

APPENDIX

APPENDIX A ^1H and ^{13}C NMR Spectra, FT-IR spectra, and HPLC

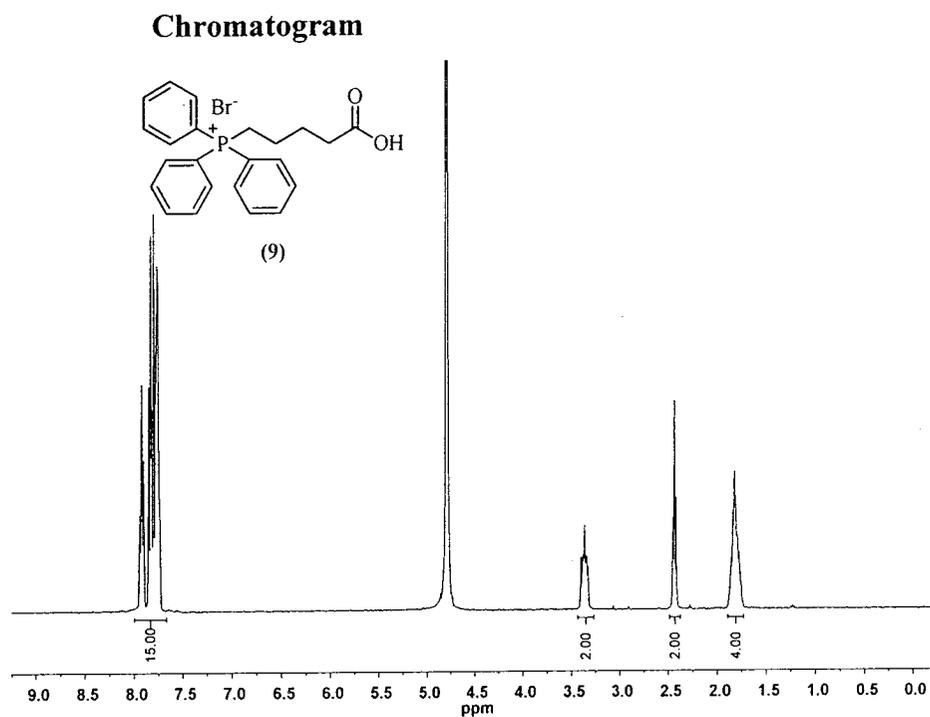


Figure 37 ^1H NMR Spectrum of (4-carboxybutyl)triphenylphosphonium bromide (9) (D_2O)

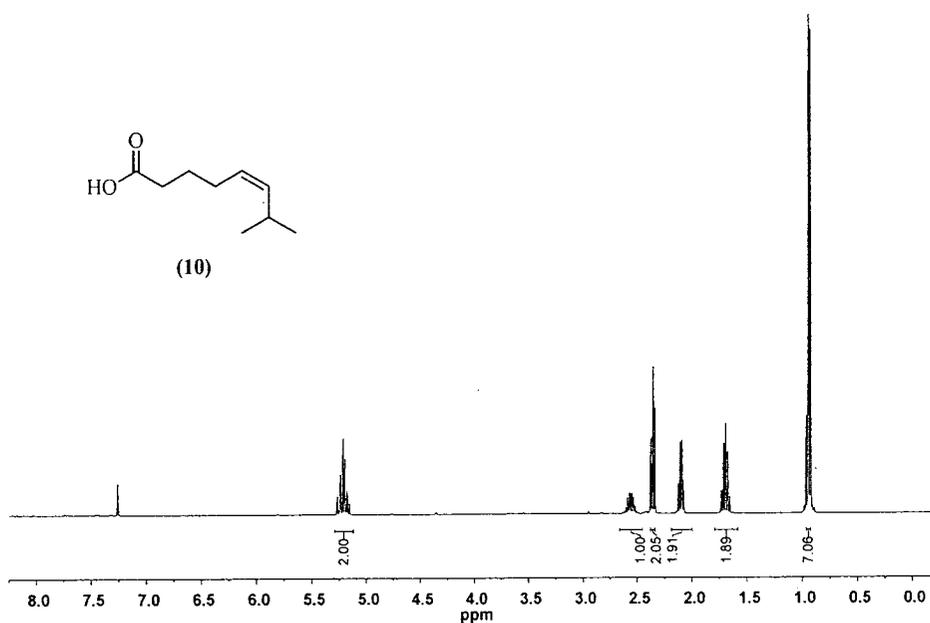


Figure 38 ^1H NMR Spectrum of (Z)-7-methyl-5-octenoic acid (10) (CDCl_3)

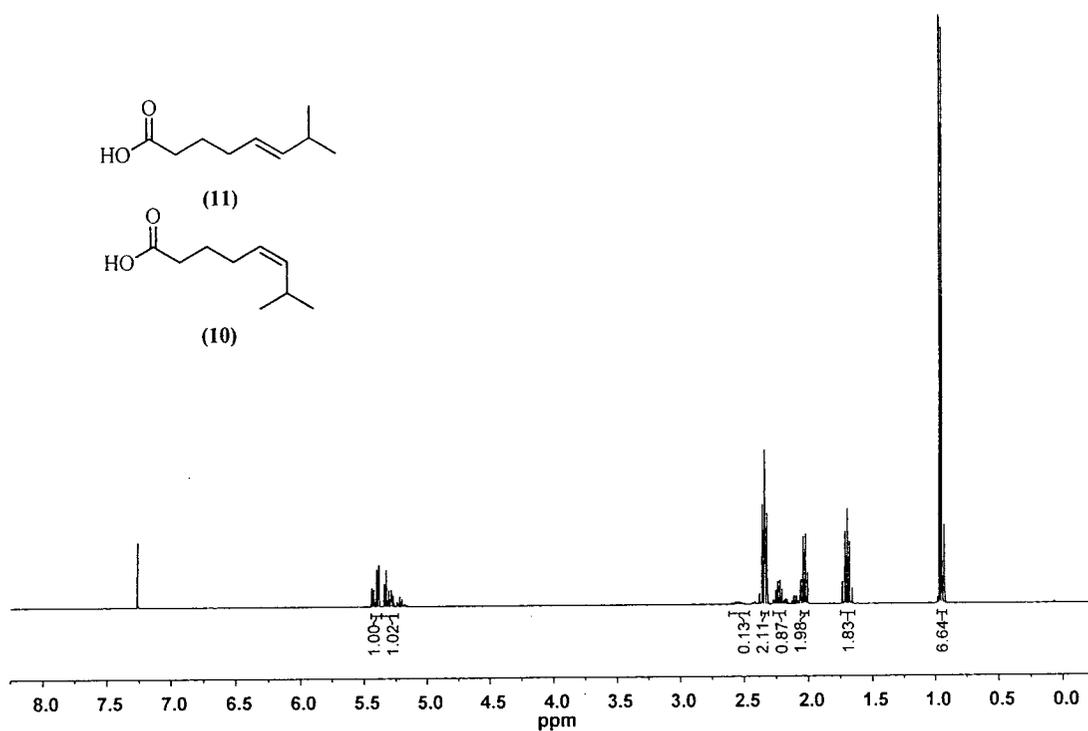


Figure 39 ¹H NMR Spectrum of (*E/Z*)-7-methyl-5-octenoic acid (11, 10) (CDCl₃)

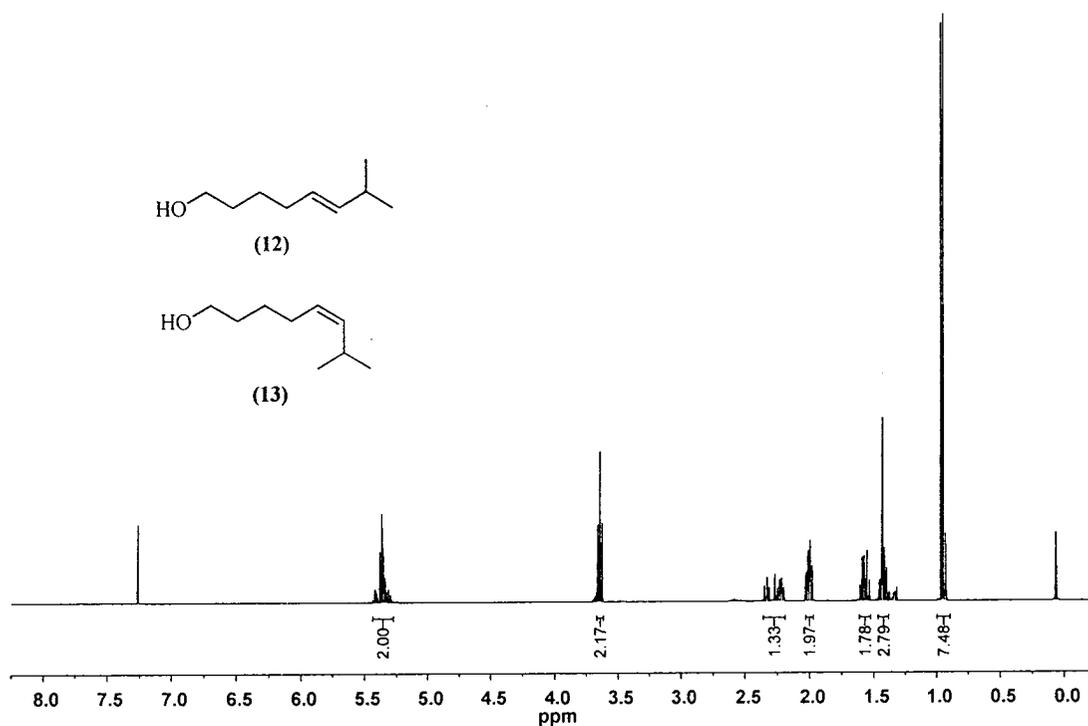


Figure 40 ¹H NMR Spectrum of (*E/Z*)-7-methyl-5-octenol (12, 13) (CDCl₃)

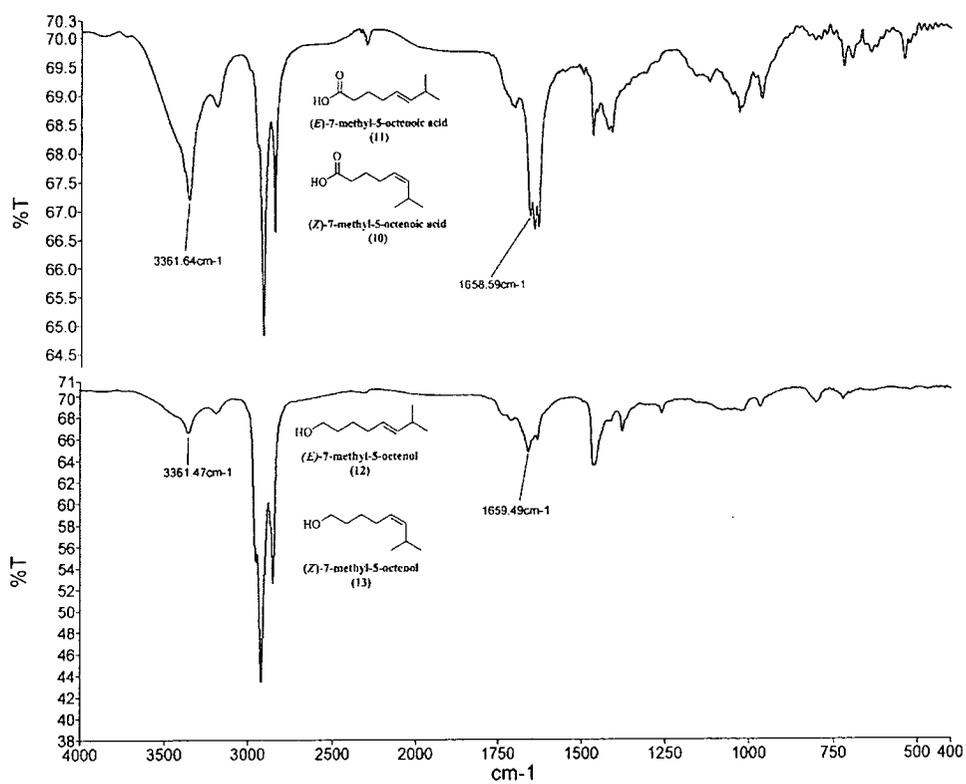


Figure 41 Overlay FT-IR spectrum of mixture (*E/Z*)-7-methyl-5-octenoic acid (11 and 10) and (*E/Z*)-7-Methyl-5-octenal (12 and 13)

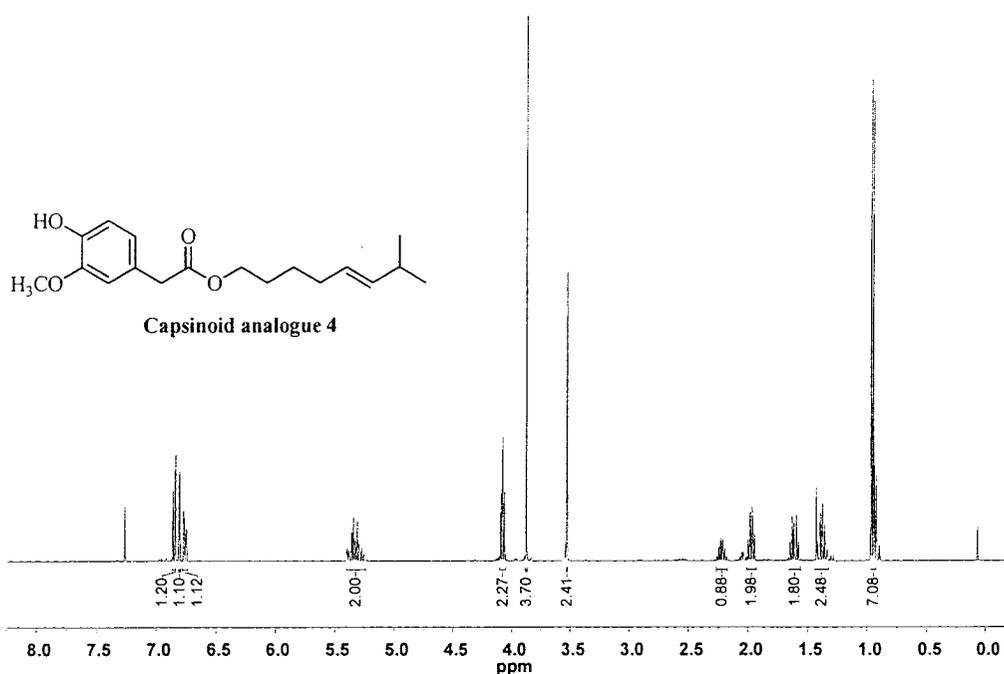


Figure 42 ¹H NMR Spectrum of capsinoid analogue 4, (CDCl₃)

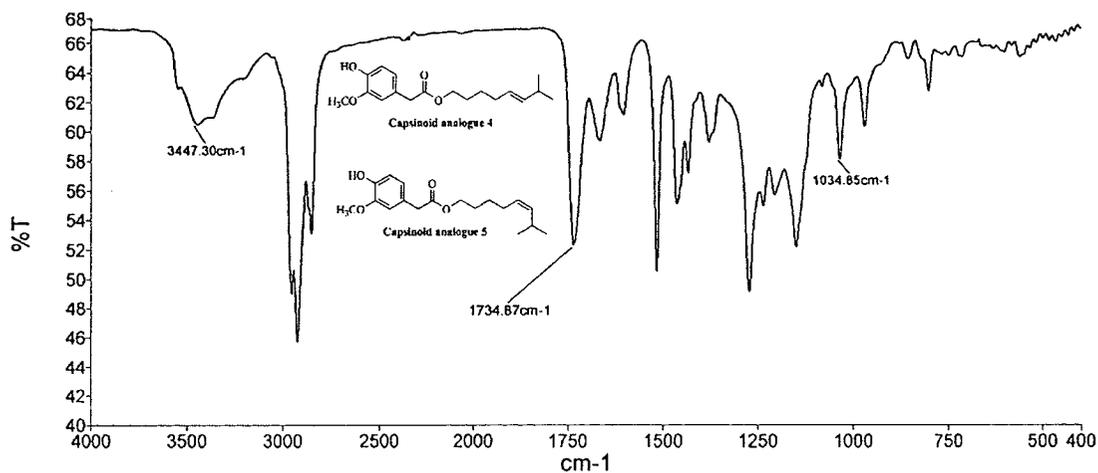


Figure 43 FT-IR spectrum of mixture capsinoid analogue 4 and 5

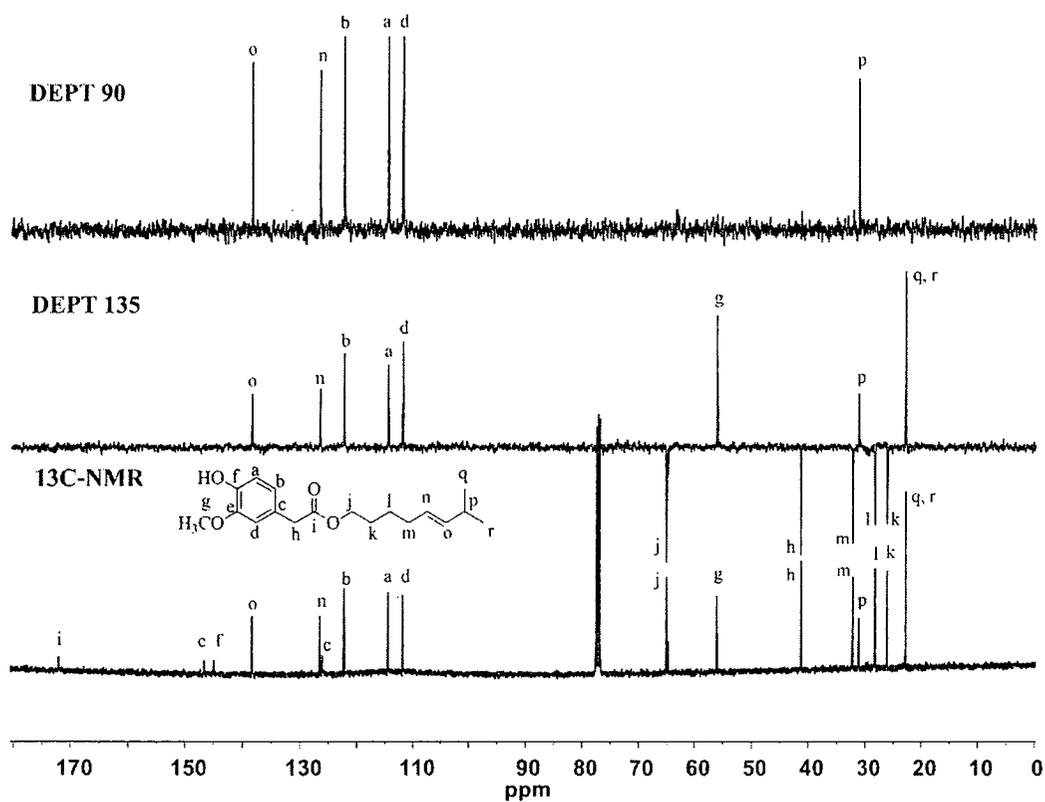


Figure 44 ^{13}C NMR Spectrum of capsinoid analogue 4, (CDCl_3)

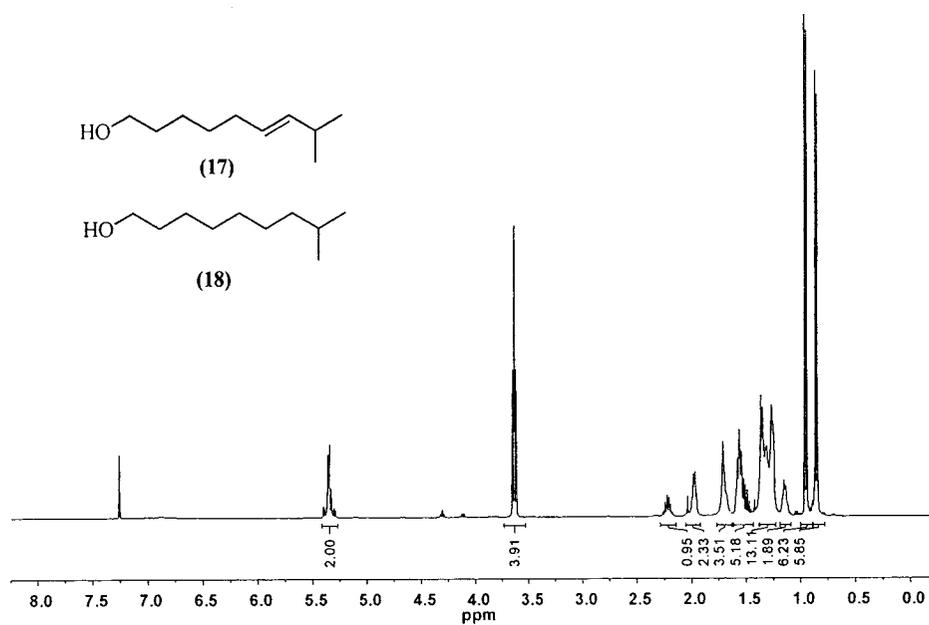


Figure 45 ^1H NMR Spectrum the mixture of (*E*)-8-methyl-6-nonenol (17) and 8-methylnonanol (18) (CDCl_3)

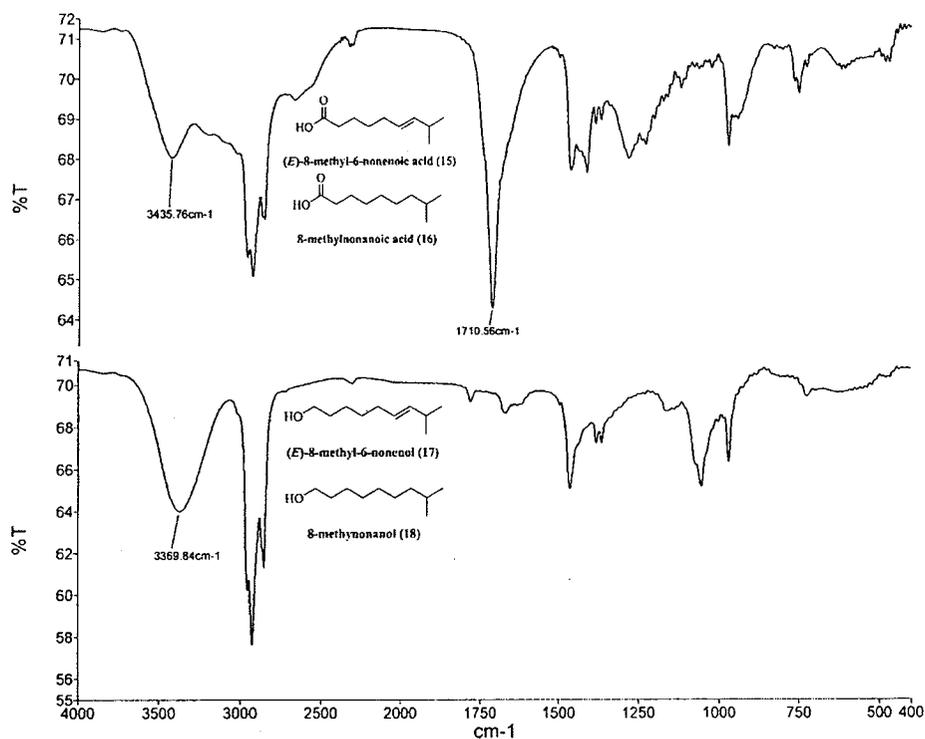


Figure 46 Overlay FT-IR spectrum of mixture (*E*)-8-methyl-6-nonenol (17) and 8-methylnonanol (18)

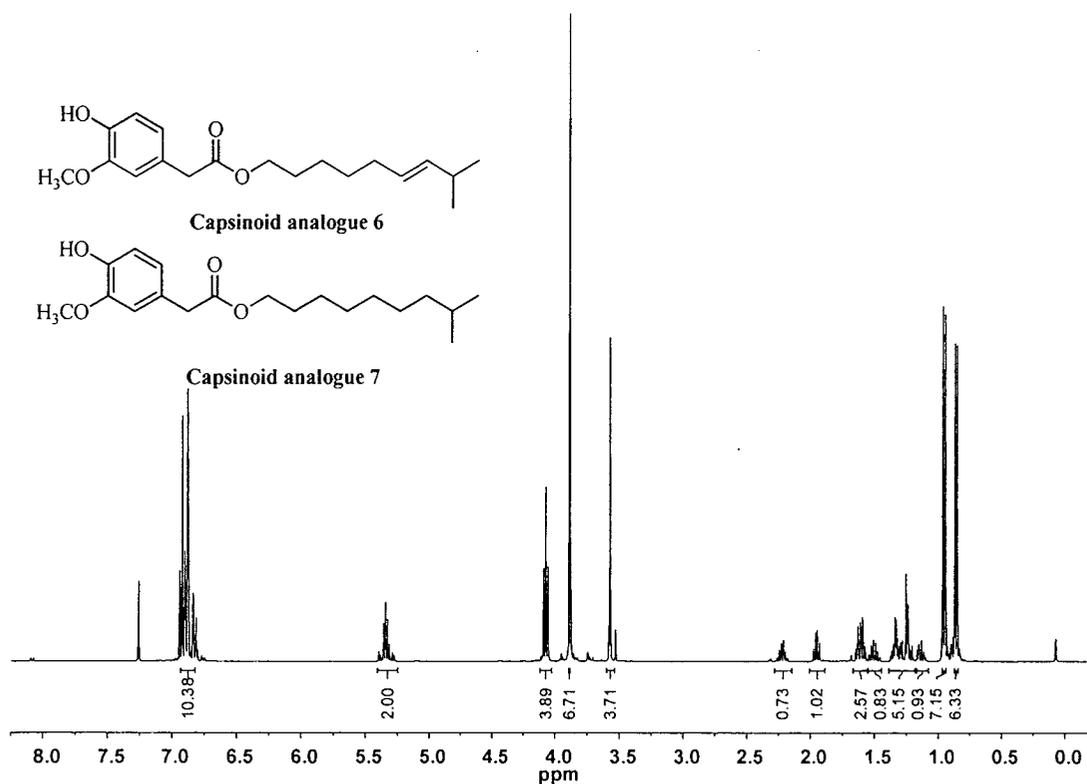


Figure 47 ¹H NMR Spectrum the mixture of capsinoid analogue 6 and 7 (CDCl₃)

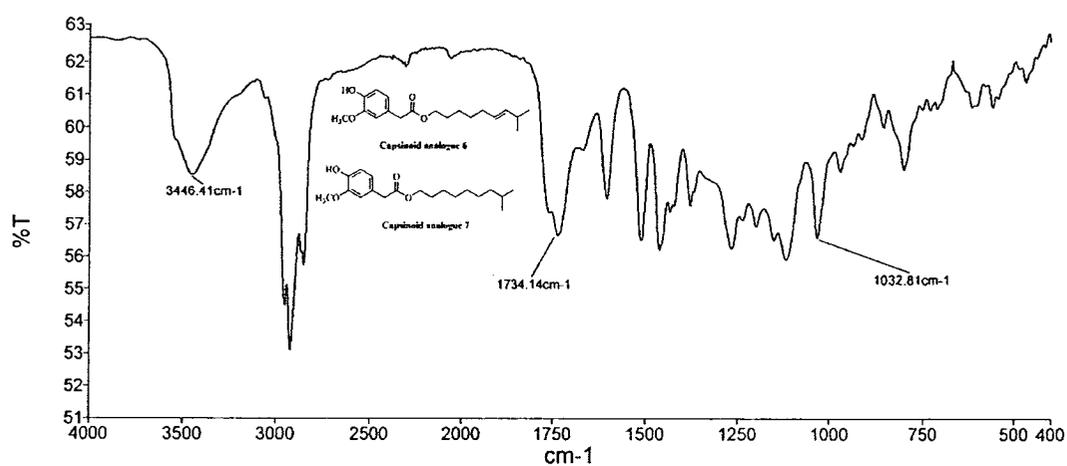


Figure 48 FT-IR spectrum of mixture capsinoid analogue 6 and 7

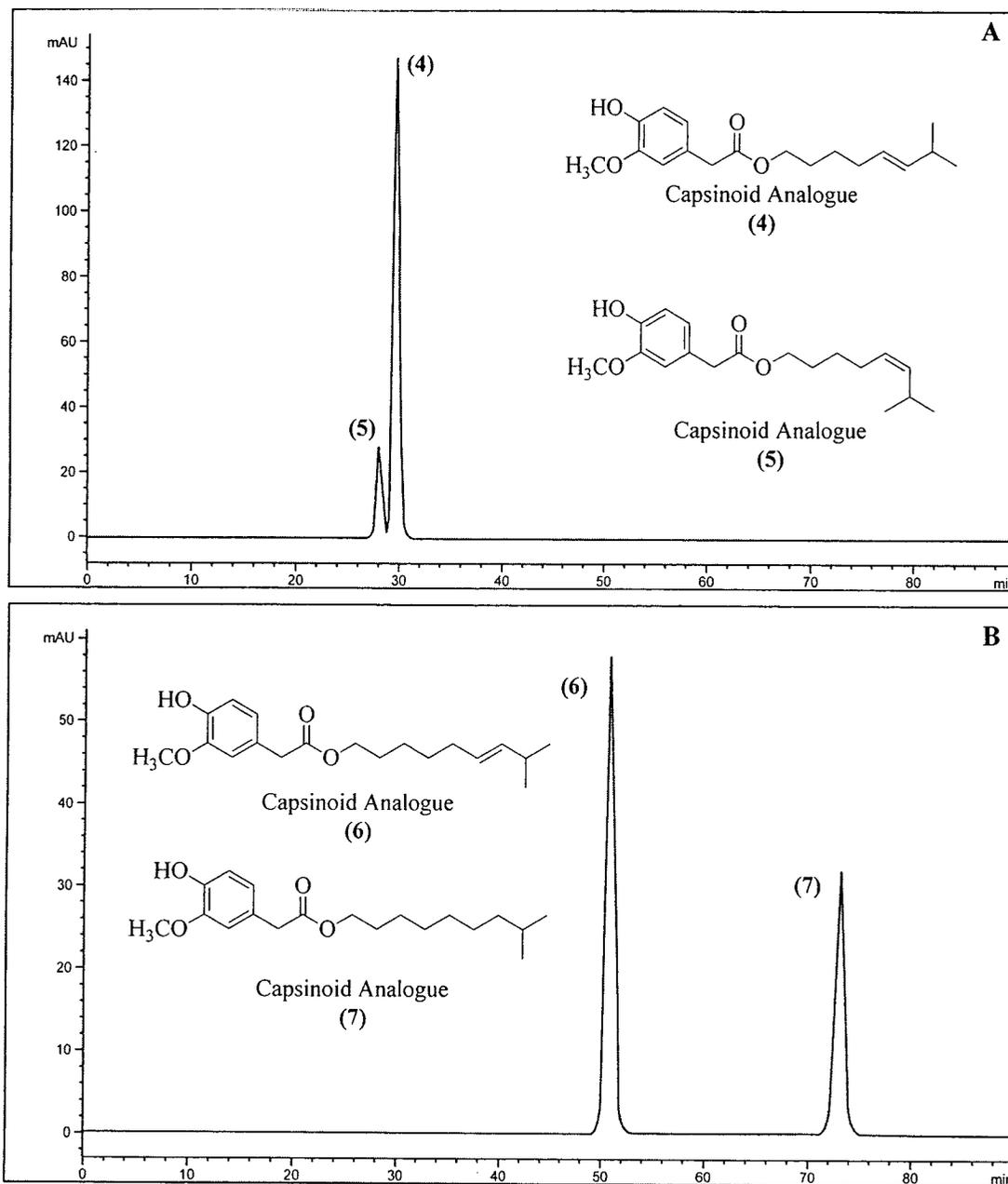
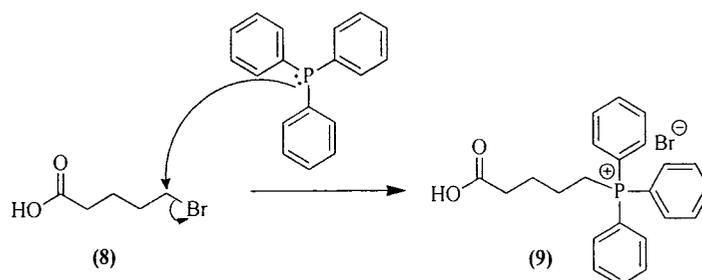
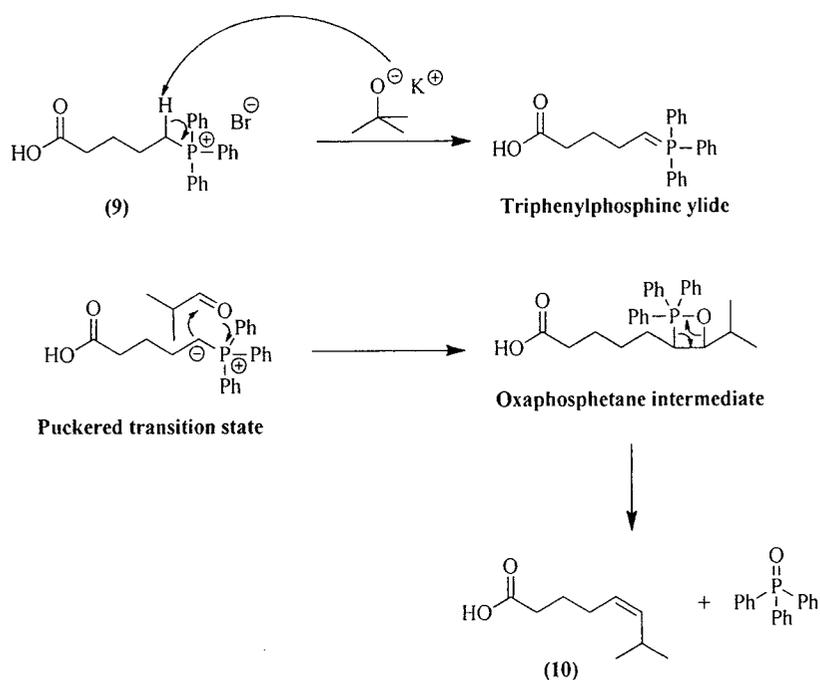


Figure 49 Chromatogram of Capsinoid analogues (4-7) at 0 hour, C18, 4.6 x 250 mm, 5 μ m, mobile phase, CH₃OH:H₂O 80:20 v/v with 0.025 AcOH, flow rate 0.5 mL/min, detection UV 280 nm

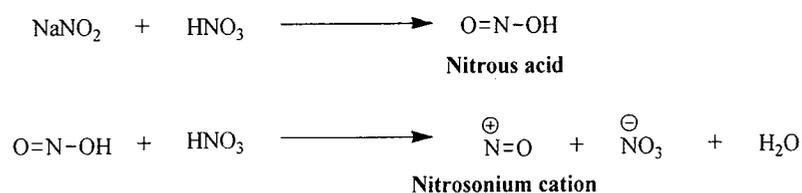
APPENDIX B Mechanism



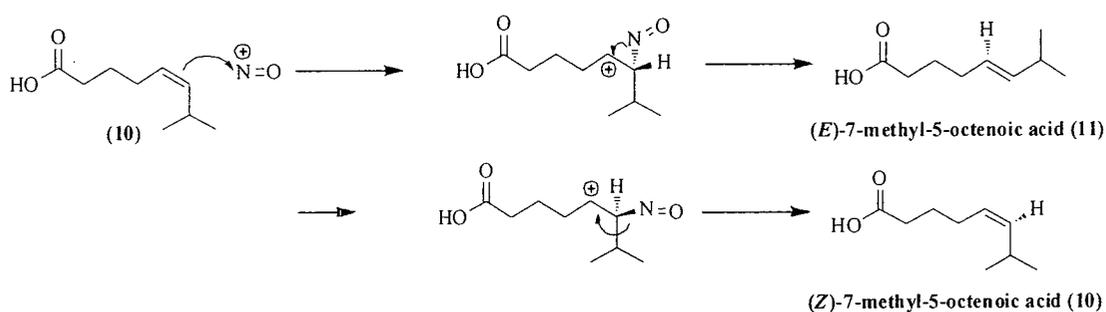
Scheme 21 S_N2 mechanism of 5-bromovaleric acid (8) via triphenylphosphine



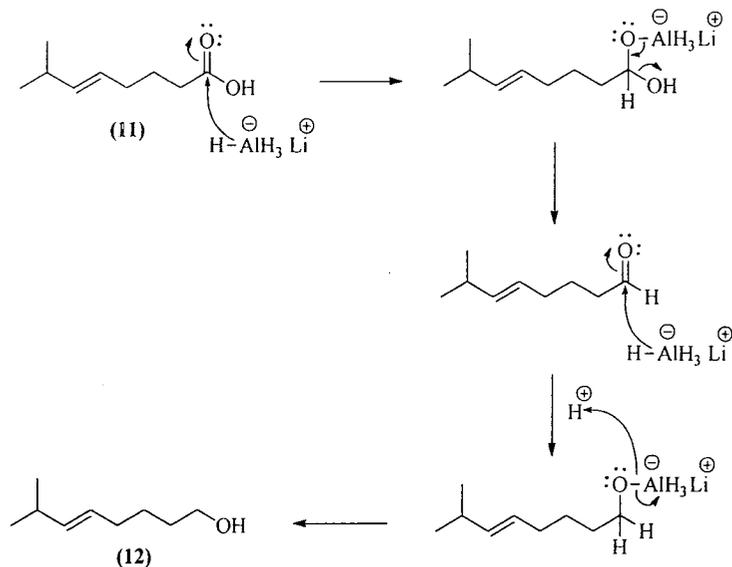
Scheme 22 Wittig reaction mechanism of phosphine ylide with isobutyraldehyde producing (*Z*)-7-methyl-5-octenoic acid (10)



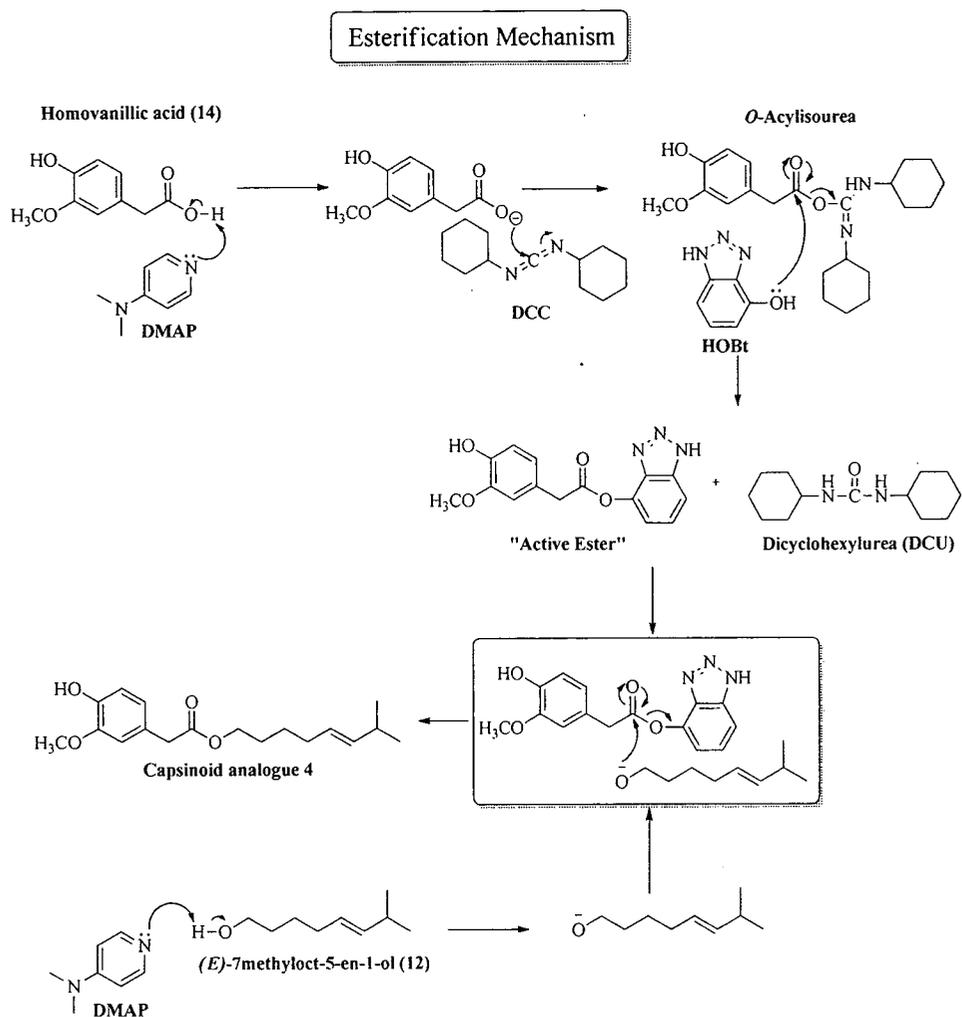
Scheme 23 Generation of nitrosonium cation from sodium nitrite and nitric acid



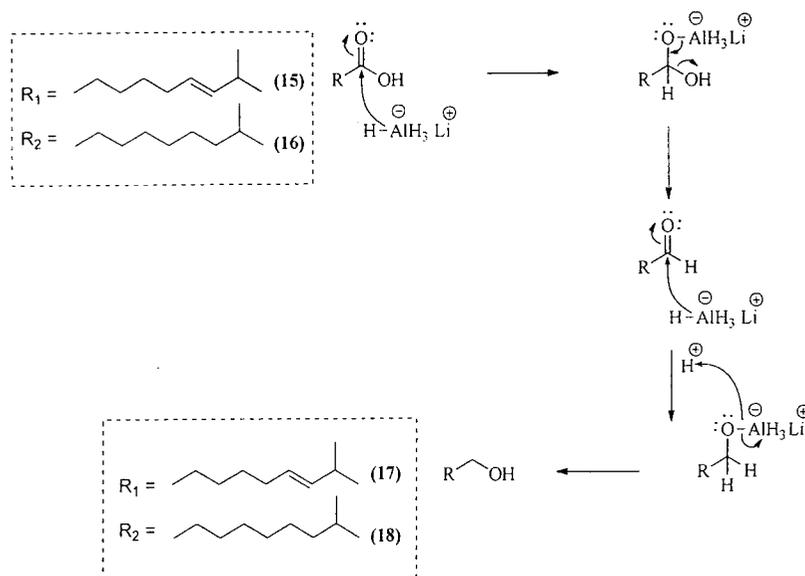
Scheme 24 Isomerization of (Z)-7-methyl-5-octenoic acid (10) to (E)-7-methyl-5-octenoic acid (11) via nitrosonium cation



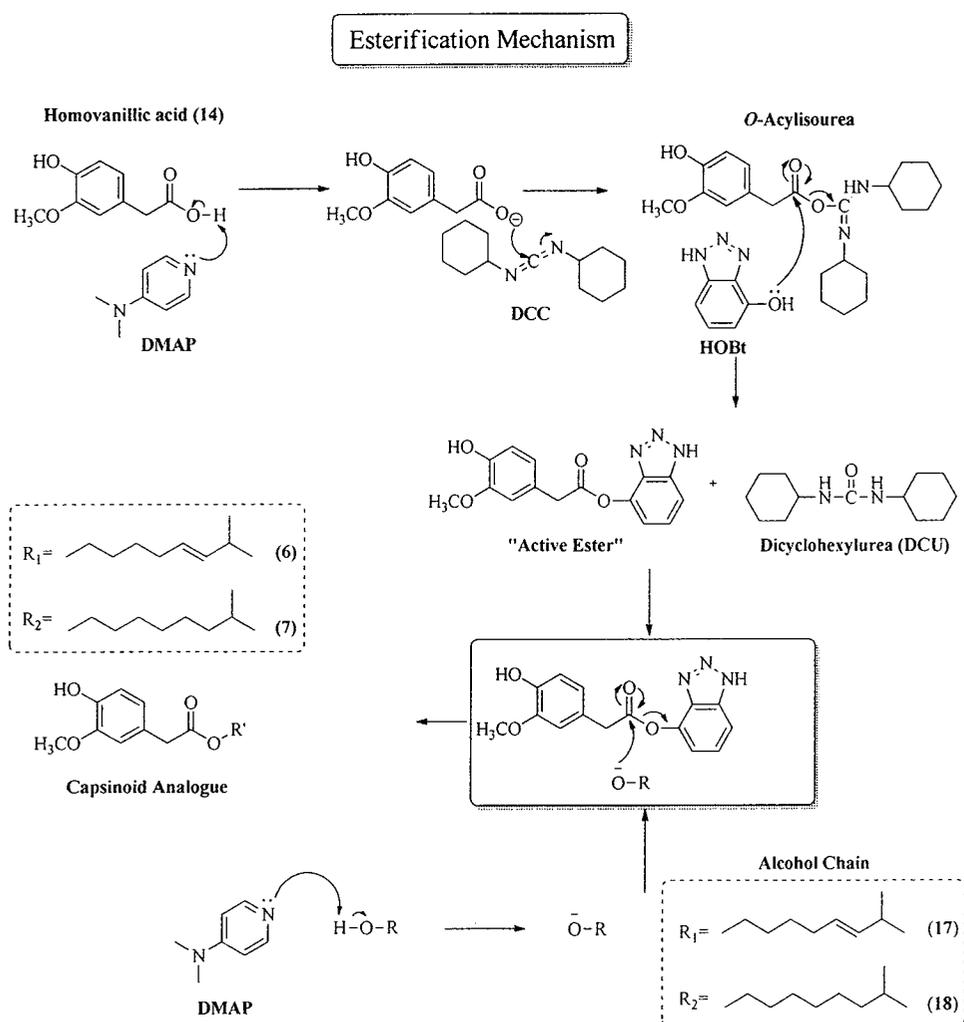
Scheme 25 Reduction mechanism of (E)-7-methyl-5-octenoic acid (11) to (E)-7-Methyl-5-octenol (12) via LiAlH_4 as a reducing agent



Scheme 26 Esterification reaction mechanism *via* DCC, DMAP and HOBt as a coupling agent



Scheme 27 Reduction mechanism of the mixture compounds (15, 16) to *(E)*-8 Methyl-6-nonenol (17) and 8-methylnonanol (18) via LiAlH_4 as a reducing agent



**Scheme 28 Esterification reaction mechanism of 6, 7
via DCC, DMAP and HOBT as a coupling agent**

APPENDIX C Calculation for percent remaining of *E/Z*-capsiate, capsinoid analogue 4, capsinoid analogue 5, capsinoid analogue 6 and capsinoid analogue 7

Table 6 Calculated percent remaining of capsinoid analogue 4

Sample	Percent remaining of capsinoid analogue 4				
	under period of time (%)				
	0 hour	6 hour	12 hour	18 hour	24 hour
n = 1	100.000	100.043	99.980	100.017	100.293
n = 2	100.000	99.999	100.046	100.255	100.169
n = 3	100.000	100.018	99.983	99.991	100.169
n = 4	100.000	100.032	100.020	100.040	100.345
average	100.000	100.023	100.007	100.076	100.211
SD	0.000	0.019	0.032	0.121	0.072

Table 7 Calculated percent remaining capsinoid analogue 5

Sample	Percent remaining of capsinoid analogue 5				
	under period of time (%)				
	0 hour	6 hour	12 hour	18 hour	24 hour
n = 1	100.000	99.939	99.943	99.951	100.090
n = 2	100.000	99.830	99.998	101.176	100.139
n = 3	100.000	99.934	100.053	100.040	100.247
n = 4	100.000	100.055	100.020	99.997	100.233
average	100.000	99.939	100.004	100.291	100.159
SD	0.000	0.092	0.046	0.591	0.081

Table 8 Calculated percent remaining capsinoid analogue 6

Sample	Percent remaining of capsinoid analogue 6 under period of time (%)				
	0 hour	6 hour	12 hour	18 hour	24 hour
n = 1	100.000	99.946	100.043	101.395	100.051
n = 2	100.000	100.122	100.159	101.016	100.082
n = 3	100.000	100.119	100.054	100.938	100.027
n = 4	100.000	99.876	100.047	100.884	100.025
average	100.000	100.016	100.076	101.116	100.046
SD	0.000	0.125	0.056	0.245	0.027

Table 9 Calculated percent remaining capsinoid analogue 7

Sample	Percent remaining of capsinoid analogue 7 under period of time (%)				
	0 hour	6 hour	12 hour	18 hour	24 hour
n = 1	100.000	99.941	100.009	100.064	99.996
n = 2	100.000	99.549	99.913	100.035	100.004
n = 3	100.000	99.019	99.937	100.040	99.982
n = 4	100.000	98.929	99.970	99.981	100.026
average	100.000	99.360	99.957	100.030	100.002
SD	0.000	0.474	0.042	0.035	0.018

Table 10 Calculated percent remaining *E*-capsiate

Sample	Percent remaining of <i>E</i> -capsiate under period of time (%)				
	0 hour	6 hour	12 hour	18 hour	24 hour
n = 1	100.000	94.707	91.632	88.230	86.513
n = 2	100.000	94.957	92.445	85.817	85.340
n = 3	100.000	90.453	89.481	88.149	85.665
n = 4	100.000	93.843	93.591	83.513	85.298
average	100.000	93.490	91.839	86.427	85.704
SD	0.000	2.080	2.121	2.242	0.563

Table 11 Calculated percent remaining *Z*-capsiate

Sample	Percent remaining of <i>Z</i> -capsiate under period of time (%)				
	0 hour	6 hour	12 hour	18 hour	24 hour
n = 1	100.000	77.063	69.598	66.231	59.969
n = 2	100.000	77.122	69.869	71.275	59.669
n = 3	100.000	72.749	69.061	65.815	59.669
n = 4	100.000	74.142	71.512	65.961	59.945
average	100.000	75.269	70.010	67.320	59.813
SD	0.000	2.181	1.056	2.642	0.167

APPENDIX D %Cell viability of Caco-2 cell *via* cytotoxicity MTT assay

Table 12 %Cell viability of Caco-2 cell *via* cytotoxicity MTT assay at 18 hours

Treatment Dose (μM)	%Cell viability of Caco-2 cell <i>via</i> cytotoxicity MTT assay					
	Capsaicin	Capsiate	Capsinoid analogue (4)	Capsinoid analogue (5)	Capsinoid analogue (6)	Capsinoid analogue (7)
0.10	83.77 \pm 0.00	98.65 \pm 0.01	99.74 \pm 0.01	90.88 \pm 0.01	99.53 \pm 0.01	98.03 \pm 0.02
1.00	82.90 \pm 0.00	98.17 \pm 0.04	93.37 \pm 0.02	94.07 \pm 0.08	99.41 \pm 0.00	92.68 \pm 0.01
10.00	71.57 \pm 0.02	88.24 \pm 0.03	95.36 \pm 0.02	91.80 \pm 0.01	94.07 \pm 0.01	95.04 \pm 0.00
100.00	54.96 \pm 0.01	84.25 \pm 0.00	93.04 \pm 0.02	91.37 \pm 0.03	94.82 \pm 0.01	100.22 \pm 0.02
200.00	49.68 \pm 0.00	76.43 \pm 0.02	82.74 \pm 0.03	83.71 \pm 0.03	90.51 \pm 0.04	96.76 \pm 0.01

n = 3, 1%PEG+DMEM us as control, capsaicin negative control, capsiate positive control

Table 13 %Cell viability of Caco-2 cell *via* cytotoxicity MTT assay at 24 hours

Treatment Dose (μM)	%Cell viability of Caco-2 cell <i>via</i> cytotoxicity MTT assay					
	Capsaicin	Capsiate	Capsinoid analogue (4)	Capsinoid analogue (5)	Capsinoid analogue (6)	Capsinoid analogue (7)
0.10	88.95 \pm 0.00	91.01 \pm 0.00	97.76 \pm 0.01	90.09 \pm 0.01	98.37 \pm 0.01	96.09 \pm 0.02
1.00	72.97 \pm 0.01	89.36 \pm 0.00	88.87 \pm 0.00	85.07 \pm 0.01	83.94 \pm 0.02	81.13 \pm 0.02
10.00	69.37 \pm 0.00	85.06 \pm 0.02	85.66 \pm 0.00	84.13 \pm 0.00	77.01 \pm 0.00	80.01 \pm 0.01
100.00	51.12 \pm 0.00	79.78 \pm 0.00	82.57 \pm 0.03	80.13 \pm 0.00	74.54 \pm 0.01	78.20 \pm 0.01
200.00	46.85 \pm 0.01	76.37 \pm 0.00	80.89 \pm 0.02	73.32 \pm 0.01	71.37 \pm 0.00	75.13 \pm 0.00

n = 3, 1%PEG+DMEM us as control, capsaicin negative control, capsiate positive control