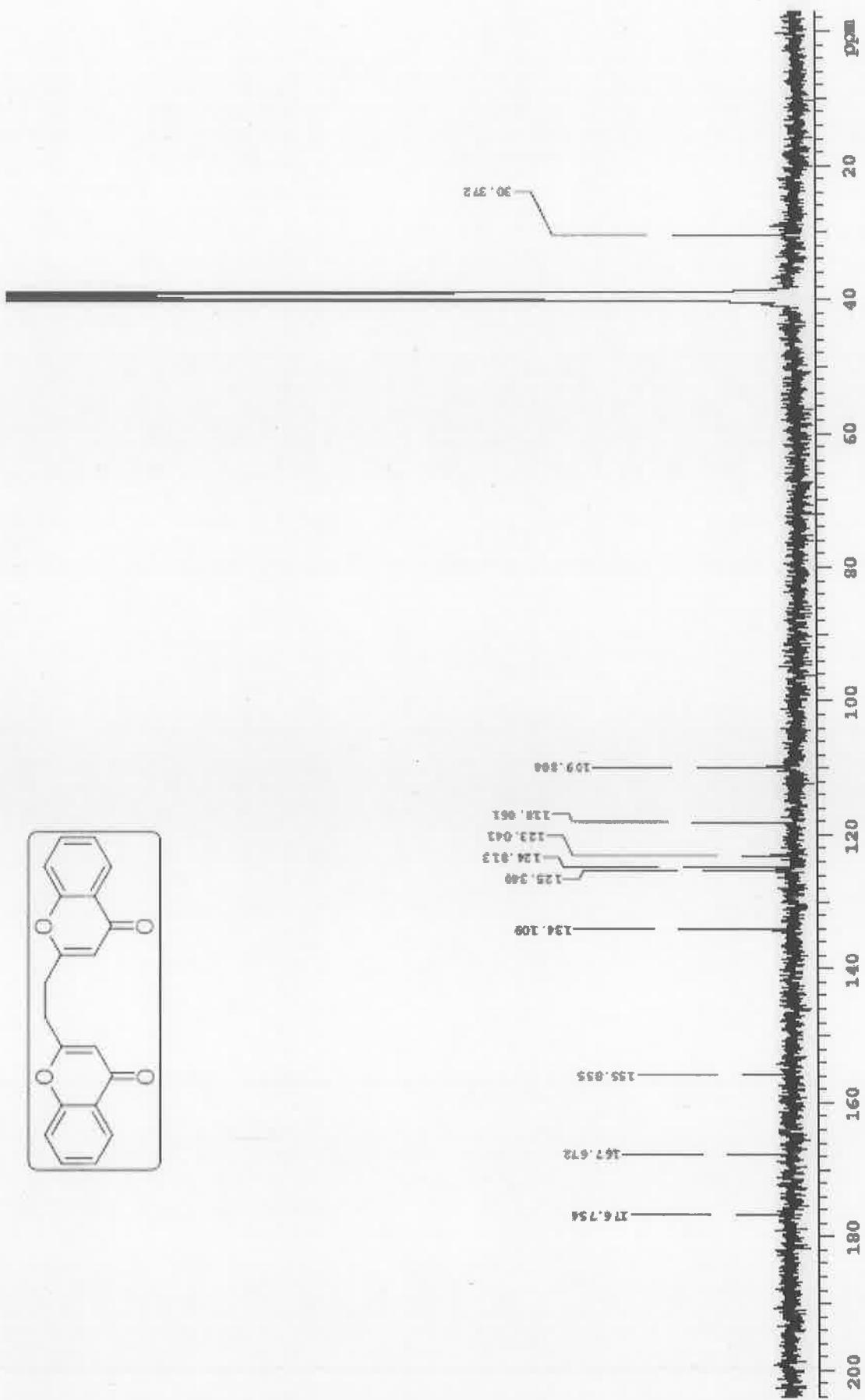
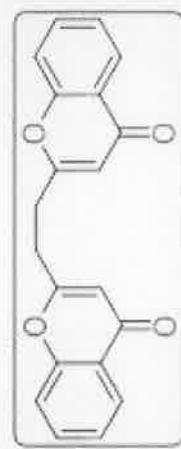


Ortep diagram of 2,2'-(ethane-1,2-diyl)bis(6-methyl-4*H*-chromen-4-one) (9h)

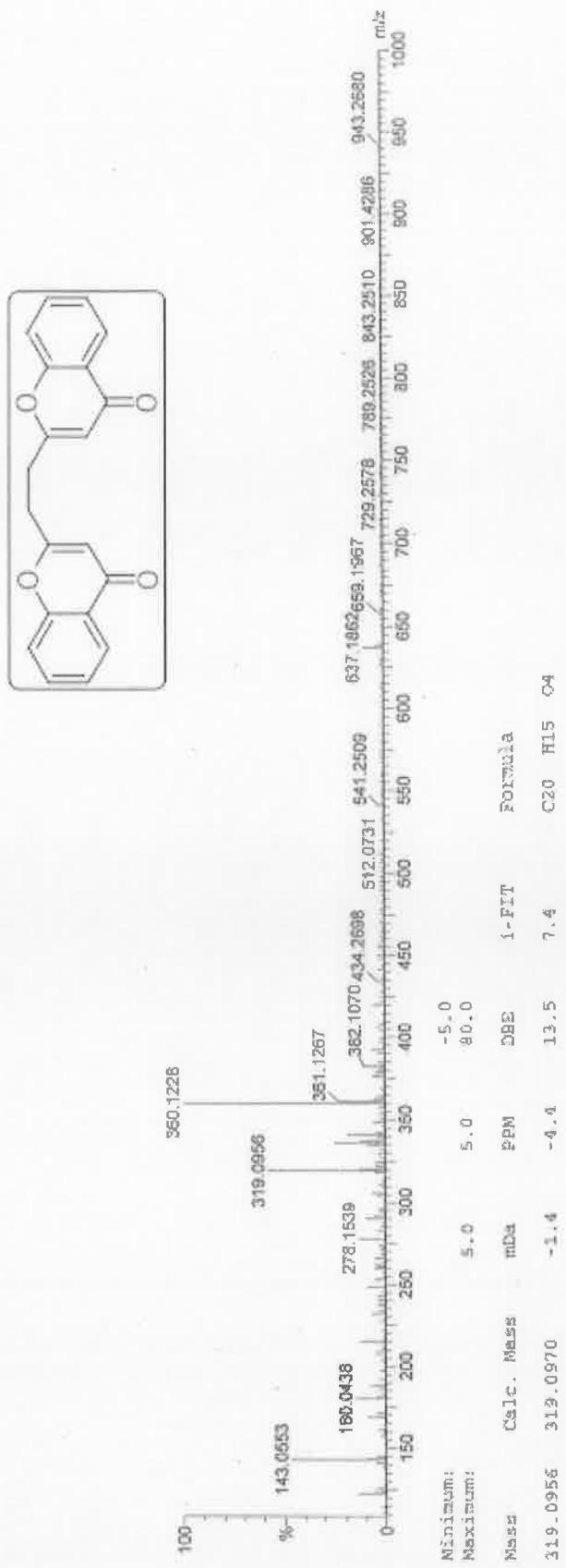
¹H NMR (DMSO-d6) of 2,2'-{(Ethane-1,2-diyl)bis(4H-chromen-4-one)} (**9a**)



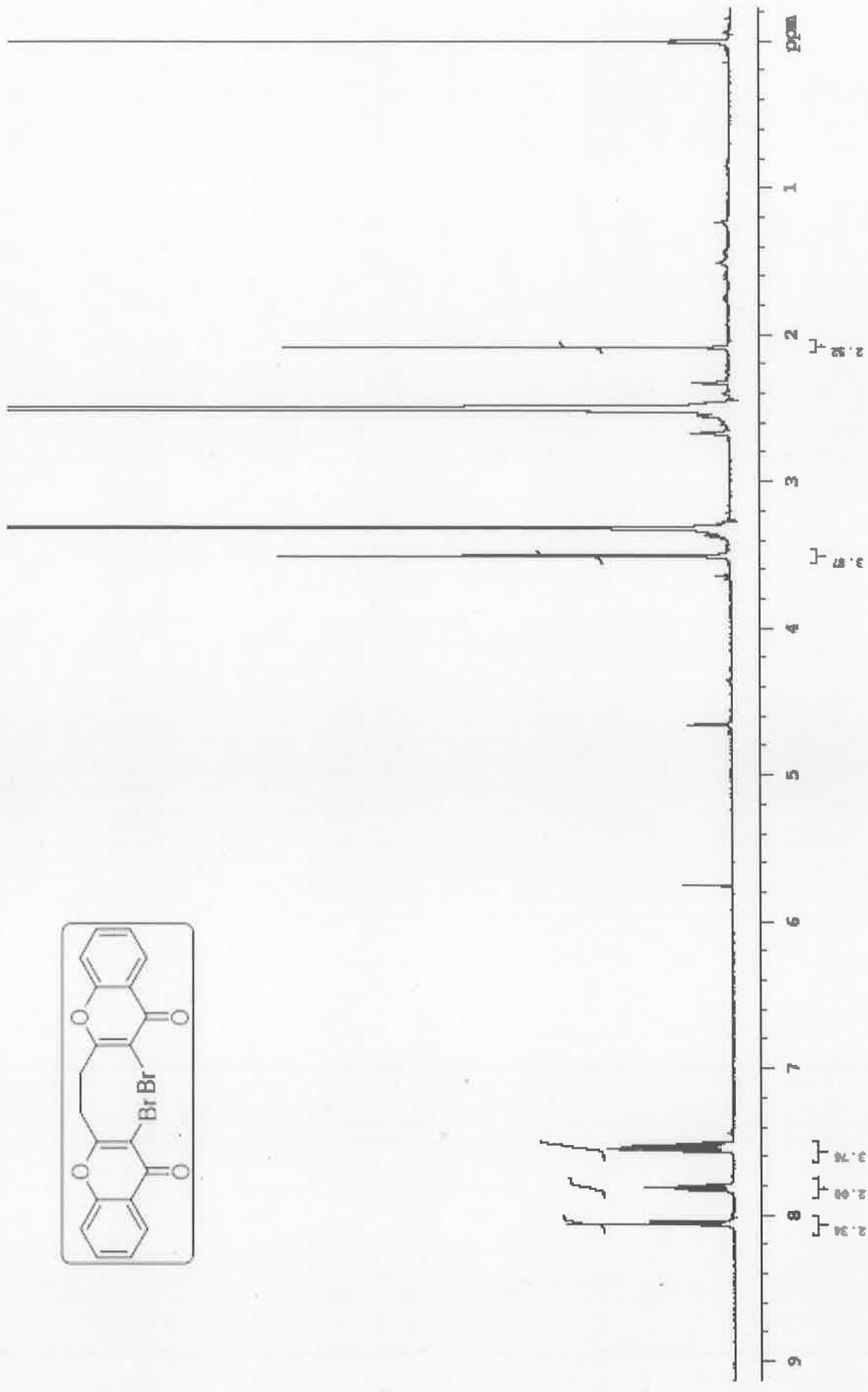
^{13}C NMR (DMSO-d₆) of 2,2'-{(Ethane-1,2-diyl)bis(4H-chromen-4-one)} (**9a**)



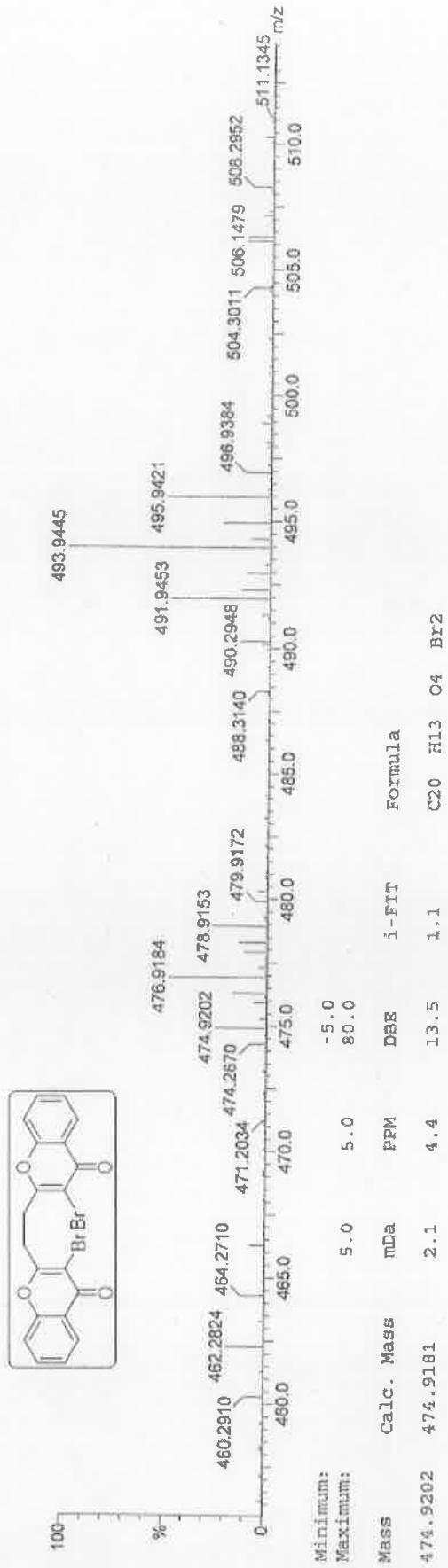
HRMS of 2,2'-(Ethane-1,2-diy)bis(4H-chromen-4-one) (**9a**)



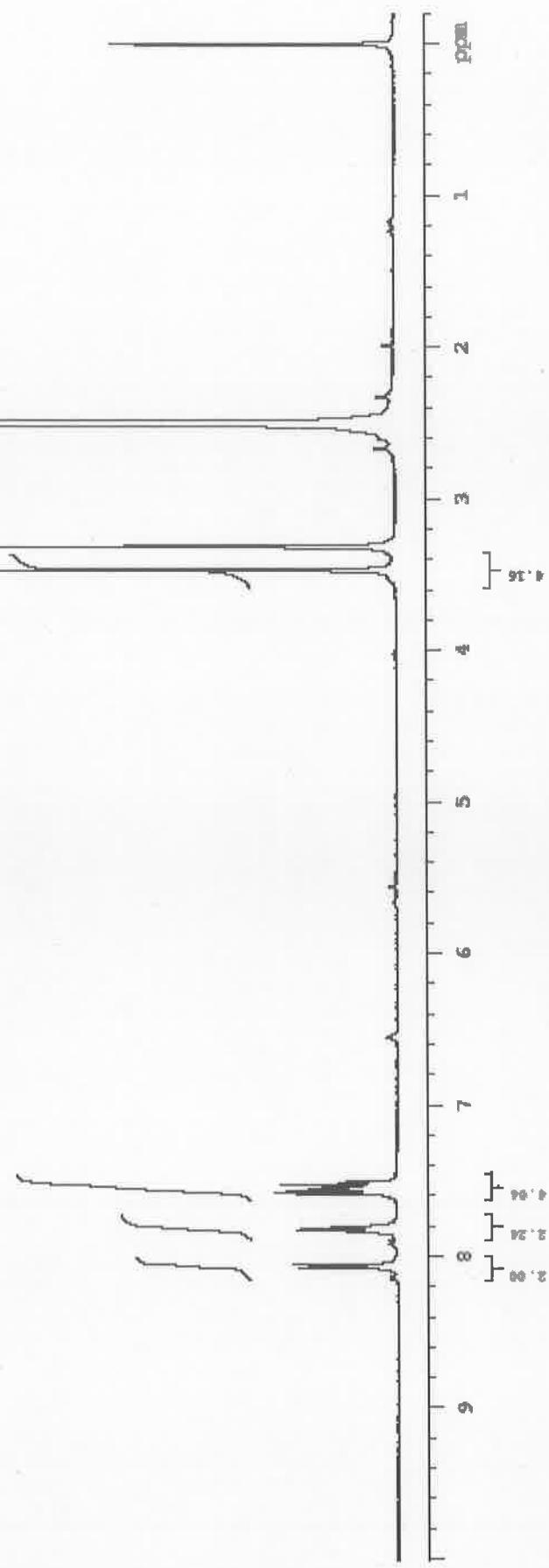
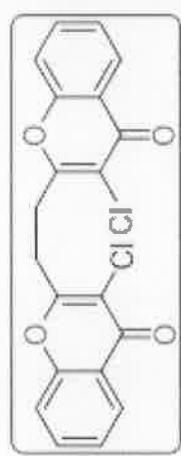
¹H NMR (DMSO-d₆) of 2,2'-(Ethane-1,2-diy)bis(3-bromo-4H-chromen-4-one) (9b)



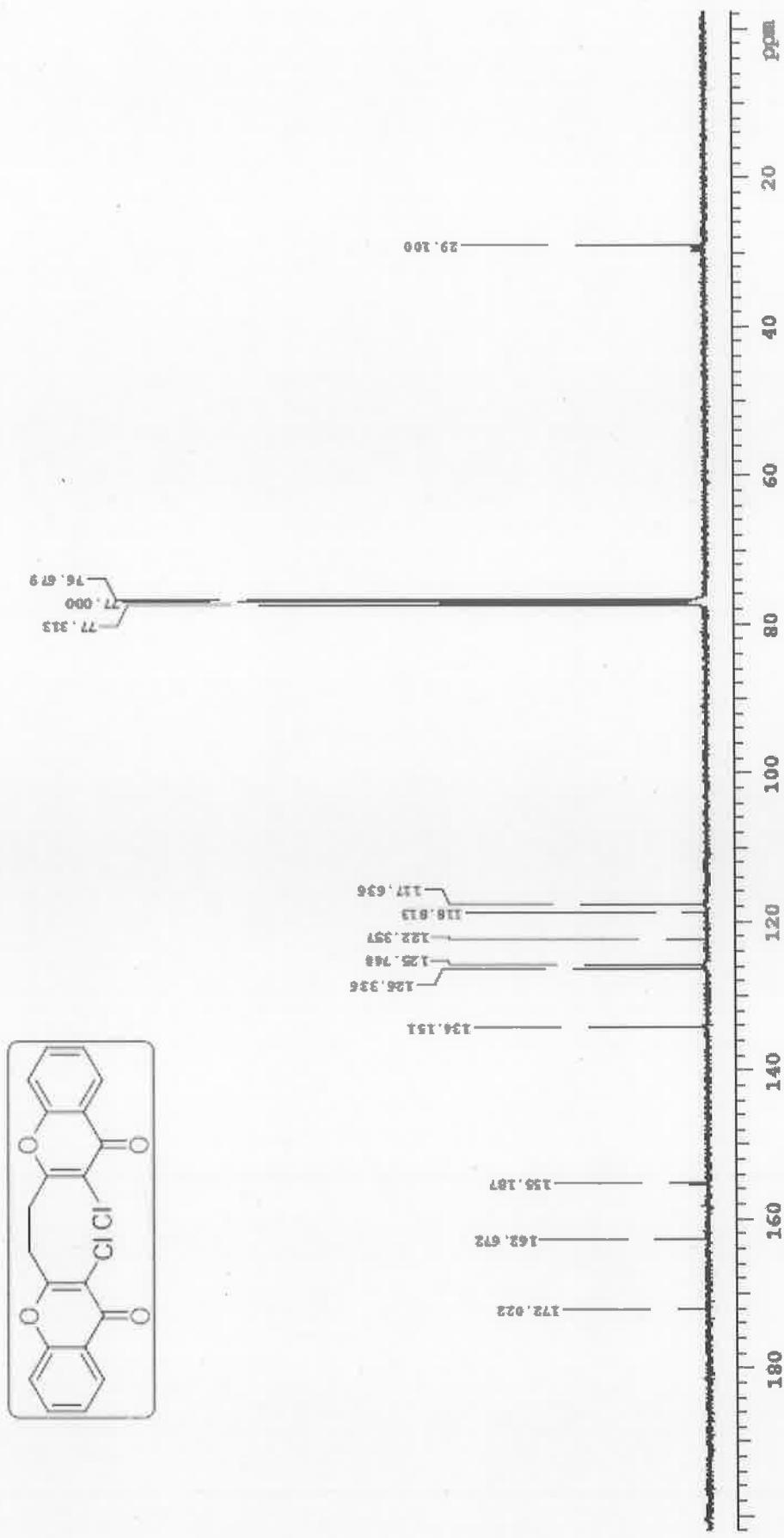
HRMS of 2,2'-(Ethane-1,2-diyl)bis(3-bromo-4H-chromen-4-one) (**9b**)



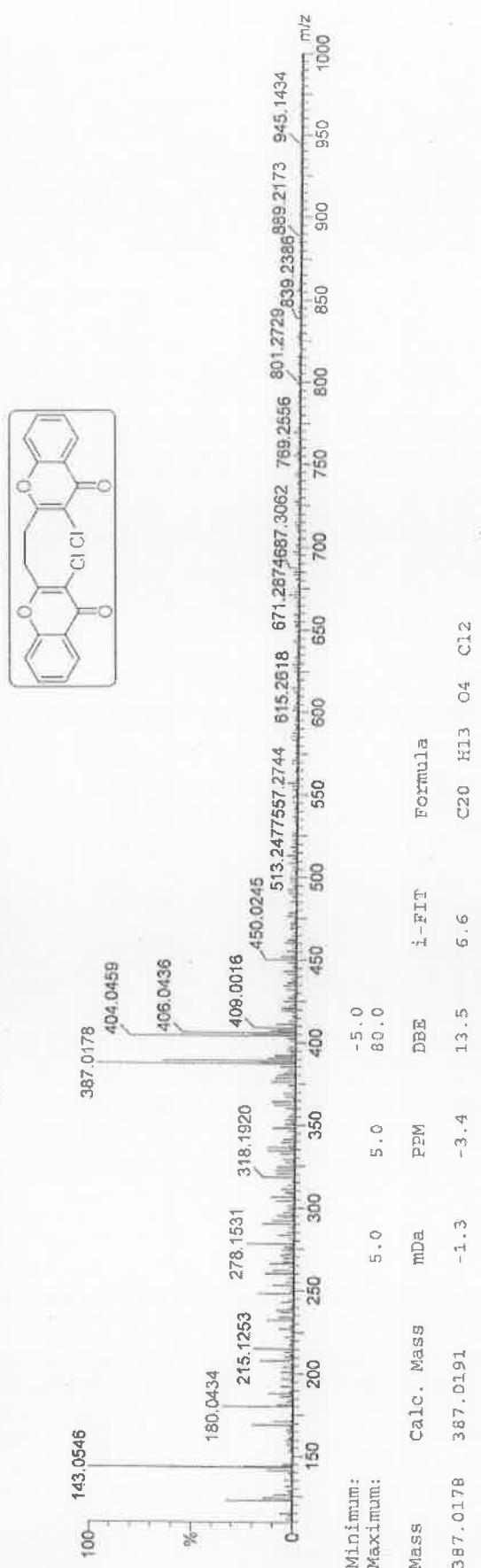
¹H NMR (DMSO-d₆) of 2,2'-(Ethane-1,2-diyl)bis(3-chloro-4H-chromen-4-one) (9c)



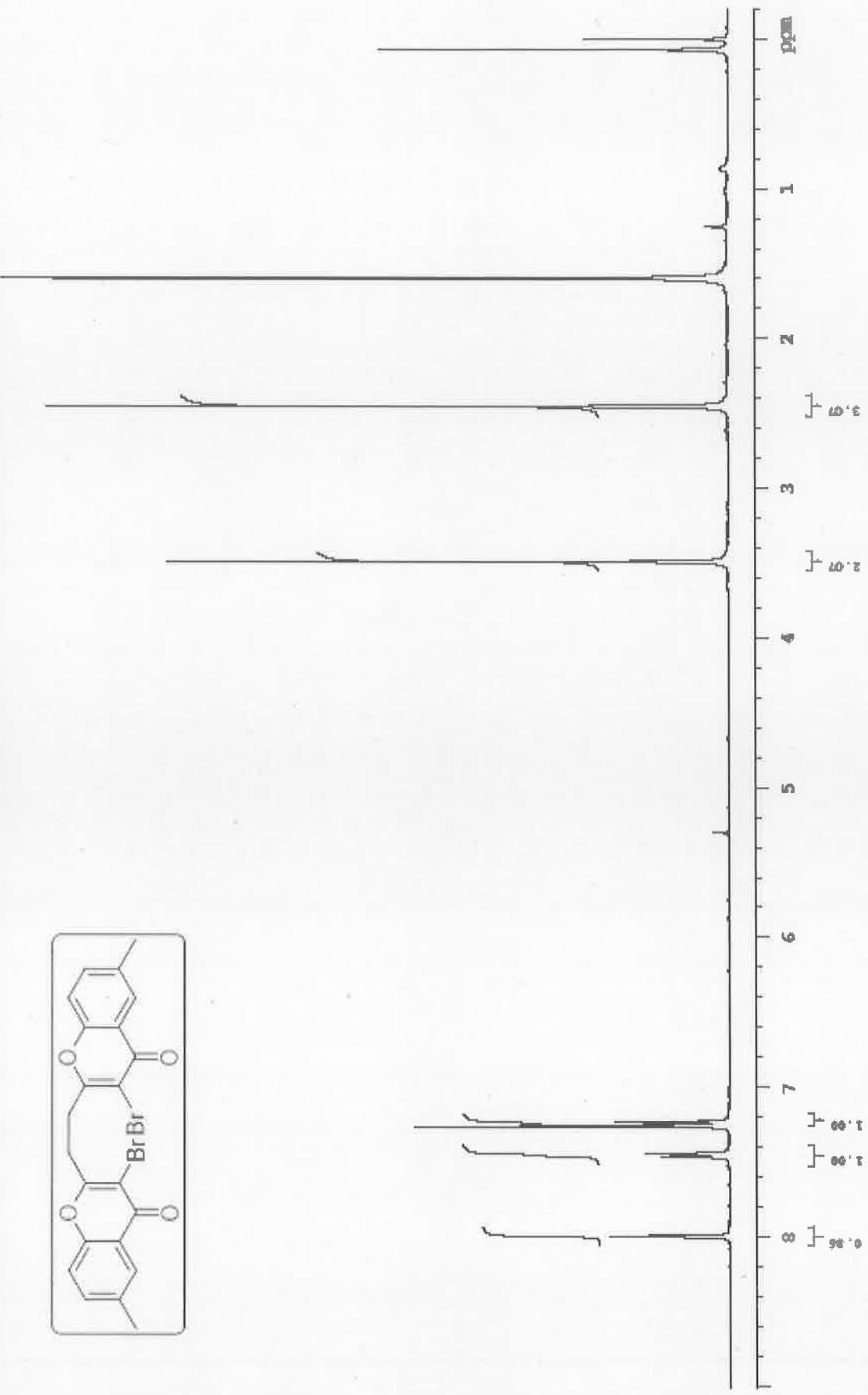
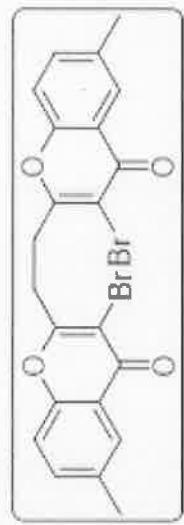
^{13}C NMR (CDCl_3) of 2,2'-(Ethane-1,2-diy)bis(3-chloro-4H-chromen-4-one) (9c)



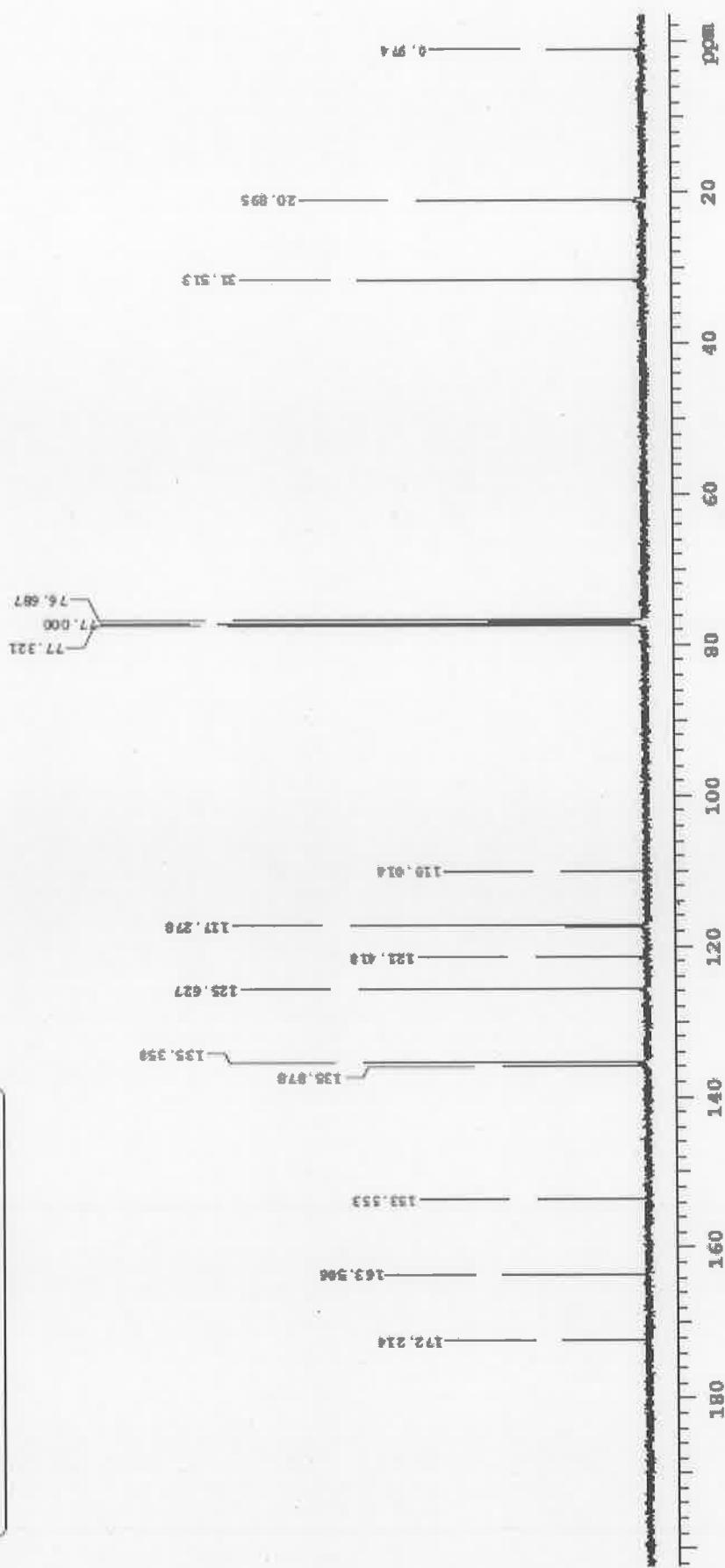
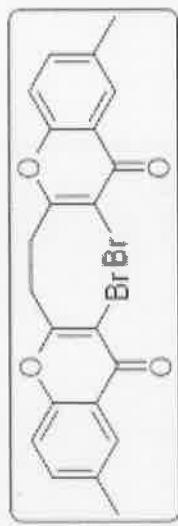
HRMS of 2,2'-(ethane-1,2-diyl)bis(3-chloro-4H-chromen-4-one) (9c)



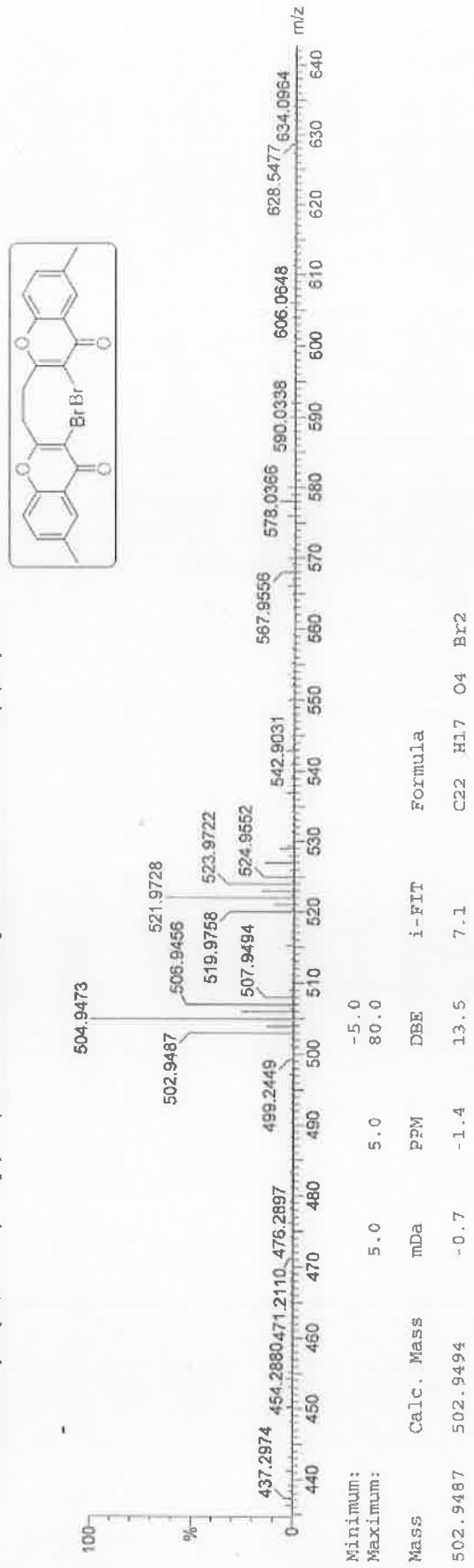
¹H NMR (CDCl_3) of 2,2'-(Ethane-1,2-diy)bis(3-bromo-6-methyl-4H-chromen-4-one) (9d)



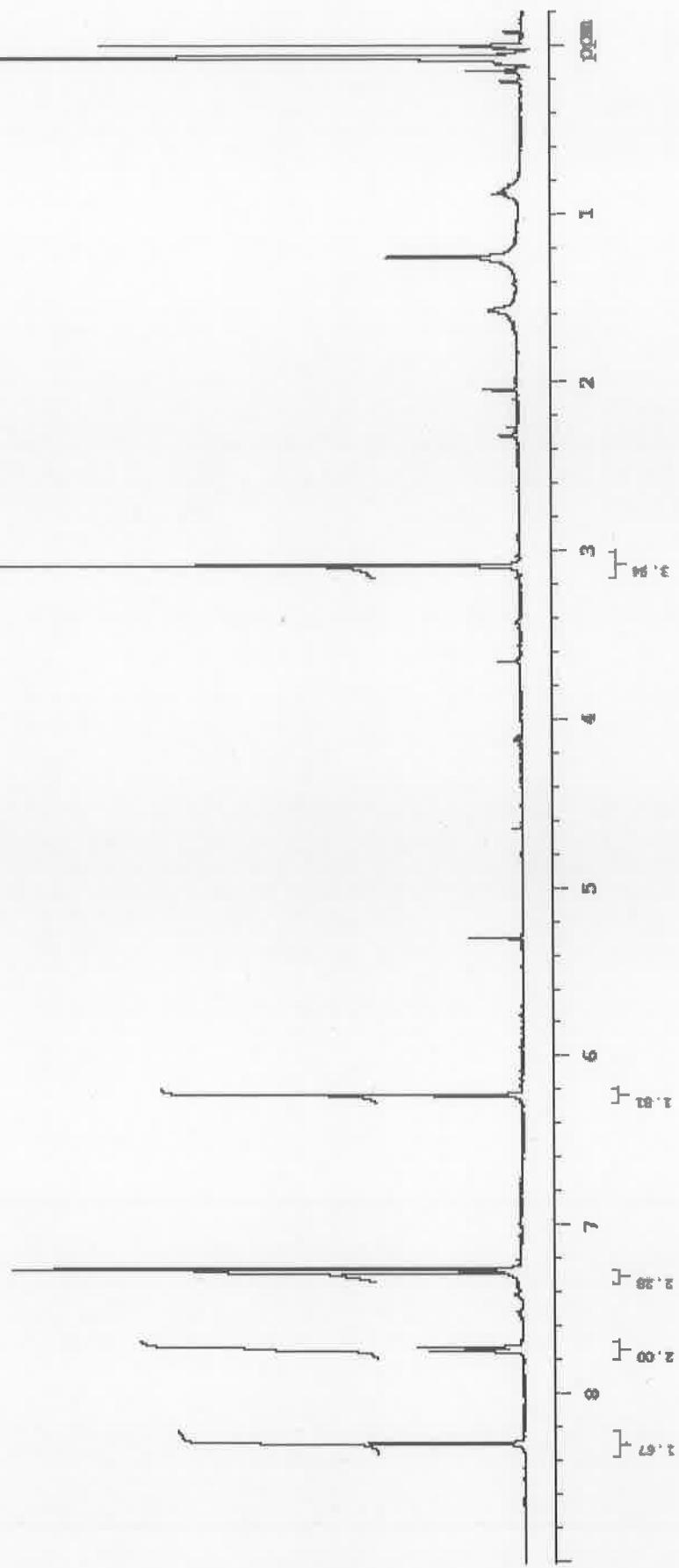
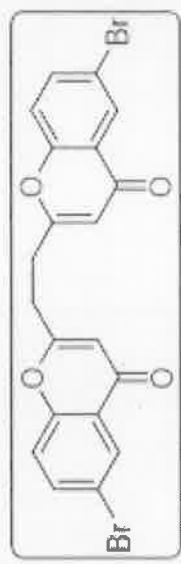
^{13}C NMR (CDCl_3) of 2,2'-(Ethane-1,2-diy)bis(3-bromo-6-methyl-4H-chromen-4-one) (9d)



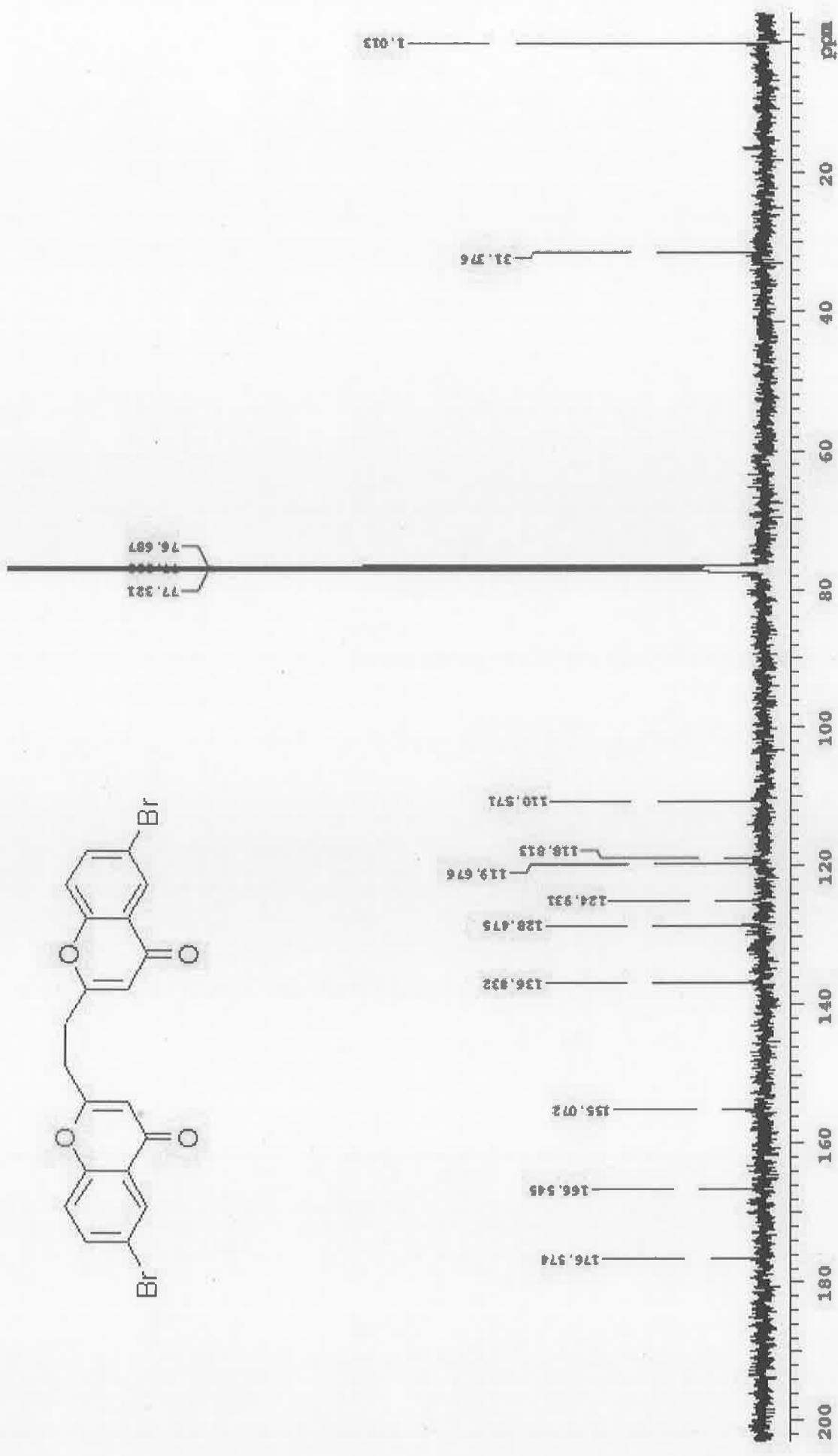
HRMS of 2,2'-(Ethane-1,2-diy)bis(3-bromo-6-methyl-4H-chromen-4-one) (**9d**)



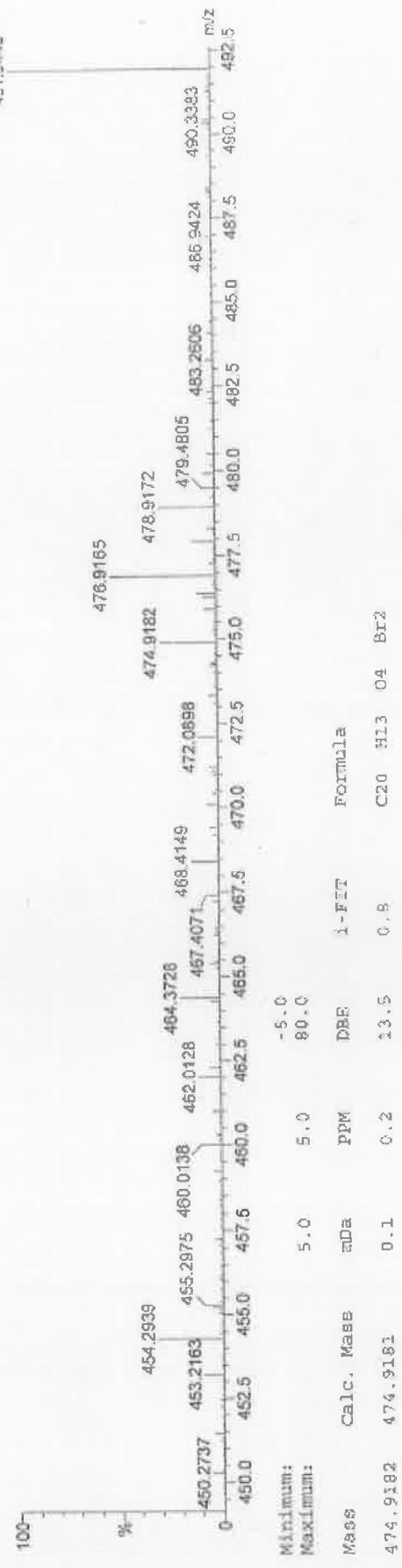
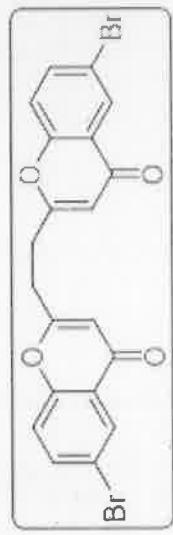
¹H NMR (CDCl_3) of 2,2'-(Ethane-1,2-diy)bis(6-bromo-4H-chromen-4-one) (9e)



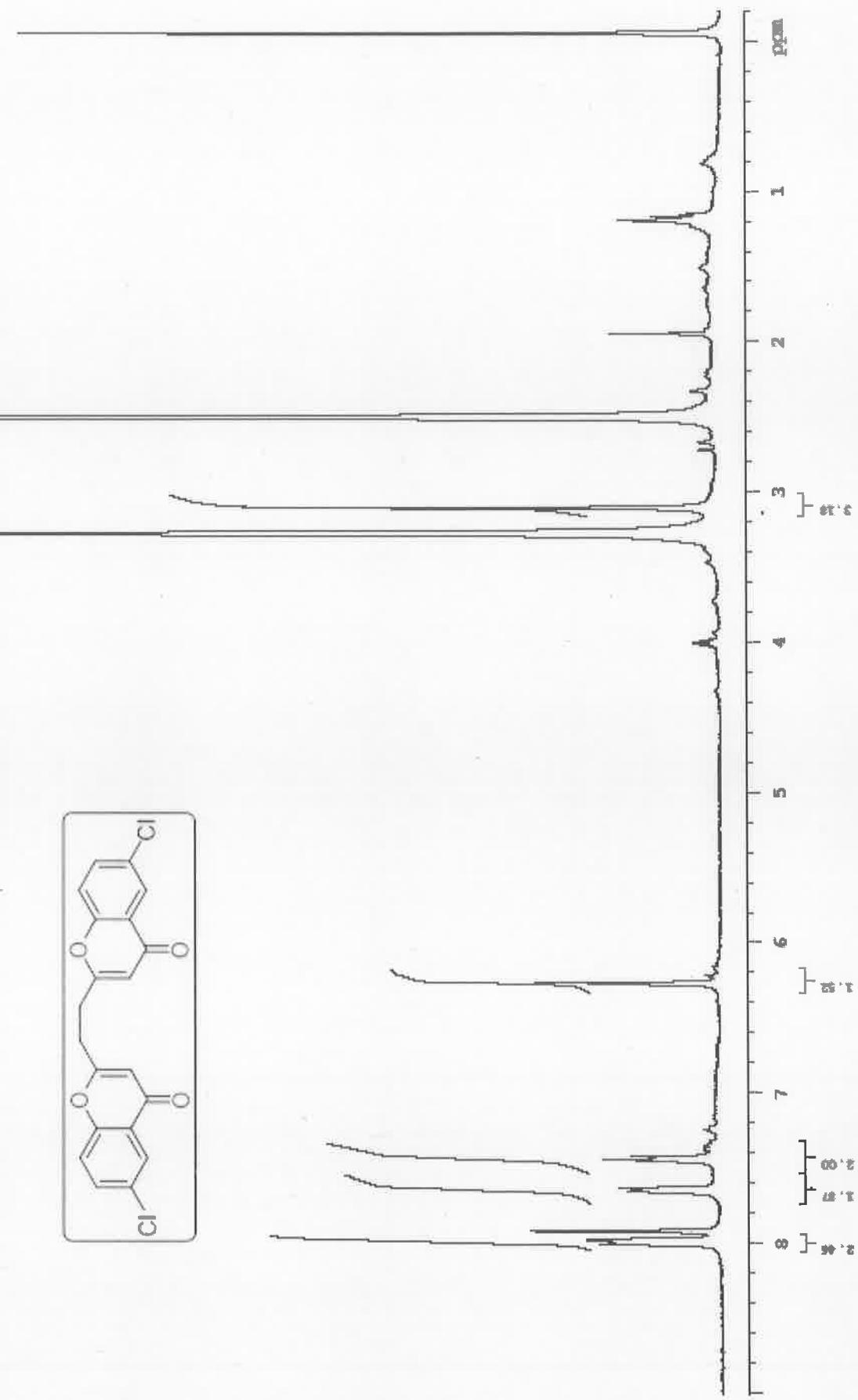
^{13}C NMR (CDCl_3) of 2,2'-(Ethane-1,2-diy)bis(6-bromo-4H-chromen-4-one) (**9e**)



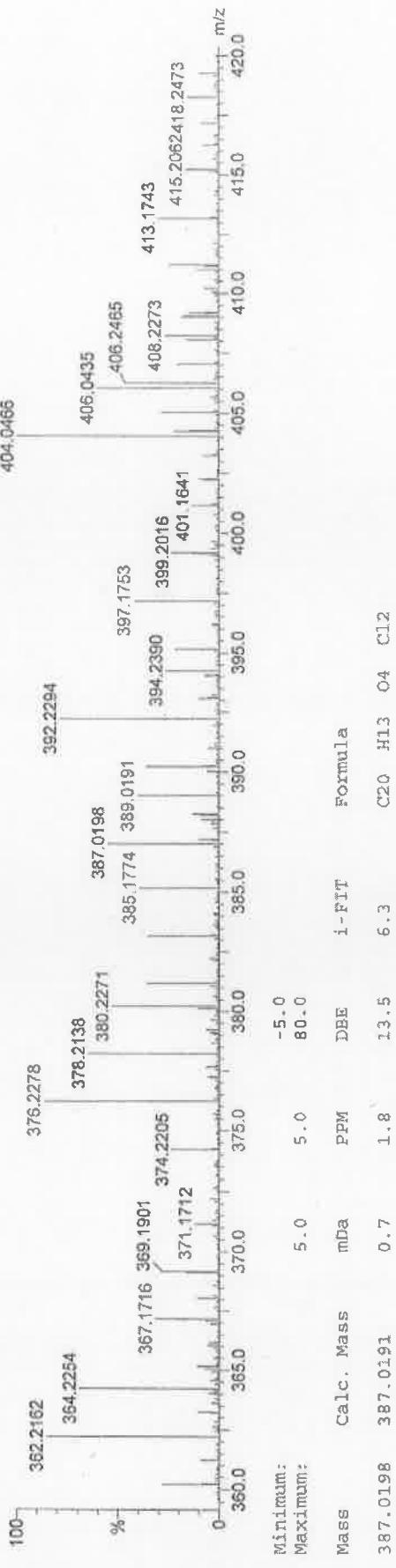
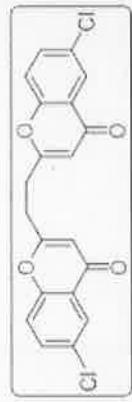
HRMS of 2,2'-(Ethane-1,2-diy)bis(6-bromo-4H-chromen-4-one) (9e)



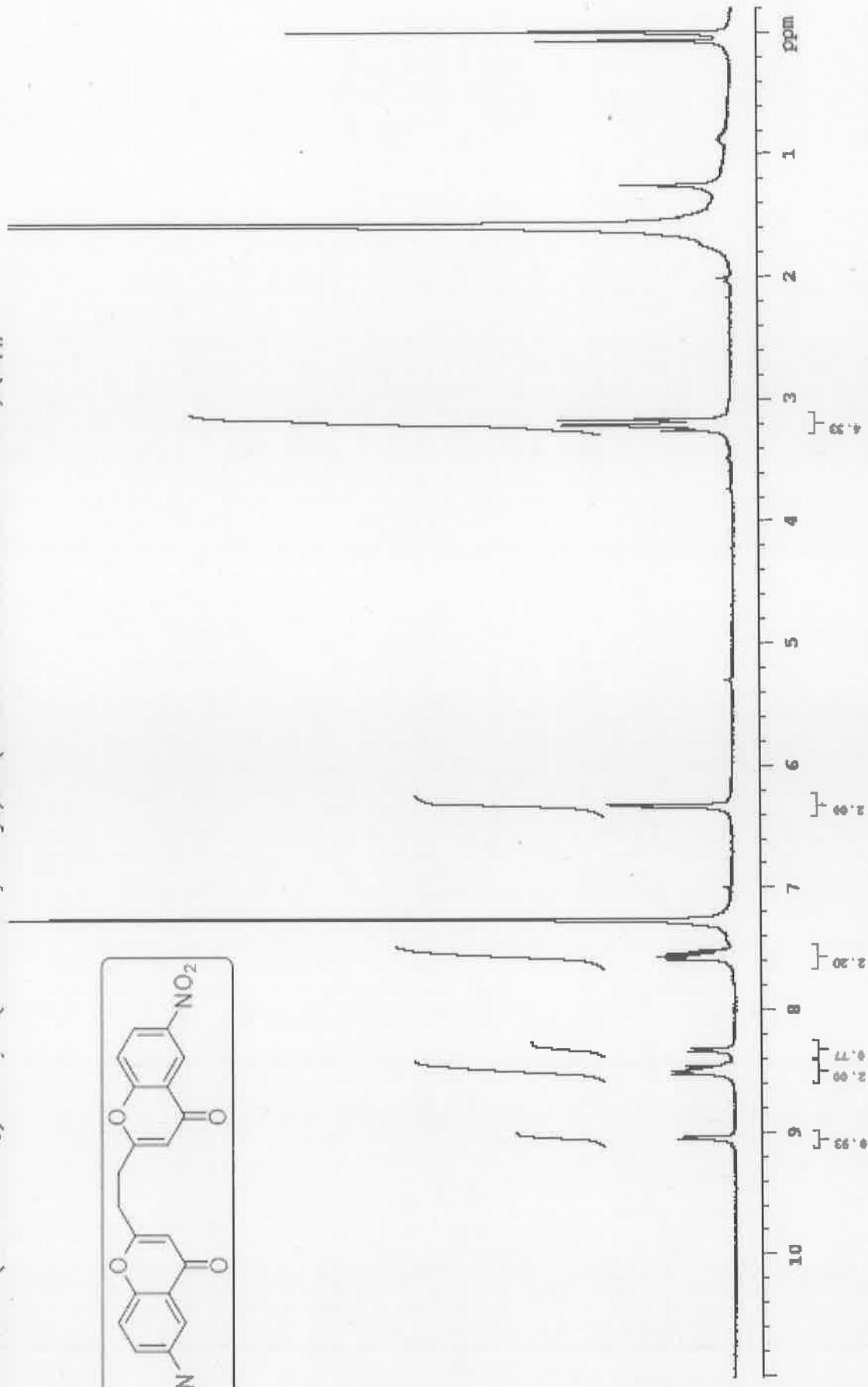
^1H NMR ($\text{CDCl}_3 + \text{DMSO-d}_6$) of 2,2'-(Ethane-1,2-diyl)bis(6-chloro-4H-chromen-4-one) (9f)



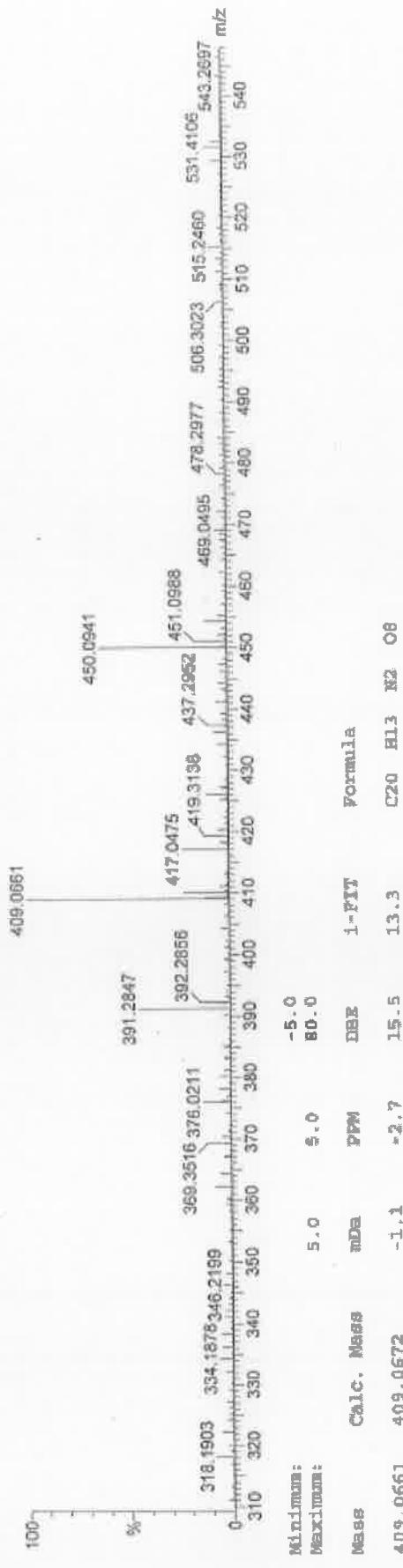
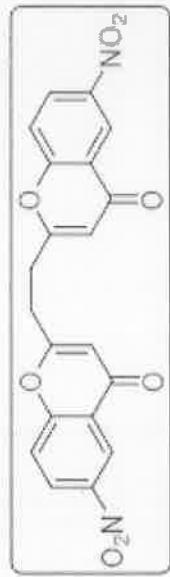
HRMS of 2,2'-(Ethane-1,2-diy)bis(6-chloro-4H-chromen-4-one) (9f)



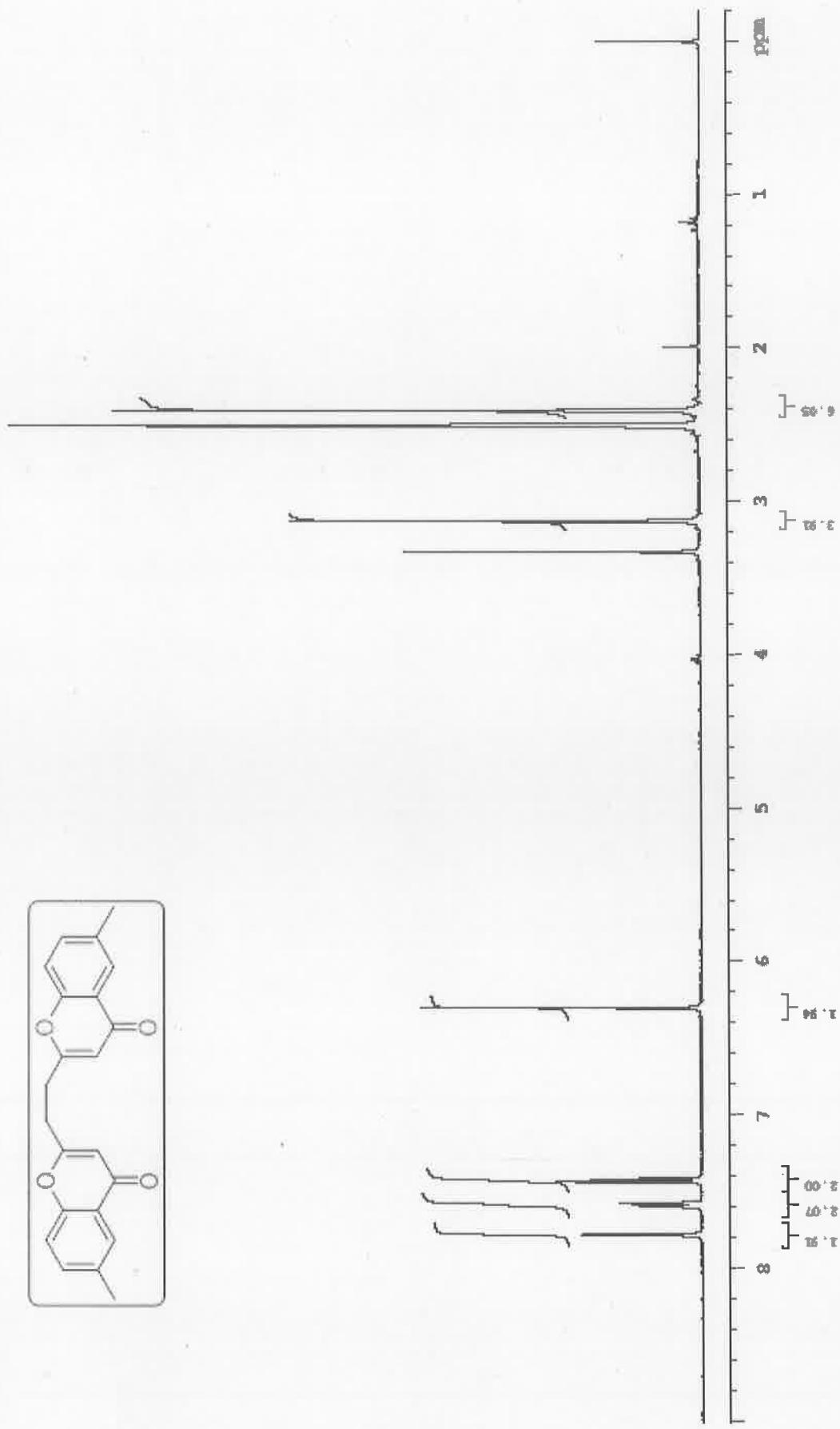
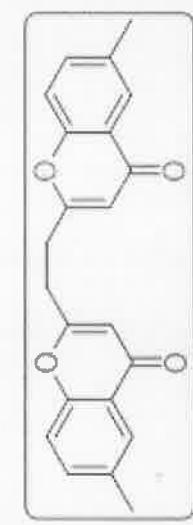
¹H NMR (DMSO-d₆) of 2,2'-(Ethane-1,2-diy)bis(6-nitro-4H-chromen-4-one) (9g)



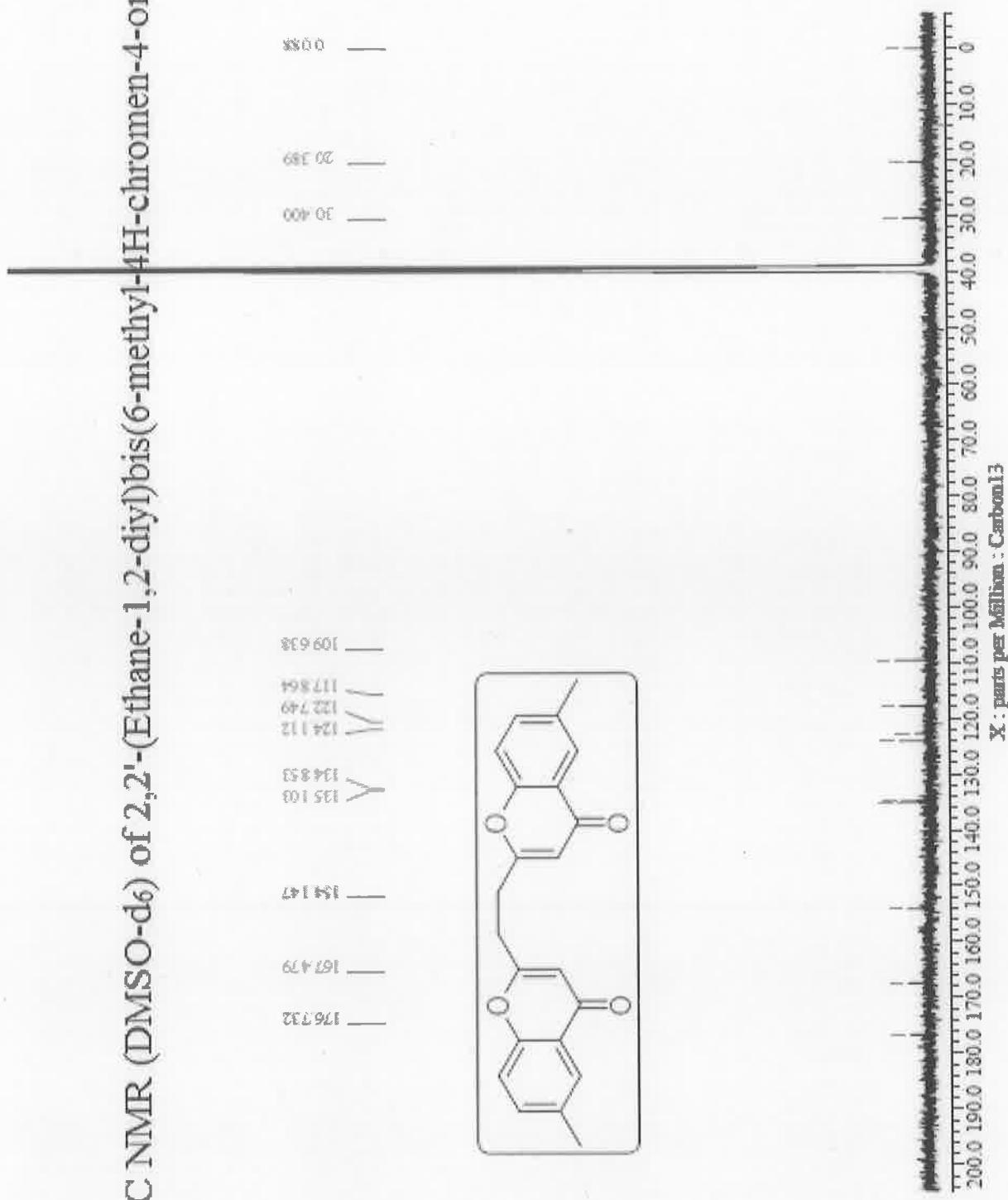
HRMS of 2,2'-(Ethane-1,2-diy)bis(6-nitro-4H-chromen-4-one) (9g)



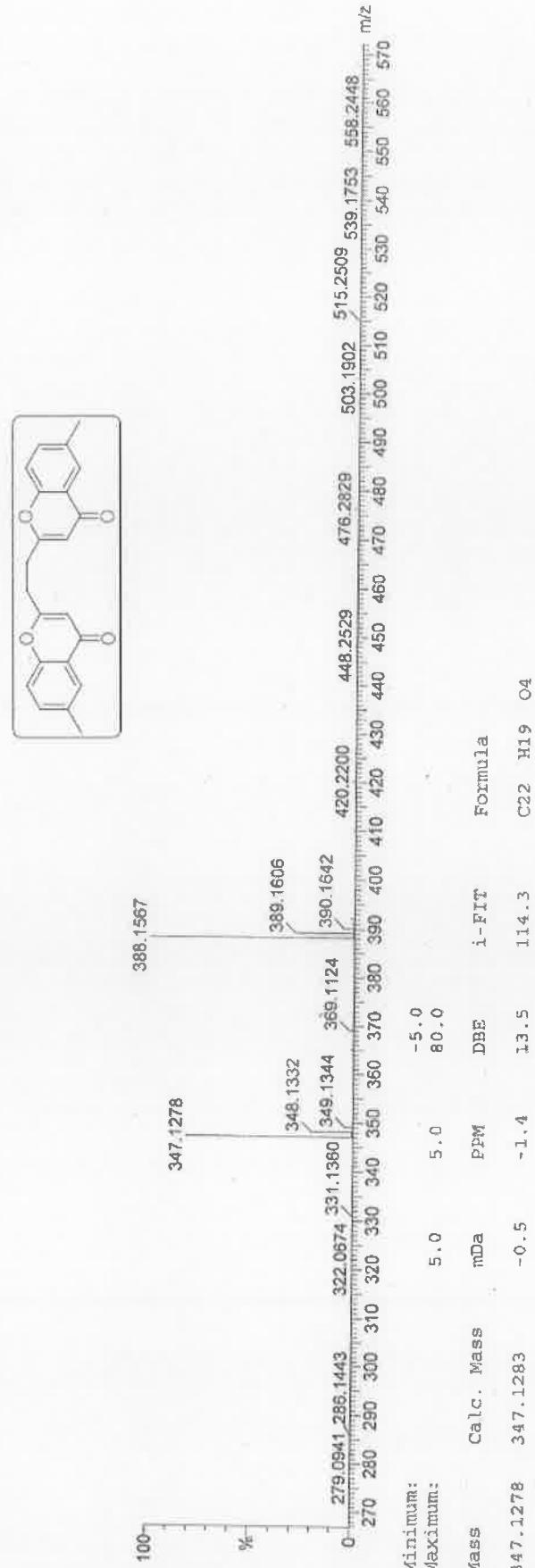
¹H NMR (DMSO-d₆) of 2,2'-(Ethane-1,2-diy)bis(6-methyl-4H-chromen-4-one) (**9h**)



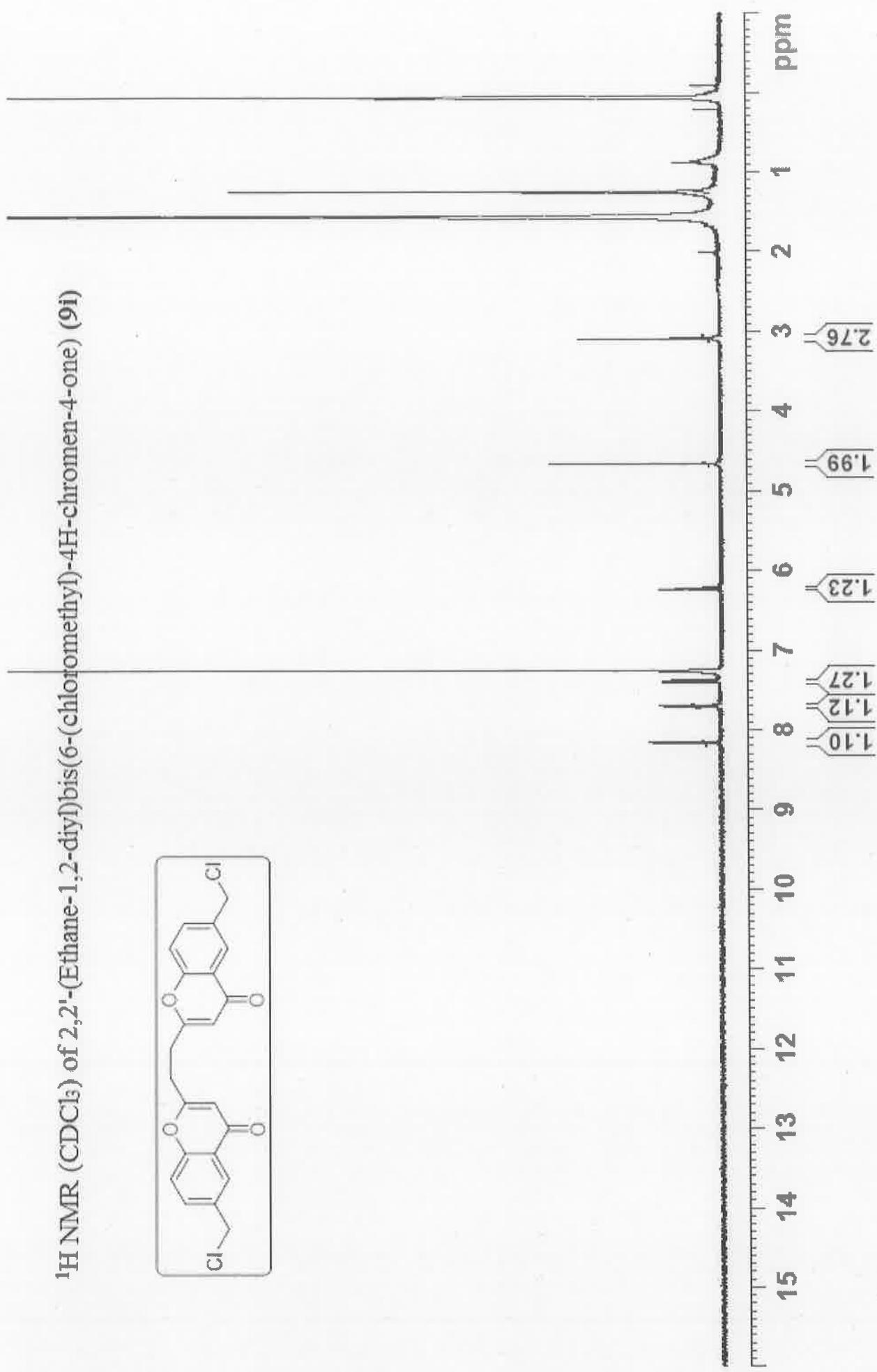
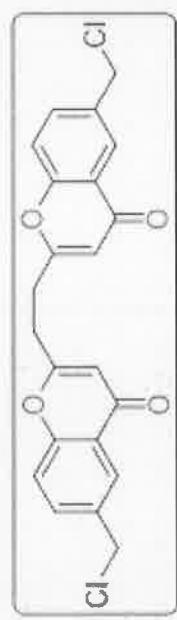
^{13}C NMR (DMSO-d₆) of 2,2'-(Ethane-1,2-diyl)bis(6-methyl-4H-chromen-4-one) (9h)



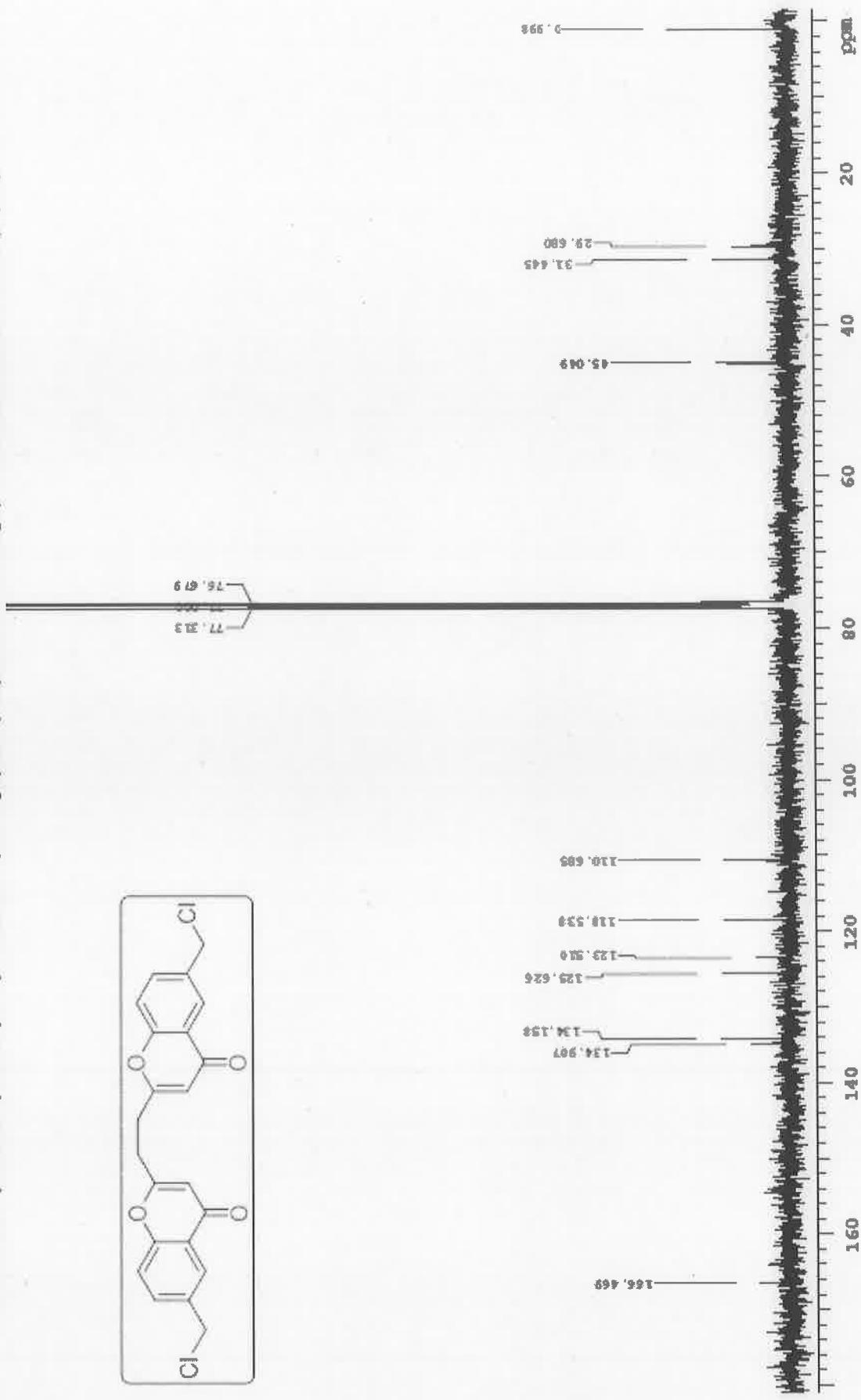
HRMS of 2,2'-(Ethane-1,2-diy)bis(6-methyl-4H-chromen-4-one) (9h)



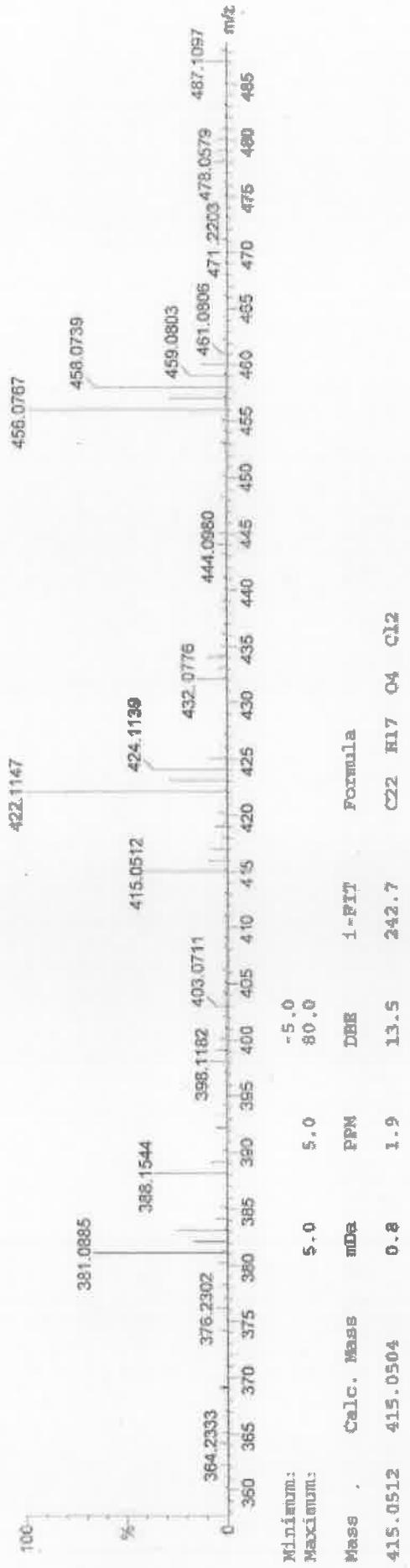
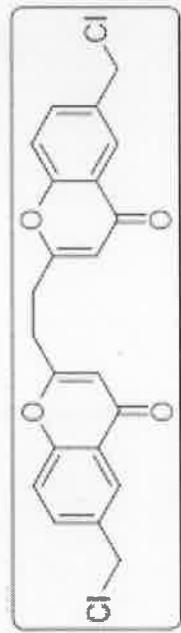
¹H NMR (CDCl_3) of 2,2'-(Ethane-1,2-diy)bis(6-(chloromethyl)-4H-chromen-4-one) (9i)



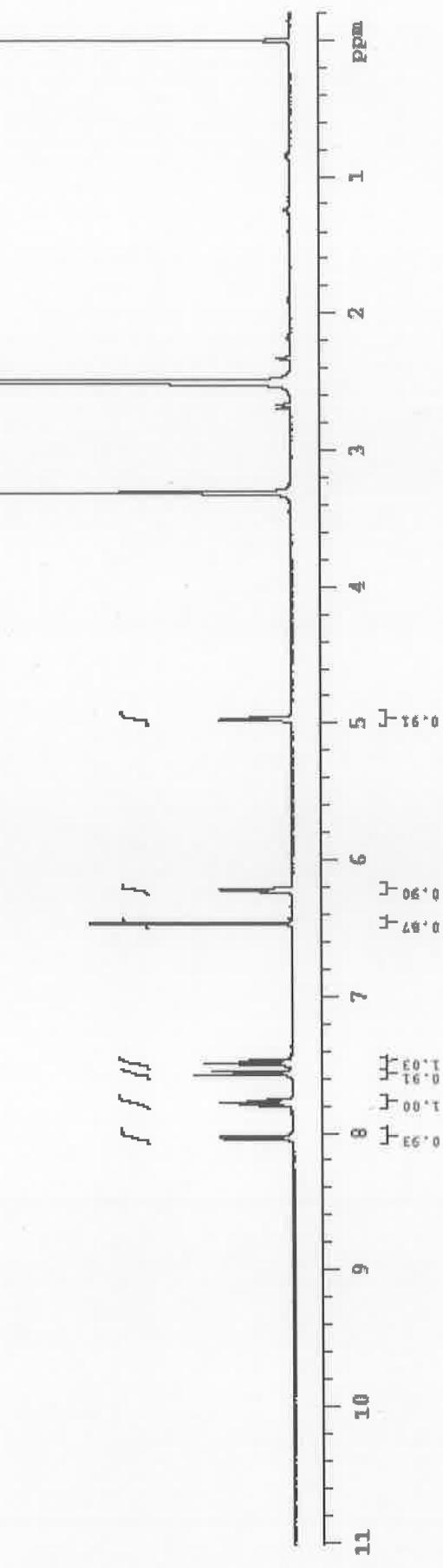
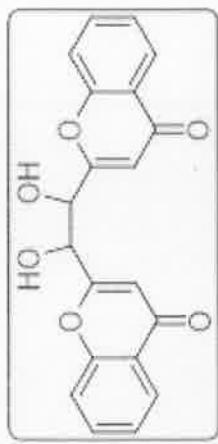
^{13}C NMR (CDCl_3) of 2,2'-(Ethane-1,2-diy)-bis(6-(chloromethyl)-4H-chromen-4-one) (**9i**)



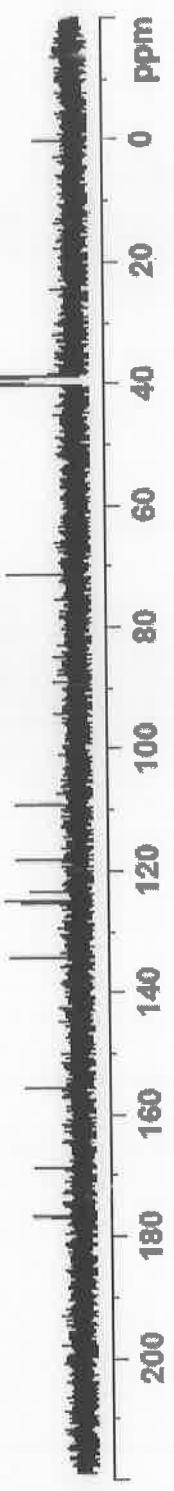
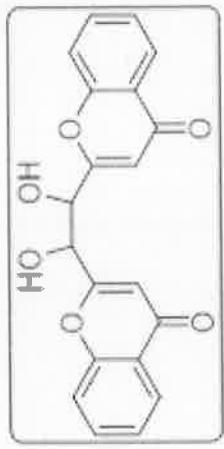
HRMS of 2,2'-(Ethane-1,2-diy)-bis(6-(chloromethyl)-4H-chromen-4-one) (9i)



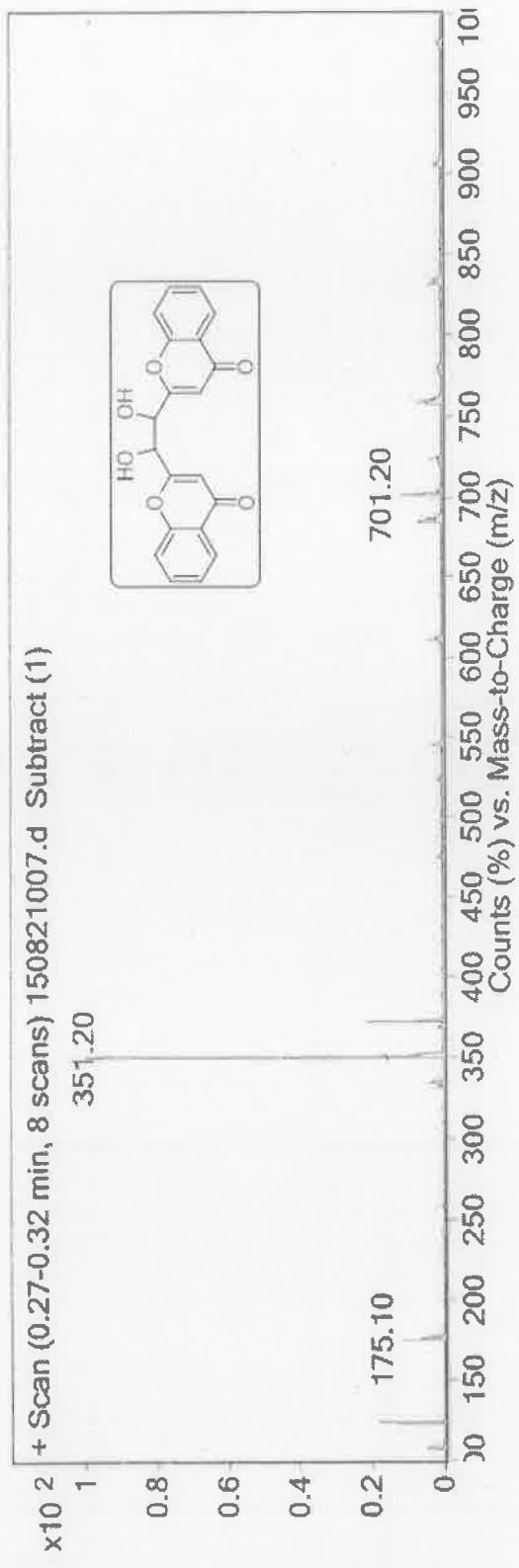
¹H NMR (CDCl_3) of 2,2'-(1,2-dihydroxyethane-1,2-diy)bis(4H-chromen-4-one) (11)



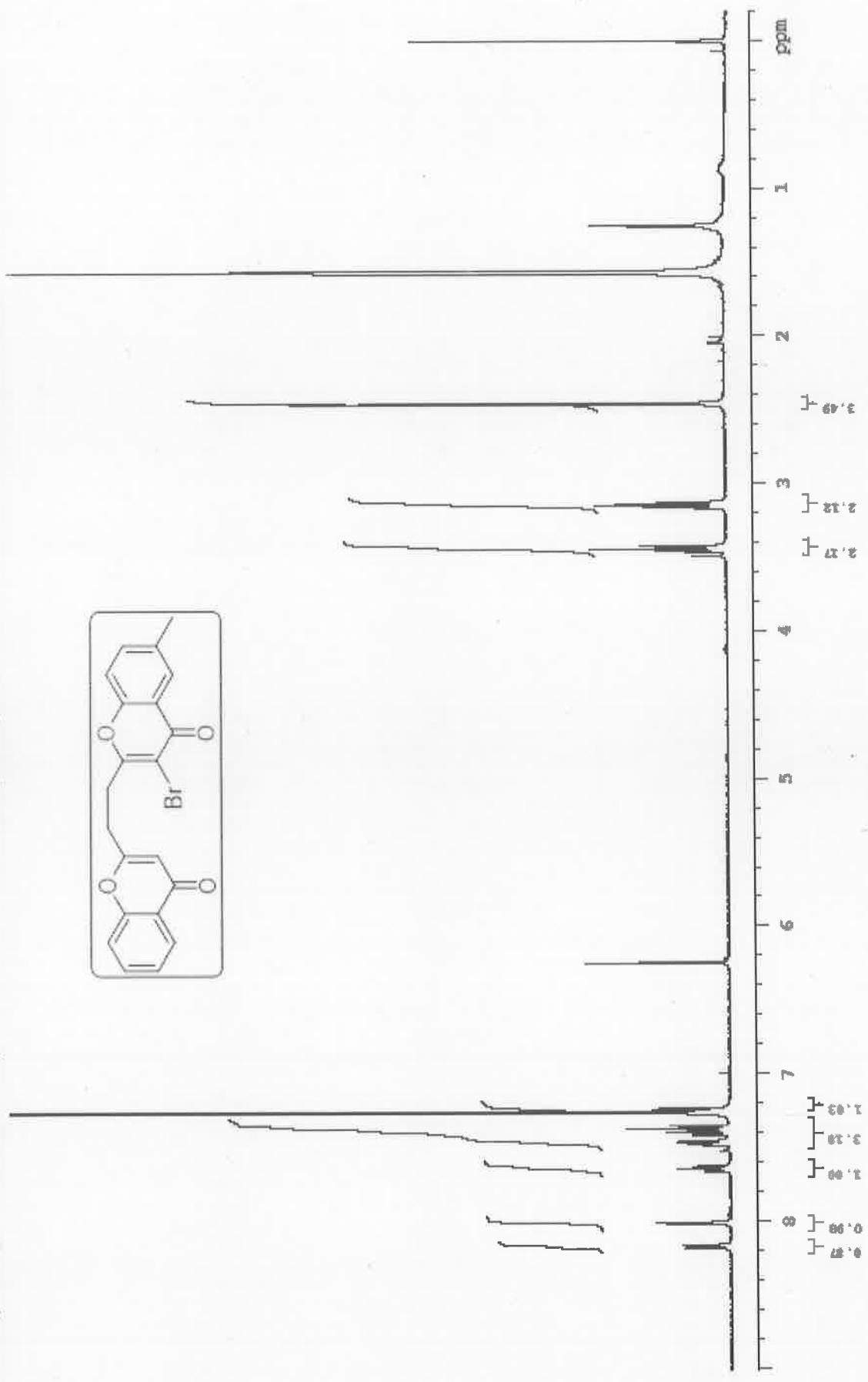
^{13}C NMR (DMSO-d_6) of 2,2'-(1,2-dihydroxyethane-1,2-diy)bis(4H-chromen-4-one) (11)



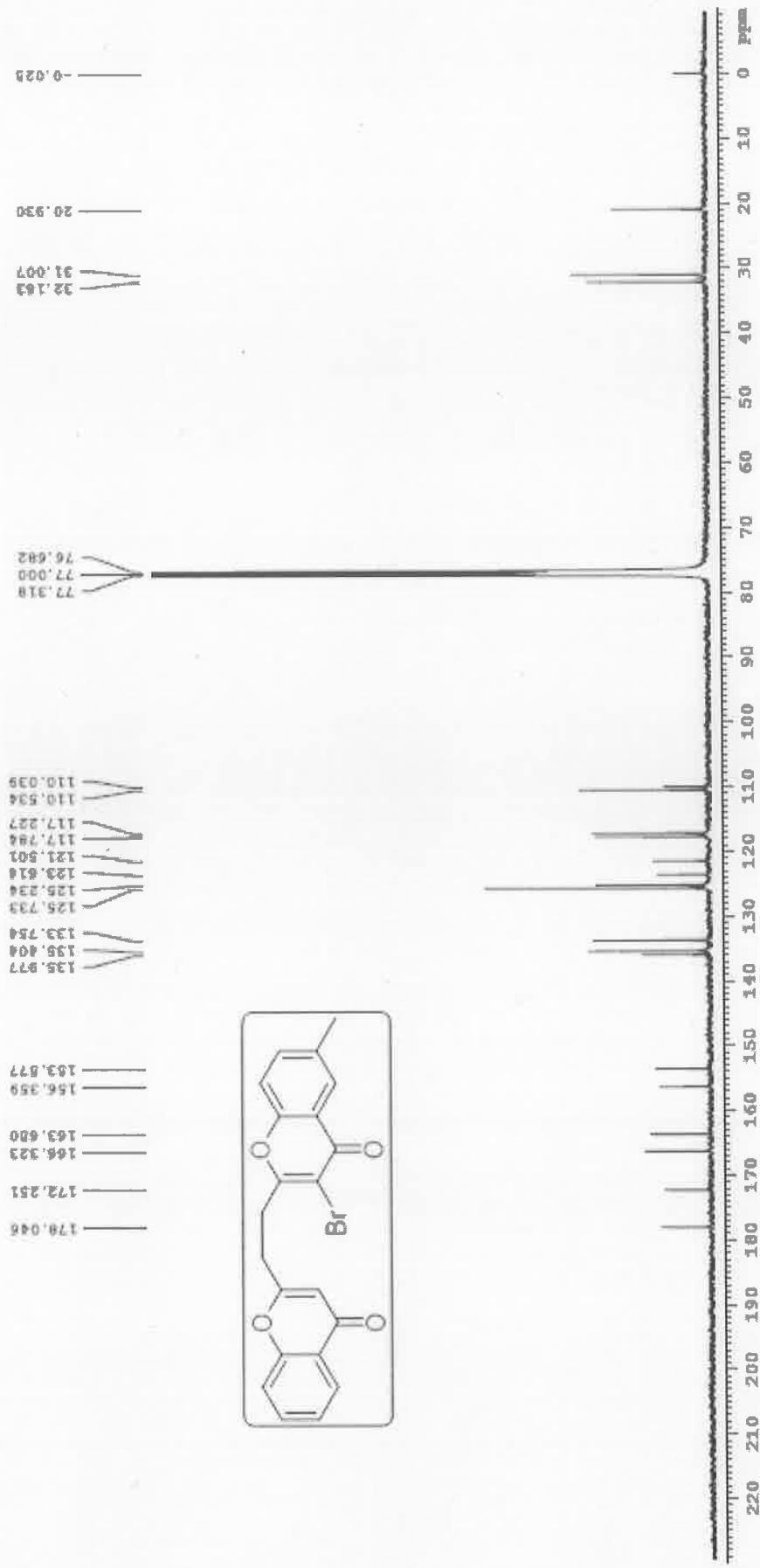
MASS of 2,2'-(1,2-dihydroxyethane-1,2-diyl)bis(4H-chromen-4-one) (11)



¹H NMR (CDCl₃ + DMSO-d₆) of 3-Bromo-6-methyl-2-(2-(4-oxo-4H-chromen-2-yl)ethyl)-4H-chromen-4-one (12)



^{13}C NMR (CDCl_3) of 3-Bromo-6-methyl-2-(2-(4-Oxo-4H-chromen-2-yl)ethyl)-4H-chromen-4-one (12)



HRMS of 3-Bromo-6-methyl-2-(2-(4-oxo-4H-chromen-2-yl)ethyl)-4H-chromen-4-one (**12**)

