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SUPPLEMENTARY MATERIAL

The preparation of macrocyclic calpain inhibitors by RCM and CM

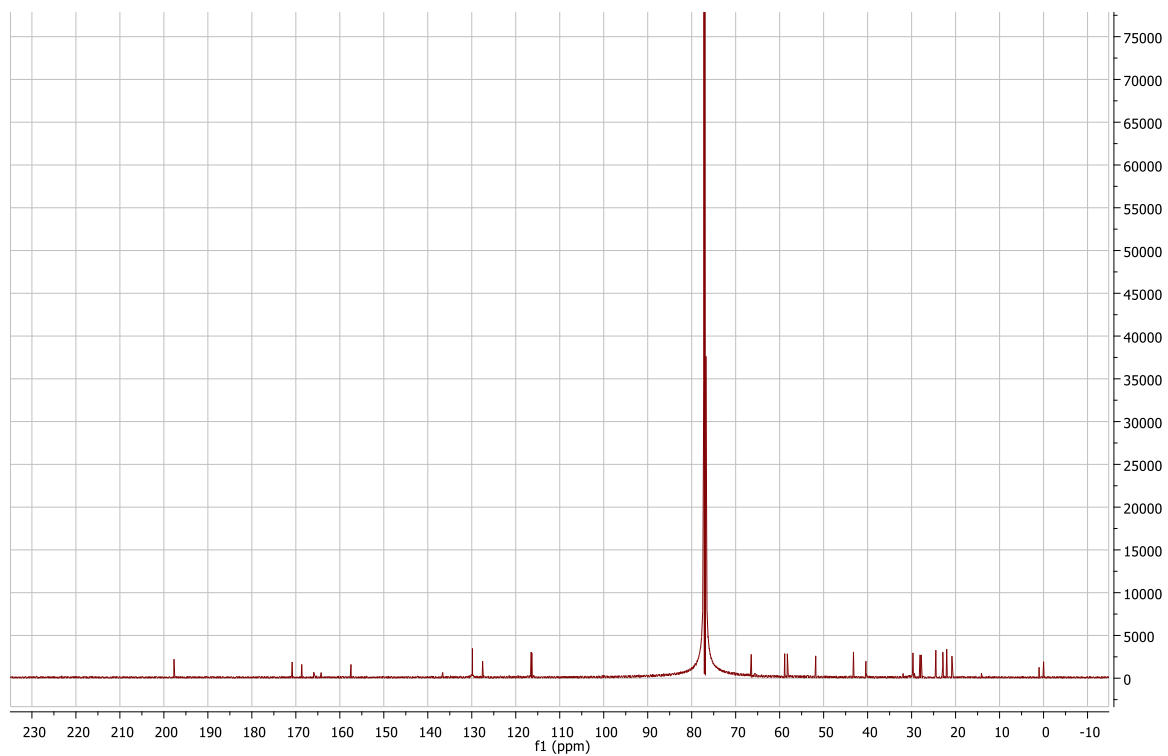
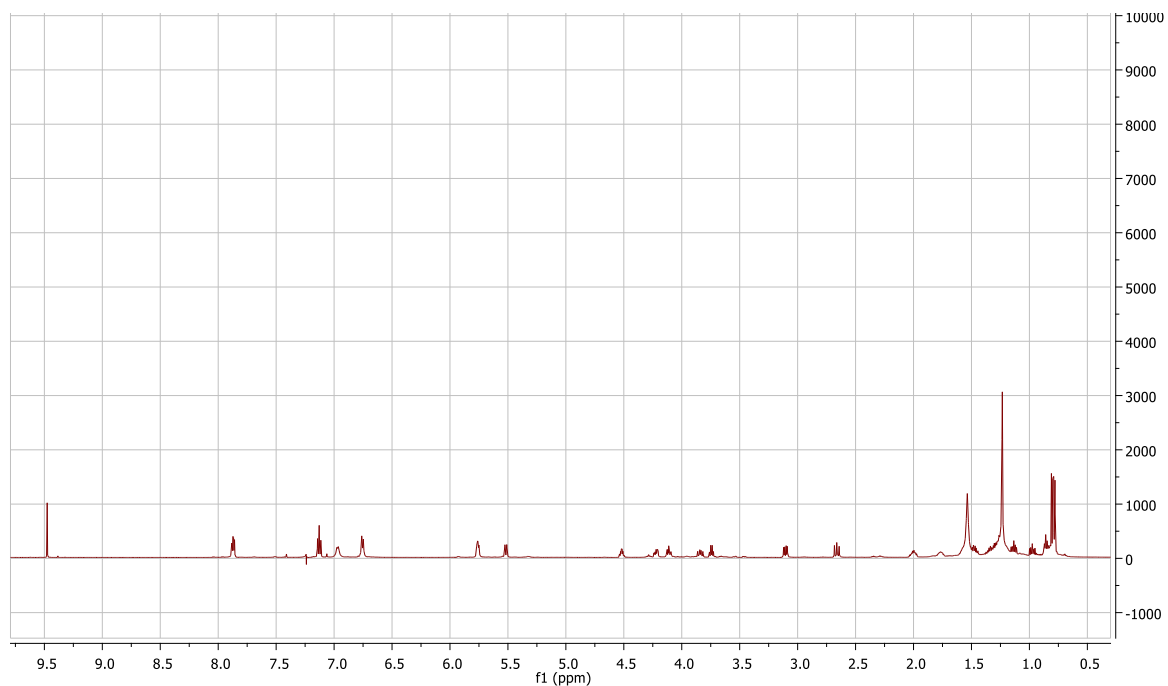
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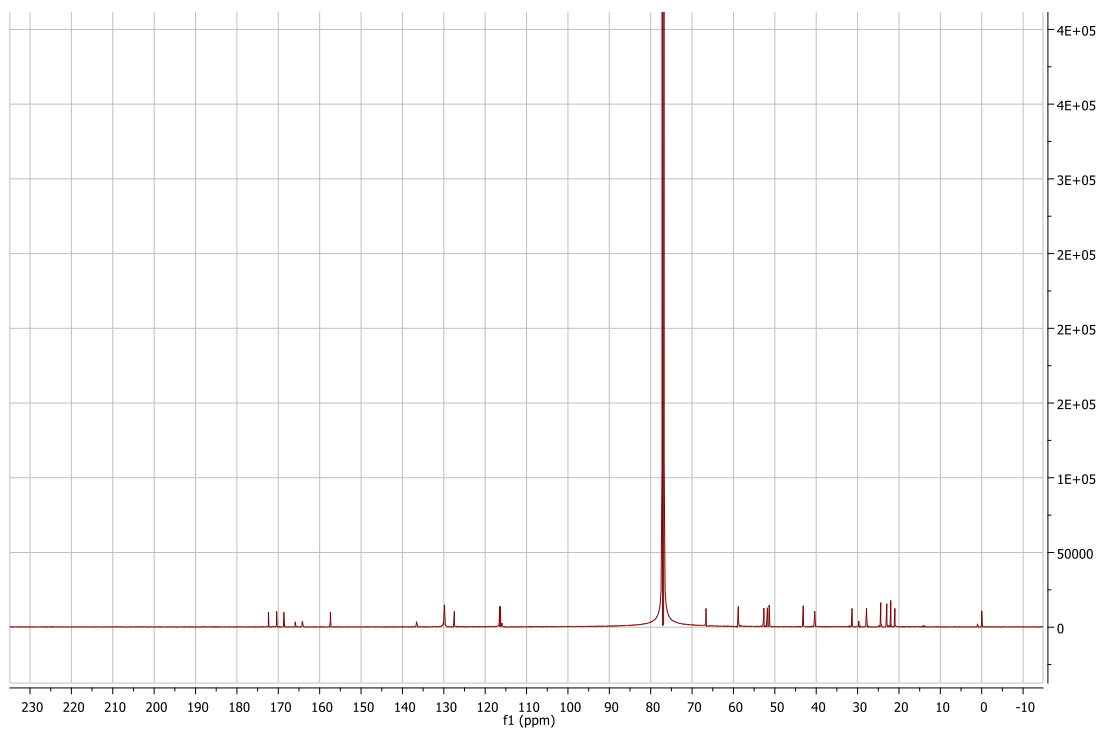
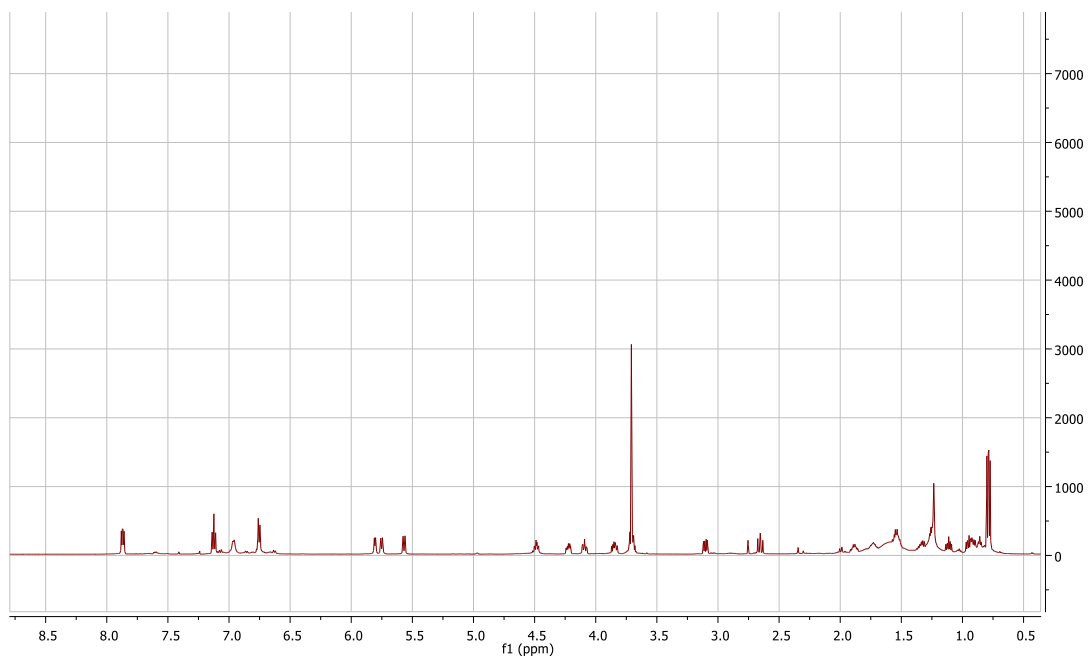
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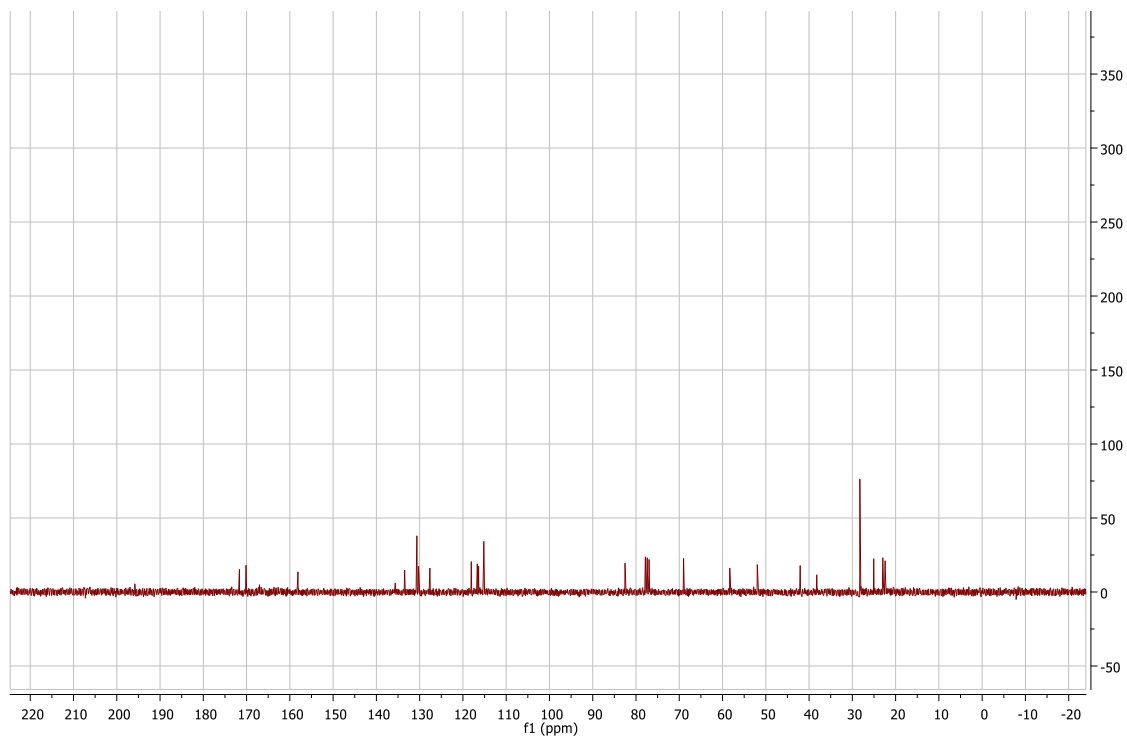
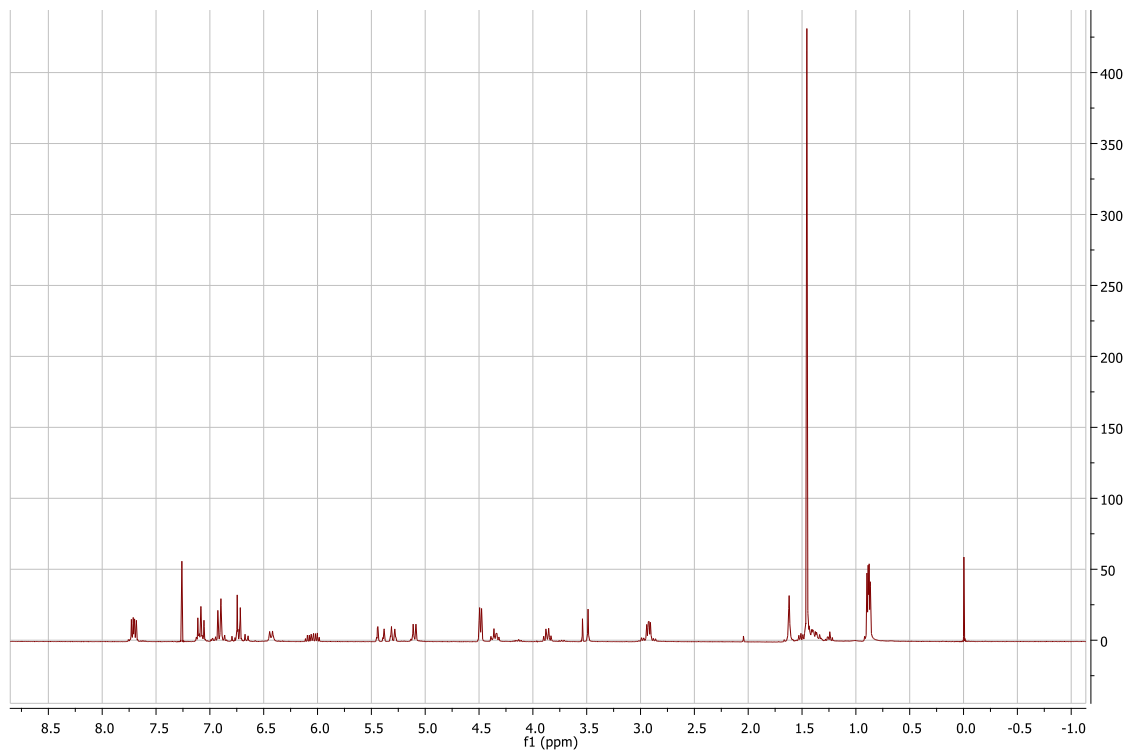
4-Fluoro-N-((7S,10S,13S)-7-formyl-10-isobutyl-9,12-dioxo-2-oxa-8,11-diaza-bicyclo[13.2.2]-nonadeca-1(18),15(19),16-trien-13-yl)-benzenesulfonamide (2)



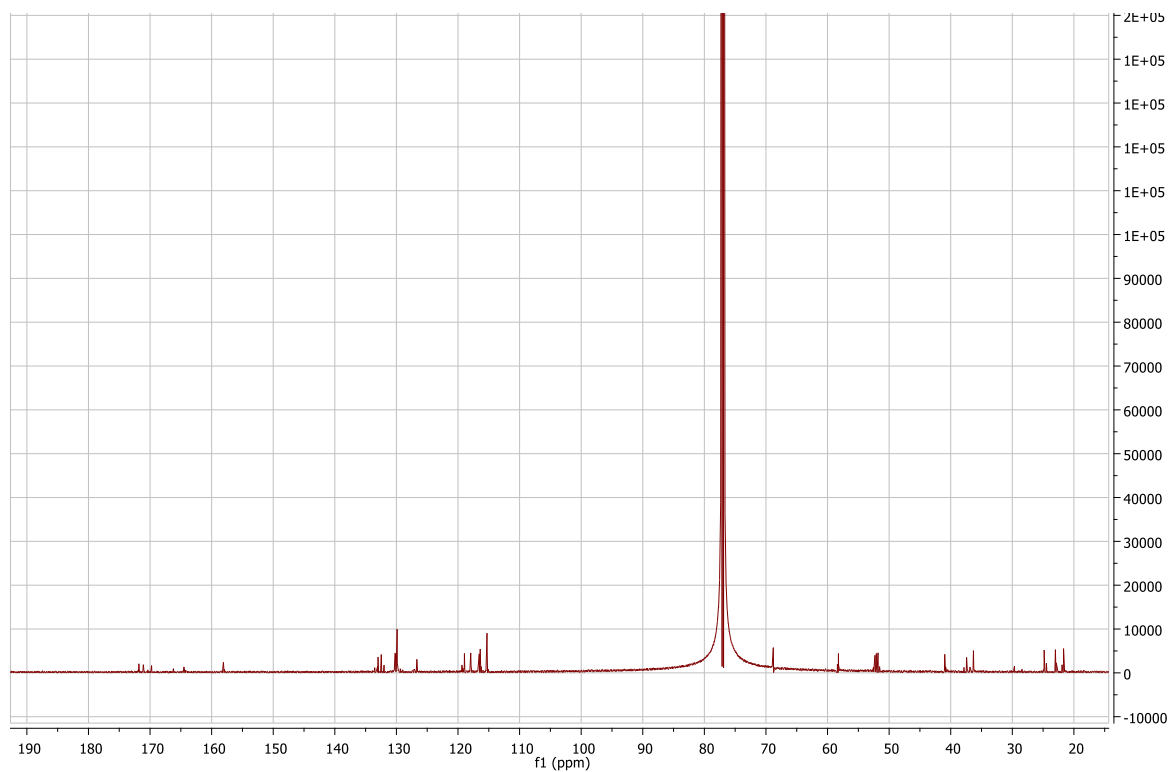
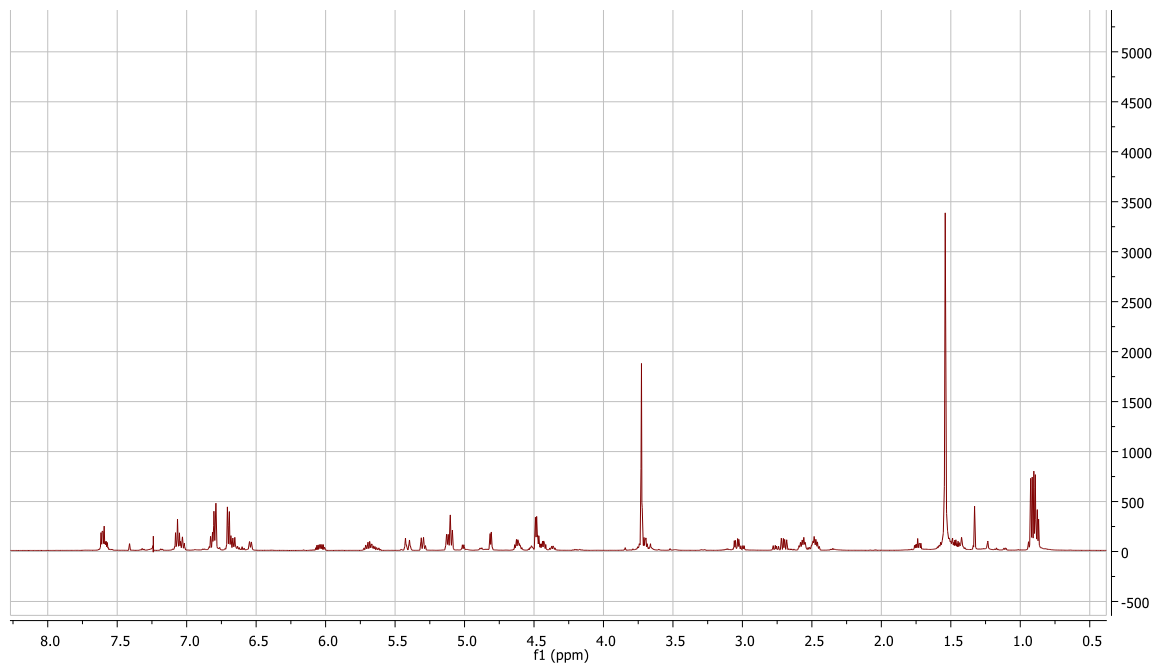
(7S,10S,13S)-13-(4-Fluoro-benzenesulfonylamino)-10-isobutyl-9,12-dioxo-2-oxa-8,11-diaza-bicyclo[13.2.2]nonadeca-1(18),15(19),16-triene-7-carboxylic acid methyl ester (3)



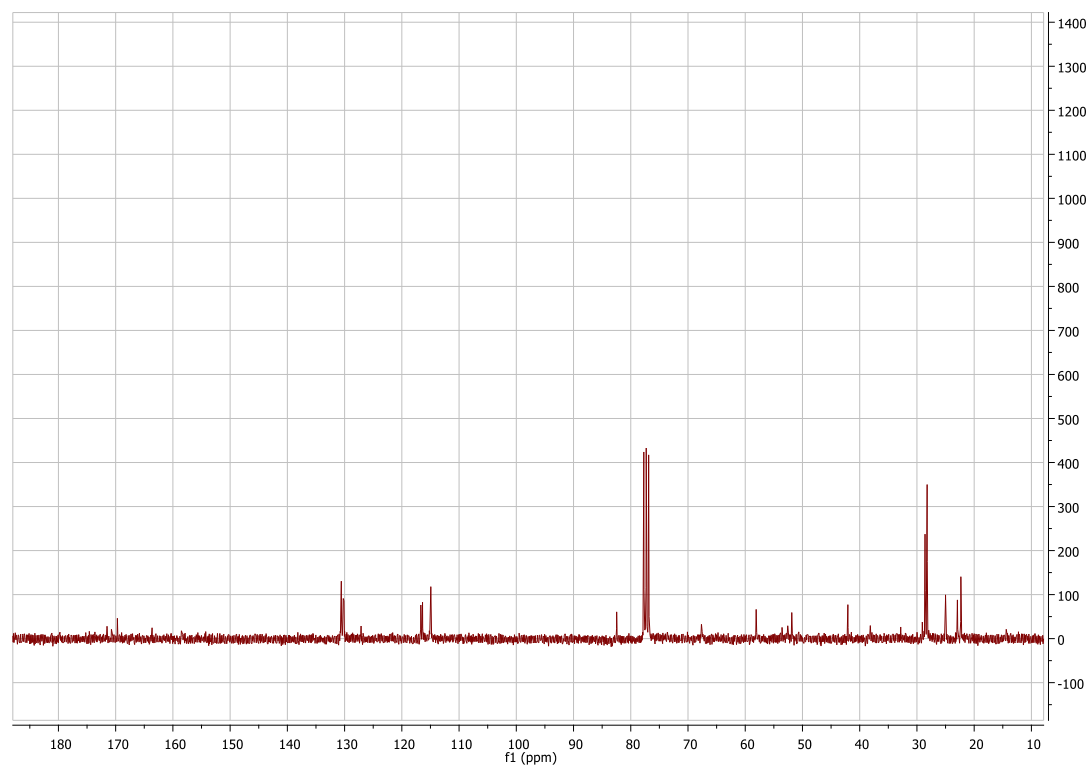
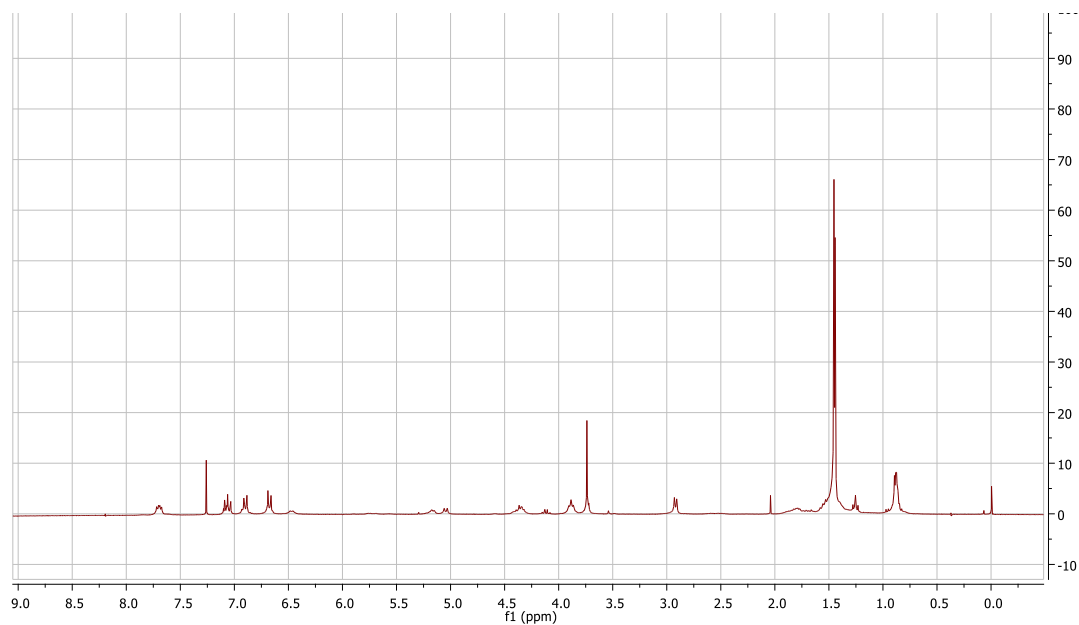
(S)-2-[(*S*)-3-(4-Allyloxy-phenyl)-2-(4-fluoro-benzenesulfonylamino)-propionylamino]-4-methyl-pentanoic acid tert-butyl ester (**6**)



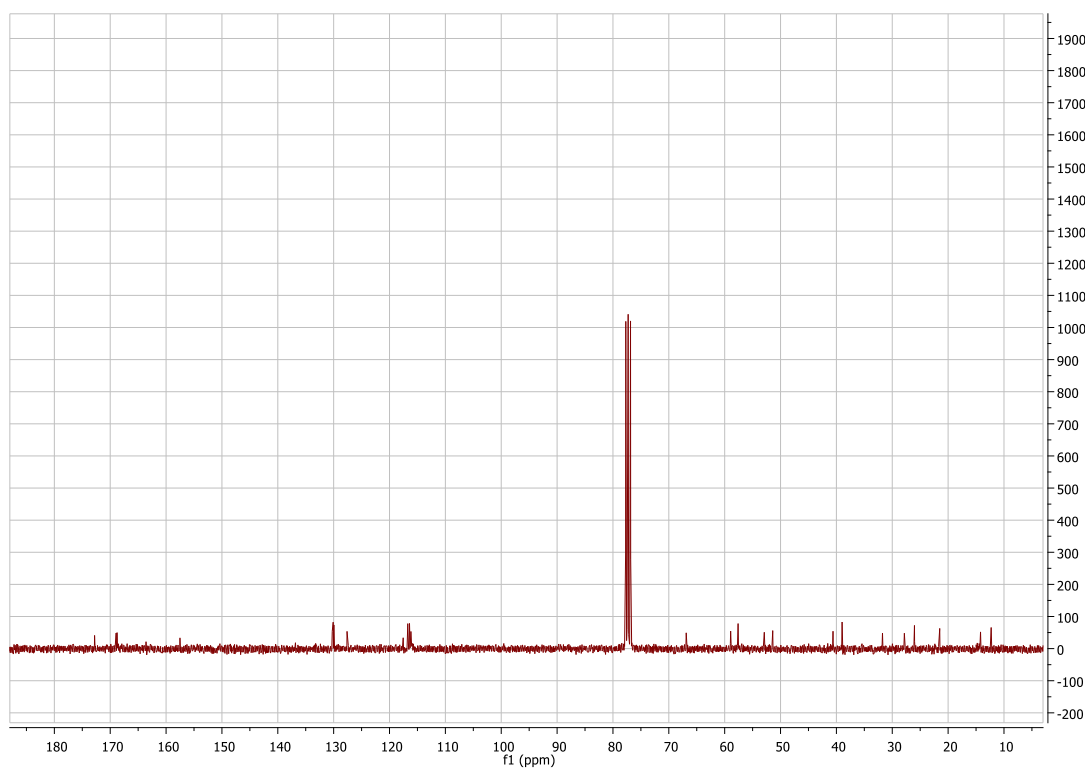
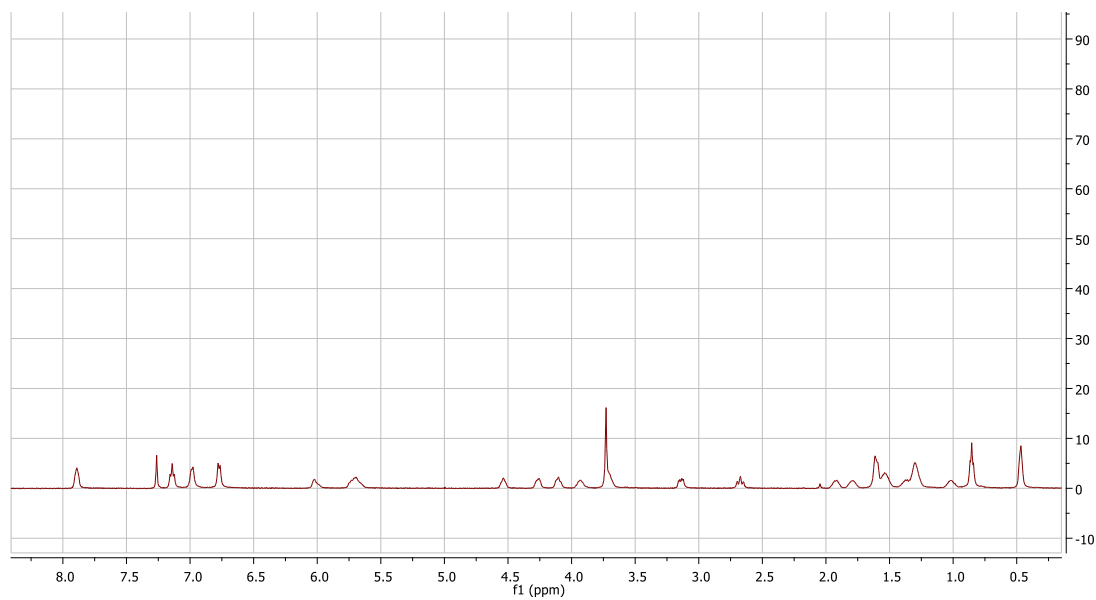
(S)-2-*{(S)*-2-*[(S)*-3-*(-4-Allyloxy-phenyl)*-2-*(4-fluoro-benzenesulfonylamino)*-*propionylamino]*-4-methyl-pentanoylamino}*}-pent-4-enoic acid methyl ester (7)*



(R)-*tert*-Butyl 6-(4-((*S*)-3-((*S*)-1-*tert*-butoxy-4-methyl-1-oxopentan-2-ylamino)-2-(4-fluorophenylsulfonamido)-3-oxopropyl)phenoxy)-2-(*tert*-butoxycarbonylamino)hexanoate (**12**)



(7S,10S,13S)-10-sec-Butyl-13-(4-fluoro-benzenesulfonylamino)-9,12-dioxo-2-oxa-8,11-diaza-bicyclo[13.2.2]nonadeca-1(18),15(19),16-triene-7-carboxylic acid methyl ester (15)



N-((7S,10S,13S)-10-sec-Butyl-7-formyl-9,12-dioxo-2-oxa-8,11-diaza-bicyclo[13.2.2]nonadeca-1(18),15(19),16-trien-13-yl)-4-fluoro-benzenesulfonamide

(16)

