

## Supporting Information

### The structural determinants for $\alpha_1$ -adrenergic/serotonin receptors activity among phenylpiperazine-hydantoin derivatives.

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† Equal contribution

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1. <sup>1</sup>H and <sup>13</sup>C NMR spectra for final compounds

Figure S1. (Z)-3-(2-hydroxy-3-(4-(2-methoxyphenyl)piperazin-1-yl)propyl)-5-(3-methoxybenzylidene)imidazolidine-2,4-dione (**4**)

$^1\text{H}$  NMR (500 MHz, DMSO- $d_6$ )

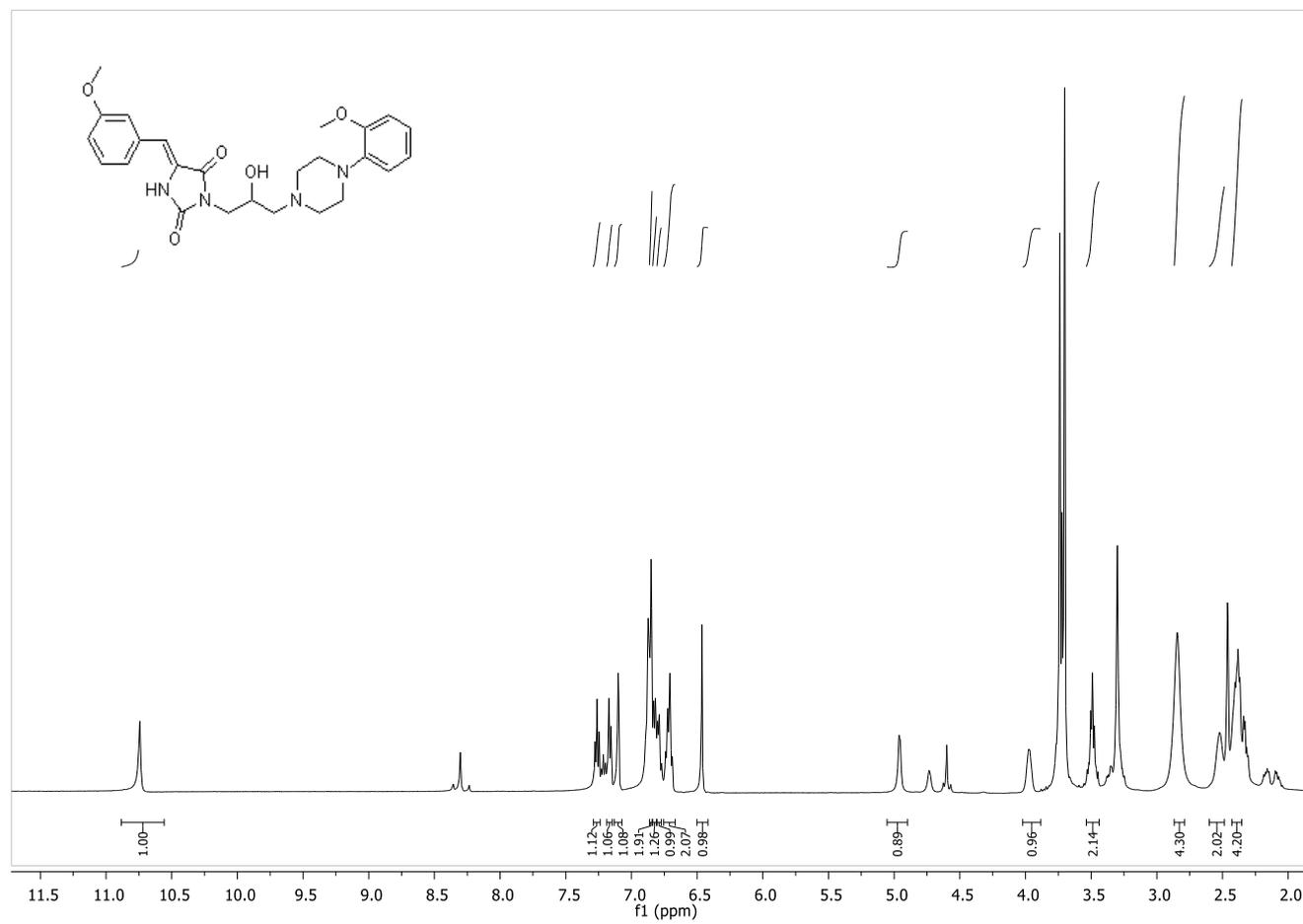


Figure S2. AD09 (Z)-3-(3-(4-(2-ethoxyphenyl)piperazin-1-yl)-2-hydroxypropyl)-5-(3-methoxybenzylidene)imidazolidine-2,4-dione (**5**)

$^1\text{H}$  NMR (500 MHz, DMSO- $d_6$ )

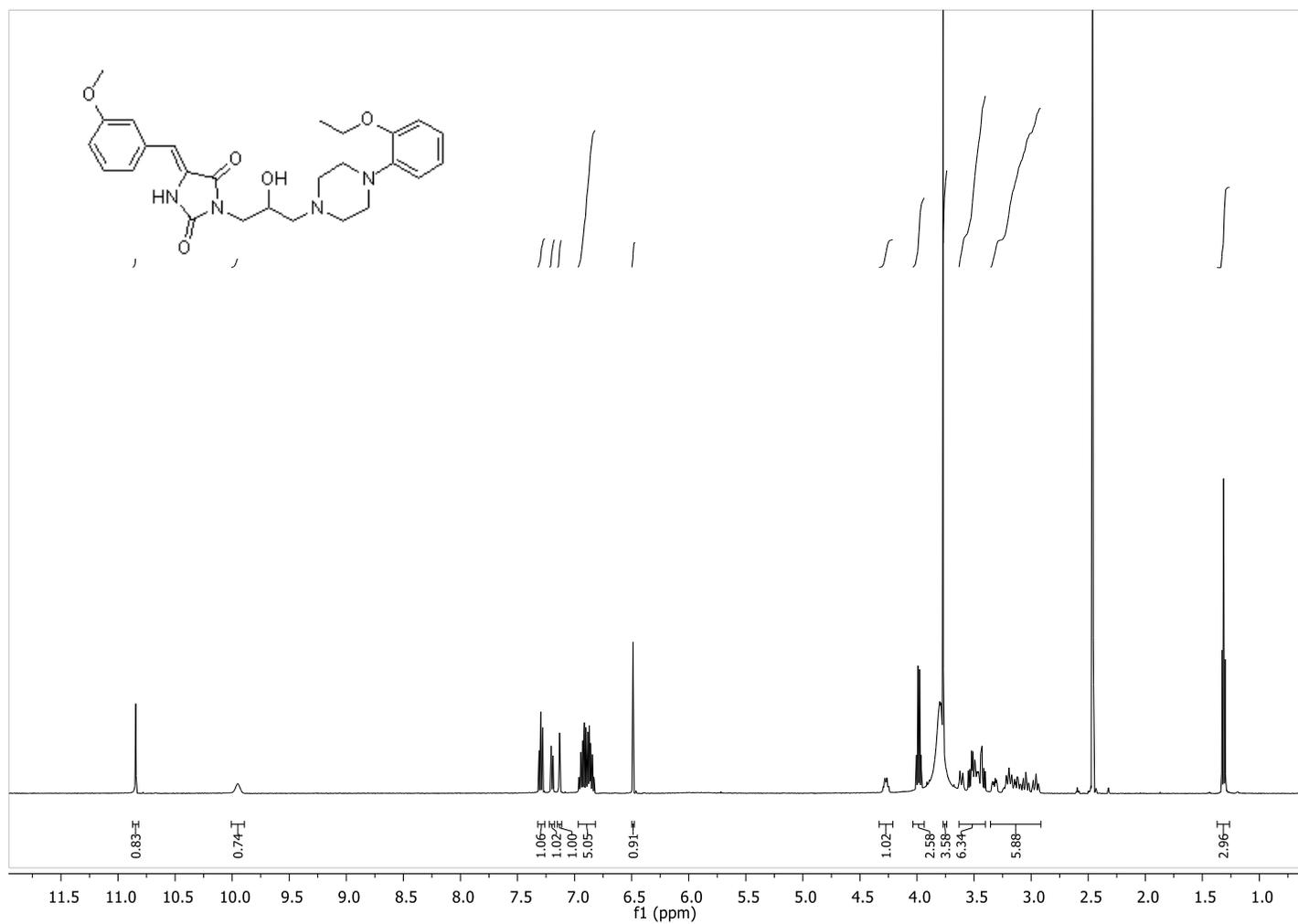


Figure S3. AD15 (*Z*)-3-(2-hydroxy-3-(4-(2-methoxyphenyl)piperazin-1-yl)propyl)-5-(4-methoxybenzylidene)imidazolidine-2,4-dione (**6**)

<sup>1</sup>H NMR (500 MHz, DMSO-d<sub>6</sub>)

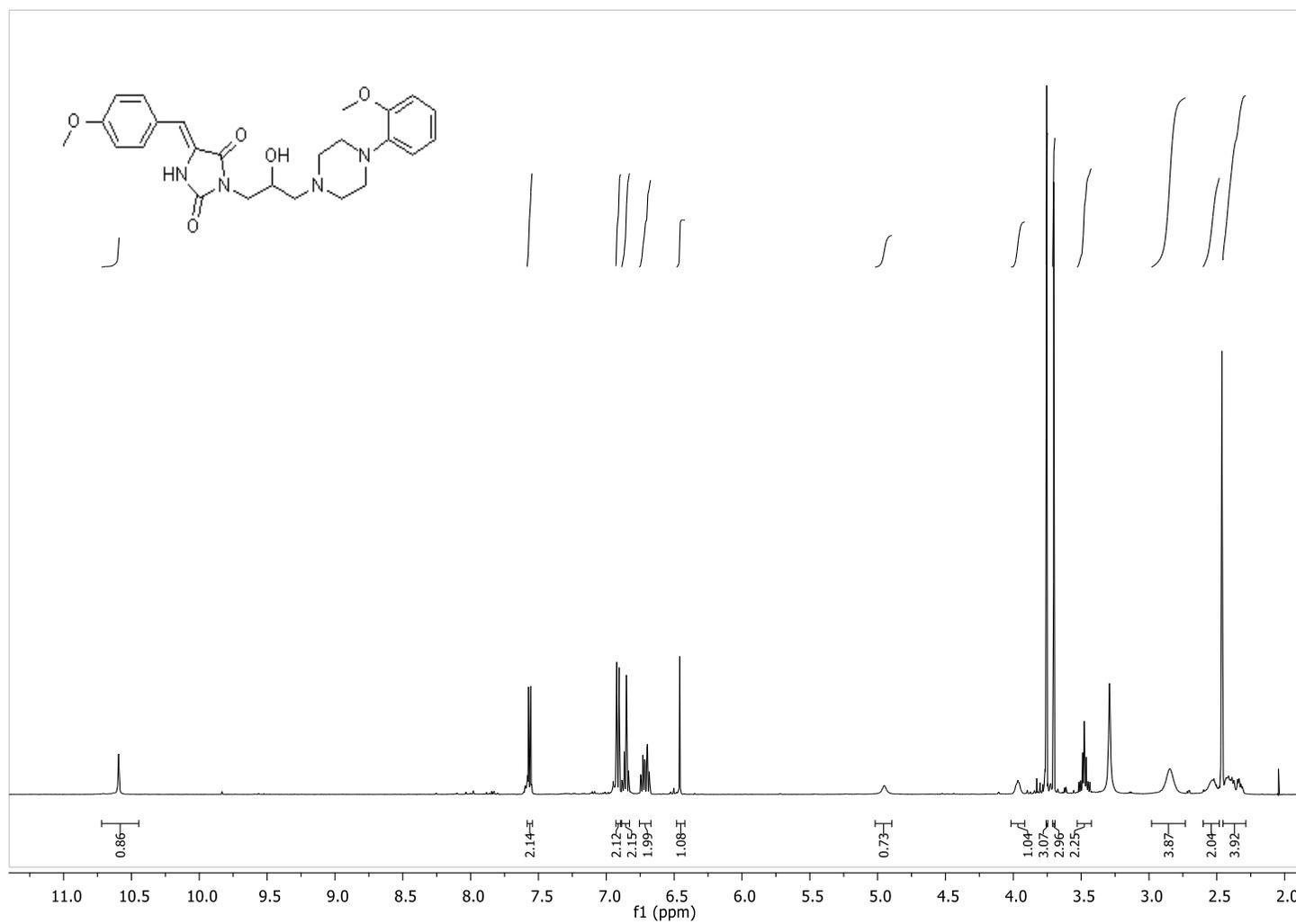
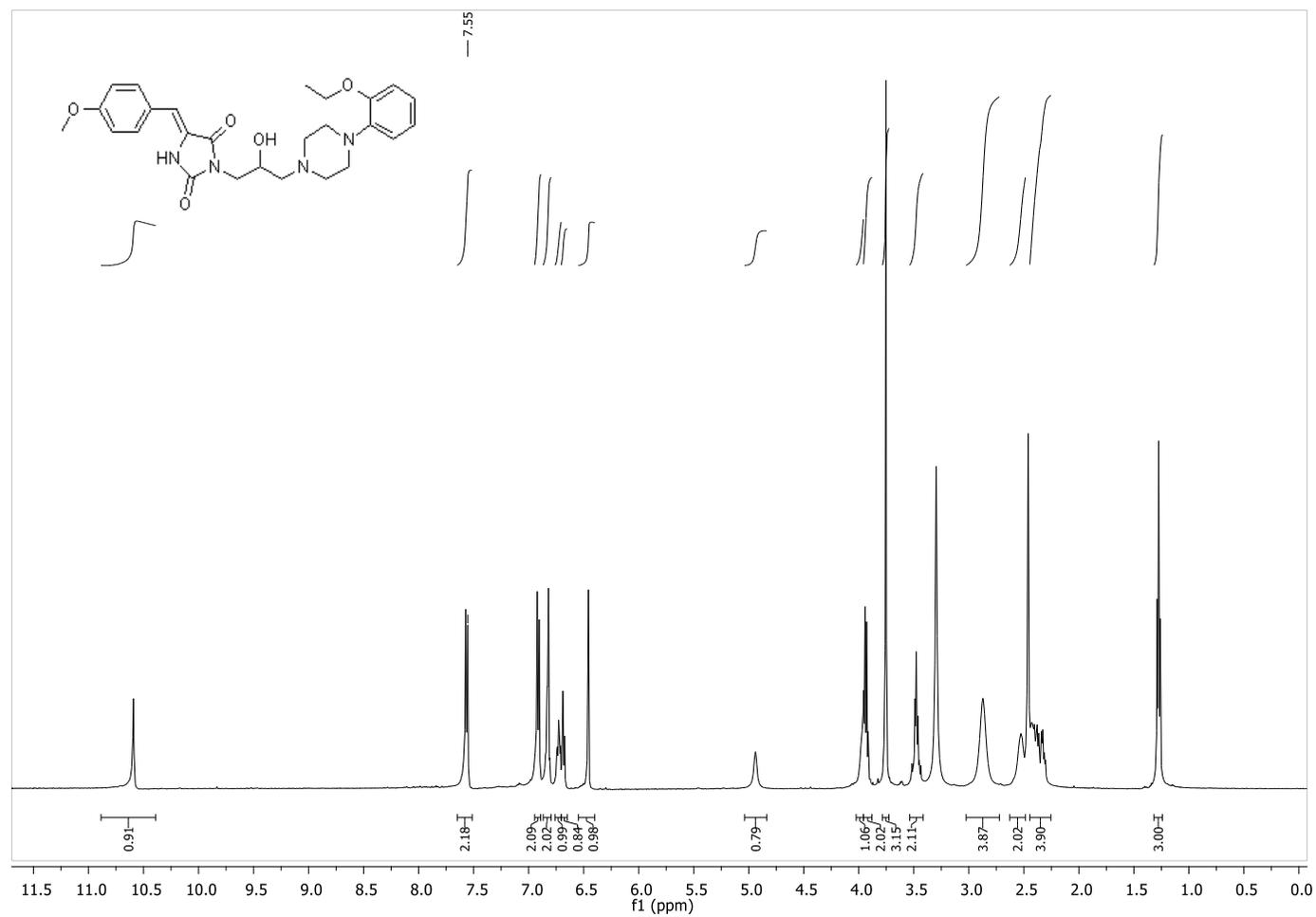


Figure S4.. AD13 (Z)-3-(3-(4-(2-ethoxyphenyl)piperazin-1-yl)-2-hydroxypropyl)-5-(4-methoxybenzylidene)imidazolidine-2,4-dione (**7**)

$^1\text{H}$  NMR (500 MHz, DMSO- $d_6$ )



$^{13}\text{C}$  NMR (125 MHz, DMSO- $d_6$ )

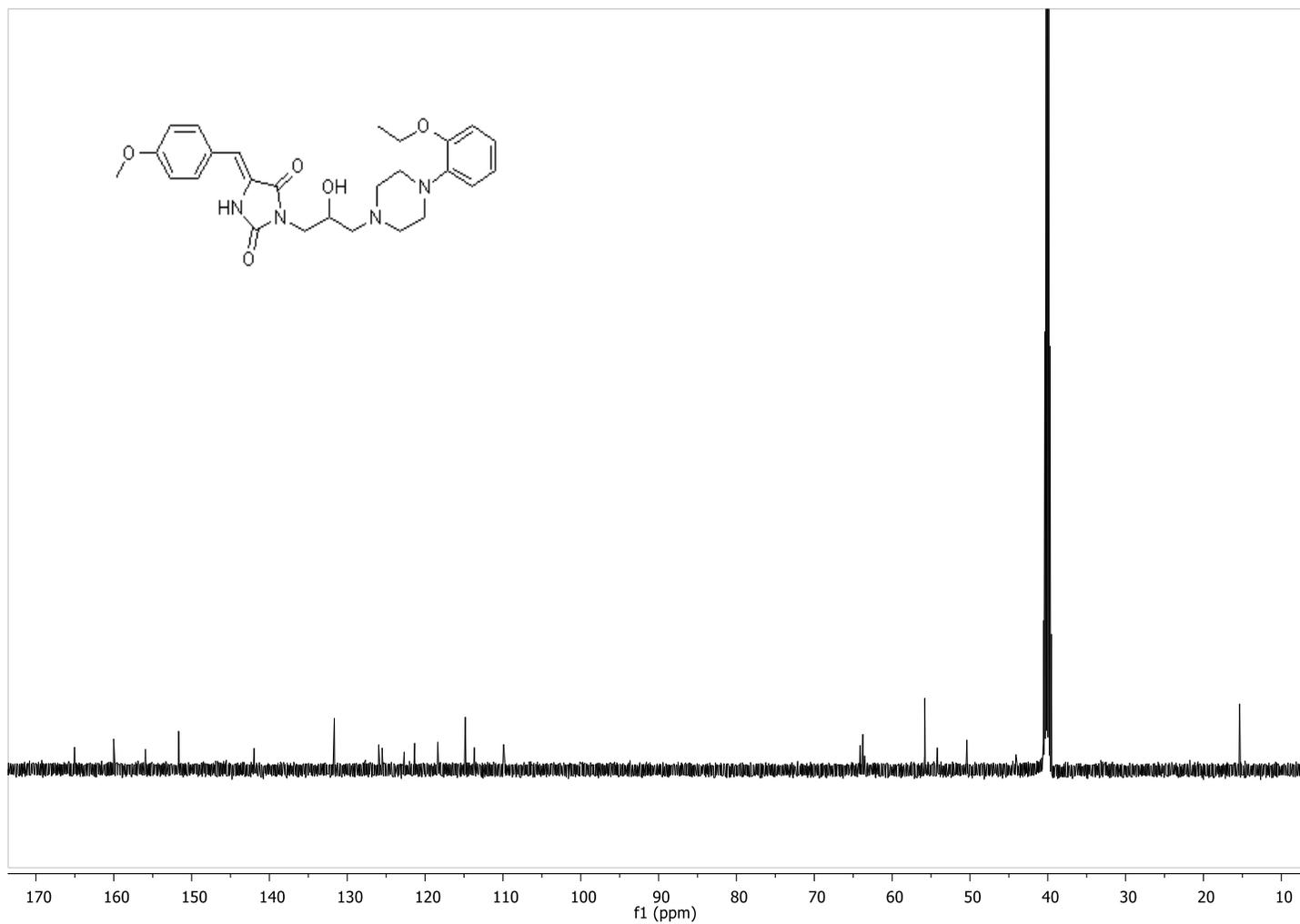
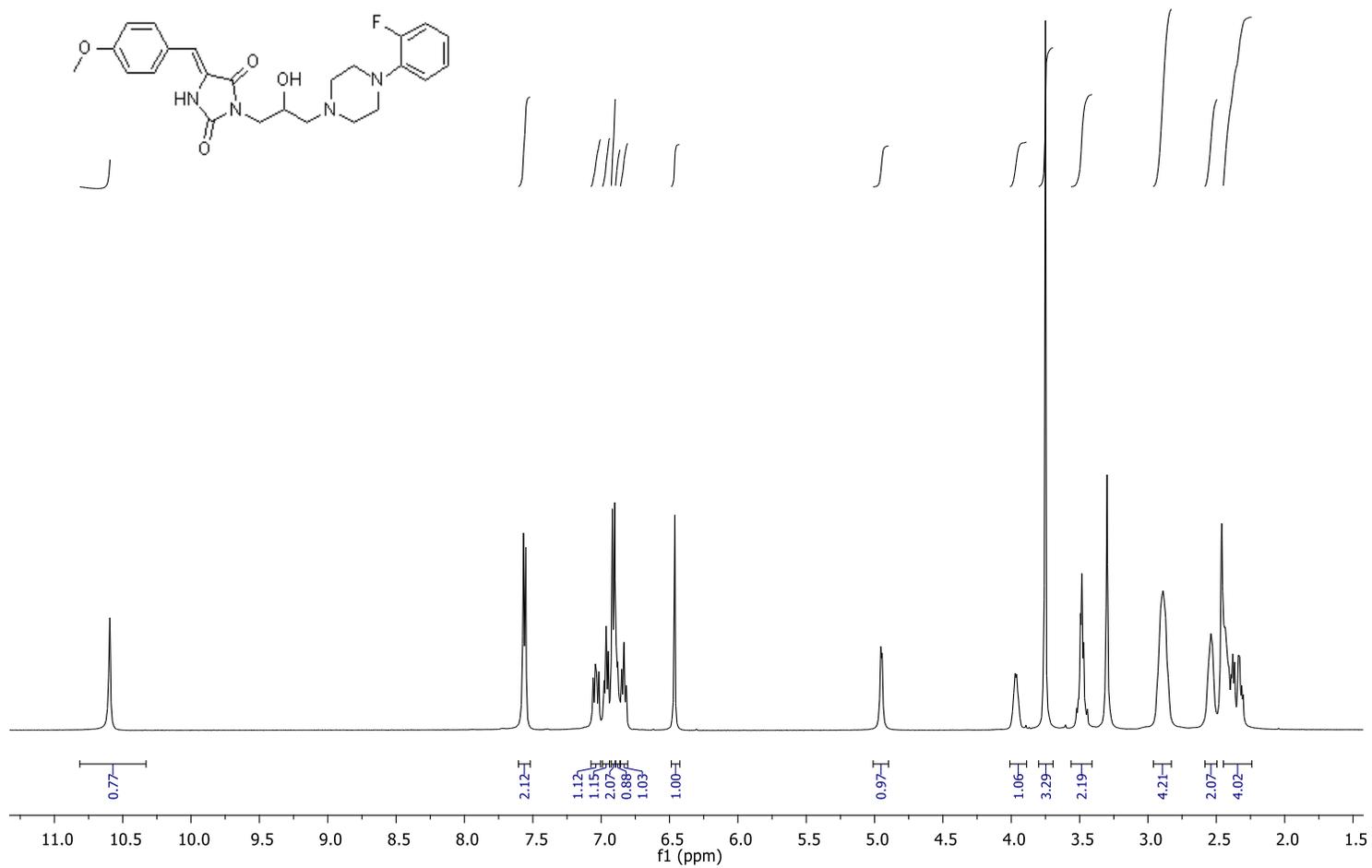


Figure S5. AD38 (*Z*)-3-(3-(4-(2-fluorophenyl)piperazin-1-yl)-2-hydroxypropyl)-5-(4-methoxybenzylidene)imidazolidine-2,4-dione (**8**)

$^1\text{H}$  NMR (500 MHz, DMSO- $d_6$ )



$^{13}\text{C}$  NMR (125 MHz, DMSO- $d_6$ )

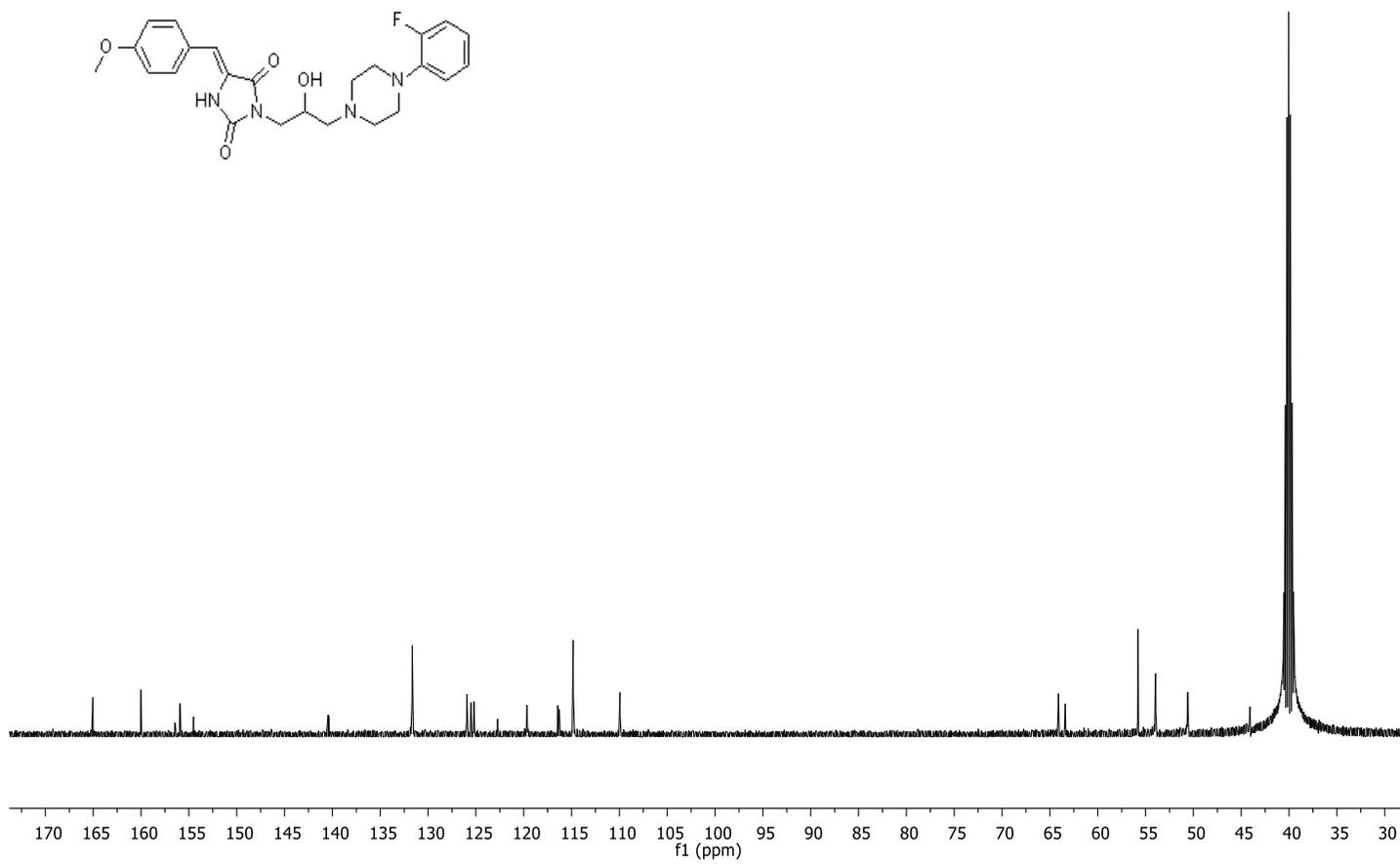
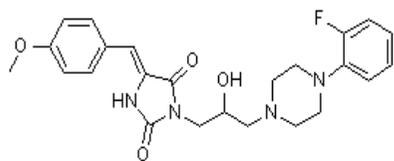


Figure S6. /AD34 (*Z*)-3-(3-(4-(2-ethoxyphenyl)piperazin-1-yl)-2-hydroxypropyl)-5-(3,4,5-trimethoxybenzylidene)imidazolidine-2,4-dione (**9**)

<sup>1</sup>H NMR (500 MHz, DMSO-d<sub>6</sub>)

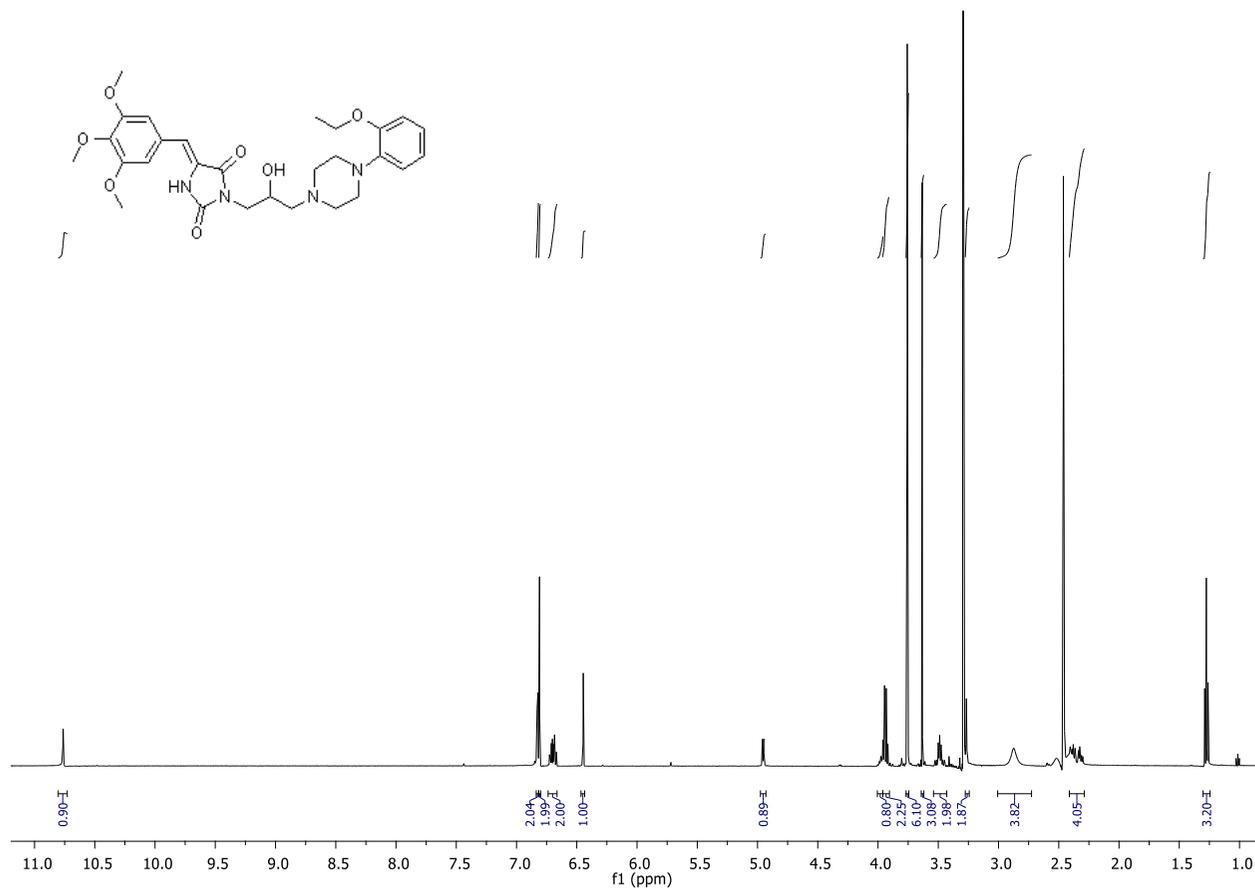


Figure S7. AD42 *(Z)*-3-(3-(4-(2-chlorophenyl)piperazin-1-yl)propyl)-5-(3-methoxybenzylidene)imidazolidine-2,4-dione (**11**)

<sup>1</sup>H NMR (500 MHz, DMSO-d<sub>6</sub>)

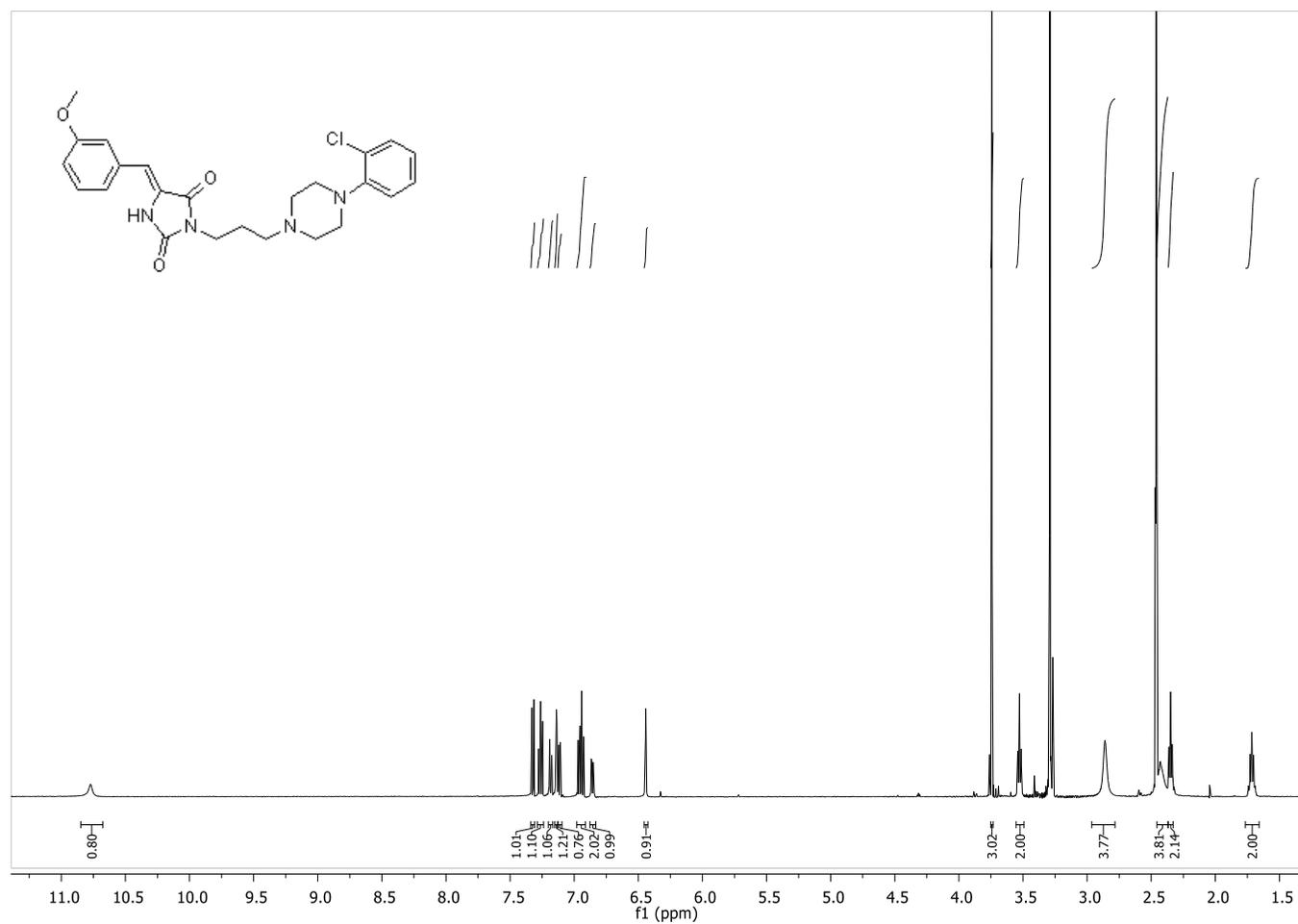


Figure S8. AD26 (Z)-5-(4-methoxybenzylidene)-3-(3-(4-(2-methoxyphenyl)piperazin-1-yl)propyl)imidazolidine-2,4-dione (**12**)

<sup>1</sup>H NMR (500 MHz, DMSO-d<sub>6</sub>)

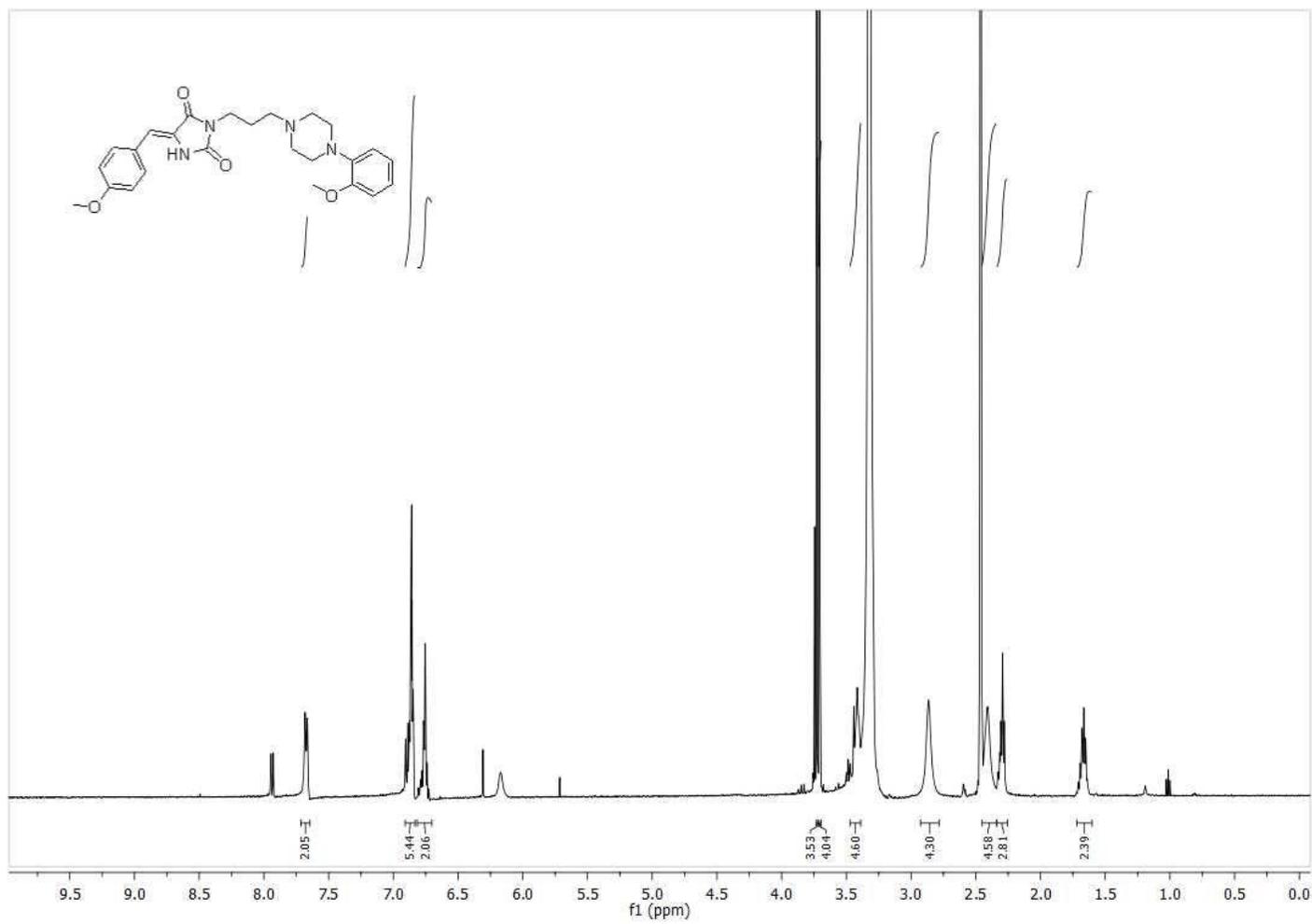


Figure S9. AD30 (*Z*)-3-(3-(4-(2-ethoxyphenyl)piperazin-1-yl)propyl)-5-(4-methoxybenzylidene)imidazolidine-2,4-dione (**13**)

<sup>1</sup>H NMR (500 MHz, DMSO-d<sub>6</sub>)

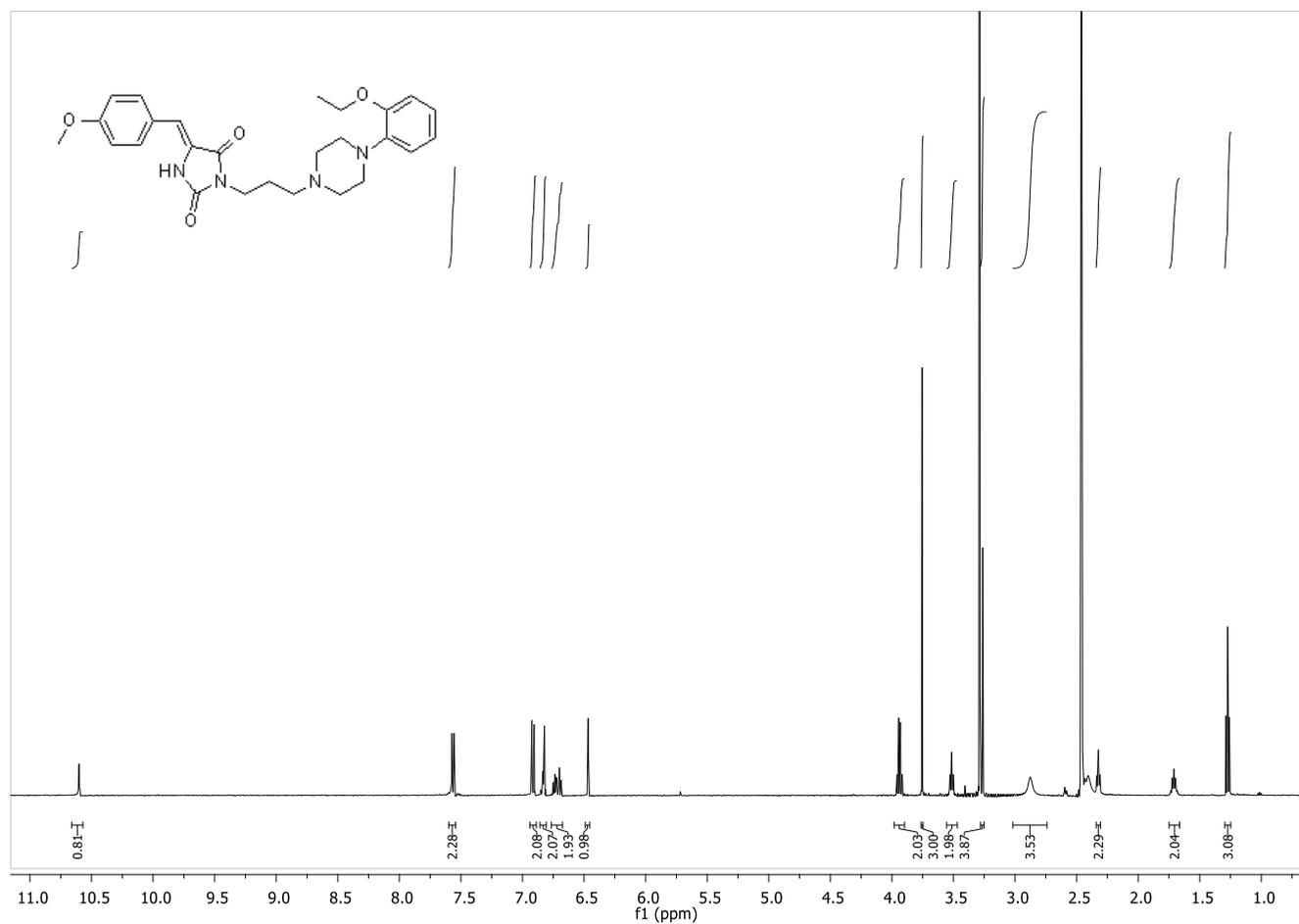


Figure S10. AD28 (Z)-3-(3-(4-(2-methoxyphenyl)piperazin-1-yl)propyl)-5-(3,4,5-trimethoxybenzylidene)imidazolidine-2,4-dione (**15**)

<sup>1</sup>H NMR (500 MHz, DMSO-d<sub>6</sub>)

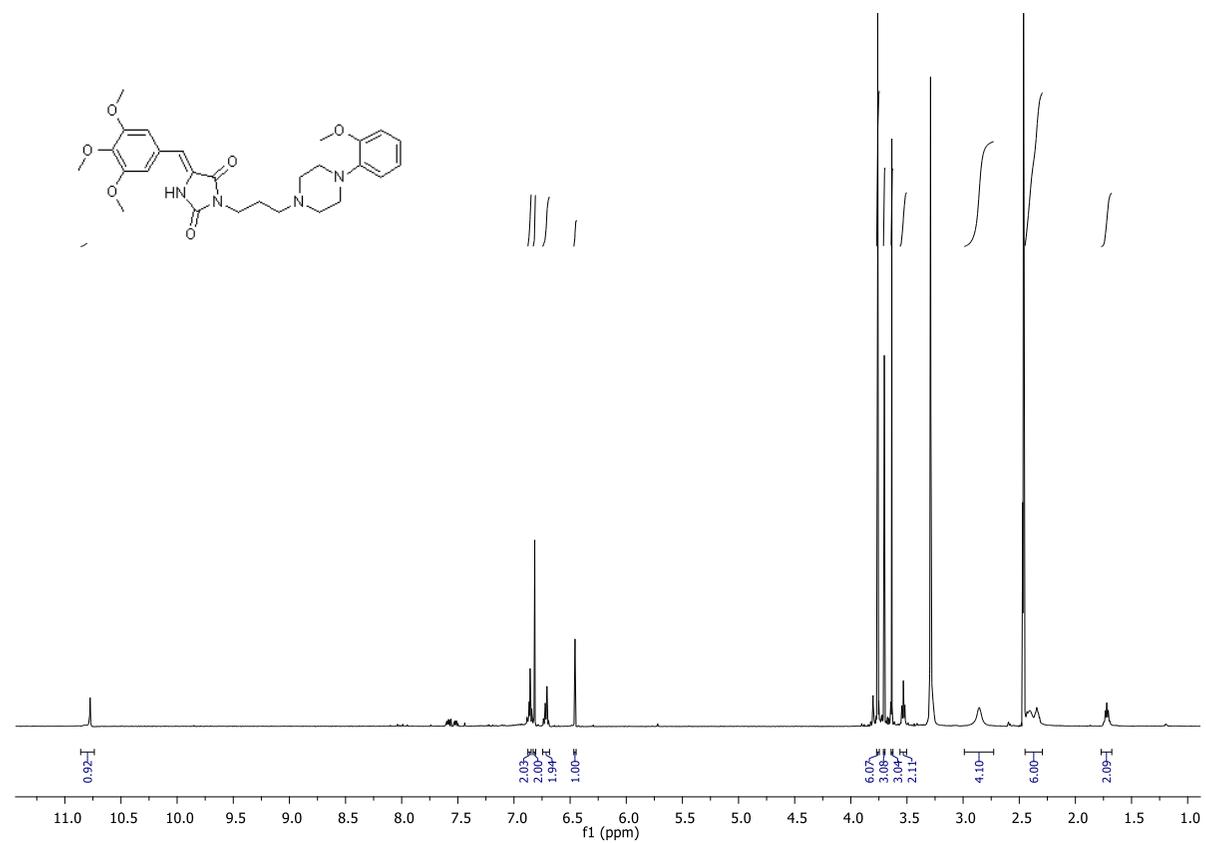


Figure S11. AD33 (Z)-3-(3-(4-(2-ethoxyphenyl)piperazin-1-yl)propyl)-5-(3,4,5-trimethoxybenzylidene)imidazolidine-2,4-dione (**16**)

$^1\text{H}$  NMR (500 MHz, DMSO- $d_6$ )

