

## Supplementary Material

### Diacetylcurcumin: Its potential antiarthritic effect on a Freund's complete adjuvant-induced murine model

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**Table S1.** The effect of curcumin and DAC on the edema induced by Freund's complete adjuvant on a murine model (acute phase).

Time (hours)	Dose (mg/kg)	Sample	% Inhibition ± SEM	Sample	% Inhibition ± SEM
4	80	Phenylbutazone	<b>53.65±0.04**</b>	Phenylbutazone	<b>58.14±0.04***</b>
4	60	Curcumin	26.56±0.04	DAC	<b>37.21±0.04*</b>
4	120	Curcumin	<b>41.67±0.03*</b>	DAC	<b>47.67±0.02**</b>
4	150	Curcumin	29.69±0.03	DAC	<b>51.55±0.02**</b>
8	80	Phenylbutazone	<b>41.43±0.04*</b>	Phenylbutazone	<b>35.48±0.02*</b>
8	60	Curcumin	14.74±0.04	DAC	20.82±0.05
8	120	Curcumin	33.47±0.04	DAC	21.85±0.03
8	150	Curcumin	29.48±0.4	DAC	29.82±0.05
24	80	Phenylbutazone	<b>35.47±0.03***</b>	Phenylbutazone	<b>35.22±0.02*</b>
24	60	Curcumin	7.65±0.04	DAC	18.51±0.03
24	120	Curcumin	<b>25.08±0.02*</b>	DAC	19.10±0.03
24	150	Curcumin	<b>26.30±0.05*</b>	DAC	15.22±0.03

\* Data were analyzed by ANOVA followed by Tukey's test.  $P < 0.05$ ,  $0.01$  and  $0.001$  (\*, \*\* and \*\*\* respectively).

**Table S2.** The effect of curcumin and DAC on the edema induced by Freund's complete adjuvant on a murine model (chronic phase).

Time (days)	Dose (mg/kg)	Curcumin experiment	% Inhibition ± SEM	Diacetylcurcumin experiment	% Inhibition ± SEM
17	80	Phenylbutazone	<b>42.7±0.04*</b>	Phenylbutazone	<b>41.39±0.05*</b>
17	60	Curcumin	6.7±0.04	DAC	17.50±0.02
17	120	Curcumin	28.1±0.05	DAC	<b>43.33±0.09*</b>
17	150	Curcumin	<b>43.9±0.07*</b>	DAC	<b>46.39±0.03**</b>
18	80	Phenylbutazone	47.29±0.04	Phenylbutazone	<b>42.75±0.05*</b>
18	60	Curcumin	20.27±0.04	DAC	28.76±0.02
18	120	Curcumin	37.83±0.06	DAC	<b>46.11±0.08*</b>
18	150	Curcumin	<b>51.35±0.07*</b>	DAC	<b>50.00±0.03**</b>
19	80	Phenylbutazone	<b>45.0±0.05*</b>	Phenylbutazone	<b>44.04±0.06**</b>
19	60	Curcumin	14.5±0.05	DAC	31.87±0.02
19	120	Curcumin	34.0±0.07	DAC	<b>45.99±0.06**</b>
19	150	Curcumin	<b>51.1±0.07*</b>	DAC	<b>50.85±0.02**</b>
20	80	Phenylbutazone	<b>46.0±0.05*</b>	Phenylbutazone	<b>43.91±0.05**</b>
20	60	Curcumin	14.6±0.05	DAC	27.41±0.02
20	120	Curcumin	33.8±0.07	DAC	<b>46.19±0.06**</b>
20	150	Curcumin	<b>49.8±0.07*</b>	DAC	<b>51.27±0.03***</b>
21	80	Phenylbutazone	<b>48.1±0.06*</b>	Phenylbutazone	<b>43.40±0.06**</b>
21	60	Curcumin	17.3±0.05	DAC	28.68±0.02
21	120	Curcumin	35.6±0.06	DAC	<b>44.42±0.07**</b>
21	150	Curcumin	<b>51.2±0.07*</b>	DAC	<b>50.76±0.03**</b>
22	80	Phenylbutazone	<b>48.6±0.05*</b>	Phenylbutazone	<b>47.31±0.04**</b>
22	60	Curcumin	16.0±0.05	DAC	<b>34.27±0.02*</b>
22	120	Curcumin	34.0±0.07	DAC	<b>47.57±0.06**</b>
22	150	Curcumin	<b>52.4±0.08*</b>	DAC	<b>50.38±0.04***</b>
23	80	Phenylbutazone	<b>49.1±0.05*</b>	Phenylbutazone	<b>44.09±0.04**</b>
23	60	Curcumin	14.7±0.05	DAC	<b>31.72±0.03*</b>
23	120	Curcumin	34.1±0.07	DAC	<b>45.16±0.07**</b>
23	150	Curcumin	<b>48.5±0.07*</b>	DAC	<b>48.66±0.04***</b>
24	80	Phenylbutazone	<b>49.3±0.05*</b>	Phenylbutazone	<b>42.63±0.05**</b>
24	60	Curcumin	15.1±0.05	DAC	<b>31.10±0.03*</b>
24	120	Curcumin	34.6±0.07	DAC	<b>39.68±0.07**</b>
24	150	Curcumin	<b>49.0±0.07*</b>	DAC	<b>47.99±0.04***</b>
25	80	Phenylbutazone	<b>48.4±0.05*</b>	Phenylbutazone	<b>42.44±0.05**</b>
25	60	Curcumin	13.8±0.05	DAC	<b>32.63±0.02*</b>
25	120	Curcumin	34.3±0.07	DAC	<b>41.11±0.06**</b>
25	150	Curcumin	<b>49.5±0.07*</b>	DAC	<b>49.07±0.04***</b>

\* Data were analyzed by ANOVA followed by Tukey's test.  $P < 0.05$ ,  $0.01$  and  $0.001$  (\*, \*\* and \*\*\* respectively).

LURMN\_0149\_curcumina1.fid  
Instituto de Química, UNAM (BQC)  
Dr. R. Enriquez / W. Meza  
Clave: curcumina  
No. de Registro: LURMN\_0149  
Experimento: 1H  
Disolvente: CDCl<sub>3</sub>  
Bruker AVANCE III HD 500 MHz  
2-septiembre-2016

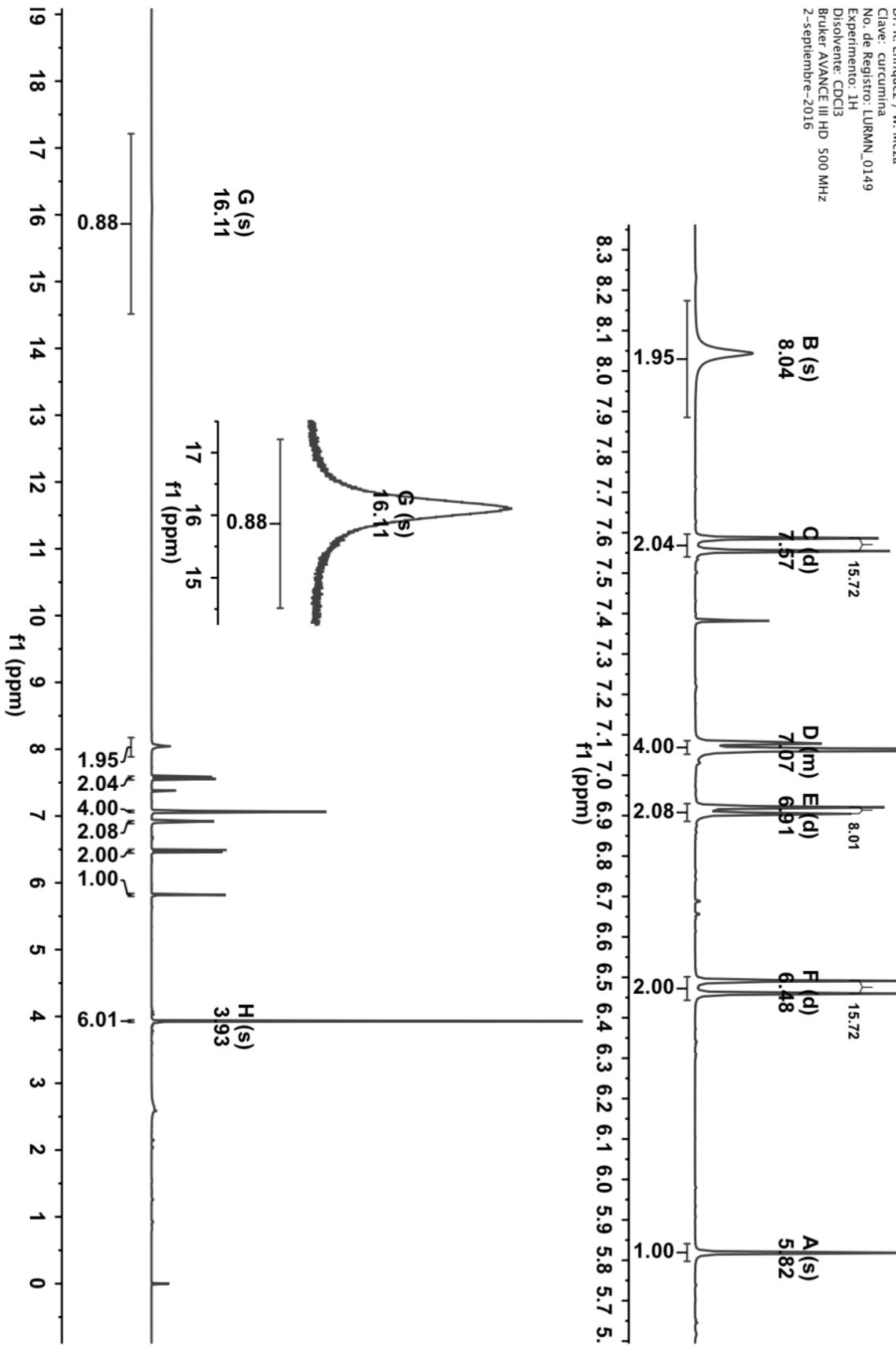


Figure S1: <sup>1</sup>H-NMR of curcumin (I).

LURMN\_0149.curcumina.4.fid  
Instituto de Química, UNAM (BQC)  
Dr. R. Enriquez / W. Meza  
Clave: curcumina  
No. de Registro: LURMN\_0149  
Experimento:  $^{13}\text{C}$   
Disolvente: CDCl<sub>3</sub>  
Bruker AVANCE III HD 500 MHz  
2-septiembre-2016

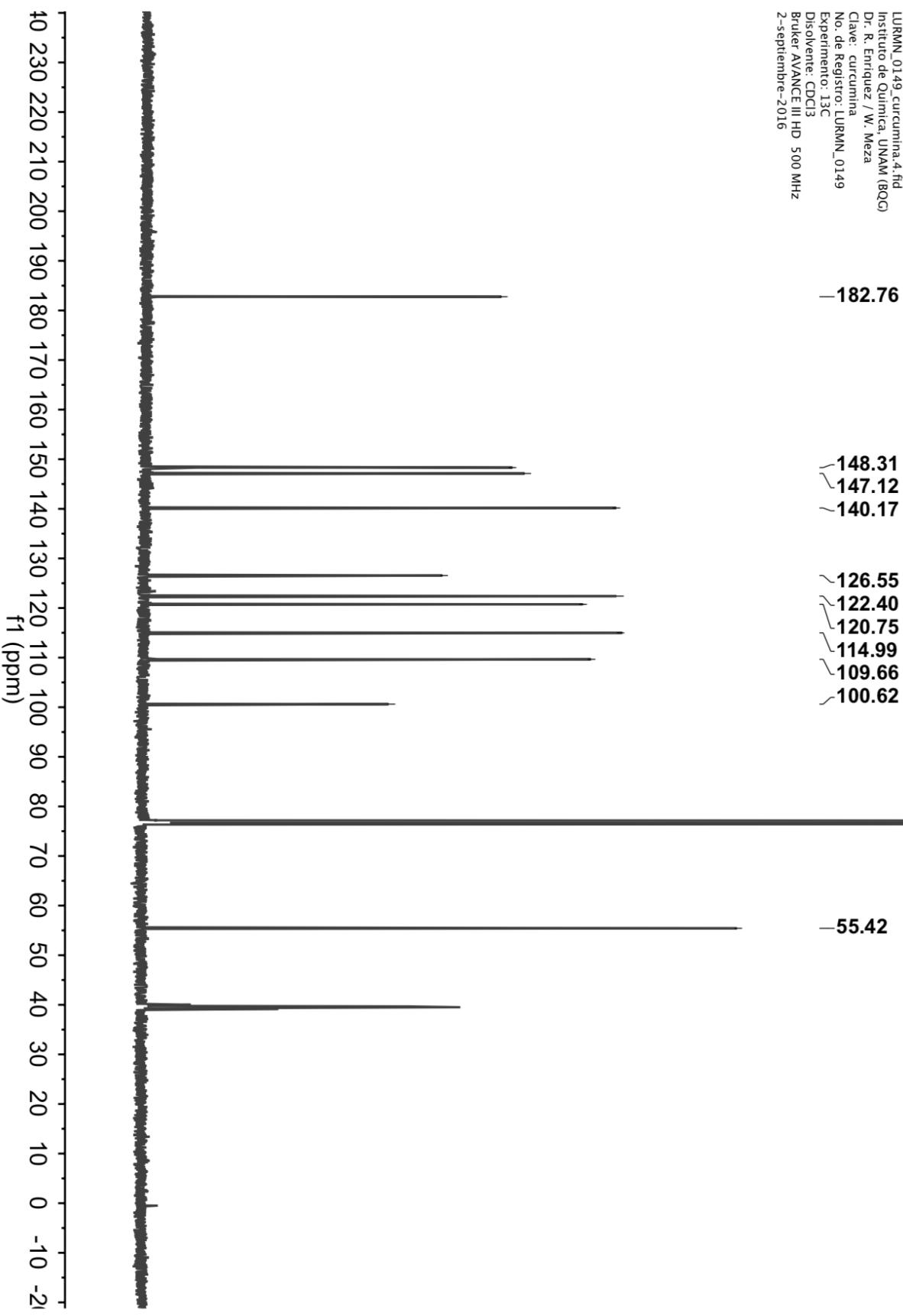


Figure S2:  $^{13}\text{C}$ -NMR of curcumin (1).

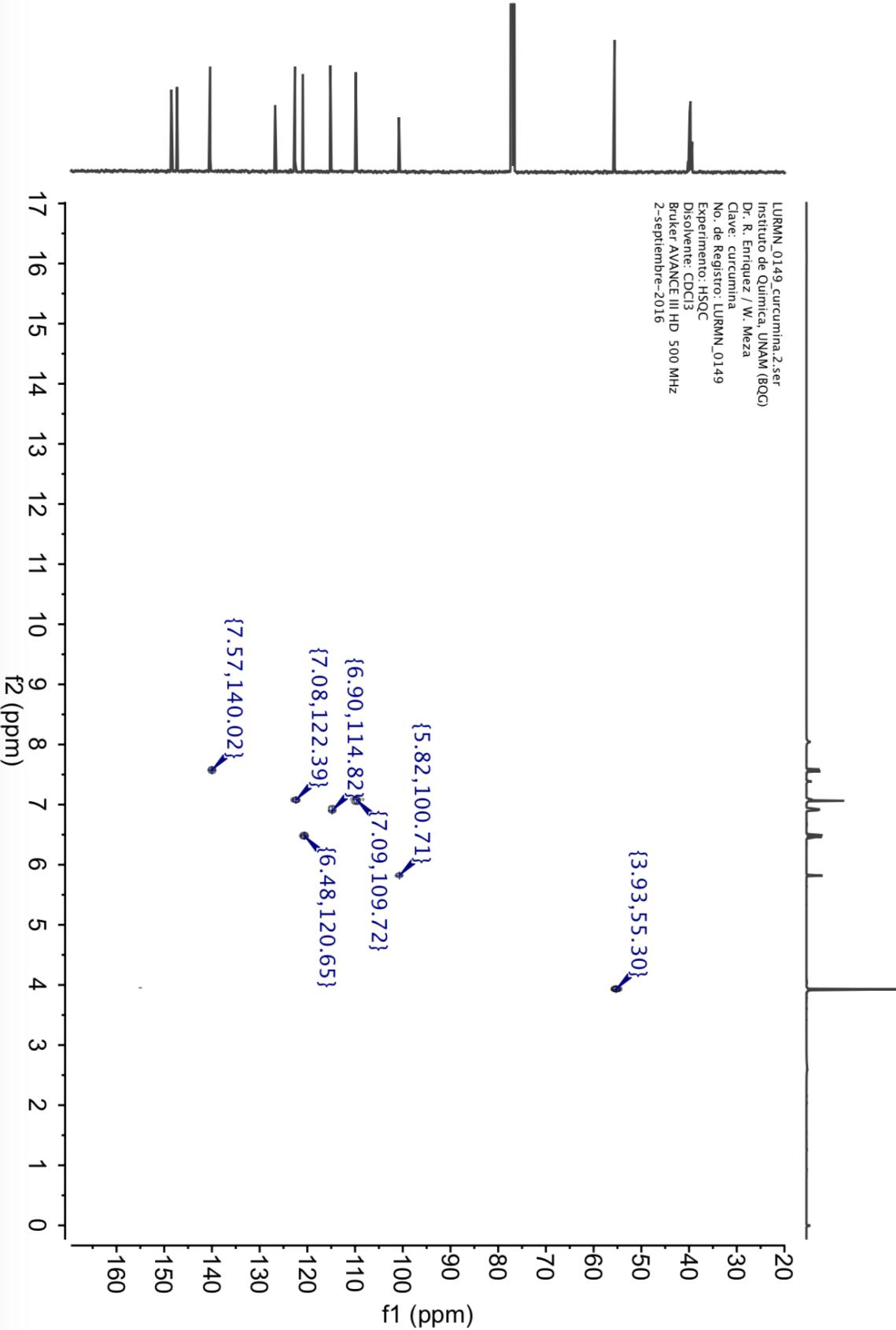


Figure S3: HSQC of curcumin (I).

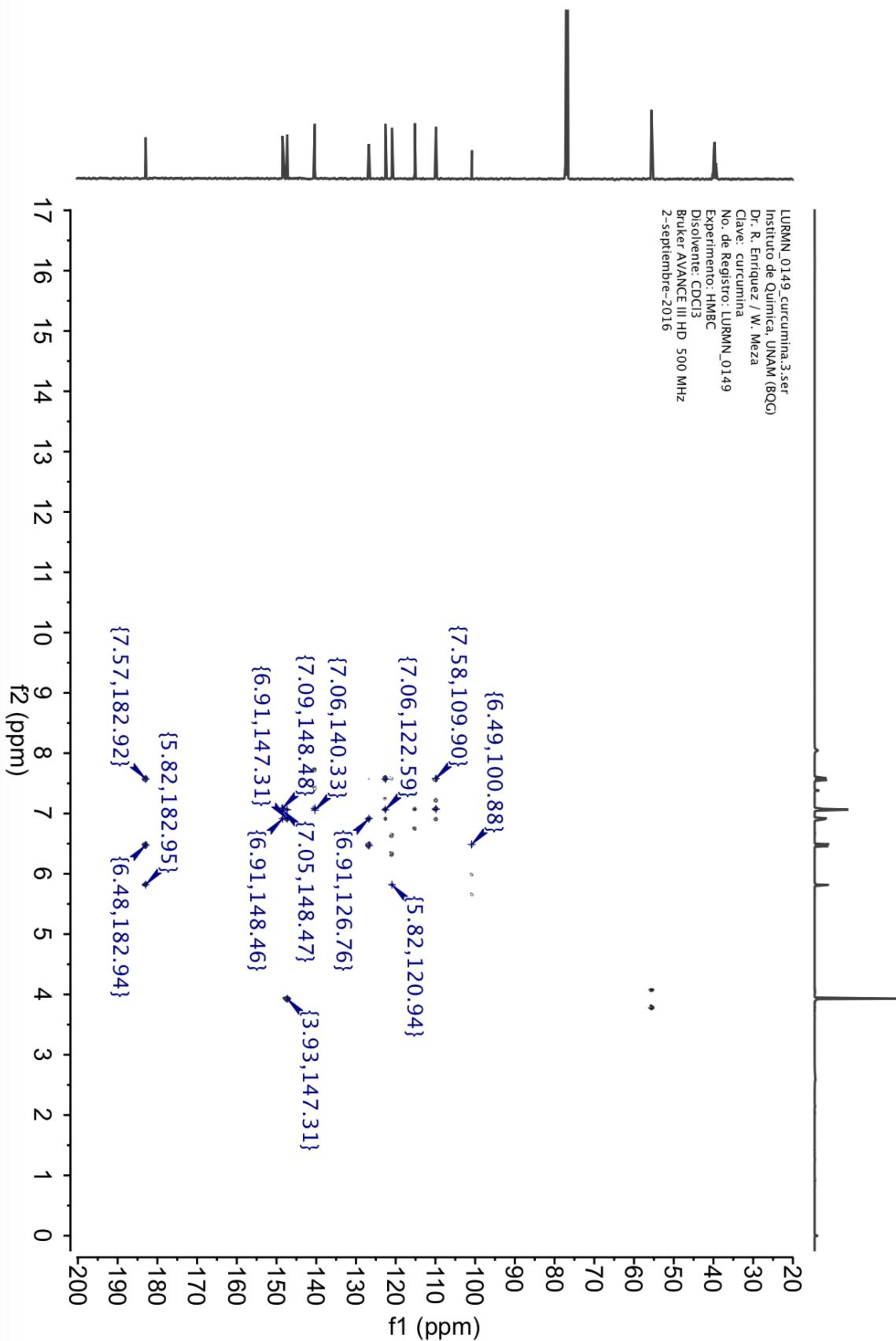


Figure S4: HMBC of curcumin (1).

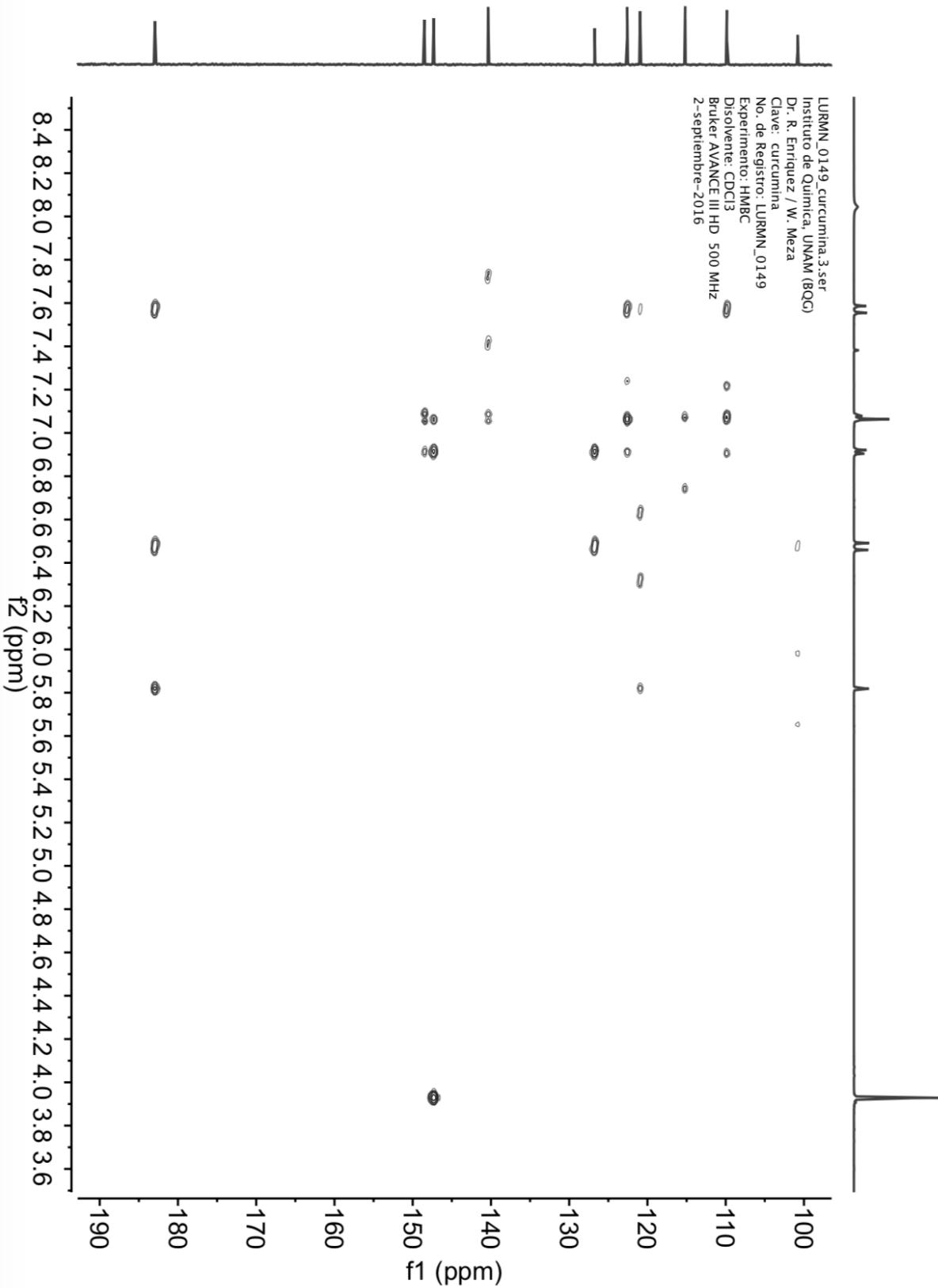


Figure S5: HMBC of curcumin (1).

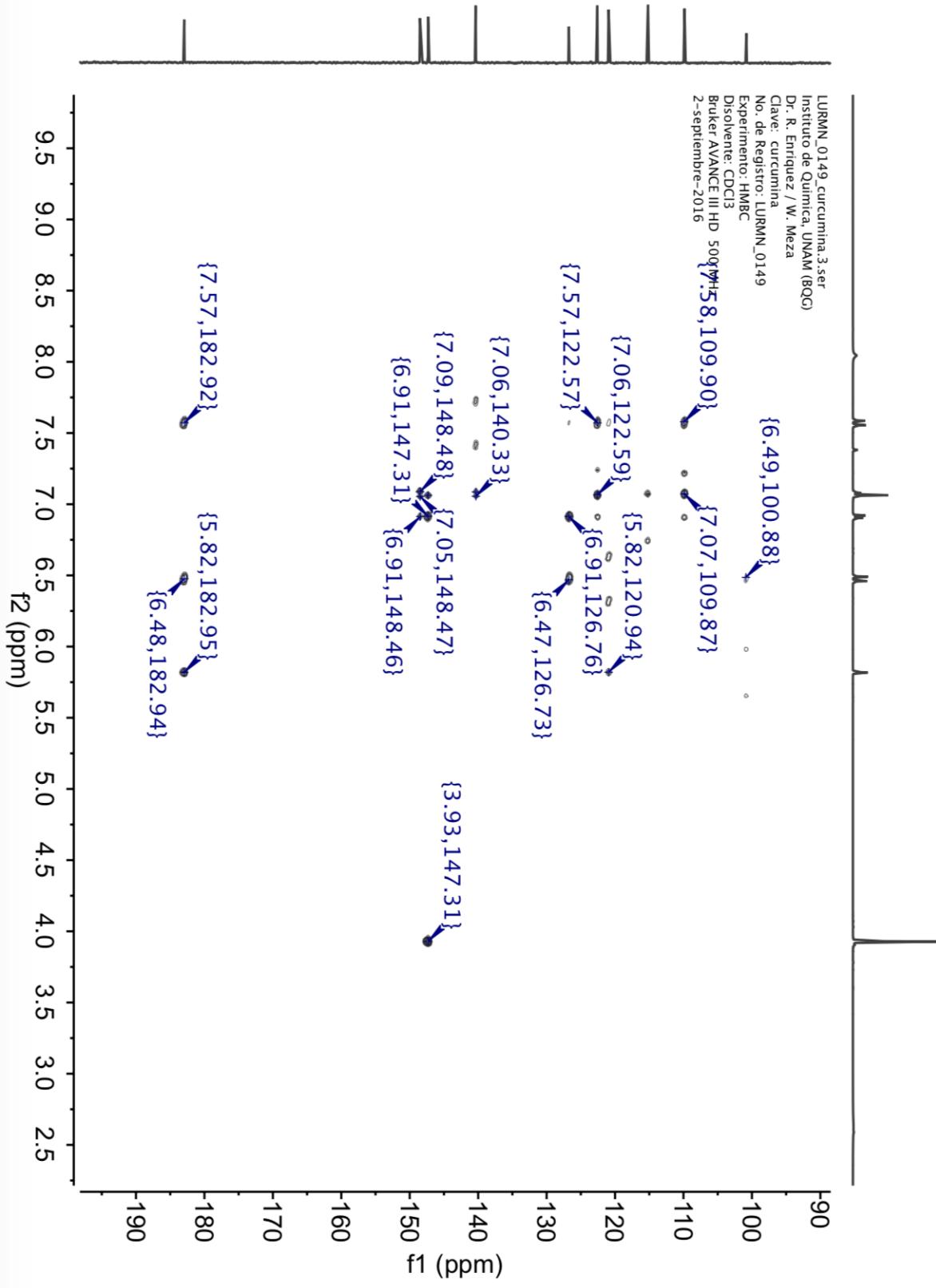


Figure S6: HMBC of curcumin (1).

LURMN\_0150.DAC.1.fid  
Instituto de Química, UNAM (BQC)  
Dr. R. Enriquez / W. Meza  
Clave: DAC  
No. de Registro: LURMN\_0150  
Experimento:  $^1\text{H}$   
Disolvente: CDCl<sub>3</sub>  
Bruker AVANCE III HD 500 MHz  
31-agosto-2016

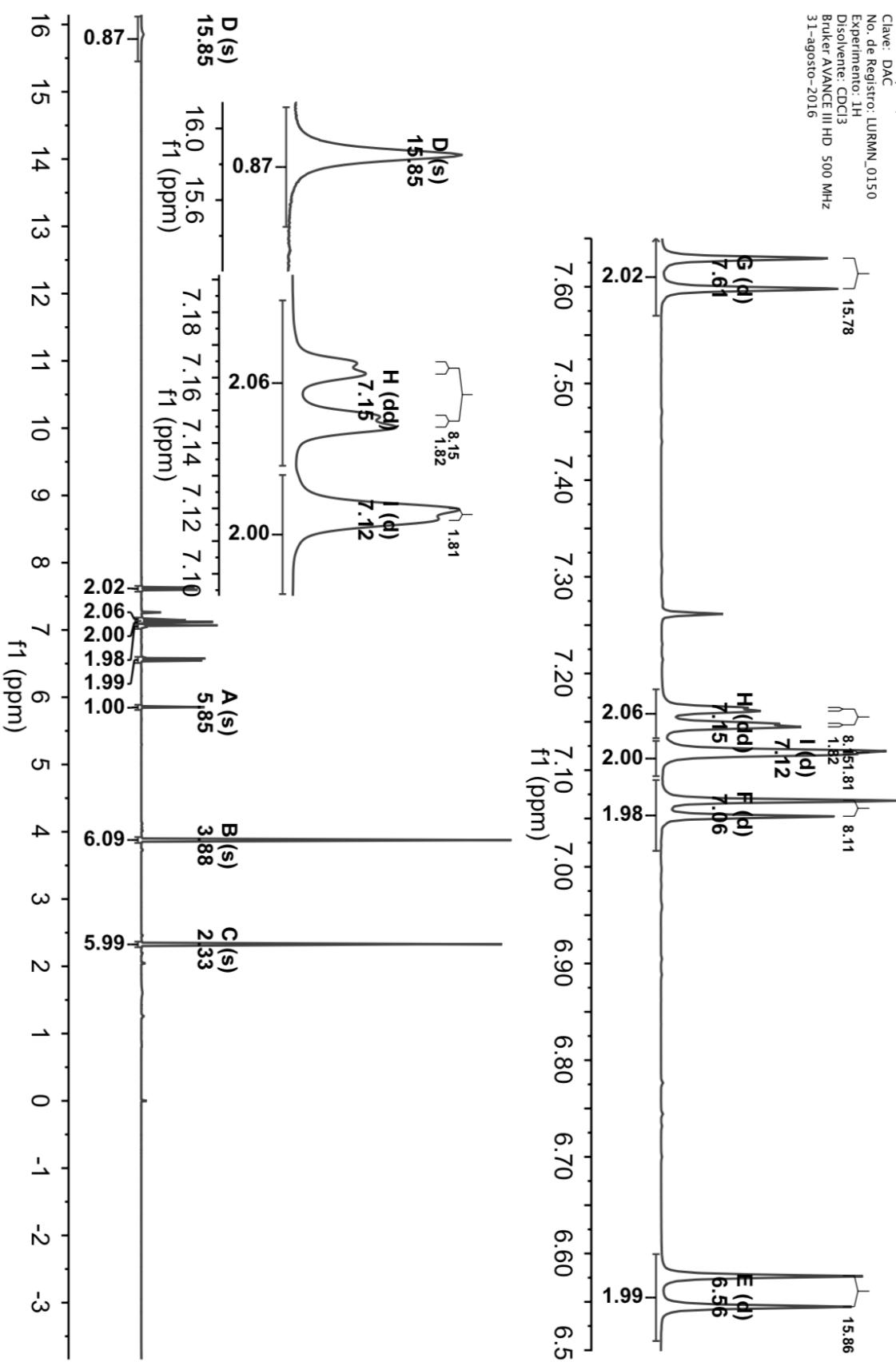


Figure S7:  $^1\text{H}$ -NMR of diacetylcurcumin (2).

LURMN\_0150.DAC.4.fid  
Instituto de Química, UNAM (B)  
Dr. R. Enríquez / W. Meza  
Clave: DAC  
No. de Registro: LURMN\_0150   **183.28**  
Experimento: 13C  
Disolvente: CDCl<sub>3</sub>  
Bruker AVANCE III HD 500 MHz  
31-agosto-2016

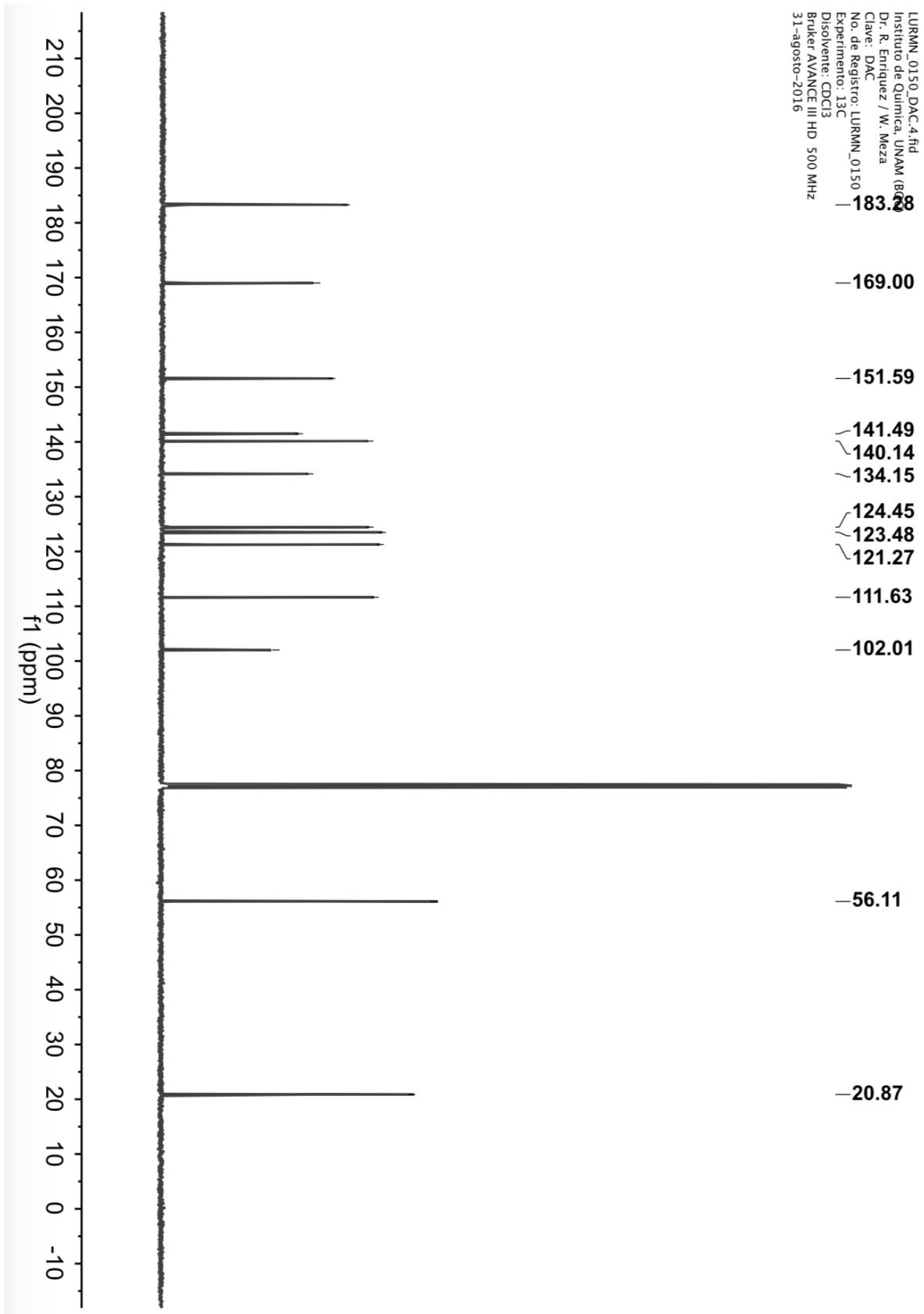


Figure S8: <sup>13</sup>C-NMR of diacetylcurcumin (2).

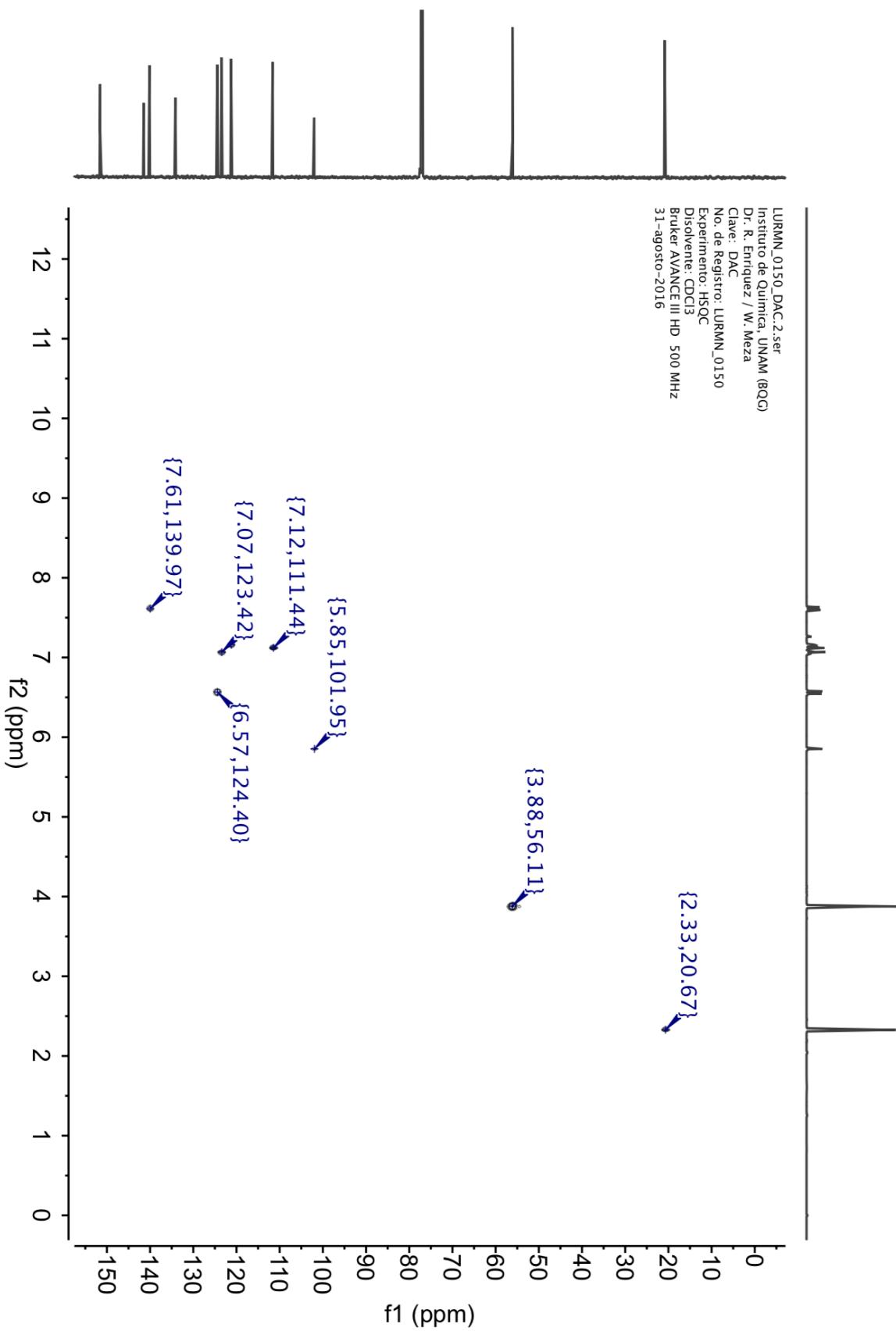


Figure S9: HSQC of diacetylcurcumin (2).

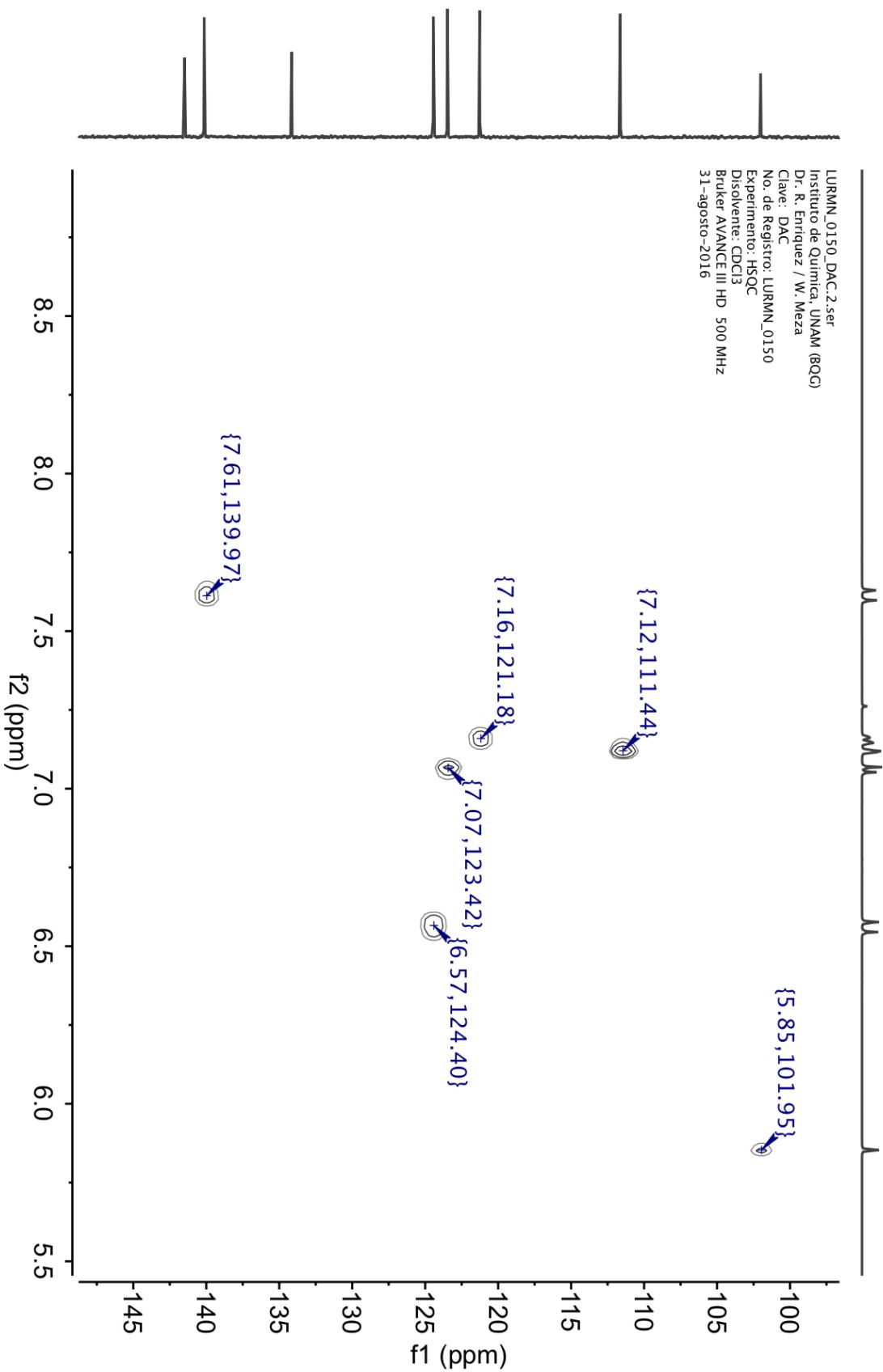


Figure S10: HSQC of diacetylcurcumin (2).

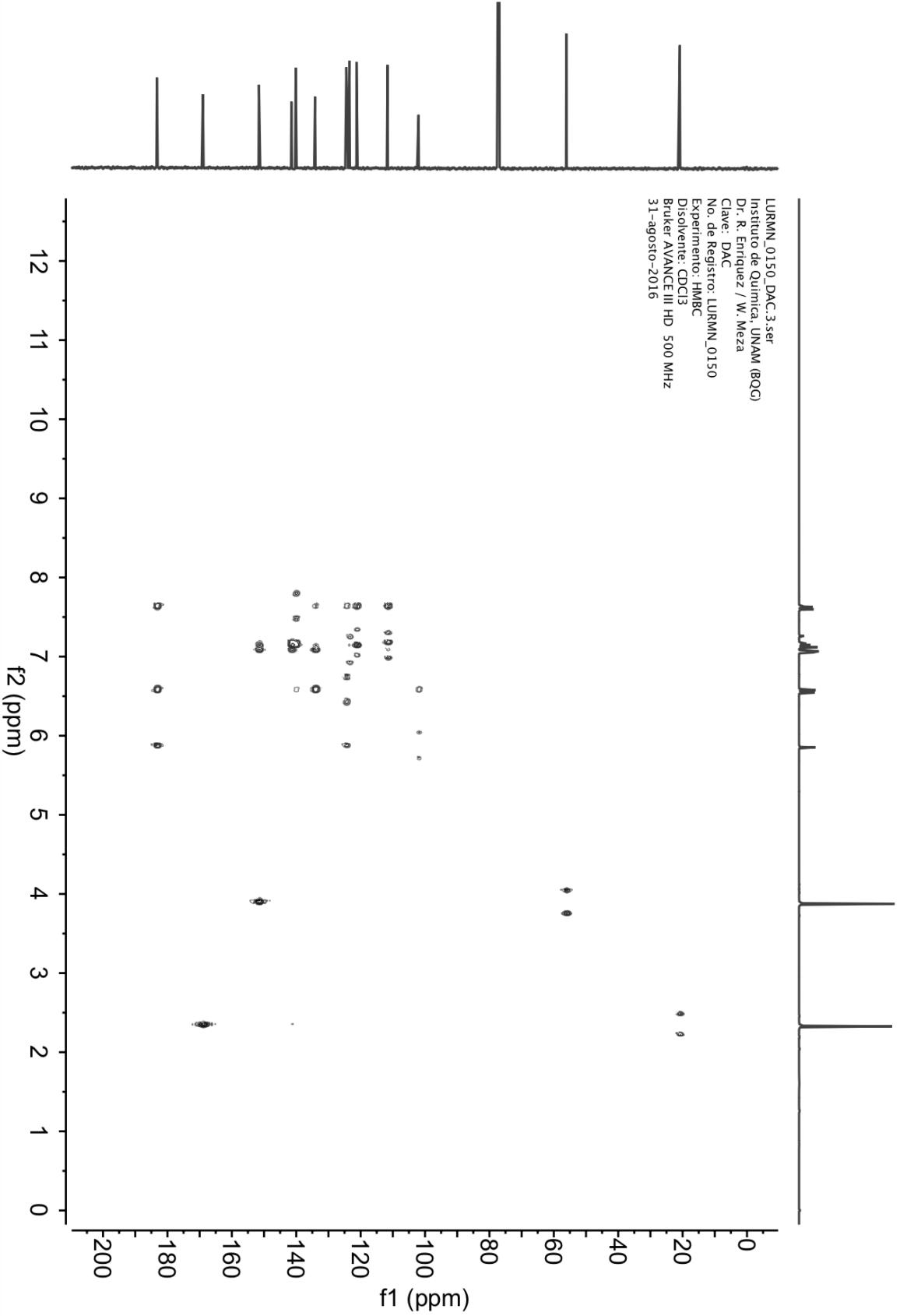


Figure S11: HMBC of diacetylcurcumin (2).

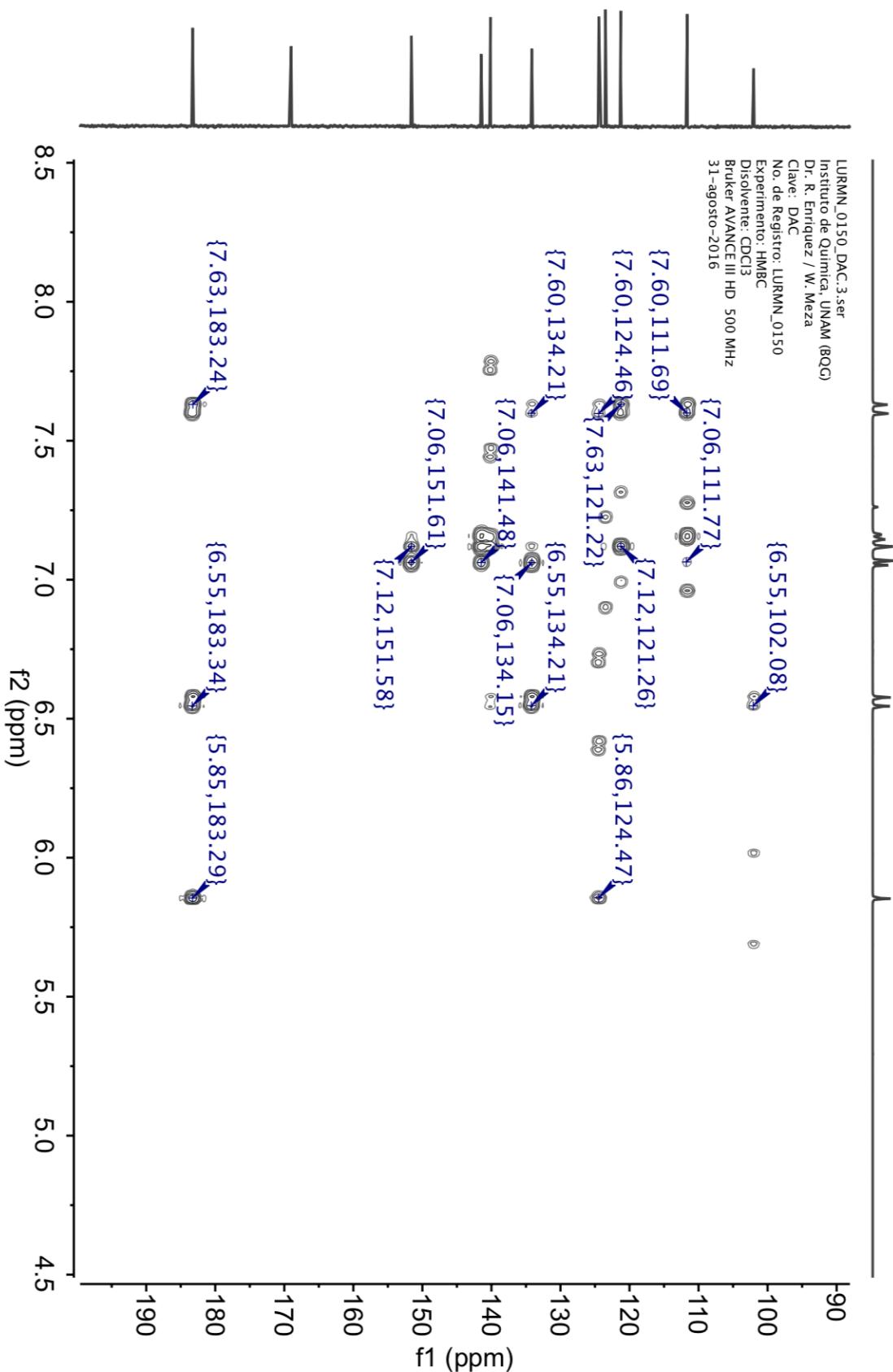


Figure S12: HMBC of diacetylcurcumin (2).