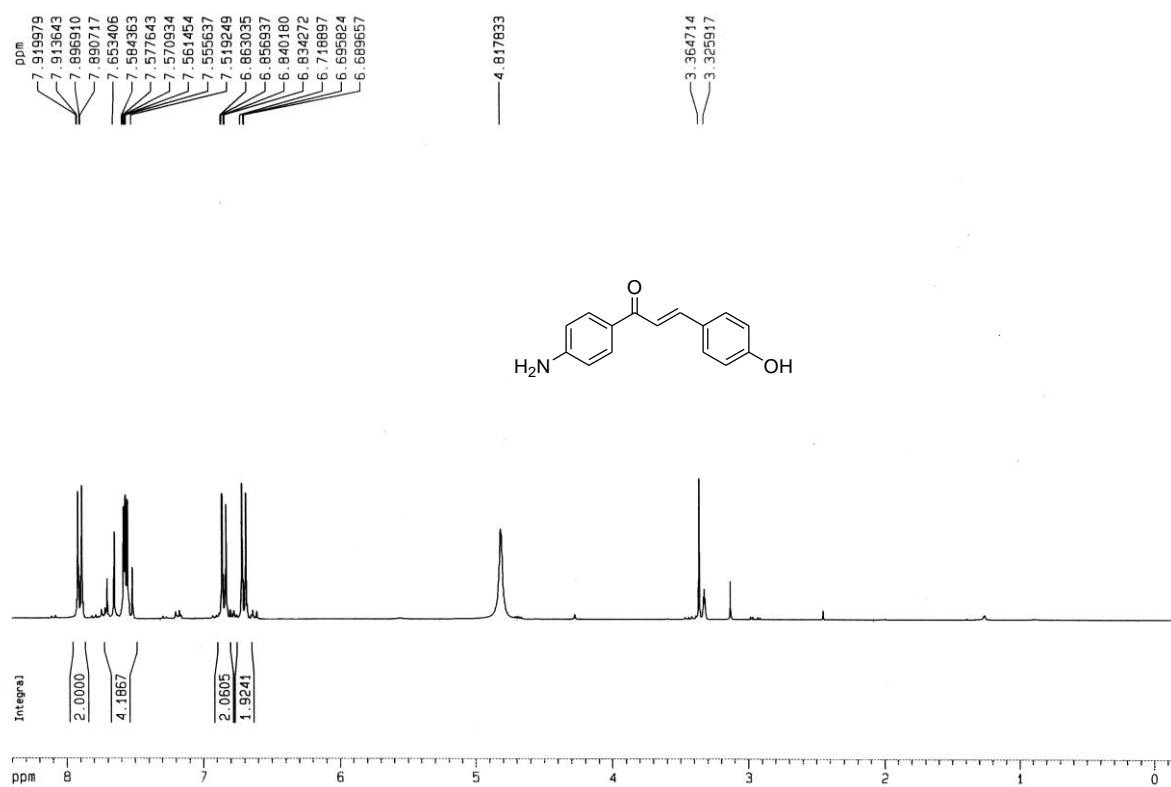
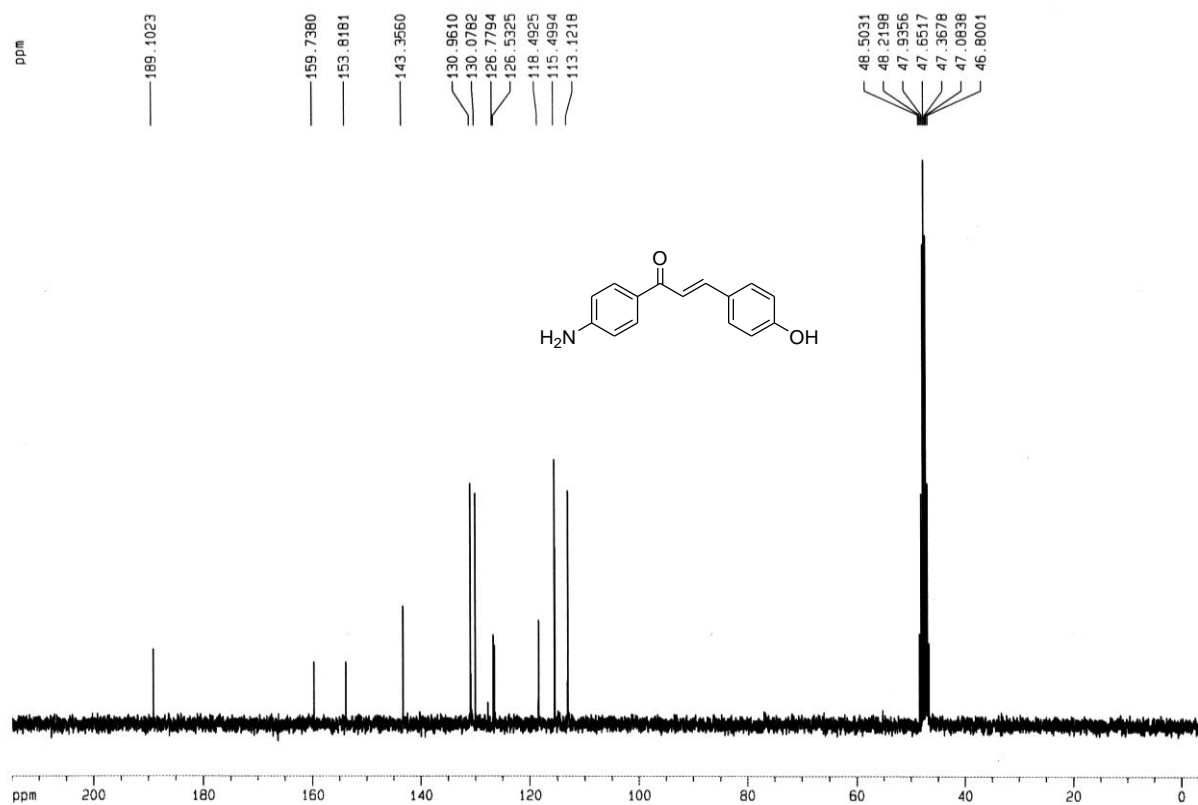
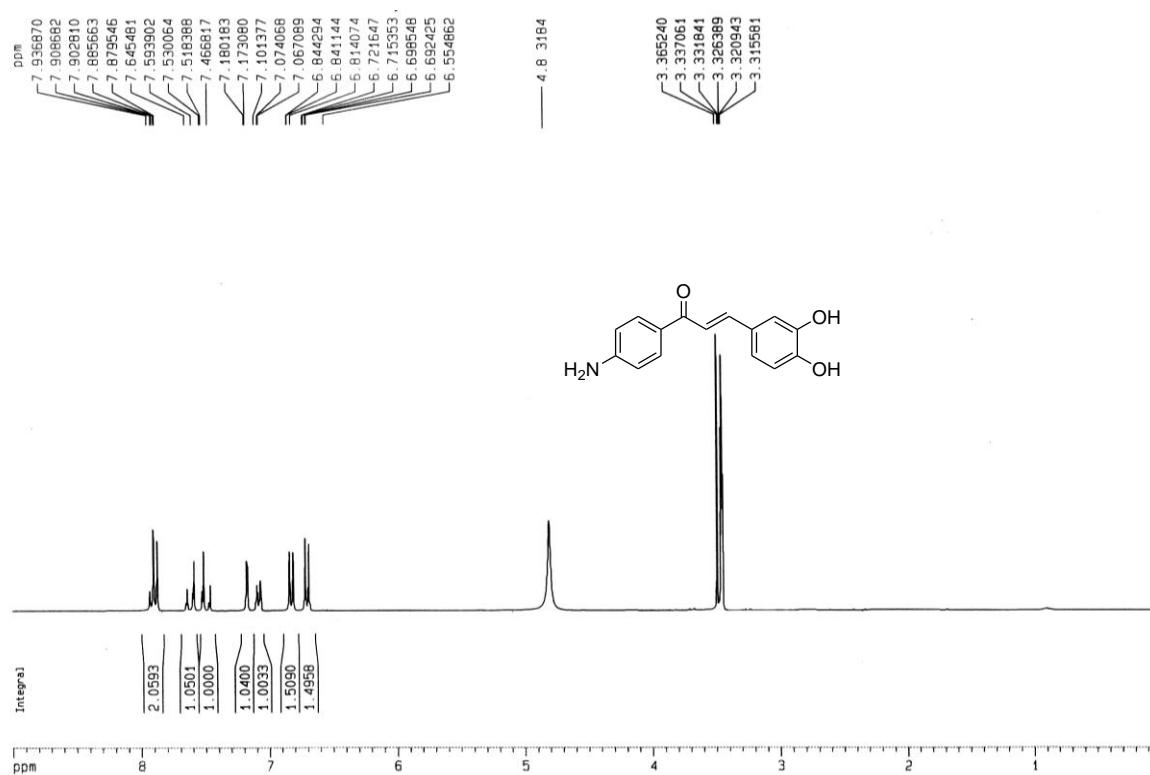
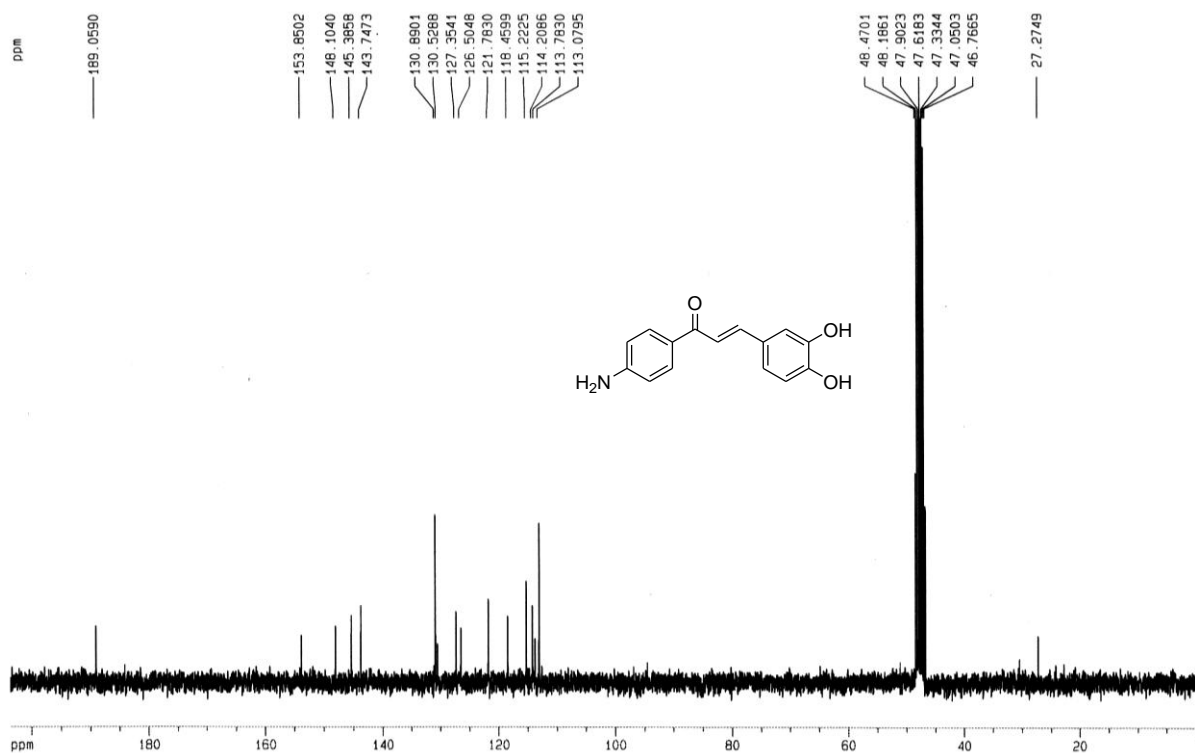


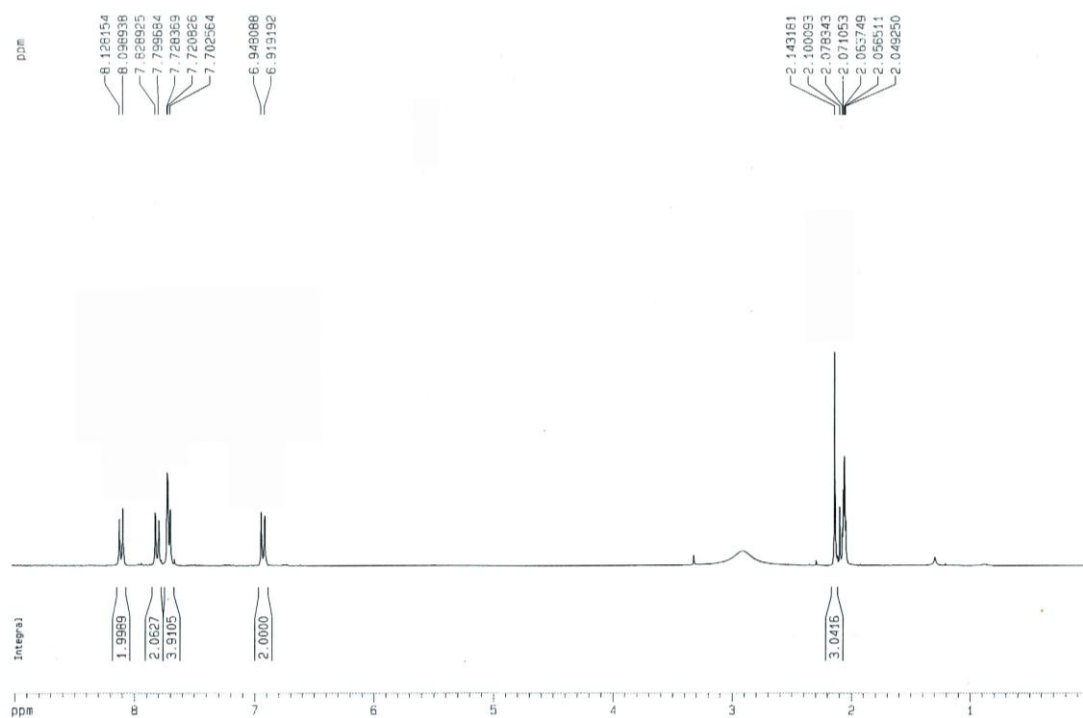
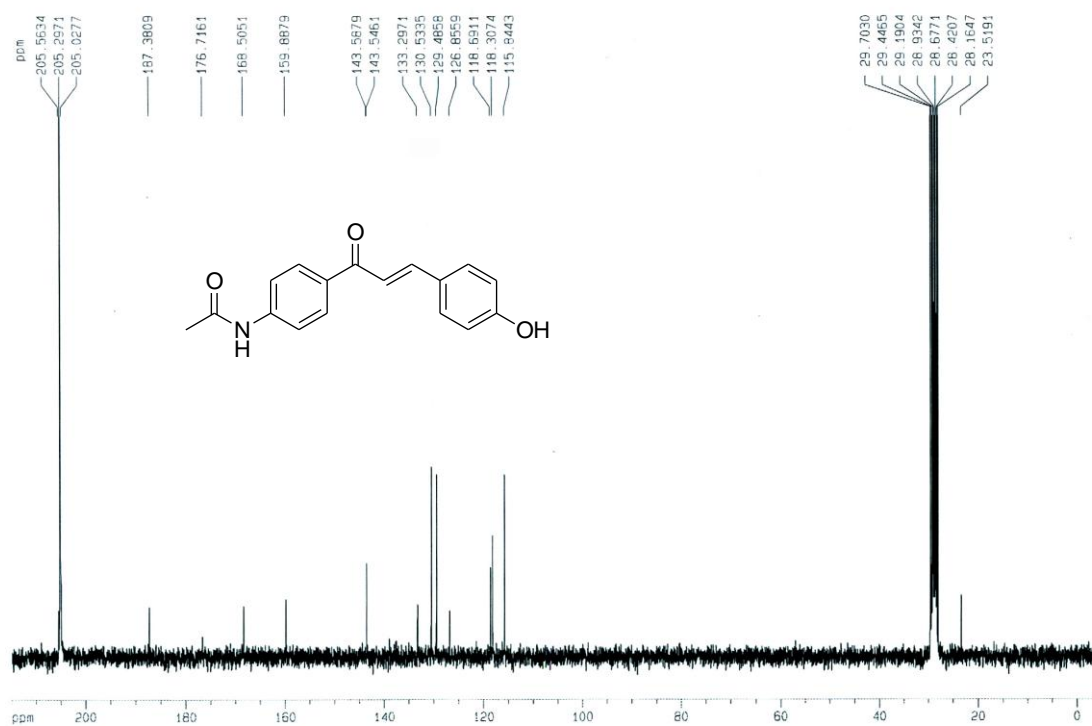
Supplementary Materials

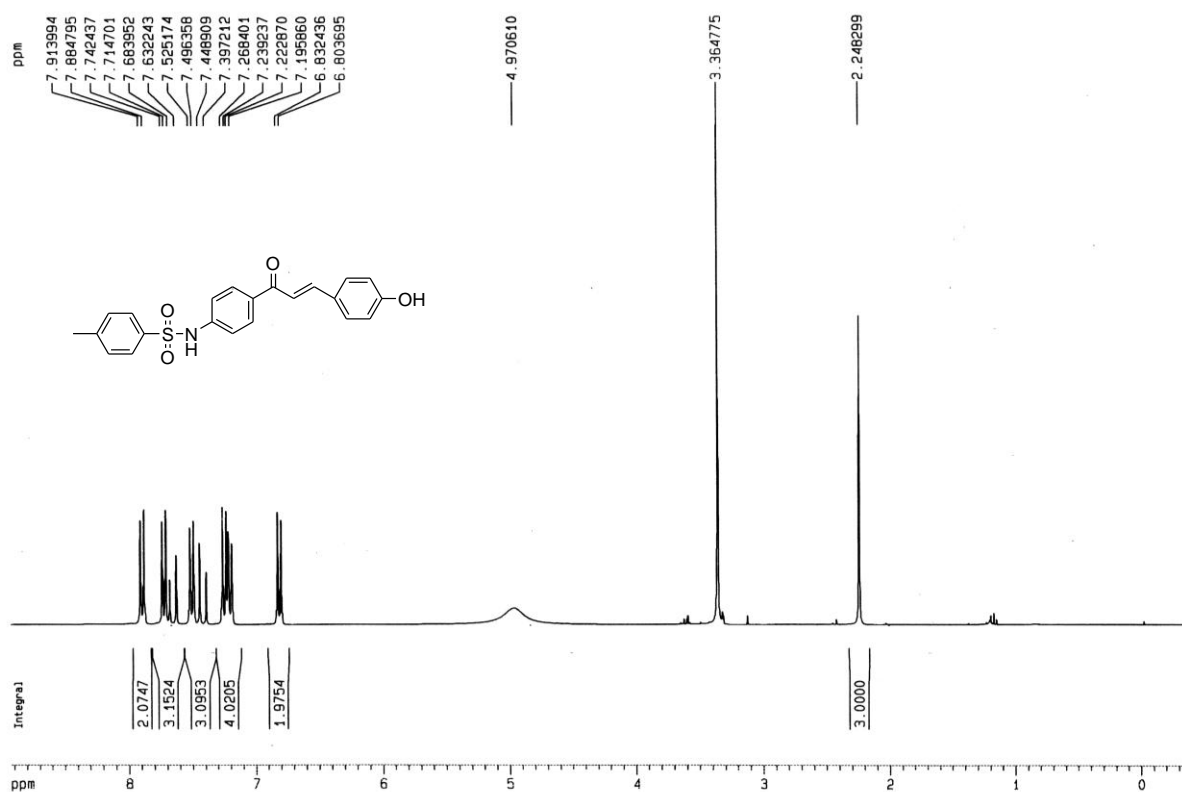
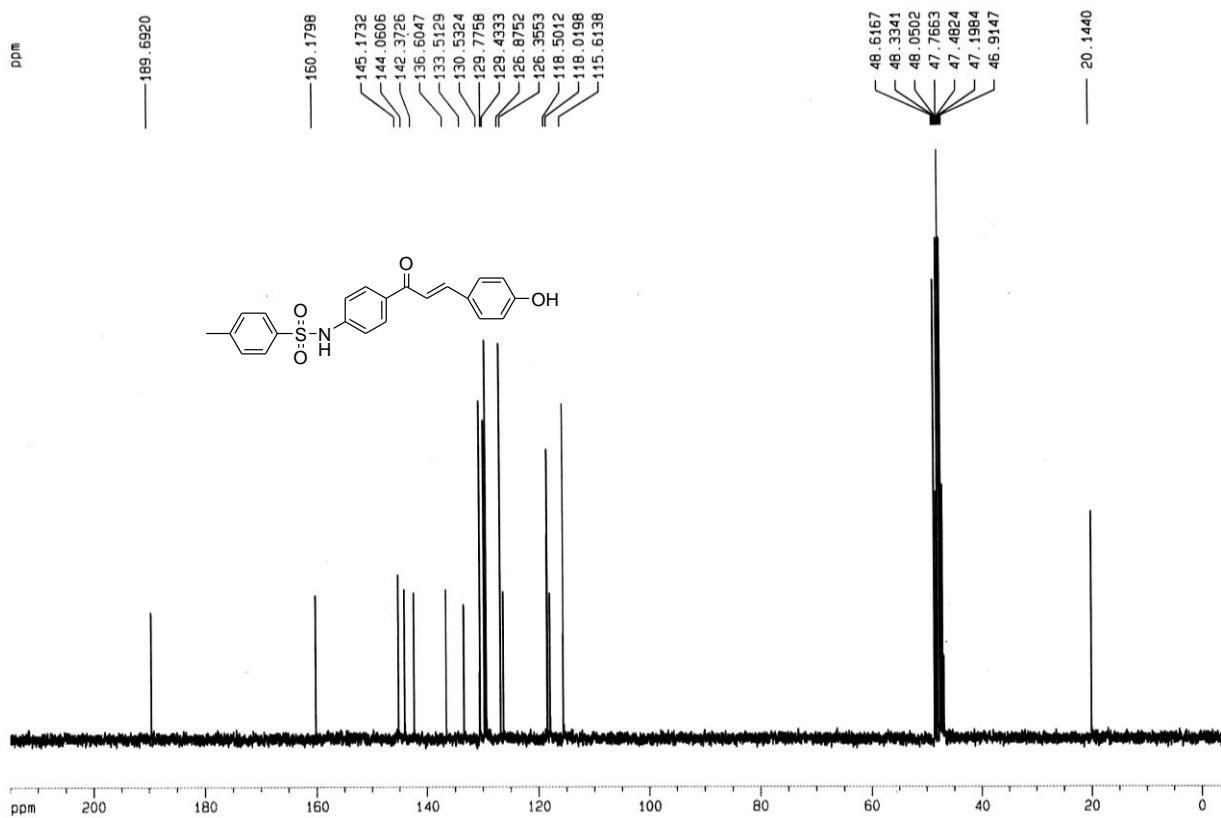
Experimental Section & Characterization Data

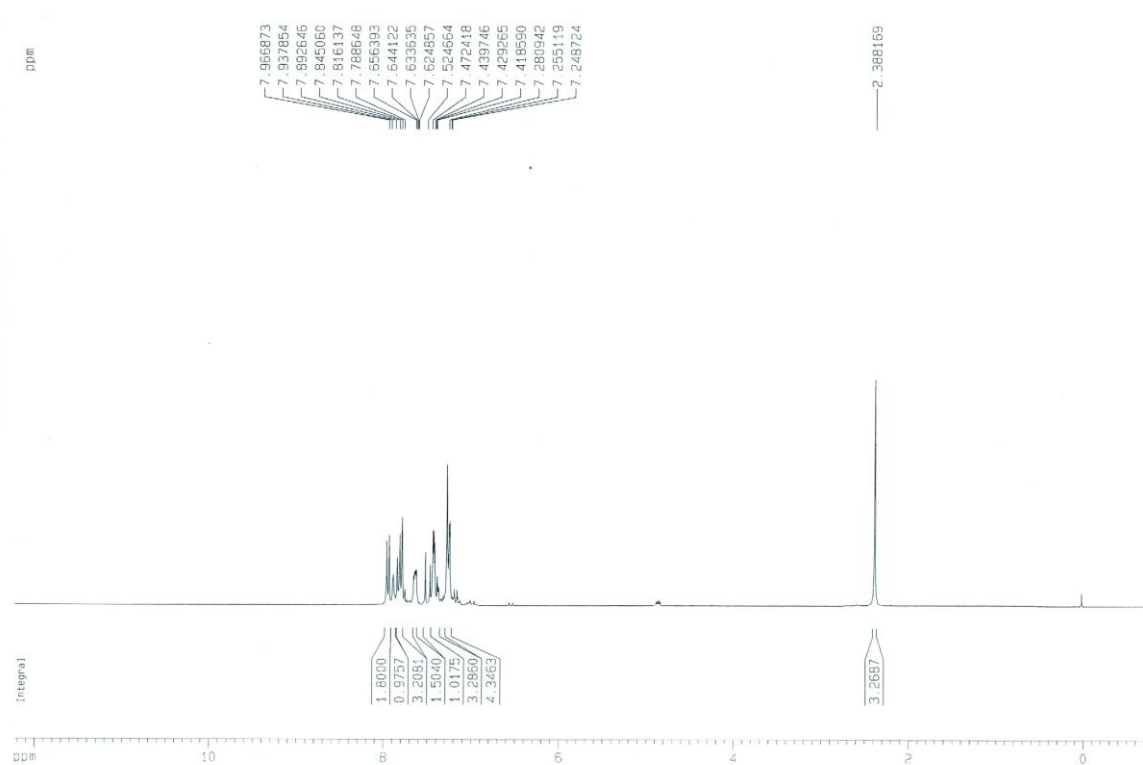
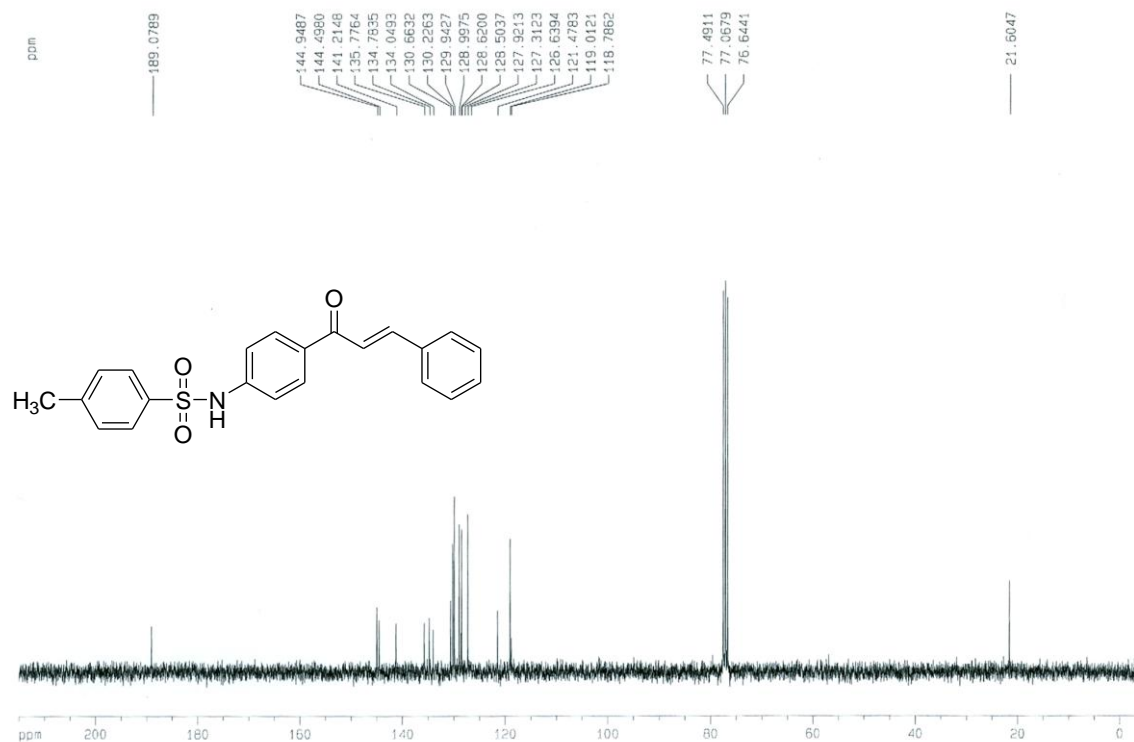
- ▶ Spectroscopic data of the compounds (1–7) -----S2–S18
- ▶ BACE1 kinetic data of the compounds (1–5) -----S19–S22
- ▶ Cholinesterase kinetic data of the compounds (4a,f, 5b,d) -----S23-S24

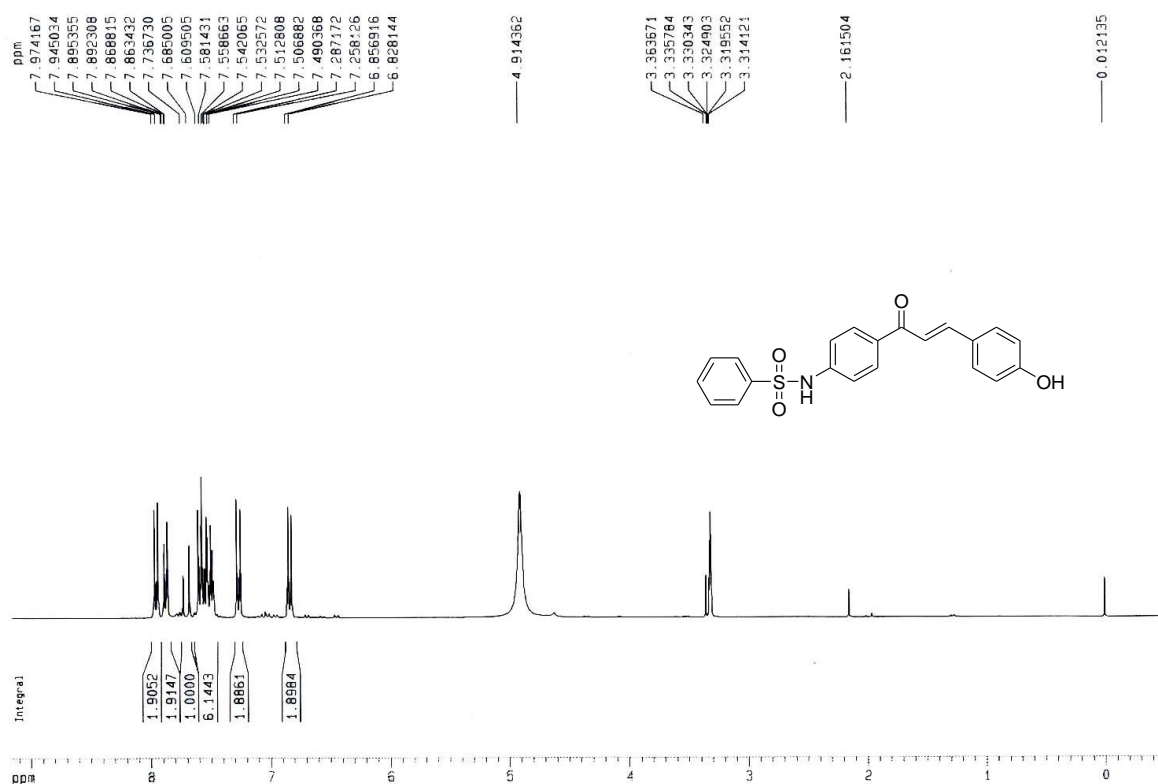
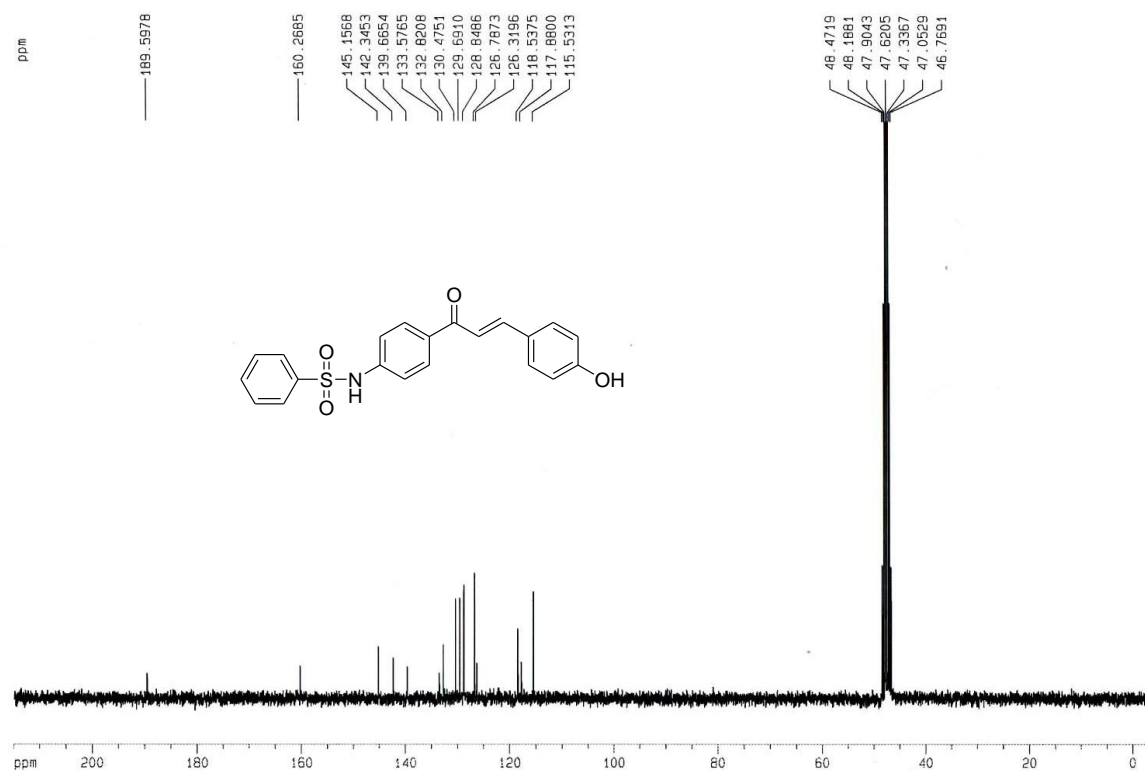
Spectra 1. $^1\text{H-NMR}$ of 4'-amino-4-hydroxychalcone (1).**Spectra 2.** $^{13}\text{C-NMR}$ of 4'-amino-4-hydroxychalcone (1).

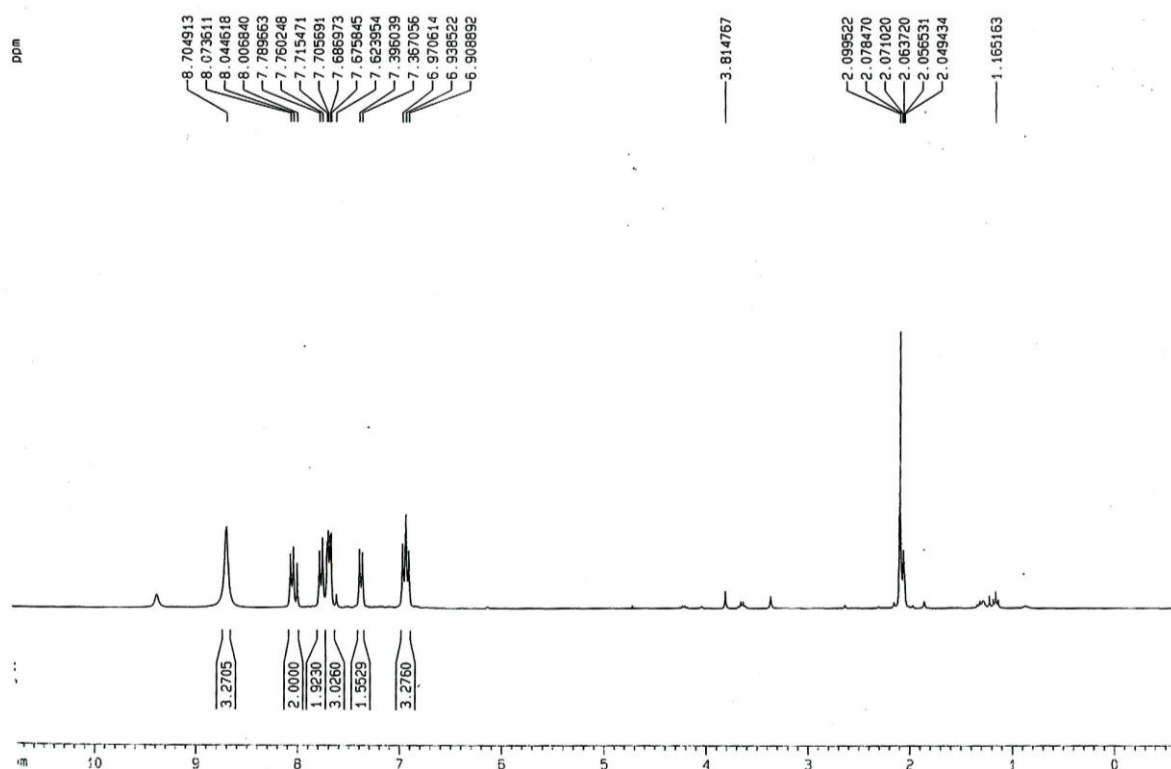
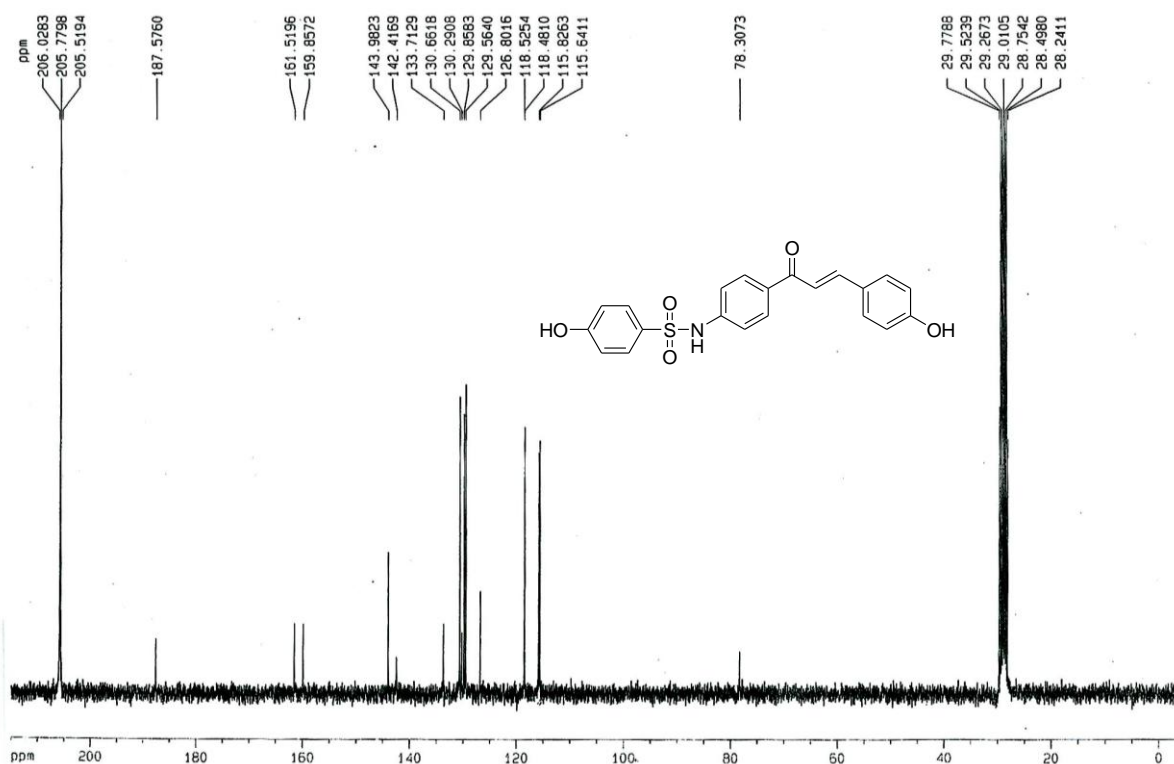
Spectra 3. $^1\text{H-NMR}$ of 4'-amino-3,4-dihydroxychalcone (2).**Spectra 4.** $^{13}\text{C-NMR}$ of 4'-amino-3,4-dihydroxychalcone (2).

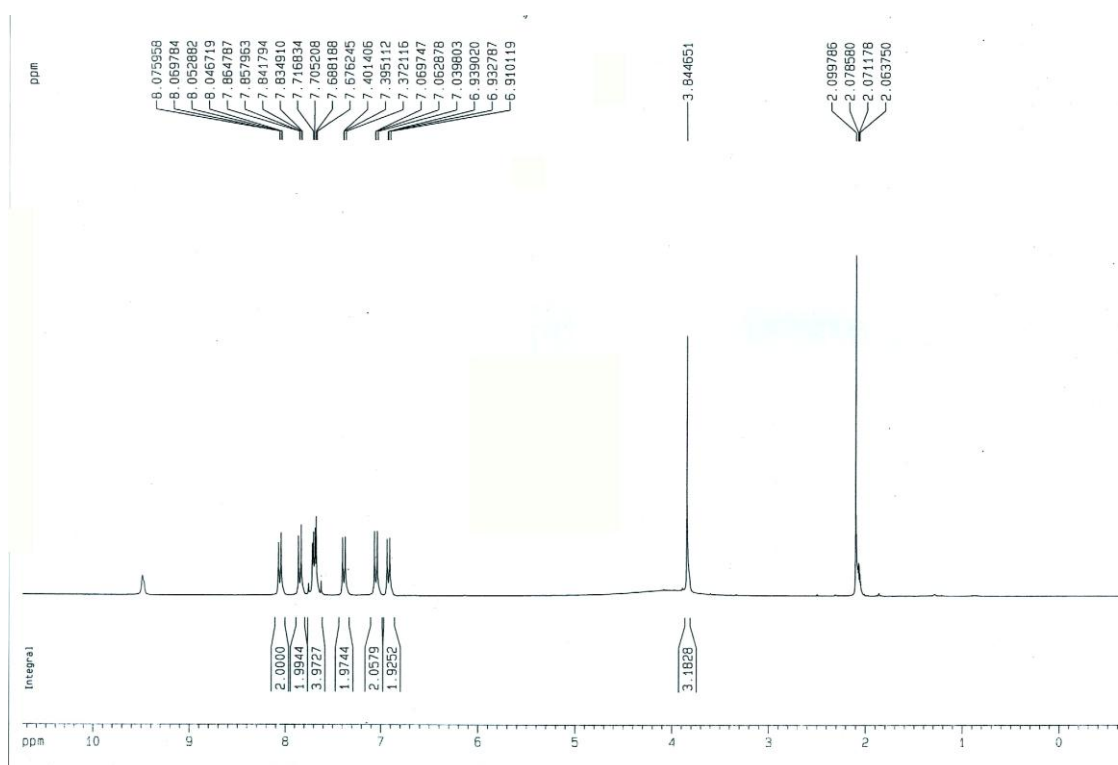
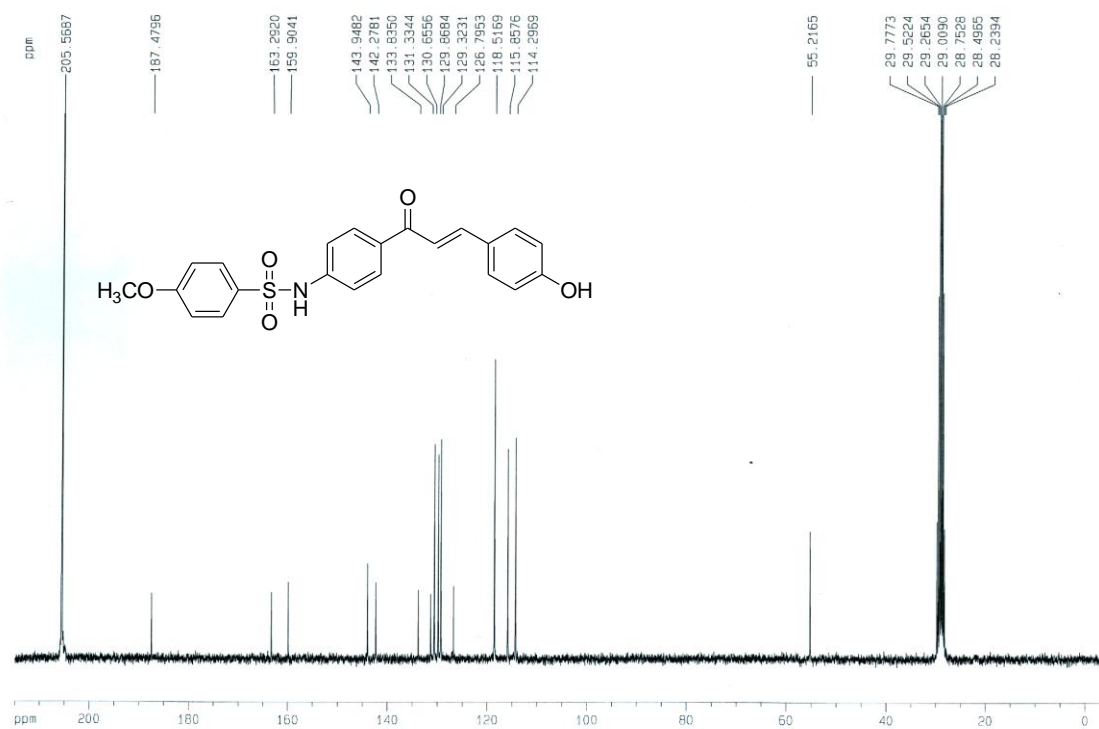
Spectra 5. $^1\text{H-NMR}$ of 4'-acetamido-4-hydroxychalcone (**3**).**Spectra 6.** $^{13}\text{C-NMR}$ of 4'-acetamido-4-hydroxychalcone (**3**).

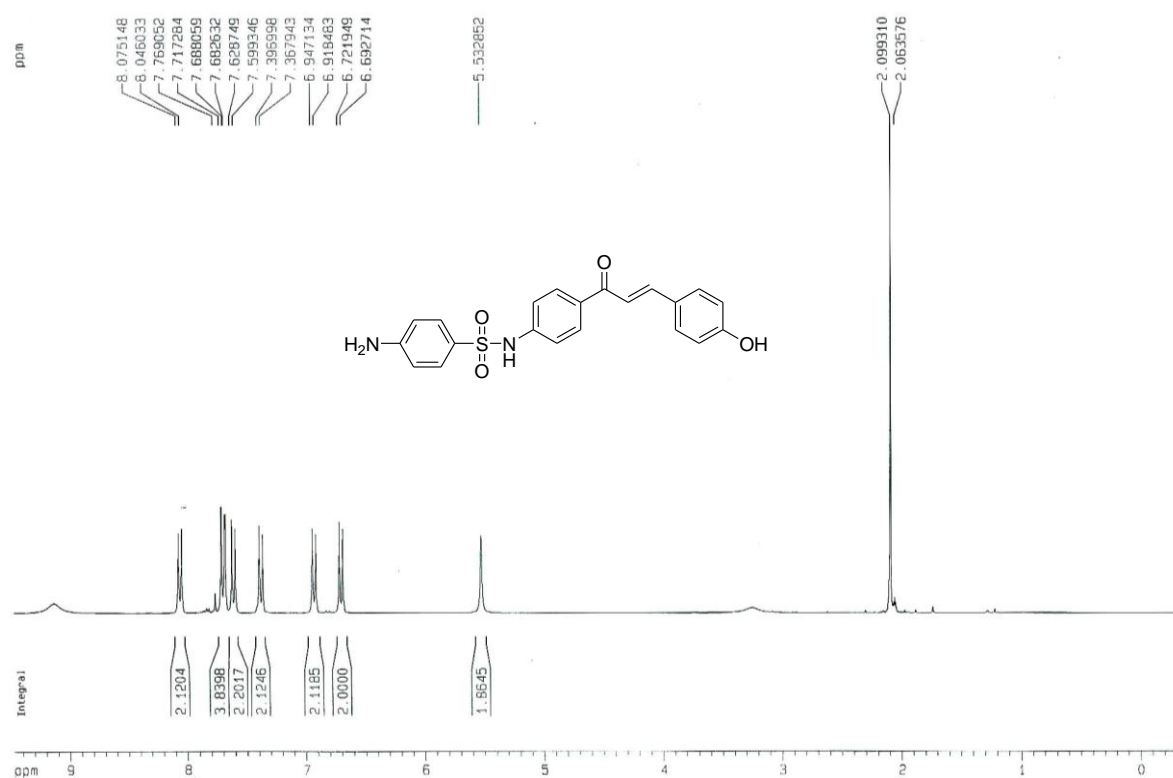
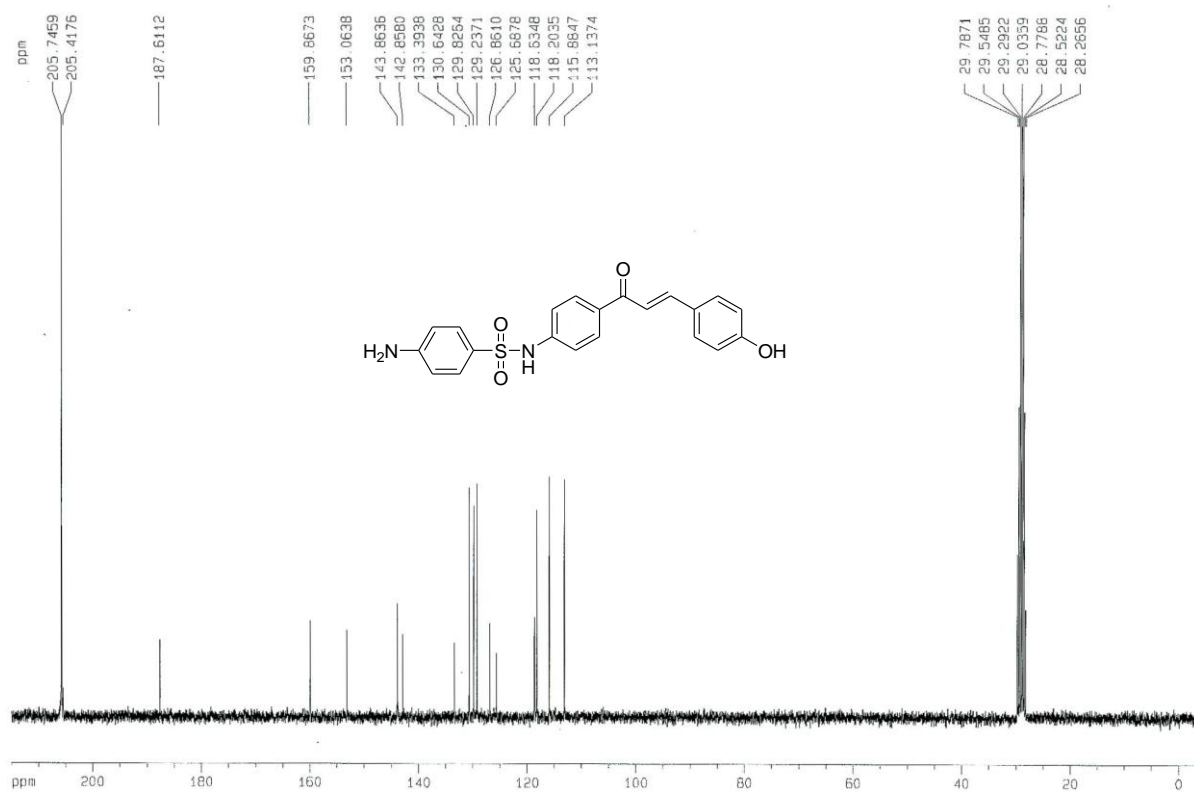
Spectra 7. $^1\text{H-NMR}$ of 4'-(*p*-toluenesulfonamide)-4-hydroxychalcone (**4a**).**Spectra 8.** $^{13}\text{C-NMR}$ of 4'-(*p*-toluenesulfonamide)-4-hydroxychalcone (**4a**).

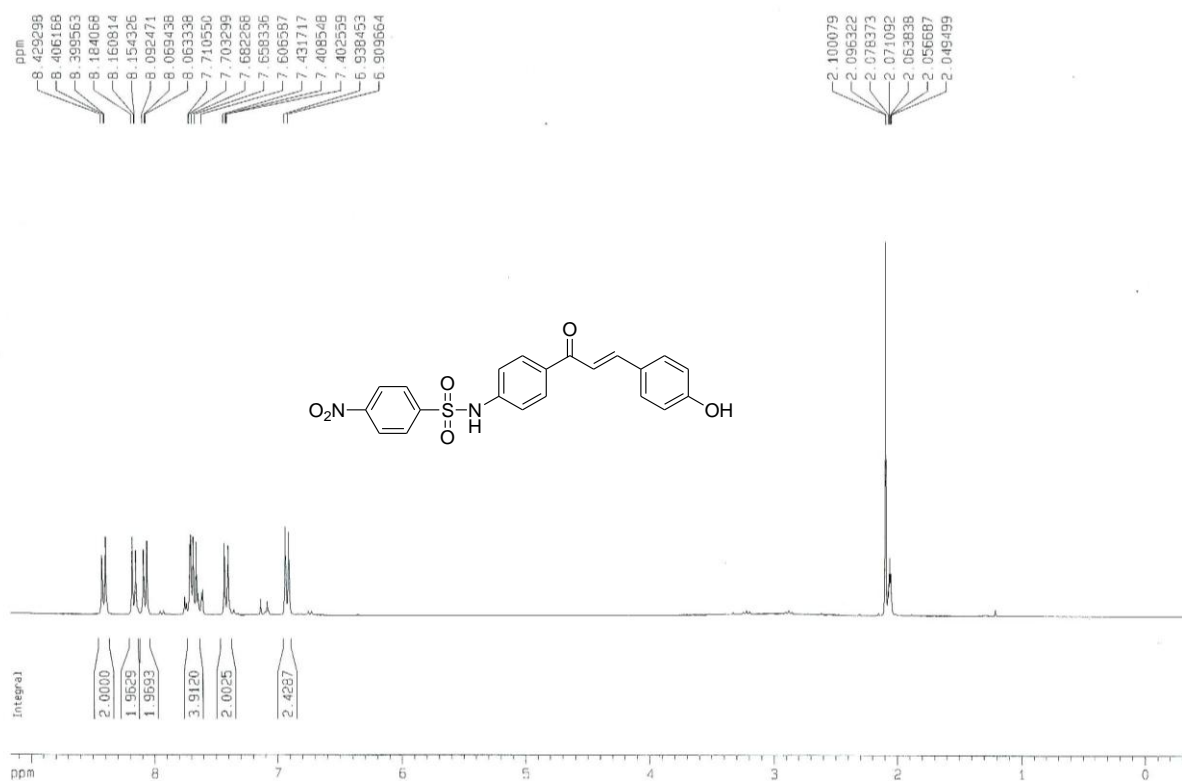
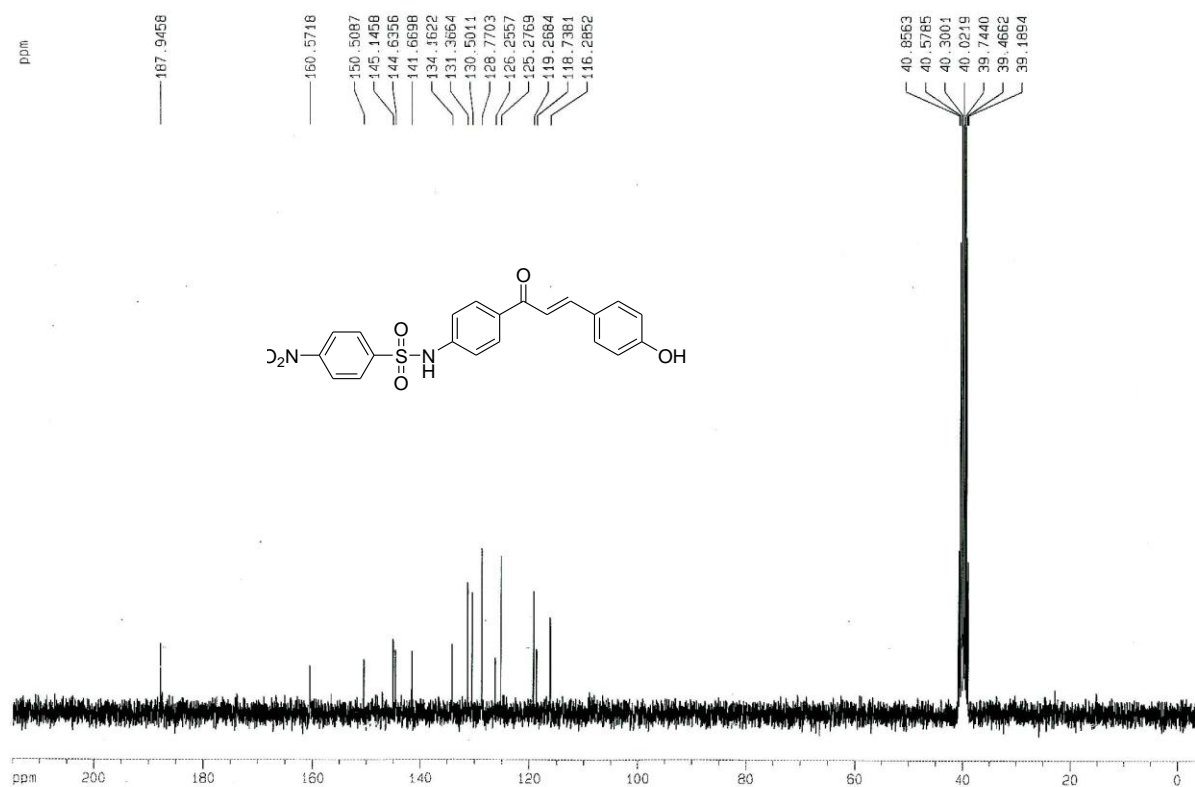
Spectra 9. $^1\text{H-NMR}$ of 4'-(4-toluenesulfonamide)-chalcone (**4b**).**Spectra 10.** $^{13}\text{C-NMR}$ of 4'-(4-toluenesulfonamide)-chalcone (**4b**).

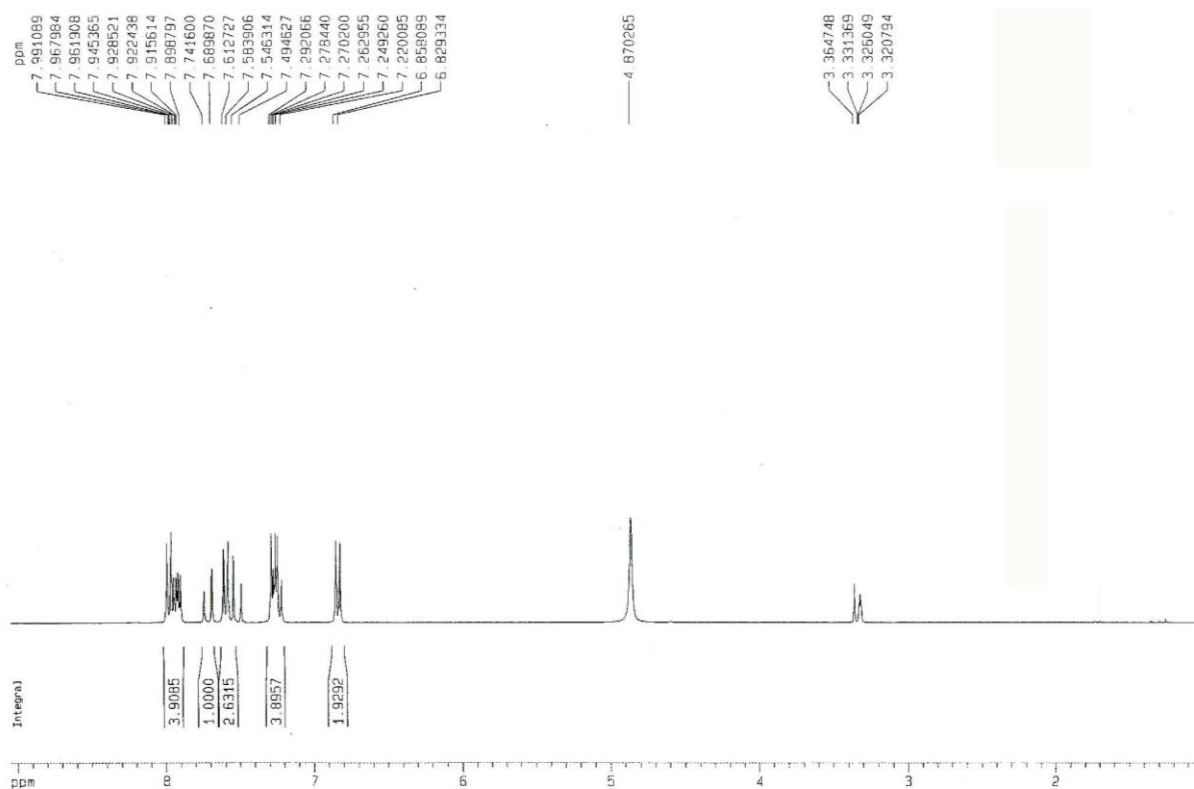
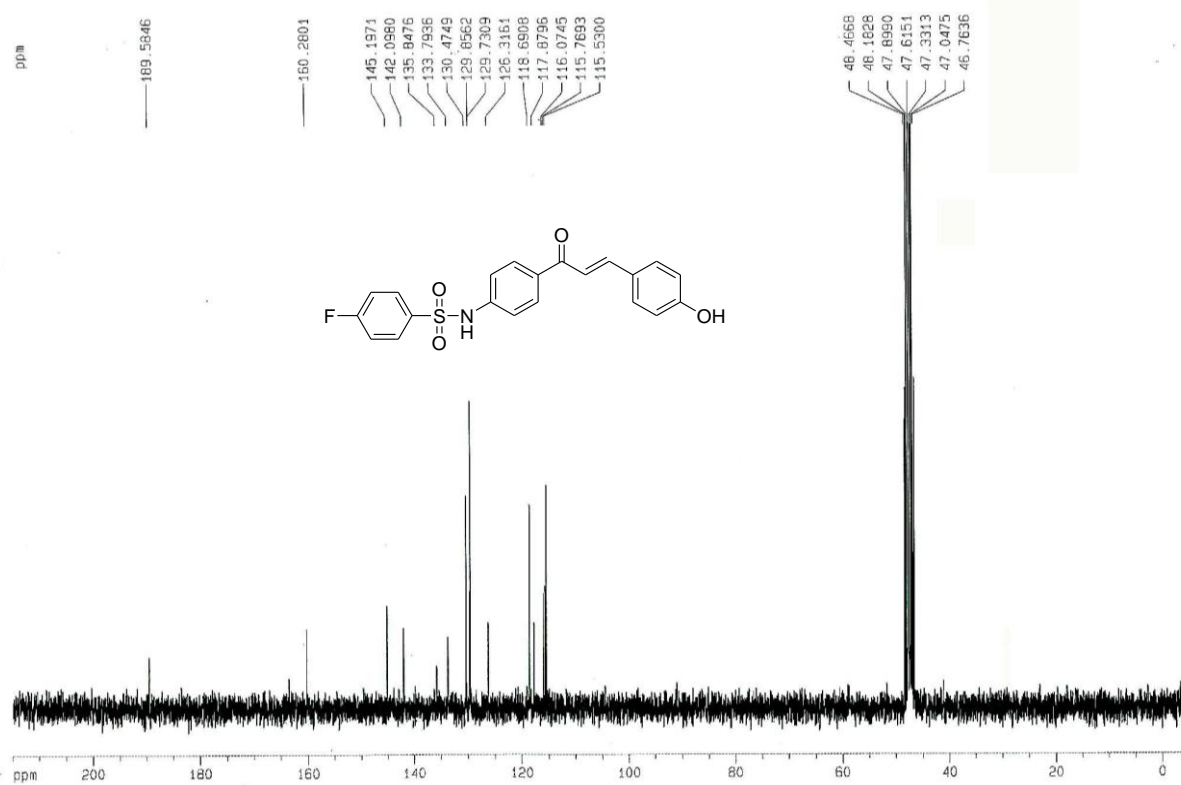
Spectra 11. $^1\text{H-NMR}$ of 4'-(benzenesulfonamide)-4-hydroxychalcone (**4c**).**Spectra 12.** $^{13}\text{C-NMR}$ of 4'-(benzenesulfonamide)-4-hydroxychalcone (**4c**).

Spectra 13. $^1\text{H-NMR}$ of 4'-(4-hydroxybenzenesulfonamide)-4-hydroxychalcone (**4d**).**Spectra 14.** $^{13}\text{C-NMR}$ of 4'-(4-hydroxybenzenesulfonamide)-4-hydroxychalcone (**4d**).

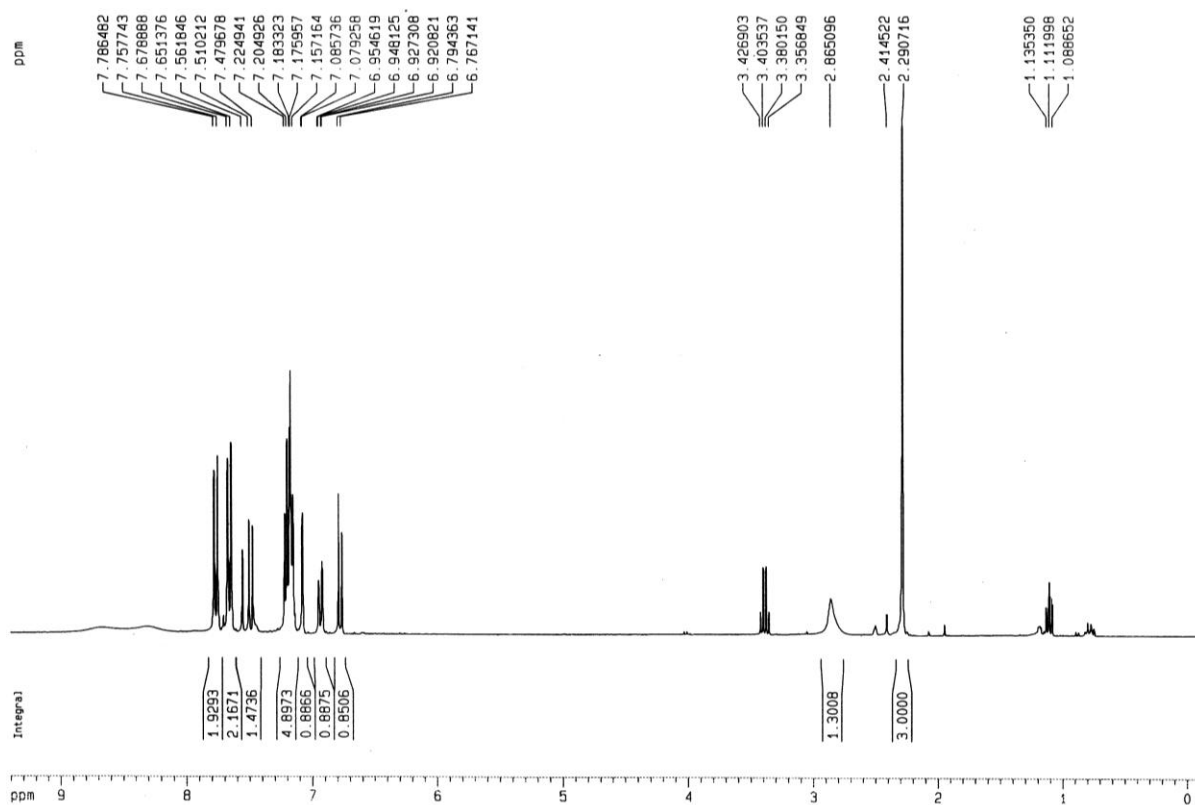
Spectra 15. $^1\text{H-NMR}$ of 4'-(4-methoxybenzenesulfonamido)-4-hydroxychalcone (**4e**).**Spectra 16.** $^{13}\text{C-NMR}$ of 4'-(4-methoxybenzenesulfonamido)-4-hydroxychalcone (**4e**).

Spectra 17. $^1\text{H-NMR}$ of 4'-(4-aminobenzensulfonamide)-4-hydroxychalcone (**4f**).**Spectra 18.** $^{13}\text{C-NMR}$ of 4'-(4-aminobenzensulfonamide)-4-hydroxychalcone (**4f**).

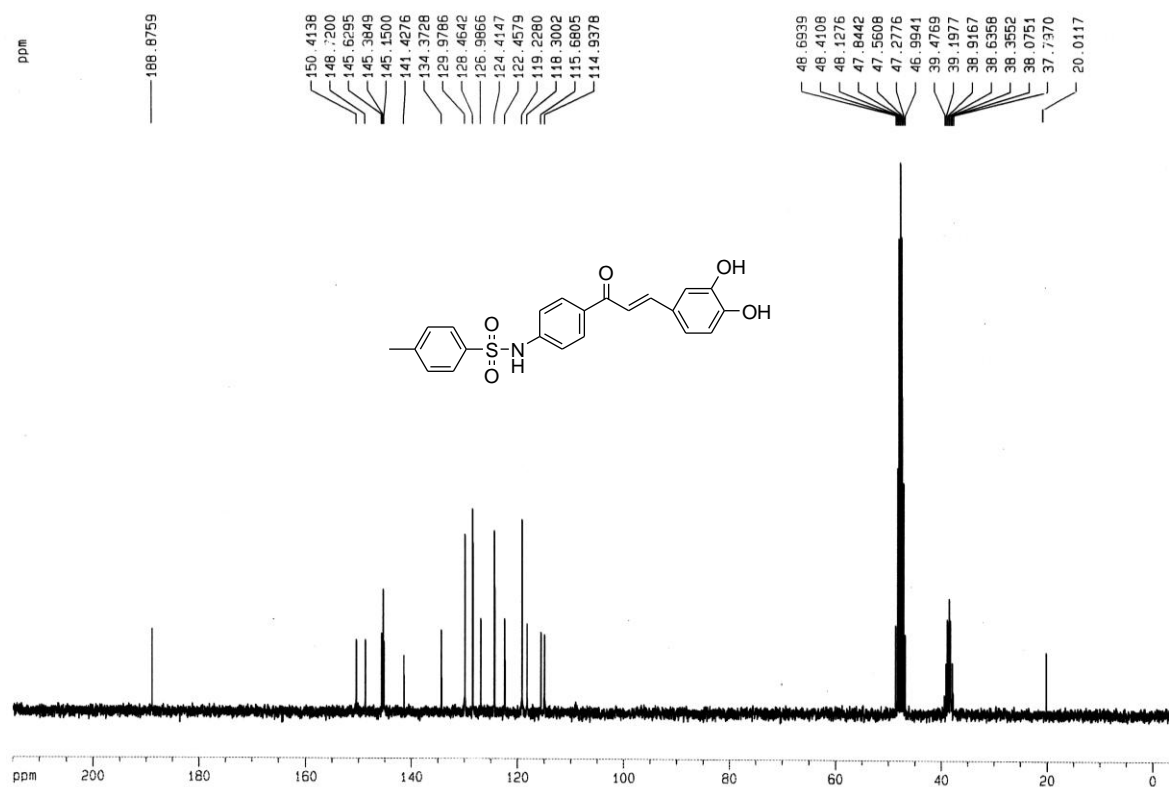
Spectra 19. $^1\text{H-NMR}$ of 4'-(4-nitrobenzenesulfonamide)-4-hydroxychalcone (**4g**).**Spectra 20.** $^{13}\text{C-NMR}$ of 4'-(4-nitrobenzenesulfonamide)-4-hydroxychalcone (**4g**).

Spectra 21. ^{13}C -NMR of 4'-(4-fluorobenzensulfonamide)-4-hydroxychalcone (**4h**).**Spectra 22.** ^{13}C -NMR of 4'-(4-fluorobenzensulfonamide)-4-hydroxychalcone (**4h**).

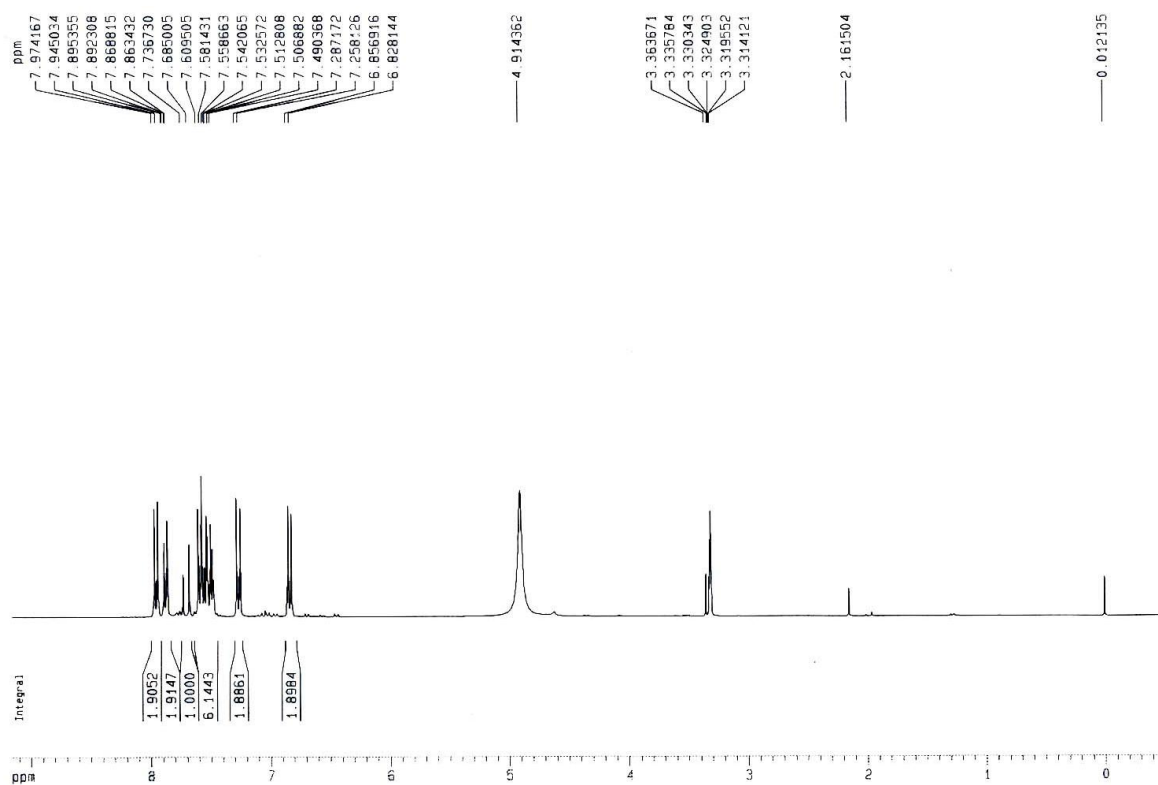
Spectra 23. $^1\text{H-NMR}$ of 4'-(*p*-toluenesulfonamide)-3,4-dihydroxychalcone (**5a**).



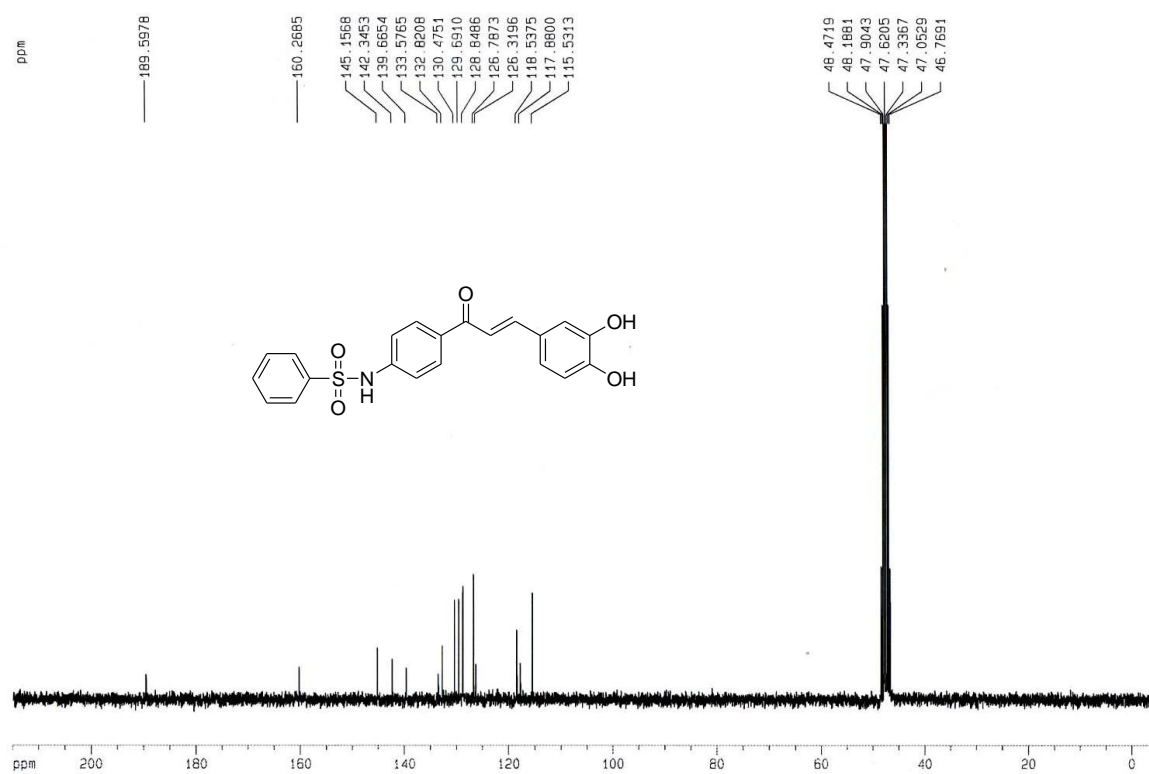
Spectra 24. $^{13}\text{C-NMR}$ of 4'-(*p*-toluenesulfonamide)-3,4-dihydroxychalcone (**5a**).

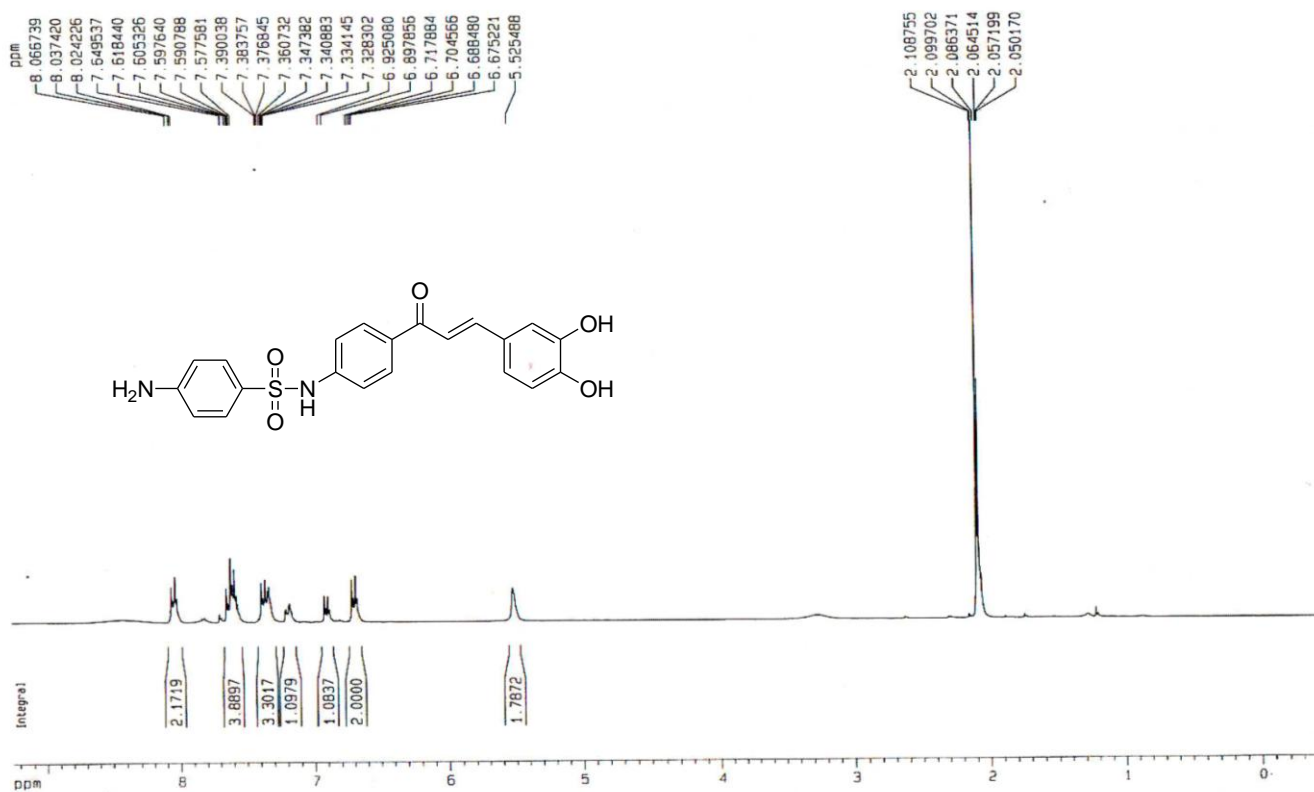
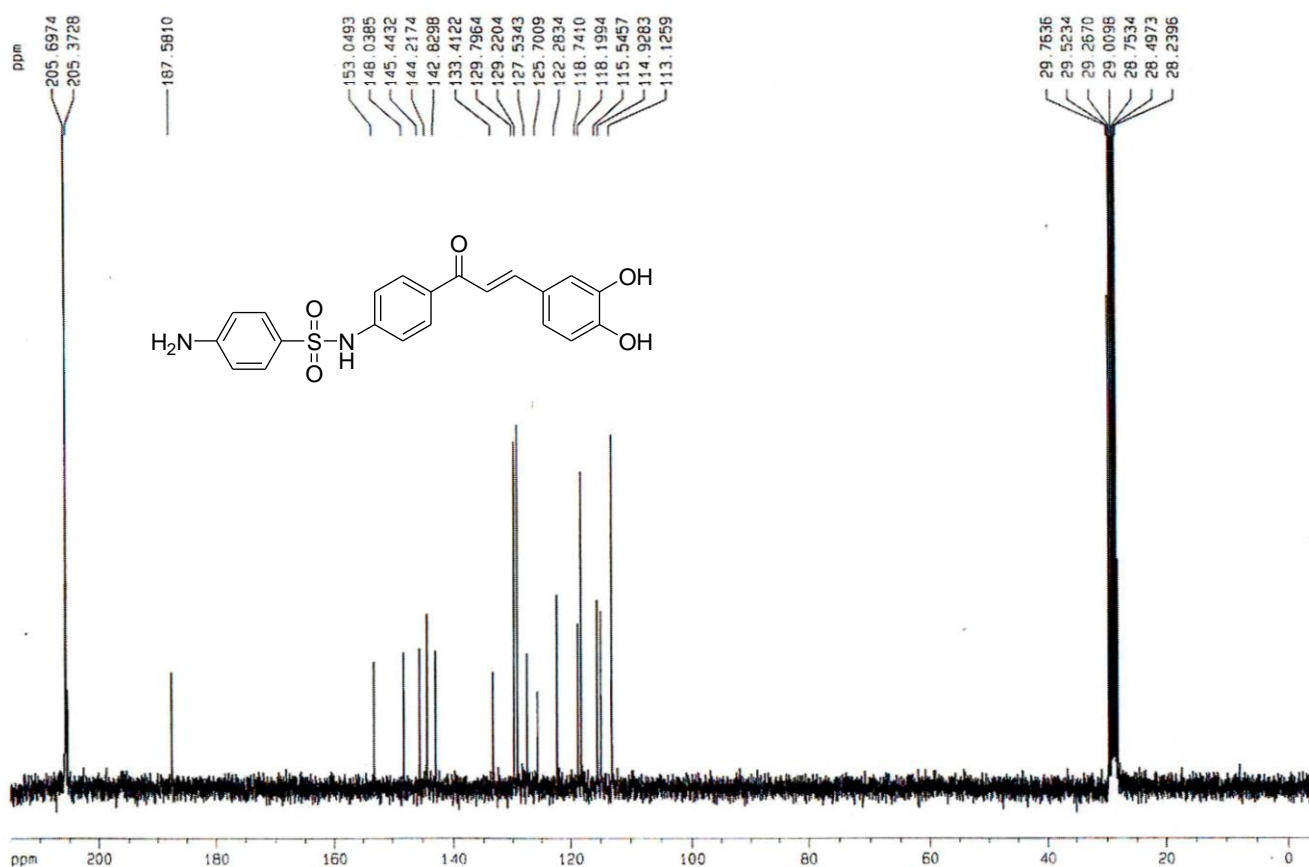


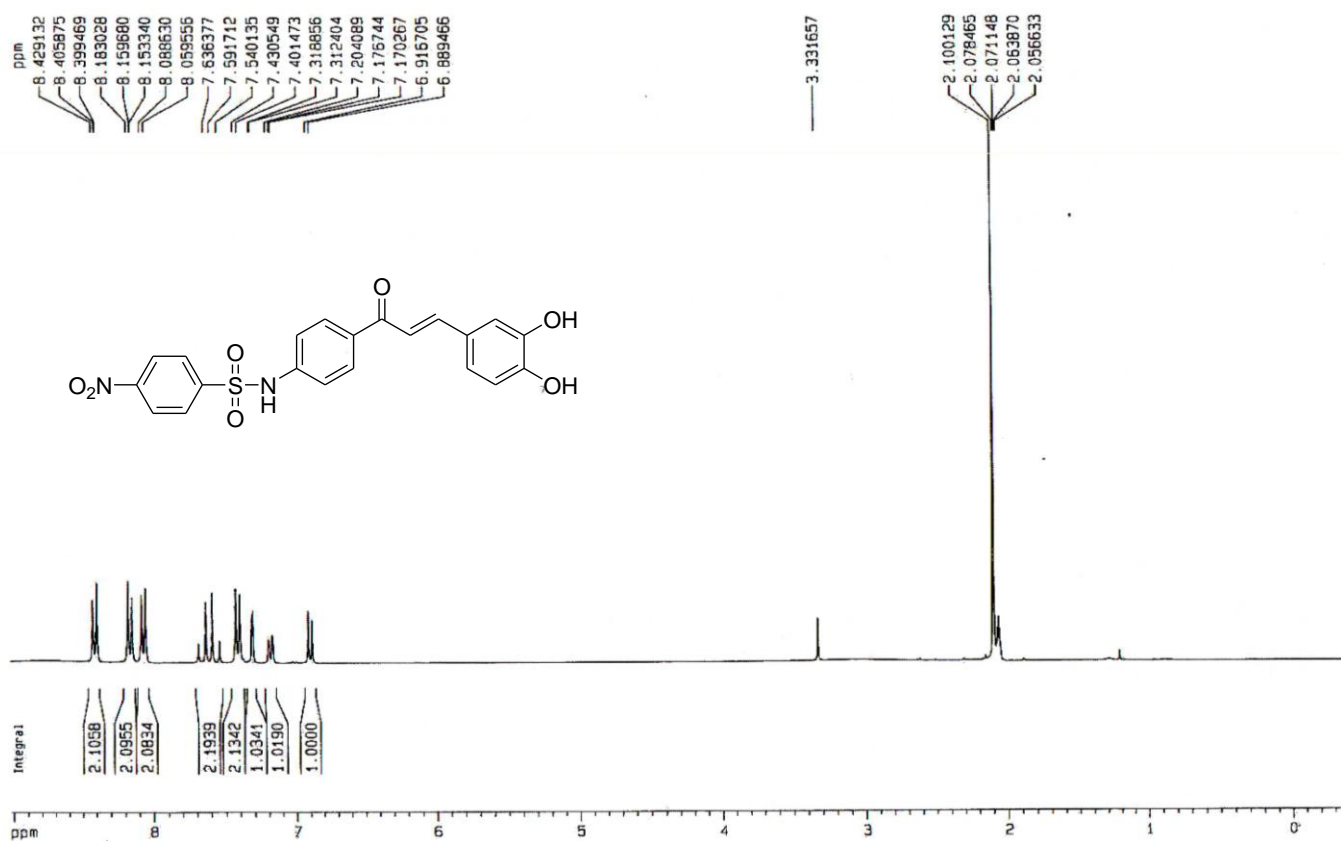
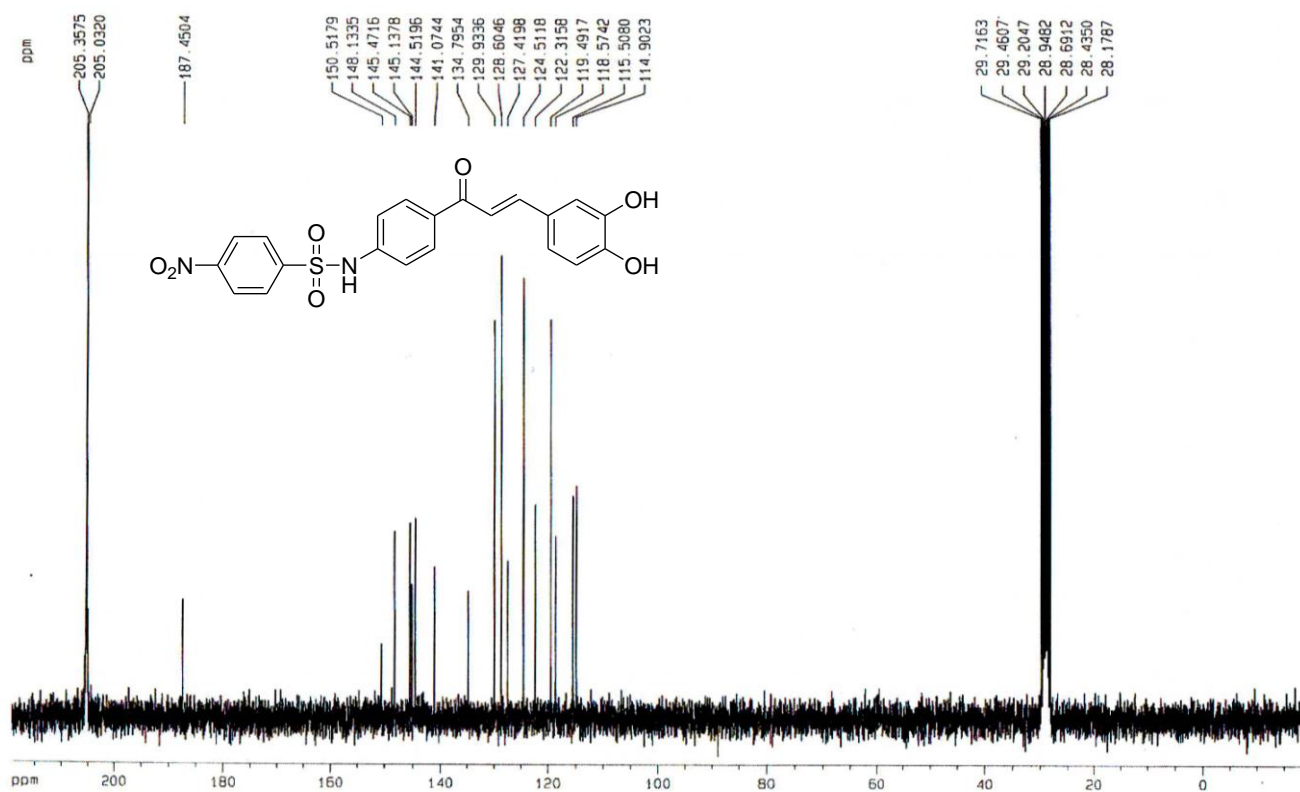
Spectra 25. $^1\text{H-NMR}$ of 4'-(benzenesulfonamide)-3,4-dihydroxychalcone (**5b**).

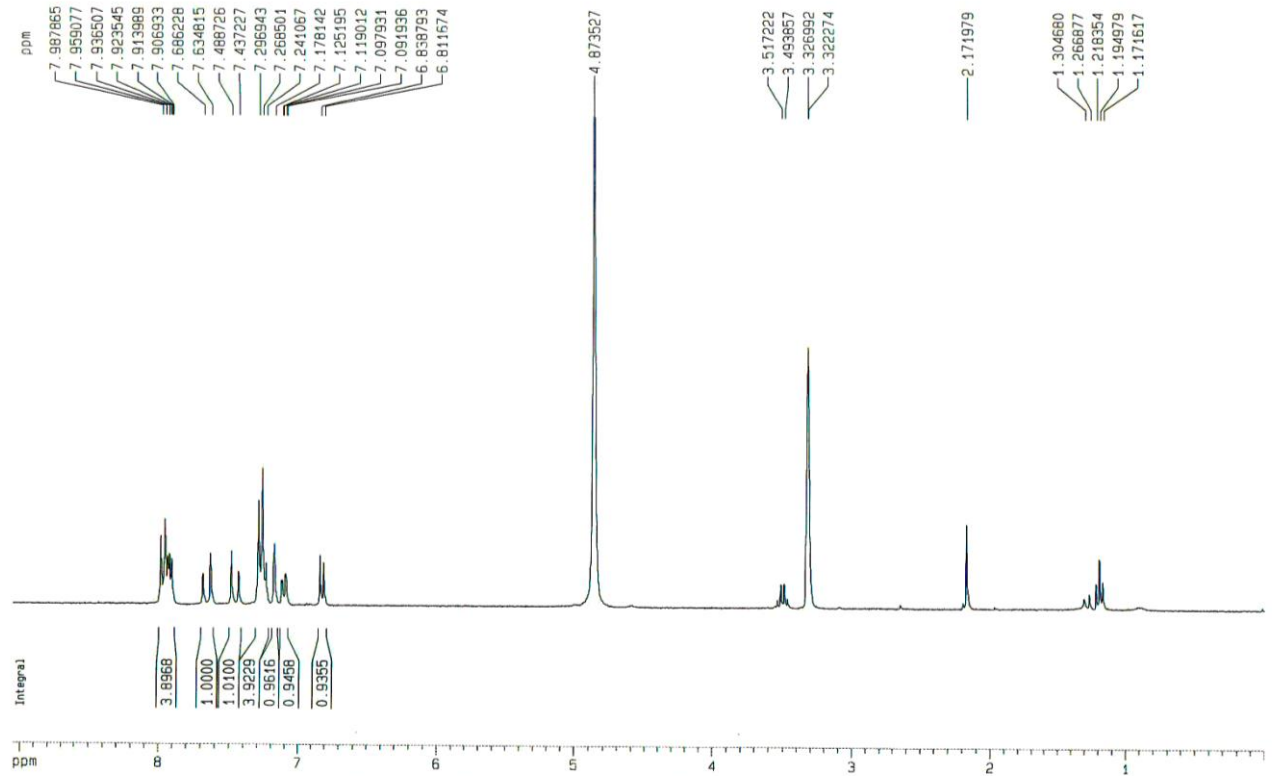
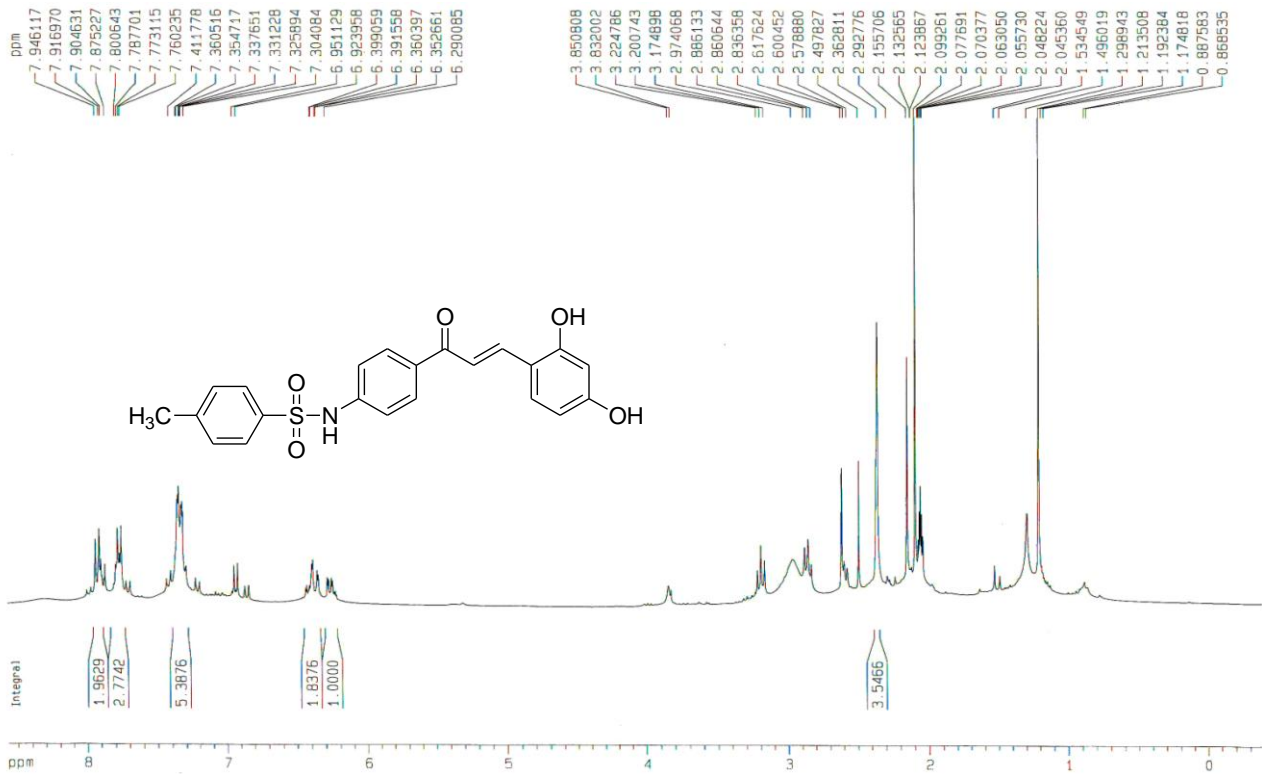


Spectra 26. $^{13}\text{C-NMR}$ of 4'-(benzenesulfonamide)-3,4-dihydroxychalcone (**5b**).



Spectra 27. $^1\text{H-NMR}$ of 4'-(4-aminobenzensulfonamide)-3,4-dihydroxychalcone (**5d**).**Spectra 28.** $^{13}\text{C-NMR}$ of 4'-(4-aminobenzensulfonamide)-3,4-dihydroxychalcone (**5d**).

Spectra 29. $^1\text{H-NMR}$ of 4'-(4-nitrobenzenesulfonamide)-3,4-dihydrochalcone (**5e**).**Spectra 30.** $^{13}\text{C-NMR}$ of 4'-(4-nitrobenzenesulfonamide)-3,4-dihydrochalcone (**5e**).

Spectra 31. $^1\text{H-NMR}$ of 4'-(4-fluoroenzenulfonamide)-3,4-dihydroxychalcone (**5f**).**Spectra 32.** $^1\text{H-NMR}$ of 4'-(*p*-toluenesulfonamide)-2,4-dihydroxychalcone (**6**).

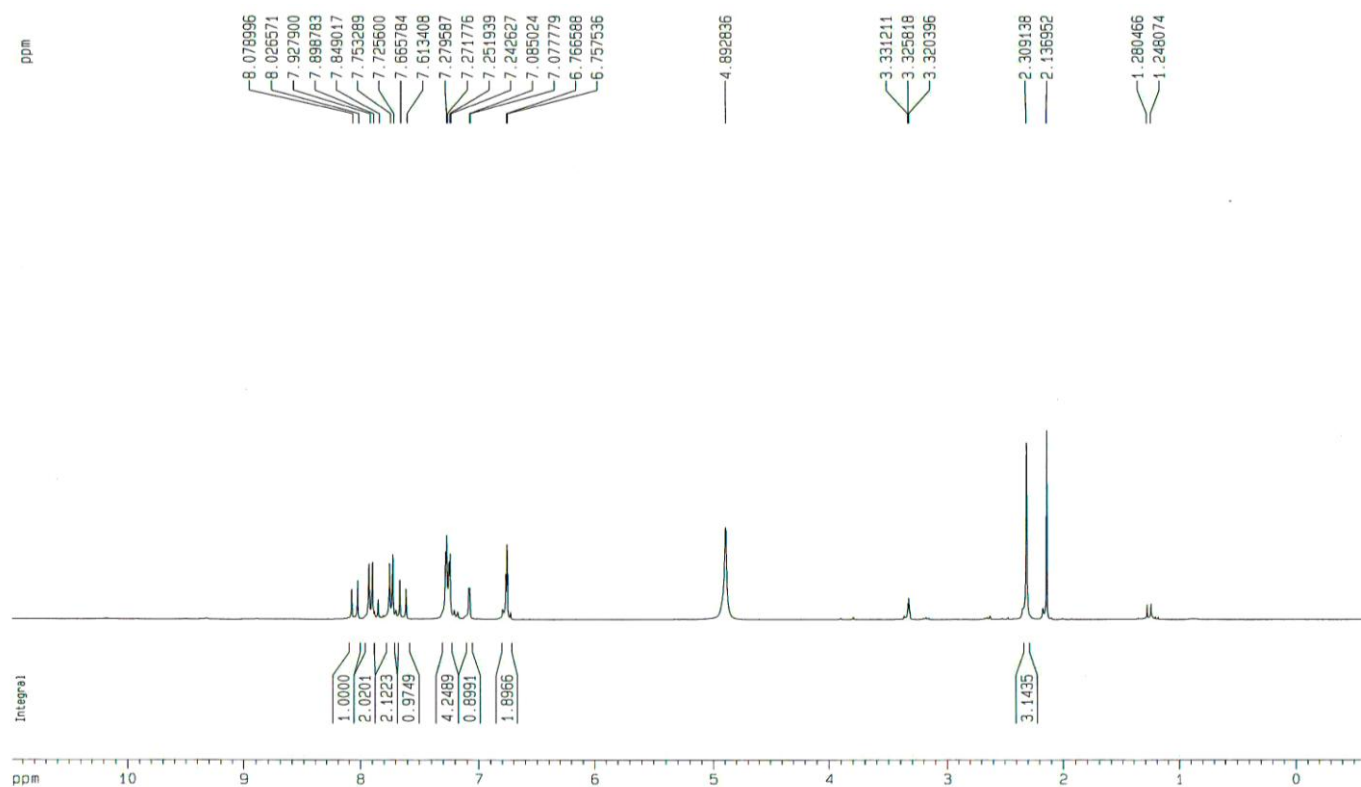
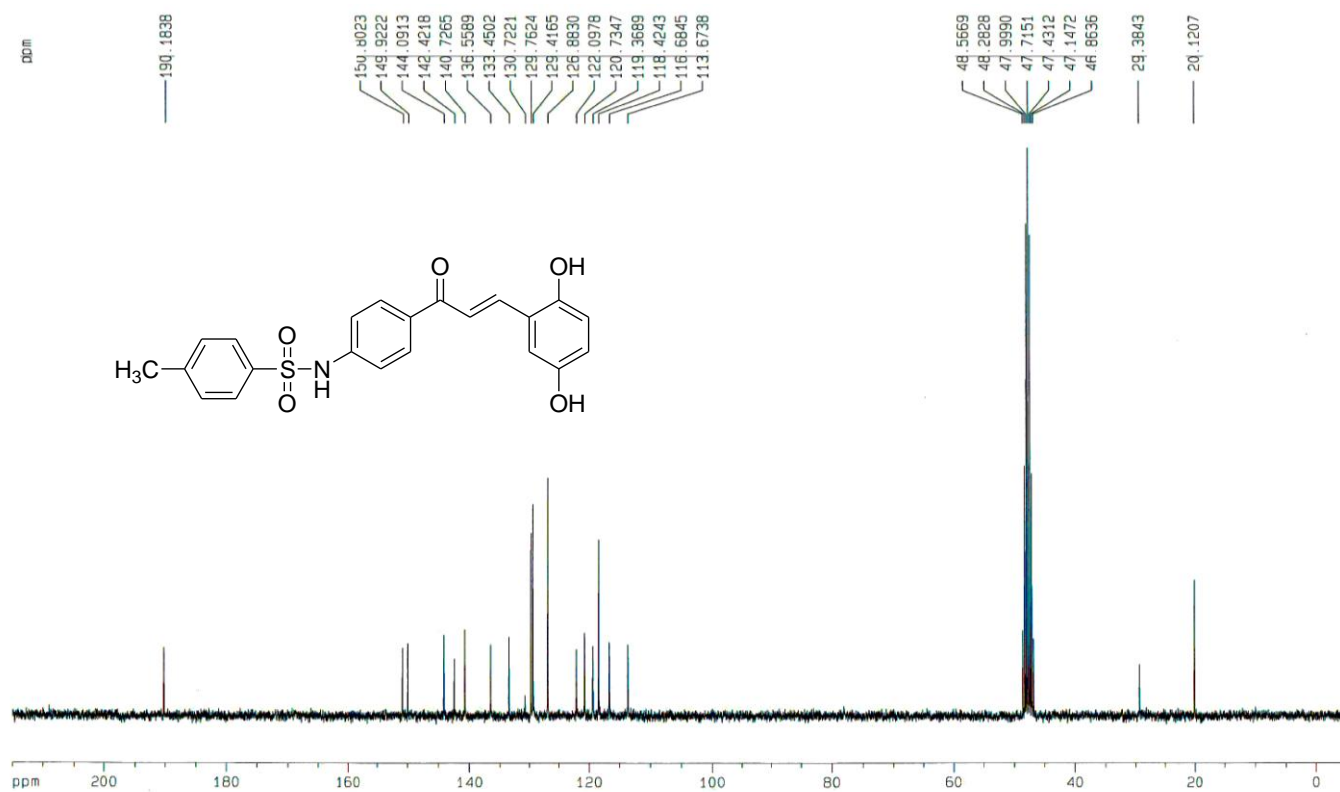
Spectra 33. $^1\text{H-NMR}$ of 4'-(*p*-toluenesulfonamide)-2,5-dihydroxychalcone (7).

Spectra 34. $^{13}\text{C-NMR}$ of 4'-(*p*-toluenesulfonamide)-2,5-dihydroxychalcone (7).


Figure 1. Effect of compounds **1**, **2**, **4a,c**, and **5d** on the hydrolytic activity of BACE1. K_m values as a function of the concentrations of **1**, **2**, **4a,c**, and **5d**. (Inset) Dependence of the values of V_{max} on the concentration of compounds **1**, **2**, **4a,c**, and **5d**.

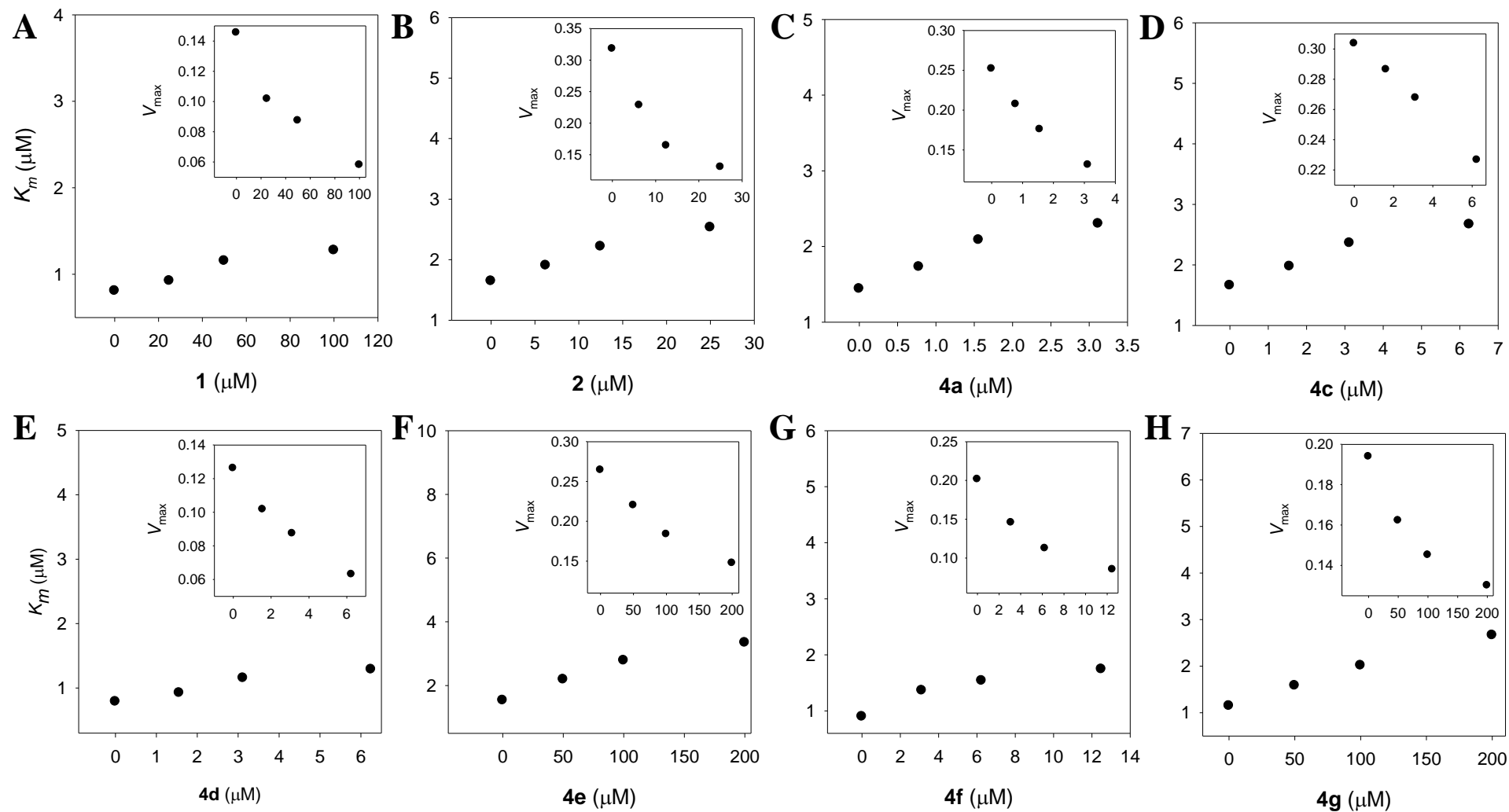


Figure 2. Effect of compounds **1**, **2**, **4a,c**, and **5d** on the hydrolytic activity of BACE1. Dixon plots for the inhibition of compounds **1**, **2**, **4a,c**, and **5d** on BACE1 catalyzed proteolysis of substrate.

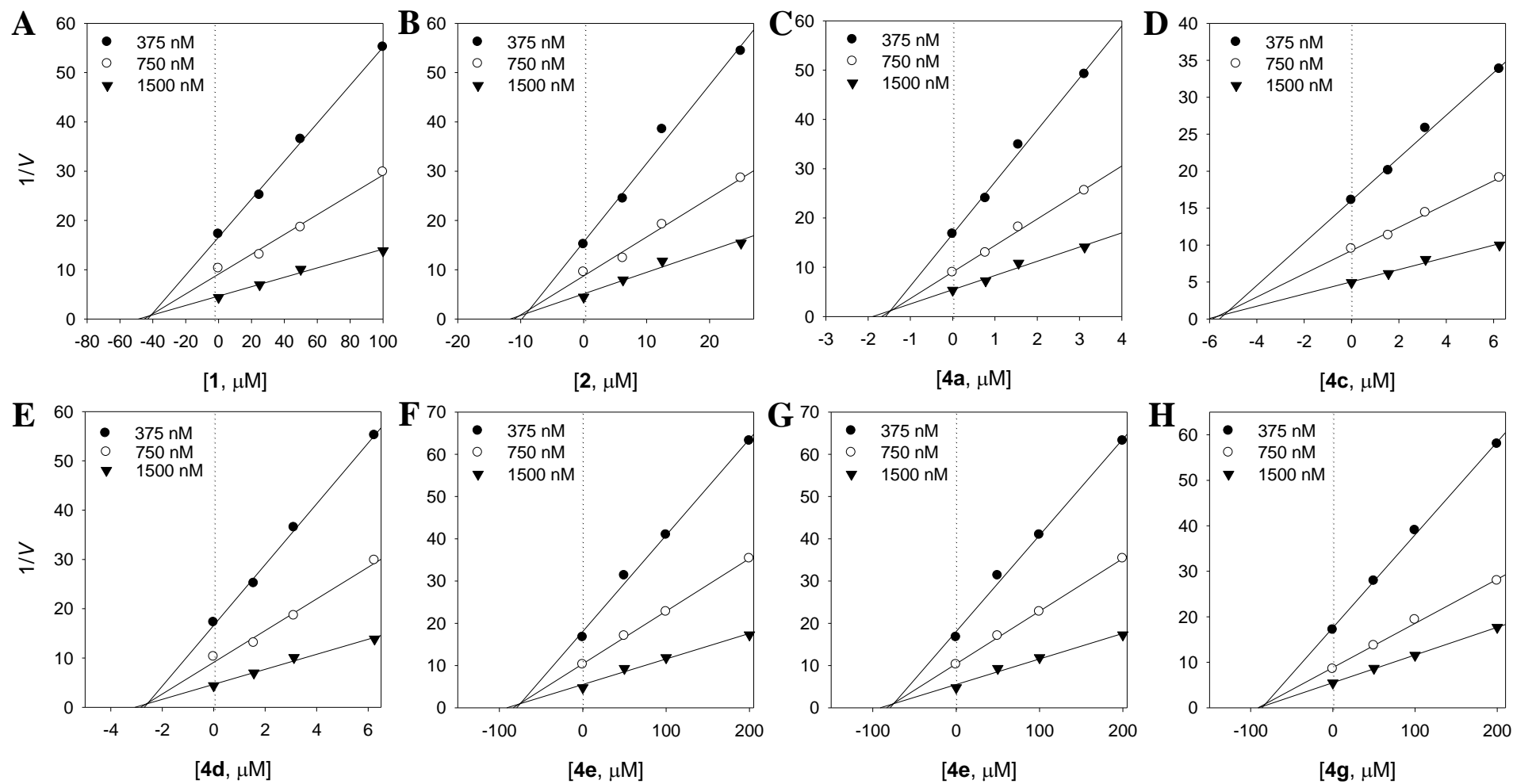


Figure 3. Effect of compounds **5a–d** on the hydrolytic activity of BACE1. K_m values as a function of the concentrations of **5a–d**. (Inset) Dependence of the values of V_{max} on the concentration of compounds **5a–5d**.

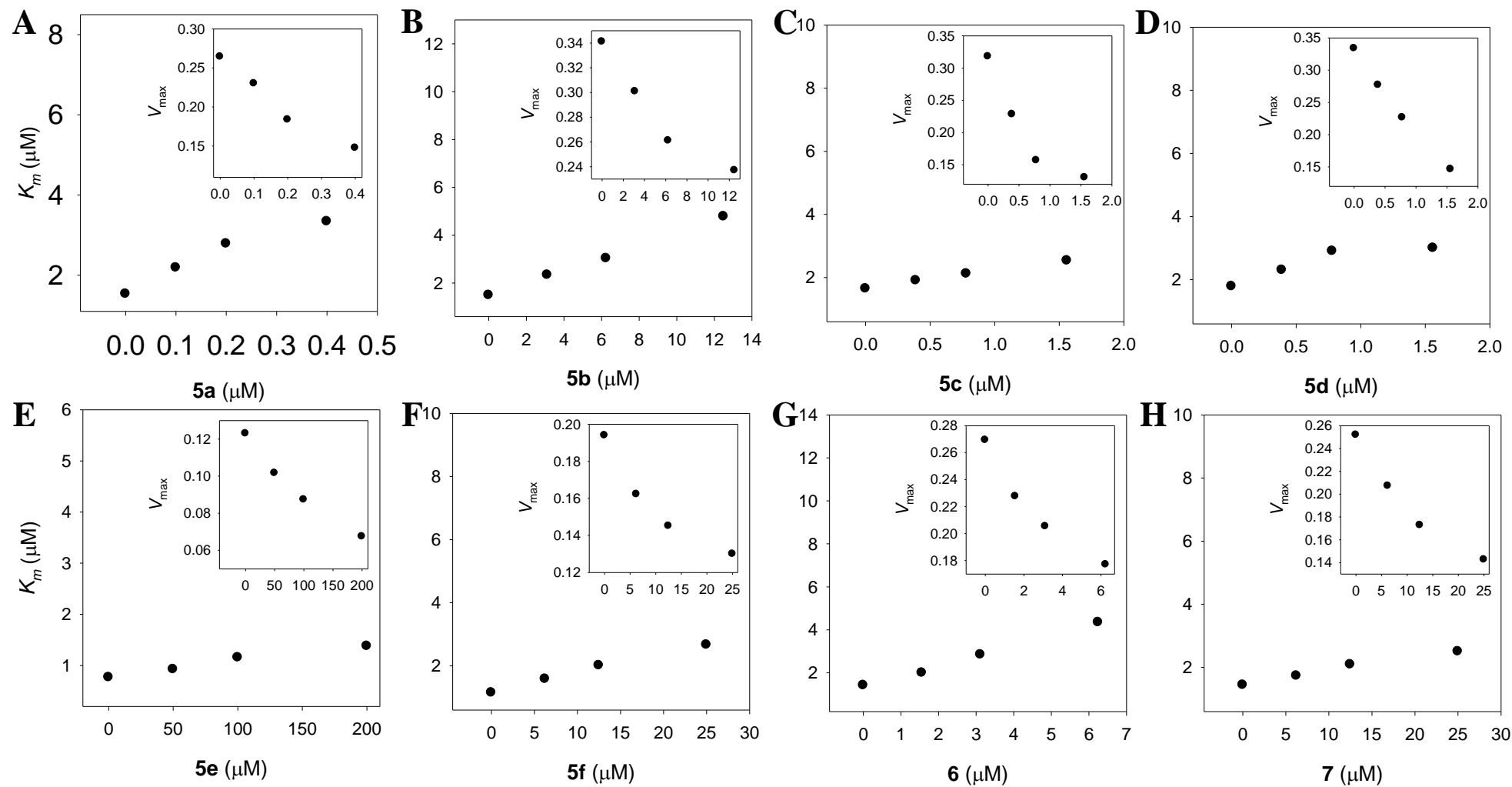


Figure 4. Effect of compounds **5a–d** on the hydrolytic activity of BACE1. Dixon plots for the inhibition of compounds **5a–d** on BACE1 catalyzed proteolysis of substrate.

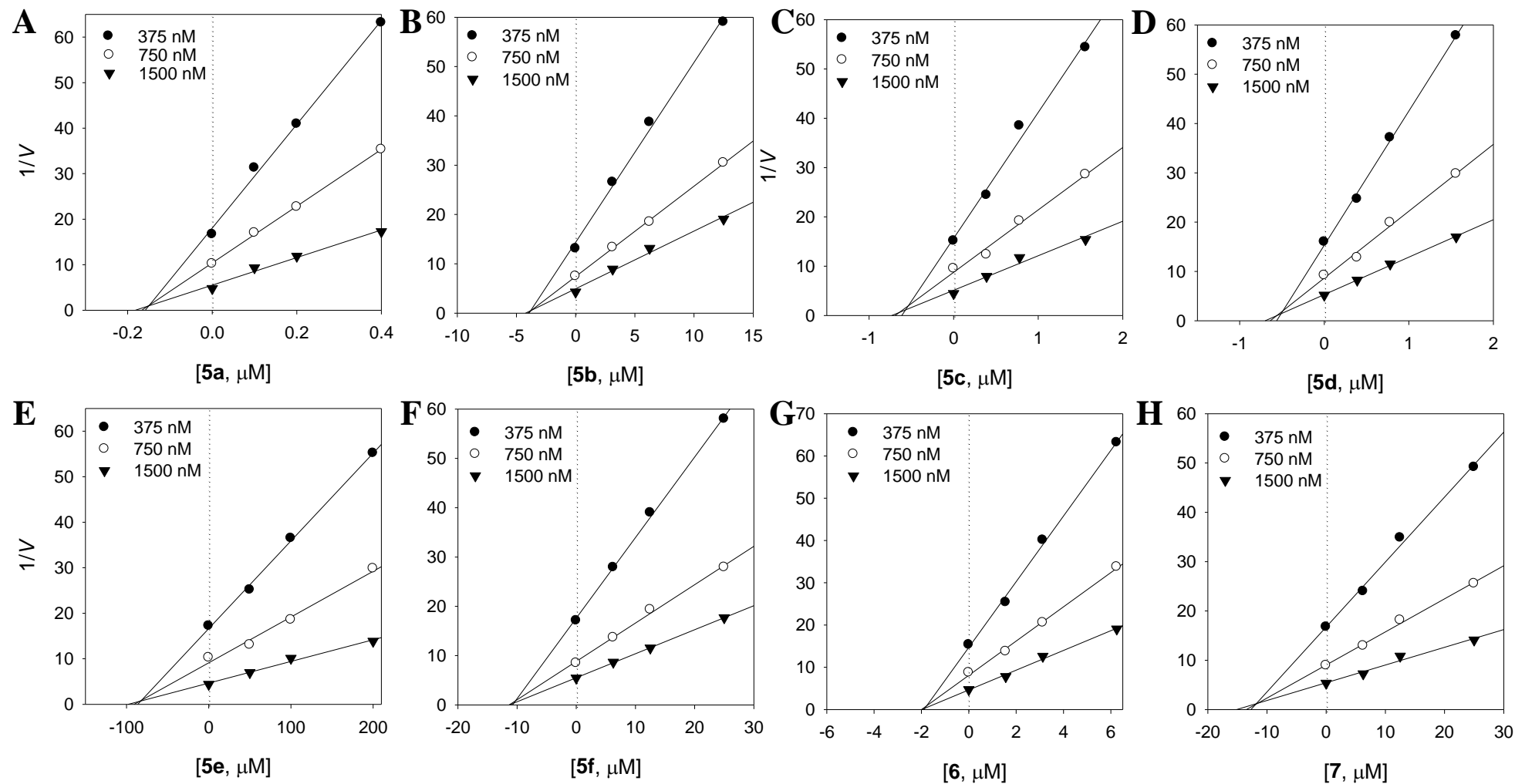


Figure 5. Acetylcholinesterase kinetic data of compounds 4a,f, and 5b,d.

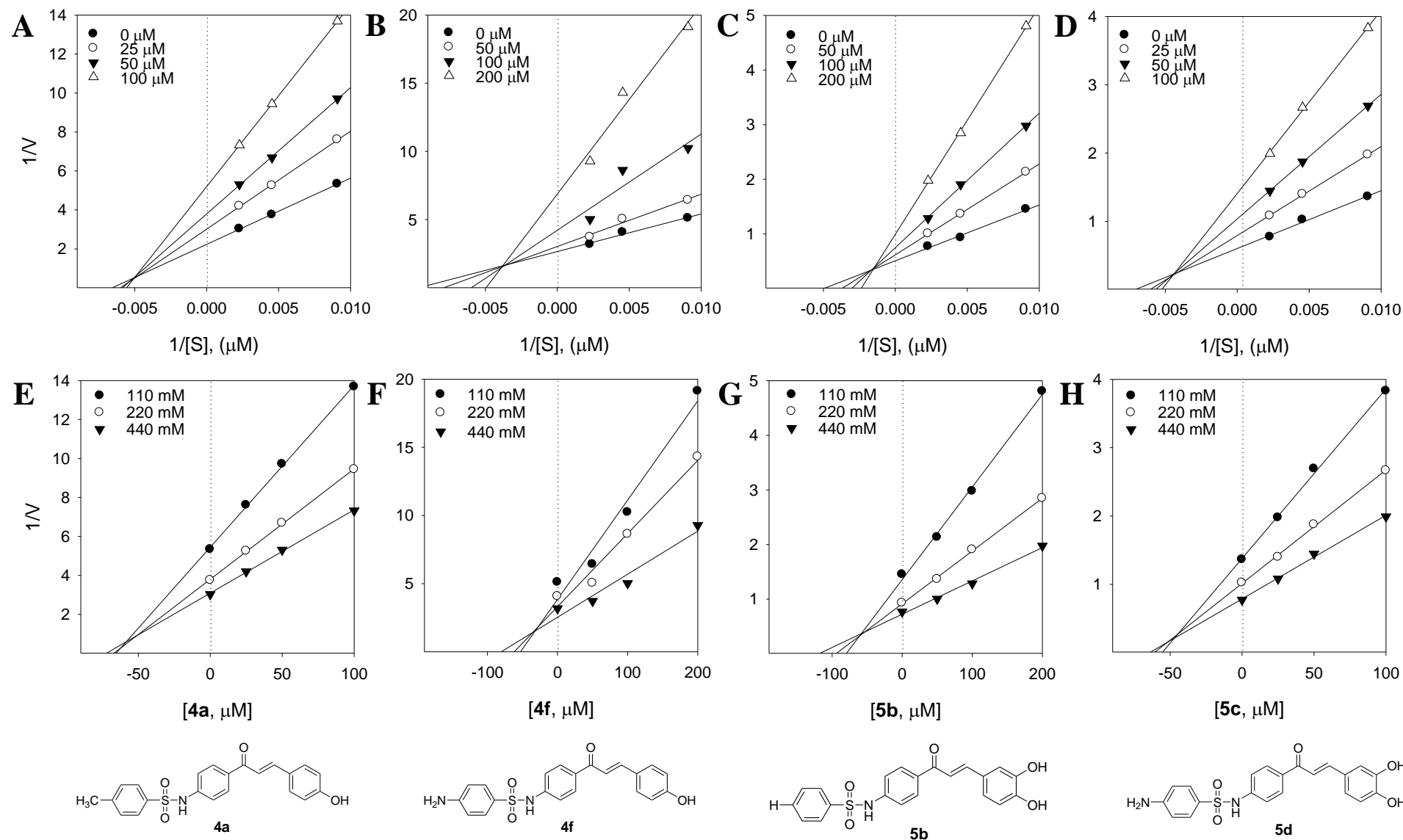


Figure 6. Butyrylcholinesterase kinetic data of compounds 4a,f, and 5b,d.

