

## Supplementary Material

### Chamigrane-type sesquiterpenes from *Laurencia dendroidea* as lead compounds against *Naegleria fowleri*

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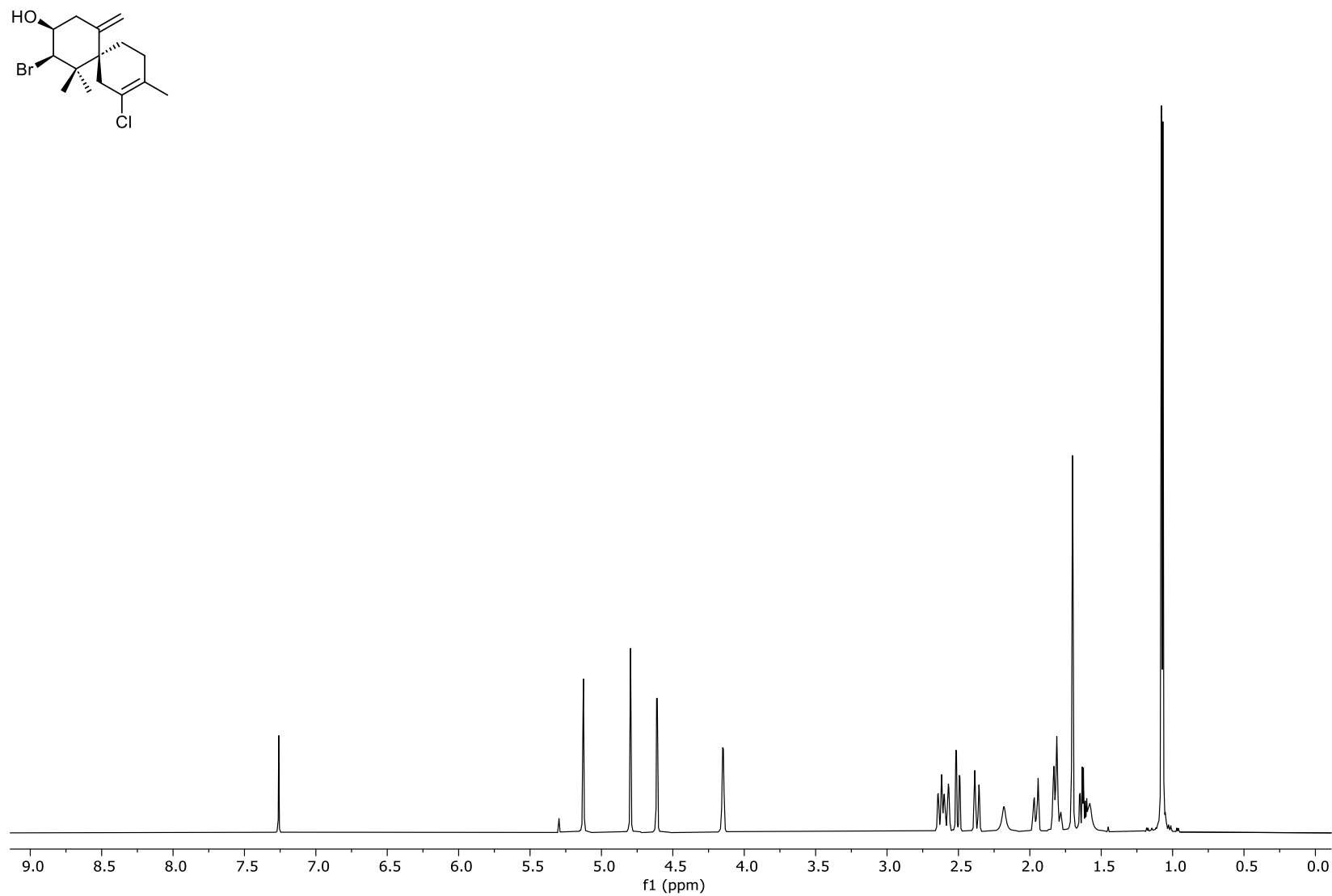
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**Figure S1:** <sup>1</sup>H NMR spectrum of (+)-elatol (**1**) at 600 MHz in CDCl<sub>3</sub>

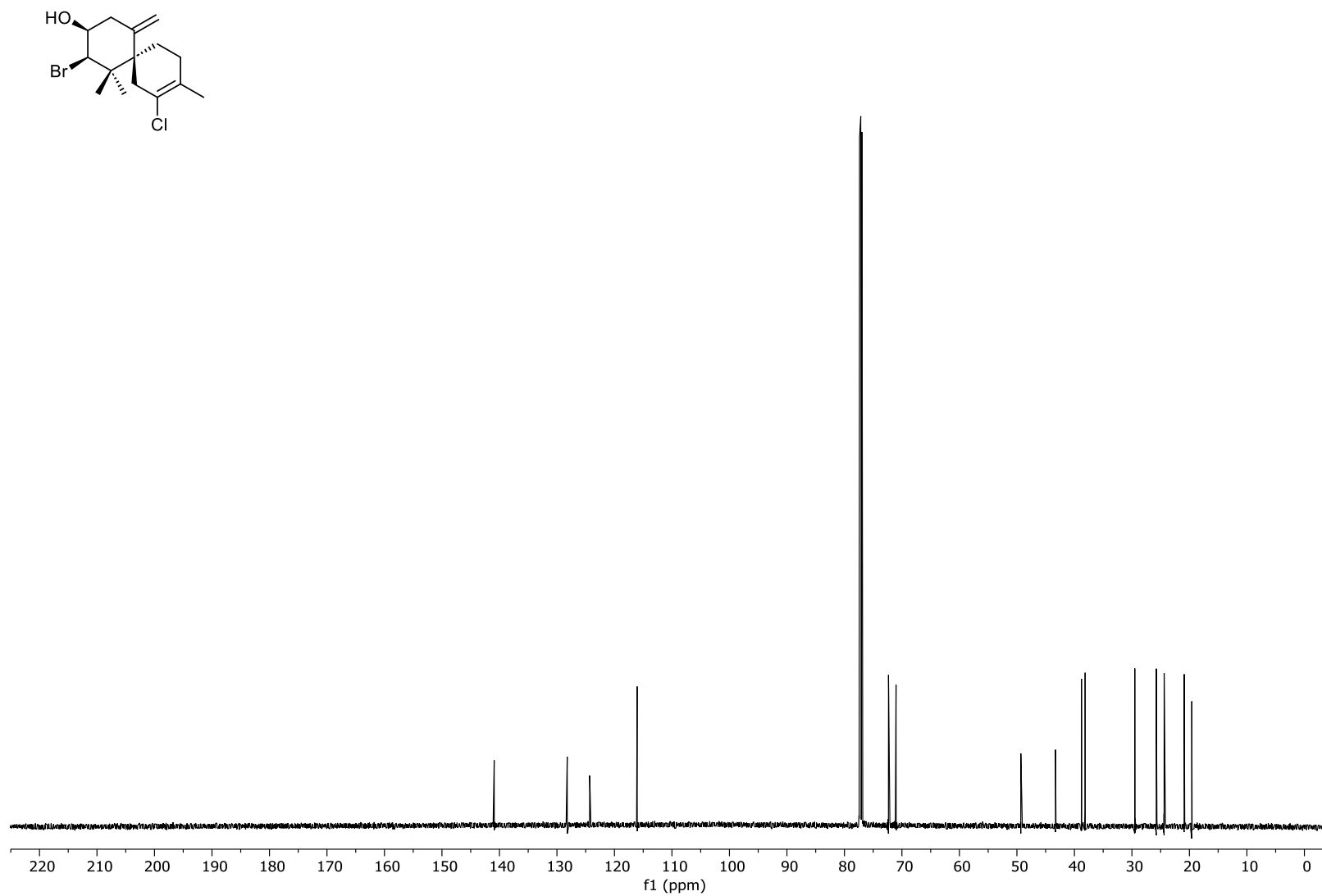
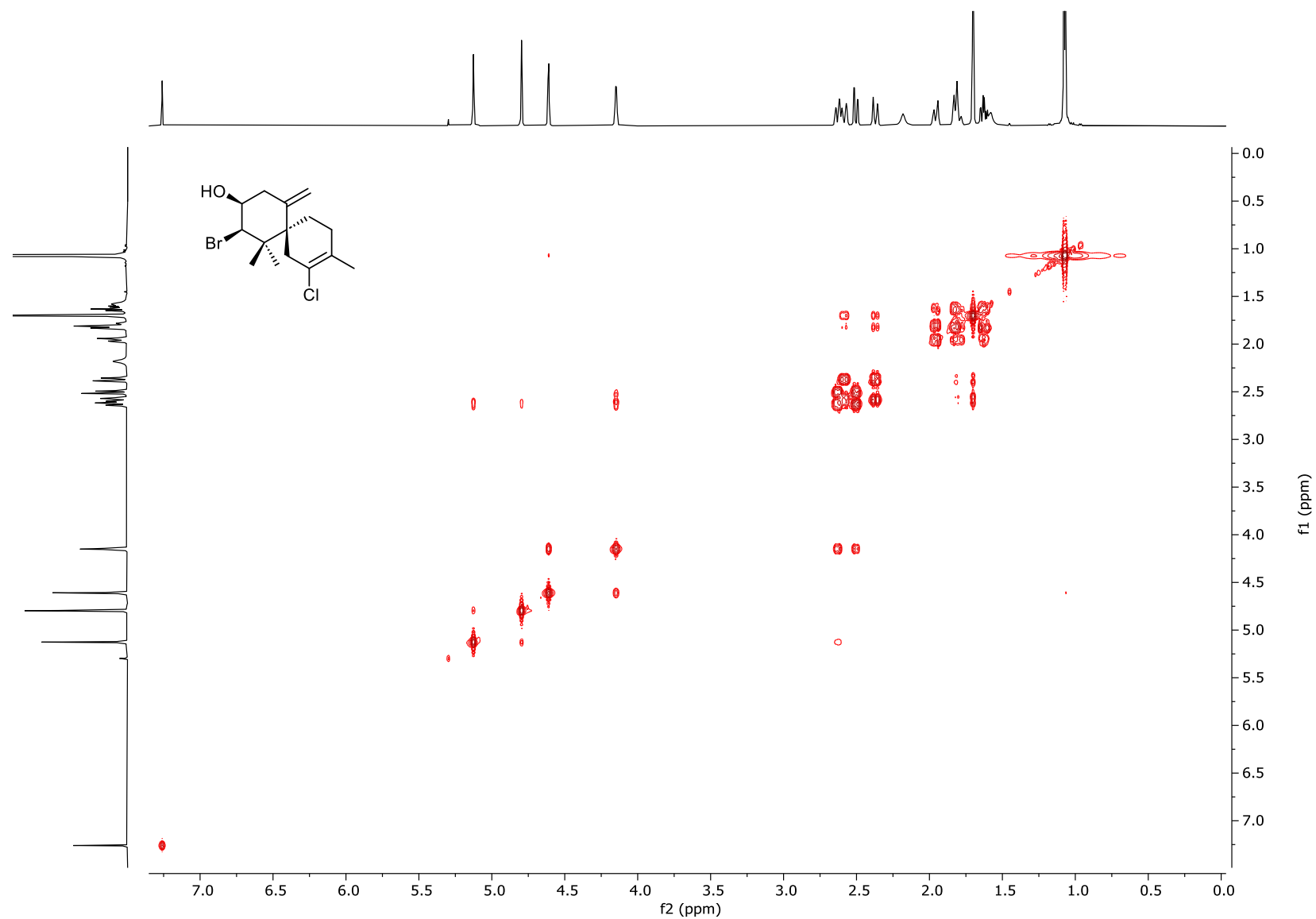
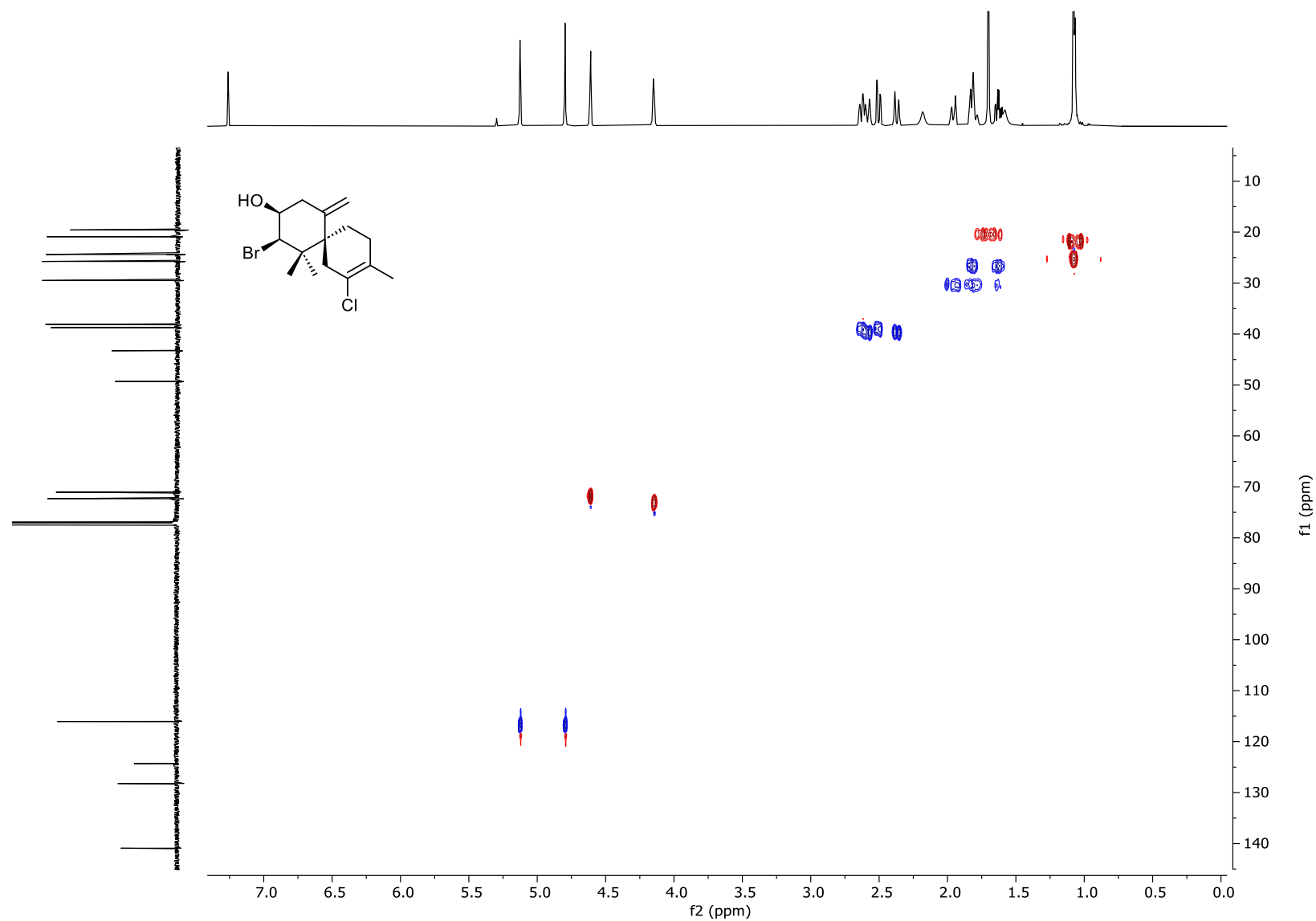


Figure S2:  $^{13}\text{C}$  NMR spectrum of (+)-elatol (1) at 600 MHz in  $\text{CDCl}_3$



**Figure S3:** COSY NMR spectrum of (+)-elatol (**1**) at 600 MHz in CDCl<sub>3</sub>



**Figure S3:** COSY NMR spectrum of (+)-elatol (**1**) at 600 MHz in  $\text{CDCl}_3$

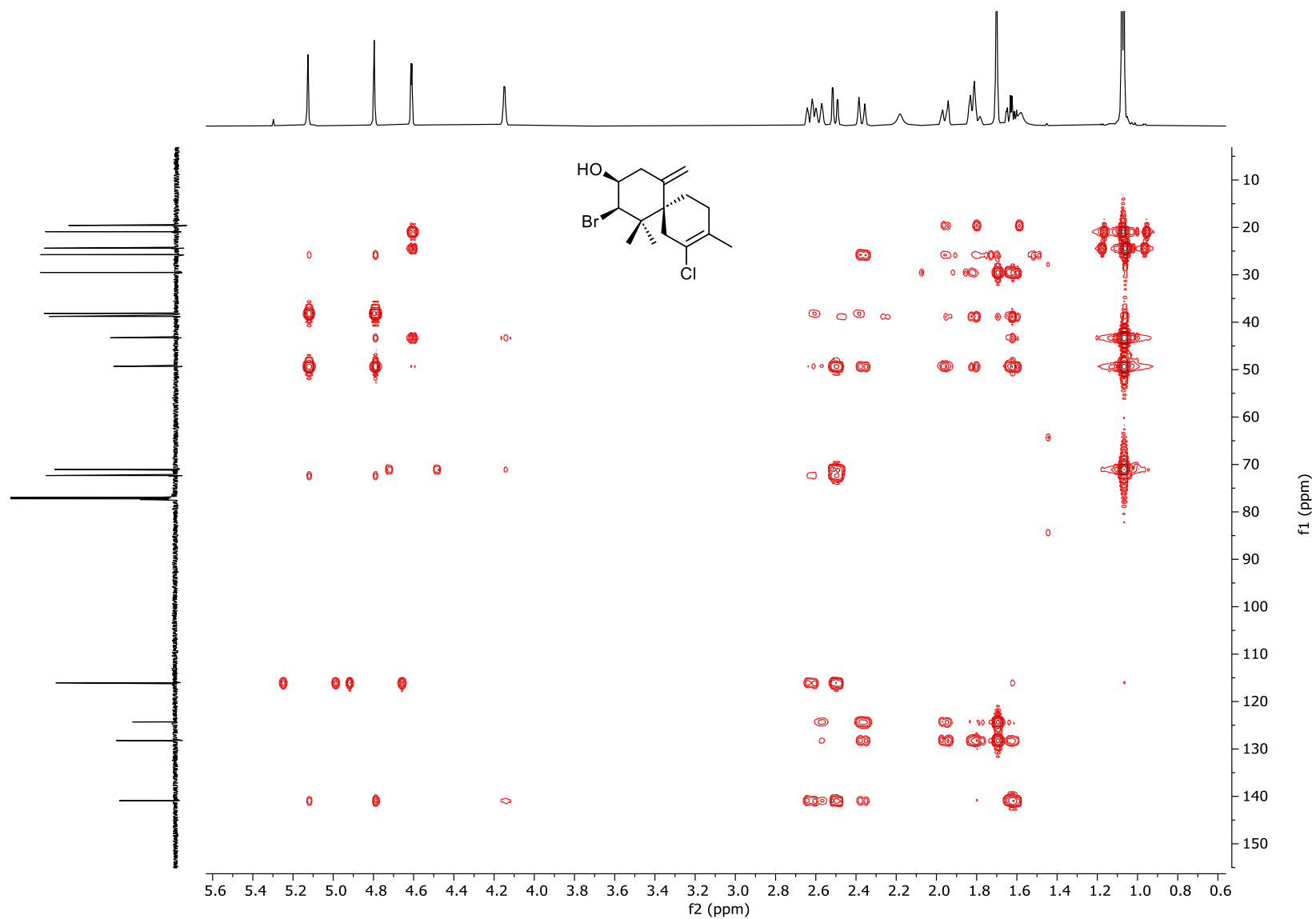
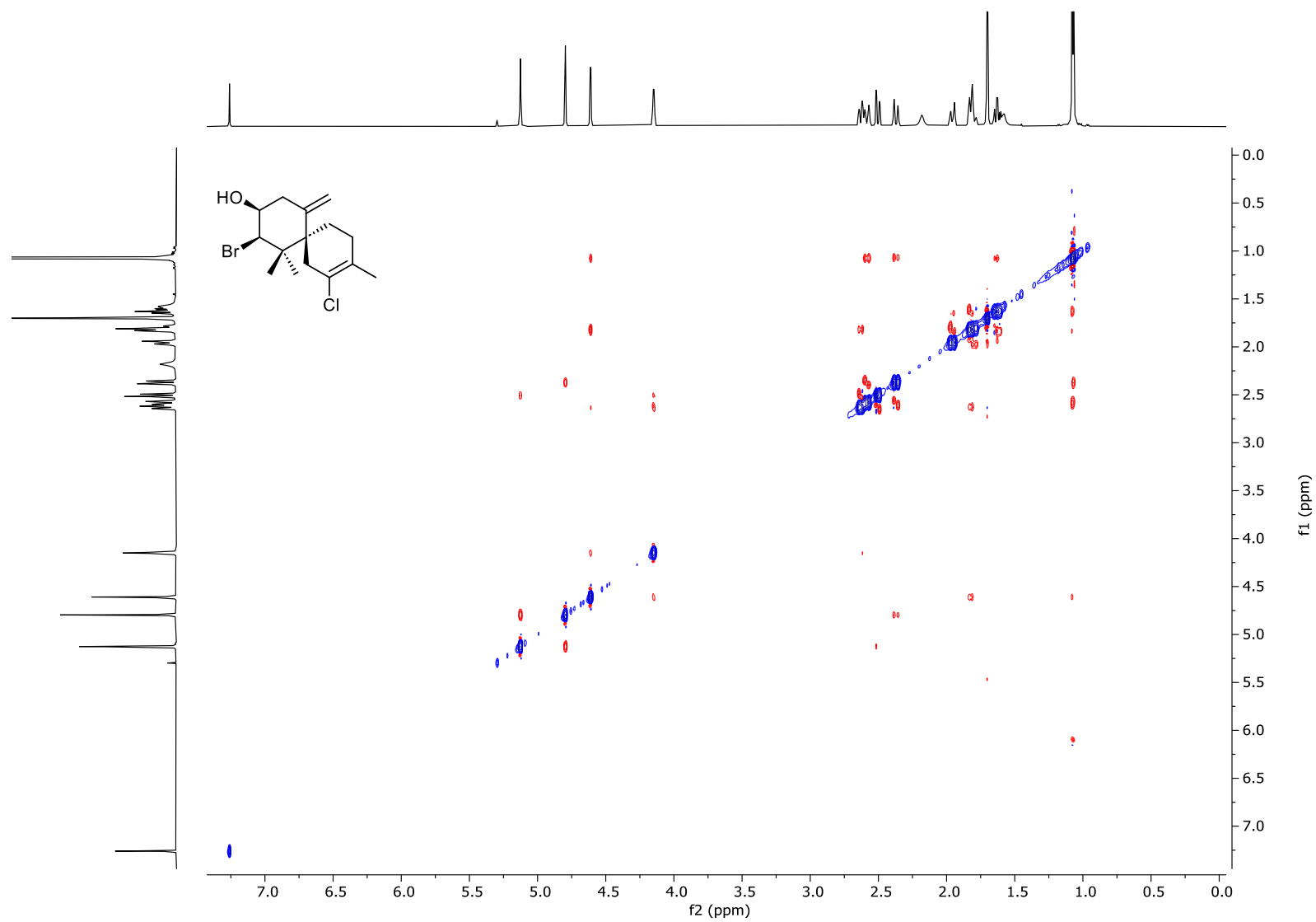
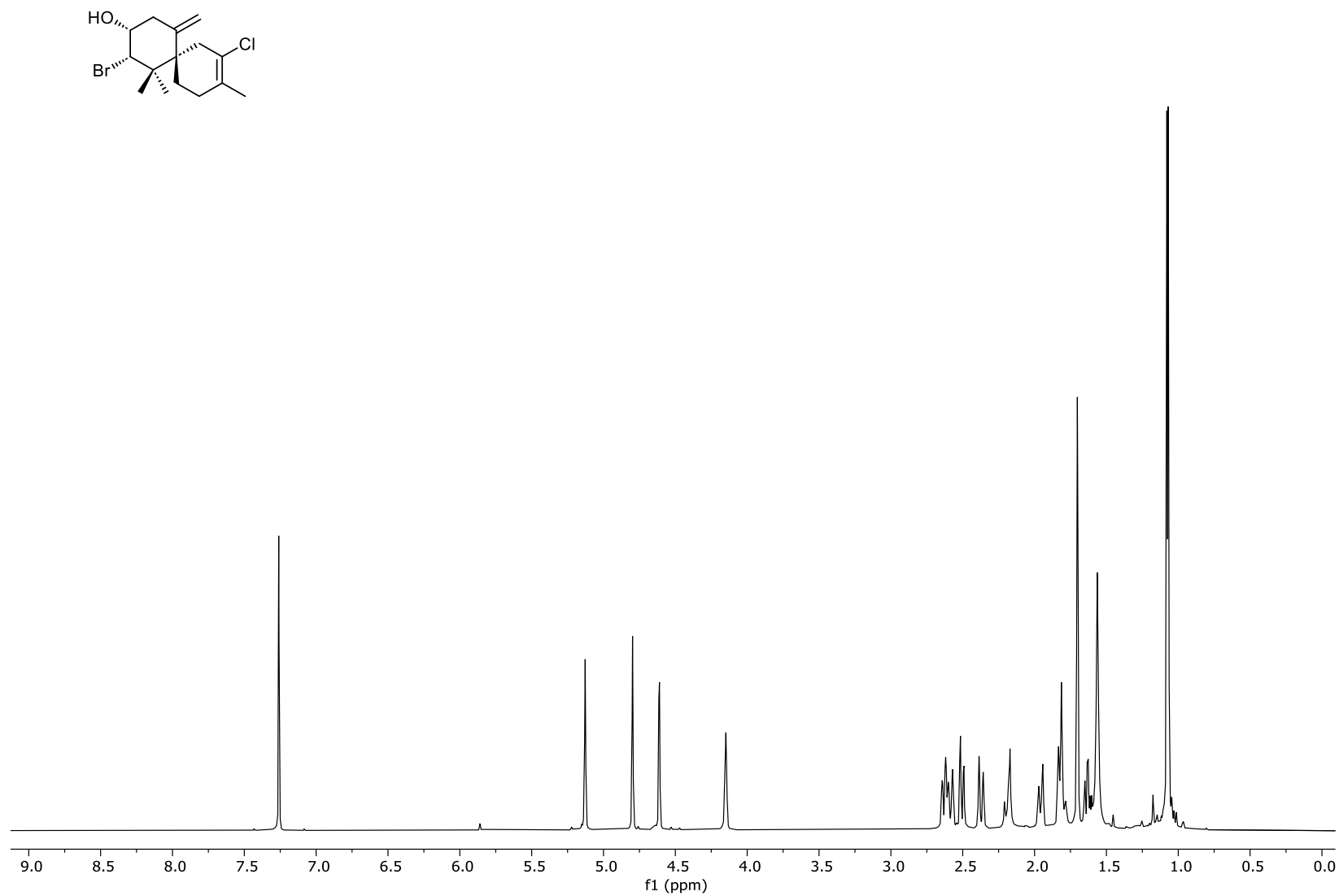


Figure S5: HMBC NMR spectrum of (+)-elatol (**1**) at 600 MHz in  $\text{CDCl}_3$



**Figure S6:** ROESY spectrum of (+)-elatol (**1**) at 600 MHz in CDCl<sub>3</sub>





**Figure S7:**  $^1\text{H}$  NMR spectrum of (-)-elatol (**2**) at 600 MHz in  $\text{CDCl}_3$

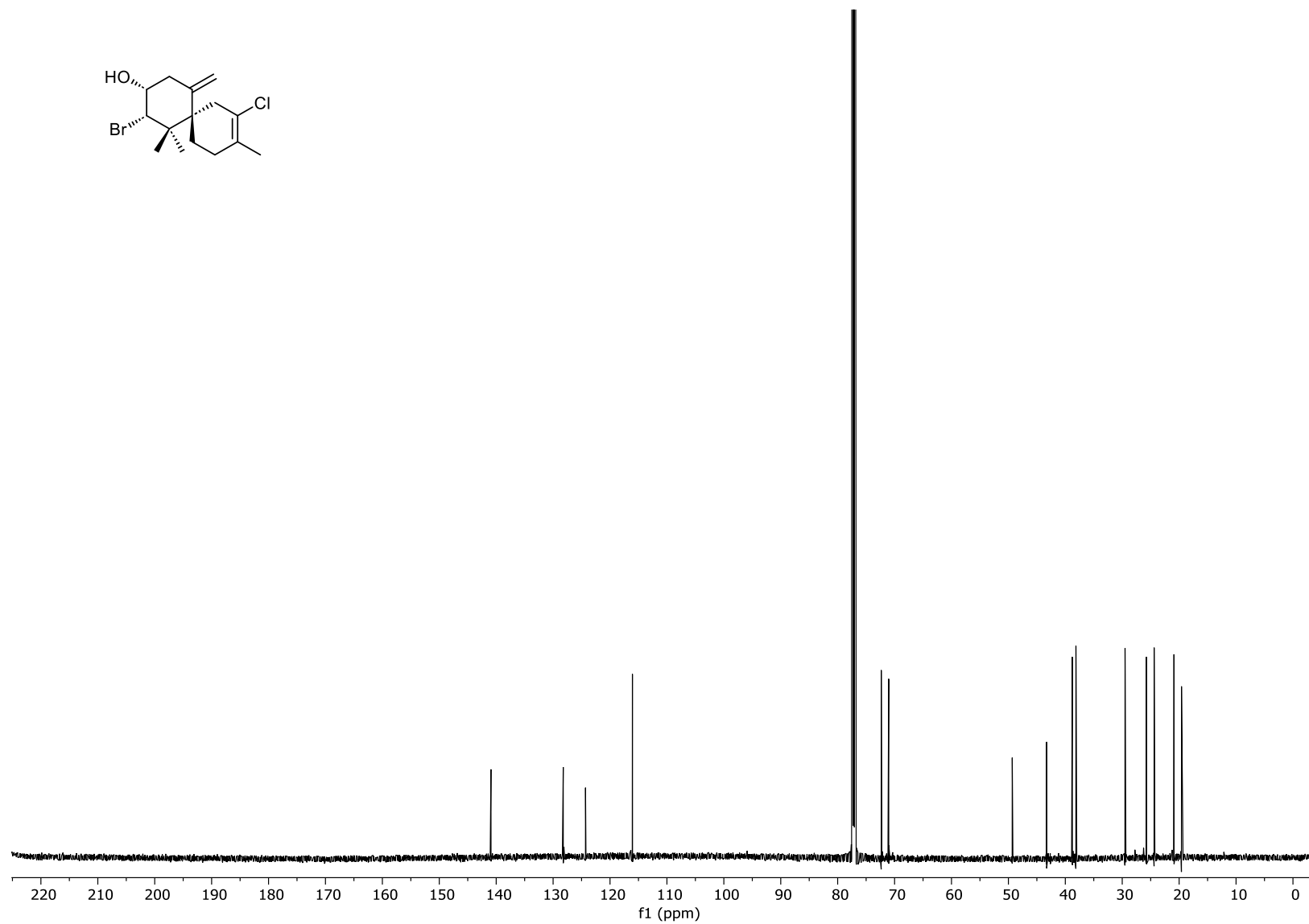
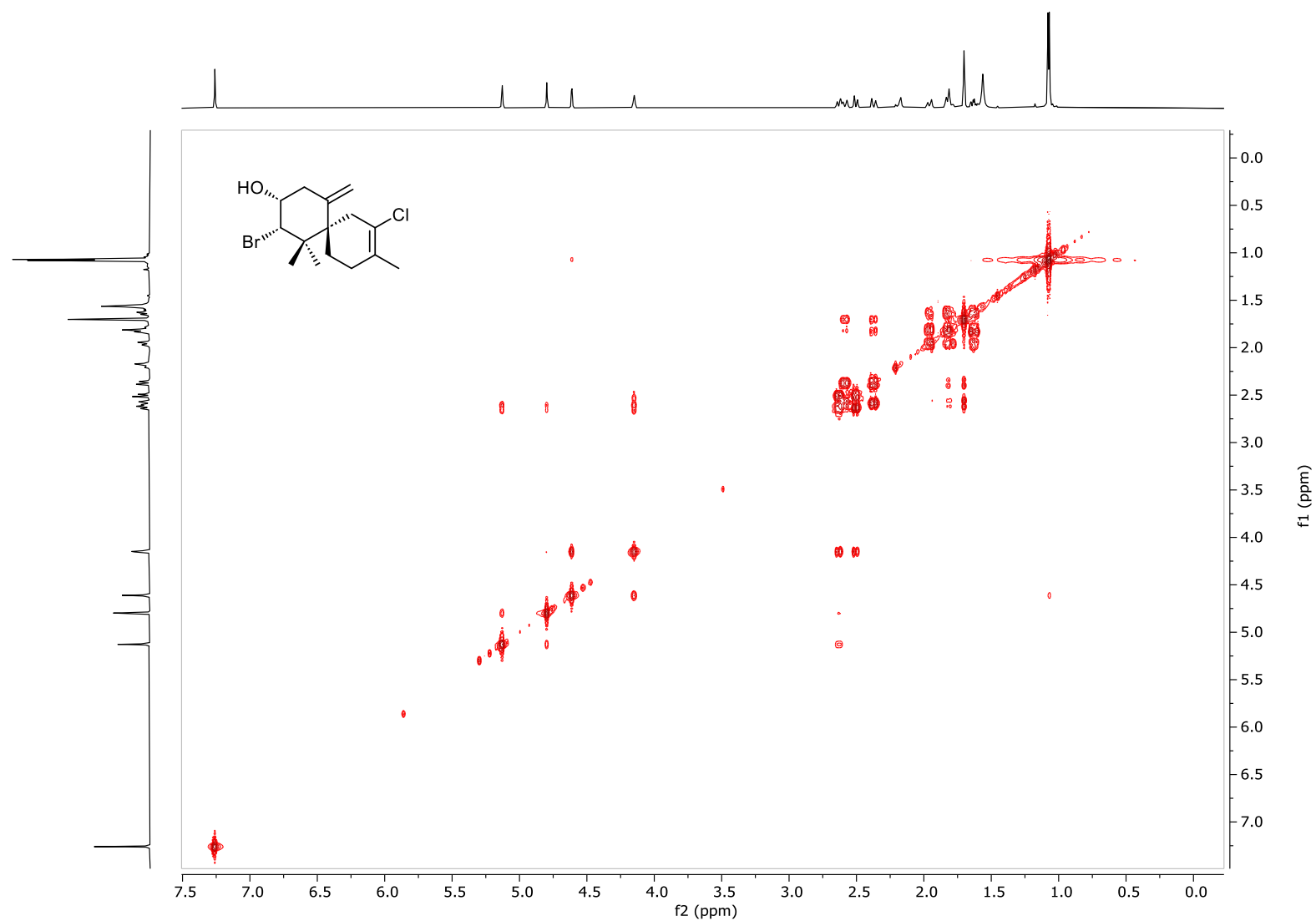
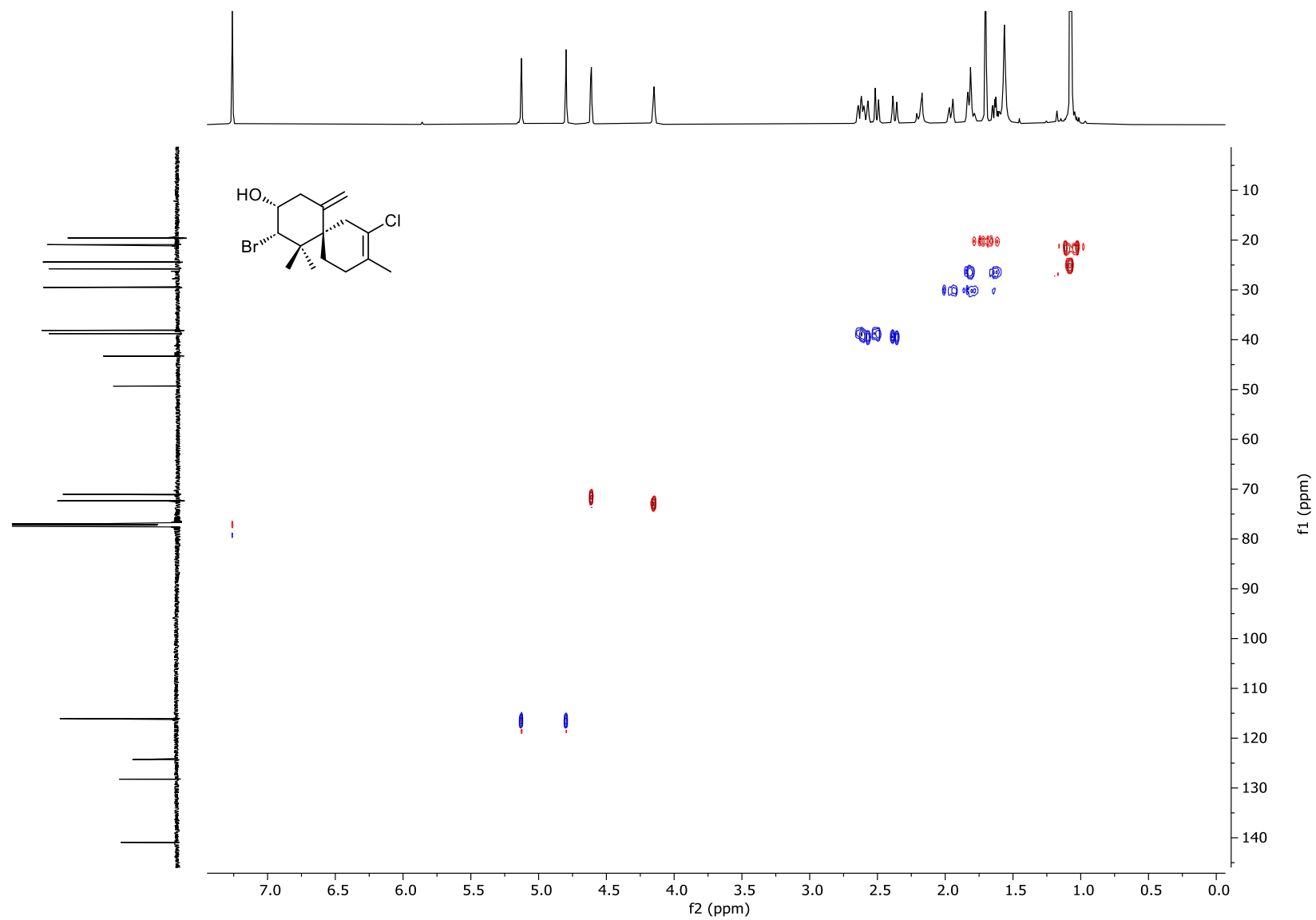


Figure S8:  $^{13}\text{C}$  NMR spectrum of (-)-elatol (2) at 600 MHz in  $\text{CDCl}_3$



**Figure S9:** COSY NMR spectrum of (-)-elatol (**1**) at 600 MHz in CDCl<sub>3</sub>



**Figure S10:** HSQC NMR spectrum of (-)-elatol (**2**) at 600 MHz in CDCl<sub>3</sub>

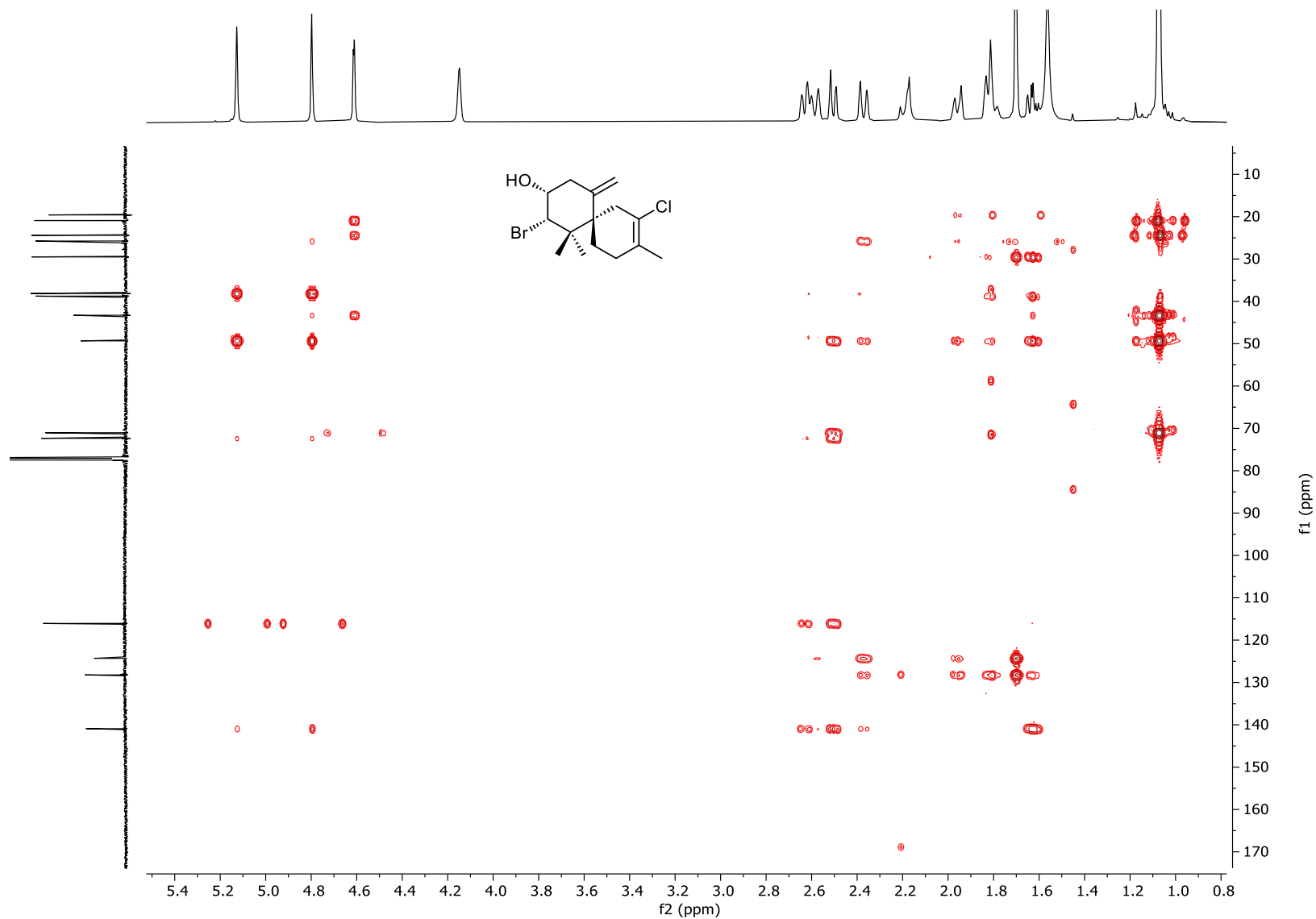


Figure S11: HMBC NMR spectrum of (-)-elatol (2) at 600 MHz in CDCl<sub>3</sub>

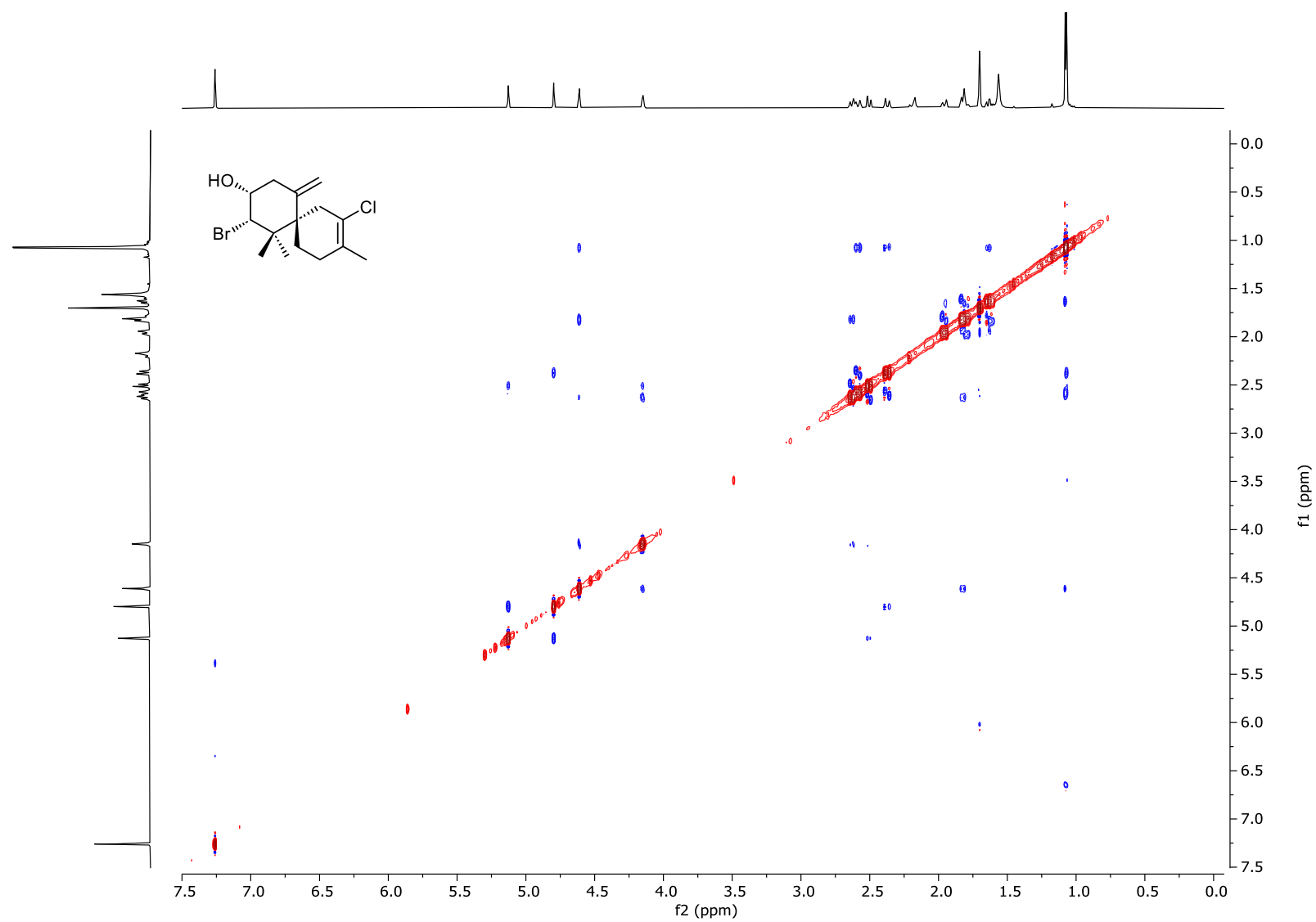
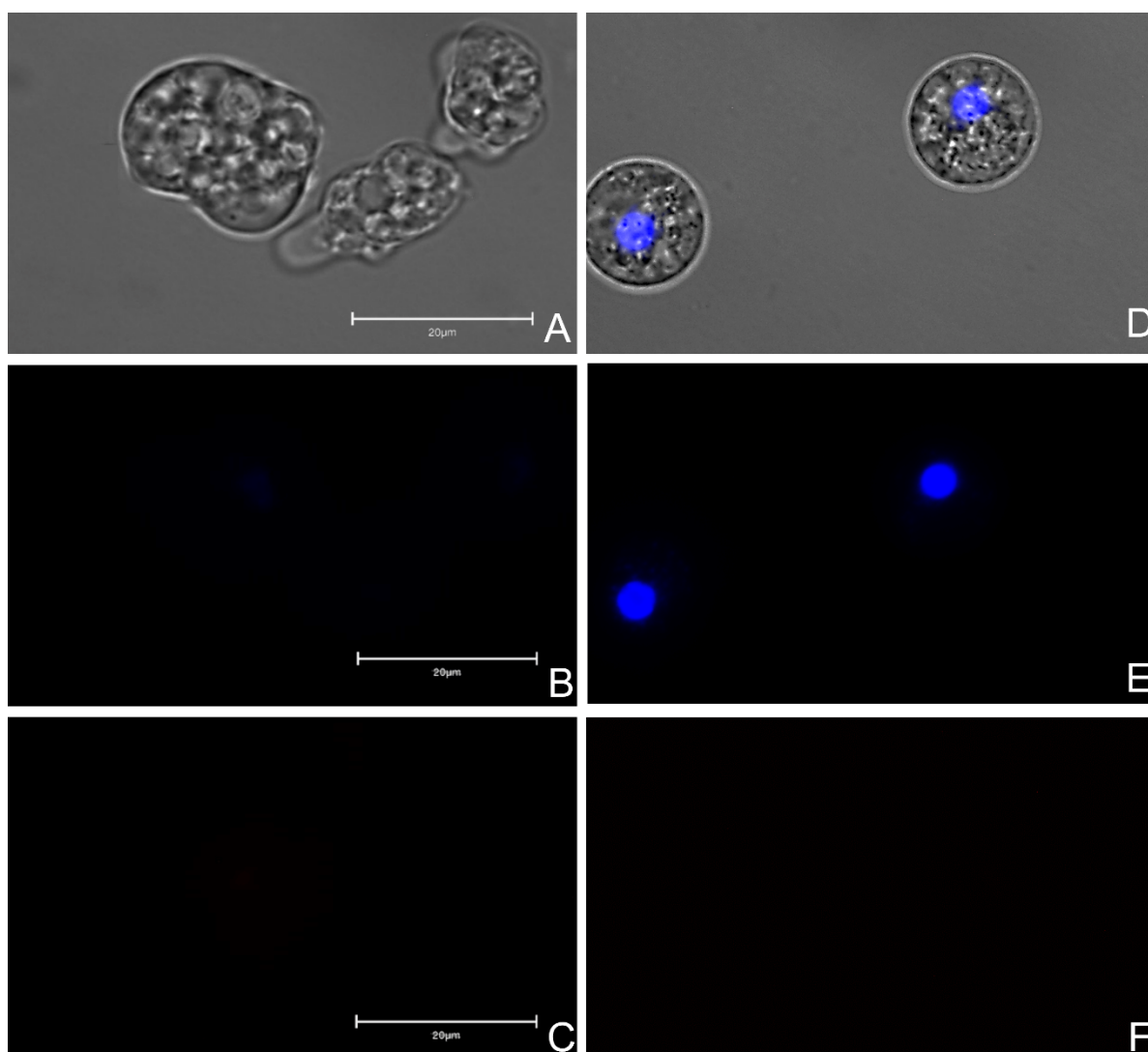
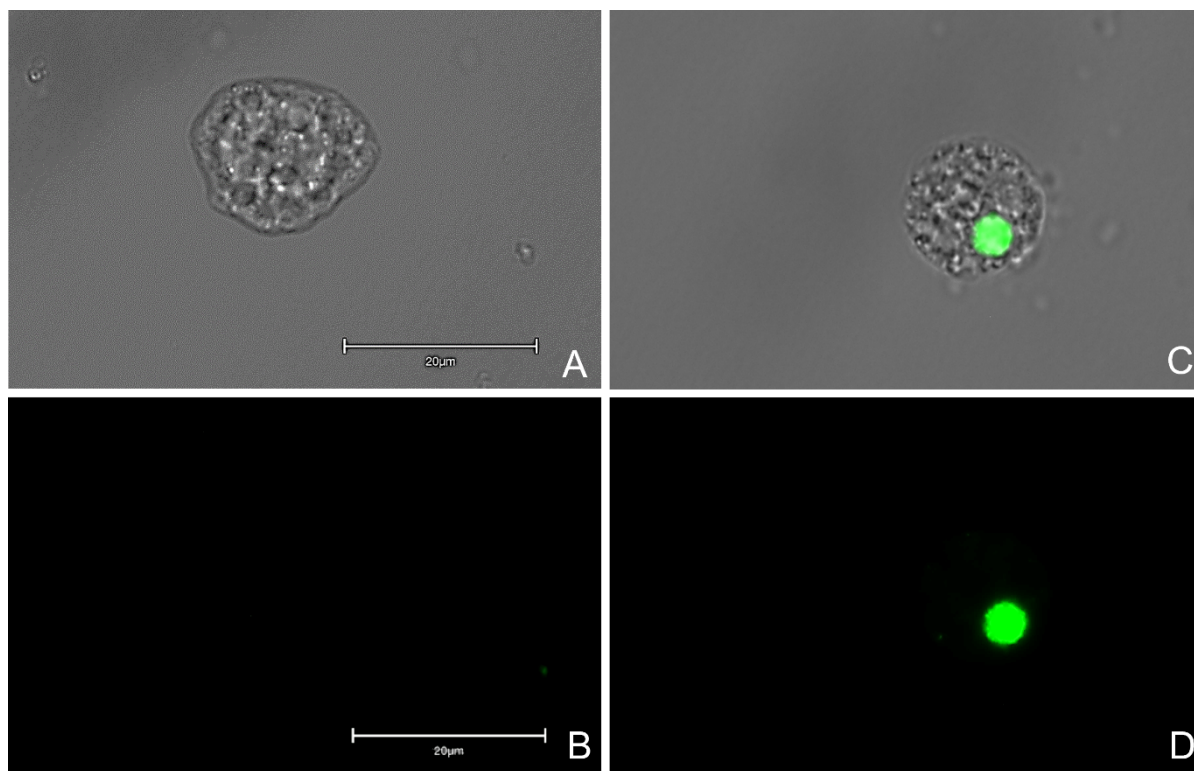


Figure S12: ROESY spectrum of (-)-elatol (**2**) at 600 MHz in CDCl<sub>3</sub>

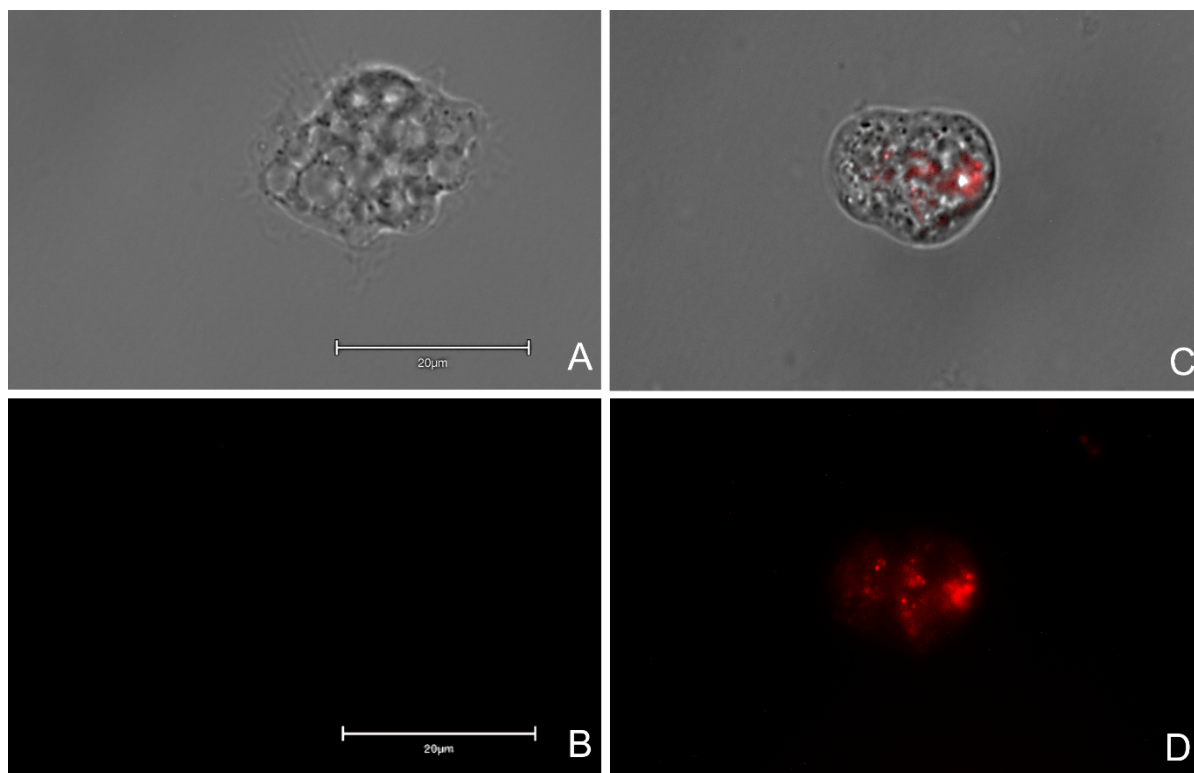


**Figure S13.** Higher magnification of Figure 2 (x100). (D-F) images show early apoptotic cells with intense blue fluorescence and no red fluorescence. Overlay channel (A and D); Hoechst channel (B and E), and propidium iodide channel (C and F). Images are representative of the cell population observed in the performed experiments. Images were obtained using an EVOS M500 Cell Imaging System, Life Technologies, Spain. (Scale bar: 20 μm)

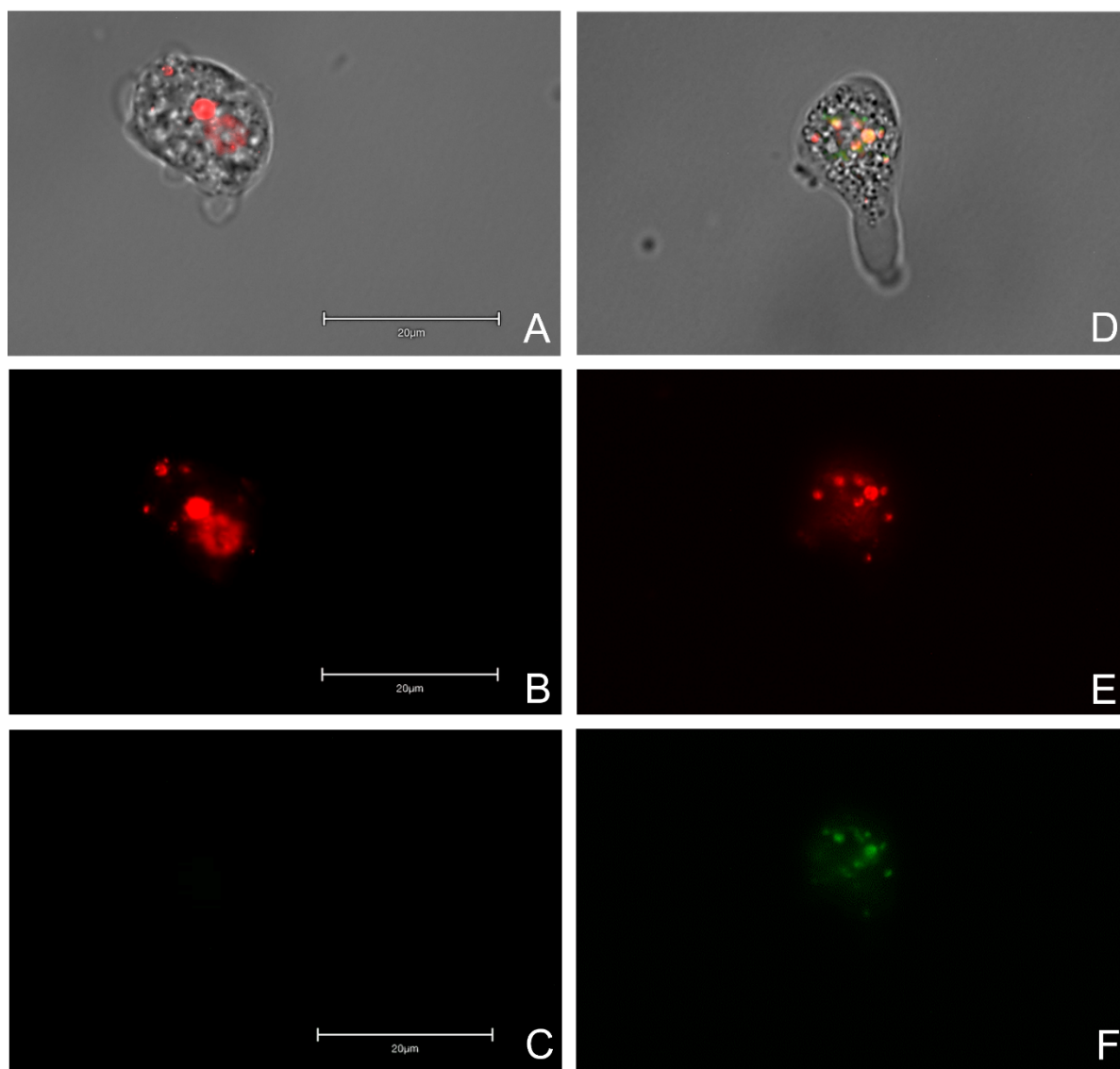


**Figure S14.** Higher magnification of Figure 3 (x100). Cells treated with the IC<sub>90</sub> of (+)- Elatol showing green fluorescence(**C** and **D**), negative control (**A** and **B**). Overlay channel (**A** and **C**) and Sytox green channel (**B** and **D**). Images are representative of the cell population observed in the performed experiments. Images were obtained using an EVOS M500 Cell Imaging System, Life Technologies, Spain.





**Figure S15.** Higher magnification of Figure 4 (x100). Cells treated with the IC<sub>90</sub> of (+)- Elatol showing red fluorescence (**C** and **D**), negative control (**A** and **B**). Overlay channel (**A** and **C**) and CellROX® channel (**B** and **D**). Images are representative of the cell population observed in the performed experiments. Images were obtained using an EVOS M500 Cell Imaging System, Life Technologies, Spain.



**Figure S16.** Higher magnification of Figure 5 (x100). Cells treated with the IC<sub>90</sub> of (+)- Elatol showing red fluorescence (**C** and **D**), negative control (**A** and **B**). Overlay channel (**A** and **C**) and CellROX® channel (**B** and **D**). Images are representative of the cell population observed in the performed experiments. Images were obtained using an EVOS M500 Cell Imaging System, Life Technologies, Spain.