



Chemical Constituents from the Bark of *Artocarpus elasticus*

Aeesoh Yanya

**A Thesis Submitted in Partial Fulfillment of the Requirements
for the Degree of Master of Science in Organic Chemistry**

Prince of Songkla University

2009

Copyright of Prince of Songkla University

Thesis Title Chemical Constituents from the Bark of *Artocarpus elasticus*
Author Miss Aeesoh Yanya
Major Program Organic Chemistry

Major Advisor:

.....
(Assoc. Prof. Dr. Wilawan Mahabusarakam)

Examining Committee:

.....Chairperson
(Assoc. Prof. Dr. Kan Chantrapomma)

Co-advisor:

.....
(Assoc. Prof. Chanita Ponglimanont)

.....
(Assoc. Prof. Dr. Wilawan Mahabusarakam)

.....
(Assoc. Prof. Chanita Ponglimanont)

.....
(Dr. Suda Chakthong)

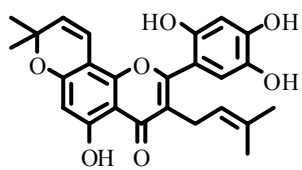
The Graduate School, Prince of Songkla University, has approved this thesis as partial fulfillment of the requirements for the Master of Science Degree in Organic Chemistry

.....
(Assoc. Prof. Dr. Krerckchai Thongnoo)
Dean of Graduate School

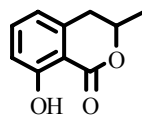
ชื่อวิทยานิพนธ์	องค์ประกอบทางเคมีจากเปลือกต้นกะออก (<i>Artocarpus elasticus</i>)
ผู้เขียน	นางสาวอวีเชาะส์ ยานยา
สาขาวิชา	เคมีอินทรีย์
ปีการศึกษา	2551

บทคัดย่อ

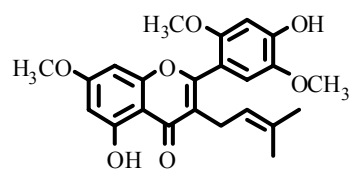
การศึกษาองค์ประกอบทางเคมีของเปลือกต้นกะออก (*Artocarpus elasticus*) แยกได้สารประกอบที่ยังไม่มีรายงานการวิจัย 6 สาร ซึ่งเป็นสารประกอบที่เกิดจากการเรียงตัวใหม่ของสารประเภท prenylated flavone คือ 5-hydroxy-2-(4-hydroxy-2,5-dimethoxyphenyl)-7-methoxy-3-(3-methylbut-2-enyl)-4*H*-chromen-4-one (AE3) อนุพันธ์ของสารประกอบประเภท furanodihydrobenzoxanthone (AE7) อนุพันธ์ของสารประกอบประเภท quinonobenzoxanthone (AE11 และ AE13) 1,3,4,8-tetrahydroxy-10-methoxy-5-(prop-1-en-2-yl)-5*H*-benzo[*c*]xanthen-7-(6*H*)-one (AE12) 5-hydroxy-8,8-dimethyl-2-(2,4-dihydroxyphenyl)-4*H*,8*H*-benzo[1,2-*b'*]dipyrans-4-one (AE15) นอกจากนี้ยังได้สารที่มีรายงานวิจัยแล้ว 10 สาร ได้แก่ 5-hydroxy-8,8-dimethyl-3-(3-methyl-2-butenyl)-2-(2,4,5-trihydroxyphenyl)-4*H*,8*H*-benzo[1,2-*b*:3,4-*b'*]dipyrans-4-one (AE1) 8-hydroxy-3-methylisochroman-1-one (AE2) 6,7-dihydro-5,9,14-trihydroxy-11-methoxy-3,3-dimethyl-6-(1-methylethyl)-3*H*,8*H*-[1]benzopyrano[7,6-*c*]xanthen-8-one (AE4) 12-acetyl-6-hydroxy-3,3,9,9-tetramethyl-3*H*,7*H*,furo[3,4-*b*]pyrano[3,2-*h*]xanthene-7,11(9*H*)-dione (AE5) 6,7-dihydro-5,9,11,14-tetrahydroxy-3,3-dimethyl-6-(1-methylethenyl)-(-)-3*H*,8*H*-pyrano[3',2':4,5]benzo[1,2-*c*]xanthen-8-one (AE6) 5*a*,6-dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-9-(3-methyl-2-buten-1-yl)-5*H*,7*H*,11*H*-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one (AE8) (3,4,5-trimethoxyphenyl)methanol (AE9) 5*a*,6-dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-5*H*,7*H*,11*H*-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one (AE10) 5*a*,6-dihydro-1,3,8-trihydroxy-10-methoxy-5,5-dimethyl-5*H*,7*H*-benzofuro[3,4-*bc*]xanthen-7-one (AE14) และ 2-(2,4-dihydroxyphenyl)-5,7-dihydroxy-4*H*-chromen-4-one (AE16) โครงสร้างของสารประกอบเหล่านี้วิเคราะห์โดยใช้ข้อมูลทางสเปกโทรสโกปี UV IR NMR MS และ เปรียบเทียบกับสารที่มีรายงานการวิจัยแล้ว



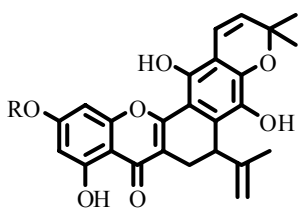
AE1



AE2

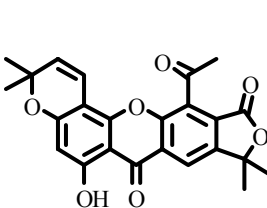


AE3

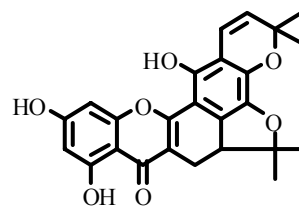


AE4: R = CH₃

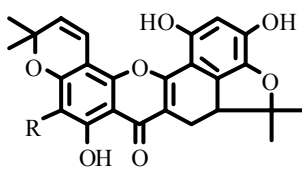
AE6: R = H



AE5

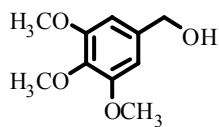


AE7

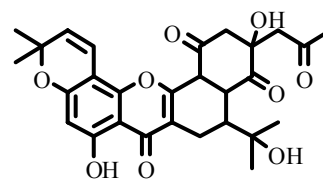


AE8 : R = prenyl

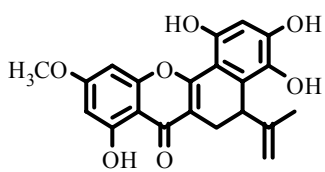
AE10: R = H



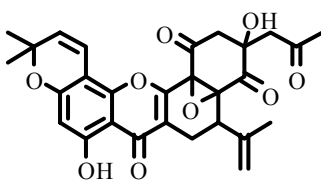
AE9



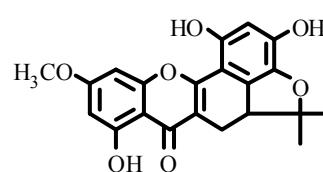
AE11



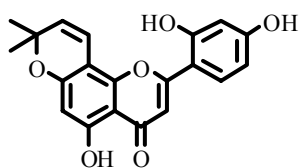
AE12



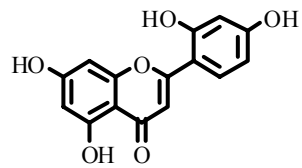
AE13



AE14



AE15

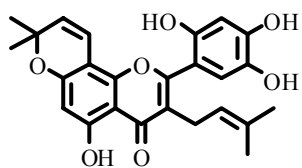


AE16

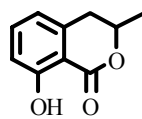
Thesis Title Chemical Constituents from the Bark of *Artocarpus elasticus*
Author Miss Aeesoh Yanya
Major Program Organic Chemistry
Acedemic Year 2008

ABSTRACT

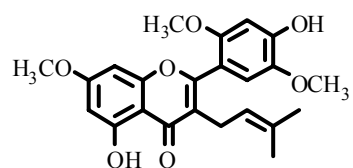
Investigation of the chemical constituents from the bark of *Artocarpus elasticus* yielded six modified prenylated flavones: 5-hydroxy-2-(4-hydroxy-2,5-dimethoxyphenyl)-7-methoxy-3-(3-methylbut-2-enyl)-4*H*-chromen-4-one (**AE3**), furanodihydrobenzoxanthone derivative (**AE7**), quinonobenzoxanthone derivatives (**AE11** and **AE13**), 1,3,4,8-tetrahydroxy-10-methoxy-5-(prop-1-en-2-yl)-5*H*-benzo[*c*]xanthen-7-(6*H*)-one (**AE12**), 5-hydroxy-8,8-dimethyl-2-(2,4-dihydroxyphenyl)-4*H*, 8*H*-benzo[1,2-*b'*]dipyran-4-one (**AE15**). Ten known compounds were also obtained: 5-hydroxy-8,8-dimethyl-3-(3-methyl-2-butenyl)-2-(2,4,5-trihydroxyphenyl)-4*H*, 8*H* benzo[1,2-*b*:3,4-*b'*]dipyran-4-one (**AE1**), 8-hydroxy-3-methylisochroman-1-one (**AE2**), 6,7-dihydro-5,9,14-trihydroxy-11-methoxy-3,3-dimethyl-6-(1-methylethyl)-3*H*, 8*H*-[1]benzopyrano[7,6-*c*]xanthen-8-one (**AE4**), 12-acetyl-6-hydroxy-3,3,9,9-tetramethyl-3*H*, 7*H*, furo[3,4-*b*]pyrano[3,2-*h*]xanthen-7,11(9*H*)-dione (**AE5**), 6,7-dihydro-5,9,11,14-tetrahydroxy-3,3-dimethyl-6-(1-methylethenyl)-(-)-3*H*, 8*H*-pyrano[3',2':4,5]benzo[1,2-*c*]xanthen-8-one (**AE6**), 5a,6-dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-9-(3-methyl-2-buten-1-yl)-5*H*, 7*H*, 11*H*-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one (**AE8**), (3,4,5-trimethoxyphenyl)methanol (**AE9**), 5a,6-dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-5*H*, 7*H*, 11*H*-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one (**AE10**), 5a,6-dihydro-1,3,8-trihydroxy-10-methoxy-5,5-dimethyl-5*H*, 7*H*-benzofuro[3,4-*bc*]xanthen-7-one (**AE14**) and 2-(2,4-dihydroxyphenyl)-5,7-dihydroxy-4*H*-chromen-4-one (**AE16**). Their structures were determined on the basis of UV, IR, NMR MS and by comparison of their spectroscopic data with those reported.



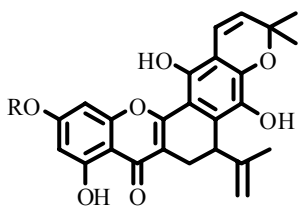
AE1



AE2

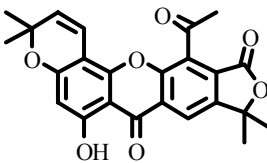


AE3

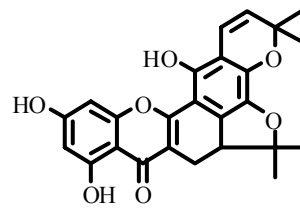


AE4: R = CH₃

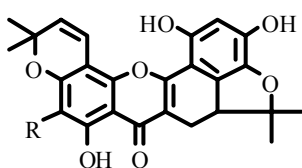
AE6: R = H



AE5

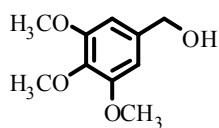


AE7

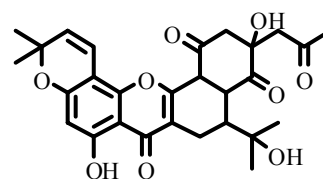


AE8 : R = prenyl

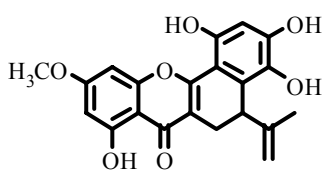
AE10: R = H



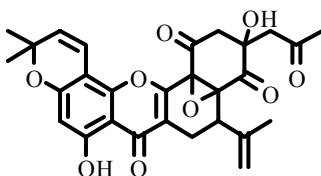
AE9



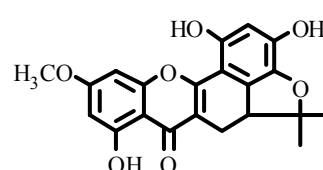
AE11



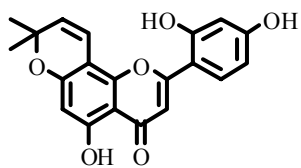
AE12



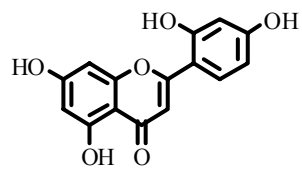
AE13



AE14



AE15



AE16

ACKNOWLEDGEMENTS

I wish to express my deepest and sincere gratitude to my supervisor, Associate Professor Dr. Wilawan Mahabusarakam, for her valuable instruction, expert guidance, excellent suggestion and kindness. I would also like to express my appreciation to Assoc. Prof. Chanita Ponglimanont my co-advisor, for correction of my thesis.

My sincere thanks are expressed to Associate Professor Dr. Saowalak Phongpaijit and Associate Professor Ramida Watanapokasin for bioactivity testing and Mr. Charernsak Saewai for plant identification. I would like to thank Mr. Asadhawut Hiranrat for his encouragement, assistance and guidance. In addition, I would like to sincerely thank Mr. Kreangsak Wicheansang for plant material, my family and friends for their love that supports me.

This work was partly supported by Center of Excellence for Innovation in Chemistry (PERCH-CIC), Commission on Higher Education, Ministry of Education, the Graduate School, Prince of Songkla University and Natural Products Research Center, Prince of Songkla University.

I would like to express my appreciation to the staffs of the Department of Chemistry, Faculty of Science, Prince of Songkla University for making this thesis possible.

Aeesoh Yanya

THE RELEVANCE OF THE RESEARCH WORK TO THAILAND

The purpose of this research is to investigate the chemical constituents of *Artocarpus elasticus*. It is a part of the basic research on the utilization of Thai medicinal plants. This research will contribute significantly to scientific basis of traditional medicine. Fourteen prenylated flavones type pure compounds, mullein and (3,4,5-trimethoxyphenyl)methanol have been isolated from this plant. Artonin E showed strong anti-*S. aureus* ATCC25923 and MRSA SK1 with MIC 4 and 8 $\mu\text{g/ml}$, respectively. Moreover, many compounds of prenylated flavones type have been reported to have cytotoxic, antimicrobial and antioxidation activities. Further study on the biological activity of the isolated compounds should be performed which can lead to active compounds. Therefore Thai plant can be utilized as a natural resource of potential drugs.

CONTENTS

	page
Abstract (in Thai)	iii
Abstract (in English)	v
Acknowledgements	vii
The relevance of the research work to Thailand	viii
Contents	ix
List of Tables	xi
List of Illustrations	xiii
Abbreviations and Symbols	xx
Chapter	
1. Introduction	1
1.1 Introduction	1
1.2 Review of Literatures	4
1.2.1 The Chemical Constituents and Biological Activities of <i>Artocarpus</i> genus	4
1.3 Objective	34
2. Experimental	35
2.1 General Method	35
2.2 Plant Material	35
2.3 Extraction and Isolation	36
2.3.1 Purification of dichloromethane extract	37
2.3.1.1 Purification of fraction B	40
2.3.1.2 Purification of fraction C	44
2.3.1.3 Purification of fraction D	48
2.3.2 Purification of acetone extract	55
3. Results and Discussion	56
3.1 Structure Determination	56
AE1	57
AE2	59

CONTENTS (Continued)

	page
AE3	61
AE4	64
AE5	68
AE6	71
AE7	73
AE8	77
AE9	81
AE10	83
AE11	86
AE12	90
AE13	93
AE14	96
AE15	98
AE16	100
3.2 Evaluation of Biological activities of crude extracts	102
3.2.1 Antimicrobial activity	102
3.2.2 Cytotoxic activity	104
3.3 Review of biological activities of the known compounds obtained from this study	104
3.4 Biogenesis of flavonoids and related compounds from <i>Artocapus elasticus</i>	106
References	110
Appendix	117
Vitae	182

LIST OF TABLES

Table		Page
1	Compounds from the <i>Artocarpus</i> genus	7
2	Physical characteristic and weight of the fractions obtained from QCC of crude CH ₂ Cl ₂ extract	37
3	Physical characteristic and weight of the fractions obtained from CC of fraction B	40
4	Physical characteristic and weight of the fractions obtained from CC of fraction C	44
5	Physical characteristic and weight of the fractions obtained from CC of fraction D	48
6	¹³ C, ¹ H and HMBC spectral data of AE1	58
7	¹³ C, ¹ H and HMBC spectral data of AE2	60
8	¹ H- ¹ H COSY spectral data of AE2	60
9	¹³ C, ¹ H and HMBC spectral data of AE3	63
10	¹ H- ¹ H COSY spectral data of AE3	63
11	¹³ C, ¹ H and HMBC spectral data of AE4	66
12	¹ H- ¹ H COSY spectral data of AE4	67
13	¹³ C, ¹ H and HMBC spectral data of AE5	69
14	¹³ C and ¹ H spectral data of artanol B	70
15	¹³ C, ¹ H and HMBC spectral data of AE6	72
16	¹³ C, ¹ H and HMBC spectral data of AE7	75
17	¹ H- ¹ H COSY spectral data of AE7	76
18	¹³ C, ¹ H and HMBC spectral data of AE8	79
19	¹ H- ¹ H COSY spectral data of AE8	80
20	¹³ C, ¹ H and HMBC spectral data of AE9	82
21	¹³ C, ¹ H and HMBC spectral data of AE10	84

LIST OF TABLES (Continued)

Table		Page
22	¹ H- ¹ H COSY spectral data of AE10	85
23	¹³ C, ¹ H and HMBC spectral data of AE11	88
24	¹ H- ¹ H COSY spectral data of AE11	89
25	¹³ C, ¹ H and HMBC spectral data of AE12	92
26	¹ H- ¹ H COSY spectral data of AE12	92
27	¹³ C, ¹ H and HMBC spectral data of AE13	94
28	¹ H- ¹ H COSY spectral data of AE13	95
29	¹³ C, ¹ H and HMBC spectral data of AE14	97
30	¹ H- ¹ H COSY spectral data of AE14	97
31	¹³ C, ¹ H and HMBC spectral data of AE15	99
32	¹ H- ¹ H COSY spectral data of AE15	101
33	¹³ C, ¹ H and HMBC spectral data of AE16	101
34	¹ H- ¹ H COSY spectral data of AE16	101
35	Antibacterial activity of crude extracts from the bark of <i>A. elasticus</i>	102
36	Antiyeast and Antifungal activity of crude extracts from the bark of <i>A. elasticus</i>	103
37	Antibacterial activity of compounds isolated from the bark of <i>A. elasticus</i>	103
38	Cytotoxic activity of crude extracts from the bark of <i>A. elasticus</i>	104

LIST OF ILLUSTRATIONS

Scheme		Page
1	Extraction of crude extracts from the bark of <i>A. elasticus</i>	36
2	Isolation of compounds AE2-AE15 from CH ₂ Cl ₂ extract of the bark of <i>A. elasticus</i>	38
3	Isolation of compounds AE16 from acetone extract of the bark of <i>A. elasticus</i>	55
4	Relationships of prenylated flavonoids from <i>A. elasticus</i>	106
5	Proposed biogenetic route of AE7	107
6	Proposed biogenetic route of the furanodihydrobenzoxanthone types of compounds	108
Figure		
1	<i>Artocarpus elasticus</i>	3
2	The structure of regular flavonoids in <i>Artocarpus</i>	4
3	The structure of modified flavonoid in <i>Artocarpus</i>	5
4	The structure of flavonoid-derived xanthenes in <i>Artocarpus</i>	5
5	Flavone structure	6
6	UV (CH ₃ OH) spectrum of AE1	118
7	FT-IR (Neat) spectrum of AE1	118
8	¹ H NMR (300 MHz) (CDCl ₃ +DMSO- <i>d</i> ₆) spectrum of AE1	119
9	¹³ C NMR (75 MHz) (CDCl ₃ +DMSO- <i>d</i> ₆) spectrum of AE1	119
10	DEPT 135° (CDCl ₃ +DMSO- <i>d</i> ₆) spectrum of AE1	119
11	2D HMQC spectrum of AE1	120
12	2D HMBC spectrum of AE1	120

LIST OF ILLUSTRATIONS (Continued)

Figure		Page
13	FT-IR (Neat) spectrum of AE2	121
14	¹ H NMR (300 MHz) (CDCl ₃) spectrum of AE2	122
15	¹ H- ¹ H COSY spectrum of AE2	122
16	¹³ C NMR (75 MHz) (CDCl ₃) spectrum of AE2	123
17	DEPT 135° (CDCl ₃) spectrum of AE2	123
18	DEPT 90° (CDCl ₃) spectrum of AE2	123
19	2D HMQC spectrum of AE2	124
20	2D HMBC spectrum of AE2	124
21	EI-MS spectrum of AE3	125
22	FT-IR (Neat) spectrum of AE3	126
23	¹ H NMR (300 MHz) (CDCl ₃) spectrum of AE3	126
24	NOEDIFF spectrum of AE3 after irradiation	127
25	¹ H- ¹ H COSY spectrum of AE3	127
26	¹³ C NMR (75 MHz) (CDCl ₃) spectrum of AE3	128
27	DEPT 135° (CDCl ₃) spectrum of AE3	128
28	2D HMQC spectrum of AE3	129
29	2D HMBC spectrum of AE3	129
30	UV (EtOH) spectrum of AE4	130
31	FT-IR (Neat) spectrum of AE4	130
32	¹ H NMR (300 MHz) (CDCl ₃) spectrum of AE4	131
33	¹ H- ¹ H COSY spectrum of AE4	131
34	¹³ C NMR (75 MHz) (CDCl ₃) spectrum of AE4	132
35	DEPT 135° (CDCl ₃) spectrum of AE4	132
36	2D HMQC spectrum of AE4	133
37	2D HMBC spectrum of AE4	133

LIST OF ILLUSTRATIONS (Continued)

Figure		Page
38	UV (EtOH) spectrum of AE5	134
39	FT-IR (Neat) spectrum of AE5	134
40	¹ H NMR (300 MHz) (CDCl ₃) spectrum of AE5	135
41	¹ H- ¹ H COSY spectrum of AE5	135
42	¹³ C NMR (75 MHz) (CDCl ₃) spectrum of AE5	136
43	2D HMQC spectrum of AE5	137
44	2D HMBC spectrum of AE5	137
45	UV (CH ₃ OH) spectrum of AE6	138
46	FT-IR (Neat) spectrum of AE6	138
47	¹ H NMR (300 MHz) (CDCl ₃ +Acetone- <i>d</i> ₆) spectrum of AE6	139
48	¹³ C NMR (75 MHz) (CDCl ₃ +Acetone- <i>d</i> ₆) spectrum of AE6	139
49	DEPT 135° (CDCl ₃ +Acetone- <i>d</i> ₆) spectrum of AE6	139
50	2D HMQC spectrum of AE6	140
51	2D HMBC spectrum of AE6	140
52	EI-MS spectrum of AE7	141
53	UV (EtOH) spectrum of AE7	142
54	FT-IR (Neat) spectrum of AE7	142
55	¹ H NMR (300 MHz) (CDCl ₃ +Acetone- <i>d</i> ₆) spectrum of AE7	143
56	¹ H- ¹ H COSY spectrum of AE7	143
57	¹³ C NMR (75 MHz) (CDCl ₃ +Acetone- <i>d</i> ₆) spectrum of AE7	144
58	DEPT 135° (CDCl ₃ +Acetone- <i>d</i> ₆) spectrum of AE7	144

LIST OF ILLUSTRATIONS (Continued)

Figure		Page
59	2D HMQC spectrum of AE7	145
60	2D HMBC spectrum of AE7	145
61	UV (CH ₃ OH) spectrum of AE8	146
62	FT-IR (Neat) spectrum of AE8	146
63	¹ H NMR (300 MHz) (CDCl ₃ +DMSO- <i>d</i> ₆) spectrum of AE8	147
64	¹ H- ¹ H COSY spectrum of AE8	147
65	¹³ C NMR (75 MHz) (CDCl ₃ +DMSO- <i>d</i> ₆) spectrum of AE8	148
66	DEPT 135° (CDCl ₃ +DMSO- <i>d</i> ₆) spectrum of AE8	148
67	DEPT 90° (CDCl ₃ +DMSO- <i>d</i> ₆) spectrum of AE8	148
68	2D HMQC spectrum of AE8	149
69	2D HMBC spectrum of AE8	149
70	¹ H NMR (500 MHz) (CDCl ₃) spectrum of AE9	150
71	¹³ C NMR (125 MHz) (CDCl ₃) spectrum of AE9	151
72	DEPT 135° (CDCl ₃) spectrum of AE9	151
73	DEPT 90° (CDCl ₃) spectrum of AE9	151
74	2D HMQC spectrum of AE9	152
75	2D HMBC spectrum of AE9	152
76	UV (EtOH) spectrum of AE10	153
77	FT-IR (Neat) spectrum of AE10	153
78	¹ H NMR (300 MHz) (CDCl ₃ +DMSO- <i>d</i> ₆) spectrum of AE10	154
79	¹ H- ¹ H COSY spectrum of AE10	154
80	¹³ C NMR (75 MHz) (CDCl ₃ +DMSO- <i>d</i> ₆) spectrum of AE10	155

LIST OF ILLUSTRATIONS (Continued)

Figure		Page
81	DEPT 135° (CDCl ₃ +DMSO- <i>d</i> ₆) spectrum of AE10	155
82	DEPT 90° (CDCl ₃ +DMSO- <i>d</i> ₆) spectrum of AE10	155
83	2D HMQC spectrum of AE10	156
84	2D HMBC spectrum of AE10	156
85	UV (EtOH) spectrum of AE11	157
86	FT-IR (Neat) spectrum of AE11	157
87	¹ H NMR (300 MHz) (CDCl ₃) spectrum of AE11	158
88	¹ H- ¹ H COSY spectrum of AE11	158
89	¹³ C NMR (75 MHz) (CDCl ₃) spectrum of AE11	159
90	2D HMQC spectrum of AE11	160
91	2D HMBC spectrum of AE11	160
92	EI-MS spectrum of AE12	161
93	UV (EtOH) spectrum of AE12	162
94	FT-IR (Neat) spectrum of AE12	162
95	¹ H NMR (300 MHz) (Acetone- <i>d</i> ₆) spectrum of AE12	163
96	¹ H- ¹ H COSY spectrum of AE12	163
97	¹³ C NMR (75 MHz) (Acetone- <i>d</i> ₆) spectrum of AE12	164
98	DEPT 135° (Acetone- <i>d</i> ₆) spectrum of AE12	164
99	DEPT 90° (Acetone- <i>d</i> ₆) spectrum of AE12	165
100	2D HMQC spectrum of AE12	165
101	2D HMBC spectrum of AE12	165
102	EI-MS spectrum of AE13	166
103	UV (EtOH) spectrum of AE13	167
104	FT-IR (Neat) spectrum of AE13	167
105	¹ H NMR (300 MHz) (CDCl ₃) spectrum of AE13	168

LIST OF ILLUSTRATIONS (Continued)

Figure		Page
106	^1H - ^1H COSY spectrum of AE13	168
107	^{13}C NMR (75 MHz) (CDCl_3) spectrum of AE13	169
108	DEPT 135° (CDCl_3) spectrum of AE13	169
109	DEPT 90° (CDCl_3) spectrum of AE13	169
110	2D HMQC spectrum of AE13	170
111	2D HMBC spectrum of AE13	170
112	UV (CH_3OH) spectrum of AE14	171
113	FT-IR (Neat) spectrum of AE14	171
114	^1H NMR (300 MHz) ($\text{CDCl}_3+\text{DMSO-}d_6$) spectrum of AE14	172
115	^1H - ^1H COSY spectrum of AE14	172
116	^{13}C NMR (75 MHz) ($\text{CDCl}_3+\text{DMSO-}d_6$) spectrum of AE14	173
117	DEPT 135° ($\text{CDCl}_3+\text{DMSO-}d_6$) spectrum of Ae14	173
118	2D HMQC spectrum of AE14	174
119	2D HMBC spectrum of AE14	174
120	UV (EtOH) spectrum of AE15	175
121	FT-IR (Neat) spectrum of AE15	175
122	^1H NMR (300 MHz) ($\text{Acetone-}d_6$) spectrum of AE15	176
123	^1H - ^1H COSY spectrum of AE15	176
124	^{13}C NMR (75 MHz) ($\text{Acetone-}d_6$) spectrum of AE15	177
125	2D HMQC spectrum of AE15	178
126	2D HMBC spectrum of AE15	178
127	^1H NMR (300 MHz) ($\text{Acetone-}d_6$) spectrum of AE16	179
128	^1H - ^1H COSY spectrum of AE16	179

LIST OF ILLUSTRATIONS (Continued)

Figure		Page
129	^{13}C NMR (75 MHz) (Acetone- d_6) spectrum of AE16	180
130	DEPT 135° (Acetone- d_6) spectrum of AE16	180
131	DEPT 90° (Acetone- d_6) spectrum of AE16	180
132	2D HMQC spectrum of AE16	181
133	2D HMBC spectrum of AE16	181

LIST OF ABBREVIATIONS AND SYMBOLS

<i>s</i>	=	singlet
<i>d</i>	=	doublet
<i>t</i>	=	triplet
<i>m</i>	=	multiplet
<i>dd</i>	=	doublet of doublet
<i>br s</i>	=	broad singlet
<i>g</i>	=	gram
kg	=	kilogram
mg	=	milligram
%	=	percent
nm	=	nanometer
m.p.	=	melting point
cm ⁻¹	=	reciprocal centimeter (wave number)
δ	=	chemical shift relative to TMS
<i>J</i>	=	coupling constant
λ_{\max}	=	maximum wavelength
ν	=	absorption frequencies
ϵ	=	molar extinction coefficient
°C	=	degree celcius
MHz	=	Megahertz
ppm	=	part per million
IR	=	Infrared
UV	=	Ultraviolet-Visible
NMR	=	Nuclear Magnetic Resonance
2D NMR	=	Two Dimentional Nuclear Magnetic Resonance
COSY	=	Correlated Spectroscopy

LIST OF ABBREVIATIONS AND SYMBOLS (Continued)

DEPT	=	Distortionless Enhancement by Polarization Transfer
------	---	---

HMBC	=	Heteronuclear Multiple Bond Correlation
HMQC	=	Heteronuclear Multiple Quantum Coherence
CC	=	column chromatography
TMS	=	tetramethylsilane
Acetone- <i>d</i> ₆	=	deuteroacetone
DMSO- <i>d</i> ₆	=	deuterodimethylsulphoxide
CDCl ₃	=	deuteriochloroform
MeOH	=	Methanol
CH ₂ Cl ₂	=	Dichloromethane
TLC	=	Thin-Layer Chromatography
MIC	=	Minimum Inhibition Concentration

CHAPTER 1

INTRODUCTION

1.1 Introduction

Nowadays, most of people have extensively concern about their health due to various types of pollutions which occur through atmosphere, food, water and also unhealthy eating habits which result in many types of diseases such as coronary thrombosis, diabetes, hypertension, cancer, alzheimer, cataract, rheumatism, progeria syndrome *etc.*. Thus, the discovery of new drug or the development in nutrient supplement has very high competition between the suppliers within country and also abroad. Thailand is a tropical country which contains many kinds of herbal plants that promise to cure many diseases, therefore Thai scientists have realized that it is necessary to conduct a research and analyze on new substances from the herbal plant which have pharmacological and biological activities. The study of the chemical composition in Thai herbal plant is really important. The information from the study show that the plant in the genus *Artocarpus* contains really high percentage of flavonoid groups which is categorized as one type of phenolic compound. This compound is known to potentially have antioxidant, antimutagen, antitumor, anti-inflammatory and anticarcinogenic properties. Previous study by Euis H. Hakim reported that “*Artocarpus* species contain phenolic compounds, including isoprenylated flavonoids, stilbenoids and 2-arylbenzofurans” (Hakim, *et al.*, 2006). Therefore, we were interested to study the chemical composition of the herbal plant in the genus *Artocarpus* which have been traditionally used by local people in many areas of Thailand and many other countries in Southeast Asia. Even though, the chemical composition, biological activity as well as pharmacological activity of *Artocarpus* genus have been studied worldwide in the past but only few researches about *Artocarpus elasticus* are conducted. Hence, the search for new bioactive compounds is really important as it can provide basic information for future study.

The *Artocarpus* genus is widely distributed throughout Thailand. It belongs to the mulberry family Moraceae and several members of the genus encompass approximately 60 species of trees that thrive throughout tropical Asia. This genus is known to be rich in prenylated flavonoids and their derivatives (Aida, *et al.*, 1997). A number of these trees are historically reputed to possess medicinal properties and are utilized as folk medicines in Taiwan, Thailand, and Indonesia (Nicolaou, *et al.*, 2008). *Artocarpus elasticus*, is locally known as 'Ka-ok' in Thailand. The use of ground-bark to allay backache and the latex to treat dysentery prompted us to examine *Artocarpus* species further for bioactive substances. However, there are only a few reports on the chemical constituents, we are therefore motivated to investigate its constituents in detail.

Artocarpus elasticus are found wild in the southern part of Thailand. *A. elasticus* is a large tree, grow to a height of 25-40 metre, spreading branches and a straight trunk with bark, outer bark is smooth and dark brown color while inner bark is light brown. The leaves are large 12-30 c.m., wide 20-55 c.m., bright-green and glossy on the upper surface, stiff hair on the underside, the petiole 5-10 c.m. long. Flowers bear a multitude of tiny flowers. Fruits are cylindrical-shaped, 5.5 c.m. wide, 12 c.m. long, have harsh, sand paper-like rind; generally the rind is green at first, turning yellow-brown when ripe. It produce flower during December-March, fruit will be mature during July-October. For geographic distribution it is found in the forest which located near canal and is also found in the southern area of Thailand. Moreover, it has also been found in other countries such as Myanmar, Malaysia, Indonesia. It has the sticky, milky latex which has a property like glue so it is used for trapping animal such as birds.



Figure 1 *Artocarpus elasticus*

1.2 Review of Literatures

1.2.1 The Chemical Constituents and Biological Activity of *Artocarpus* genus.

Prenylated flavonoids, stilbenoids and 2-arylbenzofuran were the major components isolated from *Artocarpus* genus. The flavonoid constituents may be further classified according to their skeletons, as chalcones, flavanones, flavones, flavan-3-ol, and 3-prenylflavones. Furthermore, there are classes of modified flavonoids, which can be regarded as cyclized derivatives of 3-prenylflavones: oxepinoflavones, pyranoflavones, dihydrobenzoxanthenes, furanodihydrobenzoxanthenes, and pyranodihydrobenzoxanthenes, and classes of flavonoid-derived xanthenes, which can be regarded as rearranged flavonoids: quinonobenzoxanthenes, cyclopentenoxanthenes, xanthonolides, dihydroxanthone, and cyclopentenochromone (Hakim, *et al.*, 2006). The structures of various classes of the regular, modified, and rearranged flavonoids were summarized in **Figures 2, 3 and 4**.

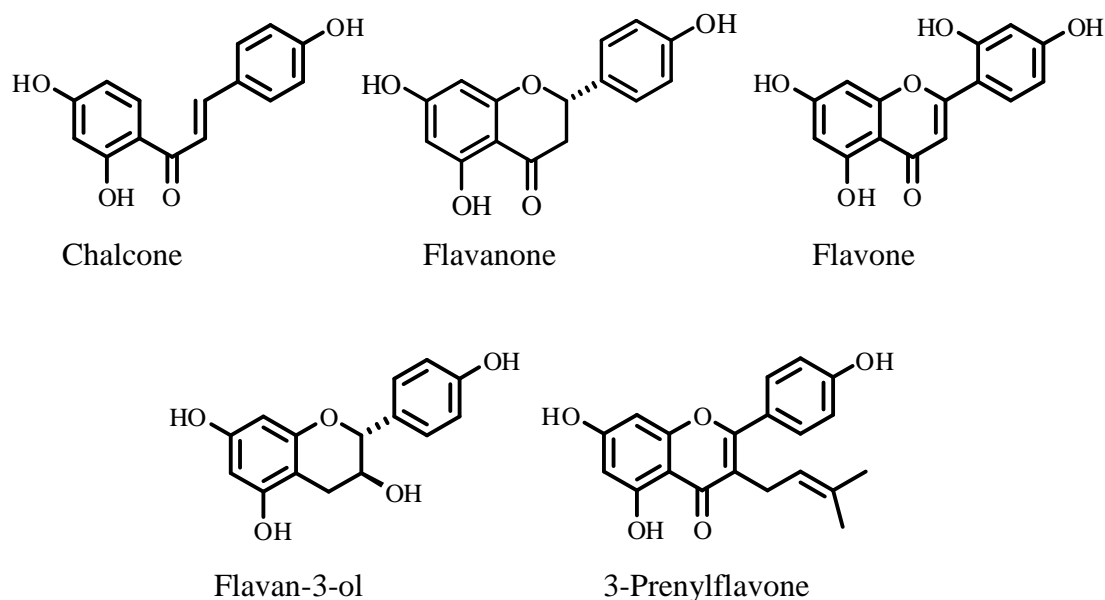


Figure 2 The structures of regular flavonoids in *Artocarpus*

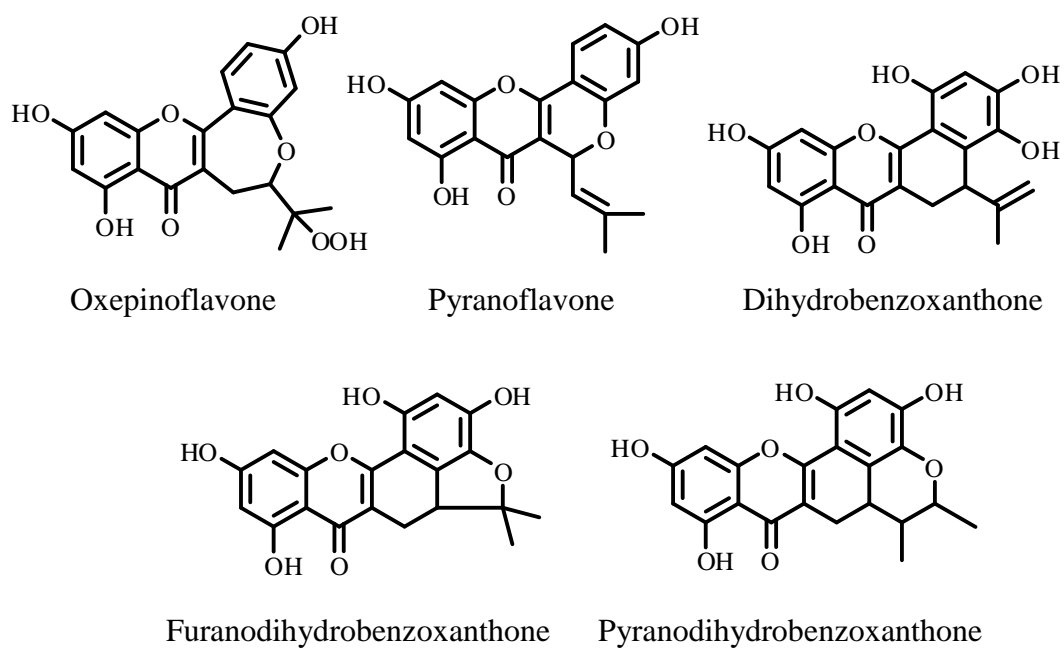


Figure 3 The structures of modified flavonoids in *Artocarpus*

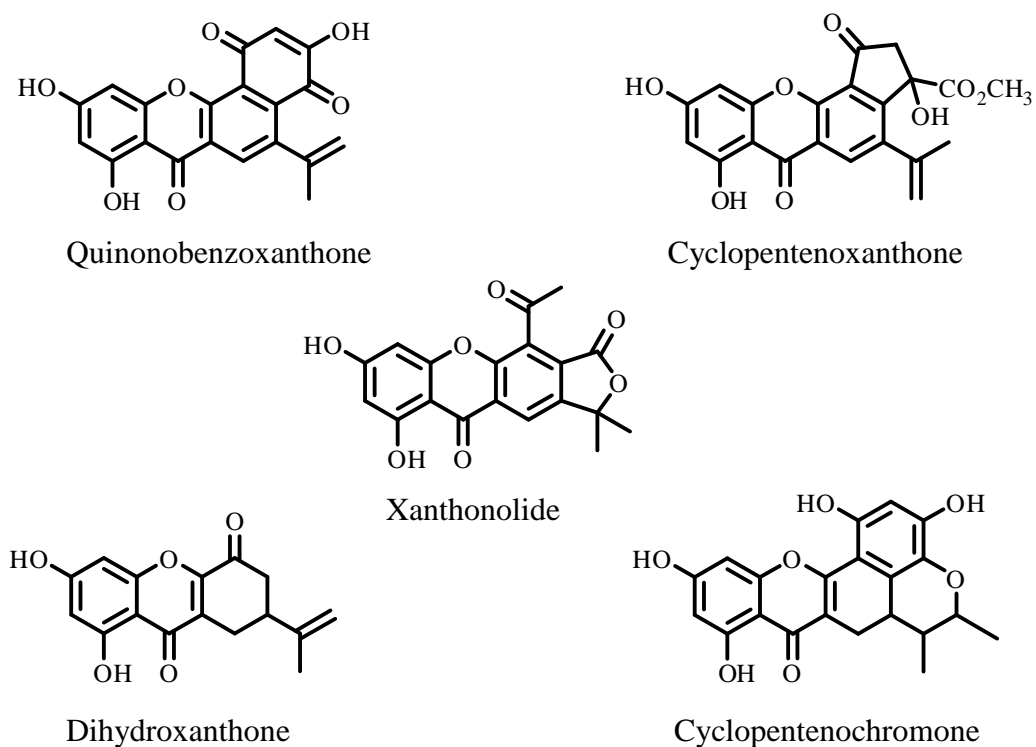


Figure 4 The structures of flavonoid-derived xanthenes in *Artocarpus*

In the previous report, prenylated flavones which have been found in *Artocarpus* genus, most frequently the hydroxyl groups are replaced (occasionally with methoxyl groups) in the C-5,7,2',4' or C-5,7,2',4',5' positions, and the isoprenyl group (occasionally geranyl group) at C-3, C-3,6, or C-3,6,8 positions in the A-ring. In the modified flavonoids, the isoprenyl substituent at C-3 position is always found in the form of a carbocyclic ring or an oxygen-bearing ring fused with rings B and C. In the rearranged-flavonoid group of compounds, the original ring B of the modified-flavonoid skeleton is found in the form of quinonoid moiety, or a five-membered ring, or in an opened, rearranged product.

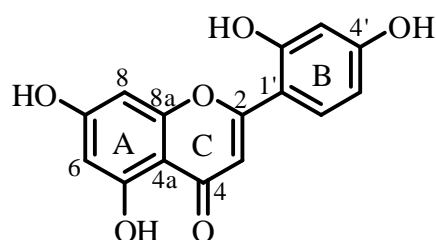


Figure 5 Flavone structure

Previously, a series of prenylated flavonoids were isolated from *Artocarpus* species, some of which showed interesting biological activities. These include cytotoxicity, antiplatelet action, antibacterial activities (Sultanbawa, *et al.*, 1989), strong radical scavenging properties towards the DPPH radical (Jayasinghe, *et al.*, 2008), and inhibitors of ROS and NO production (Cerqueira, *et al.*, 2008). A series of weakly cytotoxic (Wang, *et al.*, 2004) and antimalarial (Boonlaksiri, *et al.*, 2000) prenylated stilbenes and their derivatives were revealed.

The chemical constituents isolated from the *Artocarpus* genus were summarized in **Table 1** (The literature survey from SciFinder Scholar database).

Table 1 Compounds from the *Artocarpus* genus

Compounds	Bibliography
<p><i>I. A. altilis</i></p> <p>bud covers</p> <p>cycloaltilisin 6; 6b</p> <p>cycloaltilisin 7; 1d</p> <p>root bark</p> <p>friedelan-3β-ol; 1i</p> <p>friedelin; 2i</p> <p>artocarpin; 15e</p> <p>leaves</p> <p>1-(2,4-dihydroxyphenyl)-3-[8-hydroxy-2-methyl-2-(4-methyl-3-pentenyl)-2<i>H</i>-1-benzopyran-5-yl]-1-propanone; 1b</p> <p>1-(2,4-dihydroxyphenyl)-3-{4-hydroxy-6,6,9-trimethyl-6a,7,8,10a-tetrahydro-6<i>H</i>-dibenzo[<i>b,d</i>]pyran-5-yl}-1-propanone; 2b</p> <p>2-geranyl-2',3,4,4'-tetrahydroxydihydrochalcone; 3b</p> <p>1-(2,4-dihydroxyphenyl)-3-[3,4-dihydro-3,8-dihydroxy-2-methyl-2-(4-methyl-3-pentenyl)-2<i>H</i>-1-benzopyran-5-yl]-1-propanone; 4b</p> <p>1-(2,4-dihydroxyphenyl)-3-[8-hydroxy-2-methyl-2-(3,4-epoxy-4-methyl-1-pentenyl)-2<i>H</i>-1-benzopyran-5-yl]-1-propanone; 5b</p> <p>2'-geranyl-3',4',7-trihydroxyflavanone; 2d</p> <p>cycloaltilisin 6; 6b</p> <p>1-(2,4-dihydroxyphenyl)-3-[8-hydroxy-2-methyl-2-(4-hydroxy-4-methyl-2-pentenyl)-2<i>H</i>-1-benzopyran-5-yl]-1-propanone; 7b</p> <p>2-[6-hydroxy-3,7-dimethylocta-2(<i>E</i>),7-dienyl]-2',3,4,4'-tetrahydroxydihydrochalcone; 8b</p> <p>fruits</p> <p>oxyresveratrol; 6g</p> <p>artocarbene; 7g</p>	<p>Patil, <i>et al.</i>, 2002</p> <p>Fun, <i>et al.</i>, 2007</p> <p>Chantrapromma, <i>et al.</i>, 2007</p> <p>Wang, <i>et al.</i>, 2007</p> <p>Amarasinghe, <i>et al.</i>, 2008</p>

Table 1 (continue)

Compounds	Bibliography
<p>fruits</p> <p>moracin M; 4a</p> <p>norartocarpanone; 10d</p> <p>norartocarpetin; 4e</p> <p>isoartocarpesin; 2e</p> <p>3β-acetoxyolean-12-en-11-one; 4i</p> <p>cycloartenyl acetate; 3i</p> <p>sitosterol β-D-glucopyranoside; 1h</p>	<p>Amarasinghe, <i>et al.</i>, 2008</p>
<p>2. A. chama</p> <p>roots</p> <p>artochamin A; 49e</p> <p>artochamin B; 45e</p> <p>artochamin C; 26e</p> <p>artochamin D; 19e</p> <p>artochamin E; 56e</p> <p>artocarpin; 15e</p> <p>cycloartocarpin; 43e</p> <p>cudraflavone A; 54e</p> <p>artonin A; 75e</p> <p>artonin U; 8e</p> <p>cycloartobiloxanthone; 76e</p> <p>artonin E; 24e</p> <p>artocarpetin A; 5e</p>	<p>Wang, <i>et al.</i>, 2004</p>
<p>3. A. champeden</p> <p>tree bark</p> <p>cyclochampedol; 41e</p> <p>cycloeucalenol; 5i</p> <p>cycloartenone; 6i</p>	<p>Achmad, <i>et al.</i>, 1996</p>

Table 1 (continue)

Compounds	Bibliography
<p>tree bark</p> <p>artocarpin; 15e</p> <p>heteroflavanone A; 9d</p> <p>root bark</p> <p>artoindonesianin A; 79e</p> <p>artoinin A; 75e</p> <p>root trunk</p> <p>artoindonesianin B; 84e</p> <p>heartwood</p> <p>artoindonesianin Q; 14e</p> <p>artoindonesianin R; 13e</p> <p>artoindonesianin S; 57e</p> <p>artoindonesianin T; 58e</p> <p>artoindonesianin A-2; 39e</p> <p>artoindonesianin A-3; 61e</p> <p>artoinin B; 63e</p> <p>heterophyllin; 28e</p> <p>cudraflavone C; 16e</p>	<p>Parenti, <i>et al.</i>, 1998</p> <p>Hakim, <i>et al.</i>, 1999</p> <p>Hakim, <i>et al.</i>, 1999</p> <p>Syah, <i>et al.</i>, 2002</p> <p>Syah, <i>et al.</i>, 2006</p>
<p>4. A. communis</p> <p>bark</p> <p>artoinin E; 24e</p> <p>artoinin F; 77e</p> <p>cycloartobiloxanthone; 76e</p> <p>artoinol A; 2c</p> <p>artoinol B; 1j</p> <p>artoinol C; 67e</p> <p>artoinol D; 68e</p> <p>artoinol E; 65e</p>	<p>Hano, <i>et al.</i>, 1990</p> <p>Aida, <i>et al.</i>, 1997</p>

Table 1 (continue)

Compounds	Bibliography
<p>bark</p> <p>artonin K; 73e</p> <p>artobiloxanthone; 60e</p> <p>roots</p> <p>artocommunol CA; 52e</p> <p>artocommunol CB; 38e</p> <p>artocommunol CC; 88e</p> <p>artocommunol CD; 37e</p> <p>artocommunol CE; 32e</p> <p>cyclomorusin; 51e</p> <p>root bark</p> <p>cycloartomunin; 50e</p> <p>cycloartomunoxanthone; 78e</p> <p>dihydrocycloartomunin; 47e</p> <p>cudraflavone A; 54e</p> <p>artomunoxanthone; 59e</p> <p>artomunoxanthentrione; 3j</p> <p>cyclomulberrin; 46e</p> <p>cyclocommunol; 40e</p> <p>cyclocommunin; 44e</p> <p>dihydroisocycloartomunin; 48e</p> <p>heartwood</p> <p>3'',3''-dimethylpyrano[1',4']2,4,2'-trihydroxychalcone; 9b</p> <p>cycloartocarpin; 43e</p> <p>cudraflavone A; 54e</p> <p>isobacachalcone; 10b</p> <p>morachalcone A; 11b</p> <p>gemichalcone B; 20b</p> <p>artoindonesianin E; 11d</p>	<p>Aida, <i>et al.</i>, 1997</p> <p>Chan, <i>et al.</i>, 2003</p> <p>Lin, <i>et al.</i>, 1991</p> <p>Shieh, <i>et al.</i>, 1992</p> <p>Lin, <i>et al.</i>, 1992</p> <p>Han, <i>et al.</i>, 2006</p>

Table 1 (continue)

Compounds	Bibliography
<p>heartwood gemichalcone C; 21b artocarpin; 15e cudraflavone C; 16e licoflavone C; 6e (2<i>S</i>)-euchrenone a₇; 4d</p>	<p>Han, <i>et al.</i>, 2006</p>
<p>5. <i>A. elasticus</i> heartwood artocarpone A; 12e norartocarpin; 18e artocarpin; 15e artocarpone B; 71e cycloartocarpin; 43e wood artelastin; 42e artelastochromene; 53e artelasticin; 20e artelastinin; 86e artelastofuran; 36e cyclocommunin; 44e carpelastofuran; 87e roots bark artelastocarpin; 85e artelasticinol; 30e cycloartelastoxanthendiol; 81e cycloartelastoxanthone; 82e artelastoxanthone (7-demethylartonol E); 66e artonol A; 2c artelastoheterol; 31e</p>	<p>Kijjoa, <i>et al.</i>, 1996</p> <p>Kijjoa, <i>et al.</i>, 1996</p> <p>Kijjoa, <i>et al.</i>, 1998</p> <p>Ko, <i>et al.</i>, 2005</p>

Table 1 (continue)

Compounds	Bibliography
<p>6. <i>A. gomezianus</i> not specified cyclomorusin; 51e cycloartocarpin; 43e artocarpin; 15e norartocarpetin; 4e cudraflavone C; 16e</p> <p>roots artogomezianol; 2g andalasin A; 1g</p> <p>tree bark artoindonesianin N; 3g artoindonesianin O; 3a oxyresveratrol; 6g</p>	<p>Likhitwitayawuid, <i>et al.</i>, 2000</p> <p>Likhitwitayawuid, <i>et al.</i>, 2001</p> <p>Hakim, <i>et al.</i>, 2002</p>
<p>7. <i>A. heterophyllus</i> root bark heteroflavanone A; 9d heteroflavanone B; 5d artoinin A; 75e artoinin B; 63e artoinin C; 22b artoinin D; 23b artoinin I; 11e artoinin J; 72e artoinin K; 73e artoinin L; 74e heterophyllol; 8g heteroflavanone C; 6d cudraflavone A; 54e</p>	<p>Lu, <i>et al.</i>, 1993</p> <p>Aida, <i>et al.</i>, 1993</p> <p>Aida, <i>et al.</i>, 1993</p> <p>Lin, <i>et al.</i>, 1993</p> <p>Lin, <i>et al.</i>, 1995</p>

Table 1 (continue)

Compounds	Bibliography
<p>root bark</p> <p>cycloheterophyllin; 55e</p> <p>artocarpetin B; 7e</p> <p>heteroartonin A; 22e</p> <p>kuwanon T; 23e</p> <p>heartwood of the root</p> <p>artocapanone A; 8d</p> <p>cycloartocarpin; 43e</p> <p>artocarpanone; 7d</p> <p>artocarpetin; 3e</p> <p>norartocarpetin; 4e</p> <p>artocarpin; 15e</p> <p>artocarpesin; 1e</p> <p>dihydromorin; 14d</p>	<p>Chung, <i>et al.</i>, 1995</p> <p>Lin, <i>et al.</i>, 1995</p>
<p>8. <i>A. incisus</i></p> <p>heartwood</p> <p>artocarbene; 7g</p> <p>leaves</p> <p>3-geranyl-2,3',4,4'-tetrahydroxychalcone; 13b</p>	<p>Shimizu, <i>et al.</i>, 1997</p> <p>Shimizu, <i>et al.</i>, 1997</p>
<p>9. <i>A. lakoocha</i></p> <p>roots</p> <p>lakoochin A; 1a</p> <p>lakoochin B; 2a</p>	<p>Puntumchai, <i>et al.</i>, 2004</p>
<p>10. <i>A. lanceifolius</i></p> <p>heartwood</p> <p>artoindonesianin G; 33e</p> <p>artoindonesianin H; 35e</p> <p>artoindonesianin I; 34e</p> <p>artelastofuran; 36e</p>	<p>Syah, <i>et al.</i>, 2001</p>

Table 1 (continue)

Compounds	Bibliography
artelastacin; 20e tree bark artoidonesianin P; 80e artoidonesianin V; 62e artobiloxanthone; 60e cycloartobiloxanthone; 76e artanol B; 1j	Hakim, <i>et al.</i> , 2002
11. A. nobilis bark artobiloxanthone; 60e cycloartobiloxanthone; 76e artobilochromen; 10e leaves xanthoangelol; 12b xanthoangelol B; 15b 2',4',4-trihydroxy-3'-[2-hydroxy-7-methyl-3-methylene-6-octaenyl]chalcone; 17b 3-geranyl-2,3',4,4'-tetrahydroxychalcone; 13b 2',3,4,4'-tetrahydroxy-3'-[6-hydroxy-3,7-dimethyl-2(<i>E</i>),7-octadienyl]chalcone; 16b fruits 2,4,4'-trihydroxy-3-[(2 <i>E</i>)-5-methoxy-3,7-dimethylocta-2,6-dienyl]chalcone; 14b 1-(3,4-dihydro-3,5-dihydroxy-2-methyl-2(3-methyl-2-buteryl)-2 <i>H</i> -1-benzopyran-6-yl)-3-(4-hydroxyphenyl)-2(<i>E</i>)-propen-1-one; 18b 8-geranyl-3',4',7-trihydroxyflavone; 9e 3'-geranyl-4',5,7-trihydroxyflavanone; 12d	Sultanbawa, <i>et al.</i> , 1989 Jayasinghe, <i>et al.</i> , 2004 Jayasinghe, <i>et al.</i> , 2006

Table 1 (continue)

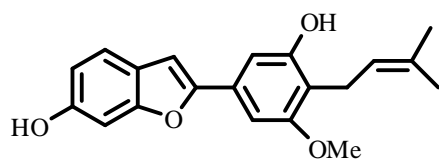
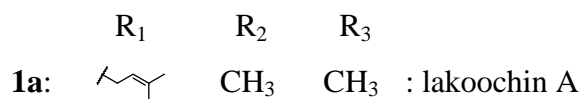
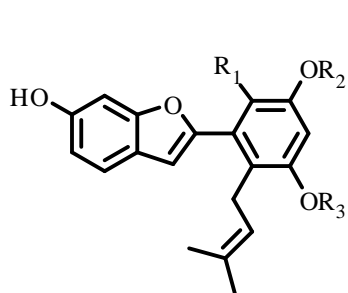
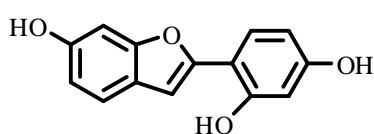
Compounds	Bibliography
<p>fruits</p> <p>xanthoangelol; 12b</p> <p>xanthoangelol B; 15b</p> <p>3-geranyl-2,3',4,4'-tetrahydrochalcone; 13b</p> <p>lespeol; 19b</p> <p>8-geranyl-4',7-dihydroxyflavone; 3d</p> <p>isonymphaeol-B; 13d</p> <p>root bark</p> <p>artoinin E 2'-methylether; 25e</p> <p>isoartoinin E 2'-methylether; 29e</p> <p>dihydroisoartoinin E 2'-methylether; 17e</p> <p>artoinin V 2'-methylether; 21e</p> <p>artobiloxanthone; 60e</p> <p>artoinin E; 24e</p> <p>cycloartobiloxanthone; 76e</p>	<p>Jayasinghe, <i>et al.</i>, 2006</p> <p>Jayasinghe, <i>et al.</i>, 2008</p>
<p>12. A. rigida</p> <p>bark</p> <p>artoinin M; 83e</p> <p>artoinin N; 64e</p> <p>artoinin O; 69e</p> <p>artoinin P; 70e</p>	<p>Hano, <i>et al.</i>, 1993</p>
<p>13. A. rigidus subsp. rigidus</p> <p>root bark</p> <p>7-demethylartoinol E; 66e</p> <p>artorigidusin; 1c</p> <p>artoinol B; 1j</p> <p>artoinin F; 77e</p> <p>cycloartobiloxanthone; 76e</p>	<p>Namdaung, <i>et al.</i>, 2006</p>

Table 1 (continue)

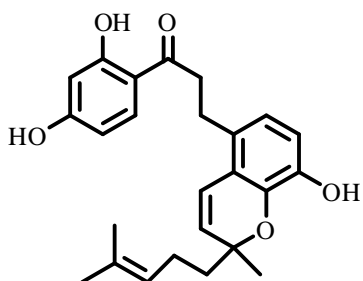
Compounds	Bibliography
morin; 1f 14. A. teysmanii root bark artoindonesianin C; 2j artoindonesianin U; 27e cycloartobiloxanthone; 76e artonin J; 72e <i>trans</i> -4-isopentenyl-3,5,2',4'-tetrahydroxystilbene; 4g <i>trans</i> -4-(3-methyl-E-but-1-enyl)-3,5,2',4'-tetrahydroxystilbene; 5g	Makmur, <i>et al.</i> , 1999

Structures of compounds from *Artocarpus* genus

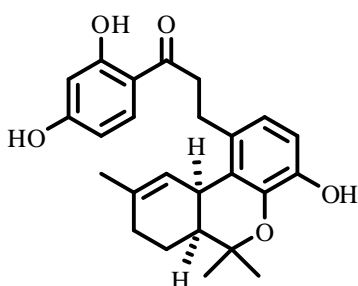
a. 2-arylbenzofurans

**3a:** artoindonesianin O**4a:** moracin M

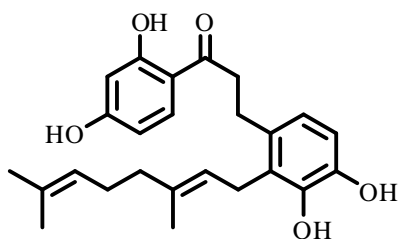
b. Chalcones



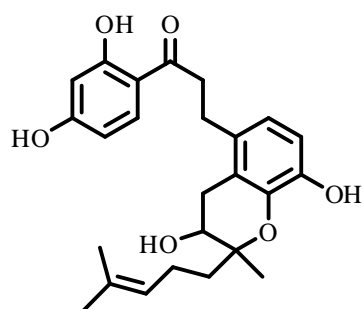
1b: 1-(2,4-dihydroxyphenyl)-3-[8-hydroxy-2-methyl-2-(4-methyl-3-pentenyl)-2*H*-1-benzopyran-5-yl]-1-propanone



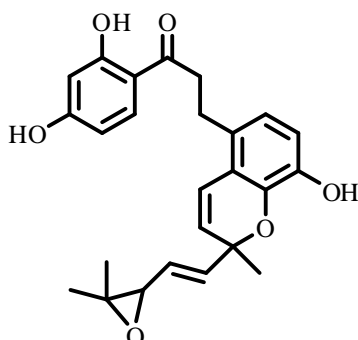
2b: 1-(2,4-dihydroxyphenyl)-3-{4-hydroxy-6,6,9-trimethyl-6a,7,8,10a-tetrahydro-6*H*-dibenzo[*b,d*]pyran-5-yl}-1-propanone



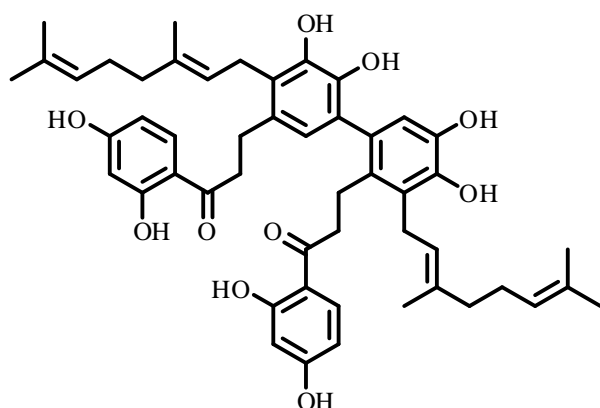
3b: 2-geranyl-2',3,4,4'-tetrahydroxydihydrochalcone



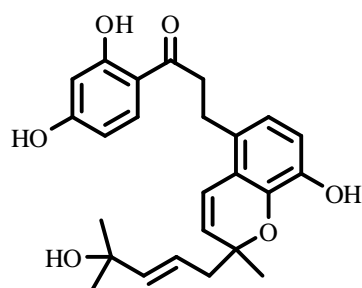
4b: 1-(2,4-dihydroxyphenyl)-3-[3,4-dihydro-3,8-dihydroxy-2-methyl-2-(4-methyl-3-pentenyl)-2*H*-1-benzopyran-5-yl]-1-propanone



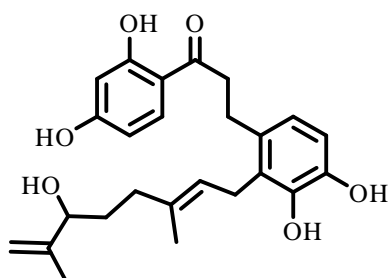
5b: 1-(2,4-dihydroxyphenyl)-3-[8-hydroxy-2-methyl-2-(3,4-epoxy-4-methyl-1-pentenyl)-2*H*-1-benzopyran-5-yl]-1-propanone



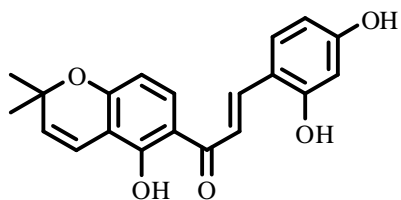
6b: cycloaltilisin 6



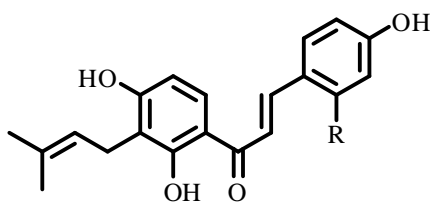
7b: 1-(2,4-dihydroxyphenyl)-3-[8-hydroxy-2-methyl-2-(4-hydroxy-4-methyl-2-pentenyl)-2*H*-1-benzopyran-5-yl]-1-propanone



8b: 2-[6-hydroxy-3,7-dimethylocta-2(*E*),7-dienyl]-2',3,4,4'-tetrahydroxydihydrochalcone

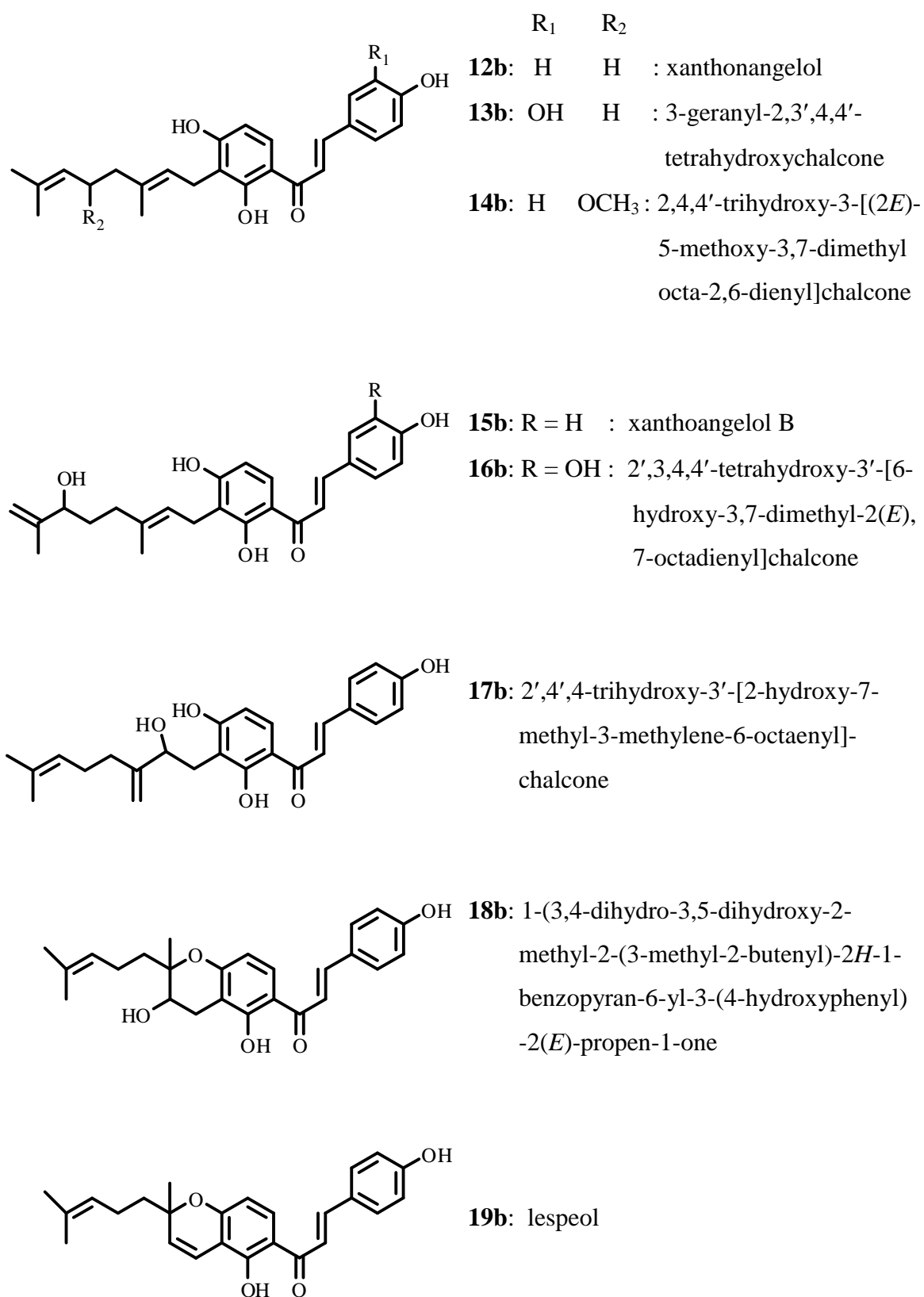


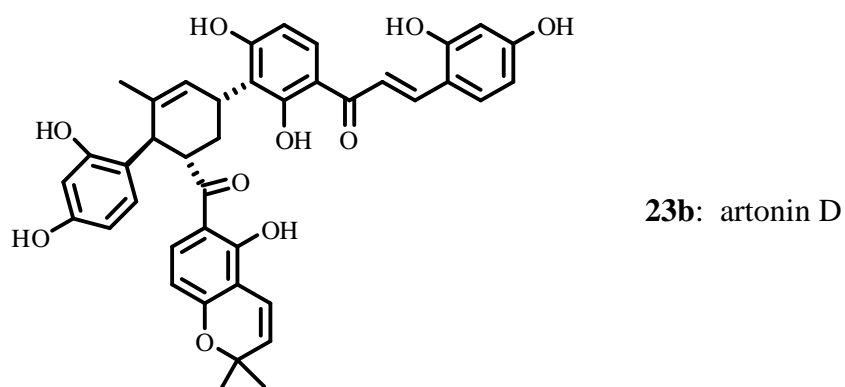
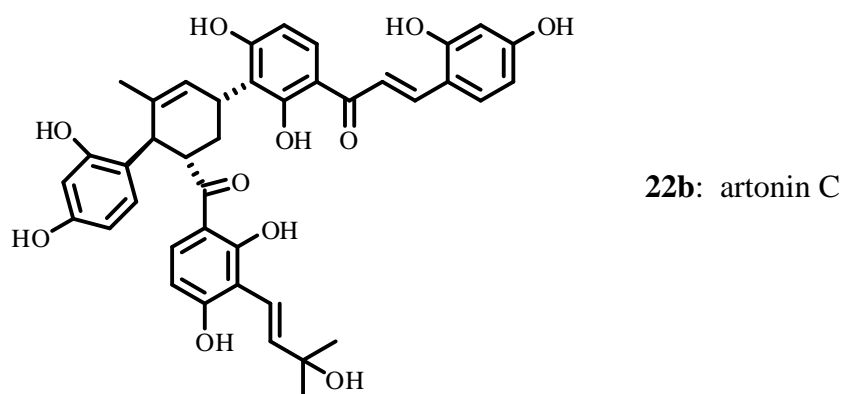
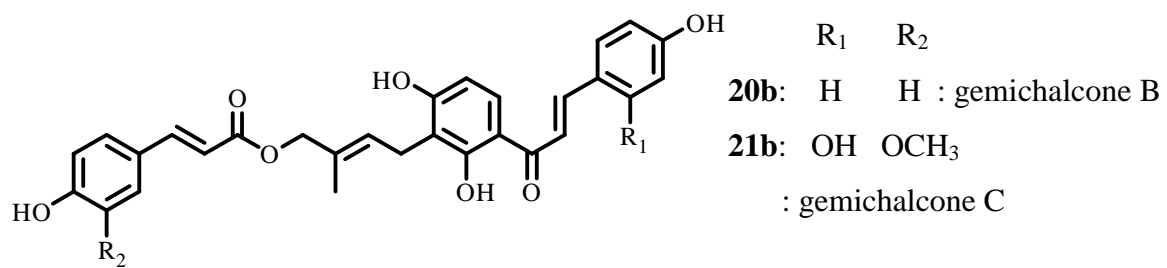
9b: 3', 3''-dimethylpyrano[3',4']2,4,2'-trihydroxychalcone



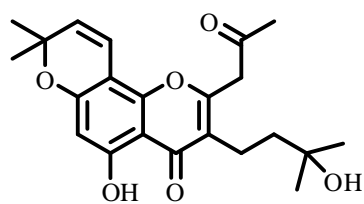
10b: R = H : isobacachalcone

11b: R = OH : morachalcone A

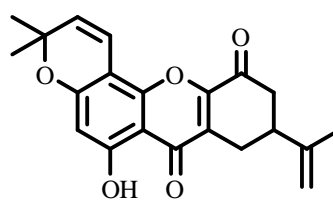




c. Chromones

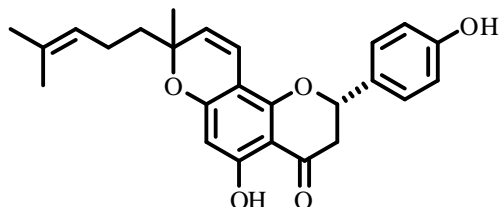


1c: artorigidusin

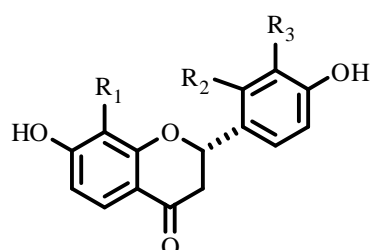


2c: artonol A

d. Flavanones

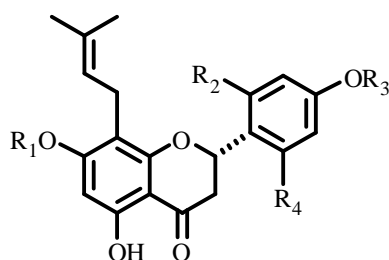


1d : cycloaitilisin 7



	R ₁	R ₂	R ₃	
2d :	H		OH	: 2'-geranyl-3',4',7-trihydroxyflavanone

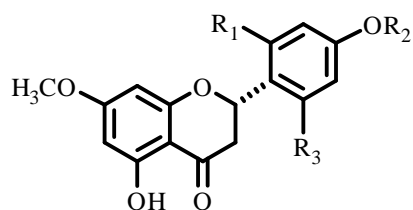
3d :		H	H	: 8-geranyl-4',7-dihydroxyflavanone
-------------	--	---	---	-------------------------------------



	R ₁	R ₂	R ₃	R ₄	
4d :	H	OH	H	H	: (2 <i>S</i>)-euchrenone a ₇

5d :	CH ₃	OCH ₃	CH ₃	OCH ₃	: heteroflavanone B
-------------	-----------------	------------------	-----------------	------------------	---------------------

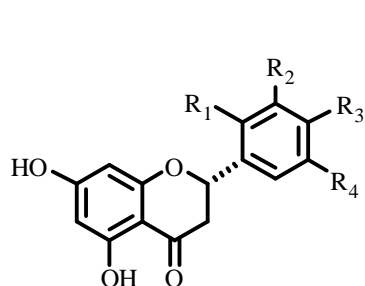
6d :	H	OCH ₃	CH ₃	OCH ₃	: heteroflavanone C
-------------	---	------------------	-----------------	------------------	---------------------



	R ₁	R ₂	R ₃	
7d :	H	H	H	: artocarpanone

8d :	H	CH ₃	H	: artocarpanone A
-------------	---	-----------------	---	-------------------

9d :	CH ₃	CH ₃	OCH ₃	: heteroflavanone A
-------------	-----------------	-----------------	------------------	---------------------

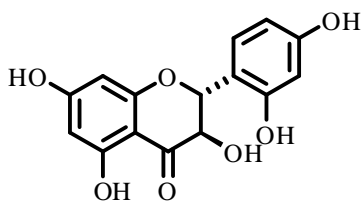
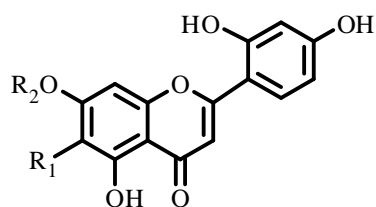


	R ₁	R ₂	R ₃	R ₄	
10d :	OH	H	OH	H	: norartocarpanone

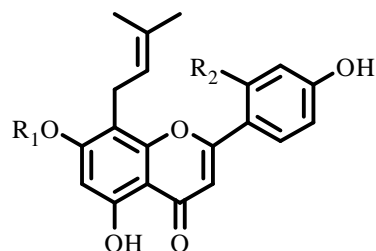
11d :	OCH ₃	H	CH ₃	OCH ₃	: artoindonesianin E
--------------	------------------	---	-----------------	------------------	----------------------

12d :	H		H	H	: 3'-geranyl-4',5,7-trihydroxyflavanone
--------------	---	--	---	---	---

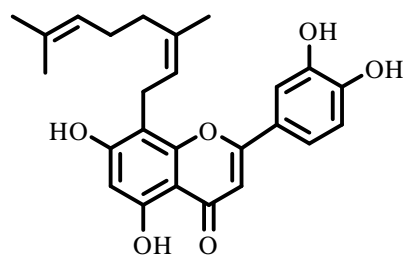
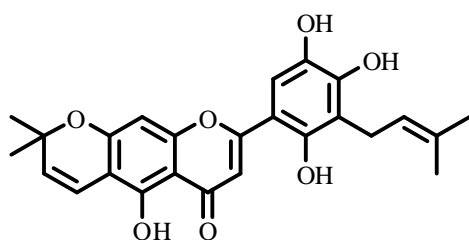
13d :	OH	OH	OH		: isonymphaeol-B
--------------	----	----	----	--	------------------

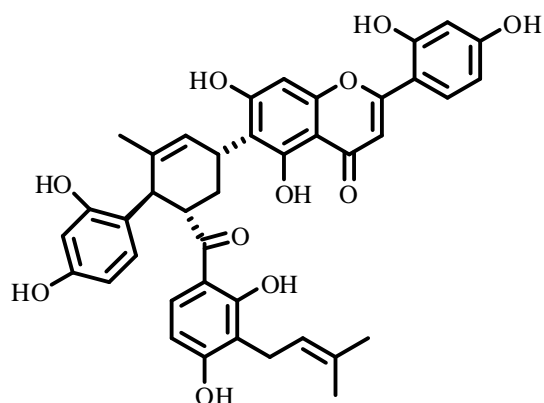
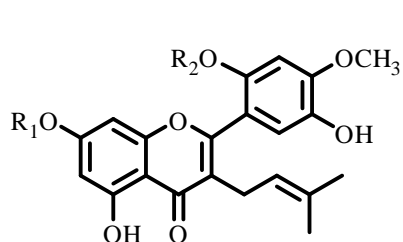
**14d:** dihydromorin**e. Flavones**

	R ₁	R ₂	
1e:		H	: artocarpesin
2e:		H	: isoartocarpesin
3e:	H	CH ₃	: artocarpetin
4e:	H	H	: norartocarpetin

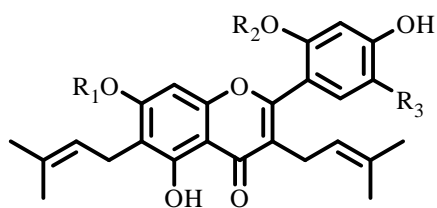


	R ₁	R ₂	
5e:	CH ₃	OH	: artocarpetin A
6e:	H	H	: licoflavone C
7e:	CH ₃	OCH ₃	: artocarpetin B
8e:	CH ₃	H	: artonin U

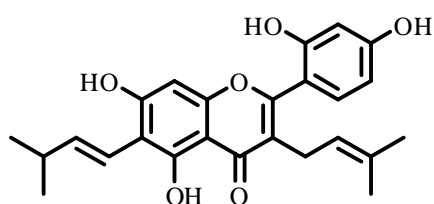
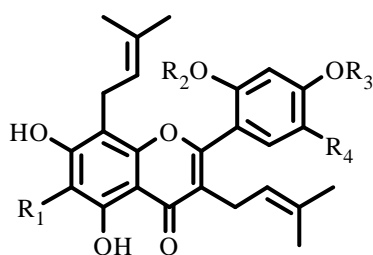
**9e:** 8-geranyl-3',4',7-trihydroxyflavone**10e:** artobilochromen


**11e:** artonin I

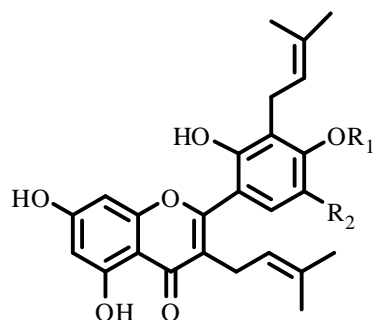
	R ₁	R ₂	
12e:	H	H	: artocarpone A
13e:	H	CH ₃	: artocarpone R
14e:	CH ₃	H	: artocarpone Q



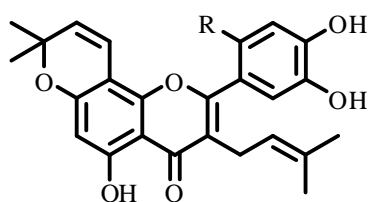
	R ₁	R ₂	R ₃	
15e:	CH ₃	H	H	: artocarpin
16e:	H	H	H	: cudraflavone C
17e:	CH ₃	CH ₃	OH	: dihydroisoartocarpin E 2'-methylether

**18e:** norartocarpin

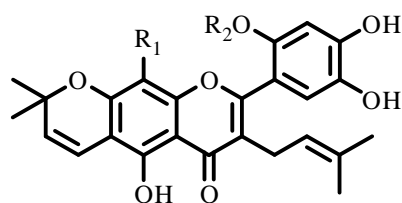
	R ₁	R ₂	R ₃	R ₄	
19e:	H	H	CH ₃	OH	: artochamin D
20e:		H	H	H	: artelasticin
21e:	H	CH ₃	H	OH	: artonin V 2'-methylether



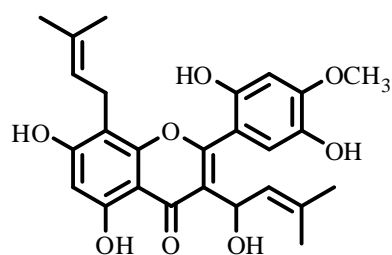
- | | R ₁ | R ₂ | |
|-------------|-----------------|----------------|-------------------|
| 22e: | CH ₃ | OH | : heteroartonin A |
| 23e: | H | H | : kuwanon T |



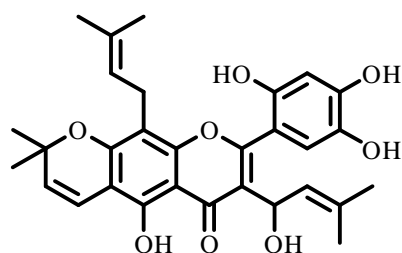
- | | | |
|-------------|----------------------|----------------------------|
| 24e: | R = OH | : artonin E |
| 25e: | R = OCH ₃ | : artonin E 2'-methylether |
| 26e: | R = H | : artochamin C |



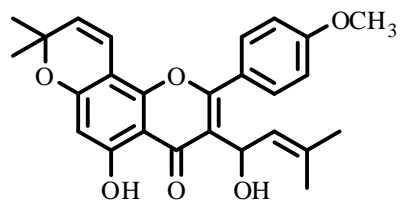
- | | R ₁ | R ₂ | |
|-------------|----------------|-----------------|-------------------------------|
| 27e: | | H | : artoindonesianin U |
| 28e: | | H | : heterophyllin |
| 29e: | H | CH ₃ | : isoartonin E 2'-methylether |



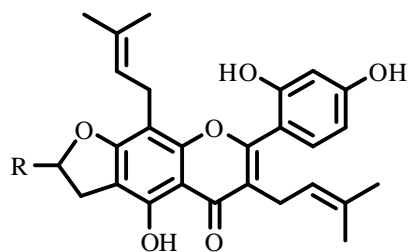
30e: artelastincol





31e: artelasheterol

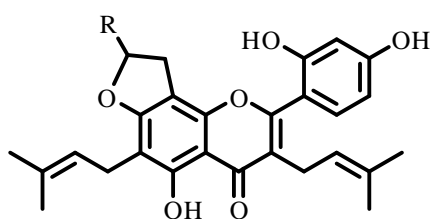



32e: artocommunol CE




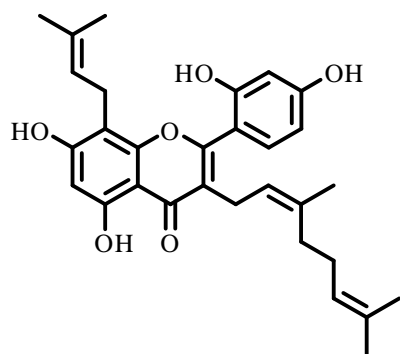
33e: R =  : artindonesianin G

34e: R =  : artindonesianin I

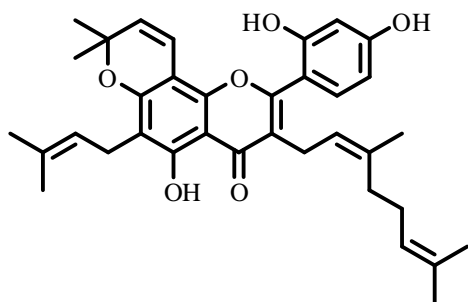


35e: R =  : artindonesianin H

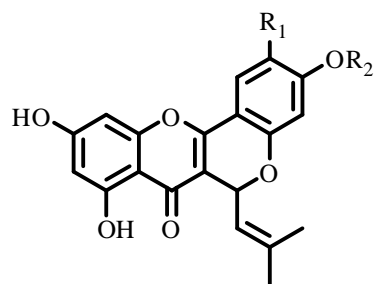
36e: R =  : artelastofuran



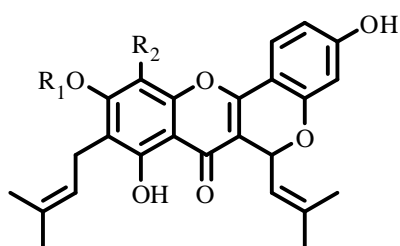
37e: artocommunol CD



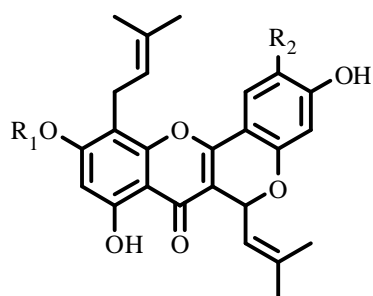
38e: artocommunol CB



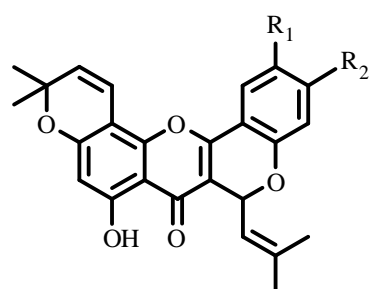
	R ₁	R ₂	
39e:	OH	CH ₃	: artoidonesianin A-2
40e:	H	H	: cyclocommunol
41e:	H	OH	: cyclochampedol



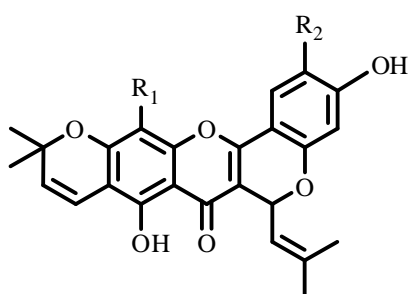
	R ₁	R ₂	
42e:	H		: artelastin
43e:	CH ₃	H	: cycloartocarpin
44e:	H	H	: cyclocommunin



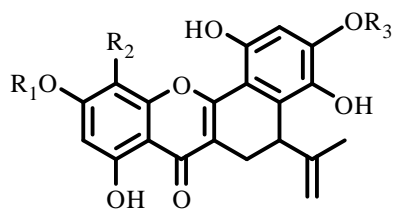
	R ₁	R ₂	
45e:	H	H	: artochamin B
46e:	H	H	: cyclomulberrin
47e:	CH ₃	H	: dihydrocycloartomunin
48e:	H	OCH ₃	: dihydroisocycloartomunin



	R ₁	R ₂	
49e:	OH	OH	: artochamin A
50e:	OH	OCH ₃	: cycloartomunin
51e:	H	OH	: cyclomorusin
52e:	H	OCH ₃	: artocommunol CA



	R ₁	R ₂	
53e:		H	: artelastochromene
54e:	H	H	: cudraflavone A
55e:		OH	: cycloheterophyllin

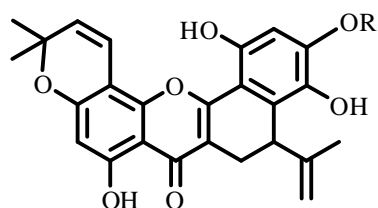


R₁ R₂ R₃

56e: H  H : artochamin E

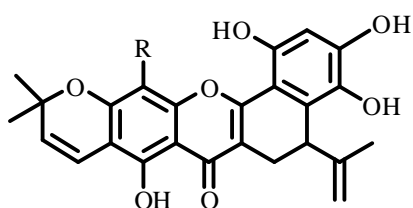
57e: CH₃ H CH₃ : artoindonesianin S

58e: H H CH₃ : artoindonesianin T

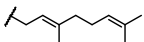



59e: R = CH₃ : artomunoxanthone

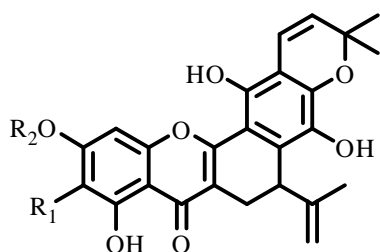
60e: R = OH : artobiloxanthone




61e: R = H : artoindonesianin A-3

62e: R =  : artoindonesianin V

63e: R =  : artonin B

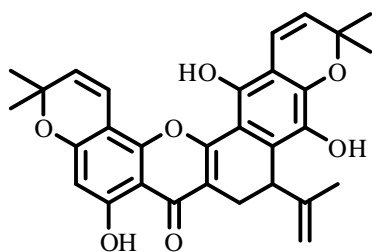


R₁ R₂

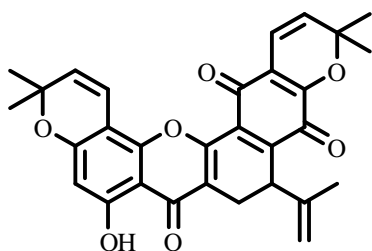
64e: H  : artonin N

65e: H OH : artonol E

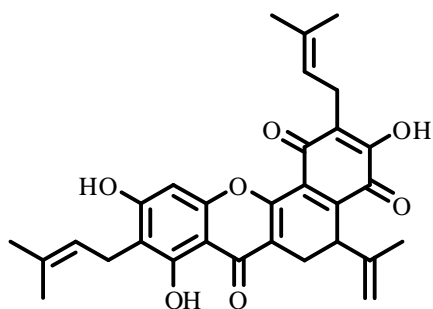
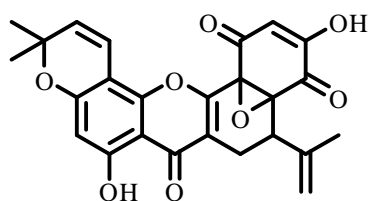
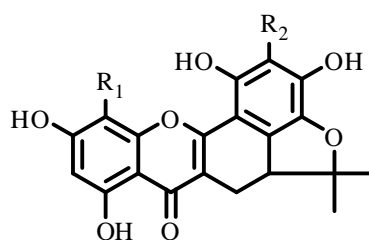
66e: H H : 7-demethylartonol E





67e: artonol C

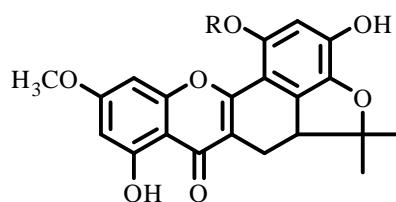
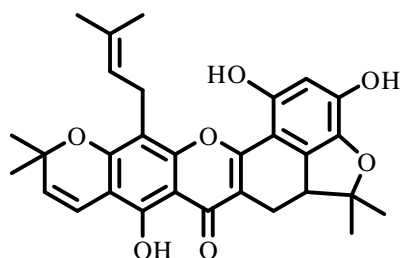
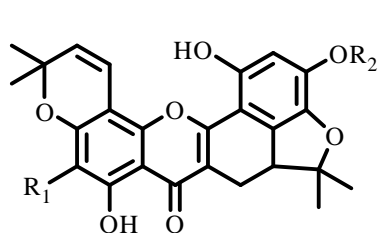


68e: artonol D

**69e:** artonin O**70e:** artonin P

71e: R_1 R_2
 H : artocarpone B

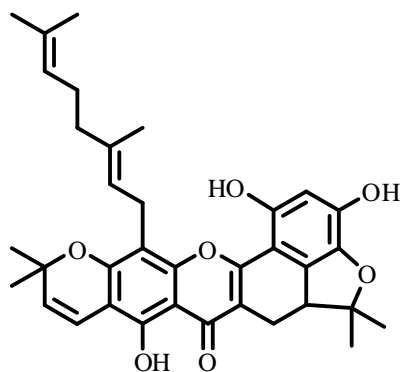
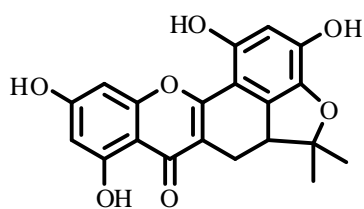
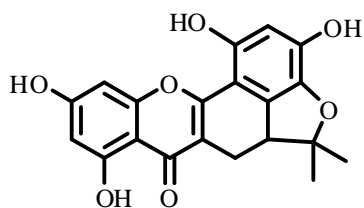
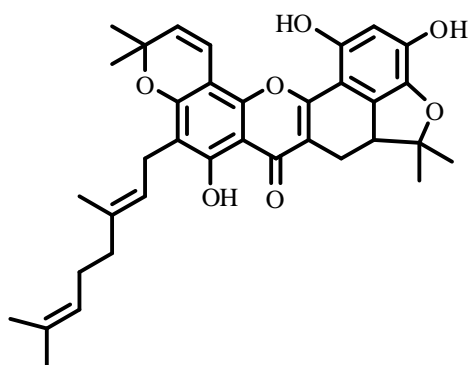
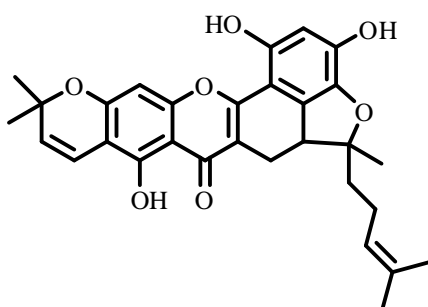
72e: H  : artonin J

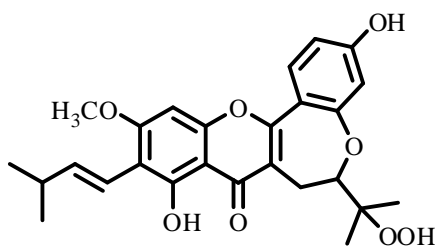
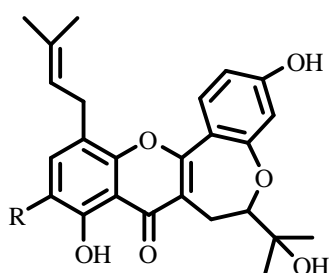


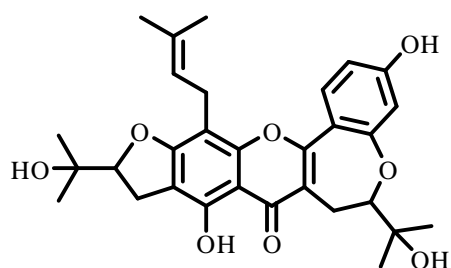
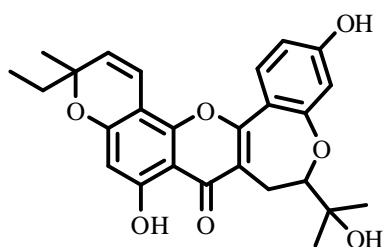
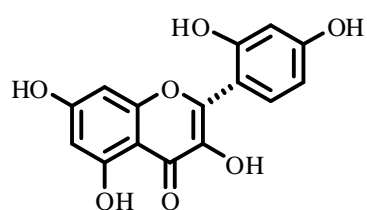
**73e:** R = H : artonin K**74e:** R = CH₃ : artonin L**75e:** artonin A

76e: R_1 R_2
 H H : cycloartobiloxanthone

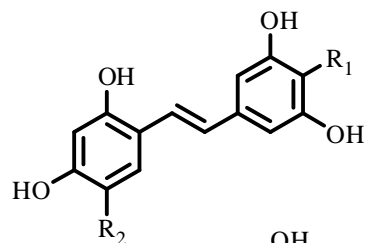
77e:  H : artonin F

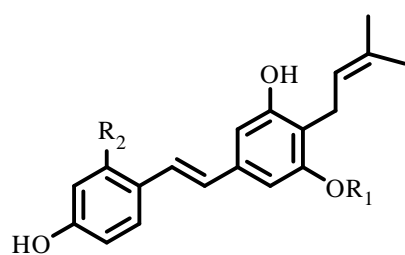
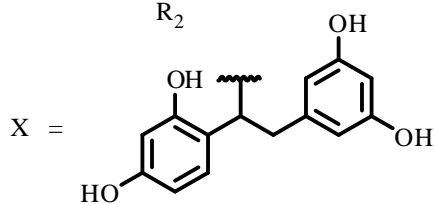
78e: H CH₃ : cycloartomunoxanthone

**79e:** artoindonesianin A**80e:** artoindonesianin P**81e:** cycloartelastoxanthendiol**82e:** cycloartelastoxanthone**83e:** artonin M

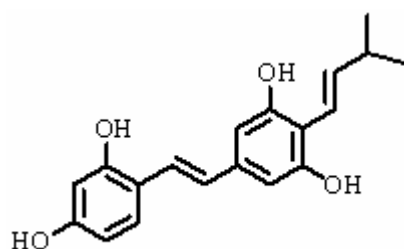
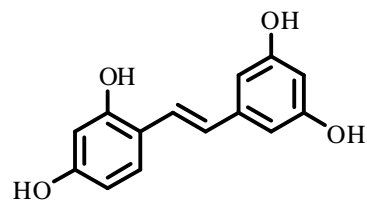
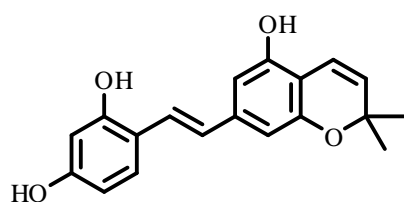
**84e:** artoindonesianin B**85e:** R =  : artelastocarpin**86e:** R =  : artelastinin**87e:** carpelastofuran**88e:** artoccommunol CC**f. Flavonol****1f:** morin

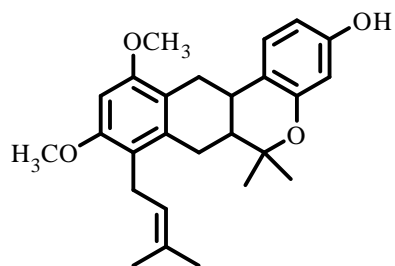
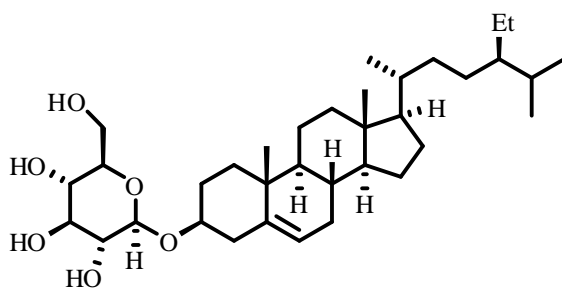
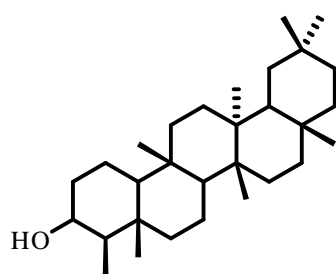
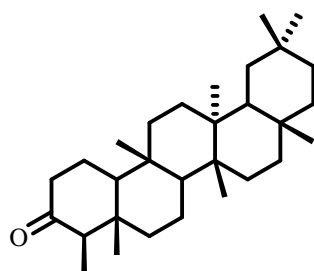
g. Stilbenes

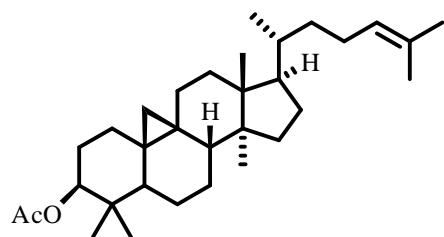
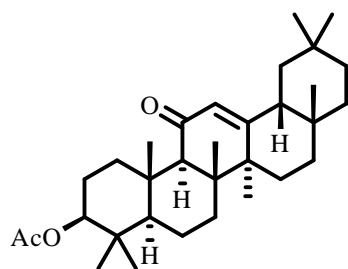
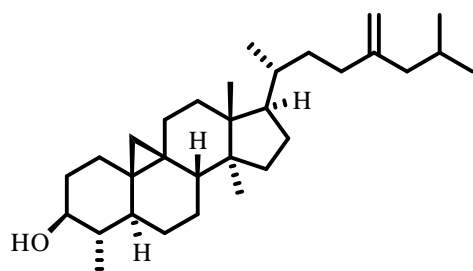
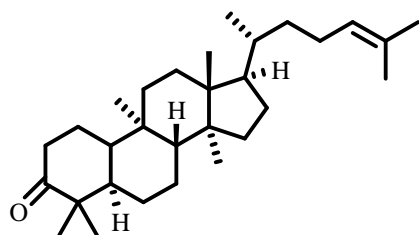
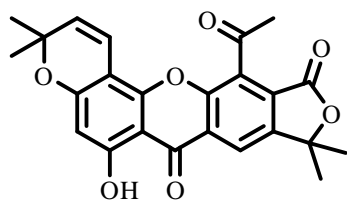

 $R_1 \quad R_2$
1g: X H : andalasin A

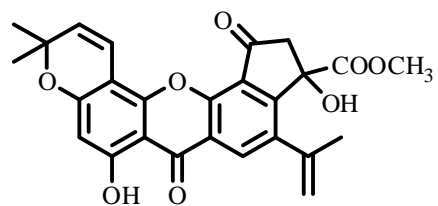
2g: H X : artogomezianol

 $R_1 \quad R_2$
3g: CH₃ H : artoindonesianin N

4g: H OH

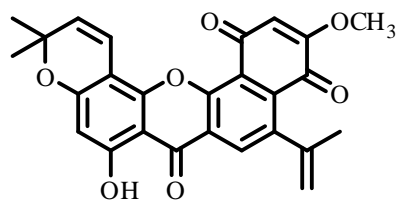
: *trans*-4-isopentenyl-3,5,2',4'-tetrahydroxystilbene

5g: *trans*-4-(3-methyl-*E*-but-1-enyl)-3,5,2',4'-tetrahydroxystilbene

6g: oxyresveratrol

7g: artocarbene

**8g:** heterophyllol**h. Steroids****1h:** sitosterol β -D-glucopyranoside**i. Triterpenes****1i:** friedelan-3 β -ol**2i:** friedelin

**3i:** cycloartenyl acetate**4i:** 3 β -acetoxylean-12-en-11-one**5i:** cycloeucalenol**6i:** cycloartenone**j. Xanthenes****1j:** artonol B



2j: artoindonesianin C



3j: artomunoxanthentrione

1.3 Objective

The objective of this work was to investigate the chemical constituents from the bark of *A. elasticus*.

CHAPTER 2

EXPERIMENTAL

2.1 General Method

Column chromatography was performed on silica gel 100 (70-230 Mesh ASTM, Merck) or SephadexTM LH-20 (Amersham Biosciences, Sweden). Quick column chromatography utilized silica gel 60 (230-400 Mesh ASTM, Merck). Aluminum sheets of silica gel 60 F₂₅₄ (layer thickness 0.2 mm, Merck) were used for thin-layer chromatography (TLC) and the compounds were visualized under ultraviolet light. Solvents for extraction and chromatography were distilled at their boiling ranges prior to use. Melting points were determined on the Fisher-John melting point apparatus. Ultraviolet spectra were measured with UV-160A spectrophotometer (SHIMADZU). Principle bands (λ_{\max}) were recorded as wavelengths (nm) and $\log \epsilon$ in MeOH and CH₂Cl₂ solution. Infrared spectra (IR) were obtained on a FT-IR spectrum BX spectrophotometer and were recorded in wave number (cm⁻¹). ¹H- and ¹³C-Nuclear magnetic resonance spectra were recorded on an FT-NMR Bruker Ultra ShieldTM 300 MHz spectrometer. Spectra were recorded in CDCl₃, DMSO and were recorded as δ value in ppm downfield from TMS (internal standard δ 0.00). Low and high resolution mass spectra were recorded on a MAT 95 XL at Scientific Equipment Center, Prince of Songkla University.

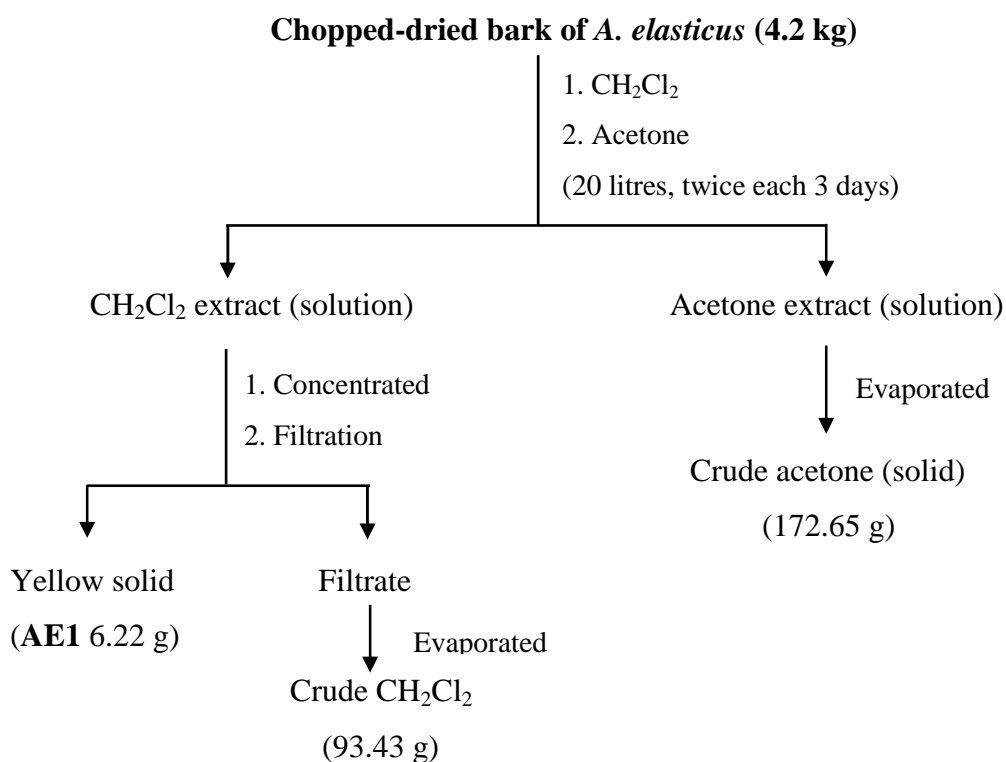
2.2 Plant Material

The bark of *A. elasticus* (Moraceae) was collected from Amphur Kuraburi, Phang Nga Province in the southern part of Thailand, in July 2007. Identification was made by Mr. Charernsak Saewai, Department of Biology, Faculty of Science, Prince of Songkla University. The specimen (A. Yanya 1Phang-nga: Kuraburi 2/4/2009) have been deposited in the Herbarium of Department of Biology,

Faculty of Science, Prince of Songkla University, Hatyai, Songkhla, Thailand.

2.3 Extraction and isolation

Chopped-dried bark of *A. elasticus* (4.2 kg) was immersed at room temperature in dichloromethane and acetone (each extract 3 days), successively. The solution of dichloromethane extract was concentrated under reduced pressure to produce a yellow solid upon standing overnight at room temperature. The solid **AE1** (6.22 g) was collected by filtration and the filtrate was further evaporated to give the dark-red viscous liquid (93.43 g). The solution of acetone extract was evaporated to give dark-red solid (172.65 g). The process of extraction was shown in **Scheme 1**.



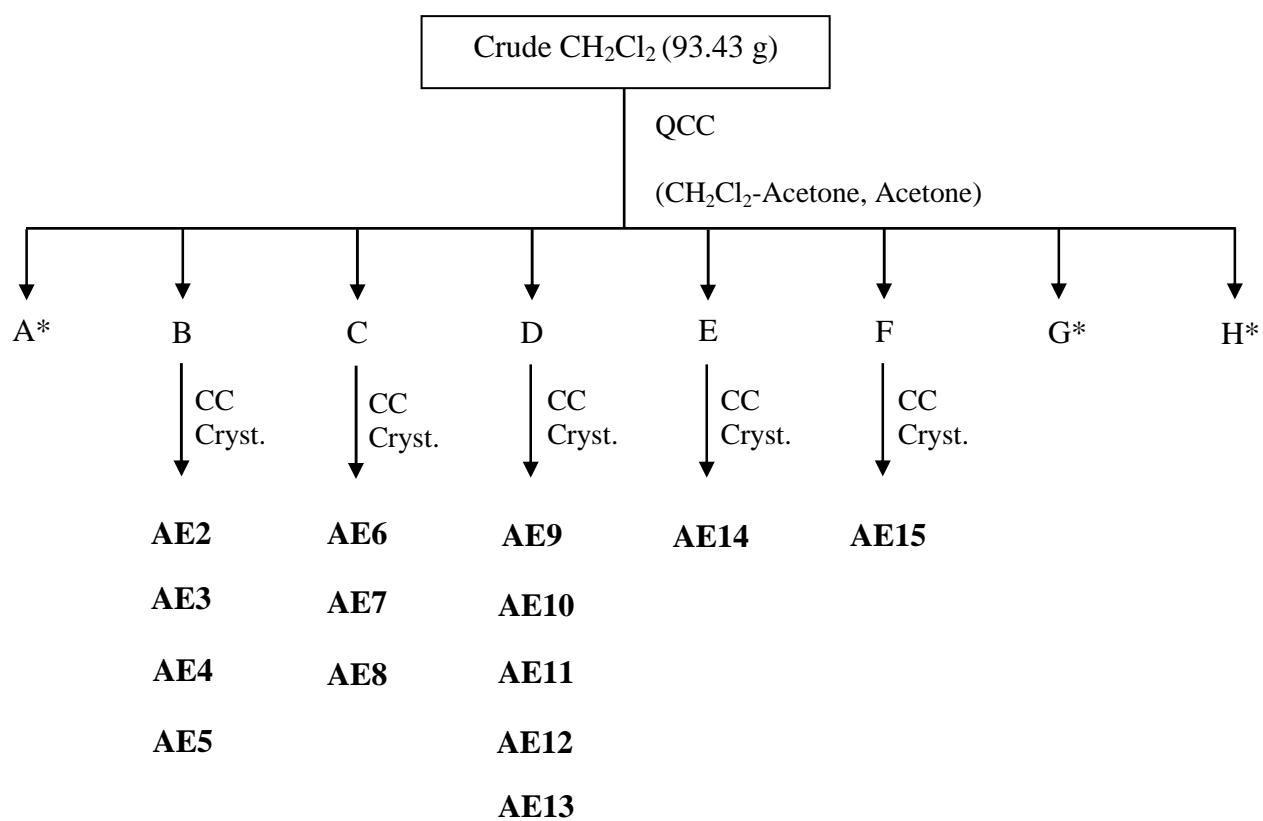
Scheme 1 Extraction of the crude extracts from the bark of *A. elasticus*

2.3.1 Purification of dichloromethane extract of the bark of *A. elasticus*

The solution of dichloromethane extract was concentrated under reduced pressure to produce a yellow solid upon standing overnight at room temperature. The solid **AE1** (6.22 g) was collected by filtration and the filtrate was further evaporated to give the dark-red viscous liquid (93.43 g). Which was subjected to quick column chromatography on silica gel, and eluted with CH₂Cl₂-acetone (13:1), and acetone (300 ml each). The fractions that showed the same major components on TLC were combined to yield fractions A-H. Further purification of each fraction gave fourteen pure compounds (**Scheme 2**).

Table 2 Physical characteristic and weight of fractions obtained from QCC of the crude CH₂Cl₂ extract

Fraction	Weight (g)	Appearance
A	15.8796	yellow gel
B	16.7920	brown gel
C	29.8941	dark-red viscous liquid
D	9.0987	dark-red viscous liquid
E	1.7163	dark-red viscous liquid
F	2.8492	dark-red viscous liquid
G	2.8205	brown solid
H	3.2393	brown solid



*No further investigation

Scheme 2 Isolation of compounds **AE2-AE15** from dichloromethane extract of the bark of *A. elasticus*

Isolation of AE1

The solution of the dichloromethane extract was concentrated under reduced pressure to produce a yellow solid upon standing overnight at room temperature. The solid **AE1** (6.22 g) was collected by filtration.

AE1

5-Hydroxy-8,8-dimethyl-3-(3-methyl-2-butenyl)-2-(2,4,5-trihydroxyphenyl)-4*H*,8*H*-benzo[1,2-*b*:3,4-*b'*]dipyran-4-one (artoinin E)

Melting point: 217-219 °C

UV (CH₃OH) λ_{\max} nm (log ϵ): 224 (4.34), 258 (4.41), 266 (4.47), 271 (4.47), 302 (3.90), 352 (3.89)

IR (Neat) ν_{\max} (cm⁻¹): 3359 (O-H stretching), 1653 (C=O stretching)

¹H-NMR (CDCl₃+DMSO-*d*₆, 300 MHz) δ (ppm): 13.21 (1H, *s*, 5-OH), 8.56 (1H, *s*, OH), 8.38 (1H, *s*, OH), 7.54 (1H, *s*, OH), 6.79 (1H, *s*, H-6'), 6.62 (1H, *d*, *J* = 9.9 Hz, H-14), 6.19 (1H, *s*, H-6), 6.58 (1H, *s*, H-3'), 5.48 (1H, *d*, *J* = 9.9 Hz, H-15), 5.12 (1H, *t*, *J* = 6.6 Hz, H-10), 3.14 (2H, *d*, *J* = 6.6 Hz, H-9), 1.61 (3H, *s*, 13-CH₃), 1.47 (3H, *s*, 12-CH₃), 1.44 (6H, *s*, 17-CH₃ and 18-CH₃)

¹³C NMR (CDCl₃+DMSO-*d*₆, 75 MHz) δ (ppm): δ 182.5 (C-4), 161.5 (C-5), 161.2 (C-2), 158.8 (C-7), 152.4 (C-8a), 148.8 (C-2'), 147.9 (C-4'), 137.6 (C-5'), 132.0 (C-11), 126.5 (C-15), 121.5 (C-10), 120.8 (C-3), 116.2 (C-6'), 115.2 (C-14), 110.7 (C-1'), 105.0 (C-4a), 104.0 (C-3'), 100.8 (C-8), 99.2 (C-6), 17.5 (C-12), 77.7 (C-16), 28.0 (C-17 and C-18), 25.7 (C-13), 24.2 (C-9)

2.3.1.1 Purification of fraction B

Fraction B (16.7920 g) was chromatographed on silica gel 100 (hexane-dichloromethane, 3:1) to give fractions B1-B14.

Table 3 Physical characteristic and weight of fractions obtained from CC of fraction B

Fraction	Weight (g)	Appearance
B1	0.1571	white solid mixed with cream gel
B2	0.0209	colorless gel
B3	0.4449	yellow gel
B4	1.1679	yellow gel
B5	2.9514	yellow gel
B6	8.6561	yellow gel
B7	0.1761	orange viscous liquid
B8	0.4108	yellow-brown viscous liquid
B9	0.3249	brown solid mixed with yellow viscous liquid
B10	0.2635	brown viscous liquid
B11	0.4416	brown viscous liquid
B12	0.2287	dark red viscous liquid
B13	0.2754	dark red viscous liquid
B14	0.2177	dark red viscous liquid

Isolation of AE2

Fraction B8 (0.4108 g) was further purified by column chromatography over silica gel and eluted with hexane-acetone (20:1) solvent system. The subfractions containing similar components were combined to give fractions B8.1-B8.6. **AE2** (55.8 mg) was obtained as a yellow gum from fraction B8.4.

AE2

8-Hydroxy-3-methylisochroman-1-one (mullein)

$[\alpha]_D^{31} -72^\circ$ (*c* 0.07, CHCl₃)

IR (Neat) ν_{\max} (cm⁻¹): 3438 (O-H stretching), 1680 (C=O stretching)

¹H-NMR (CDCl₃, 300 MHz) δ (ppm): 10.95 (1H, *s*, 8-OH), 7.32 (1H, *t*, *J* = 7.9 Hz, H-6), 6.80 (1H, *d*, *J* = 7.9 Hz, H-7), 6.61 (1H, *d*, *J* = 7.9 Hz, H-5), 4.65 (1H, *sext*, *J* = 6.6 Hz, H-3), 2.85 (2H, *d*, *J* = 6.6 Hz, H-4), 1.47 (3H, *d*, *J* = 6.6 Hz, 9-CH₃)

¹³C NMR (CDCl₃, 75 MHz) δ (ppm): δ 169.9 (C-1), 162.2 (C-8), 139.4 (C-4a), 136.1 (C-6), 117.9 (C-5), 116.2 (C-7), 108.3 (C-8a), 76.1 (C-3), 34.6 (C-4), 20.7 (C-9)

Isolation of AE3

Fraction B12 (0.2287 g) was purified by a silica gel column with hexane-acetone (7:1) as an eluent to provide six fractions (B12.1-B12.6). A yellow gum of **AE3** (6.6 mg) was obtained from the fraction B12.6.

AE3

5-Hydroxy-2-(4-hydroxy-2,5-dimethoxyphenyl)-7-methoxy-3-(3-methylbut-2-enyl)-4*H*-chromen-4-one

IR (Neat) ν_{\max} (cm⁻¹): 3445 (O-H stretching), 1653 (C=O stretching)

^1H NMR (CDCl_3 , 300 MHz) δ (ppm): 13.00 (1H, *s*, 5-OH), 6.83 (1H, *s*, H-6'), 6.67 (1H, *s*, H-3'), 6.35 (2H, *s*, H-6 and H-8), 5.93 (1H, *s*, 4'-OH), 5.11 (1H, *t*, $J = 6.6$ Hz, H-10), 3.86 (3H, *s*, 5'-OCH₃), 3.83 (3H, *s*, 7-OCH₃), 3.75 (3H, *s*, 2'-OCH₃), 3.04 (2H, *d*, $J = 6.6$ Hz, H-9), 1.62 (3H, *s*, 13-CH₃), 1.42 (3H, *s*, 12-CH₃)

^{13}C NMR (CDCl_3 , 75 MHz) δ (ppm): 182.4 (C-4), 165.3 (C-7), 162.2 (C-5), 160.7 (C-2), 158.2 (C-8a), 148.5 (C-4'), 140.3 (C-5'), 132.0 (C-11), 121.7 (C-10), 121.4 (C-3), 112.9 (C-6'), 112.7 (C-1'), 105.5 (C-4a), 99.7 (C-3'), 97.8 (C-6), 92.0 (C-8), 56.7 (7-OCH₃), 56.2 (5'-OCH₃), 55.7 (2'-OCH₃), 25.7 (C-13), 24.2 (C-9), 17.6 (C-12)

EI-MS m/z (% relative intensity): 412 [M^+] (13), 411 (47), 396 (4), 381 (35), 368 (100), 356 (8), 338 (19), 324 (36), 296 (10), 282 (3), 256 (2), 242 (2), 190 (3), 180 (5), 166 (17), 163 (4), 149 (3), 128 (2), 69 (6)

HREI-MS m/z : 412.1542 for $\text{C}_{23}\text{H}_{24}\text{O}_7$ (calcd. 412.1522)

Isolation of AE4 and AE5

Fraction B13 (0.2754 g) was further purified by crystallization from hexane-dichloromethane (1:1) to give a yellow solid of **AE4** (11.9 mg) upon standing at room temperature. The filtrate of B13 was rechromatographed using hexane-acetone (7:1) as eluent to give eight fractions (B13.1-B13.8). The fraction B13.6 gave an orange solid of **AE5** (3.8 mg).

AE4

6,7-Dihydro-5,9,14-trihydroxy-11-methoxy-3,3-dimethyl-6-(1-methylethyl)-3*H*,8*H*-[1]benzopyrano[7,6-*c*]xanthen-8-one (artanol E)

Melting point: 205-207 °C

UV (EtOH) λ_{max} nm (log ϵ): 214 (2.81), 271 (2.79), 304 (2.41), 378 (2.58)

IR (Neat) ν_{max} (cm^{-1}): 3406 (O-H stretching), 1653 (C=O stretching)

^1H NMR (CDCl_3 , 300 MHz) δ (ppm): 12.99 (1H, *s*, 5-OH), 7.76 (1H, *s*, 2'-OH), 6.74 (1H, *d*, $J = 10.0$ Hz, H-14), 6.37 (2H, *s*, H-6 and H-8), 5.64 (1H, *d*,

$J = 10.0$ Hz, H-15), 5.27 (1H, *s*, 5'-OH), 4.71 (1H, *s*, H $_{\beta}$ -12), 4.35 (1H, *s*, H $_{\alpha}$ -12), 3.96 (1H, *d*, $J = 6.9$ Hz, H-10), 3.86 (3H, *s*, 7-OCH $_3$), 3.40 (1H, *dd*, $J = 16.2, 1.5$ Hz, H $_{\beta}$ -9), 2.54 (1H, *dd*, $J = 16.2, 6.9$ Hz, H $_{\alpha}$ -9), 1.85 (3H, *s*, 13-CH $_3$), 1.52 (3H, *s*, 17-CH $_3$), 1.49 (3H, *s*, 18-CH $_3$)

^{13}C NMR (CDCl $_3$, 75 MHz) δ (ppm): 180.1 (C-4), 165.1 (C-7), 162.4 (C-5), 159.9 (C-2), 155.7 (C-8a), 145.0 (C-2'), 144.3 (C-11), 143.9 (C-4'), 135.6 (C-5'), 128.6 (C-15), 126.7 (C-6'), 116.3 (C-14), 111.9 (C-12), 111.7 (C-3), 108.8 (C-3'), 105.1 (C-1'), 105.0 (C-4a), 98.2 (C-6), 92.1 (C-8), 78.5 (C-16), 55.8 (7-OCH $_3$), 36.6 (C-10), 28.3 (C-17), 28.2 (C-18), 21.6 (C-9, C-13)

AE5

12-Acetyl-6-hydroxy-3,3,9,9-tetramethyl-3*H*,7*H*,furo[3,4-*b*]pyrano[3,2-*h*]xanthen
7,11(9*H*)-dione (artanol B)

Melting point: 190-193 °C

UV (EtOH) λ_{max} nm (log ϵ): 207 (2.49), 241 (2.41), 275 (2.25), 285 (2.24), 314 (1.92), 338 (1.89), 411 (1.54)

IR (Neat) ν_{max} (cm $^{-1}$): 3455 (O-H stretching), 1731, 1653 (C=O stretching)

^1H NMR (CDCl $_3$, 300 MHz) δ (ppm): 12.50 (1H, *s*, 1-OH), 8.31 (1H, *s*, H-8), 6.60 (1H, *d*, $J = 10.1$ Hz, H-10), 6.32 (1H, *s*, H-2), 5.63 (1H, *d*, $J = 10.1$ Hz, H-11), 2.82 (3H, *s*, 5-COCH $_3$), 1.76 (6H, *s*, 17-CH $_3$ and 18-CH $_3$), 1.50 (6H, *s*, 13-CH $_3$ and 14-CH $_3$)

^{13}C NMR (CDCl $_3$, 75 MHz) δ (ppm): 198.5 (5-COCH $_3$), 179.1 (C-9), 169.4 (C-15), 163.2 (C-1), 162.3 (C-3), 152.5 (C-4a and C-4b), 148.6 (C-7), 131.0 (C-5), 128.2 (C-11), 125.2 (C-6 and 8a), 119.2 (C-8), 114.2 (C-10), 103.5 (C-9a), 101.4 (C-4), 100.2 (C-2), 86.6 (C-16), 79.1 (C-12), 32.3 (5-COCH $_3$), 28.5 (C-13 and C-14), 27.6 (C-17 and C-18)

2.3.1.2 Purification of fraction C

Fraction C (29.8941 g) was chromatographed on silica gel 100 (hexane-acetone, 20:1 to 2.5:1) to give fractions C1-C12.

Table 4 Physical characteristic and weight of fractions obtained from CC of fraction C

Fraction	Weight (g)	Appearance
C1	3.6894	yellow gel
C2	3.0701	green gel
C3	3.6262	yellow viscous liquid
C4	3.2734	green viscous liquid
C5	0.6268	red-brown viscous liquid
C6	0.3340	red-brown viscous liquid
C7	1.3897	red-brown viscous liquid
C8	2.3683	red-brown viscous liquid
C9	2.4563	red-brown viscous liquid
C10	1.6307	red-brown viscous liquid
C11	1.6454	dark brown viscous liquid
C12	0.3377	dark-brown viscous solid

Isolation of AE6

Fraction C7 (1.3897 g) was chromatographed on column chromatography and elution was conducted with hexane-acetone (7:1) to afford 10 portions (C7.1-C7.10). Fraction C7.2 was rechromatographed on column chromatography and eluted with hexane-acetone (7:1) to give a red-brown gum **AE6** (9.1 mg).

AE6

6,7-Dihydro-5,9,11,14-tetrahydroxy-3,3-dimethyl-6-(1-methylethenyl)-(-)-3*H*,8*H*-pyrano[3',2':4,5]benzo[1,2-*c*]xanthen-8-one (artelastoxanthone or 7-demethylartanol E)

UV (CH₃OH) λ_{\max} nm (log ϵ): 263 (4.39), 269 (4.36), 307 (3.71), 379 (4.11)

IR (Neat) ν_{\max} (cm⁻¹): 3402 (O-H stretching), 1655 (C=O stretching)

¹H NMR (CDCl₃+Acetone-*d*₆, 300 MHz) δ (ppm): 12.98 (1H, *s*, 5-OH), 7.78 (1H, *s*, 2'-OH), 6.74 (1H, *d*, *J* = 10.0 Hz, H-14), 6.40 (1H, *d*, *J* = 1.8 Hz, H-8), 6.35 (1H, *d*, *J* = 1.8 Hz, H-6), 5.64 (1H, *d*, *J* = 10.0 Hz, H-15), 5.46 (1H, *s*, 5'-OH), 4.71 (1H, *s*, H _{β} -12), 4.34 (1H, *s*, H _{α} -12), 3.96 (1H, *d*, *J* = 6.9 Hz, H-10), 3.39 (1H, *dd*, *J* = 16.2, 1.5 Hz, H _{β} -9), 2.53 (1H, *dd*, *J* = 16.2, 6.9 Hz, H _{α} -9), 1.81 (3H, *s*, 13-CH₃), 1.52 (3H, *s*, 17-CH₃), 1.49 (3H, *s*, 18-CH₃)

¹³C NMR (CDCl₃, 75 MHz) δ (ppm): 180.0 (C-4), 163.0 (C-7), 162.4 (C-5), 159.7 (C-2), 155.9 (C-8a), 144.9 (C-2'), 144.3 (C-11), 143.8 (C-4'), 135.6 (C-5'), 128.5 (C-15), 126.7 (C-6'), 116.3 (C-14), 111.7 (C-12), 111.5 (C-3), 108.8 (C-3'), 105.2 (C-1'), 104.5 (C-4a), 99.8 (C-6), 93.6 (C-8), 78.3 (C-16), 36.5 (C-10), 28.2 (C-17), 28.1 (C-18), 21.6 (C-13), 21.5 (C-9)

Isolation of AE7

Fraction C7.3 was further purified by crystallization from hexane-dichloromethane-acetone (1:1:1) upon standing at room temperature to give yellow solid of **AE7** (4.9 mg).

AE7

New furanodihydrobenzoxanthone derivative

$[\alpha]_{\text{D}}^{26.7} -18.5^\circ$ (*c* 0.03, MeOH)

Melting point: 287-289 °C

UV (EtOH) λ_{max} nm (log ϵ): 214 (2.81), 253 (2.70), 273 (2.82), 288 (2.69), 308 (2.45), 375 (2.62)

IR (Neat) ν_{max} (cm^{-1}): 3442 (O-H stretching), 1630 (C=O stretching)

^1H NMR (CDCl_3 +Acetone- d_6 , 300 MHz) δ (ppm): 13.00 (1H, *s*, 5-OH), 9.42 (1H, *s*, 7-OH), 7.17 (1H, *s*, 2'-OH), 6.69 (1H, *d*, $J = 10.2$ Hz, H-14), 6.47 (1H, *d*, $J = 2.1$ Hz, H-8), 6.31 (1H, *d*, $J = 2.1$ Hz, H-6), 5.57 (1H, *d*, $J = 10.2$ Hz, H-15), 3.39 (1H, *dd*, $J = 15.3, 7.2$ Hz, H-10), 3.21 (1H, *dd*, $J = 15.3, 7.2$ Hz, H $_{\beta}$ -9), 2.40 (1H, *t*, $J = 15.3$ Hz, H $_{\alpha}$ -9), 1.68 (1H, *s*, 13-CH $_3$), 1.50 (3H, *s*, 17-CH $_3$), 1.47 (3H, *s*, 18-CH $_3$), 1.35 (3H, *s*, 12-CH $_3$)

^{13}C NMR (CDCl_3 +DMSO- d_6 , 75 MHz) δ (ppm): 185.4 (C-4), 168.7 (C-7), 167.5 (C-5), 165.1 (C-2), 161.5 (C-8a), 149.6 (C-2'), 147.2 (C-4'), 142.6 (C-5'), 136.4 (C-6'), 133.2 (C-15), 121.9 (C-14), 117.4 (C-3), 116.1 (C-3'), 109.4 (C-4a), 108.5 (C-1'), 104.9 (C-6), 99.0 (C-8), 98.7 (C-11), 82.7 (C-16), 51.5 (C-10), 33.2 (C-13), 33.1 (C-17), 33.0 (C-18), 27.6 (C-12), 24.9 (C-9)

EI-MS m/z (% relative intensity): 434 [M^+] (15), 433 (50), 418 (100), 416 (17), 391 (7), 390 (23), 375 (22), 372 (8), 348 (16), 344 (5), 330 (4), 292 (3), 254 (4), 209 (3), 188 (7), 164 (3), 152 (4), 138 (1), 115 (2), 69 (2)

HREI-MS m/z : 434.1377 for C $_{25}$ H $_{22}$ O $_7$ (calcd. 434.1366)

Isolation of AE8

Fraction C7.4 containing one major component was further purified by crystallization from hexane-dichloromethane-acetone (1:1:1). A yellow solid of **AE8** (18.8 mg) which formed was filtered.

AE8

5a,6-Dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-9-(3-methyl-2-buten-1-yl)-5*H*,7*H*,11*H*-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one (artonin F)

Melting point: 249-251 °C

UV (CH₃OH) λ_{\max} nm (log ϵ): 236 (4.46), 256 (4.42), 265 (4.41), 278 (4.50), 335 (4.09), 390 (4.27)

IR (Neat) ν_{\max} (cm⁻¹): 3371 (O-H stretching), 1629 (C=O stretching)

¹H NMR (CDCl₃+DMSO-*d*₆, 300 MHz) δ (ppm): 13.42 (1H, *s*, 5-OH), 9.23 (1H, *s*, 4'-OH), 7.78 (1H, *s*, 2'-OH), 6.76 (1H, *d*, *J* = 9.9 Hz, H-19), 6.38 (1H, *s*, H-3'), 5.57 (1H, *d*, *J* = 9.9 Hz, H-20), 5.24 (1H, *t*, *J* = 7.2 Hz, H-15), 3.41 (1H, *dd*, *J* = 15.0, 7.2 Hz, H-10), 3.33 (1H, *d*, *J* = 7.2 Hz, H-14), 3.23 (1H, *dd*, *J* = 15.0, 7.2 Hz, H _{β} -9), 2.40 (1H, *t*, *J* = 15.0 Hz, H _{α} -9), 1.81 (3H, *s*, 18-CH₃), 1.68 (3H, *s*, 17-CH₃), 1.66 (3H, *s*, 13-CH₃), 1.46 (6H, *s*, 22-CH₃ and 23-CH₃), 1.35 (3H, *s*, 12-CH₃)

¹³C NMR (CDCl₃, 75 MHz) δ (ppm): 180.7 (C-4), 160.3 (C-2), 158.7 (C-5), 156.4 (C-7), 150.1 (C-2'), 149.2 (C-8a), 146.4 (C-4'), 137.2 (C-5'), 131.8 (C-6'), 131.3 (C-16), 127.1 (C-20), 122.1 (C-15), 115.3 (C-19), 112.5 (C-6), 111.5 (C-3), 104.6 (C-3'), 104.3 (C-4a), 103.4 (C-1'), 100.5 (C-8), 93.5 (C-11), 77.3 (C-21), 46.6 (C-10), 28.1 (C-13), 28.0 (C-22 and C-23), 25.8 (C-18), 22.7 (C-12), 21.3 (C-14), 19.9 (C-9), 17.9 (C-17)

2.3.1.3 Purification of fraction D

Fraction D (9.0987 g) was chromatographed on silica gel 100 (hexane-acetone, 20:1 to 2:1) to give fractions D1-D15.

Table 5 Physical characteristic and weight of fractions obtained from CC of fraction D

Fraction	Weight (g)	Appearance
D1	0.2134	yellow gel
D2	0.1456	orange gel
D3	0.0961	orange gel
D4	0.2567	orange gel
D5	0.3564	yellow gel
D6	0.2689	brown-yellow viscous liquid
D7	0.0670	brown viscous liquid
D8	0.0921	brown viscous liquid
D9	0.1587	black-brown viscous liquid
D10	0.1109	black-brown viscous liquid
D11	0.4822	black-brown viscous liquid
D12	0.8762	black-brown viscous liquid
D13	2.2525	black-brown viscous liquid
D14	1.5685	black-brown solid
D15	0.6456	black solid

Isolation of AE9 and AE10

Fraction D11 (0.4822 g) was further separated by column chromatography over silica gel and eluted with hexane-acetone (7:1) solvent system. The fractions containing similar components were combined into eleven fractions (D11.1-D11.11). The fraction D11.8 was rechromatographed using hexane-acetone (5:1) as an eluent to afford red-brown gum **AE9** (1.7 mg). The fraction D11.9 which contained one major component was further purified by crystallization from hexane-acetone (7:1). A yellow solid of **AE10** (22.5 mg) which formed was filtered.

AE9

(3,4,5-Trimethoxyphenyl)methanol

^1H NMR (CDCl_3 , 500 MHz) δ (ppm): 6.61 (2H, *s*, H-2 and H-6), 4.65 (2H, *s*, H-7), 3.88 (6H, *s*, 3-OCH₃ and 5-OCH₃), 3.85 (3H, *s*, 4-OCH₃)

^{13}C NMR (CDCl_3 , 75 MHz) δ (ppm): 153.5 (C-3 and C-5), 137.5 (C-4), 135.5 (C-1), 103.9 (C-2 and C-6), 65.6 (C-7), 60.9 (4-OCH₃), 56.1 (3-OCH₃ and 5-OCH₃)

AE10

5a,6-Dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-5*H*,7*H*,11*H*-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one (cycloartobiloxanthone)

Melting point: 284-285 °C

UV (EtOH) λ_{max} nm (log ϵ): 228 (2.92), 257 (2.88), 273 (2.94), 312 (2.56), 330 (2.52), 391 (2.72)

IR (Neat) ν_{max} (cm^{-1}): 3405 (O-H stretching), 1642 (C=O stretching)

^1H NMR (CDCl_3 +DMSO-*d*₆, 300 MHz) δ (ppm): 13.22 (1H, *s*, 5-OH), 9.26 (1H, *s*, 4'-OH), 8.00 (1H, *s*, 2'-OH), 6.78 (1H, *d*, $J = 10.0$ Hz, H-14), 6.38 (1H, *s*, H-3'), 6.23 (1H, *s*, H-6), 5.56 (1H, *d*, $J = 10.0$ Hz, H-15), 3.38 (1H, *dd*, $J = 15.0, 7.2$ Hz, H-10), 3.21 (1H, *dd*, $J = 15.0, 7.2$ Hz, H $_{\beta}$ -9), 2.40 (1H, *t*, $J = 15.0$ Hz, H $_{\alpha}$ -9), 1.66 (3H, *s*, 13-CH₃), 1.46 (6H, *s*, 17-CH₃ and 18-CH₃), 1.34 (3H, *s*, 12-CH₃)

^{13}C NMR (CDCl_3 +DMSO-*d*₆, 75 MHz) δ (ppm): 180.6 (C-4), 161.5 (C-5), 160.7 (C-2), 158.7 (C-7), 150.9 (C-8a), 150.3 (C-2'), 146.5 (C-4'), 137.2

(C-5'), 131.9 (C-6'), 127.3 (C-15), 115.0 (C-14), 111.6 (C-3), 104.7 (C-4a and C-3'), 103.4 (C-1'), 101.0 (C-8), 99.9 (C-6), 93.5 (C-11), 77.8 (C-16), 46.6 (C-10), 28.1 (C-13, C-17 and C-18), 22.6 (C-12), 19.8 (C-9)

Isolation of AE11

Fraction D12 (0.8762 g) was further purified by column chromatography over silica gel and eluted with hexane-acetone (4:1) solvent system. The fractions containing similar components were combined into thirteen fractions. Crystallization of the eighth fraction from hexane-acetone gave a yellow solid of **AE11** (3.1 mg).

AE11

New quinonobenzoxanthone derivative

$[\alpha]_{\text{D}}^{26.3} -18.2^\circ$ (*c* 0.045, MeOH)

Melting point: 179-180 °C

UV (EtOH) λ_{max} nm (log ϵ): 217 (2.84), 229 (2.86), 245 (2.88), 267 (2.98), 303 (2.47), 354 (2.28)

IR (Neat) ν_{max} (cm^{-1}): 3433 (O-H stretching), 1720, 1647, 1584 (C=O stretching)

^1H NMR (CDCl_3 , 300 MHz) δ (ppm): 12.56 (1H, *s*, H-5), 6.77 (1H, *d*, $J = 10.0$ Hz, H-14), 6.27 (1H, *s*, H-6), 5.59 (1H, *d*, $J = 10.0$ Hz, H-15), 4.30 (1H, *s*, 4'-OH), 3.49 (1H, *s*, 11-OH), 3.50 (1H, *t*, $J = 2.1$ Hz, H-6'), 3.00 (1H, *bs*, H-19), 2.96 (1H, *dd*, $J = 16.8, 5.7$ Hz, H-9), 2.93 (1H, *bs*, H-19), 2.92 (1H, *m*, H-10), 2.67 (1H, *bs*, H-3'), 2.63 (1H, *bs*, H-3'), 2.62 (1H, *d*, $J = 2.1$ Hz, H-1'), 2.58 (1H, *dd*, $J = 16.8, 5.7$ Hz, H-9), 2.27 (1H, *s*, H-21), 1.48 (3H, *s*, 17-CH₃), 1.46 (3H, *s*, 18-CH₃), 1.20 (3H, *s*, 13-CH₃), 0.90 (3H, *s*, 12-CH₃)

^{13}C NMR (CDCl_3 , 75 MHz) δ (ppm): 207.6 (C-20), 205.5 (C-5'), 198.8 (C-2'), 180.9 (C-4), 161.6 (C-5), 159.7 (C-7), 156.9 (C-2), 151.8 (C-8a), 127.4 (C-15), 115.9

(C-3), 114.8 (C-14), 104.9 (C-4a), 101.2 (C-8), 100.3 (C-6), 78.2 (C-16), 75.6 (C-4'), 74.5 (C-11), 47.8 (C-6'), 46.9 (C-19), 46.9 (C-1'), 43.5 (C-3'), 32.0 (C-21), 32.8 (C-10), 31.9 (C-13), 28.3 (C-17), 28.2 (C-18), 25.3 (C-12), 22.6 (C-9)

Isolation of AE12 and AE13

Fraction D13 (2.2525 g) was further purified by crystallization from hexane-acetone (1:1) upon standing at room temperature to give a brown-yellow solid of **AE12** (29.5 mg) and the filtrate (2.2230 g). The filtrate was purified by column chromatography over silica gel and eluted with mixed solvent of hexane-acetone (7:1 to 3:1) to give thirteen fractions (D13.1-D13.13). Fraction D13.12 was further purified by crystallization from acetone to give a yellow solid of **AE13** (18.8 mg) upon standing at room temperature.

AE12

1,3,4,8-Tetrahydroxy-10-methoxy-5-(prop-1-en-2-yl)-5*H*-benzo[*c*]xanthen-7-(6*H*)-one

$[\alpha]_D^{26} -22^\circ$ (*c* 0.045, MeOH)

Melting point: 242-243 °C

UV (EtOH) λ_{\max} nm (log ϵ): 214 (2.77), 228 (2.67), 261 (2.77), 314 (2.36), 380 (2.66)

IR (Neat) ν_{\max} (cm⁻¹): 3180 (O-H stretching), 1653 (C=O stretching)

¹H NMR (Acetone *d*₆, 300 MHz) δ (ppm): 13.23 (1H, *s*, 5-OH), 8.27 (1H, *s*, OH), 6.65 (1H, *d*, *J* = 2.4 Hz, H-8), 6.52 (1H, *s*, H-3'), 6.30 (1H, *d*, *J* = 2.4 Hz, H-6), 4.65 (1H, *s*, H _{β} -12), 4.29 (1H, *s*, H _{α} -12), 4.00 (1H, *d*, *J* = 6.3 Hz, H-10), 3.89 (3H, *s*, 7-OCH₃), 3.41 (1H, *dd*, *J* = 15.9, 1.8 Hz, H _{β} -9), 2.46 (1H, *dd*, *J* = 15.9, 6.3 Hz, H _{α} -9), 1.78 (3H, *s*, 13-CH₃)

¹³C NMR (Acetone *d*₆, 75 MHz) δ (ppm): 180.1 (C-4), 165.1 (C-7), 161.9 (C-5), 161.0 (C-2), 156.7 (C-8a), 150.5 (C-2'), 150.3 (C-4'), 144.4 (C-11), 136.0 (C-5'),

128.3 (C-6'), 110.9 (C-12), 110.6 (C-3), 105.5 (C-1'), 104.7 (C-4a), 102.9 (C-3'), 97.7 (C-6), 92.2 (C-8), 55.4 (7-OCH₃), 37.0 (C-10), 21.4 (C-9), 21.0 (C-13)

EI-MS m/z (% relative intensity): 382 [M⁺] (18), 381 (68), 366 (20), 364 (18), 353 (5), 341 (13), 340 (56), 338 (100), 311 (6), 310 (25), 294 (7), 266 (4), 241 (5), 217 (3), 200 (4), 188 (4), 166 (10), 161 (3), 115 (3), 91 (1), 77 (1), 68 (4)

HREI-MS m/z : 382.1068 for C₂₁H₁₈O₇ (calcd. 382.1053)

AE13

New quinonobenzoxanthone derivative

$[\alpha]_D^{25.8}$ -32 ° (c 0.025, MeOH)

Melting point: 202-203 °C

UV (EtOH) λ_{\max} nm (log ϵ): 212 (2.73), 232 (2.83), 269 (2.97), 308 (2.35), 327 (2.20), 361 (2.07)

IR (Neat) ν_{\max} (cm⁻¹): 3426 (O-H stretching), 1753, 1722, 1697, 1657 (C=O stretching)

¹H NMR (CDCl₃, 300 MHz) δ (ppm): 12.42 (1H, *s*, 5-OH), 6.78 (1H, *d*, $J = 10.2$ Hz, H-14), 6.28 (1H, *s*, H-6), 5.64 (1H, *s*, 4'-OH), 5.60 (1H, *d*, $J = 10.2$ Hz, H-15), 4.98 (1H, *s*, H _{β} -12), 4.91 (1H, *d*, $J = 0.9$ Hz, H _{α} -12), 3.57 (1H, *d*, $J = 19.4$ Hz, H-3'), 3.47 (1H, *d*, $J = 19.4$ Hz, H-3'), 3.38 (1H, *d*, $J = 7.2$ Hz, H-10), 3.25 (1H, *d*, $J = 15.8$ Hz, H _{β} -19), 3.03 (1H, *d*, H _{α} -19), 2.89 (1H, *d*, $J = 17.1$ Hz, H _{β} -9), 2.49 (1H, *dd*, $J = 17.1, 7.2$ Hz, H _{α} -9), 2.43 (3H, *s*, 21-CH₃), 1.67 (3H, *s*, 13-CH₃), 1.49 (3H, *s*, 17-CH₃), 1.46 (3H, *s*, 18-CH₃)

¹³C NMR (CDCl₃, 75 MHz) δ (ppm): 213.3 (C-20), 198.7 (C-5'), 195.6 (C-2'), 181.4 (C-4), 162.8 (C-5), 161.3 (C-7), 155.2 (C-2), 153.4 (C-8a), 145.1 (C-11), 129.1 (C-15), 117.8 (C-12), 117.3 (C-3), 116.3 (C-14), 106.6 (C-4a), 103.1 (C-8), 102.0 (C-6), 83.6 (C-4'), 79.9 (C-16), 72.0 (C-6'), 61.9 (C-1'), 50.9 (C-3'), 41.7 (C-19), 41.1 (C-10), 33.3 (C-21), 29.9 (C-17), 29.7 (C-18), 24.3 (C-9), 22.5 (C-13)

EI-MS m/z (% relative intensity): 506 [M^+] (9), 505 (27), 490 (100), 462 (3), 446 (6), 433 (13), 432 (34), 418 (14), 362 (19), 336 (7), 306 (3), 292 (6), 280 (2), 203 (1), 202 (13), 176 (2), 134 (2), 68 (2)

HREI-MS m/z : 506.1572 for $C_{28}H_{26}O_9$ (calcd. 506.1577)

Isolation of AE14

Fraction E (1.7163 g) was further separated by column chromatography over silica gel and eluted with hexane-acetone (4:1) solvent system. The fractions containing similar components were combined into twenty fractions (E1-E20). Crystallization of the fraction E13 from acetone gave yellow solid of **AE14** (4.3 mg).

AE14

5a,6-Dihydro-1,3,8-trihydroxy-10-methoxy-5,5-dimethyl-5*H*,7*H*-benzofuro[3,4-*bc*]xanthen-7-one (arntonin K)

Melting point: 287-288 °C

UV (CH₃OH) λ_{\max} nm (log ϵ): 225 (4.55), 266 (4.84), 270 (4.83), 304 (4.15), 362 (4.15)

IR (Neat) ν_{\max} (cm⁻¹): 3369 (O-H stretching), 1629 (C=O stretching)

¹H NMR (CDCl₃+DMSO-*d*₆, 300 MHz) δ (ppm): 13.17 (1H, *s*, 5-OH), 9.71 (1H, *s*, 4'-OH), 8.38 (1H, *s*, 2'-OH), 6.56 (1H, *d*, $J = 2.1$ Hz, H-8), 6.39 (1H, *s*, H-3'), 6.32 (1H, *d*, $J = 2.1$ Hz, H-6), 3.88 (3H, *s*, 7-OCH₃), 3.38 (1H, *dd*, $J = 15.0, 6.9$ Hz, H-10), 3.22 (1H, *dd*, $J = 15.0, 6.9$ Hz, H _{β} -9), 2.40 (1H, *t*, $J = 15.0$ Hz, H _{α} -9), 1.67 (3H, *s*, 13-CH₃), 1.35 (3H, *s*, 12-CH₃)

¹³C NMR (CDCl₃+DMSO-*d*₆, 75 MHz) δ (ppm): 185.2 (C-4), 169.6 (C-7), 166.5 (C-5), 165.9 (C-2), 161.3 (C-8a), 155.3 (C-2'), 151.5 (C-4'), 141.9 (C-5'), 136.6 (C-6'), 116.4 (C-3), 110.5 (C-4a), 109.6 (C-3'), 108.2 (C-1'), 102.8 (C-6), 97.8 (C-11), 97.1 (C-8), 60.5 (7-OCH₃), 51.4 (C-10), 32.9 (C-13), 27.4 (C-12), 24.6 (C-9)

Isolation of AE15

Fraction F (2.8492 g) was further separated by column chromatography over SephadexTM LH-20 and eluted with methanol. The fractions containing similar components were combined into sixteen fractions (F1-F16). The fraction F11 was further purified by column chromatography over silica gel and eluted with hexane-acetone (3:1) solvent system to give **AE15** as a pale yellow solid (7.0 mg).

AE15

5-Hydroxy-8,8-dimethyl-2-(2,4-dihydroxyphenyl)-4*H*,8*H*-benzo[1,2-*b'*]dipyran-4-one

Melting point: 189-191 °C

UV (EtOH) λ_{\max} nm (log ϵ): 212 (2.66), 230 (2.58), 254 (2.44), 273 (2.52), 287 (2.51), 307 (2.38), 357 (2.54)

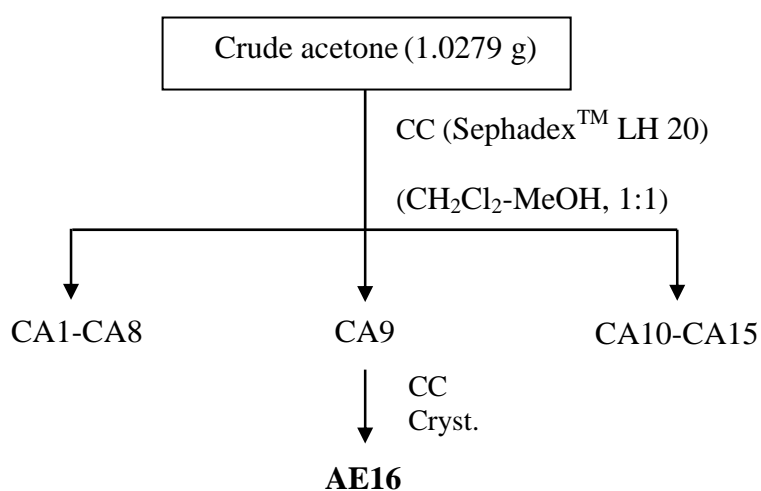
IR (Neat) ν_{\max} (cm⁻¹): 3379 (O-H stretching), 1651 (C=O stretching)

¹H NMR (Acetone-*d*₆, 300 MHz) δ (ppm): 13.53 (1H, *s*, 5-OH), 7.83 (1H, *d*, *J* = 8.7 Hz, H-6'), 7.15 (1H, *s*, H-3), 6.67 (1H, *d*, *J* = 10.2 Hz, H-9), 6.58 (1H, *d*, *J* = 2.1 Hz, H-3'), 6.54 (1H, *dd*, *J* = 8.7, 2.1 Hz, H-5'), 6.44 (1H, *s*, H-6), 5.74 (1H, *d*, *J* = 10.2 Hz, H-10), 1.47 (6H, *s*, 12-CH₃ and 13-CH₃)

¹³C NMR (Acetone-*d*₆, 75 MHz) δ (ppm): 182.8 (C-4), 162.3 (C-2), 161.9 (C-4'), 159.1 (C-7), 158.8 (C-2'), 157.1 (C-5), 156.3 (C-8a), 129.9 (C-6'), 128.1 (C-10), 115.0 (C-9), 108.2 (C-5'), 107.5 (C-3), 104.9 (C-4a, C-8, C-1'), 103.5 (C-3'), 94.6 (C-6), 78.3 (C-11), 27.6 (C-12 and C-13).

2.3.2 Purification of acetone extract

Acetone extract (1.0279 g) was separated by column chromatography over SephadexTM LH 20 and eluted with CH₂Cl₂-MeOH (1:1) solvent system. On the basis of their TLC characteristic, fractions which contained the same major component were combined to give fractions (CA1-CA15). Fraction CA9 was rechromatographed on column chromatography and eluted with the mixed solvent of hexane-acetone (2.5:1) to give five fractions. Crystallization of the third fraction in hexane-acetone (2.5:1) gave a pale creamy solid of **AE16** (1.8 mg).



Scheme 3 Isolation of compounds **AE16** from acetone extract of the bark of *A. elasticus*

AE16

2-(2,4-Dihydroxyphenyl)-5,7-dihydroxy-4*H*-chromen-4-one

¹H NMR (Acetone-*d*₆, 500 MHz) δ (ppm): 13.00 (1H, *s*, 5-OH), 9.95 (1H, *s*, OH), 9.90 (1H, *s*, OH), 9.42 (1H, *s*, OH), 7.70 (1H, *d*, *J* = 8.5 Hz, H-6'), 6.96 (1H, *s*, H-3), 6.48 (1H, *d*, *J* = 2.5 Hz, H-3'), 6.42 (1H, *dd*, *J* = 8.5, 2.5 Hz, H-5'), 6.36 (1H, *d*, *J* = 2.5 Hz, H-6), 6.10 (1H, *d*, *J* = 2.5 Hz, H-8)

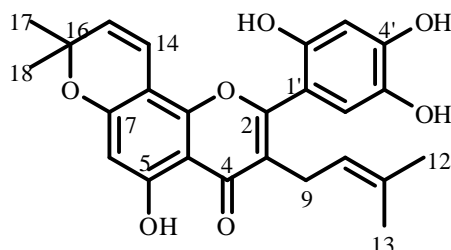
¹³C NMR (Acetone-*d*₆, 125 MHz) δ (ppm): 182.6 (C-4), 164.0 (C-7), 162.4 (C-8a), 162.0 (C-5), 161.8 (C-2'), 158.5 (C-2), 158.0 (C-4'), 129.9 (C-6'), 108.2 (C-3), 107.7 (C-5'), 107.6 (C-1'), 103.5 (C-4a), 103.4 (C-3'), 98.5 (C-8), 93.6 (C-6)

CHAPTER 3

RESULTS AND DISCUSSION

3.1 Structural Determination

The bark of *Artocarpus elasticus* was extracted with dichloromethane and acetone, successively. Separation of the dichloromethane extract by column chromatography produced 5-hydroxy-8,8-dimethyl-3-(3-methyl-2-butenyl)-2-(2,4,5-trihydroxyphenyl)-4*H*,8*H*-benzo[1,2-*b*:3,4-*b'*]dipyran-4-one (**AE1**), 8-hydroxy-3-methylisochroman-1-one (**AE2**), 5-hydroxy-2-(4-hydroxy-2,5-dimethoxyphenyl)-7-methoxy-3-(3-methylbut-2-enyl)-4*H*-chromen-4-one (**AE3**), 6,7-dihydro-5,9,14-trihydroxy-11-methoxy-3,3-dimethyl-6-(1-methylethyl)-3*H*,8*H*-[1]benzopyrano[7,6-*c*]xanthen-8-one (**AE4**), 12-acetyl-6-hydroxy-3,3,9,9-tetramethyl-3*H*,7*H*,furo[3,4-*b*]pyrano[3,2-*h*]xanthen-7,11(9*H*)-dione (**AE5**), 6,7-dihydro-5,9,11,14-tetrahydroxy-3,3-dimethyl-6-(1-methylethenyl)-(-)-3*H*,8*H*-pyrano[3',2':4,5]benzo[1,2-*c*]xanthen-8-one (**AE6**), new furanodihydrobenzoxanthone derivative (**AE7**), 5a,6-dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-9-(3-methyl-2-buten-1-yl)-5*H*,7*H*,11*H*-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one (**AE8**), (3,4,5-trimethoxyphenyl)methanol (**AE9**), 5a, 6-dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-5*H*,7*H*,11*H*-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one (**AE10**), new quinonobenzoxanthone derivative (**AE11**), 1,3,4,8-tetrahydroxy-10-methoxy-5-(prop-1-en-2-yl)-5*H*-benzo[*c*]xanthen-7-(6*H*)-one (**AE12**), new quinonobenzoxanthone derivative (**AE13**), 5a,6-dihydro-1,3,8-trihydroxy-10-methoxy-5,5-dimethyl-5*H*,7*H*-benzofuro[3,4-*bc*]xanthen-7-one (**AE14**) and 5-hydroxy-8,8-di-methyl-2-(2,4-dihydroxyphenyl)-4*H*,8*H*-benzo[1,2-*b'*]dipyran-4-one (**AE15**), whereas purification of acetone extract gave one compound: 2-(2,4-dihydroxyphenyl)-5,7-dihydroxy-4*H*-chromen-4-one (**AE16**). Their structures were elucidated by 1D and 2D NMR spectroscopic data.

AE1**5-Hydroxy-8,8-dimethyl-3-(3-methyl-2-butenyl)-2-(2,4,5-trihydroxyphenyl)-4*H*,8*H*-benzo[1,2-b:3,4-*b'*]dipyrans-4-one**

AE1 is a brown-yellow solid, m.p. 217-219 °C. The UV spectrum exhibited the absorption bands at 224, 258, 266, 271, 302 and 352 nm. The IR spectrum showed the absorption bands of a hydroxyl group at 3359 cm^{-1} and carbonyl group at 1653 cm^{-1} . The ^1H NMR spectrum (**Table 6**) showed signals of a hydrogen-bonded hydroxyl group (δ 13.21, *s*, 5-OH), three non-bonded hydroxyl groups (δ 8.56, 8.38, and 7.54), and three isolated aromatic protons (δ 6.19, *s*, H-6; δ 6.58, *s*, H-3' and δ 6.79, *s*, H-6'). The doublet signals of vinylic protons at δ 5.48 (H-15) and δ 6.62 (H-14) and singlet signal of methyl groups at δ 1.44 (CH_3 -17 and CH_3 -18) were assigned for those of a 2, 2-dimethylchromene ring. The correlations of H-14 to C-7, C-8, C-8a; H-15 to C-8 and H-6 to C-7, C-8 correctly determined that the chromene ring was at C-7 and C-8 position. The presence of a prenyl group was observed from the characteristic signals at δ 3.14 (*d*, H-9); δ 5.12 (*t*, H-10); δ 1.47 (*s*, CH_3 -12) and δ 1.61 (*s*, CH_3 -13). This side chain was placed at C-3 according to the HMBC correlation of H-9 to C=O (δ 182.5) and C-2. These assignments were in agreement with a previously isolated compound, **artonin E** (Hano, *et al.*, 1990).

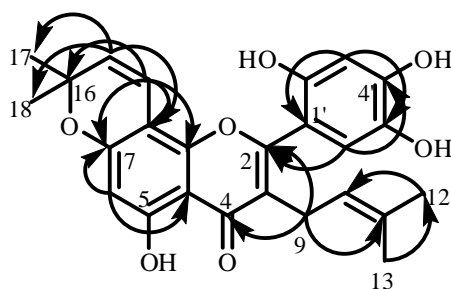
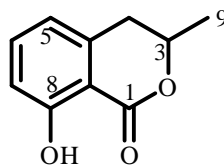
Major HMBC of **AE1**

Table 6 ^{13}C , ^1H and HMBC spectral data of **AE1**

Position	δ_{C} (C-Type)	δ_{H} (mult, J_{Hz})	HMBC
2	161.2 (C)	-	-
3	120.8 (C)	-	-
4	182.5 (C=O)	-	-
4a	105.0 (C)	-	-
5	161.5 (C)	-	-
6	99.2 (CH)	6.19 (1H, <i>s</i>)	C-4a, C-5, C-7, C-8
7	158.8 (C)	-	-
8	100.8 (C)	-	-
8a	152.4 (C)	-	-
9	24.2 (CH ₂)	3.14 (2H, <i>d</i> , 6.6)	C-2, C-3, C-4, C-10, C-11
10	121.5 (CH)	5.12 (1H, <i>t</i> , 6.6)	-
11	132.0 (C)	-	-
12	17.5 (CH ₃)	1.47 (3H, <i>s</i>)	C-10, C-11, C-13
13	25.7 (CH ₃)	1.61 (3H, <i>s</i>)	C-10, C-11, C-12
14	115.2 (CH)	6.62 (1H, <i>d</i> , 9.9)	C-7, C-8, C-8a, C-16
15	126.5 (CH)	5.48 (1H, <i>d</i> , 9.9)	C-8, C-16, C-17, C-18
16	77.7 (C)	-	-
17	28.0 (CH ₃)	1.44 (3H, <i>s</i>)	C-15, C-16, C-18
18	28.0 (CH ₃)	1.44 (3H, <i>s</i>)	C-15, C-16, C-17
1'	110.7 (C)	-	-
2'	148.8 (C)	-	-
3'	104.0 (CH)	6.58 (1H, <i>s</i>)	C-1', C-2', C-5'
4'	147.9 (C)	-	-
5'	137.6 (C)	-	-
6'	116.2 (CH)	6.79 (1H, <i>s</i>)	C-2, C-4', C-5'
5-OH	-	13.21 (1H, <i>s</i>)	C-4a, C-5, C-6
*OH	-	8.56 (1H, <i>s</i>)	-
*OH	-	8.38 (1H, <i>s</i>)	-
*OH	-	7.54 (1H, <i>s</i>)	-

* the position not identified

AE2: 8-Hydroxy-3-methylisochroman-1-one

AE2 is a yellow gum, $[\alpha]_D^{31} -72^\circ$ (c 0.07, CHCl_3). The IR spectrum indicated the presence of O-H stretching at 3438 cm^{-1} and C=O stretching at 1680 cm^{-1} . The ^1H NMR spectral data (**Table 7**) showed signals of a chelated hydroxyl proton at δ 10.95 (8-OH, *s*), and three coupled aromatic protons H-7, H-6 and H-5 at δ 6.80 (*d*), δ 7.32 (*t*), δ 6.61 (*d*). The spectrum further showed a doublet signal of methylene protons at δ 2.85 (H-4), a sextet signal of a methine proton at δ 4.65 (H-3) and a doublet signal of methyl protons at δ 1.47 (H-9). The ^1H - ^1H COSY correlation of H-3 to H-4 and H-3 to CH_3 -9 confirmed the connection of partial structure ($\text{CH}_2\text{-CH-CH}_3$). The HMBC correlations of H-3 to C-4a and H-4 to C-8a, C-5 suggested the point of attachment of C-3, C-4 and C-4a of aromatic ring. Moreover, the HMBC spectral data also showed the correlation of H-3 to 1-C=O. The ^{13}C NMR spectrum showed signals of carbonyl carbon at δ 169.9 (1-C=O), methyl carbon at δ 20.7, methylene carbons at δ 34.6, four methine carbons at δ 136.1, 117.9, 116.2 and 76.1, and three quaternary carbons at δ 162.2, 139.4 and 108.3. The chemical shift value of carbonyl carbon (δ 169.9) indicated that it was carbonyl of ester group. The HMBC experiment also confirmed the assignments structure of **AE2** as 8-hydroxy-3-methylisochroman-1-one, its optical rotation was corresponded to the R-(-)-mellein (Dimitriadis, *et al.*, 1997).

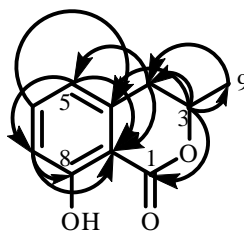
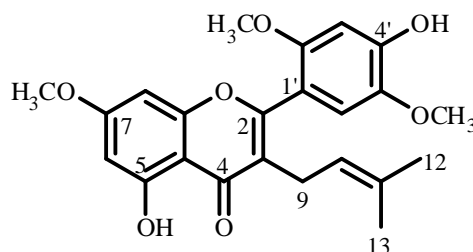
Major HMBC of **AE2**

Table 7 ^{13}C , ^1H and HMBC spectral data of **AE2**

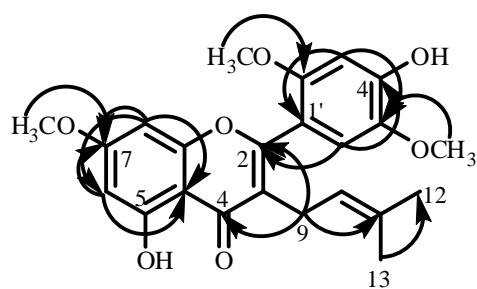
Position	δ_{C} (C-Type)	δ_{H} (<i>mult</i> , J_{Hz})	HMBC
1	169.9 (C=O)	-	-
3	76.1 (CH)	4.65 (1H, <i>sext</i> , 6.6)	C-1, C-4, C-4a, C-9
4	34.6 (CH ₂)	2.85 (2H, <i>d</i> , 6.6)	C-3, C-5, C-8a, C-9
4a	139.4 (C)	-	-
5	117.9 (CH)	6.61 (1H, <i>d</i> , 7.9)	C-4a, C-6, C-7, C-8a
6	136.1 (CH)	7.32 (1H, <i>t</i> , 7.9)	C-4a
7	116.2 (CH)	6.80 (1H, <i>d</i> , 7.9)	C-5, C-8, C-8a
8	162.2 (C)	-	-
8a	108.3 (C)	-	-
9	20.7 (CH ₃)	1.47 (3H, <i>d</i> , 6.6)	C-3, C-4
8-OH	-	10.9 (OH, <i>s</i>)	C-6, C-7, C-8, C-8a

Table 8 ^1H - ^1H COSY spectral data of **AE2**

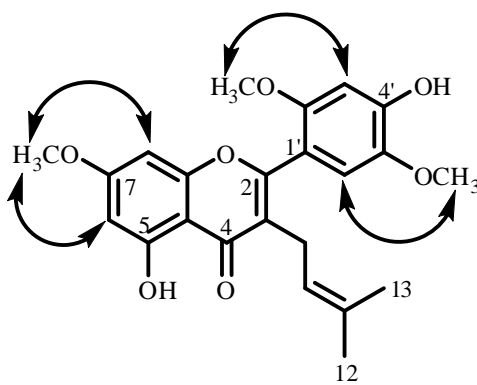
Proton (δ_{ppm})	\longleftrightarrow	Correlated proton (δ_{ppm})
H-3 (4.65)		H-4 (2.85), H-9 (1.47)
H-4 (2.85)		H-3 (4.65)
H-5 (6.61)		H-6 (7.32)
H-6 (7.32)		H-5 (6.61), H-7 (6.80)
H-7 (6.80)		H-6 (7.32)
H-9 (1.47)		H-3 (4.65)

AE3**5-Hydroxy-2-(4-hydroxy-2, 5-dimethoxyphenyl)-7-methoxy-3-(3-methylbut-2-enyl)-4H-chromen-4-one**

AE3 is a yellow gum. Its molecular formula of $C_{23}H_{24}O_7$ was established on the basis of mass spectrum, EI-MS ($[M]^+$ m/z 412.1542). The IR spectrum showed the absorption bands of a hydroxyl group at 3445 cm^{-1} and carbonyl group at 1653 cm^{-1} . The ^1H NMR showed singlet signals of a hydrogen-bonded hydroxyl group (5-OH) at δ 13.00, non-bonded hydroxyl group (4'-OH) at δ 5.93 and three methoxyl groups (7-OCH₃, 2'-OCH₃, and 5'-OCH₃) at δ 3.83, 3.75, and 3.86. The singlet signals at δ 6.67 and 6.83 were assigned for aromatic protons H-3' and H-6' whereas singlet signals at δ 6.35 with integration of two protons was proposed for *meta*-aromatic protons H-6 and H-8. The presence of a prenyl group was observed from the characteristic signals of methylene protons (H-9, δ 3.04, *d*), a methine proton (H-10, δ 5.11, *t*, 6.6 Hz), methyl protons (CH₃-12, δ 1.42, *s* and CH₃-13, δ 1.62, *s*). This side chain was placed at C-3 position due to the HMBC correlation of H-9 to the C=O (δ 182.4) and C-2 (δ 160.7). The position of 7-OCH₃ was confirmed by HMBC correlation of methyl protons at δ 3.83, H-6 and H-8 to C-7 and the differential NOE technique by irradiation of the signal of H-6 and H-8 which enhanced the signal of 7-OCH₃. Furthermore, the locations of two methoxyl groups were assigned at C-2' and C-5' which were supported by the differential NOE technique; irradiation of a H-3' and H-6' enhanced the signals of 2'-OCH₃ and 5'-OCH₃. The HMBC experiment confirmed the structure of **AE3** as a new 3-prenylflavone derivative, 5-hydroxy-2-(4-hydroxy-2,5-dimethoxyphenyl)-7-methoxy-3-(3-methylbut-2-enyl)-4H-chromen-4-one.



Major HMBC of AE3



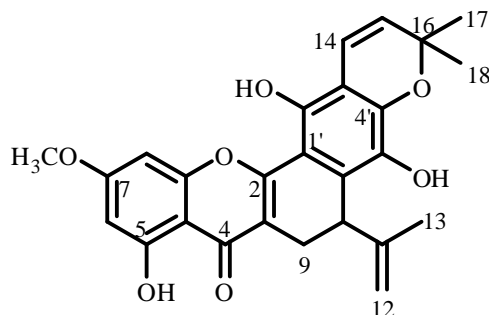
NOE of AE3

Table 9 ^{13}C , ^1H and HMBC spectral data of **AE3**

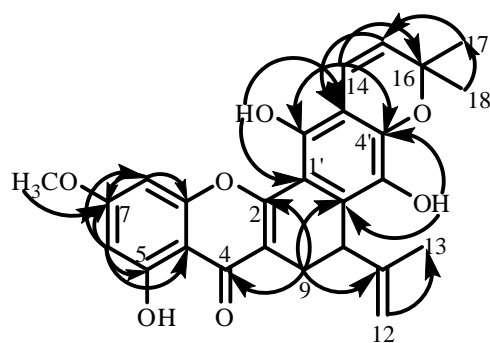
Position	δ_{C} (C-Type)	δ_{H} (mult, J_{Hz})	HMBC
2	160.7 (C)	-	-
3	121.4 (C)	-	-
4	182.4 (C=O)	-	-
4a	105.5 (C)	-	-
5	162.2 (C)	-	-
6	97.8 (CH)	6.35 (1H, <i>s</i>)	C-4, C-4a, C-5, C-7, C-8
7	165.3 (C)	-	-
8	92.0 (CH)	6.35 (1H, <i>s</i>)	C-4, C-4a, C-7, C-6, C-8a
8a	158.2 (C)	-	-
9	24.2 (CH ₂)	3.04 (2H, <i>d</i> , 6.6)	C-2, C-3, C-4, C-11
10	121.7 (CH)	5.11 (1H, <i>t</i> , 6.6)	-
11	132.0 (C)	-	-
12	17.6 (CH ₃)	1.42 (3H, <i>s</i>)	C-9, C-10, C-11, C-13
13	25.7 (CH ₃)	1.62 (3H, <i>s</i>)	C-10, C-11, C12
1'	112.7 (C)	-	-
2'	152.3 (C)	-	-
3'	99.7 (CH)	6.67 (1H, <i>s</i>)	C-1', C-2', C-4', C-5'
4'	148.5 (C)	-	-
5'	140.3 (C)	-	-
6'	112.9 (CH)	6.83 (1H, <i>s</i>)	C-2, C-4', C-5'
5-OH	-	13.00 (1H, <i>s</i>)	C-4a, C-5, C-6
4'-OH	-	5.93 (1H, <i>s</i>)	C-4', C-5'
7-OCH ₃	56.7 (OCH ₃)	3.83 (3H, <i>s</i>)	C-7
2'-OCH ₃	55.7 (OCH ₃)	3.75 (3H, <i>s</i>)	C-2'
5'-OCH ₃	56.2 (OCH ₃)	3.86 (3H, <i>s</i>)	C-5'

Table 10 ^1H - ^1H COSY spectral data of **AE3**

Proton (δ_{ppm})	↔	Correlated proton (δ_{ppm})
H-9 (3.04)		H-10 (5.11)

AE4**6,7-Dihydro-5,9,14-trihydroxy-11-methoxy-3,3-dimethyl-6-(1-methylethyl)-3*H*,8*H*-[1]benzopyrano[7,6-*c*]xanthen-8-one**

AE4 is a yellow solid, mp 205-207 °C. The UV spectrum exhibited absorption maxima at 214, 271, 304 and 378 nm. The IR spectrum showed the stretching band of O-H at 3406 cm^{-1} and C=O at 1653 cm^{-1} . The ^1H NMR showed signals of three hydroxyl groups (δ 12.99, 5-OH; δ 7.76, 2'-OH; δ 5.27, 5'-OH), two *meta* aromatic protons (δ 6.37, H-6 and H-8), and a methoxyl group (δ 3.86, 7-OCH₃). The characteristic signals of a 2,2-dimethylchromene ring were shown at δ 5.64 (*d*, H-15), δ 6.74 (*d*, H-14), δ 1.49 (*s*, CH₃-18), and δ 1.52 (*s*, CH₃-17). It was placed at C-3' and C-4' position due to the HMBC correlation (**Table 11**) of H-14 to C-2', C-3', C-4' and of H-15 to C-3'. The ^1H NMR spectrum further showed an ABX system signal of non-equivalent methylene protons H _{α} -9 (δ 2.54, *dd*, $J = 16.2, 6.9$ Hz), H _{β} -9 (δ 3.40, *dd*, $J = 16.2, 1.5$ Hz) and a methine proton H-10 (δ 3.96, *d*, $J = 6.9$ Hz). The signal of non-equivalent vinylic protons (δ 4.35, *s*, H _{α} -12 and δ 4.71, *s*, H _{β} -12) and methyl protons (δ 1.85, *s*, H-13), corresponding to an isopropenyl group, were shown in the spectrum. The 3J HMBC correlations of H-10 to C-3, C-12, C-1', C-5' suggested the point of attachment of C-10 to isopropenyl group and to C-6' of the aromatic ring. This evidence indicated that the cyclic was formed between C-3 and C-6' position, whereas the isoprenyl group was linked at C-10. The constant value 6.9 Hz of H _{β} -9 and H-10 suggested the *trans*-axial position of these two protons, consequently the isopropenyl group was in agreement. These signals are the characteristic signals of a dihydrobenzoxanthone skeleton (Hakim, *et al.*, 2006). The spectral data and assignments corresponded to the previously isolated, **artanol E** (Aida, *et al.*, 1997).



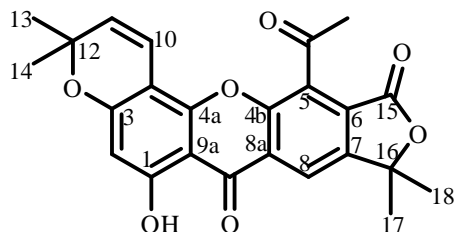
Major HMBC of AE4

Table 11 ^{13}C , ^1H and HMBC spectral data of **AE4**

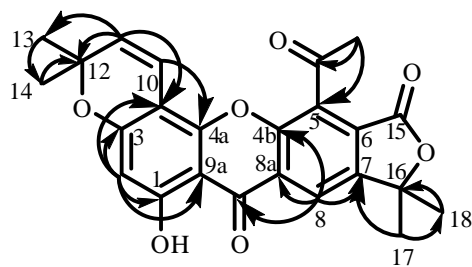
Position	δ_{C} (C-Type)	δ_{H} (<i>mult</i> , J_{Hz})	HMBC
2	159.9 (C)	-	-
3	111.7 (C)	-	-
4	180.1 (C=O)	-	-
4a	105.0 (C)	-	-
5	162.4 (C)	-	-
6	98.2 (CH)	6.37 (1H, <i>s</i>)	C-4, C-4a, C-5, C-7, C-8
7	165.1 (C)	-	-
8	92.1 (CH)	6.37 (1H, <i>s</i>)	C-4, C-4a, C-5, C-6, C-7, C-8a
8a	155.7 (C)	-	-
9	21.6 (CH ₂)	3.40 (1H _{β} , <i>dd</i> , 16.2, 1.5) 2.54 (1H _{α} , <i>dd</i> , 16.2, 6.9)	C-2, C-3, C-4, C-6', C-10, C-11 C-2, C-3, C-4, C-6', C-10, C-11
10	36.6 (CH)	3.96 (1H, <i>d</i> , 6.9)	C-1', C-5', C-6', C-9, C-11, C-12
11	144.3 (C)	-	-
12	111.9 (CH ₂)	4.71 (1H _{β} , <i>s</i>) 4.35 (1H _{α} , <i>s</i>)	C-9 C-9, C-10, C-11
13	21.6 (CH ₃)	1.85 (3H, <i>s</i>)	C-10, C-11, C-12
14	116.3 (CH)	6.74 (1H, <i>d</i> , 10.0)	C-2', C-3', C-4', C-16
15	128.6 (CH)	5.64 (1H, <i>d</i> , 10.0)	C-3', C-16, C-17
16	78.5 (C)	-	-
17	28.3 (CH ₃)	1.52 (3H, <i>s</i>)	C-15, C-16, C-18
18	28.2 (CH ₃)	1.49 (3H, <i>s</i>)	C-15, C-16, C-17
1'	105.1 (C)	-	-
2'	145.0 (C)	-	-
3'	108.8 (C)	-	-
4'	143.9 (C)	-	-
5'	135.6 (C)	-	-
6'	126.7 (C)	-	-
5-OH	-	12.99 (1H, <i>s</i>)	C-4a, C-5, C-6
2'-OH	-	7.76 (1H, <i>s</i>)	C-1', C-2', C-3'
5'-OH	-	5.27 (1H, <i>s</i>)	C-4', C-5', C-6'
7-OCH ₃	55.8 (OCH ₃)	3.86 (3H, <i>s</i>)	C-7

Table 12 ^1H - ^1H COSY spectral data of **AE4**

Proton (δ_{ppm})	\longleftrightarrow	Correlated proton (δ_{ppm})
H $_{\alpha}$ -9 (2.54)		H $_{\beta}$ -9 (3.40), H-10 (3.96)
H $_{\beta}$ -9 (3.40)		H $_{\alpha}$ -9 (2.54), H-10 (3.96)
H-10 (3.96)		H $_{\alpha}$ -9 (2.54), H $_{\beta}$ -9 (3.40)
H-14 (6.74)		H-15 (5.64)

AE5**12-Acetyl-6-hydroxy-3,3,9,9-tetramethyl-3*H*,7*H*,furo[3,4-*b*]pyrano[3,2-*h*]xanthene-7,11(9*H*)-dione**

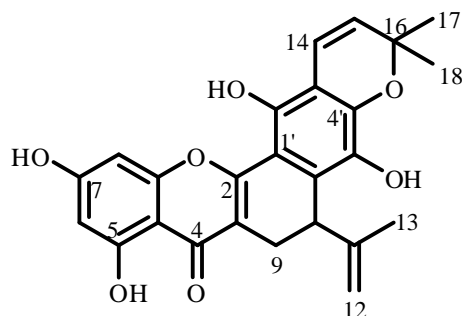
AE5 is an orange solid, m.p. 190-193 °C. The UV spectrum showed specific absorptions with maxima at 207, 240, 275, 285, 314, 338 and 411 nm. The IR spectrum showed the stretching bands of O-H at 3455 cm^{-1} and C=O at 1731 and 1653 cm^{-1} . The ^1H NMR spectrum exhibited a sharp singlet signal of a hydroxyl proton which formed an intramolecular hydrogen bond to a carbonyl group at δ 12.50 and signals of two isolated aromatic protons at δ 6.32 (s, H-2) and δ 8.31 (s, H-8). The HMBC correlations of H-2 to C-1, C-3, C-4, C-9a and of H-8 to C-4b, C-6, C-8a, C-9, C-16 confirmed the location of aromatic protons at C-2 and C-8 position, respectively. The presence of characteristic signals of 2,2-dimethylchromene ring were observed from doublet signal of methylene protons at δ 6.60 (H-10) and 5.63 (H-11) and a singlet signal with integration of six protons at δ 1.50 (CH_3 -13 and CH_3 -14). The location of the chromene moiety at C-3 and C-4 was supported by HMBC correlations of H-11 to C-3, C-4, C-4a; H-12 to C-4, and of H-2 to C-1, C-3, C-4. The methyl protons of an acetyl group were indicated from the proton signal at δ 2.82 (s) and carbon signal of C=O at δ 198.5. It was placed at C-5 position due to HMBC correlation of $-\text{COCH}_3$ to C-5. Moreover, the signals of two equivalent methyl groups at δ 1.76 (s, CH_3 -17 and CH_3 -18) and the presence of a carbonyl carbon resonance in lactone moiety at δ 169.4 suggested the xanthonolide skeleton. The HMBC correlations of H₃-17 (H₃-18) to C-16 and C-7 suggested the point of attachment of C-16, C-17 (C-18) and C-7 of aromatic ring. The lactone functionality was assigned from the carbonyl carbon signal at δ 169.4. These spectral data corresponded to those of **artanol B**, which was first isolated from *Artocarpus communis* (Aida, *et al.*, 1997).

Major HMBC of **AE5****Table 13** ^{13}C , ^1H and HMBC spectral data of **AE5**

Position	δ_{C} (C-Type)	δ_{H} (mult, J_{Hz})	HMBC
1	163.2 (C)	-	-
2	100.2 (CH)	6.32 (1H, s)	C-1, C-3, C-4, C-9a
3	162.3 (C)	-	-
4	101.4 (C)	-	-
4a	152.5 (C)	-	-
4b	152.5 (C)	-	-
5	131.0 (C)	-	-
6	125.2 (C)	-	-
7	148.6 (C)	-	-
8	119.2 (CH)	8.31 (1H, s)	C-4b, C-6, C-8a, C-9, C-16
8a	125.2 (C)	-	-
9	179.1 (C=O)	-	-
9a	103.5 (C)	-	-
10	114.2 (CH)	6.60 (1H, d, 10.1)	C-3, C-4, C-4a, C-12
11	128.2 (CH)	5.63 (1H, d, 10.1)	C-4, C-12, C13, C-14
12	79.1 (C)	-	-
13	28.5 (CH ₃)	1.50 (3H, s)	C-11, C-12, C-14
14	28.5 (CH ₃)	1.50 (3H, s)	C-11, C-12, C-13
15	169.4 (C=O)	-	-
16	86.6 (C)	-	-
17	27.6 (CH ₃)	1.76 (3H, s)	C-7, C-16, C-18
18	27.6 (CH ₃)	1.76 (3H, s)	C-7, C-16, C-17
5-C $\underline{\text{O}}$ CH ₃	198.5 (C=O)	-	-
5-CO $\underline{\text{C}}$ H ₃	32.3 (CH ₃)	2.82 (3H, s)	C-5, 5-C $\underline{\text{O}}$ CH ₃
1-OH	-	12.50 (1H, s)	C-1, C-2, C-9a

Table 14 ^{13}C and ^1H spectral data of **artonol B**

Position	δ_{C} (C-Type)	δ_{H} (mult, J_{Hz})
1	163.2 (C)	-
2	100.2 (CH)	6.31 (s)
3	162.3 (C)	-
4	101.4 (C)	-
4a	151.3 (C)	-
4b	151.1 (C)	-
5	130.2 (C)	-
6	126.5 (C)	-
7	148.6 (C)	-
8	119.2 (CH)	8.30 (s)
8a	125.2 (C)	-
9	179.2 (C=O)	-
9a	103.6 (C)	-
10	114.2 (CH)	6.60 (1H, <i>d</i> , 10.1)
11	128.2 (CH)	5.63 (1H, <i>d</i> , 10.1)
12	79.1 (C)	-
13	28.5 (CH)	1.50 (3H, <i>s</i>)
14	28.5 (CH)	1.50 (3H, <i>s</i>)
15	166.7 (C=O)	-
16	86.6 (C)	-
17	27.6 (CH ₃)	1.76 (3H, <i>s</i>)
18	27.6 (CH ₃)	1.76 (3H, <i>s</i>)
5- <u>C</u> OCH ₃	198.5 (C)	-
5-CO <u>C</u> H ₃	32.3 (CH)	2.81 (3H, <i>s</i>)
1-OH	-	12.49 (1H, <i>s</i>)

AE6**6,7-Dihydro-5,9,11,14-tetrahydroxy-3,3-dimethyl-6-(1-methylethenyl)-(-)-3*H*,8*H*-Pyrano[3',2':4,5]benzo[1,2-*c*]xanthen-8-one**

AE6 is a red-brown gum. The UV spectrum showed the absorption bands at 263, 269, 307 and 379. The IR spectrum indicated the presence of O-H stretching at 3402 cm^{-1} and C=O stretching at 1655 cm^{-1} . The ^1H NMR spectral data showed the signals of a chelated hydroxyl proton 5-OH (δ 12.98), two phenolic hydroxyl groups 2'-OH and 5'-OH (δ 7.78 and 5.46), and meta-coupled aromatic protons H-6 and H-8 (δ 6.35 and 6.40) and the signals of protons of the 2,2-dimethylchromene ring (δ 1.52, 17-CH₃; δ 1.49, 18-CH₃; δ 5.64, H-15 and δ 6.74, H-14). The spectrum also showed the characteristic signals of a dihydrobenzoxanthone skeleton (H _{α} -9, δ 2.53; H _{β} -9, δ 3.39; H-10, δ 3.96; H _{α} -12, δ 4.34; H _{β} -12, δ 4.71 and 13-CH₃, δ 1.81). The chemical shift values and coupling patterns of all proton signals were similar to those of relevant protons of **artanol E**. The difference was the disappearance of a singlet signal of a methoxyl group at δ 3.86 in **AE6**. It was then proposed as **7-demethylartanol E** (Namdaung, *et al.*, 2006) or **artelastoxanthone** (Ko, *et al.*, 2005). The assignment was also confirmed by the HMBC experiment (**Table 15**).

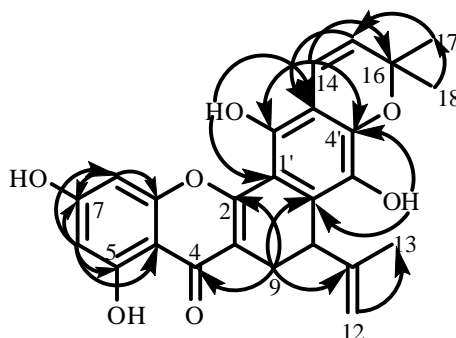
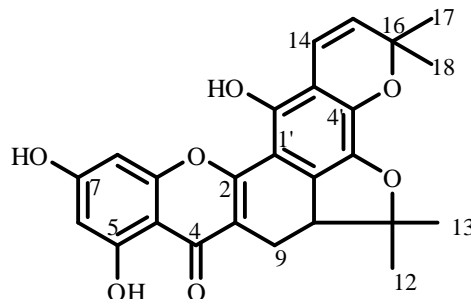
Major HMBC of **AE6**

Table 15 ^{13}C , ^1H and HMBC spectral data of **AE6**

Position	δ_{C} (C-Type)	δ_{H} (<i>mult</i> , J_{Hz})	HMBC
2	159.7 (C)	-	-
3	111.5 (C)	-	-
4	180.0 (C=O)	-	-
4a	104.5 (C)	-	-
5	162.4 (C)	-	-
6	99.8 (CH)	6.35 (1H, <i>d</i> , 1.8 Hz)	C-4a, C-5, C-7, C-8
7	163.0 (C)	-	-
8	93.6 (CH)	6.40 (1H, <i>d</i> , 1.8 Hz)	C-4a, C-6, C-7, C-8a
8a	155.9 (C)	-	-
9	21.5 (CH ₂)	3.39 (1H _{β} , <i>dd</i> , 16.2, 1.5) 2.53 (1H _{α} , <i>dd</i> , 16.2, 6.9)	C-2, C-3, C-4, C-6', C-10, C-11 C-2, C-3, C-10, C-11
10	36.5 (CH)	3.96 (1H, <i>d</i> , 6.9)	C-1', C-3, C-5', C-6', C-9, C-11, C-12, C-13
11	144.3 (C)	-	-
12	111.7 (CH ₂)	4.71 (1H _{β} , <i>s</i>) 4.34 (1H _{α} , <i>s</i>)	C-10, C-13 C-10, C-13
13	21.6 (CH ₃)	1.81 (3H, <i>s</i>)	C-10, C-11, C-12
14	116.3 (CH)	6.74 (1H, <i>d</i> , 10.0)	C-2', C-3', C-4', C-16
15	128.5 (CH)	5.64 (1H, <i>d</i> , 10.0)	C-3', C-16, C-17, C-18
16	78.3 (C)	-	-
17	28.2 (CH ₃)	1.52 (3H, <i>s</i>)	C-15, C-16, C-18
18	28.1 (CH ₃)	1.49 (3H, <i>s</i>)	C-15, C-16, C-17
1'	105.2 (C)	-	-
2'	144.9 (C)	-	-
3'	108.8 (C)	-	-
4'	143.8 (C)	-	-
5'	135.6 (C)	-	-
6'	126.7 (C)	-	-
5-OH	-	12.98 (1H, <i>s</i>)	C-4, C-4a, C-5, C-6
2'-OH	-	7.78 (1H, <i>s</i>)	C-1', C-2', C-3'
5'-OH	-	5.46 (1H, <i>s</i>)	C-4', C-5', C-6'

AE7**New furanodihydrobenzoxanthone derivative**

AE7 is a yellow solid, m.p. 287-289 °C. The molecular formula, $C_{25}H_{22}O_7$, was deduced from the mass spectrum, EI-MS ($[M]^+$ m/z 434.1377). The UV spectrum showed maximum absorption bands at 214, 253, 273, 289, 308 and 375 nm. The IR spectrum showed the stretching of O-H (3442 cm^{-1}) and C=O (1630 cm^{-1}). The ^1H NMR spectrum exhibited a singlet signal of a chelated hydroxyl proton (δ 13.00, 5-OH) and two singlet signals of non-chelated hydroxyl protons (δ 9.42, 7-OH) and δ 7.17 (2'-OH). The appearance of two meta-coupled signals at δ 6.31 and 6.47 with coupling constant of 2.1 Hz were assigned for aromatic protons H-6 and H-8. The HMBC correlation of H-6 to C-4a, C-5, C-7, C-8, and H-8 to C-4a, C-6, C-7, C-8a were also confirmed the positions of H-6 and H-8. Two singlet signals of two methyl groups (δ 1.35, *s*, CH₃-12 and δ 1.68, *s*, CH₃-13), ABX system signals of a methine proton (δ 3.39, *dd*, H-10) and methylene protons (δ 3.21, *dd*, H _{β} -9 and 2.40, *t*, H _{α} -9) were in agreement with the characteristic signals of a furanodihydrobenzoxanthone skeleton (Hakim, *et al.*, 2006). The HMBC correlations of H-10 to C-3, C-9, C-11, C-12, C-13, C-6' suggested that the cyclic was formed between C-3 and C-6' whereas the furan moiety was formed at C-5' and C-6' of the aromatic ring. The remaining signals were assigned for 2,2-dimethylchromene ring of which two vicinal protons H-14 and H-15 appeared as two doublet signals at δ 6.69 and 5.57, and two germinal methyl groups resonating at δ 1.50 (H₃-17) and δ 1.47 (H₃-18). The correlations of H-14 to C-2', C-5'; of H-15 to C-3', and of 2'-OH to C-1', C-2', C-3' correctly determined that the chromene ring was at C-3' and C-4' position. Therefore a new furanodihydrobenzoxanthone derivative was assigned for **AE7**.

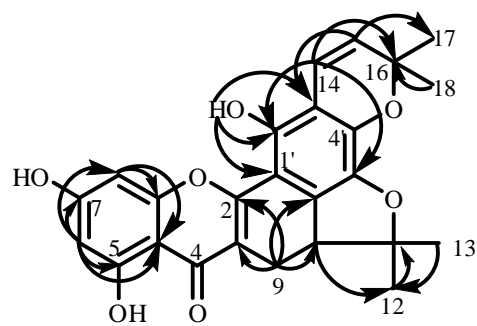
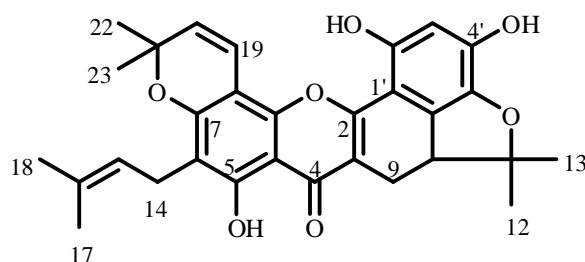
Major HMBC of **AE7**

Table 16 ^{13}C , ^1H and HMBC spectral data of **AE7**

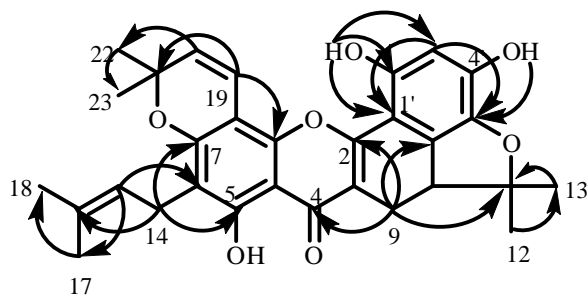
Position	δ_{C} (C-Type)	δ_{H} (<i>mult</i> , J_{Hz})	HMBC
2	165.1 (C)	-	-
3	117.4 (C)	-	-
4	185.4 (C=O)	-	-
4a	109.4 (C)	-	-
5	167.5 (C)	-	-
6	104.9 (CH)	6.31 (1H, <i>d</i> , 2.1)	C-4a, C-5, C-7, C-8
7	168.7 (C)	-	-
8	99.0 (CH)	6.47 (1H, <i>d</i> , 2.1)	C-4a, C-6, C-7, C-8a
8a	161.5 (C)	-	-
9	24.9 (CH ₂)	3.21 (1H _{β} , <i>dd</i> , 15.3, 7.2) 2.40 (1H _{α} , <i>t</i> , 15.3)	C-2, C-3, C-4, C-6', C-10 C-2, C-3, C-10, C-11
10	51.5 (CH)	3.39 (1H, <i>dd</i> , 15.3, 7.2)	C-5', C-6', C-9, C-11, C-12, C-13
11	98.7 (C)	-	-
12	27.6 (CH ₃)	1.35 (3H, <i>s</i>)	C-10, C-11
13	33.2 (CH ₃)	1.68 (3H, <i>s</i>)	C-10, C-11, C-12
14	121.9 (CH)	6.69 (1H, <i>d</i> , 10.2)	C-2', C-5', C-16
15	133.2 (CH)	5.57 (1H, <i>d</i> , 10.2)	C-3', C-16, C-17, C-18
16	82.7 (C)	-	-
17	33.1 (CH ₃)	1.50 (3H, <i>s</i>)	C-16, C-18
18	33.0 (CH ₃)	1.47 (3H, <i>s</i>)	C-16, C-17
1'	108.5 (C)	-	-
2'	149.6 (C)	-	-
3'	116.1 (C)	-	-
4'	147.2 (C)	-	-
5'	142.6 (C)	-	-
6'	136.4 (C)	-	-
5-OH	-	13.00 (1H, <i>s</i>)	C-4a, C-5, C-6
7-OH	-	9.42 (1H, <i>s</i>)	-
2'-OH	-	7.17 (1H, <i>s</i>)	C-1', C-2', C-3'

Table 17 ^1H - ^1H COSY spectral data of **AE7**

Proton (δ_{ppm})	\longleftrightarrow	Correlated proton (δ_{ppm})
H-6 (6.31)		H-8 (6.47)
H $_{\alpha}$ -9 (2.40)		H $_{\beta}$ -9 (3.21), H-10 (3.39)
H $_{\beta}$ -9 (3.21)		H $_{\alpha}$ -9 (2.40), H-10 (3.39)
H-10 (3.39)		H $_{\alpha}$ -9 (2.40), H $_{\beta}$ -9 (3.21)
H-14 (6.69)		H-15 (5.57)

AE8**5a,6-Dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-9-(3-methyl-2-buten-1-yl)-5H,7H,11H-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one**

AE8 is a yellow solid, m.p. 249-251 °C. The UV spectrum showed maximum absorptions at 236, 256, 265, 278, 335 and 390 nm. The IR spectrum showed the stretching of hydroxyl group (3371 cm^{-1}) and carbonyl group (1629 cm^{-1}). The ^1H NMR spectrum showed a singlet signal of a chelated proton 5-OH at δ 13.42 and two singlet signals of non chelated protons 2'-OH and 4'-OH at δ 7.78 and 9.23, respectively. An aromatic proton at δ 6.38 (*s*) was assigned for H-3'. The spectrum further showed the signals corresponding to furanoxanthone moiety (δ 2.40, *t*, $J = 15.0\text{ Hz}$, H $_{\alpha}$ -9; 3.23, *dd*, $J = 15.0, 7.2\text{ Hz}$, H $_{\beta}$ -9; 3.41, *dd*, $J = 15.0, 7.2\text{ Hz}$, H-10). The prenyl unit commemorated from distinctive signals of two equivalent methylene protons at δ 3.33 (*d*, H-14), olefinic proton at δ 5.24 (*t*, H-15) and two methyl protons at δ 1.68 (*s*, 17-CH $_3$) and 1.81 (*s*, 18-CH $_3$). Its location was assigned at C-6 by HMBC correlation of OH-5 and of H-15 to C-6. Moreover, a chromene ring was detected from the characteristic signals at δ 5.57 (*d*, H-20), 6.76 (*d*, H-19) and 1.46 (*s*, 22-CH $_3$ and 23-CH $_3$). The correlations of H-19 to C-8a, and of H-20 to C-8, confirmed the orientation of 2,2-dimethylchromene ring at C-7 and C-8 position. The assigned structure of **AE8** was in agreement with **artonin F** (Hano, Y. *et al.* 1990).



Major HMBC of AE8

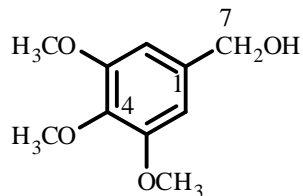
Table 18 ^{13}C , ^1H and HMBC spectral data of **AE8**

Position	δ_{C} (C-Type)	δ_{H} (mult, J_{Hz})	HMBC
2	160.3 (C)	-	-
3	111.5 (C)	-	-
4	180.7 (C=O)	-	-
4a	104.3 (C)	-	-
5	158.7 (C)	-	-
6	112.5 (C)	-	-
7	156.4 (C)	-	-
8	100.5 (CH)	-	-
8a	149.2 (C)	-	-
9	19.9 (CH ₂)	3.23 (1H _{β} , <i>dd</i> , 15.0, 7.2) 2.40 (1H _{α} , <i>t</i> , 15.0)	C-2, C-3, C-4, C-6', C-10 C-2, C-3, C-4, C-6', C-10, C-11
10	46.6 (CH)	3.41 (1H, <i>dd</i> , 15.0, 7.2)	C-1', C-5', C-6', C-9, C-11, C-12, C-13
11	93.5 (C)	-	-
12	22.7 (CH ₂)	1.35 (3H, <i>s</i>)	C-10, C-11, C-13
13	28.1 (CH ₃)	1.66 (3H, <i>s</i>)	C-10, C-12
14	21.3 (CH ₂)	3.33 (2H, <i>d</i> , 7.2)	C-5, C-6, C-7, C-15, C-16
15	122.1 (CH)	5.24 (1H, <i>t</i> , 7.2)	C-6, C-14, C-17, C-18
16	131.3 (C)	-	-
17	17.9 (CH ₃)	1.68 (3H, <i>s</i>)	C-15, C-16, C-18
18	25.8 (CH ₃)	1.81 (3H, <i>s</i>)	C-15, C-16, C-17
19	115.3 (CH)	6.76 (1H, <i>d</i> , 9.9)	C-7, C-8, C-8a, C-20, C-21
20	127.1 (CH)	5.57 (1H, <i>d</i> , 9.9)	C-8, C-21, C-22, C-23
21	77.3 (C)	-	-
22	28.0 (CH ₃)	1.46 (3H, <i>s</i>)	C-20, C-21, C-23
23	28.0 (CH ₃)	1.46 (3H, <i>s</i>)	C-20, C-21, C-22
1'	103.4 (C)	-	-
2'	150.1 (C)	-	-
3'	104.6 (C)	6.38 (1H, <i>s</i>)	C-1', C-2, C-2', C-4', C-5'
4'	146.4 (C)	-	-
5'	137.2 (C)	-	-
6'	131.8 (C)	-	-
5-OH	-	13.42 (1H, <i>s</i>)	C-4, C-4a, C-5, C-6
2'-OH	-	7.78 (1H, <i>s</i>)	C-1', C-2', C-3'
4'-OH	-	9.23 (1H, <i>s</i>)	C-5'

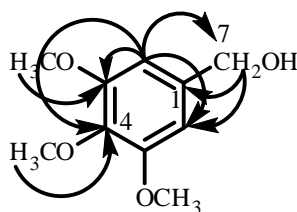
Table 19 ^1H - ^1H COSY spectral data of **AE8**

Proton (δ_{ppm})		Correlated proton (δ_{ppm})
H $_{\alpha}$ -9 (2.40)	↔	H $_{\beta}$ -9 (3.23), H-10 (3.41)
H $_{\beta}$ -9 (3.23)	↔	H $_{\alpha}$ -9 (2.40), H-10 (3.41)
H-10 (3.41)	↔	H $_{\alpha}$ -9 (2.40), H $_{\beta}$ -9 (3.23)
H-14 (3.33)	↔	H-15 (5.24), H-17 (1.68), H-18 (1.81)
H-15 (5.24)	↔	H-14 (3.33), H-17 (1.68), H-18 (1.81)
H-17 (1.68)	↔	H-14 (3.33), H-15 (5.24), H-18 (1.81)
H-18 (1.81)	↔	H-14 (3.33), H-15 (5.24), H-17 (1.68)
H-19 (6.76)	↔	H-20 (5.57)

AE9
(3,4,5-Trimethoxyphenyl)methanol



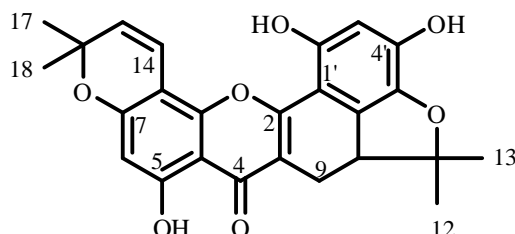
AE9 is a red-brown gum. The ^1H NMR spectrum exhibited a singlet resonance of two equivalent meta-aromatic protons at δ 6.61 (H-2 and H-6) indicating a tetrasubstituted benzene ring. The remaining proton resonances were those of a hydroxy methylene protons (δ 4.65, *s*, 2H), a methoxyl group (δ 3.85, *s*, 6H) and two equivalent methoxyl group (δ 3.88, *s*, 6H). These substituent groups were placed at C-1, C-4, C-3 and C-5, respectively. The placement of 1- CH_2OH was confirmed by HMBC correlation of $-\text{CH}_2$ to C-2 and C-3. These assignment indicated that **AE9** was (3,4,5-trimethoxyphenyl)methanol.



Major HMBC of **AE9**

Table 20 ^{13}C , ^1H and HMBC spectral data of **AE9**

Position	δ_{C} (C-Type)	δ_{H} (<i>mult</i> , J_{Hz})	HMBC
1	135.5 (C)	-	-
2	103.9 (CH)	6.61 (1H, <i>s</i>)	C-3, C-4, C-5, C-6, C-7
3	153.5 (C)	-	-
4	137.5 (C)	-	-
5	153.5 (C)	-	-
6	103.9 (CH)	6.61 (1H, <i>s</i>)	C-2, C-3, C-4, C-5, C-7
7	65.6 (CH ₂)	4.65 (2H, <i>s</i>)	C-1, C-2, C-6
3-OCH ₃	56.1 (OCH ₃)	3.88 (3H, <i>s</i>)	C-3, C-5
4-OCH ₃	60.9 (OCH ₃)	3.85 (3H, <i>s</i>)	C-4
5-OCH ₃	56.1 (OCH ₃)	3.88 (3H, <i>s</i>)	C-3, C-5

AE10**5a,6-Dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-5H,7H,11H-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one**

AE10 is a yellow solid, m.p. 284-285 °C. The UV spectrum showed maximum absorption bands at 228, 257, 272, 312, 330 and 391 nm. The IR spectrum showed the absorption bands of O-H stretching at 3405 cm^{-1} , C=O stretching at 1642 cm^{-1} . The ^1H NMR spectrum indicated the presence of a chelated hydroxyl group (δ 13.22, 5-OH), an aromatic proton H-3' (δ 6.38) and a furanoxanthonoid moiety (δ 3.38, H-10; δ 3.21, H $_{\beta}$ -9; δ 2.40, H $_{\alpha}$ -9; δ 1.66, 13-CH $_3$ and δ 1.34, 12-CH $_3$). The ^1H NMR spectrum (**Table 21**) further showed a signal of an aromatic proton (δ 6.23, H-6, *s*) and a characteristic signals of a 2,2-dimethylchromene ring (δ 1.46, 17-CH $_3$ and 18-CH $_3$, 5.56, H-15 and δ 6.78, H-14). The HMBC correlations of H-6 to C-5, C-4a, C-7, C-8 and H-3' to C-1', C-2', C-4', C-5' confirmed the assignment of H-6 and H-3'. The HMBC correlations of H-14 to C-6, C-8, C-8a and of H-15 to C-8 confirmed the placement of the chromene ring at C-7 and C-8. **AE10** was identified to be **cycloartobiloxanthone** (Sultanbawa, *et al.*, 1989).

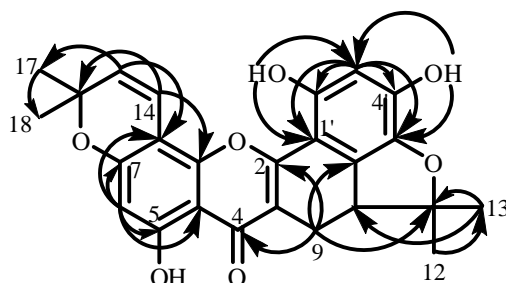
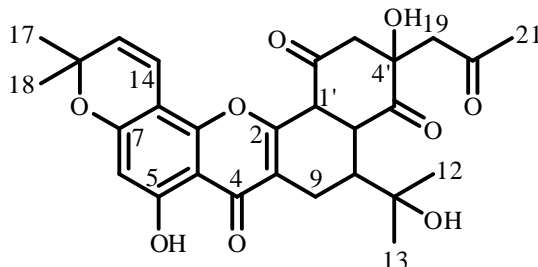
Major HMBC of **AE10**

Table 21 ^{13}C , ^1H and HMBC spectral data of **AE10**

Position	δ_{C} (C-Type)	δ_{H} (mult, J_{Hz})	HMBC
2	160.7 (C)	-	-
3	111.6 (C)	-	-
4	180.6 (C=O)	-	-
4a	104.7 (C)	-	-
5	161.5 (C)	-	-
6	99.9 (CH)	6.23 (1H, <i>s</i>)	C-4, C-4a, C-5, C-7, C-8
7	158.7 (C)	-	-
8	101.0 (C)	-	-
8a	150.9 (C)	-	-
9	19.8 (CH ₂)	3.21 (1H _{β} , <i>dd</i> , 15.0, 7.2) 2.40 (1H _{α} , <i>t</i> , 15.0)	C-2, C-3, C-4, C-6', C-10 C-2, C-3, C-4, C-6', C-10, C-11
10	46.6 (CH)	3.38 (1H, <i>dd</i> , 15.0, 7.2)	C-1', C-3, C-5', C-6', C-9, C-11, C-12, C-13
11	93.5 (C)	-	-
12	22.6 (CH ₃)	1.34 (3H, <i>s</i>)	C-10, C-11, C-13
13	28.1 (CH ₃)	1.66 (3H, <i>s</i>)	C-10, C-11, C-12
14	115.0 (CH)	6.78 (1H, <i>d</i> , 10.0)	C-6, C-8, C-8a, C-15, C-16
15	127.3 (CH)	5.56 (1H, <i>d</i> , 10.0)	C-8, C-16, C-17, C-18
16	77.8 (C)	-	-
17	28.1 (CH ₃)	1.46 (3H, <i>s</i>)	C-14, C-15, C-16, C-18
18	28.1 (CH ₃)	1.46 (3H, <i>s</i>)	C-14, C-15, C-16, C-17
1'	103.4 (C)	-	-
2'	150.3 (C)	-	-
3'	104.7 (CH)	6.38 (1H, <i>s</i>)	C-1', C-2', C-4', C-5'
4'	146.5 (C)	-	-
5'	137.2 (C)	-	-
6'	131.9 (C)	-	-
5-OH	-	13.22 (1H, <i>s</i>)	C-4a, C-4, C-5, C-6
2'-OH	-	8.00 (1H, <i>s</i>)	C-1', C-2', C-3'
4'-OH	-	9.26 (1H, <i>s</i>)	C-3', C-4', C-5'

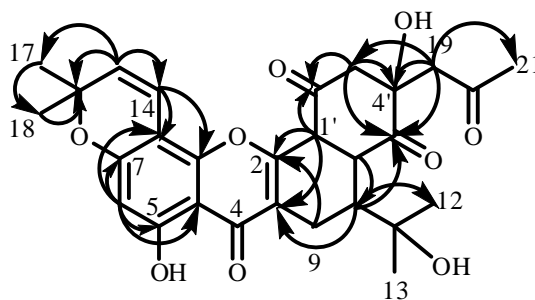
Table 22 ^1H - ^1H COSY spectral data of **AE10**

Proton (δ_{ppm})		Correlated proton (δ_{ppm})
H $_{\alpha}$ -9 (2.40)	↔	H $_{\beta}$ -9 (3.21), H-10 (3.38)
H $_{\beta}$ -9 (3.21)	↔	H $_{\alpha}$ -9 (2.40), H-10 (3.38)
H-10 (3.38)	↔	H $_{\alpha}$ -9 (2.40), H $_{\beta}$ -9 (3.21)
H-14 (6.78)	↔	H-15 (5.56)

AE11**New quinonobenzoxanthone derivative**

AE11 is a yellow solid, m.p. 199-200 °C. The UV spectrum showed the absorption bands at 217, 229, 245, 267, 303 and 354 nm. The IR spectrum indicated the presence of O-H stretching at 3433 cm^{-1} and C=O stretching at 1720, 1647 and 1584 cm^{-1} . The ^1H NMR spectral data of **AE11** in CDCl_3 (**Table 23**) exhibited the signals of a chelated phenolic hydrogen proton at δ 12.56 (*s*, 5-OH), an isolated aromatic proton at δ 6.27 (*s*, H-6) and a set of 2,2-dimethylchromene ring at δ 6.77 (*d*, H-14); 5.59 (*d*, H-15); 1.48 (*s*, 17- CH_3); 1.46 (*s*, 18- CH_3). The ^1H NMR spectra also showed the resonances of two methyl groups at δ 0.90 and 1.20 (*s*, 12- CH_3 and 13- CH_3), methylene protons at δ 2.96 and 2.58 (*dd*, $J = 16.8, 5.7$ Hz, 2H-9), three methine protons at δ 2.62 (*d*, H-1'), 2.92 (*m*, H-10), 3.50 (*t*, H-6'). ^1H - ^1H COSY correlations of H-10 to H-9, H-6' and of H-6' to H-1' confirmed the assignment of a partial structure (-C₉-C₁₀-C_{6'}-C_{1'}-). The chemical shift value of C-11 (δ 74.5) indicated that it was oxycarbon. The correlations of 2H-9 and H-1' to C-2 and C-3 supported the connection at C-3 and C-2. In addition, the resonances of signals of non-equivalent methylene protons at δ 2.67 and 2.63 (*br s*, 2H-3') showed the HMBC correlation with C-2', C-4' and C-5'. A propanoyl side chain was suggested from the proton resonances of methyl protons at δ 2.27 (CH_3 -21, *s*) and 2H-19 at δ 3.00 and 2.93 (*br s*, each). The presences of four carbonyl groups were indicated from the ^{13}C NMR spectrum. The resonance at δ 207.6 reveal the carbonyl carbon of a propanoyl side chain whereas the resonance at δ 180.9 was assigned for C-4. The quinonoid structure was implied from the carbonyl carbon resonances at δ 198.8 (C-2') and 205.5 (C-5'). The HMBC correlations of H-19 to C-3', C-4', C-5', C-20, C-21 together with 4'-OH (δ 4.30) to C-3', C-4' and C-5' indicated that the propanoyl side chain and

a hydroxyl group were attached to a quinonoid structure at C-4'. The 3J correlation of H-10 to C-3, C-5', C-11 and C-12 suggested the formation of a cyclic between C-3 and a quinonoid structure together with hydroxyl propyl group as the side chain. The chromene moiety was placed at C-7 and C-8 due to HMBC correlations of H-14 to C-7, C-8, C-8a and of H-15 to C-8. Therefore a new quinonoid dihydroxanthone structure was assigned for **AE11**.



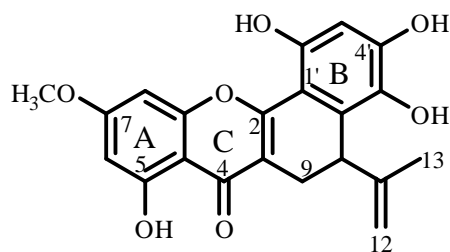
Major HMBC of **AE11**

Table 23 ^{13}C , ^1H and HMBC spectral data of **AE11**

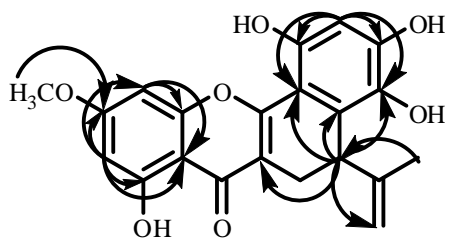
Position	δ_{C} (C-Type)	δ_{H} (<i>mult</i> , J_{Hz})	HMBC
2	156.9 (C)	-	-
3	115.9 (C)	-	-
4	180.9 (C)	-	-
4a	104.9 (C=O)	-	-
5	161.6 (C)	-	-
6	100.3 (CH)	6.27 (1H, <i>s</i>)	C-4a, C-5, C-7, C-8
7	159.7 (C)	-	-
8	101.2 (C)	-	-
8a	151.8 (C)	-	-
9	22.6 (CH ₂)	2.96 (1H, <i>dd</i> , 16.8, 5.7) 2.58 (1H, <i>dd</i> , 16.8, 5.7)	C-2, C-3, C-6', C-10 C-2, C-3, C-6', C-10
10	32.8 (CH)	2.92 (1H, <i>m</i>)	C-3, C-5', C-6', C-11, C-12
11	74.5 (C)	-	-
12	25.3 (CH ₃)	0.90 (3H, <i>s</i>)	C-10, C-11, C-13
13	31.9 (CH ₃)	1.20 (3H, <i>s</i>)	C-10, C-11, C-12
14	114.8 (CH)	6.77 (1H, <i>d</i> , 10.0)	C-7, C-8, C-8a, C-16
15	127.4 (CH)	5.59 (1H, <i>d</i> , 10.0)	C-8, C-16, C-17, C-18
16	78.2 (C)	-	-
17	28.3 (CH ₃)	1.48 (3H, <i>s</i>)	C-15, C-16, C-18
18	28.2 (CH ₃)	1.46 (3H, <i>s</i>)	C-15, C-16, C-17
19	46.9 (CH ₂)	3.00 (1H, <i>br s</i>) 2.93 (1H, <i>br s</i>)	C-3', C-4', C-5', C-20, C-21 C-3', C-4', C-5', C-20, C-21
20	207.6 (C=O)	-	-
21	32.0 (CH)	2.27 (3H, <i>s</i>)	C-19, C-20
1'	46.9 (CH)	2.62 (1H, <i>d</i> , 2.1)	C-2, C-2', C-3, C-5', C-6'
2'	198.8 (C=O)	-	-
3'	43.5 (CH ₂)	2.67 (1H, <i>br s</i>) 2.63 (1H, <i>br s</i>)	C-2', C-4', C-5' C-2', C-4', C-5'
4'	75.6 (C)	-	C-2, C-2', C-10
5'	205.5 (C=O)	-	-
6'	47.8 (CH)	3.50 (1H, <i>t</i> , 2.1)	C-2, C-2', C-10
5-OH	-	12.56 (1H, <i>s</i>)	C-4a, C-5, C-6, C-7
4'-OH	-	4.30 (1H, <i>s</i>)	C-3', C-4', C-5'
11-OH	-	3.49 (1H, <i>s</i>)	-

Table 24 ^1H - ^1H COSY spectral data of **AE11**

Proton (δ_{ppm})		Correlated proton (δ_{ppm})
H-9 (2.96)	↔	H-9 (2.58), H-10 (2.92)
H-10 (2.92)	↔	H-9 (2.96 and 2.58), H-6' (3.50)
H-14 (6.77)	↔	H-15 (5.59)
H-19 (3.00)	↔	H-19 (2.93)
H-1' (2.62)	↔	H-6' (3.50)
H-3' (2.67)	↔	H-3' (2.63)
H-6' (3.50)	↔	H-1' (2.62), H-10 (2.92)

AE12**1,3,4,8-Tetrahydroxy-10-methoxy-5-(prop-1-en-2-yl)-5H-benzo[*c*]xanthen-7-(6*H*)-one**

AE12 is a brown-yellow solid, m.p. 242-243 °C. Its molecular formula of $C_{21}H_{18}O_7$ was established on the basis of mass spectrum, EI-MS ($[M]^+$ m/z 382.1068). The UV spectrum showed maximum absorption bands at 214, 228, 261, 314 and 380 nm. The IR spectrum showed the stretching of O-H (3180 cm^{-1}), C=O (1653 cm^{-1}). The ^1H NMR spectrum (**Table 25**) exhibited signals of a hydrogen-bonded hydroxyl proton at δ 13.23 (*s*, 5-OH), a pair of meta-coupled aromatic protons at δ 6.30 and 6.65 ($J = 2.4\text{ Hz}$, H-6 and H-8), an aromatic proton at δ 6.52 (H-3') and a singlet of methoxyl group at δ 3.89 (7-OCH₃). An ABX system signal of non-equivalent methylene protons H _{α} -9 (δ 2.46, *dd*), H _{β} -9 (δ 3.41, *dd*) and a methine proton H-10 (δ 4.00, *d*) were shown in the spectrum. The spectrum further showed signals of non-equivalent vinylic protons (δ 4.29, *s*, H _{α} -12 and δ 4.65, *s*, H _{β} -12) and methyl protons (δ 1.78, *s*, H-13), corresponding to an isopropenyl group. The HMBC correlations of H-10 to C-3, C-12, C-1', C-5' and C-6' suggested that the cyclic was formed at C-3 and C-6' position, whereas the isopropenyl group was linked to the cyclic by C-10. These signals corresponded to the characteristic signals of a dihydrobenzoxanthone skeleton (Hakim, *et al.*, 2006). The 2', 4', 5'-trioxygenated pattern with H-3' resonating at δ 6.52 was proposed for B-ring based on the biogenetic pattern of constituents in *Artocarpus* genus (Hakim, *et al.*, 2006). The assigned structure was found to be the methoxy derivative of artonol E, whose methoxyl group was proposed at C-7 according to 3J correlation of OCH₃, H-6 and H-8 to C-7. Consequently, a new dihydrobenzoxanthone derivative, 1,3,4,8-tetrahydroxy-10-methoxy-5-(prop-1-en-2-yl)-5H-benzo[*c*]xanthen-7(6*H*)-one was assigned for **AE12**.



Major HMBC of **AE12**

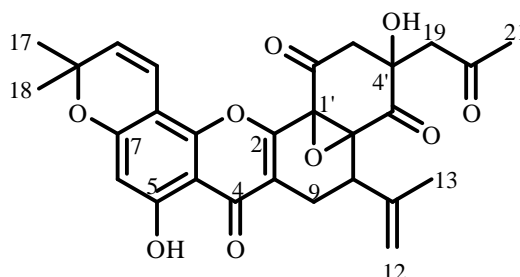
Table 25 ^{13}C , ^1H and HMBC spectral data of **AE12**

Position	δ_{C} (C-Type)	δ_{H} (mult, J_{Hz})	HMBC
2	161.0 (C)	-	-
3	110.6 (C)	-	-
4	180.1 (C=O)	-	-
4a	104.7 (C)	-	-
5	161.9 (C)	-	-
6	97.7 (CH)	6.30 (1H, <i>d</i> , 2.4)	C-4a, C-5, C-7, C-8
7	165.1 (C)	-	-
8	92.2 (CH)	6.65 (1H, <i>d</i> , 2.4)	C-4a, C-6, C-7, C-8a
8a	156.7 (C)	-	-
9	21.4 (CH ₂)	3.41 (1H _{β} , <i>dd</i> , 15.9, 1.8) 2.46 (1H _{α} , <i>dd</i> , 15.9, 6.3)	C-2, C-3, C-4, C-6', C-10, C-11 C-2, C-3, C-4, C-6', C-11
10	37.0 (CH)	4.00 (1H, <i>d</i> , 6.3)	C-3, C-5', C-6', C-9, C-11, C-12
11	144.4 (C)	-	-
12	110.9 (CH ₂)	4.65 (1H _{β} , <i>s</i>) 4.29 (1H _{α} , <i>s</i>)	C-13 C-10, C-11, C-13
13	21.0 (CH ₃)	1.78 (3H, <i>s</i>)	C-10, C-11, C-12
1'	105.5 (C)	-	-
2'	150.5 (C)	-	-
3'	102.9 (CH)	6.52 (1H, <i>s</i>)	C-1', C-2', C-4', C-5'
4'	150.3 (C)	-	-
5'	136.0 (C)	-	-
6'	128.3 (C)	-	-
7-OCH ₃	55.4 (CH ₃)	3.89 (3H, <i>s</i>)	C-7
5-OH	-	13.23 (1H, <i>s</i>)	C-4, C-4a, C-5, C-6
*OH	-	8.27 (1H, <i>s</i>)	-

* the position not identified

Table 26 ^1H - ^1H COSY spectral data of **AE12**

Proton (δ_{ppm})		Correlated proton (δ_{ppm})
H-6 (6.30)	↔	H-8 (6.65)
H _{α} -9 (2.46)	↔	H _{β} -9 (3.41), H-10 (4.00)
H _{β} -9 (3.41)	↔	H-10 (4.00), H _{α} -9 (2.46)
H-10 (4.00)	↔	H _{α} -9 (2.46), H _{β} -9 (3.41)

AE13**New quinonobenzoxanthone derivative**

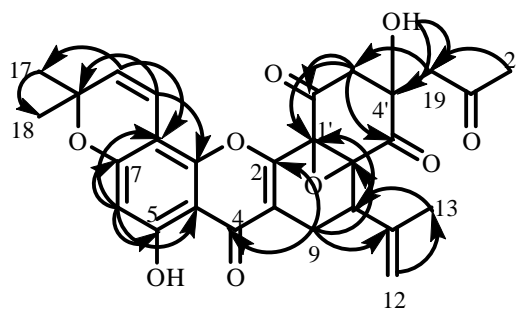
AE13 is a yellow solid, m.p. 202-203 °C. Its EI-MS exhibited a molecular ion peak at m/z 506.1572, consistent with a molecular formula of $C_{28}H_{26}O_9$. The UV spectrum showed maximum absorptions at 212, 232, 269, 308, 327 and 361 nm. The IR spectrum showed the stretching of hydroxyl (3426 cm^{-1}) and carbonyl groups (1753 , 1722 , 1697 and 1657 cm^{-1}). The ^1H NMR spectrum (**Table 27**) showed a singlet resonance of a hydrogen-bonded hydroxyl group 5-OH at δ 12.42, an aromatic proton H-6 at (δ 6.28, *s*), and protons in a 2, 2-dimethylchromene ring at δ 6.78, 5.60 (each *d*, $J = 10.2\text{ Hz}$), 1.49, 1.46 (each 3H, *s*), non-equivalent methylene protons H-3' (δ 3.47 and 3.57, *d*, $J = 19.4\text{ Hz}$ each) and a propanoyl side chain [δ 2.43 (CH_3 -21, *s*), δ 3.03 and 3.25 (2H-19, *d*)]. The spectrum further showed the resonances of an isopropenyl moiety [CH_3 -13 (δ 1.67, *s*) and 2H-12 (δ 4.98, *s* and 4.91, *d*, $J = 0.9\text{ Hz}$)], a methine proton (δ 3.38, *d*, H-10) and methylene protons (δ 2.89, *d*, H_β -9 and 2.49, *dd*, H_α -9). The quinonoid dihydroxanthone with isopropenyl side chain then was assigned for **AE13**. The oxirane ring was assignable by the oxycarbon resonances at δ 61.9 (C-1') and 72.0 (C-6'). HMBC correlation of H-3' at δ 3.57 to the carbon at δ 61.9 (C-1') and of 2H-9 at δ 2.49 and 2.89 to the carbon at δ 72.0 (C-6') confirmed the position of the oxirane ring. The proposed structure of **AE13** was in agreement with molecular ion of m/z 506.1572 ($C_{28}H_{26}O_9$). The HMBC (**Table 27**) and cosy correlations (**Table 28**) supported the assigned structure.

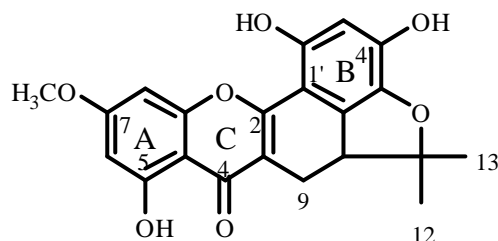
Table 27 ^{13}C , ^1H and HMBC spectral data of **AE13**

Position	δ_{C} (C-Type)	δ_{H} (<i>mult</i> , J_{Hz})	HMBC
2	155.2 (C)	-	-
3	117.3 (C)	-	-
4	181.4 (C=O)	-	-
4a	106.6 (C)	-	-
5	162.8 (C)	-	-
6	102.0 (CH)	6.28 (1H, <i>s</i>)	C-4a, C-5, C-7, C-8
7	161.3 (C)	-	-
8	103.1 (CH)	-	-
8a	153.4 (C)	-	-
9	24.3 (CH ₂)	2.49 (1H _{α} , <i>dd</i> , 17.1, 7.2) 2.89 (1H _{β} , <i>d</i> , 17.1)	C-2, C-3, C-4, C-6', C-10, C-11 C-2, C-3, C-4, C-6', C-10, C-11
10	41.1 (CH)	3.38 (1H, <i>d</i> , 7.2)	C-1', C-3, C-6', C-9, C-11, C-13
11	145.1 (C)	-	-
12	117.8 (CH ₂)	4.98 (1H _{β} , <i>s</i>) 4.91 (1H _{α} , <i>d</i> , 0.9)	C-10, C-11, C-13 C-10, C-13
13	22.5 (CH ₃)	1.67 (3H, <i>s</i>)	C-10, C-11, C-12
14	116.3 (CH)	6.78 (1H, <i>d</i> , 10.2)	C-7, C-8, C-8a, C-16
15	129.1 (CH)	5.60 (1H, <i>d</i> , 10.2)	C-8, C-16, C-17, C-18
16	79.9 (C)	-	-
17	29.9 (CH ₃)	1.49 (3H, <i>s</i>)	C-15, C-16, C-18
18	29.7 (CH ₃)	1.46 (3H, <i>s</i>)	C-15, C-16, C-17
19	41.7 (CH ₂)	3.25 (1H _{β} , <i>d</i> , 15.8) 3.03 (1H _{α} , <i>d</i> , 15.8)	C-3', C-4', C-5', C-20 C-4', C-5', C-20
20	213.3 (C=O)	-	-
21	33.3 (CH ₃)	2.43 (3H, <i>s</i>)	C-19, C-20
1'	61.9 (C)	-	-
2'	195.6 (C=O)	-	-
3'	50.9 (CH ₂)	3.57 (1H, <i>d</i> , 19.4) 3.47 (1H, <i>d</i> , 19.4)	C-1', C-2', C-5' C-2', C-4', C-5', C-19
4'	83.6 (C)	-	-
5'	198.7 (C=O)	-	-
6'	72.0 (C)	-	-
5-OH	-	12.42 (1H, <i>s</i>)	C-4, C-4a, C-5, C-6
4'-OH	-	5.64 (1H, <i>s</i>)	C-4', C-19

Table 28 ^1H - ^1H COSY spectral data of **AE13**

Proton (δ_{ppm})		Correlated proton (δ_{ppm})
H-3' (3.57)	↔	H-3' (3.47)
H $_{\alpha}$ -9 (2.49)	↔	H $_{\beta}$ -9 (2.89), H-10 (3.38)
H $_{\beta}$ -9 (2.89)	↔	H $_{\alpha}$ -9 (2.49), H-10 (3.38)
H-10 (3.38)	↔	H $_{\alpha}$ -9 (2.49), H $_{\beta}$ -9 (2.49)
H $_{\alpha}$ -12 (4.91)	↔	H $_{\beta}$ -12 (4.98), H-13 (1.67)
H $_{\beta}$ -12 (4.98)	↔	H $_{\alpha}$ -12 (4.91), H-13 (1.67)
H-13 (1.67)	↔	H $_{\alpha}$ -12 (4.91), H-12 $_{\beta}$ (4.98)
H-14 (6.78)	↔	H-15 (5.60)
H $_{\alpha}$ -19 (3.03)	↔	H $_{\beta}$ -19 (3.25)

Major HMBC of **AE13**

AE14**5a,6-Dihydro-1,3,8-trihydroxy-10-methoxy-5,5-dimethyl-5H,7H-benzofuro[3,4-*bc*]xanthen-7-one**

AE14 is a yellow solid, m.p. 287-288 °C. The UV spectrum exhibited absorption maxima at 225, 266, 270, 304 and 362 nm. The IR spectrum showed the absorption bands of hydroxyl groups at 3369 cm^{-1} and a carbonyl group at 1629 cm^{-1} . The ^1H NMR spectrum exhibited the signals of a hydrogen bonded hydroxyl group (δ 13.17, *s*, 5-OH), methoxyl protons (δ 3.88, *s*, 7-OCH₃), an isolated aromatic proton (δ 6.39, *s*, H-3') and two *meta* aromatic protons (δ 6.32, *d*, H-6 and δ 6.56, *d*, H-8). The position of *meta* protons H-6 and H-8 was confirmed by the HMBC correlations of H-6 to C-4a, C-8, and of H-8 to C-4a, C-6. The methoxyl group was assigned at C-7 by HMBC correlation of 7-OCH₃ and H-8 to C-7. The spectrum further showed the signals of two hydroxyl protons at δ 8.38 and δ 9.71 and they were located at C-2' and C-4', respectively due to HMBC correlations of 2'-OH to C-1', C-2',C-3' and of 4'-OH to C-3', C-4', C-5'. Two singlet signals of two methyl groups (δ 1.35, *s*, 12-CH₃ and δ 1.67, *s*, 13-CH₃) and an ABX system signals of a methine proton (δ 3.38, *dd*, H-10) and methylene protons (δ 3.22, *dd*, H _{β} -9 and 2.40, *t*, H _{β} -9) were assigned for the characteristic signals of a furanodihydrobenzo-xanthone skeleton. The HMBC correlations of H-10 to C-3, C-9, C-11, C-12, C-13, C-6' suggested that the cyclized prenyl moiety was located at C-3 position and was linked to ring B at C-6' position. The assignment of **AE14** was in agreement with the structure of **artoinin K** (Namdaung, *et al.*, 2006).

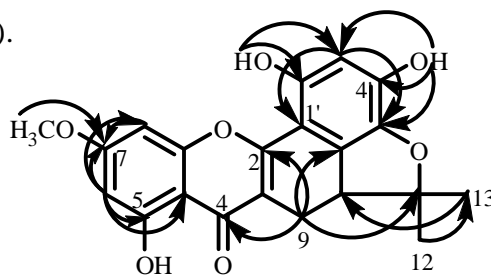
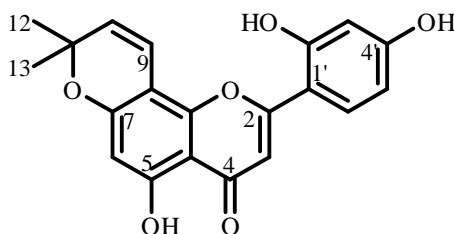
Major HMBC of **AE14**

Table 29 ^{13}C , ^1H and HMBC spectral data of **AE14**

Position	δ_{C} (C-Type)	δ_{H} (mult, J_{Hz})	HMBC
2	165.9 (C)	-	-
3	116.4 (C)	-	-
4	185.2 (C=O)	-	-
4a	110.5 (C)	-	-
5	166.5 (C)	-	-
6	102.8 (CH)	6.32 (1H, <i>d</i> , 2.1)	C-4a, C-5, C-7, C-8
7	169.6 (C)	-	-
8	97.1 (CH)	6.56 (1H, <i>d</i> , 2.1)	C-4a, C-6, C-7, C-8a
8a	161.3 (C)	-	-
9	24.6 (CH ₂)	3.22 (1H _{β} , <i>dd</i> , 15.0, 6.9) 2.40 (1H _{α} , <i>t</i> , 15.0)	C-2, C-3, C-4, C-6', C-10 C-3, C-6', C-10, C-11
10	51.4 (CH)	3.38 (1H, <i>dd</i> , 15.0, 6.9)	C-6', C-9, C-11, C-12, C-13
11	97.8 (C)	-	-
12	27.4 (CH ₃)	1.35 (3H, <i>s</i>)	C-10, C-11, C-13
13	32.9 (CH ₃)	1.67 (3H, <i>s</i>)	C-10, C-11, C-12
1'	108.2 (C)	-	-
2'	155.3 (C)	-	-
3'	109.6 (CH)	6.39 (1H, <i>s</i>)	C-1', C-2', C-4', C-5'
4'	151.5 (C)	-	-
5'	141.9 (C)	-	-
6'	136.6 (C)	-	-
7-OCH ₃	60.5 (OCH ₃)	3.88 (3H, <i>s</i>)	C-7
5-OH	-	13.17 (1H, <i>s</i>)	C-3', C-5, C-6
2'-OH	-	8.38 (1H, <i>s</i>)	C-1', C-2', C-3'
4'-OH	-	9.71 (1H, <i>s</i>)	C-3', C-4', C-5'

Table 30 ^1H - ^1H COSY spectral data of **AE14**

Proton (δ_{ppm})		Correlated proton (δ_{ppm})
H-6 (6.32)	\longleftrightarrow	H-8 (6.56)
H _{α} -9 (2.40)	\longleftrightarrow	H _{β} -9 (2.40), H-10 (3.38)
H _{β} -9 (3.22)	\longleftrightarrow	H _{α} -9 (2.40), H-10 (3.38)
H-10 (3.38)	\longleftrightarrow	H _{α} -9 (2.40), H _{β} -9 (3.22)

AE15**5-Hydroxy-8,8-dimethyl-2-(2,4-dihydroxyphenyl)-4*H*,8*H*-benzo[1,2-*b'*]dipyran-4-one**

AE15 is a pale yellow solid, m.p. 189-191 °C. The UV spectrum exhibited the absorption bands at 212, 230, 254, 272, 287, 307 and 357 nm. The IR spectrum showed the O-H stretching at 3379 cm^{-1} and the C=O stretching at 1651 and 1555 cm^{-1} . The ^1H NMR spectrum showed the characteristic resonances of a flavone proton at δ 7.15 (*s*, H-3), a hydrogen-bonded hydroxyl proton at δ 13.53 (*s*, 5-OH) and an aromatic proton at δ 6.44 (*s*, H-6). The resonances of two vinylic methine protons H-9 (δ 6.67, *d*, $J = 10.2$ Hz), H-10 (δ 5.74, *d*, $J = 10.2$ Hz), and two methyl groups 12- CH_3 (δ 1.47, *s*), 13- CH_3 (δ 1.47, *s*) revealed the presence of a 2,2-dimethylchromene ring. This moiety was placed at C-7 and C-8 of the parent structure due to HMBC correlations of H-9 to C-7, C-8a and of H-6 to C-5, C-7, C-8. A doublet at δ 6.58 ($J = 2.1$ Hz), a doublet of doublet at δ 6.54 ($J = 8.7, 2.1$ Hz) and a doublet δ 7.83 ($J = 8.7$ Hz) were in agreement with the ABX type of aromatic protons H-3', H-5' and H-6'. Thus **AE15** was assigned as 5-hydroxy-8,8-dimethyl-2-(2,4-dihydroxyphenyl)-4*H*,8*H*-benzo[1,2-*b'*]dipyran-4-one, a new flavone derivative.

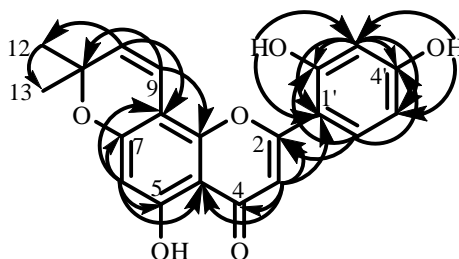
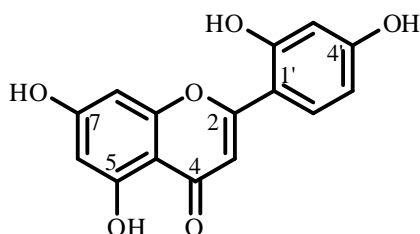
Major HMBC of **AE15**

Table 31 ^{13}C , ^1H and HMBC spectral data of **AE15**

Position	δ_{C} (C-Type)	δ_{H} (<i>mult</i> , J_{Hz})	HMBC
2	162.3 (C)	-	-
3	107.5 (CH)	7.15 (1H, <i>s</i>)	C-1', C-2, C-4, C-4a
4	182.8 (C=O)	-	-
4a	104.9 (C)	-	-
5	157.1 (C)	-	-
6	94.6 (CH)	6.44 (1H, <i>s</i>)	C-4a, C-5, C-7, C-8
7	159.1 (C)	-	-
8	104.9 (C)	-	-
8a	156.3 (C)	-	-
9	115.0 (CH)	6.67 (1H, <i>d</i> , 10.2)	C-7, C-8a, C-11
10	128.1 (CH)	5.74 (1H, <i>d</i> , 10.2)	C-8, C-11, C-12, C-13
11	78.3 (C)	-	-
12	27.6 (CH ₃)	1.47 (3H, <i>s</i>)	C-10, C-11, C-13
13	27.6 (CH ₃)	1.47 (3H, <i>s</i>)	C-10, C-11, C-12
1'	104.9 (C)	-	-
2'	158.8 (C)	-	-
3'	103.5 (CH)	6.58 (1H, <i>d</i> , 2.1)	C-2', C-4', C-5'
4'	161.9 (C)	-	-
5'	108.2 (CH)	6.54 (1H, <i>dd</i> , 8.7, 2.1)	C-2, C-2', C-4'
6'	129.9 (CH)	7.83 (1H, <i>d</i> , 8.7)	C-2, C-2', C-4'
5-OH	-	13.53 (1H, <i>s</i>)	C-4a, C-5

Table 32 ^1H - ^1H COSY spectral data of **AE15**

Proton (δ_{ppm})		Correlated proton (δ_{ppm})
H-9 (6.67)	↔	H-10 (5.74)
H-3' (6.58)	↔	H-5' (6.54)
H-5' (6.54)	↔	H-3' (6.58), H-6' (7.83)
H-6' (7.83)	↔	H-5' (6.54)

AE16**2-(2,4-Dihydroxyphenyl)-5,7-dihydroxy-4H-chromen-4-one**

AE16 is a pale creamy solid. The ^1H NMR spectrum showed the characteristic resonances of a flavone proton at δ 6.96 (s, H-3), a hydrogen-bonded hydroxyl proton at δ 13.00 (s, 5-OH), three free hydroxyl protons at δ 9.95, 9.90, 9.42, and meta aromatic protons at δ 6.36 (*d*, $J = 2.5$ Hz) and 6.10 (*d*, $J = 2.5$ Hz). The signals of doublet at δ 6.48 ($J = 2.5$ Hz), a doublet of doublet at δ 6.42 ($J = 8.5, 2.5$ Hz) and a doublet at δ 7.70 ($J = 8.5$ Hz) were in agreement with the ABX type of aromatic protons H-3', H-5' and H-6'. Thus **AE16** was assigned as 2-(2,4-dihydroxyphenyl)-5,7-dihydroxy-4H-chromen-4-one which was corresponded to norartocarpetin (Amarasinghe, *et al.*, 2008).

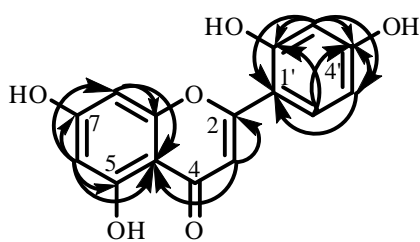
Major HMBC of **AE16**

Table 33 ^{13}C , ^1H and HMBC spectral data of **AE16**

Position	δ_{C} (C-Type)	δ_{H} (<i>mult</i> , J_{Hz})	HMBC
2	158.5 (C)	-	-
3	108.2 (CH)	6.96 (1H, <i>s</i>)	C-2, C-4a
4	182.6 (C=O)	-	-
4°	103.5 (C)	-	-
5	162.0 (C)	-	-
6	93.6 (CH)	6.36 (1H, <i>d</i> , 2.5)	C-4a, C-5, C-7, C-8
7	164.0 (C)	-	-
8	98.5 (C)	6.10 (1H, <i>d</i> , 2.5)	C-4a, C-6, C-8a
8°	162.4 (C)	-	-
1'	107.6 (C)	-	-
2'	161.8 (C)	-	-
3'	103.4 (CH)	6.48 (1H, <i>d</i> , 2.5)	C-1', C-2', C-4', C-5'
4'	158.0 (C)	-	-
5'	107.7 (CH)	6.42 (1H, <i>dd</i> , 8.5, 2.5)	C-1', C-3', C-4'
6'	129.9 (CH)	7.70 (1H, <i>d</i> , 8.5)	C-2', C-4'
5-OH	-	13.00 (1H, <i>s</i>)	C-4a, C-5, C-6
*OH	-	9.42 (<i>s</i>)	-
*OH	-	9.90 (<i>s</i>)	-
*OH	-	9.95 (<i>s</i>)	-

* the position not identified

Table 34 ^1H - ^1H COSY spectral data of **AE 16**

Proton (δ_{ppm})		Correlated proton (δ_{ppm})
H-6 (6.36)	↔	H-8 (6.10)
H-3' (6.48)	↔	H-5' (6.42)
H-5' (6.42)	↔	H-3' (6.48), H-6' (7.70)
H-6' (7.70)	↔	H-5' (6.42)

3.2 Evaluation of biological activities of the crude extracts

3.2.1 Antimicrobial activity

Dried bark of *A. elasticus* was extracted with dichloromethane and acetone to give dichloromethane extract (AeD) and acetone extract (AeA). Each extract was screened for antibacterial activity against *Staphylococcus aureus* ATCC25923 (SA), methicillin-resistant *Staphylococcus aureus* (MRSA SK1), *Pseudomonas aeruginosa* ATCC27853 (PA), and *Escherichia coli* ATCC25922, for antiyeast activity on *Candida albicans* NCPF3153 (CA), and *Cryptococcus neoformans* ATCC90113 flucytosine-resistant (CN90113), and for antifungal activity on *Microsporium gypseum* clinical isolate (M. gypseum). It was found that the CH₂Cl₂ extract showed activity with MIC 128 µg/ml for SA and MRSA SK1. The acetone extract exhibited activity with MIC 16 µg/ml. Both extracts showed no activity for PA and EC. Both extracts showed no antiyeast activity and antifungal activity. The results are shown in **Table 35** and **Table 36**

Table 35 Antibacterial activity of crude extracts from the bark of *A. elasticus*

Fractions	Antibacterial activity (MIC, µg/ml)			
	SA	MRSA SK1	PA	EC
AeD	128	128	>200	>200
AeA	16	16	>200	>200
Vancomycin	1	1	-	-
Gentamicin	-	-	1	1

Table 36 Antiyeast and Antifungal activity of crude extracts from the bark of *A. elasticus*

Fractions	Antiyeast activity (MIC, $\mu\text{g/ml}$)		Antifungal activity (MIC, $\mu\text{g/ml}$)
	CA	CN90113	<i>M. gypseum</i>
AeD	>200	>200	>200
AeA	>200	>200	>200
Amphotericin B	0.125	0.25	-
Miconazole	-	-	1

Some of the pure compounds obtained from CH_2Cl_2 extract were evaluated for their antibacterial activities against *S. aureus* ATCC25923, and MRSA SK1. The result (**Table 37**) indicated that **AE4**, **AE8**, **AE13** and **AE14** were less active than the crude extract. Whereas **AE1**, **AE10** and **AE12** were more active than the crude extract. Among the active compounds **AE1** showed the strongest inhibitory activity with a MIC value of 4 and 8 $\mu\text{g/ml}$ against *S. aureus* ATCC25923, and MRSA SK1, respectively. However it was less active than vancomycin, the standard antibiotic (MIC 1 $\mu\text{g/ml}$). **AE2**, **AE3**, **AE5**, **AE6**, **AE7**, **AE9**, **AE11**, **AE15** and **AE16** were not tested due to insufficient quantities.

Table 37 Antibacterial activity of compounds isolated from the bark of *A. elasticus*

Compound	Antibacterial activity (MIC, $\mu\text{g/ml}$)	
	SA	MRSA SK1
AE1	4	8
AE4	> 200	> 200
AE8	> 200	> 200
AE10	16	16
AE12	32	16
AE13	128	-
AE14	200	200
Vancomycin	1	1

3.2.2 Cytotoxic activity

The dichloromethane extract (AeD) and acetone extract (AeA) were further evaluated for cytotoxicity against C6 (Glial tumor), D17 (Bone cancer), OLO (Colon cancer) and PC3 Prostate cancer. According to the MIC value shown in **Table 37**, acetone extract was found to inhibit cancer cell lines with IC_{50} in the range of 300-337.5 $\mu\text{g/ml}$, whereas dichloromethane extract was found to be inactive for cytotoxic activity.

Table 38 Cytotoxic activity of crude extracts from the bark of *A. elasticus*

Fractions	Cytotoxic activity (IC_{50} $\mu\text{g/ml}$)			
	C6	D17	OLO 205	PC3
	Glial tumor	Bone cancer	Colon cancer	Prostate cancer
AeD	NA	NA	NA	NA
AeA	337.5	315	~300	NA

NA = no activity

3.3 Review of biological activities of the known compounds obtained from this study

Biological activities of some compounds isolated from this study have been previously investigated. Based on the search from SciFinder Scholar the biological activities of artonin E (**AE1**), artnol B (**AE5**), 7-demethylartanol E (**AE10**), artonin F (**AE8**), cycloartobiloxanthone (**AE10**) and norartocarpetin (**AE16**) are summarized.

Artonin E (**AE1**) is the major component of *A. elasticus*. It showed strong radical scavenging properties (DPPH radical) (Jayasinghe, *et al.*, 2008) and cytotoxicity to cell lines such as murine leukemia P388 cell line (IC_{50} 0.06 $\mu\text{g/ml}$) (Hakim, *et al.*, 2006), 1A9 (Ovarian, ED_{50} 1.25 $\mu\text{g/ml}$), MCF-7 (breast

adenocarcinoma, ED₅₀ 2.2 µg/ml), HCT-8 (ileocecal, ED₅₀ 3.3 µg/ml) and MDAMB-231 (breast adenocarcinoma, ED₅₀ 3.0 µg/ml) (Wang, *et al.*, 2004). Furthermore, it also reduced the amount of urinary protein excretion compared to nephritic mice (Fukai, *et al.*, 2003).

Artonol B (**AE5**) has been reported to show cytotoxicity against the human small cell lung cancer (NCI-H 187, IC₅₀ 1.26 µg/ml) (Namdaung, *et al.*, 2006) and murine leukemia cell (P388, IC₅₀ >100 µg/ml) (Hakim, *et al.*, 2002).

7-Demethylartonol E (**AE6**) has been tested for antiplasmodial activity against *Plasmodium falciparum* (IC₅₀ 7.9 µg/ml) and antimycobacterial activity against *Mycobacterium tuberculosis* (MIC 50 µg/ml). It was also toxic to human small cell lung cancer (NCI-H187, IC₅₀ 5.7 µg/ml) (Namdaung, *et al.*, 2006).

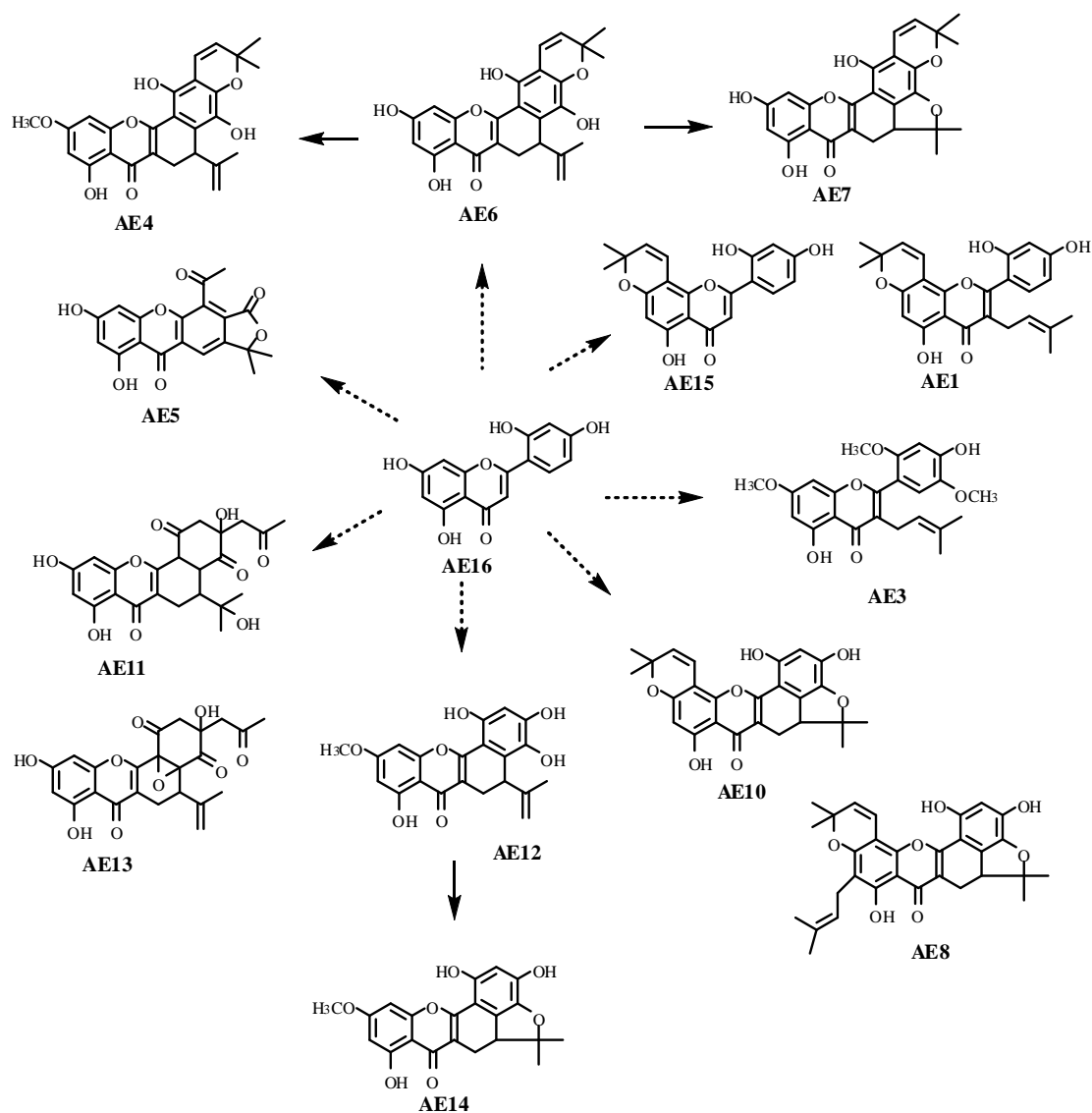
Artonin F (**AE8**) has been tested for antiplasmodial activity against *Plasmodium falciparum* (IC₅₀ 2.4 µg/ml) and antimycobacterial activity against *Mycobacterium tuberculosis* (MIC 6.25 µg/ml) (Namdaung, *et al.*, 2006).

Cycloartobiloxanthone (**AE10**) has been tested for antiplasmodial activity against *Plasmodium falciparum* (IC₅₀ 3.7 µg/ml) and antimycobacterial activity against *Mycobacterium tuberculosis* (MIC 25 µg/ml) (Namdaung, *et al.*, 2006). The radical scavenging properties (DPPH radical) were studied (Jayasinghe, *et al.*, 2008). It also exhibited cytotoxicity against human epidermoid carcinoma of the nasopharynx (KB, IC₅₀ 8.56 µg/ml), human breast cancer (BC, IC₅₀ 4.23 µg/ml) and human small cell lung cancer (NCI-H187, IC₅₀ 11.83 µg/ml).

Norartocarpetin (**AE16**) has been reported to exhibit strong toxicity against *Artemia salina* shrimp, LC₅₀ 0.05 µg/ml.

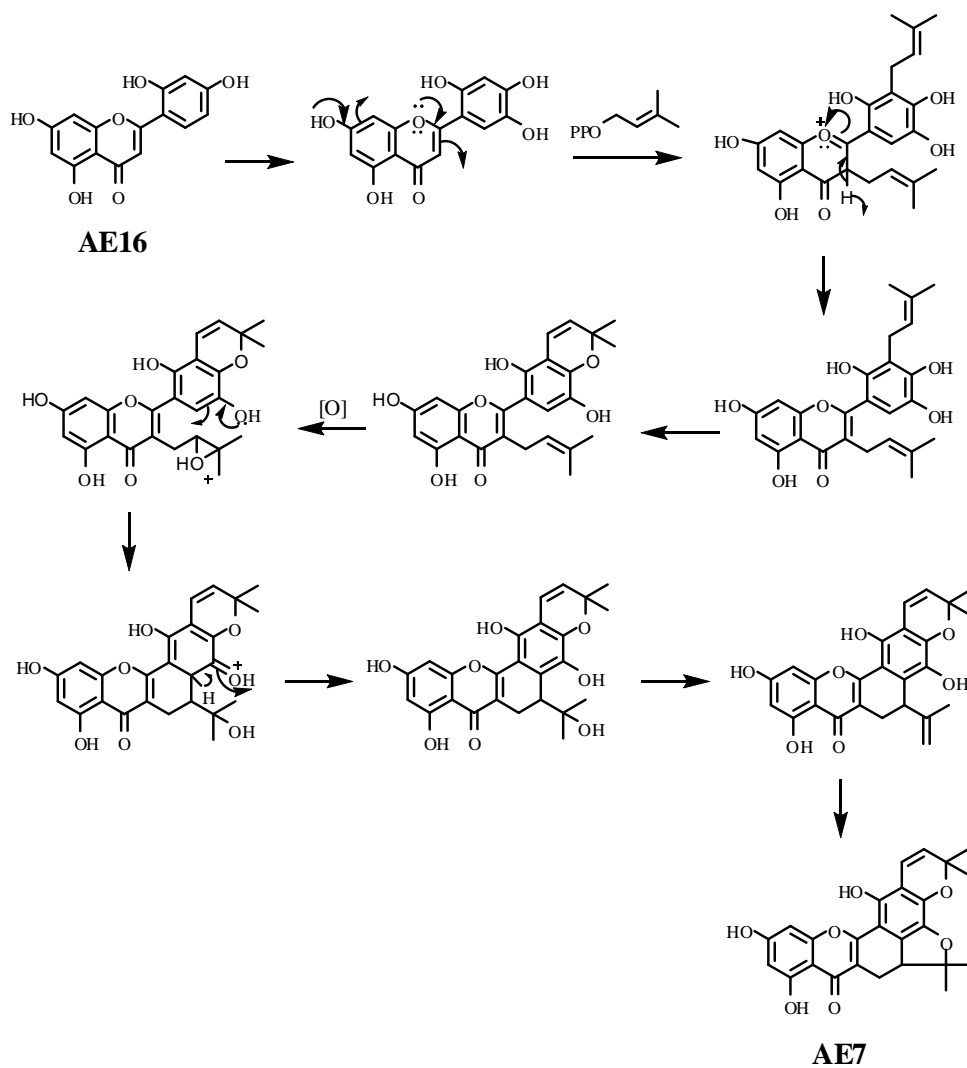
3.4 Biogenesis of flavonoids and related compounds from *Artocarpus elasticus*

Relationships between the various classes of secondary metabolites from *Artocarpus elasticus* is outlined in **Scheme 4**.



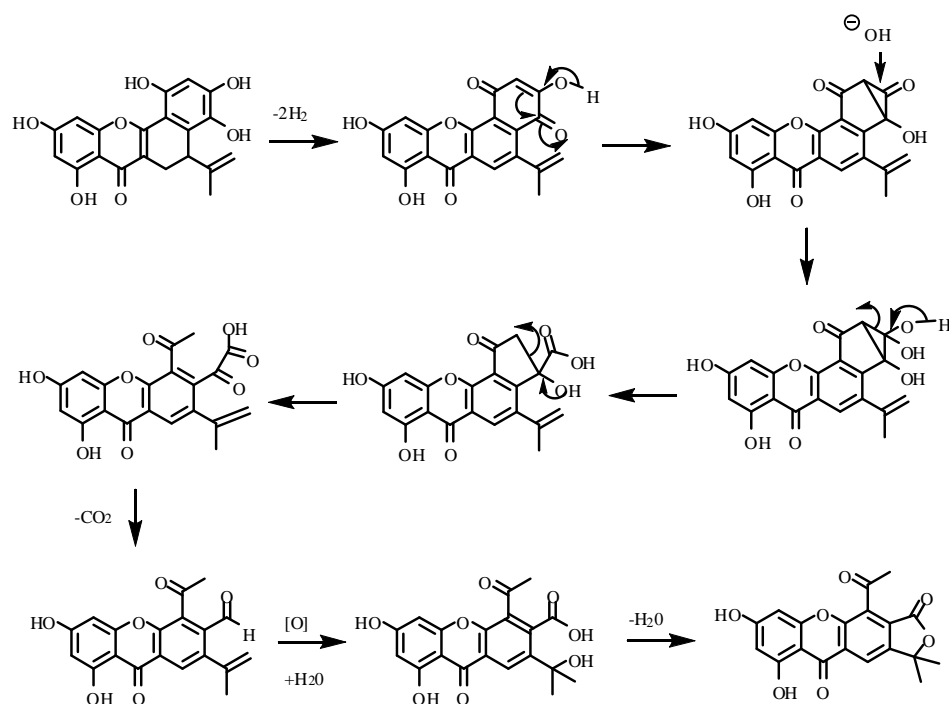
Scheme 4 Relationships of prenylated flavonoids from *A. elasticus*

Biosynthetic route for the formation of **AE7** (furanodihydrobenzoxan-
thone skeleton), a new compound, from simple flavones is proposed in **Scheme 5**
(Sultanbawa, *et al.*, 1989).



Scheme 5 Proposed biogenetic route of **AE7**

Biogenetic route of artonol B (**AE5**) (xanthonolide skeleton) has been describe by Aida, *et al.*, 1997 as shown in **scheme 6**.



Scheme 6 Proposed biogenetic route of the xanthonolide types of compounds

Conclusion

Investigation of the constituents from the bark of *A. elasticus* led to the isolation of sixteen compounds including four flavone derivatives: 5-hydroxy-8,8-dimethyl-3-(3-methyl-2-butenyl)-2-(2,4,5-trihydroxyphenyl)-4*H*,8*H*-benzo[1,2-*b*:3,4-*b'*]dipyrans-4-one (**AE1**), 5-hydroxy-2-(4-hydroxy-2,5-dimethoxyphenyl)-7-methoxy-3-(3-methylbut-2-enyl)-4*H*-chromen-4-one (**AE3**), 5-hydroxy-8,8-dimethyl-2-(2,4-dihydroxyphenyl)-4*H*,8*H*-benzo[1,2-*b'*]dipyrans-4-one (**AE15**), and 2-(2,4-dihydroxyphenyl)-5,7-dihydroxy-4*H*-chromen-4-one (**AE16**), three dihydrobenzoxanthone derivatives: 6,7-dihydro-5,9,14-trihydroxy-11-methoxy-3,3-dimethyl-6-(1-methylethyl)-3*H*,8*H*-[1]benzopyrano[7,6-*c*]xanthen-8-one (**AE4**), 6,7-dihydro-5,9,11,14-tetrahydroxy-3,3-dimethyl-6-(1-methylethenyl)-(-)-3*H*,8*H*-pyrano[3',2':4,5]benzo[1,2-*c*]xanthen-8-one (**AE6**) and 1,3,4,8-tetrahydroxy-10-methoxy-5-(prop-1-en-2-yl)-5*H*-benzo[*c*]xanthen-7-(6*H*)-one (**AE12**), four furanodihydrobenzoxanthone derivatives: (**AE7**), 5*a*,6-dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-9-(3-methyl-2-buten-1-yl)-5*H*,7*H*,11*H*-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one (**AE8**), 5*a*,6-dihydro-1,3,8-trihydroxy-5,5,11,11-tetramethyl-5*H*,7*H*,11*H*-benzofuro[3,4-*bc*]pyrano[3,2-*h*]xanthen-7-one (**AE10**) and 5*a*,6-dihydro-1,3,8-trihydroxy-10-methoxy-5,5-dimethyl-5*H*,7*H*-benzofuro[3,4-*bc*]xanthen-7-one (**AE14**), two quinonoxanthone derivatives: (**AE11**) and (**AE13**), one xanthonolide: 12-acetyl-6-hydroxy-3,3,9,9-tetramethyl-3*H*,7*H*,furo[3,4-*b*]pyrano[3,2-*h*]xanthen-7,11(9*H*)-dione (**AE5**), one dihydroisocoumarin: 8-hydroxy-3-methyl-isochroman-1-one (**AE2**), and one benzyl alcohol derivative: (3,4,5-trimethoxy phenyl) methanol (**AE9**). **AE3**, **AE7**, **AE11**, **AE12**, **AE13** and **AE15** are new compounds. **AE1**, **AE2**, **AE4**, **AE5**, **AE8**, **AE9**, **AE10**, **AE14** and **AE16** were obtained for the first time from this plant. **AE6** were previously isolated from this plant. **AE1** showed the best activity to inhibit the growth of *S. aureus* ATCC25923 and MRSA SK1 with a MIC value of 4 and 8 $\mu\text{g/ml}$. Further study on the biological activity of the isolated compound should be performed.

REFERENCES

- Achmad, S. A.; Hakim, E. H.; Juliawaty, L. D.; Makmur, L.; Suyatno; Aimi, N.; Ghisalberti, E. L. 1996. "A New Prenylated Flavone from *Artocarpus champeden*", *J. Nat. Prod.*, 59, 878-879.
- Aida, M.; Shinomiya, K.; Hano, Y.; Nomura, T. 1993. "Artonin J, K, and L, Three New Isoprenylated Flavones from the Root Bark of *Artocarpus heterophyllus* Lamk.", *Heterocycles*, 36 (3), 575-583.
- Aida, M.; Yamaguchi, N.; Hano, Y.; Nomura, T. 1997. "Artonols A, B, C, D, and E, Five New Isoprenylated Phenols from the Bark of *Artocarpus communis* Forst.", *Heterocycles*, 45 (1), 163-175.
- Amarasinghe, N. R.; Jayasinghe, L.; Hara, N.; Fujimoto, Y. 2008. "Chemical Constituents of the Fruits of *Artocarpus altilis*", *Biochemical Systematics and Ecology*, 36, 323-325.
- Arung, E. T.; Shimizu, K.; Kondo, R. 2006. "Inhibitory Effect of Artocarpanone from *Artocarpus heterophyllus* on Melanin Biosynthesis", *Biol. Pharm. Bull.*, 29 (9), 1966-1969.
- Boonlaksiri, C.; Oonant, W.; Kongsaree, P.; Kittakoop, P.; Tanticharoen, M.; Thebtaranonth, Y. 2000. "An Antimalarial Stilbene from *Artocarpus integer*", *Phytochemistry*, 54, 415-417.
- Cerqueira, F.; Cidade, H.; Ufford, L. V.; Beukelman, C.; Kijjoa, A.; Nascimento, M. S. J. 2008. "The Natural Prenylated Flavone Artelastin is an Inhibitor of ROS and NO production", *International Immunopharmacology*, 8, 597-602.

- Cerqueira, F.; Cordeiro-Da-Silva, A.; Araujo, N.; Cidade, H.; Kijjoa, A.; Nascimento. 2003. "Inhibition of Lymphocyte Proliferation by Prenylated Flavones: Artelastin as a Potent Inhibitor", *Life Sciences*, 73, 2321-2334.
- Chan, S.- C.; Ko, H.-H.; Lin, C.- N. 2003. "New Prenylflavonoids from *Artocarpus communis*", *J. Nat. Prod.*, 66, 427-430.
- Chantrapromma, S.; Boonnak, N.; Fun, H.-K.; Karalai, C. 2007. "Artocarpin dichloromethane hemisolvate", *Acta Cryst.*, E63, o1864-o1866.
- Chung, M.- I.; Lu, C.- M.; Huang, P.-L.; Lin, C.- N. 1995. "Prenylflavonoids of *Artocarpus heterophyllus*", *Phytochemistry*, 40 (4), 1279-1282.
- Dimitriadis, C.; Gill, M.; Harte, M. F. 1997. "The First Stereospecific Approach to Both Enantiomers of Mullein", *Tetrahedron: Asymmetry*, 8 (13), 2153-2158.
- Fukai, T.; Satoh, K.; Nomura, T.; Sakagami, H. 2003. "Antinephritis and Radical Scavenging Activity of Prenylflavonoids", *Fitoterapia*, 74, 720-724.
- Fun, H.-K., Boonnak, N.; Chantrapromma, S. 2007. "A Cocrystal of Friedelan-3 β -ol and Friedelin", *Acta cryst.*, E63, o2014-o2016.
- Hakim, E. H.; Achmad, S. A.; Juliawaty, L. D.; Makmur, L.; Syah, Y. M.; Aimi, N.; Kitajima, M.; Takayama, H.; Ghisalberti, E. L. 2006. "Prenylated Flavonoids and Related Compounds of the Indonesian *Artocarpus* (Moraceae)", *J. Nat. Med.*, 60, 161-184.
- Hakim, E. H.; Asnizar; Yurnawilis; Aimi, N.; Kitajima, M.; Takayama, H. 2002. "Artoindonesianin P, a New Prenylated Flavone with Cytotoxic Activity from *Artocarpus lanceifolius*", *Fitoterapia*, 73, 668-673.

- Hakim, E. H.; Fahriyati, A.; Kau, M. S.; Achmad, S. A.; Makmur, L.; Ghisalberti, E. L.; Nomura, T. 1999. "Artoindonesianins A and B, Two New Prenylated Flavones from the Root of *Artocarpus champeden*", *J. Nat. Prod.*, 62, 613-615.
- Hakim, H. H.; Juliawaty, L. D.; Syah, Y. M.; Achmad, A. S. 2005. "Molecular Diversity of *Artocarpus champeden* (Moraceae): A species Endemic to Indonesia", *Molecular Diversity*, 9, 149-158.
- Hakim, E. H.; Ulinnuha, U. Z.; Syah, Y. M.; Ghisalberti, E. L. 2002. "Artoindonesianins N and O, New Prenylated Stilbene and Prenylated Arylbenzofuran Derivatives from *Artocarpus gomezianus*", *Fitoterapia*, 73, 597-603.
- Hano, Y.; Inami, R.; Nomura, T. 1993. "Components of the Bark of *Artocarpus rigida* BL. 2.^{*1} Structures of Four New Isoprenylated Flavone Derivatives Artonins M, N, O, and P", *Heterocycles*, 35 (2), 1341-1350.
- Hano, Y.; Yamagami, Y.; Miki, M.; Isohata, R.; Nomura, T. 1990. "Artonins E and F, Two New Prenylflavones from the Bark of *Artocarpus communis* Forst.", *Heterocycles*, 31 (5), 877-882.
- Han, A.-R.; Kang, Y.-J.; Windono, T.; Lee, S. K.; Seo, E.-K. 2006. "Prenylated Flavonoids from the Heartwood of *Artocarpus communis* with Inhibitory Activity on Lipopolysaccharide-Induced Nitric Oxide Production", *J. Nat. Prod.* 69, 719-721.
- Jayasinghe, L.; Balasooriya, B. A. I. S.; Padmini, W. C.; Hara, N.; Fujimoto, Y. 2004. "Geranyl Chalcone Derivatives with Antifungal and Radical Scavenging Properties from the Leaves of *Artocarpus nobilis*", *Phytochemistry*, 65, 1287-1290.

- Jayasinghe, L.; Rupasinghe, G. K.; Hara, N.; Fujimoto, Y. 2006. "Geranylated Phenolic Constituents from the Fruits of *Artocarpus nobilis*", *Phytochemistry*, 67, 1353-1358.
- Jayasinghe, U. L. B.; Samarakoon, T. B.; Kumarihamy, B. M. M.; Hara, N.; Fujimoto, Y. 2008. "Four New Prenylated Flavonoids and Xanthenes from the Root Bark of *Artocarpus nobilis*", *Fitoterapia*, 79, 37-41.
- Kijjoa, A.; Cidade, H. M.; Gonzalez, M. J. T. G.; Afonso, C. M.; Silva, A. M. S.; Herz, W. 1998. "Further Prenylflavonoids from *Artocarpus elasticus*", *Phytochemistry*, 47 (5), 875-878.
- Kijjoa, A.; Cidade, H. M.; Pinto, M. M. M.; Gonzalez, M. J. T. G.; Anantachoke, C.; Gedris, T. E.; Herz, W. 1996. "Prenylflavonoids from *Artocarpus elasticus*", *Phytochemistry*, 43 (3), 691-694.
- Ko, H.- H.; Lu, Y.- H.; Yang, S.- Z.; Won, S.- J.; Lin, C.- N. 2005. "Cytotoxic Prenylflavonoids from *Artocarpus elasticus*", *J. Nat. Prod.*, 68, 1692-1695.
- Likhitwitayawuid, K.; Sritularak, B. 2001. "A New Dimeric Stilbene with Tyrosinase Inhibitory Activity from *Artocarpus gomezianus*", *J. Nat. Prod.*, 64, 1457-1459.
- Likhitwitayawuid, K.; Supudompol, B.; Sritularak, B.; Lipipun, V.; Rapp, K.; Schinazi, R. F. 2005. "Phenolics with Anti-HSV and Anti-HIV Activities from *Artocarpus gomezianus*, *Mallotus pallidus*, and *Triphhasia trifolia*", *Pharmaceutical Biology*, 43 (8), 651-657.
- Lin, C.- N.; Lu, C.- M.; Huang, L.- P. 1995. "Flavonoids from *Artocarpus heterophyllus*", *Phytochemistry*, 39 (6), 1447-1451.

- Lin, C.- N.; Chiu, P.- H.; Fang, S.- C.; Shieh, B.- J.; Wu, R.- R. 1996. "Revised Structure of Brousoflavonol G and the 2D NMR Spectra of Some Related Prenylflavonoids", *Phytochemistry*, 41 (4), 1215-1217.
- Lin, C.- N.; Shieh, W.- L. 1991. "Prenylflavonoids and Pyranodihydrobenzoxanthone from *Artocarpus communis*", *Phytochemistry*, 30 (5), 1669-1671.
- Lin, C.- N.; Shieh, W.- L. 1992. "Pyranoflavonoids from *Artocarpus communis*", *Phytochemistry*, 31 (8), 2922-2924.
- Lu, C.- M.; Lin, C.- N. 1993. "Two 2',4',6'-Trioxxygenated Flavanones from *Artocarpus heterophyllus*", *Phytochemistry*, 33 (4), 909-911.
- Namduang, U.; Aroonberk, N.; Suksamrarn, S.; Danwiesetkanjana, K.; Saenboonrueng, J.; Arjchomphu, W.; Suksamrarn, A. 2006. "Bioactive Constituents of the Root of *Artocarpus rigidus* subsp. *Rigidus*", *Chem. Pharm. Bull.*, 54 (10), 1433-1436.
- Parenti, P.; Pizzigoni, A.; Hanozet, G.; Hakim, E.- H.; Makmur, L.; Achmad, S.- A.; Giordana, B. 1998. "A New Prenylated Flavone from *Artocarpus champeden* Inhibits the K⁺-Dependent Amino Acid Transport in *Bombyx mori* Midgut", *Biochemical and Biophysical Research communications*, 244, 445-448.
- Patil, A. D.; Freyer, A. J.; Killmer, L.; Offen, P.; Taylor, P. B.; Votta, B. J.; Johnson, R. K. 2002. "A New Dimeric Dihydrochalcone and New Prenylated Flavone from the Bud Covers of *Artocarpus altilis*: Potent Inhibitors of Cathepsin K", *J. Nat. Prod.*, 65, 624-627.
- Pedro, M.; Ferreira, M. M.; Cidade, H.; Kijjoa, A.; Bronze-da-Rocha, E.; Nascimento, M. S. J. 2005. "Artelastin is a Cytotoxic Prenylated Flavone that Disturbs Microtubules and Interferes with DNA replication in MCF-7 human breast cancer cells", *Life Sciences*, 77, 293-311.

- Puntunchai, A.; Kittakoop, P.; Rajviroongit, S.; Vimuttipong, S.; Likhitwitayawuid, K.; Thebtaranonth, Y. 2004. "Lakoochins A and B, New Antimycobacterial Stilbene Derivatives from *Artocarpus lakoocha*", *J. Nat. Prod.*, 67, 485-486.
- Shieh, W.- L.; Lin, C.- N. 1992. "A Quinonoid Pyranobenzoxanthone and Pyranodihydrobenzoxanthone from *Artocarpus communis*", *Phytochemistry*, 31(1), 364-367.
- Shimizu, K.; Kondo, R.; Sakai, K. 1997. "A Stilbene Derivative from *Artocarpus incisus*", *Phytochemistry*, 45 (6), 1297-1298.
- Shimizu, K.; Kondo, R.; Sakai, K.; Buabarn, S.; Dilokkunanant, U. 2000. "A Geranylated Chalcone with 5 α -Reductase Inhibitory Properties from *Artocarpus incisus*", *Phytochemistry*, 54, 737-739.
- Sultanbawa, M. U. S.; Surendrakumar, S. 1989. "Two Pyranodihydrobenzoxanthones from *Artocarpus nobilis*", *Phytochemistry*, 28 (2), 599-605.
- Syah, Y. M.; Achmad, S. A.; Ghisalberti, E. L.; Hakim, E. H.; Makmur, L.; Mujahidin, D. 2001. "Artoindonesianins G-I, Three New Isoprenylated Flavones from *Artocarpus lanceifolius*", *Fitoterapia*, 72, 765-773.
- Syah, Y. M.; Achmad, S. A.; Ghisalberti, E. L.; Hakim, E. H.; Makmur, L.; Mujahidin, D. 2002. "Artoindonesianins Q-T, Four Isoprenylated Flavones from *Artocarpus champeden* Spreng. (Moraceae)", *Phytochemistry*, 61, 949-953.
- Syah, Y. M.; Achmad, S. A.; Ghisalberti, E. L.; Hakim, E. H.; Mujahidin, D. 2002. "Two New Cytotoxic Isoprenylated Flavones, Artoindonesianins U and V, From the Heartwood of *Artocarpus champeden*", *Fitoterapia*, 75, 134-140.

- Syah, Y. M.; Juliawaty, L. D.; Achmad, S. A.; Hakim, E. H.; Ghisalberti, E. L. 2006. "Cytotoxic Prenylated Flavones from *Artocarpus champeden*", *J. Nat. Med.*, 60, 308-312.
- Wang, Y.- H.; Hou, A.- J.; Chen, L.; Chen, D.- F.; Sun, H.- D.; Zhao, Q.- S.; Bastow, K. F.; Yuka, N.; Wang, X.- H.; Lee, K.-H. 2004. "New Isoprenylated Flavones, Artochamins A-E, and Cytotoxic Principles from *Artocarpus chama*", *J. Nat. Prod.*, 67, 757-761.
- Wang, Y.; Xu, K.; Lin, L.; Pan, Y.; Zheng, X. 2007. "Geranyl Flavonoids from the Leaves of *Artocarpus altilis*", *Phytochemistry*, 68, 1300-1306.
- Wei, B. - L.; Weng, J.- R.; Chiu, P. - H.; Hung, C. - F.; Wang, J. - P.; Lin, C. - N. 2005. "Antiinflammatory Flavonoids from *Artocarpus heterophyllus* and *Artocarpus communis*", *J. Agric. Food Chem.*, 53, 3867-3871.
- Widyawaruyanti, A.; Subehan; Surya, K. K.; Awale, S.; Nindatu, M.; Zaini, N.- C.; Syafruddin, D.; Asih, P. B. S.; Tezuka, Y.; Kadota, S. 2007. "New Prenylated Flavones from *Artocarpus champeden*, and Their Antimalarial Activity in Vitro", *J. Nat. Med.*, 61, 410-413.

APPENDIX

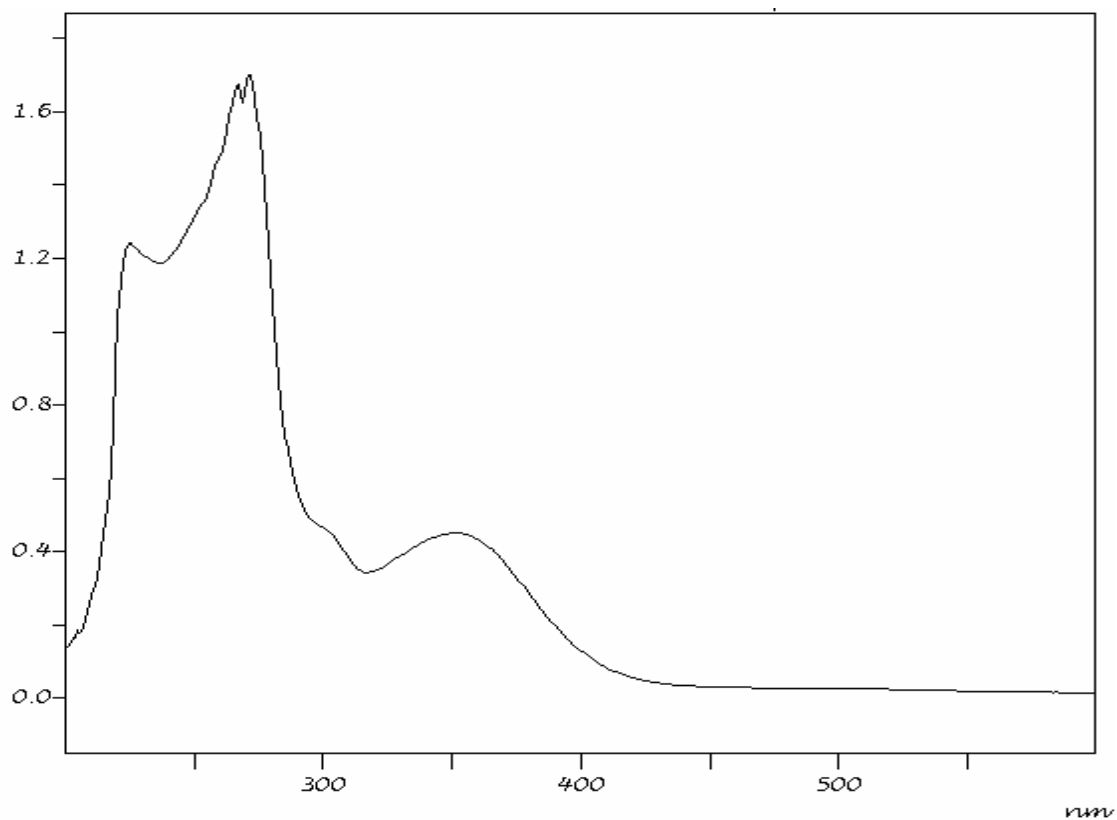


Figure 6 UV (CH₃OH) spectrum of AE1

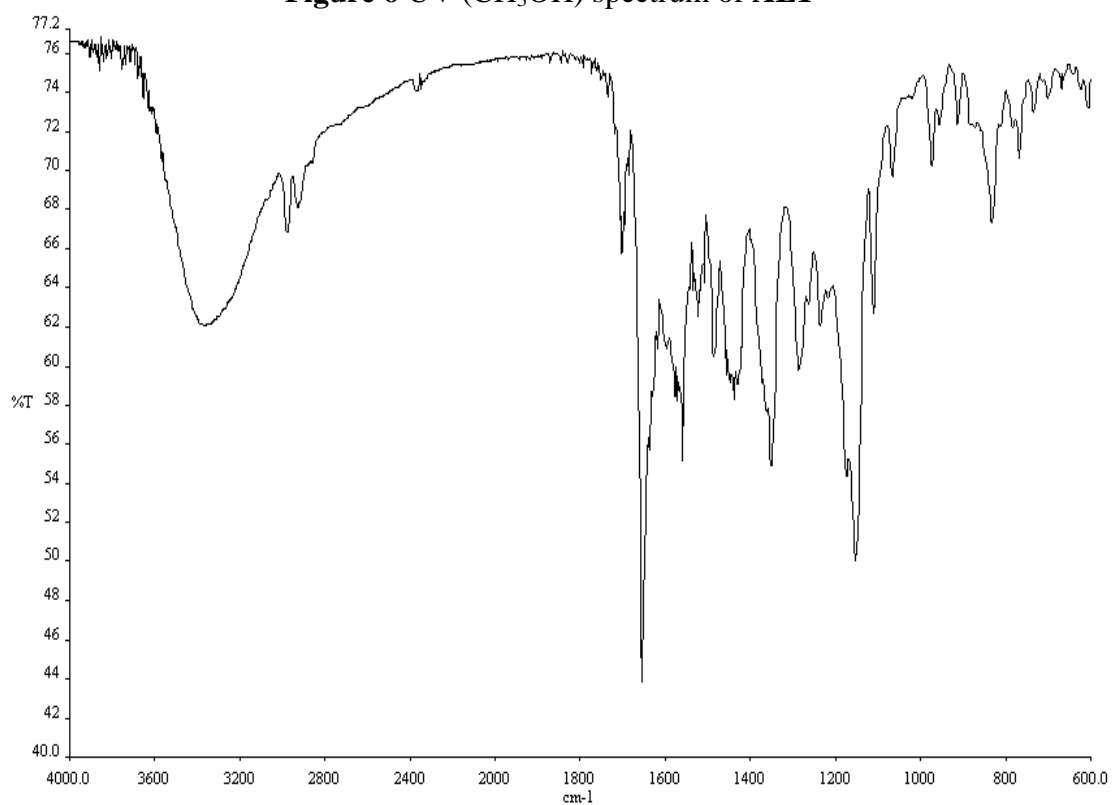


Figure 7 FT-IR (Neat) spectrum of AE1

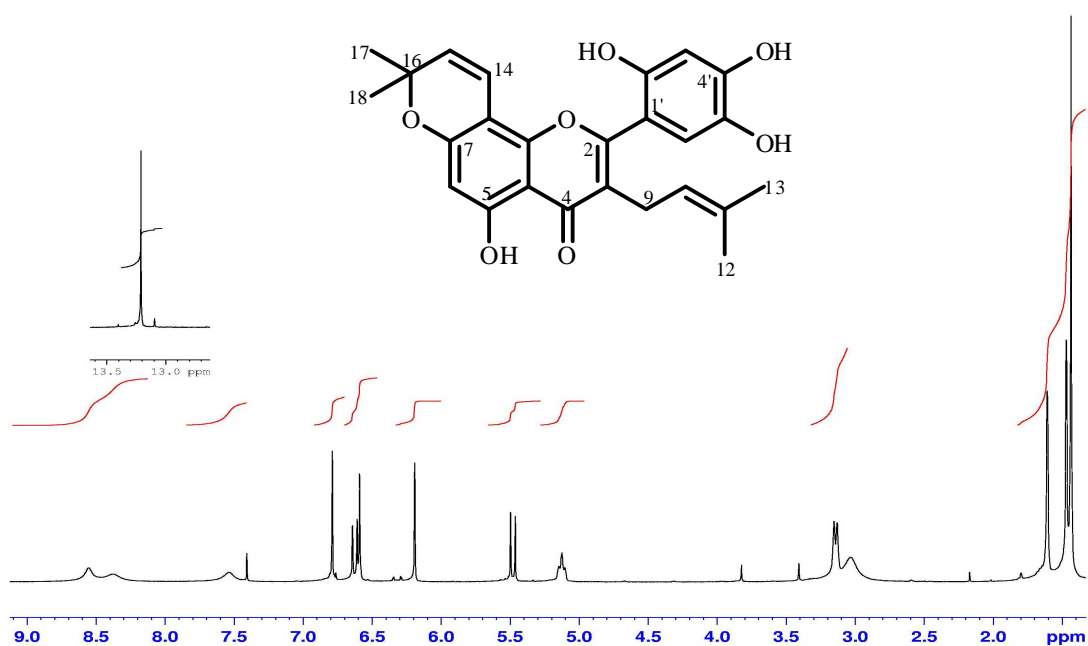


Figure 8 ^1H NMR (300 MHz) ($\text{CDCl}_3 + \text{DMSO-}d_6$) spectrum of AE1

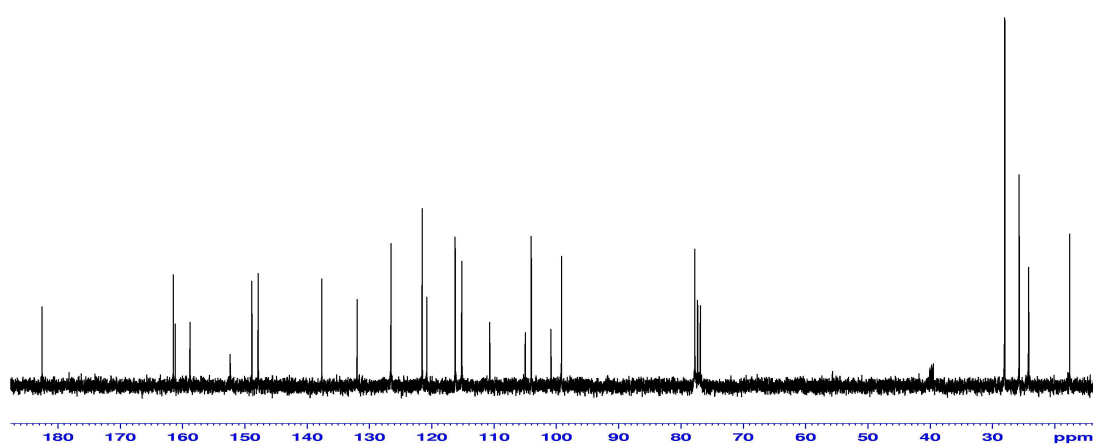


Figure 9 ^{13}C NMR (75 MHz) ($\text{CDCl}_3 + \text{DMSO-}d_6$) spectrum of AE1

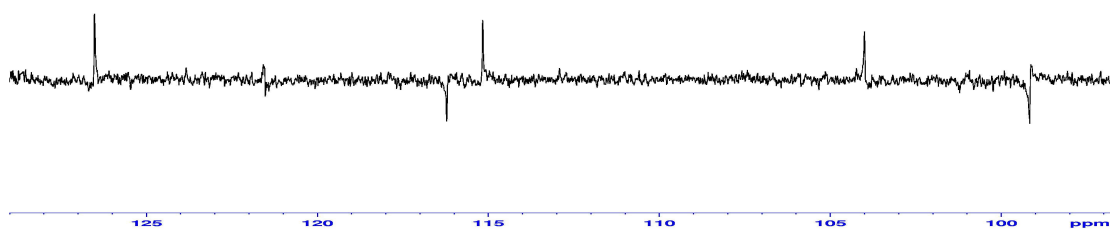


Figure 10 DEPT 135° ($\text{CDCl}_3 + \text{DMSO-}d_6$) spectrum of AE1

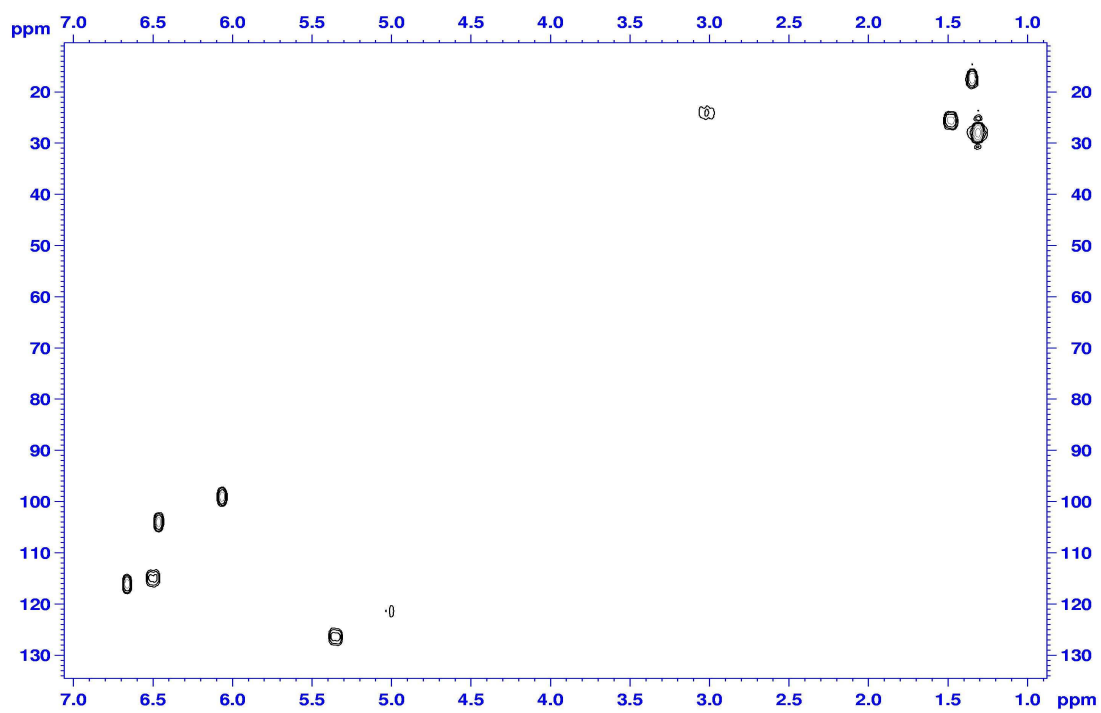


Figure 11 2D HMQC spectrum of AE1

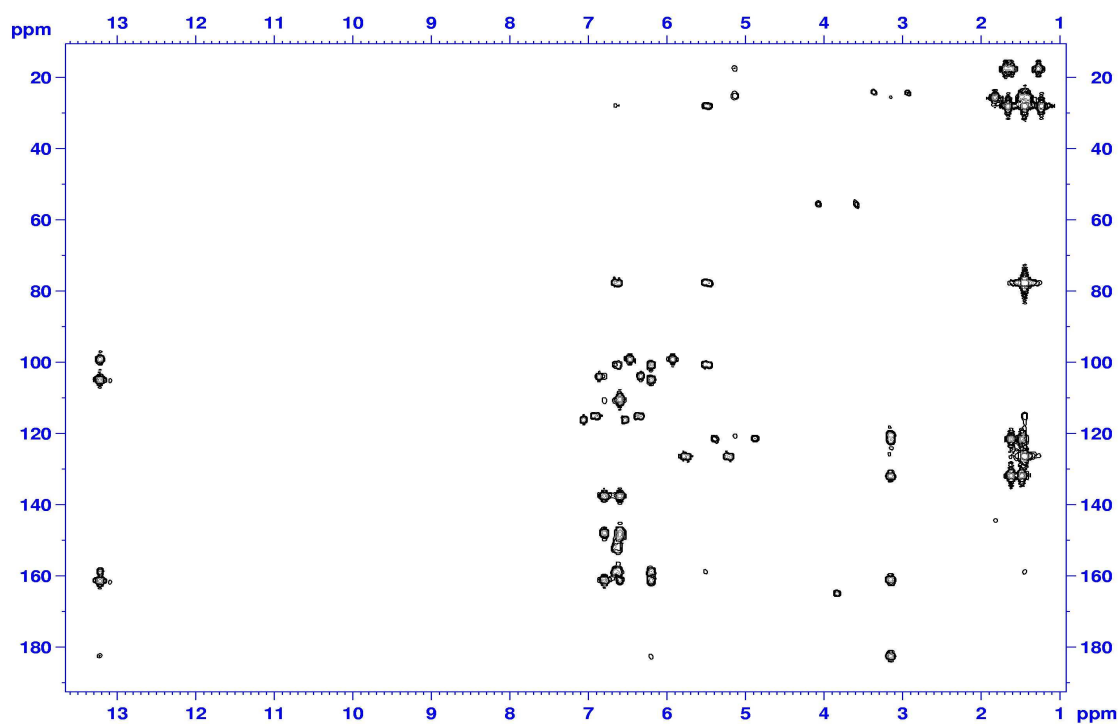


Figure 12 2D HMBC spectrum of AE1

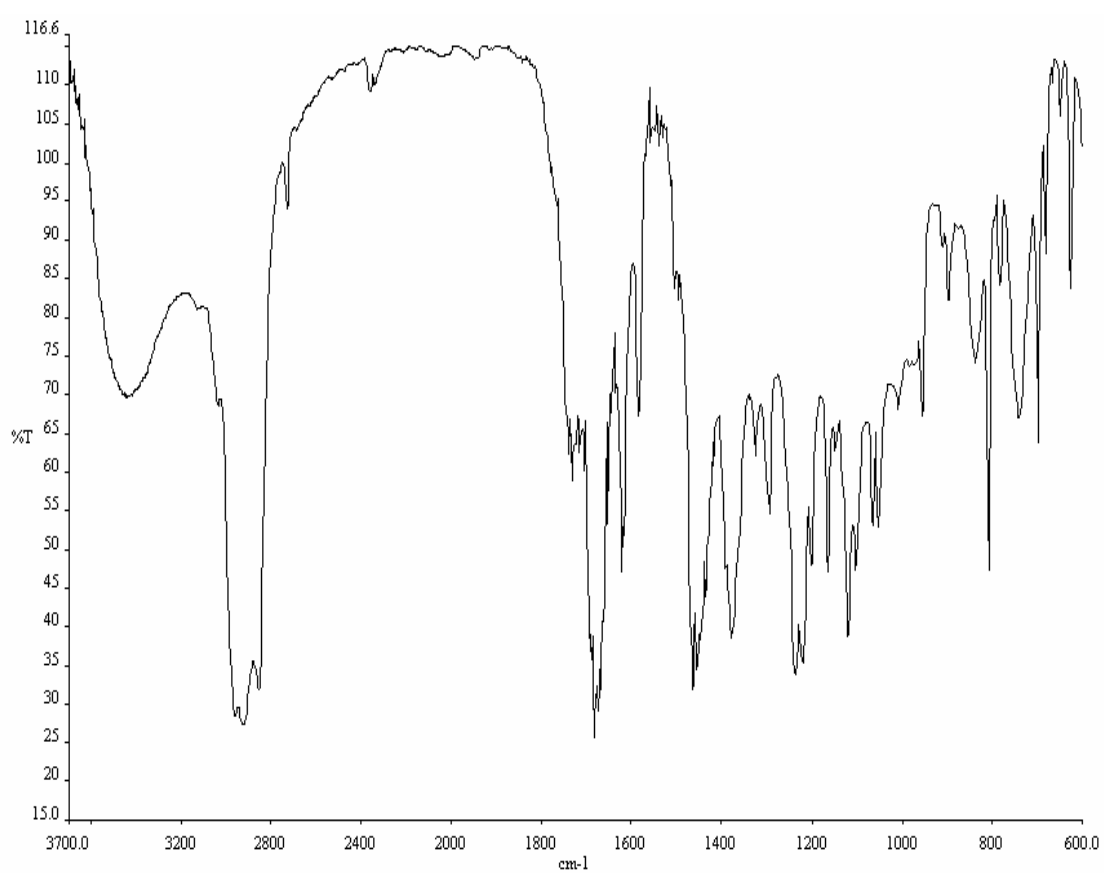


Figure 13 FT-IR (Neat) spectrum of **AE2**

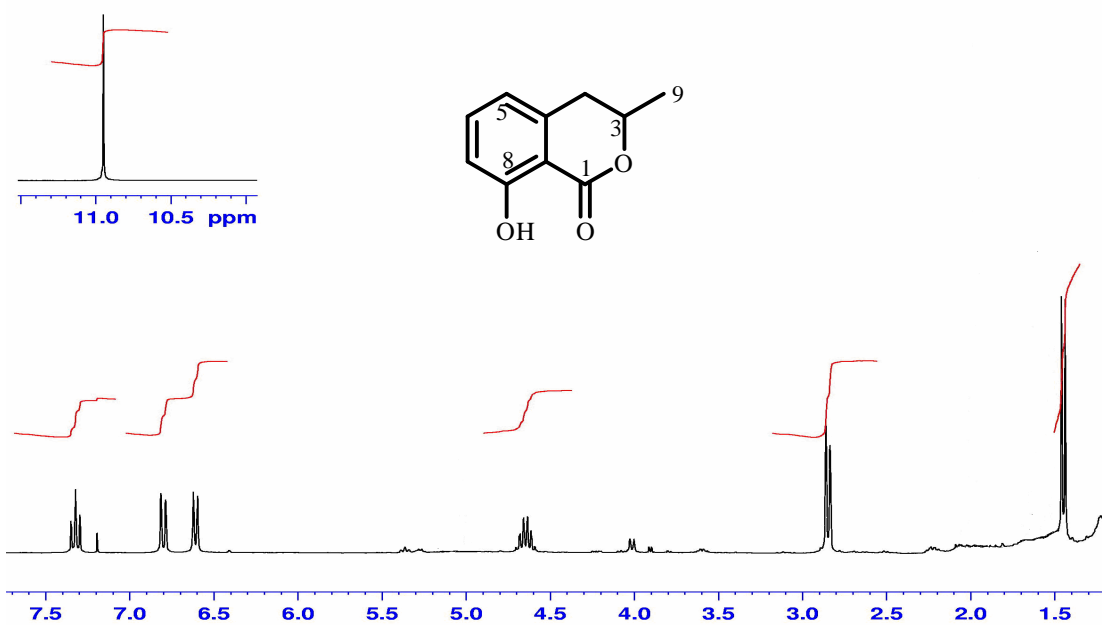


Figure 14 ^1H NMR (300 MHz) (CDCl_3) spectrum of AE2

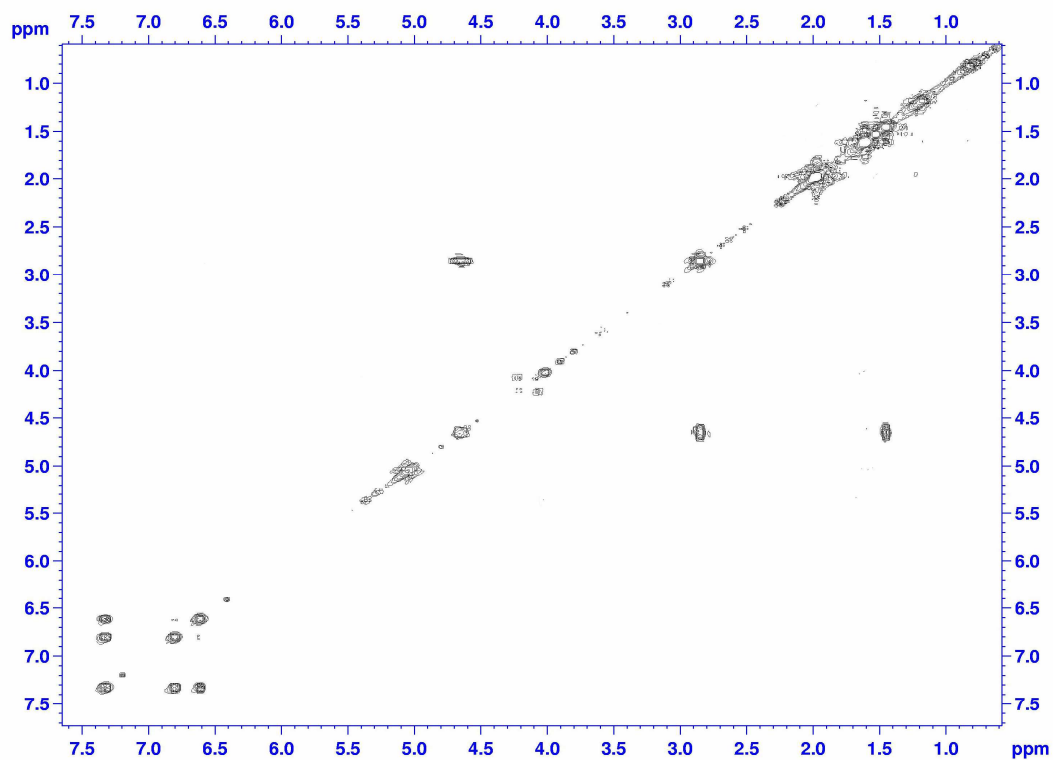


Figure 15 ^1H - ^1H COSY spectrum of AE2

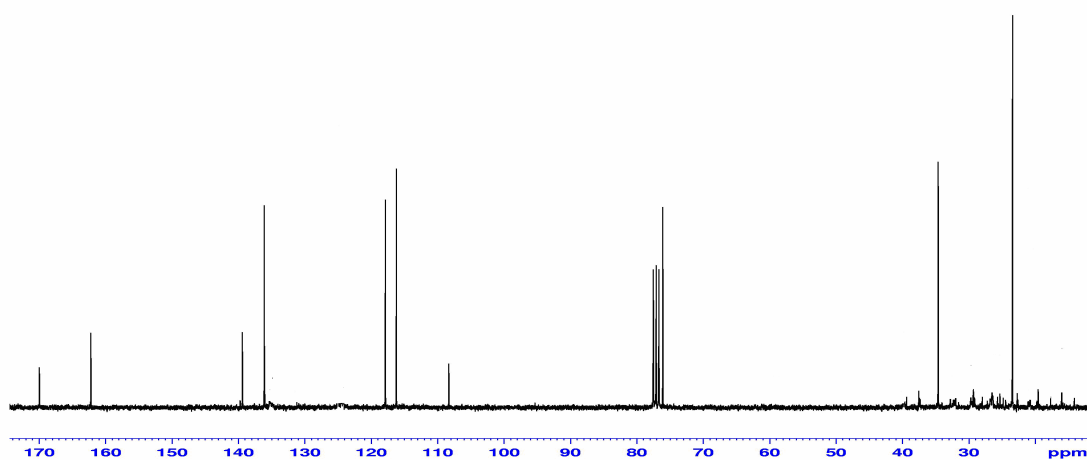


Figure 16 ^{13}C NMR (75 MHz) (CDCl_3) spectrum of AE2

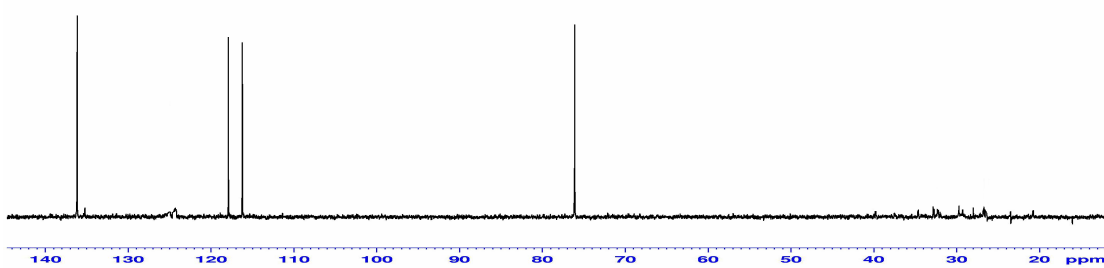


Figure 17 DEPT 135° (CDCl_3) spectrum of AE2

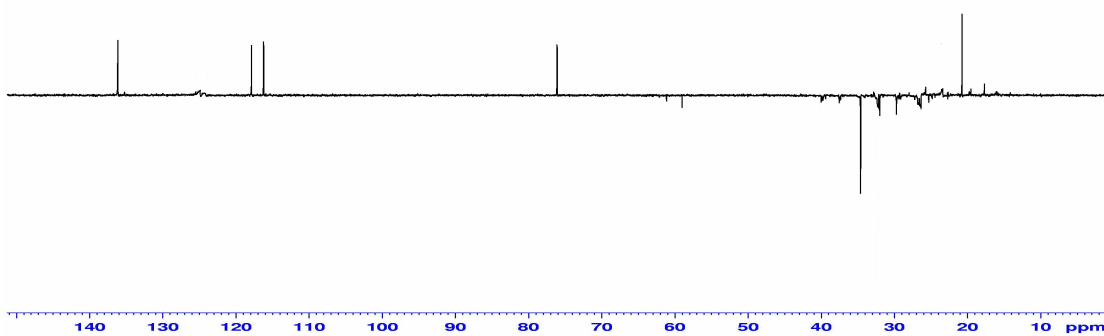


Figure 18 DEPT 90° (CDCl_3) spectrum of AE2

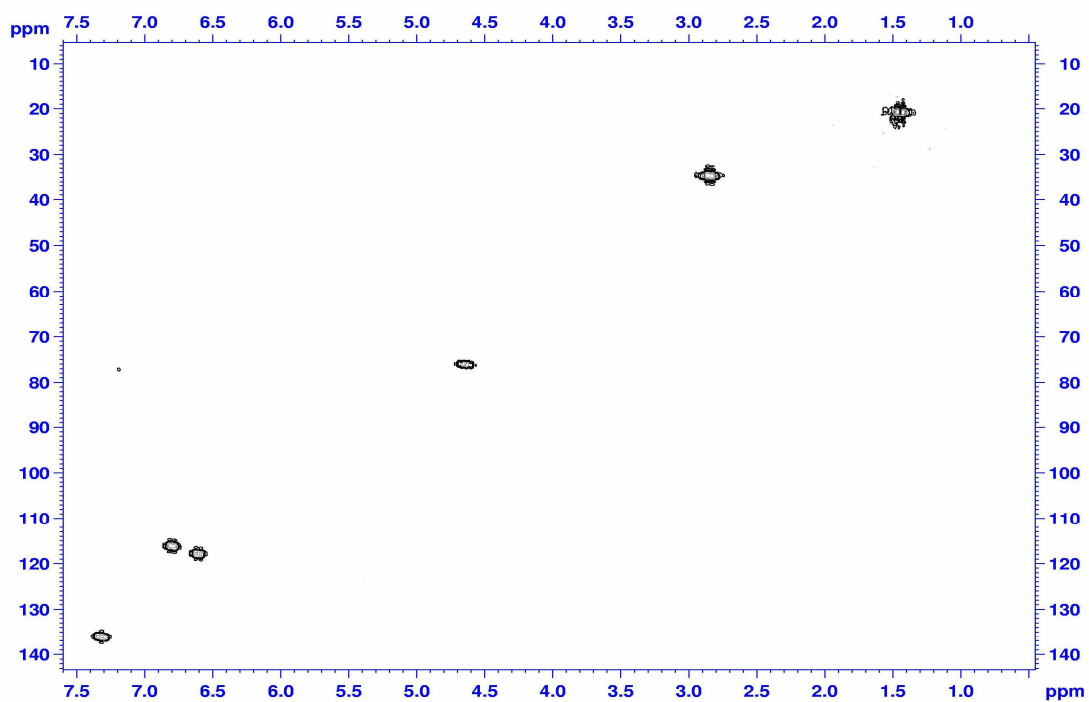


Figure 19 2D HMQC spectrum of AE2

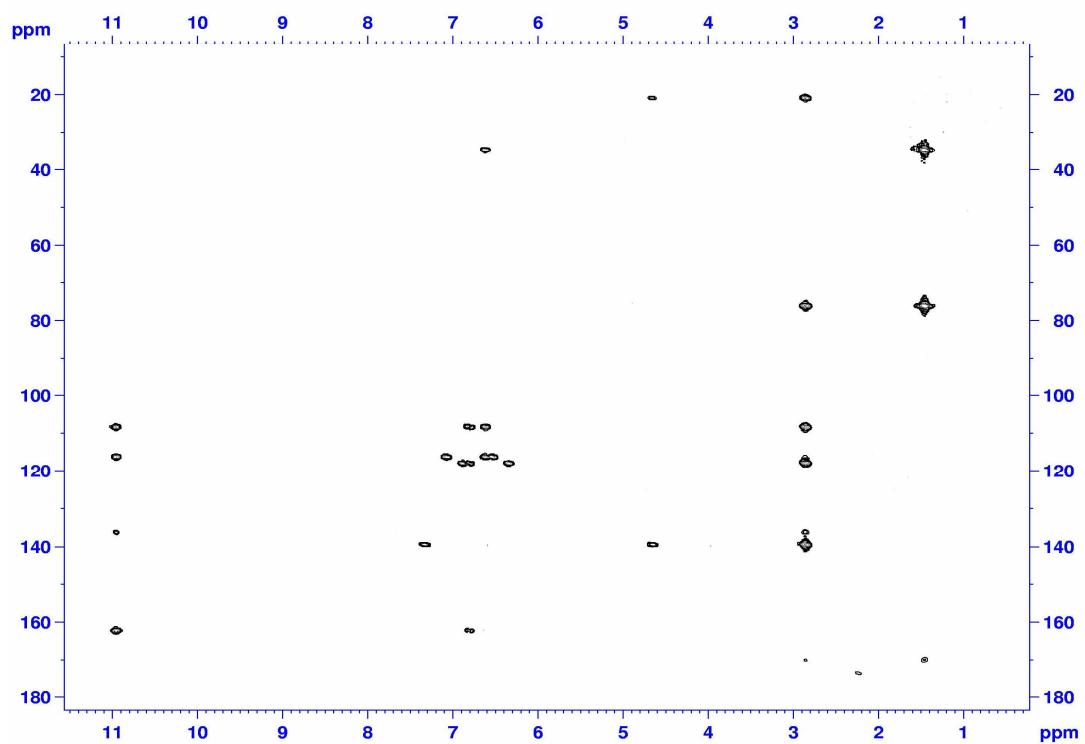


Figure 20 2D HMBC spectrum of AE2

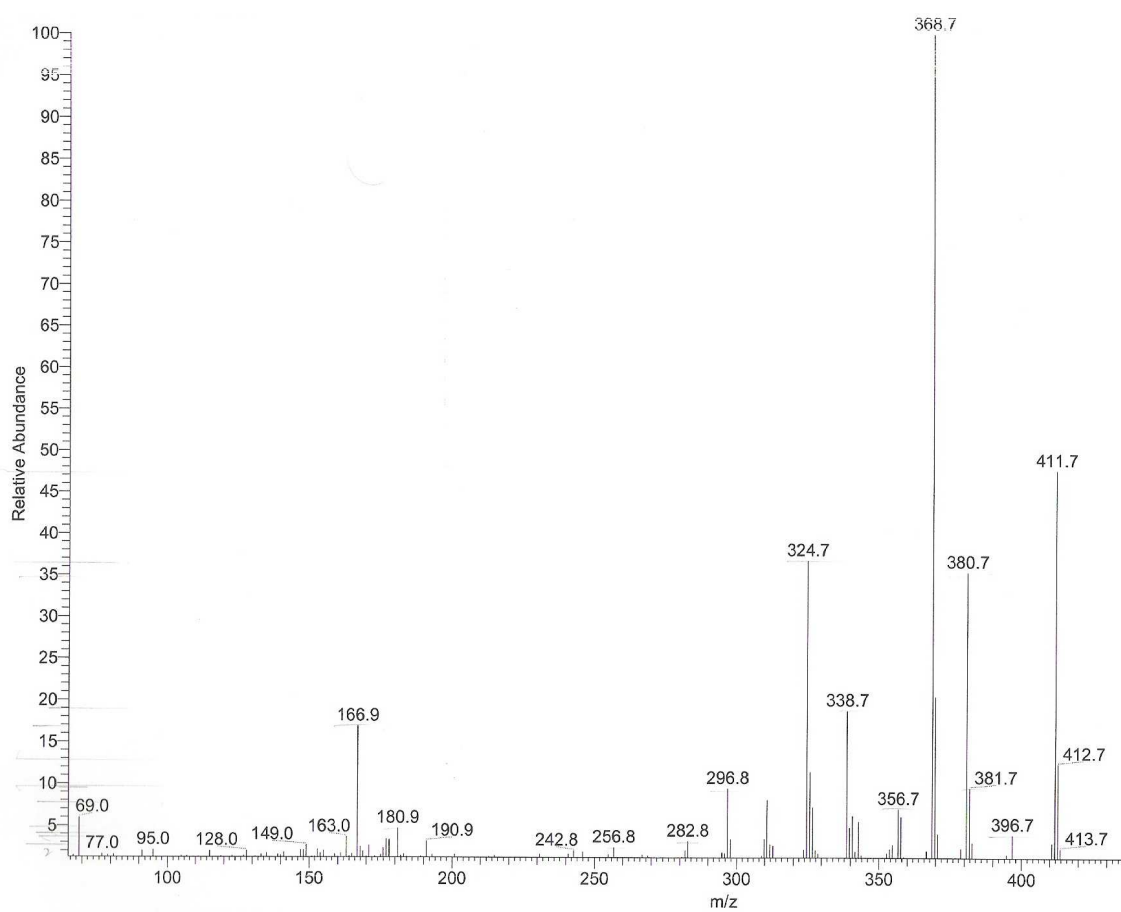


Figure 21 EI-MS spectrum of AE3

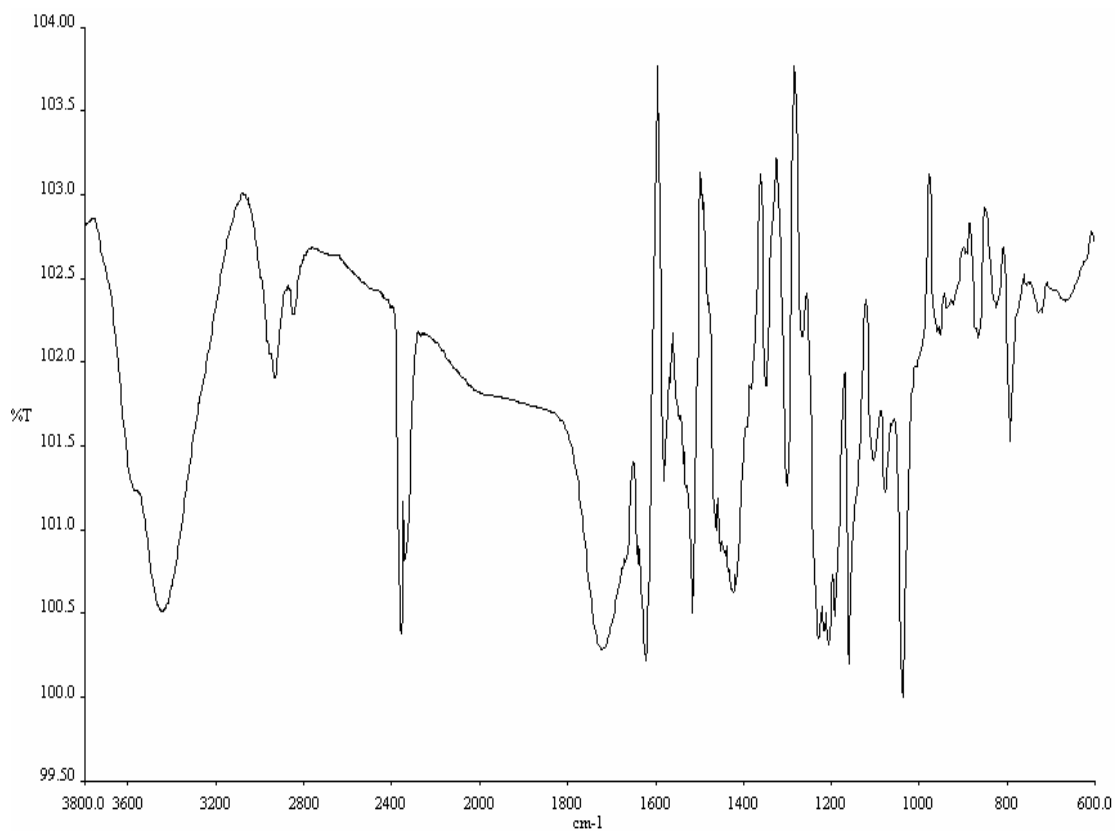


Figure 22 FT-IR (Neat) spectrum of **AE3**

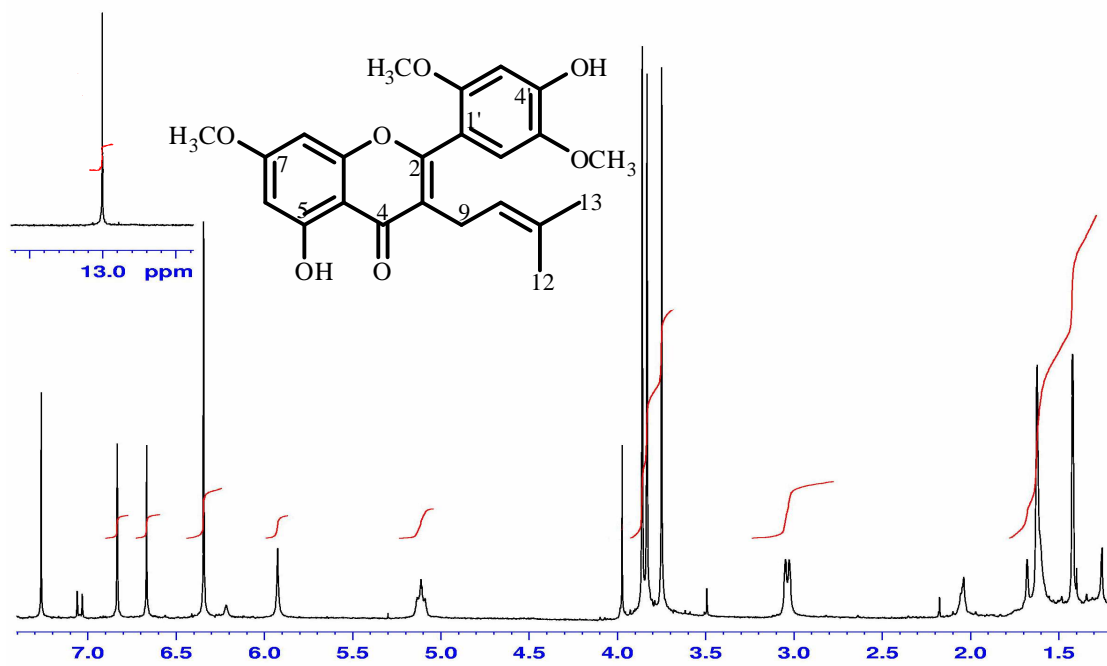


Figure 23 ¹H NMR (300 MHz) (CDCl₃) spectrum of **AE3**

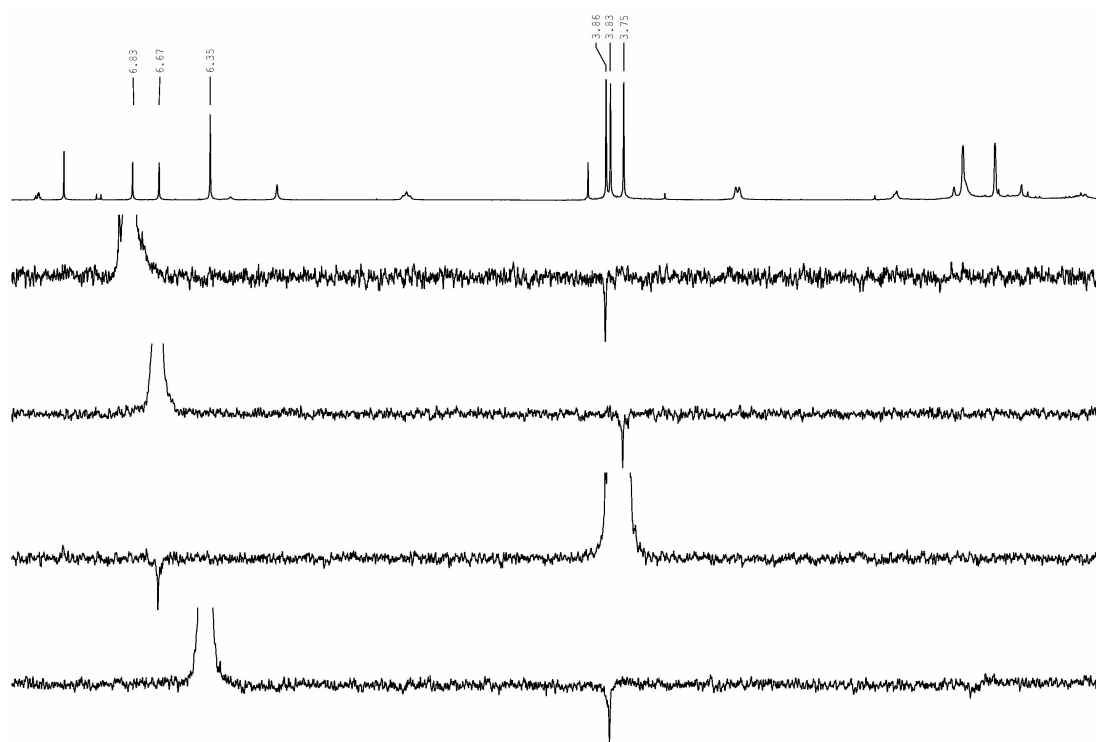


Figure 29 NOEDIFF spectrum of AE3 after irradiation

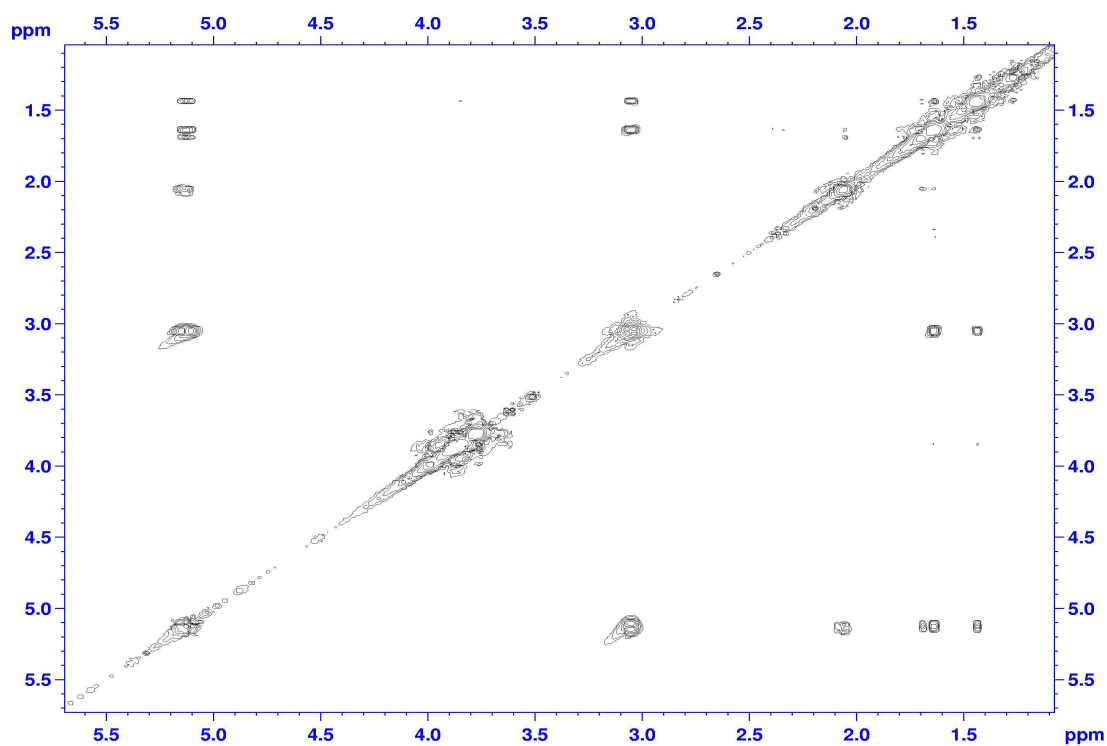


Figure 25 ^1H - ^1H COSY spectrum of AE3

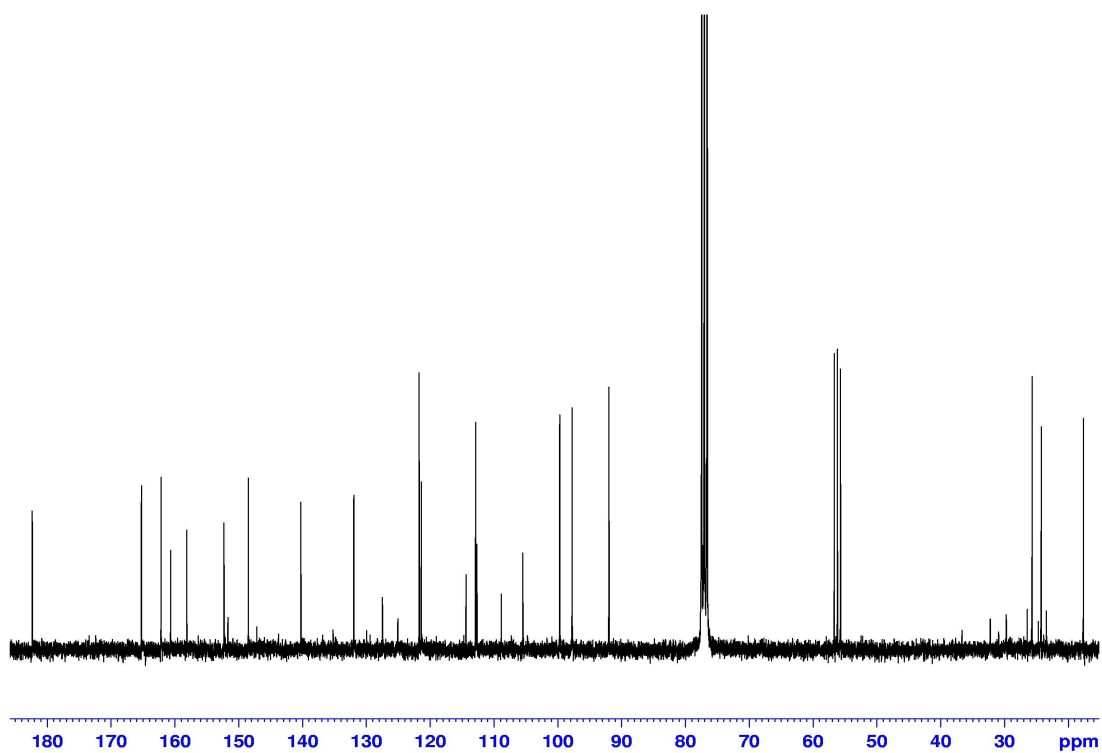


Figure 26 ^{13}C NMR (75 MHz) (CDCl_3) spectrum of AE3

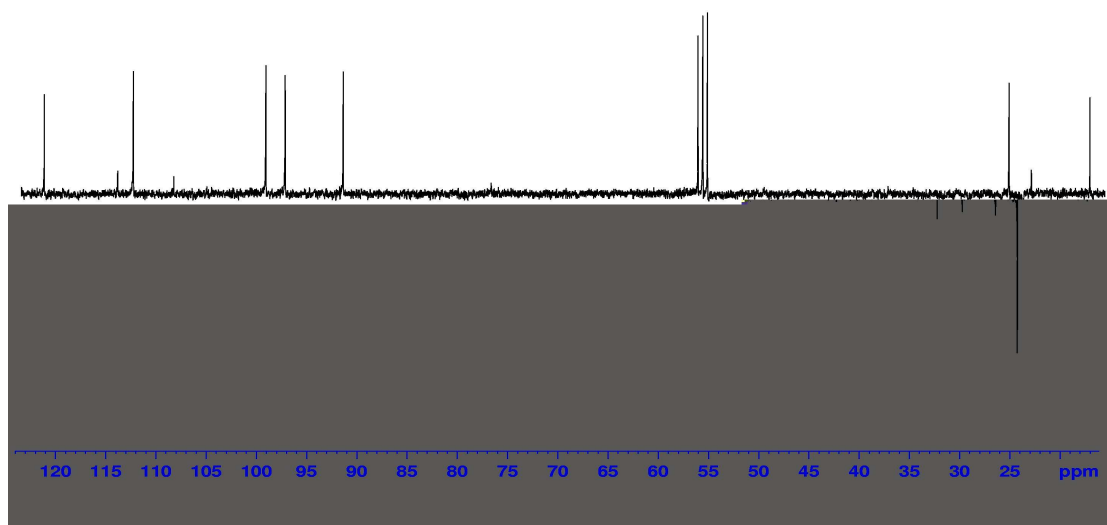


Figure 27 DEPT 135° (CDCl_3) spectrum of AE3

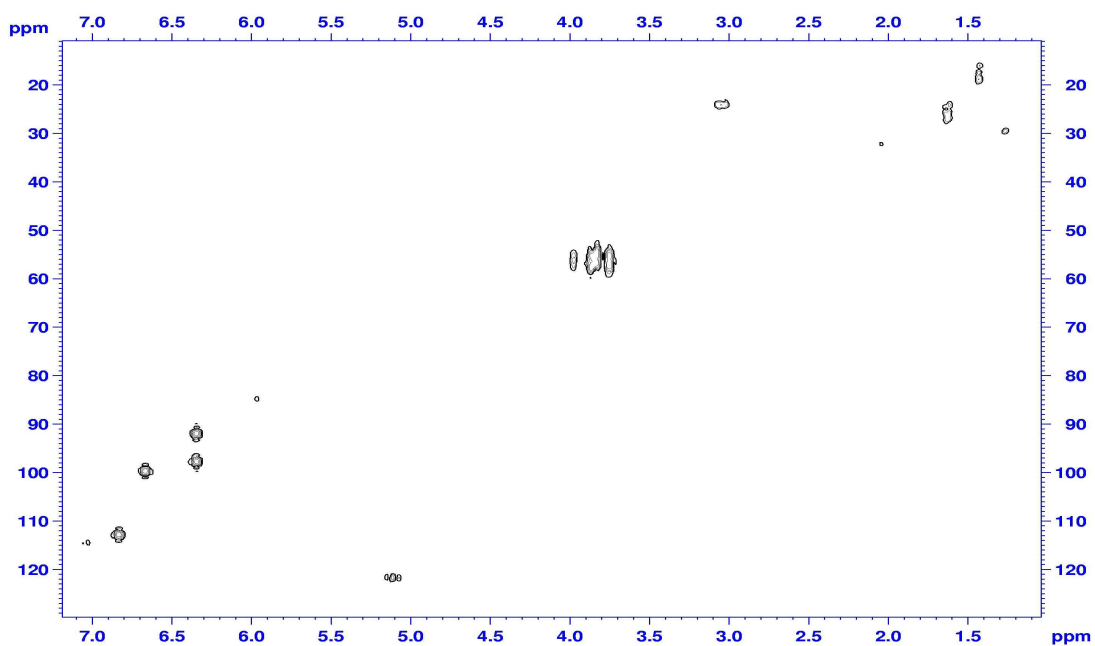


Figure 28 2D HMQC spectrum of AE3

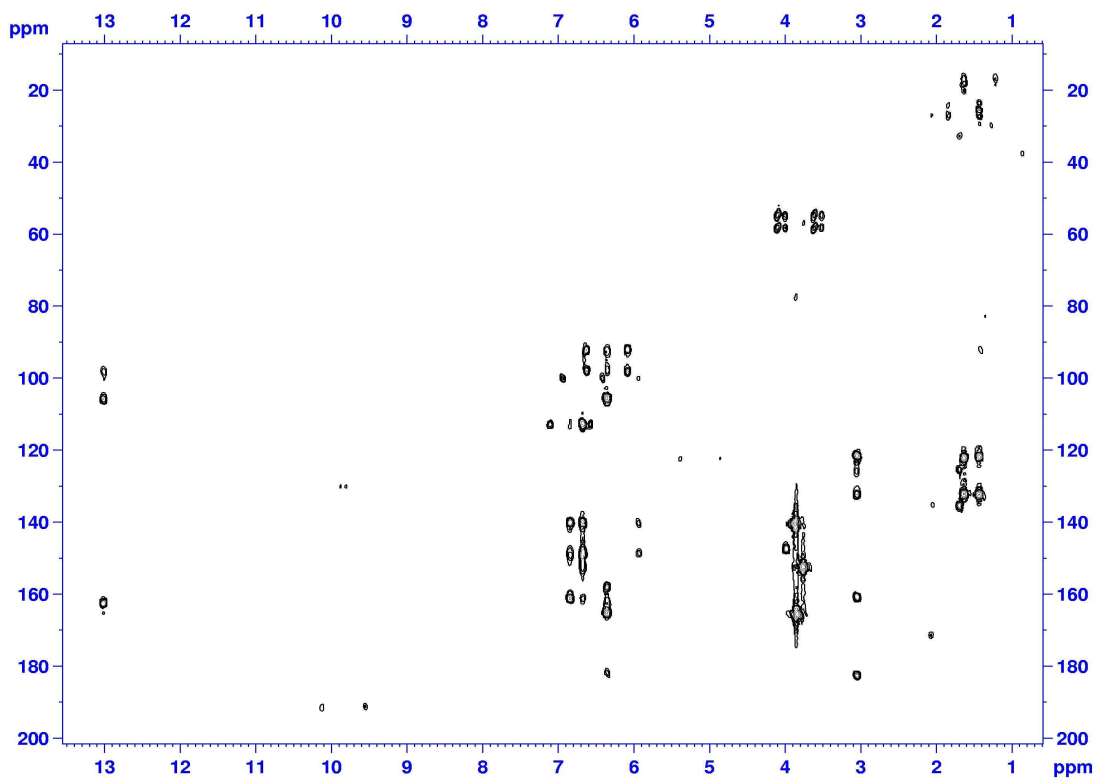


Figure 29 2D HMBC spectrum of AE3

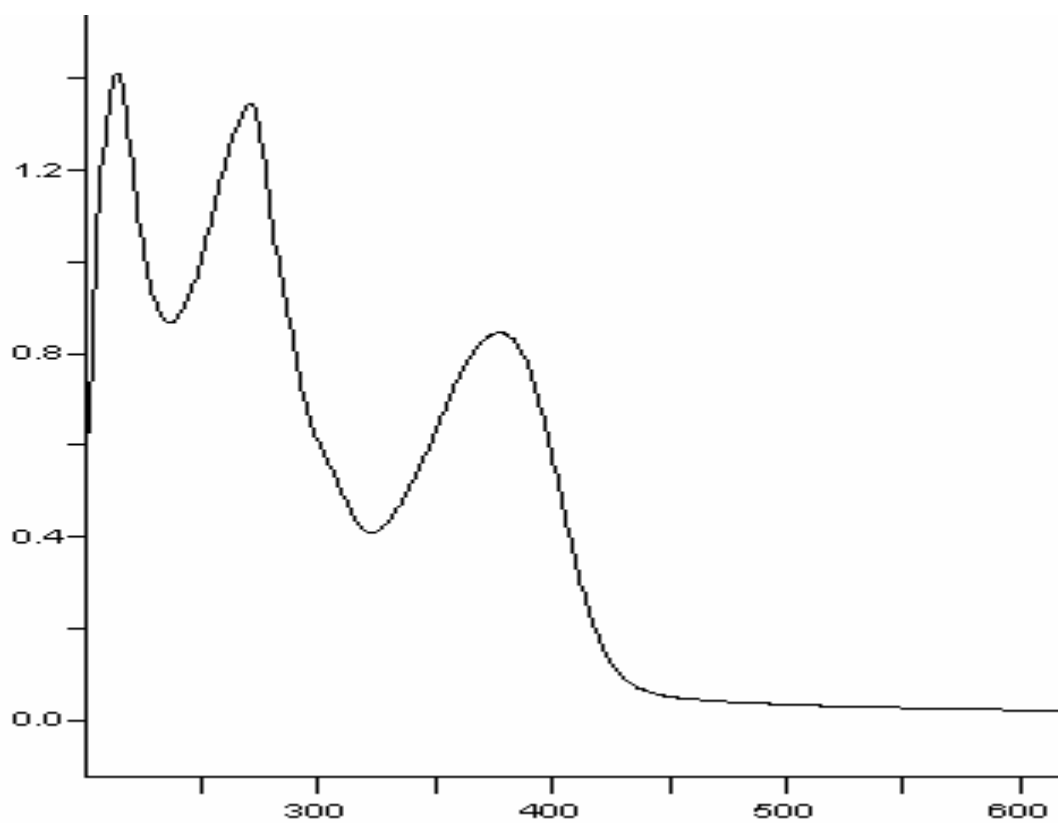


Figure 30 UV (EtOH) spectrum of **AE4**

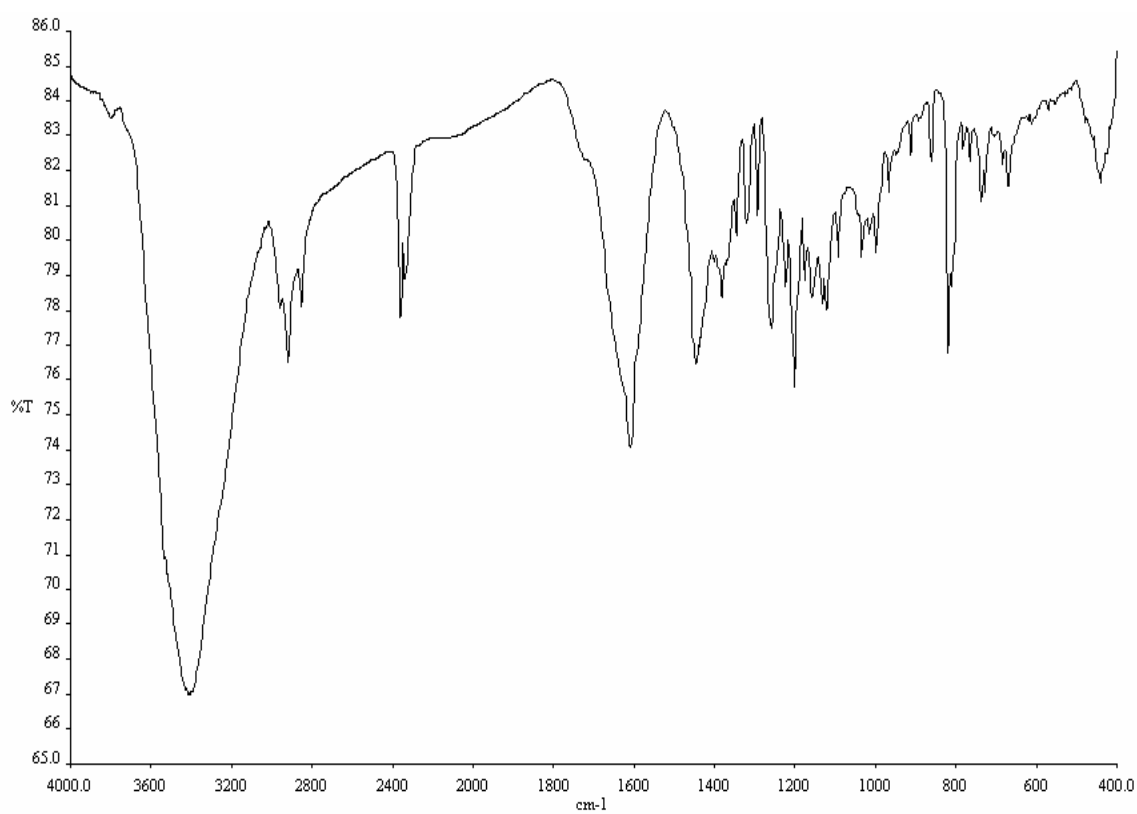


Figure 31 FT-IR (Neat) spectrum of **AE4**

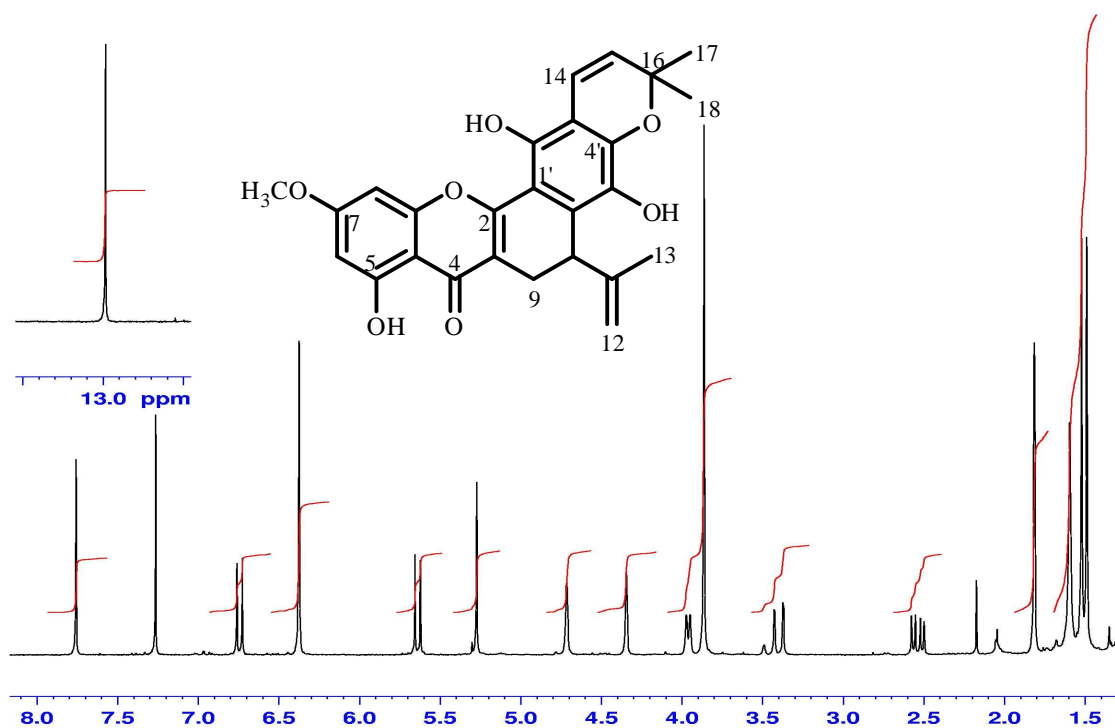


Figure 32 ^1H NMR (300 MHz) (CDCl_3) spectrum of AE4

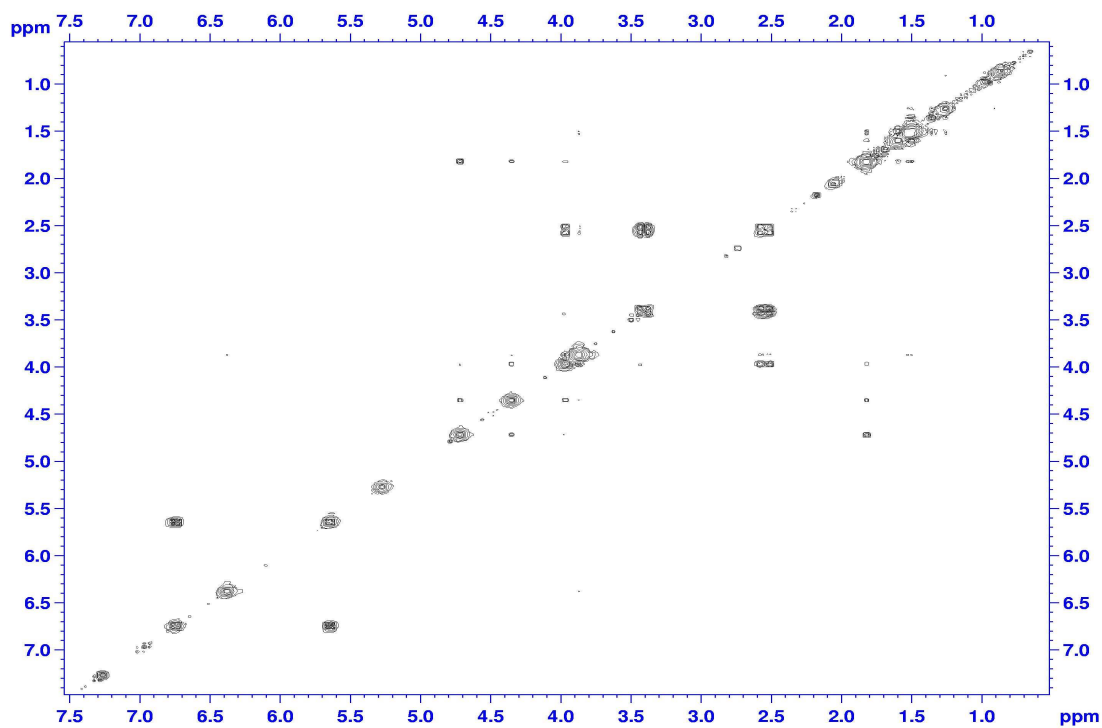


Figure 33 ^1H - ^1H COSY spectrum of AE4

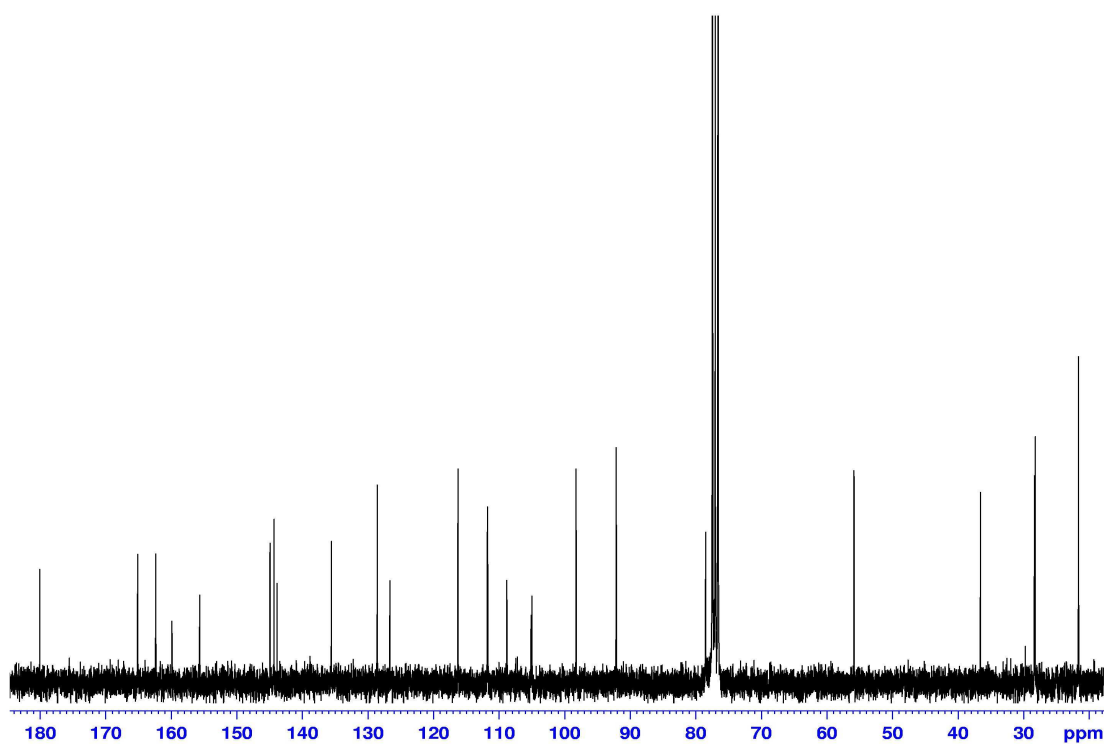


Figure 34 ^{13}C NMR (75 MHz) (CDCl_3) spectrum of **AE4**

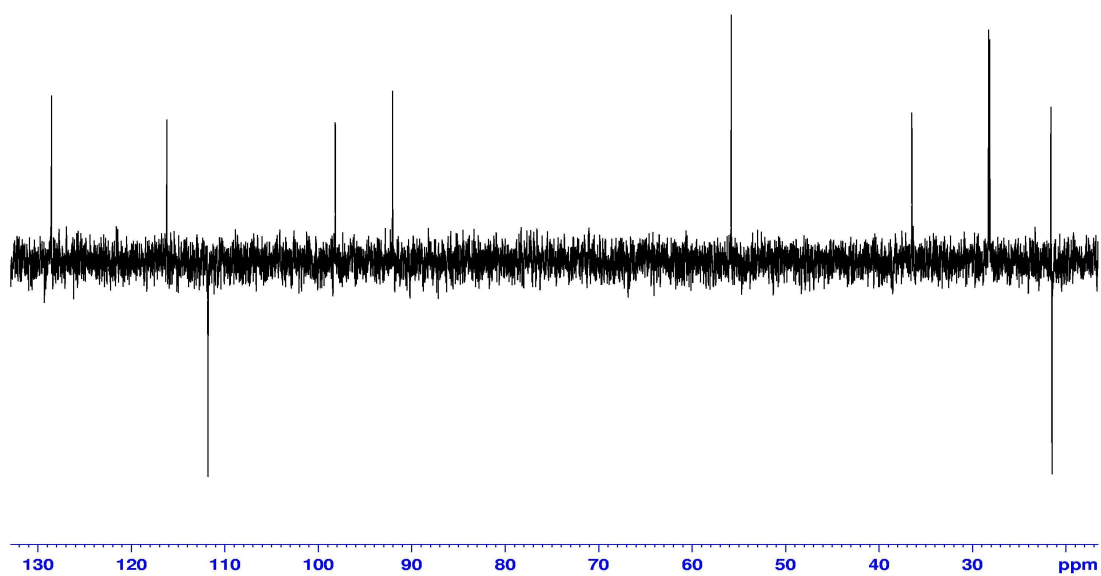


Figure 35 DEPT 135° (CDCl_3) spectrum of **AE4**

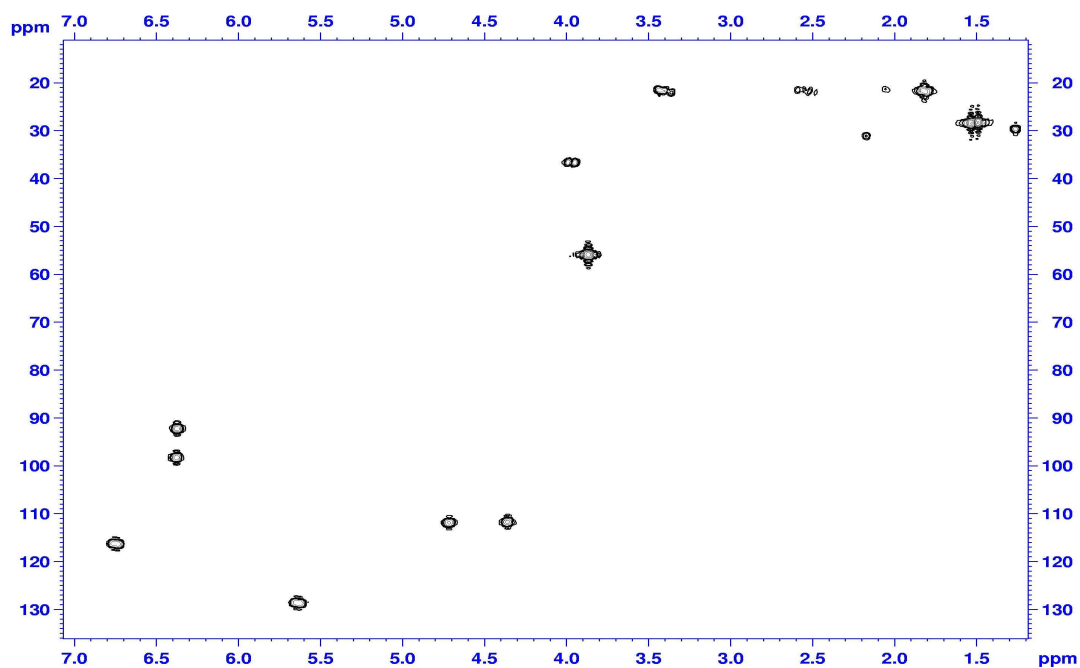


Figure 36 2D HMQC spectrum of AE4

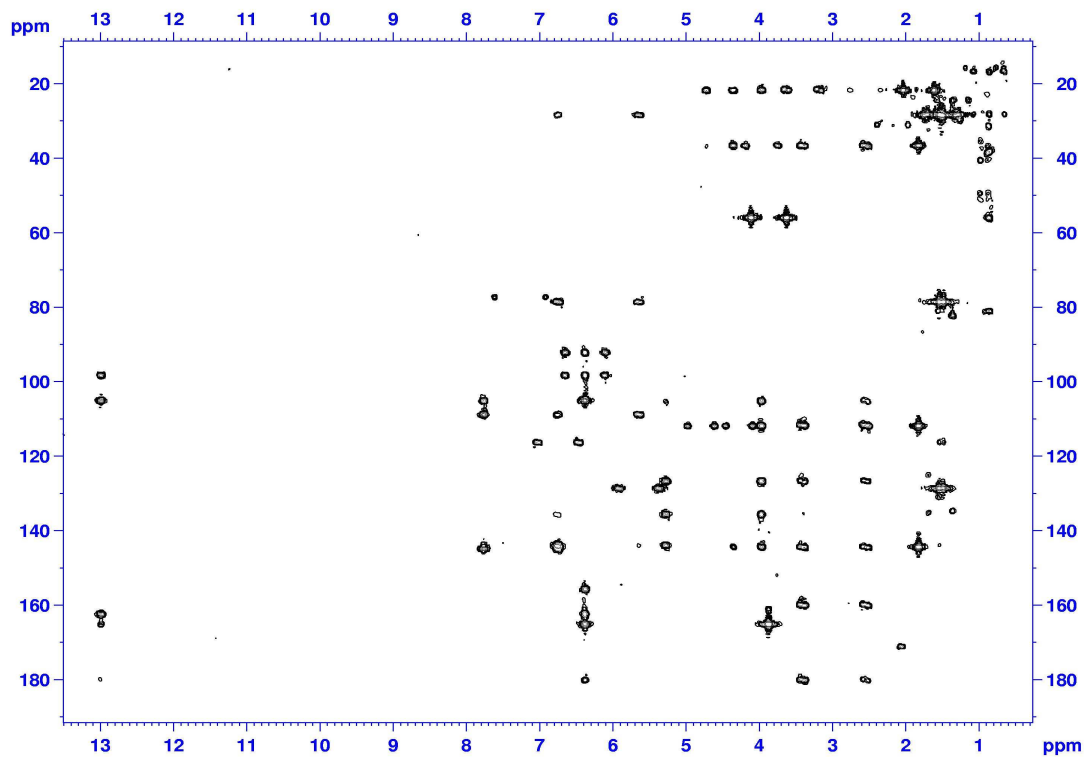


Figure 37 2D HMBC spectrum of AE4

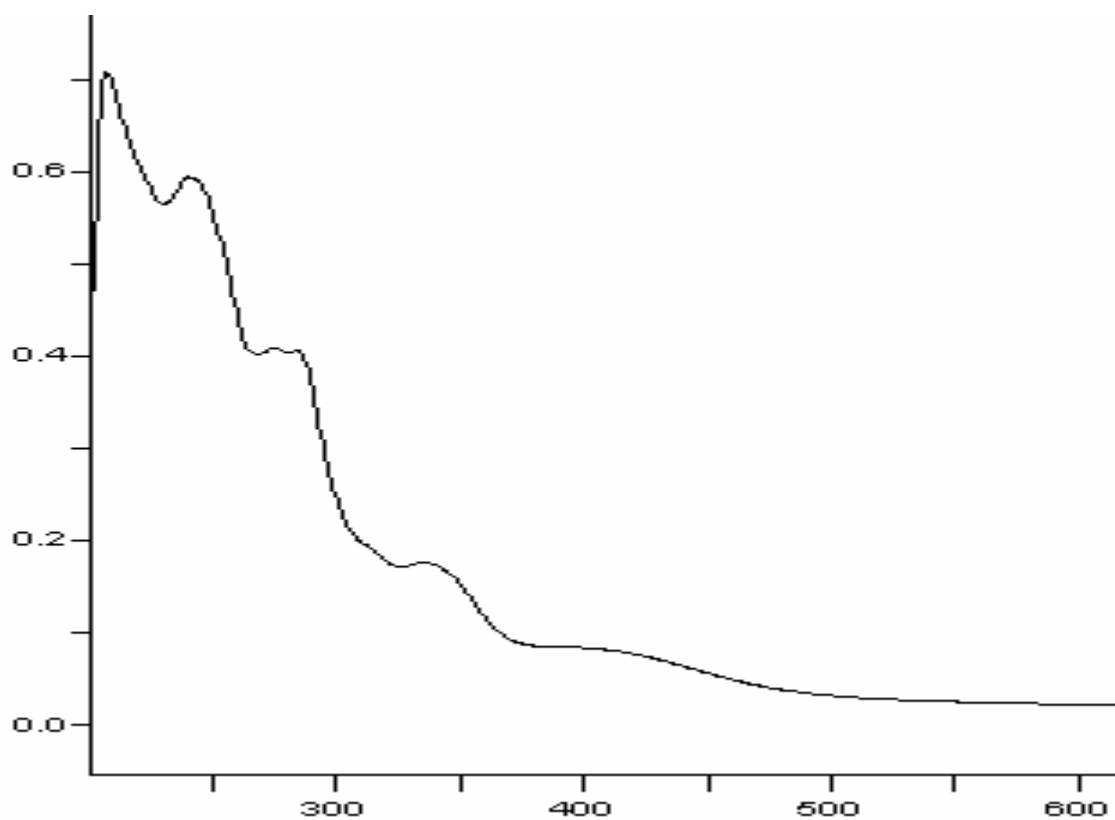


Figure 38 UV (EtOH) spectrum of **AE5**

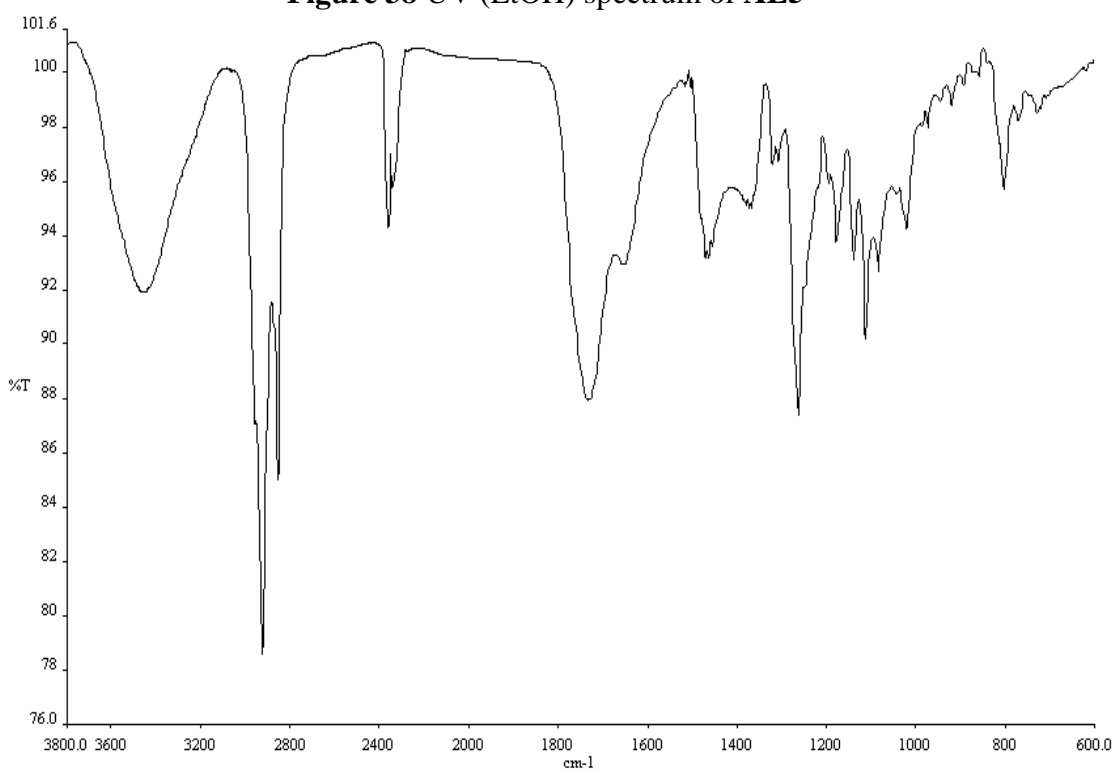


Figure 39 FT-IR (Neat) spectrum of **AE5**

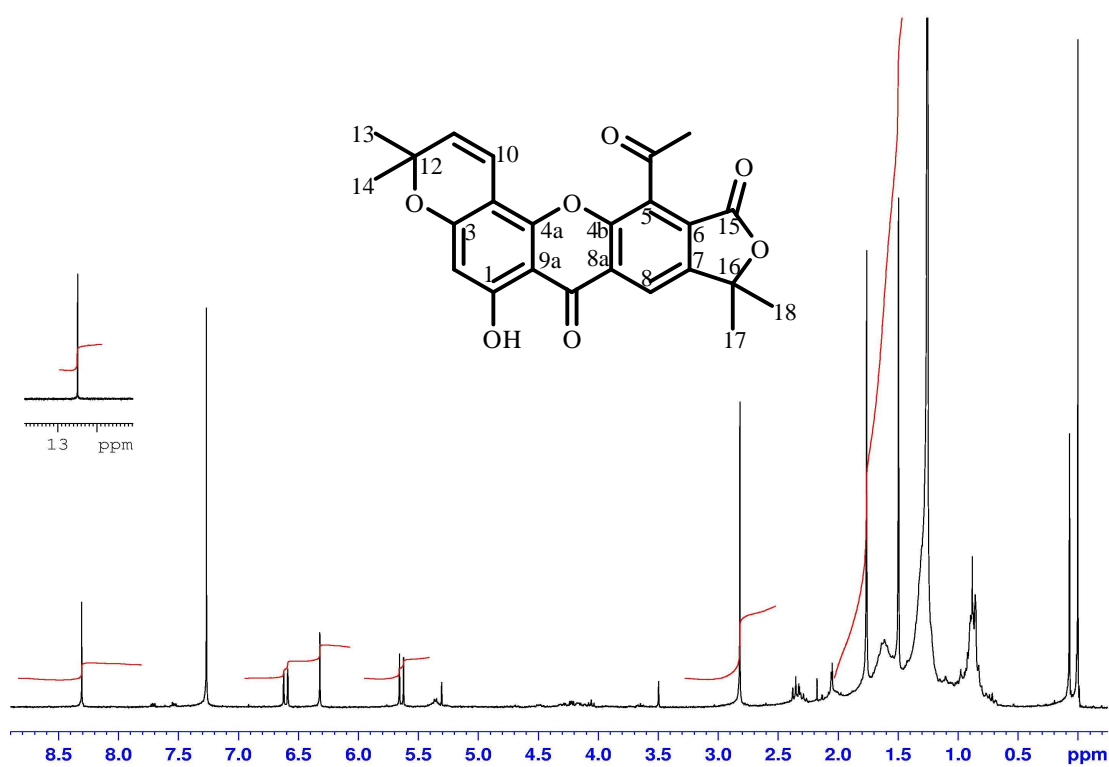


Figure 40 ^1H NMR (300 MHz) (CDCl_3) spectrum of AE5

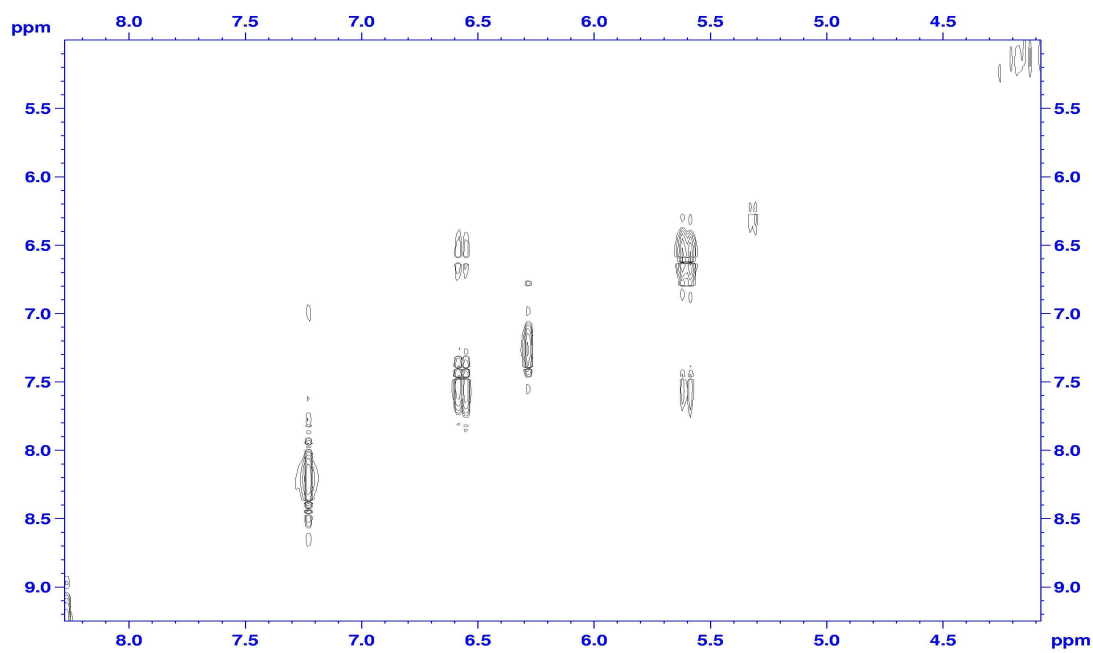


Figure 41 ^1H - ^1H COSY spectrum of AE5

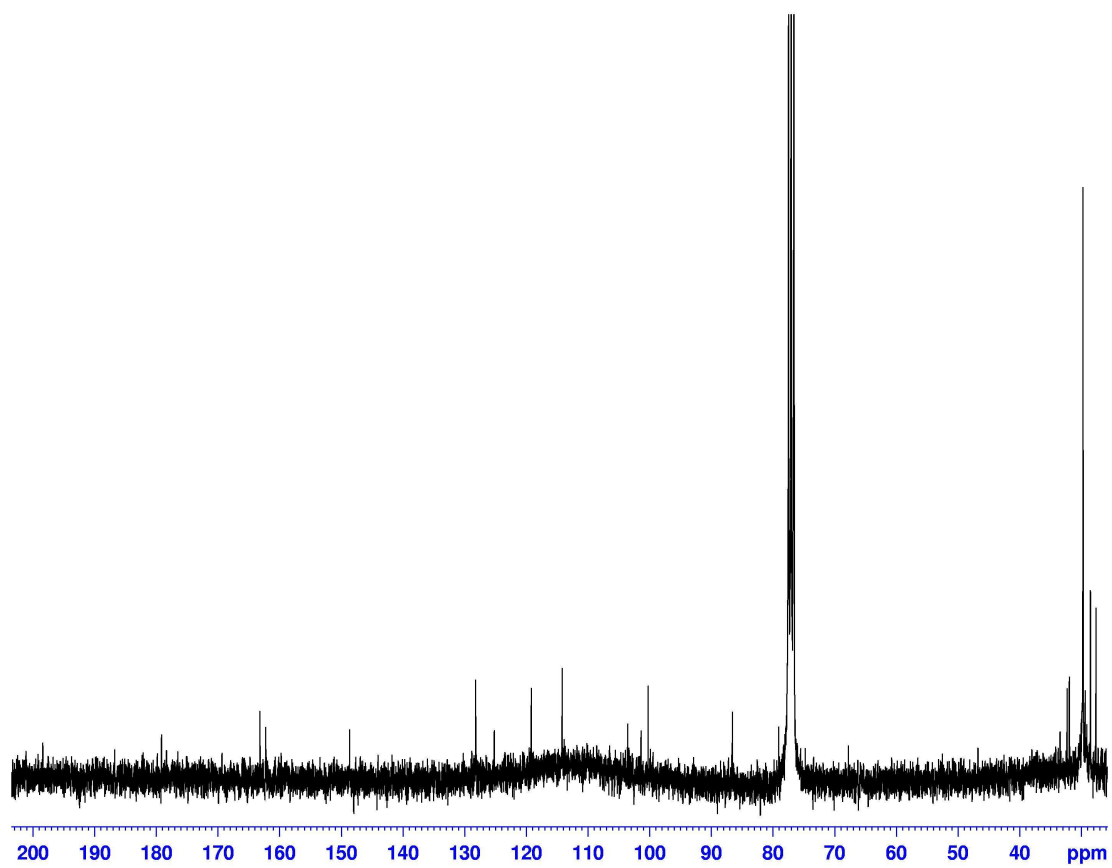


Figure 42 ^{13}C NMR (75 MHz) (CDCl_3) spectrum of AE5

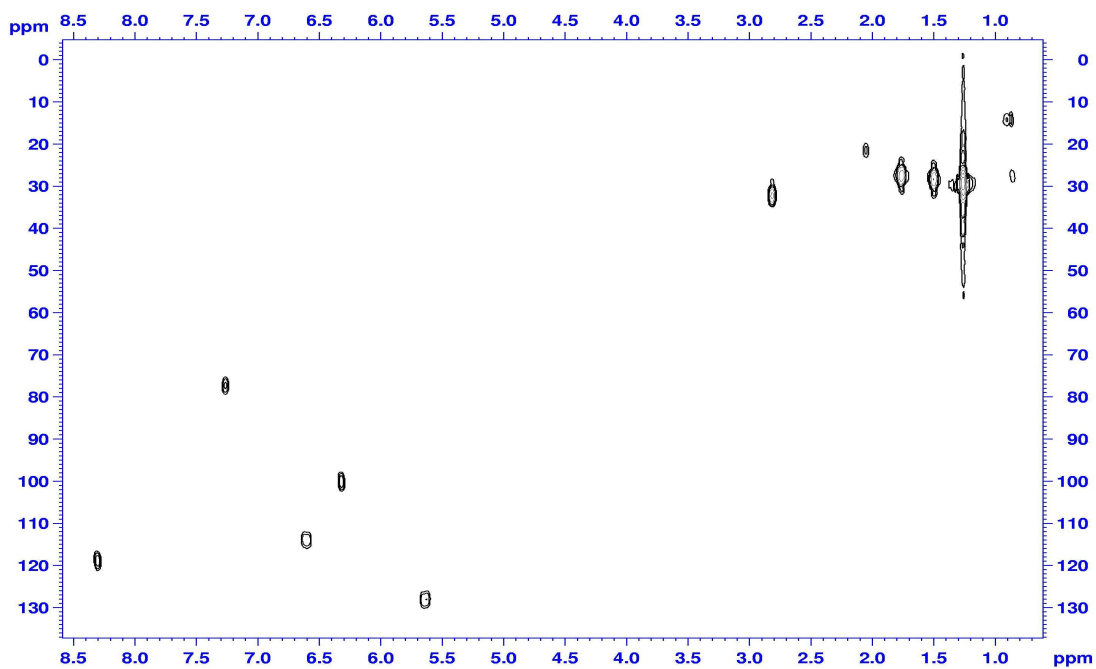


Figure 43 2D HMQC spectrum of AE5

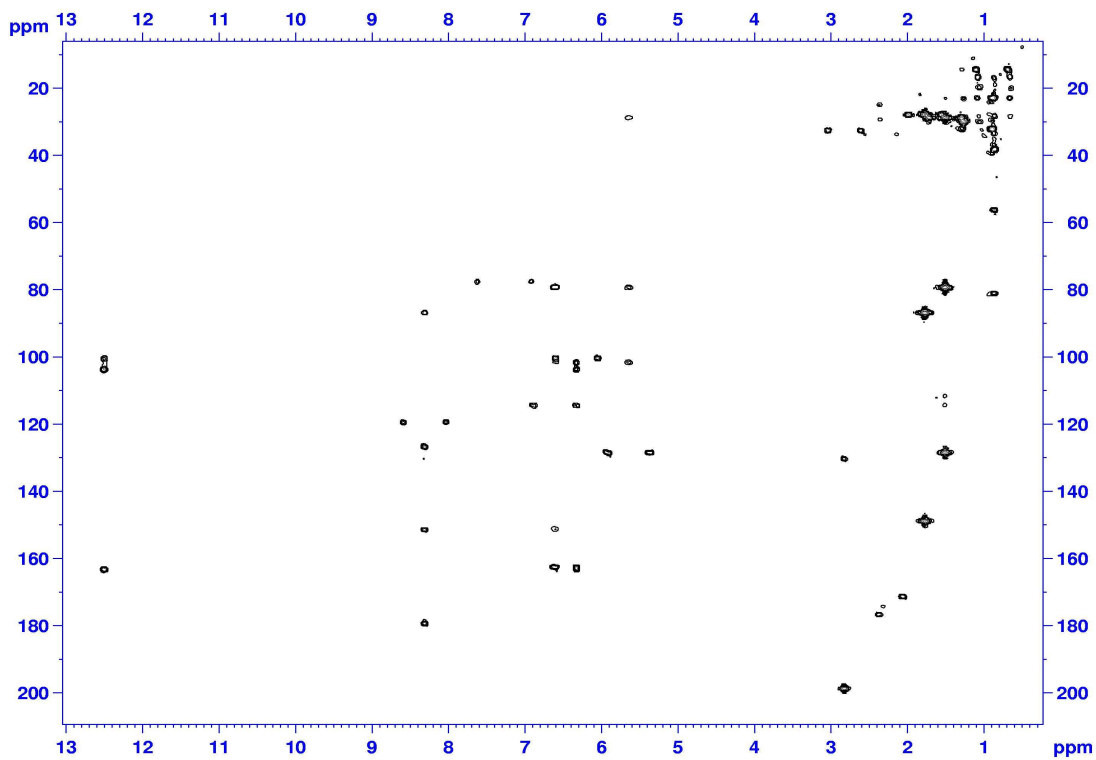


Figure 44 2D HMBC spectrum of AE5

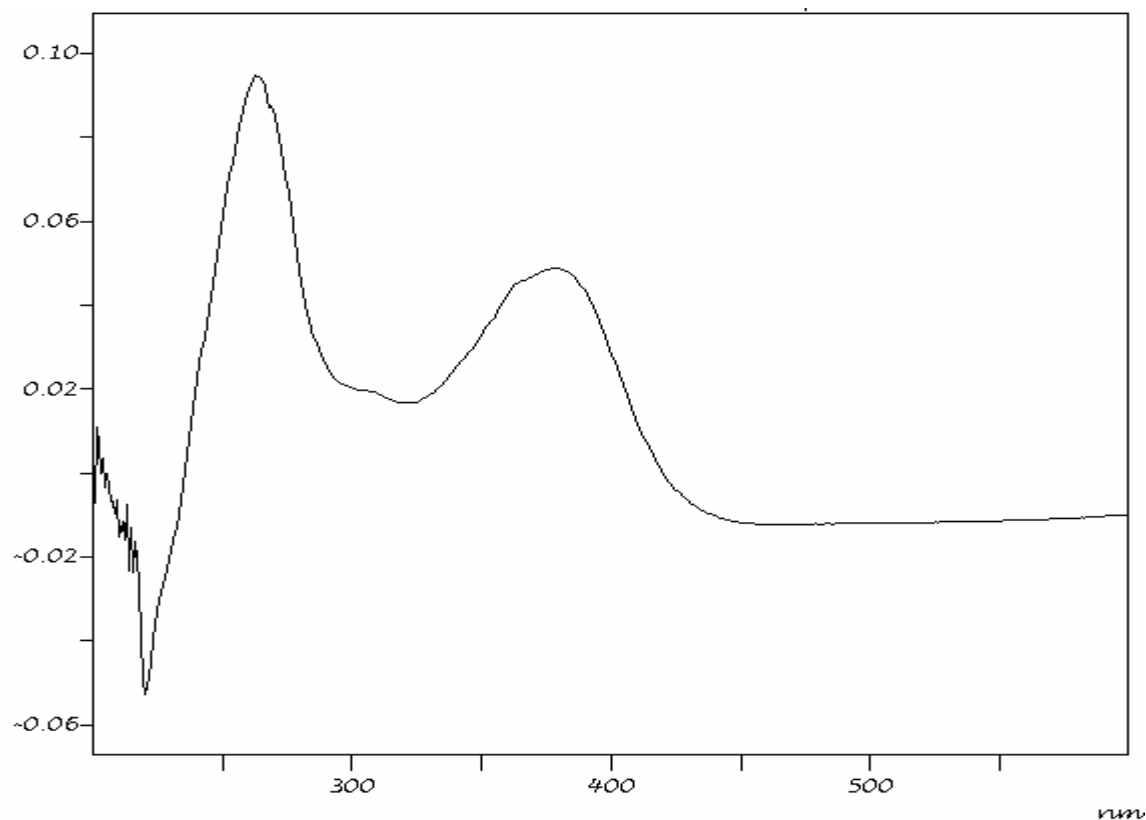


Figure 45 UV (CH₃OH) spectrum of AE6



Figure 46 FT-IR (Neat) spectrum of AE6

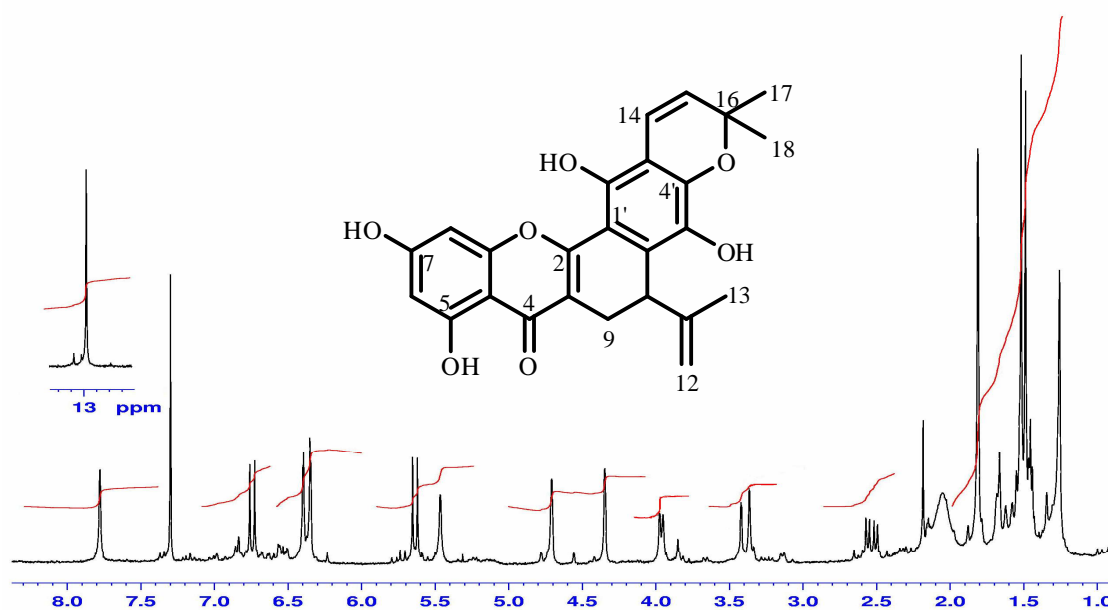


Figure 47 ^1H NMR (300 MHz) ($\text{CDCl}_3 + \text{Acetone-}d_6$) spectrum of AE6

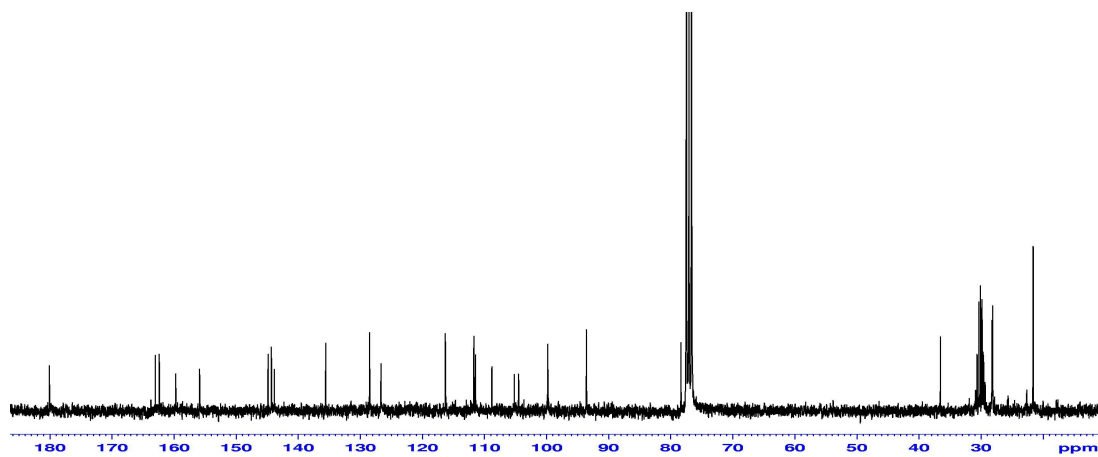


Figure 48 ^{13}C NMR (75 MHz) ($\text{CDCl}_3 + \text{Acetone-}d_6$) spectrum of AE6

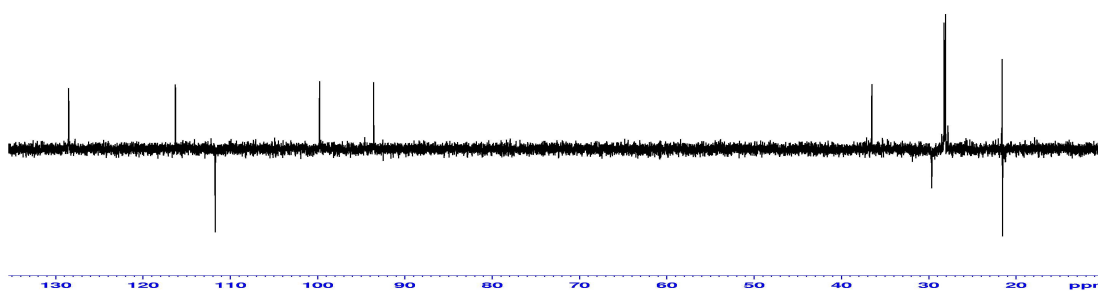


Figure 49 DEPT 135° ($\text{CDCl}_3 + \text{Acetone-}d_6$) spectrum of AE6

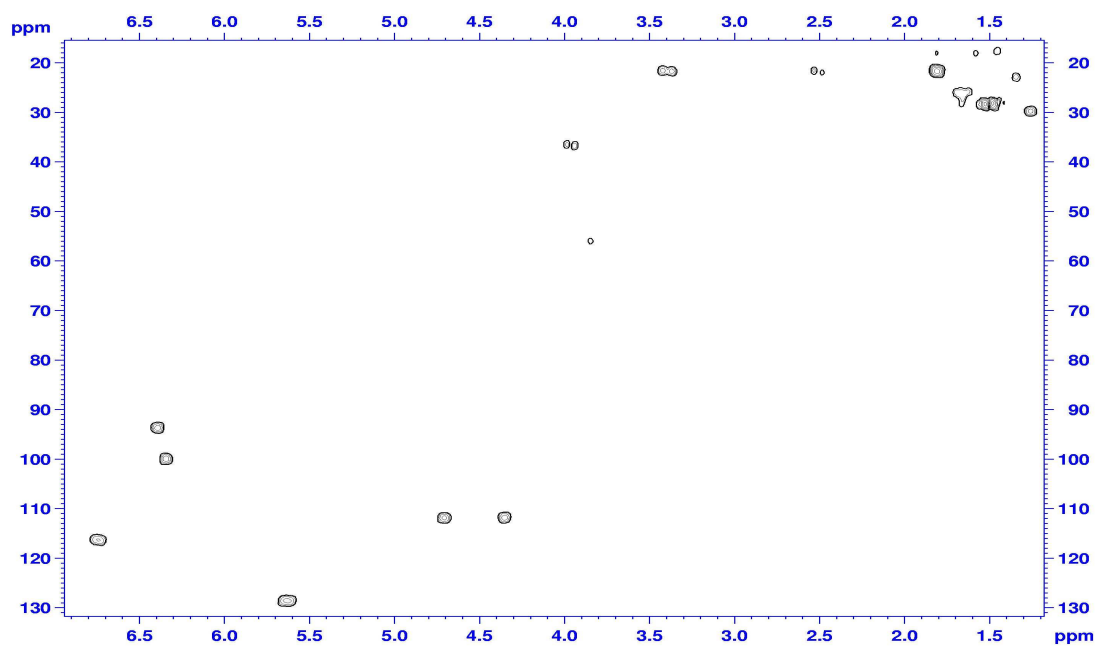


Figure 50 2D HMQC spectrum of AE6

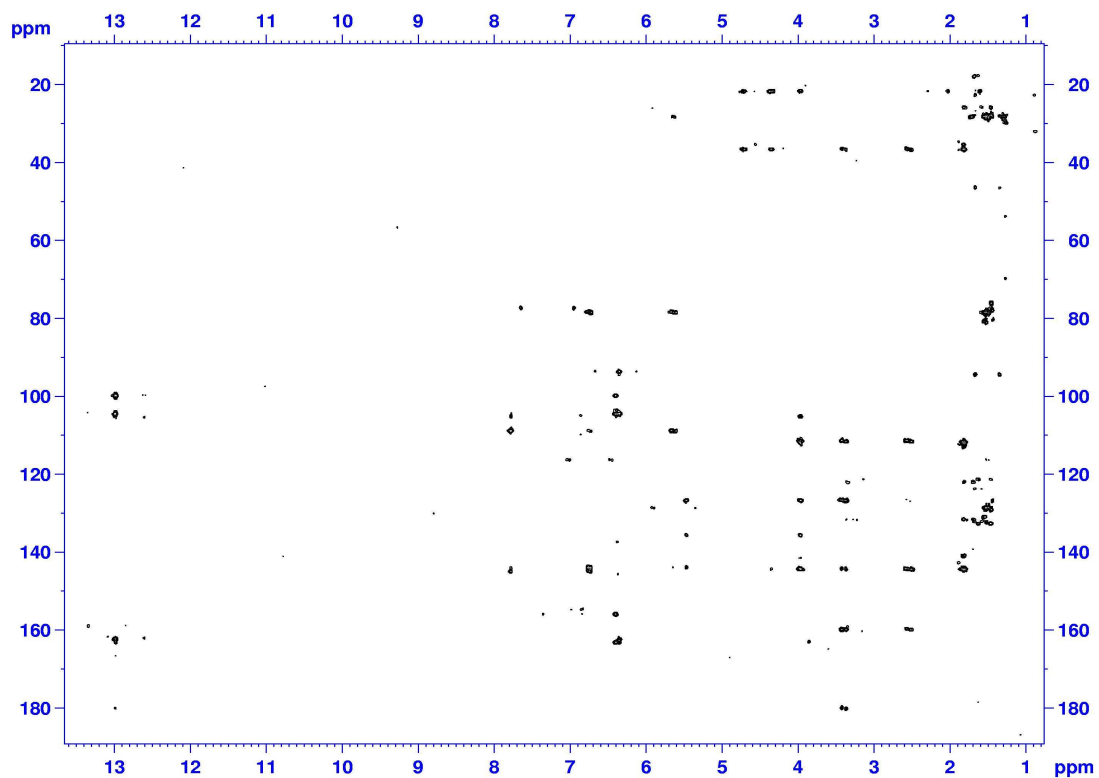


Figure 51 2D HMBC spectrum of AE6

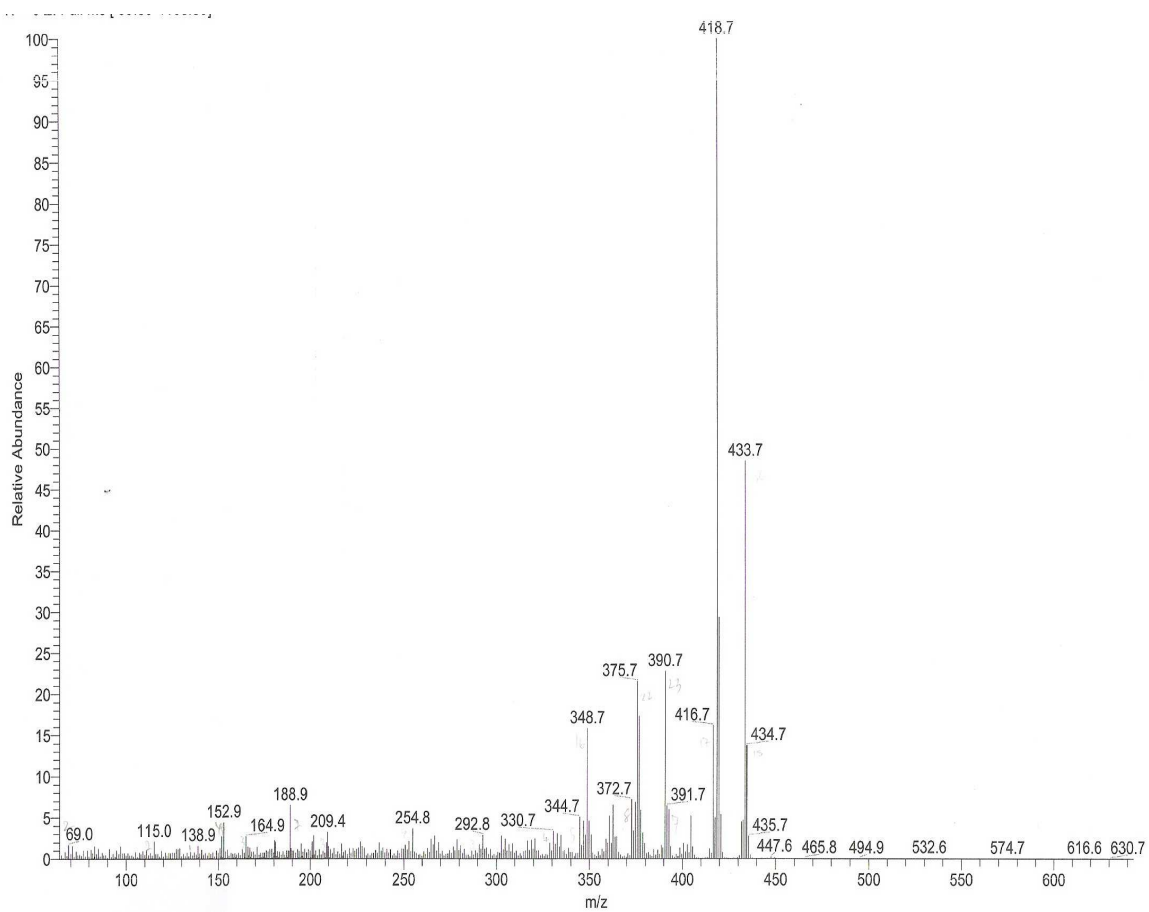


Figure 52 EI-MS spectrum of **AE7**

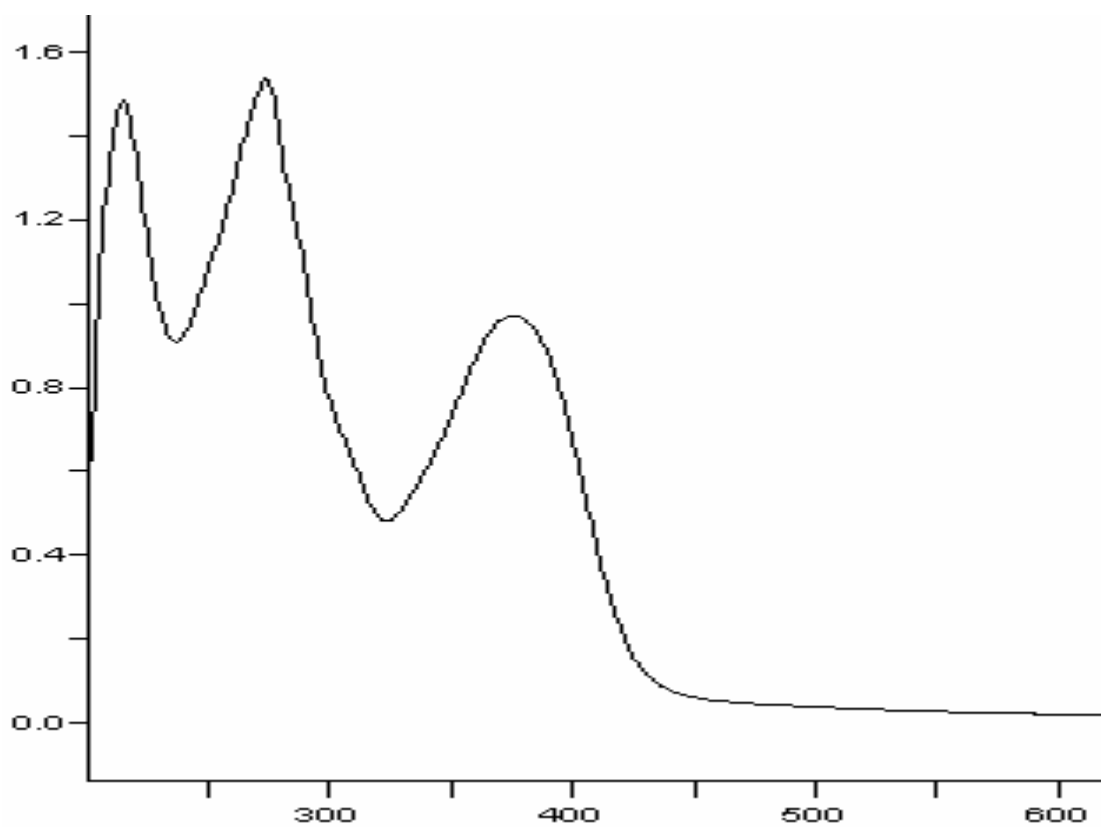


Figure 53 UV (EtOH) spectrum of AE7

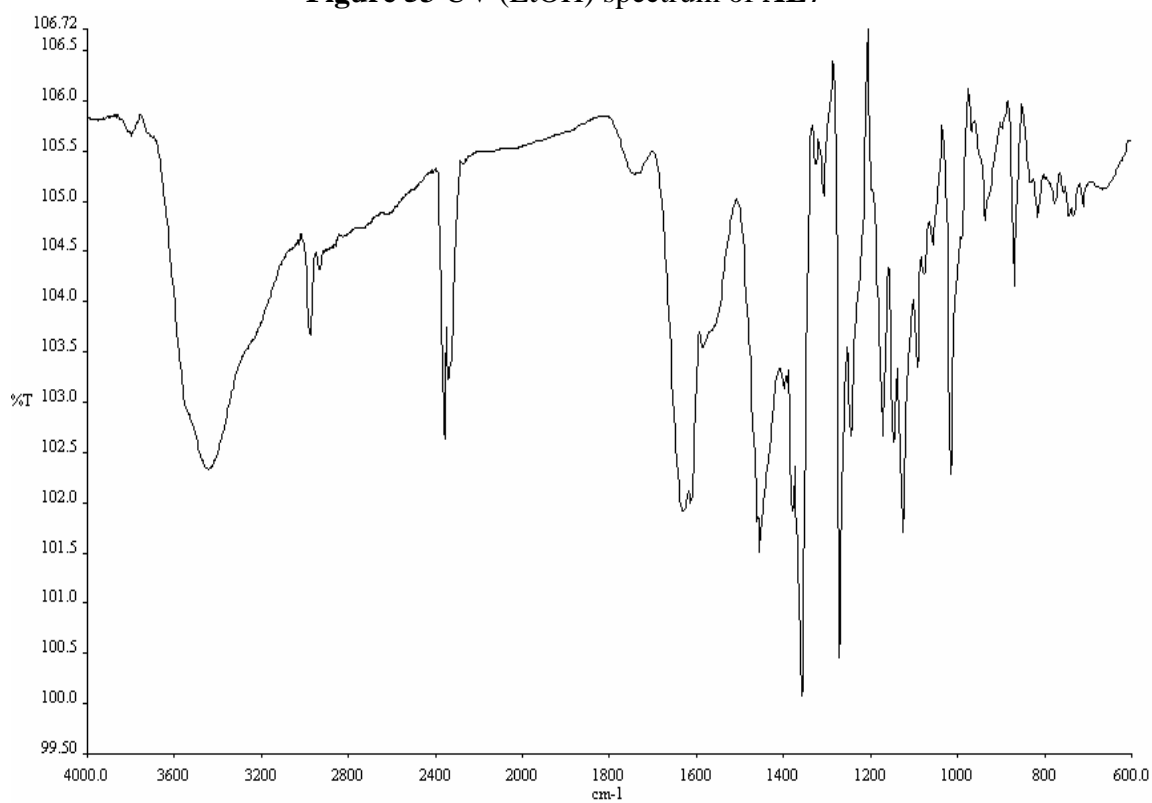


Figure 54 FT-IR (Neat) spectrum of AE7

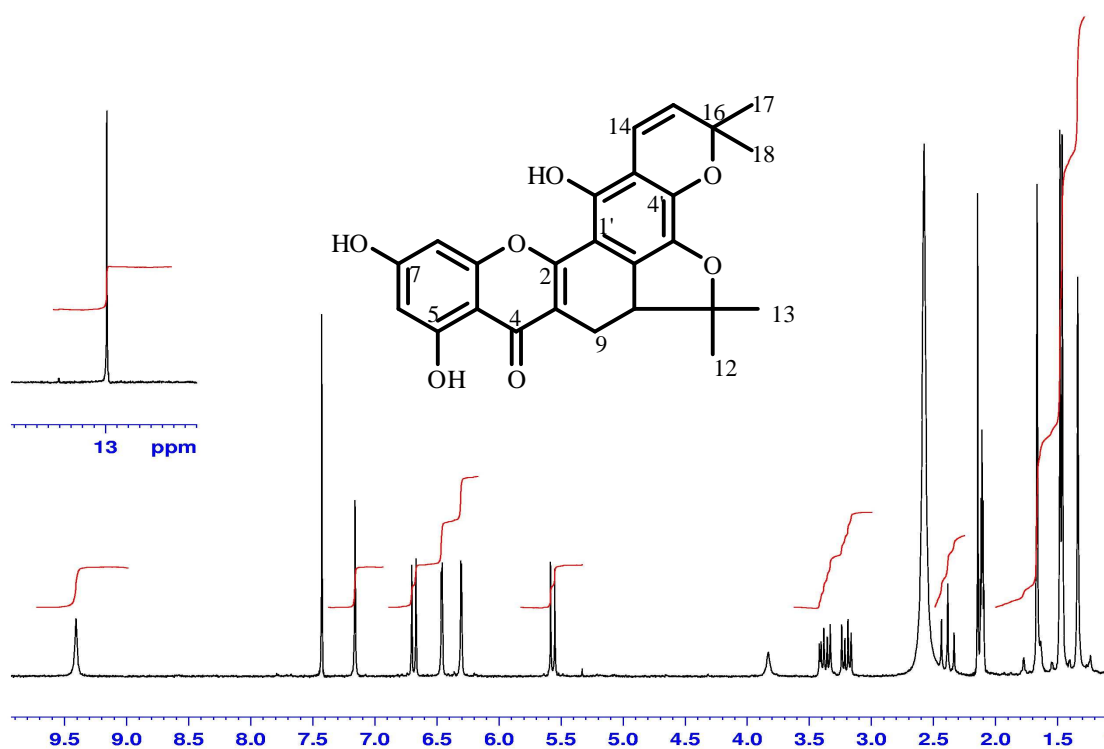


Figure 55 ^1H NMR (300 MHz) ($\text{CDCl}_3 + \text{Acetone-}d_6$) spectrum of AE7

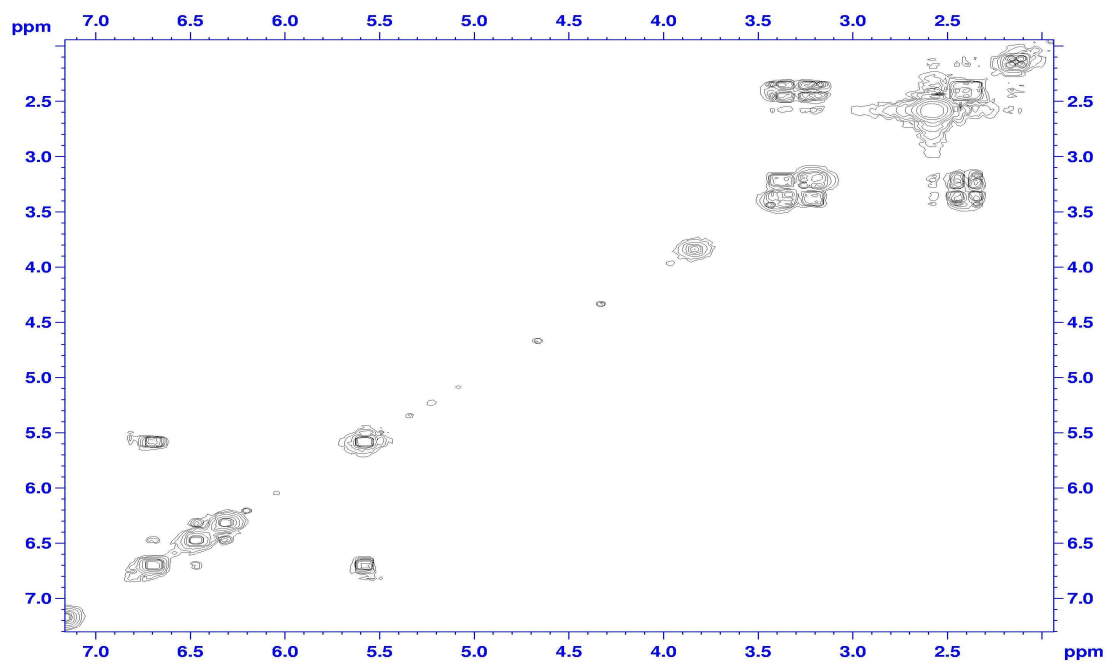


Figure 56 ^1H - ^1H COSY spectrum of AE7

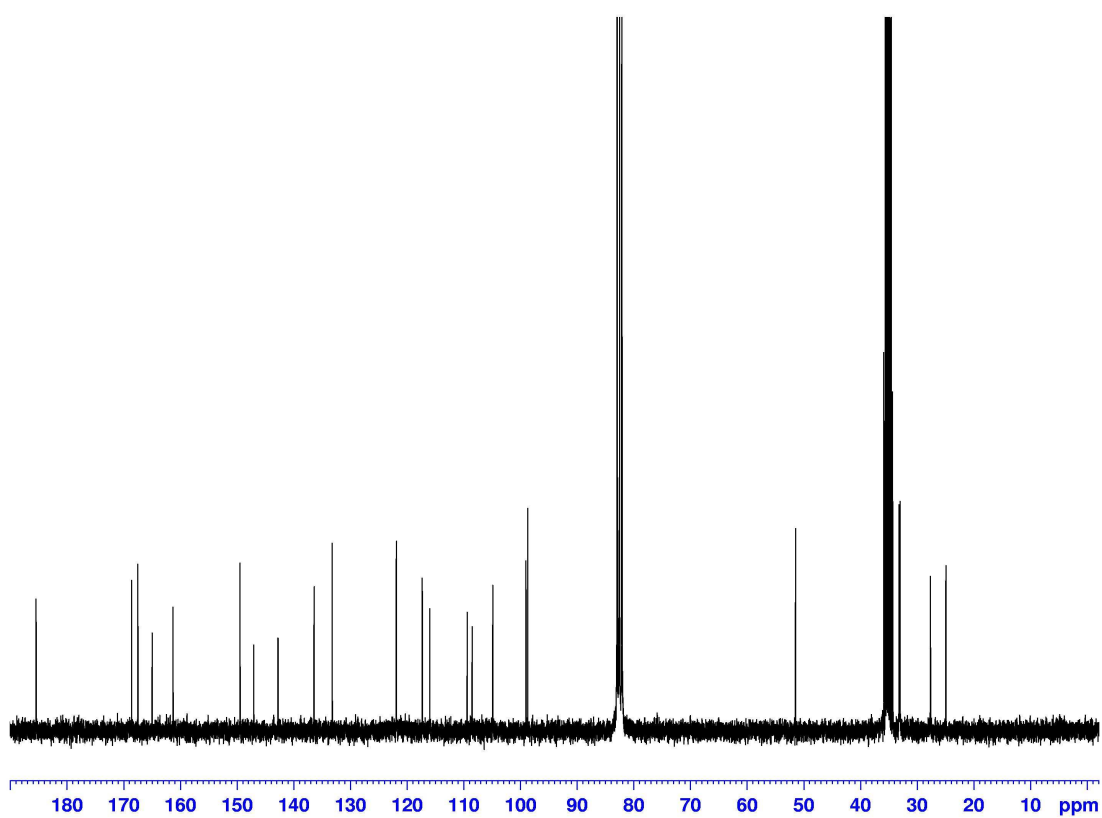


Figure 57 ^{13}C NMR (75 MHz) (CDCl_3 +Acetone- d_6) spectrum of **AE7**

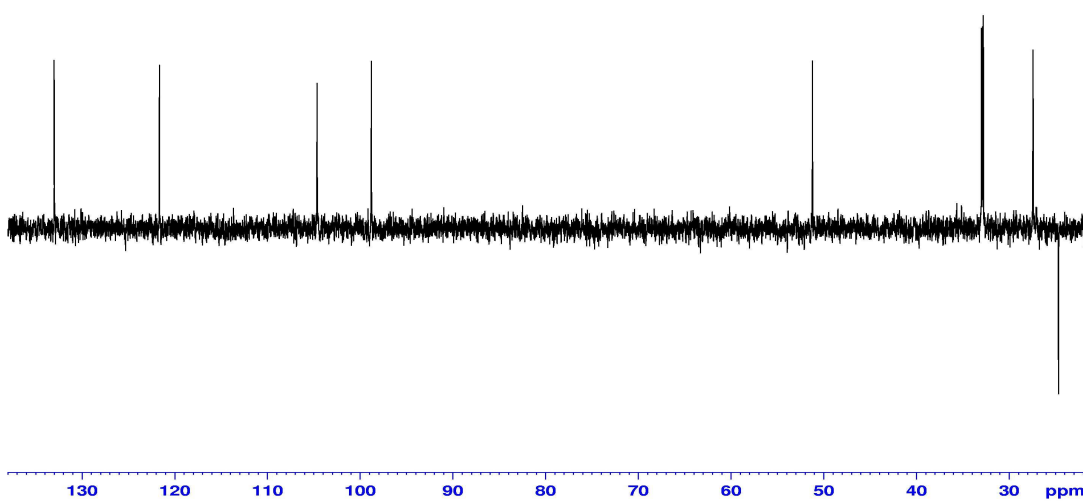


Figure 58 DEPT 135° (CDCl_3 +Acetone- d_6) spectrum of **AE7**

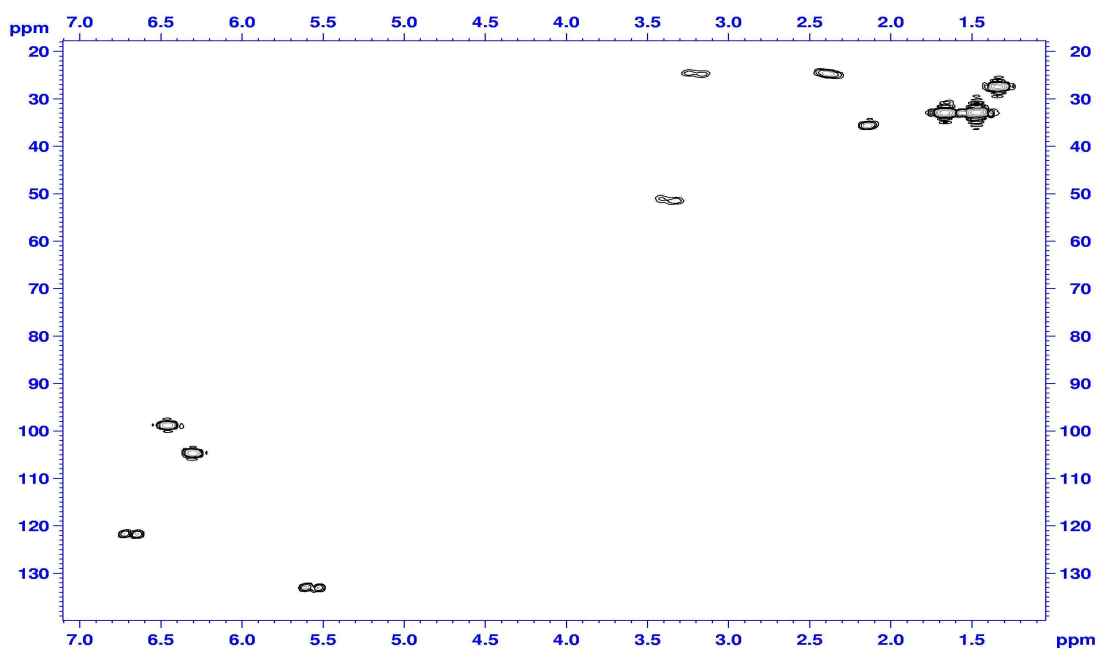


Figure 59 2D HMQC spectrum of AE7

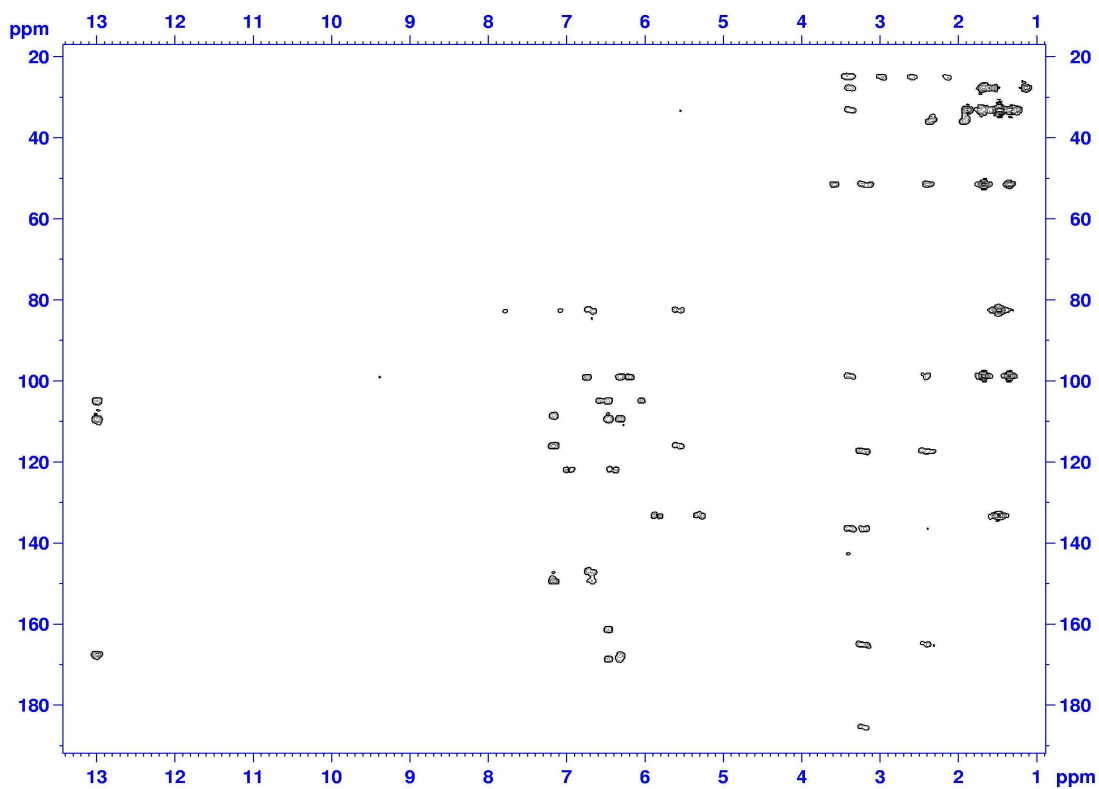


Figure 60 2D HMBC spectrum of AE7

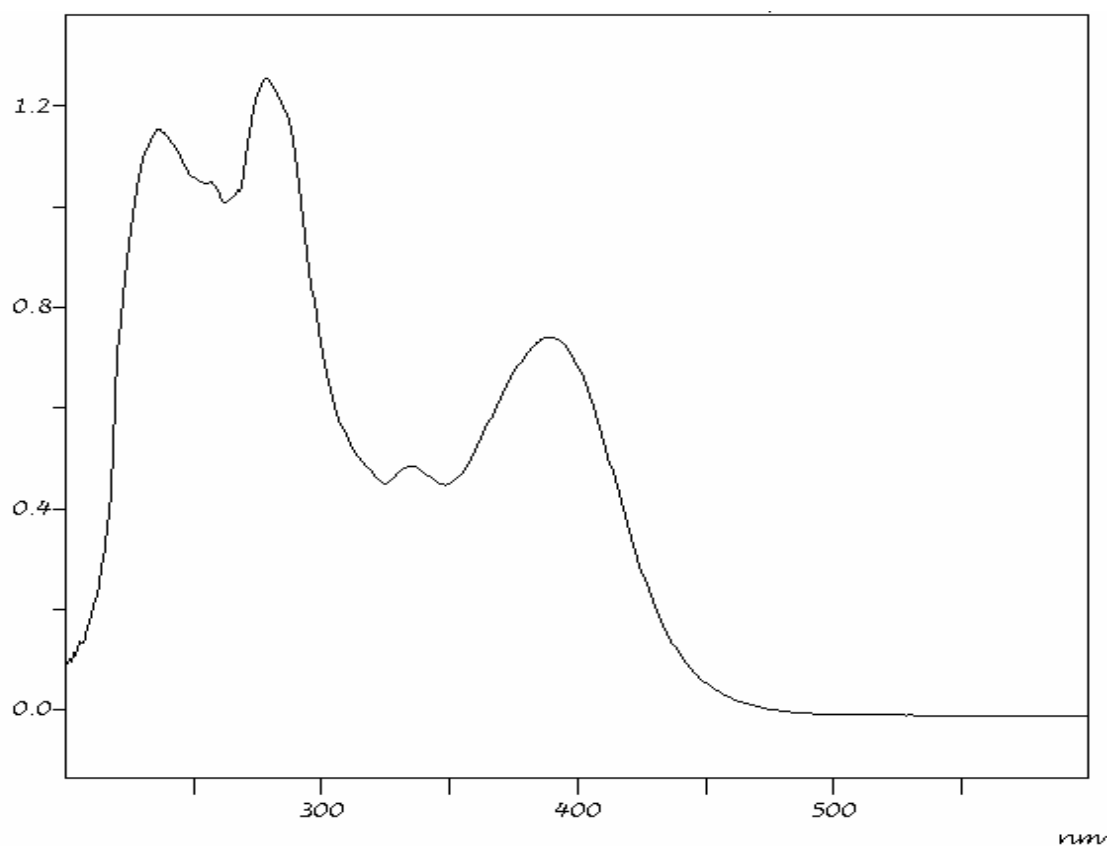


Figure 61 UV (CH₃OH) spectrum of AE8

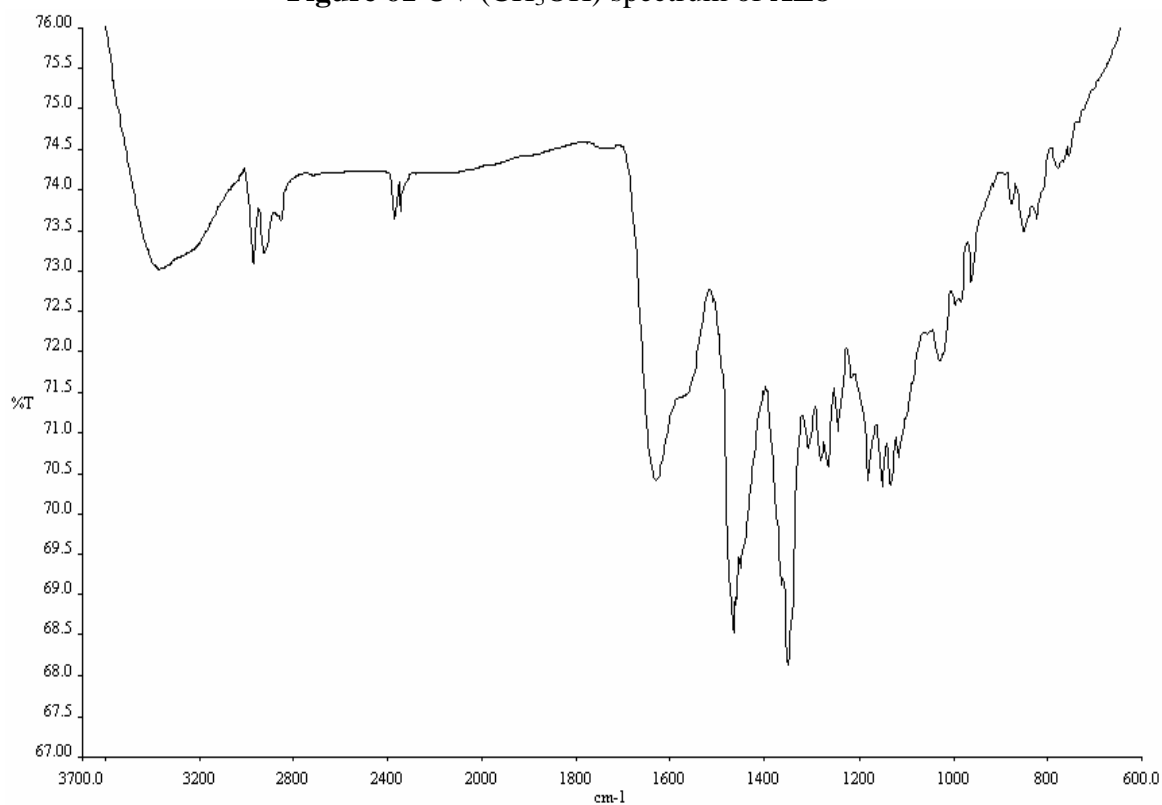


Figure 62 FT-IR (Neat) spectrum of AE8

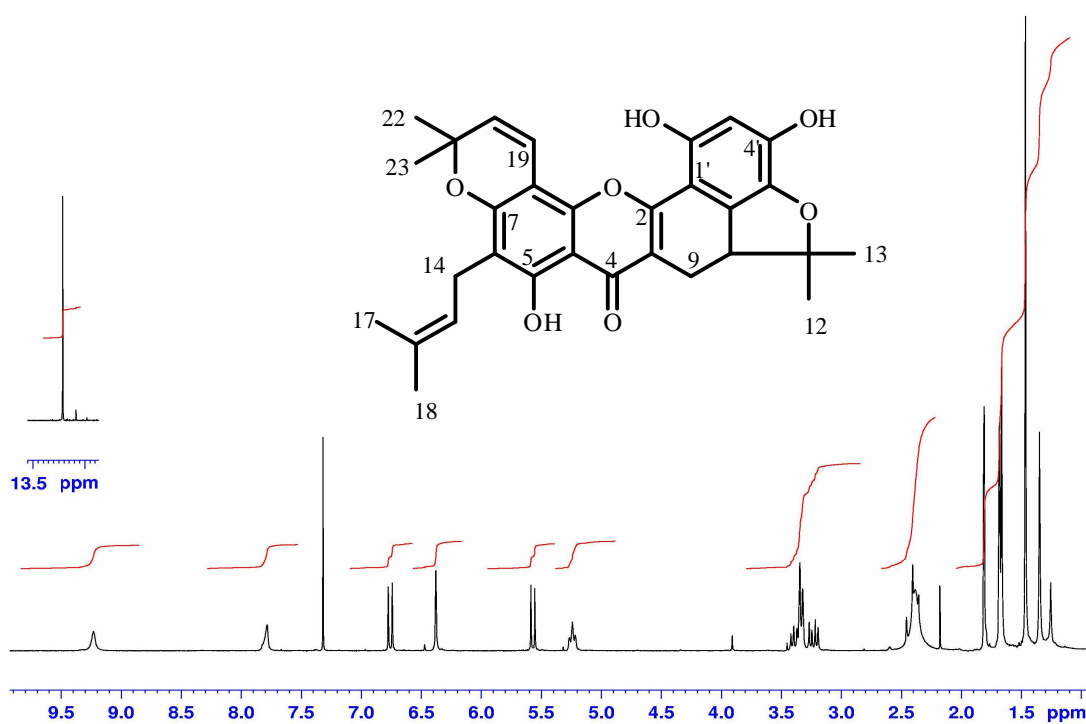


Figure 63 ^1H NMR (300 MHz) ($\text{CDCl}_3 + \text{DMSO-}d_6$) spectrum of AE8

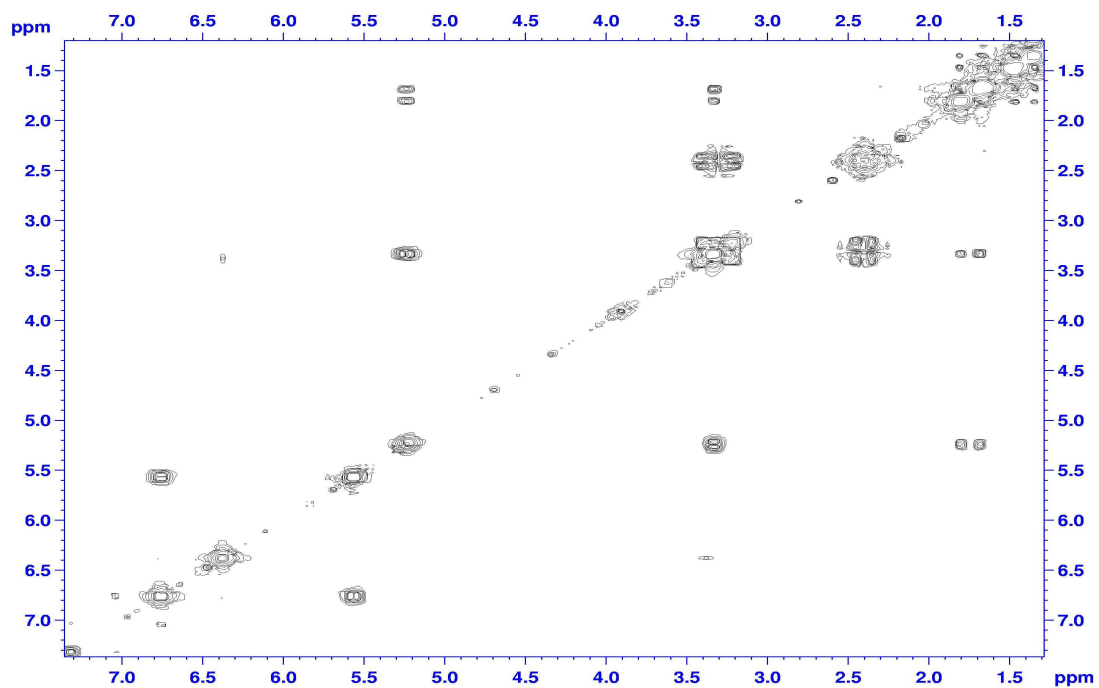


Figure 64 ^1H - ^1H COSY spectrum of AE8

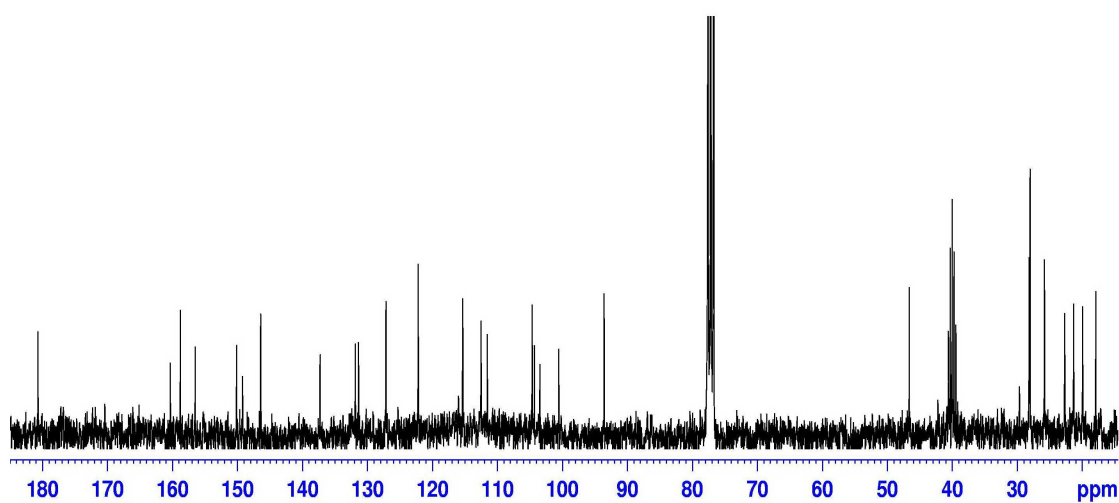


Figure 65 ^{13}C NMR (75 MHz) ($\text{CDCl}_3 + \text{DMSO-}d_6$) spectrum of **AE8**

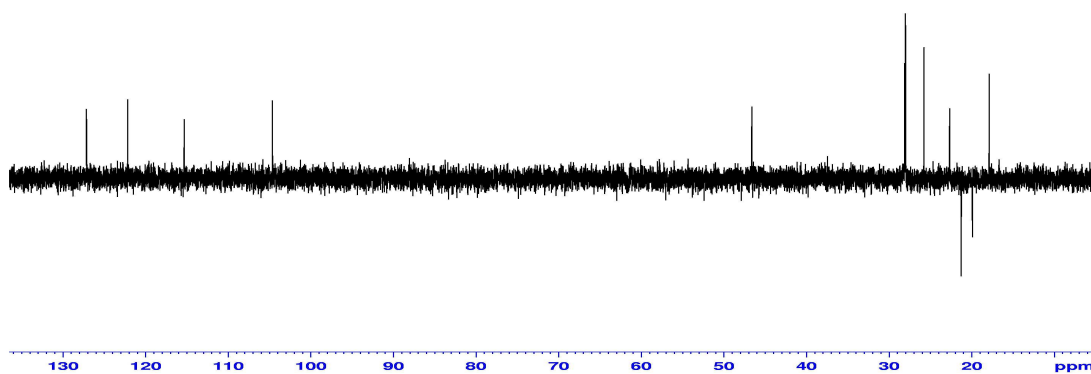


Figure 66 DEPT 135° ($\text{CDCl}_3 + \text{DMSO-}d_6$) spectrum of **AE8**

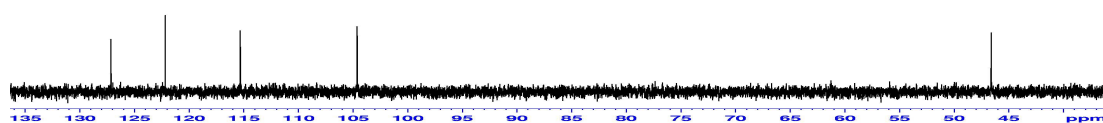


Figure 67 DEPT 90° ($\text{CDCl}_3 + \text{DMSO-}d_6$) spectrum of **AE8**

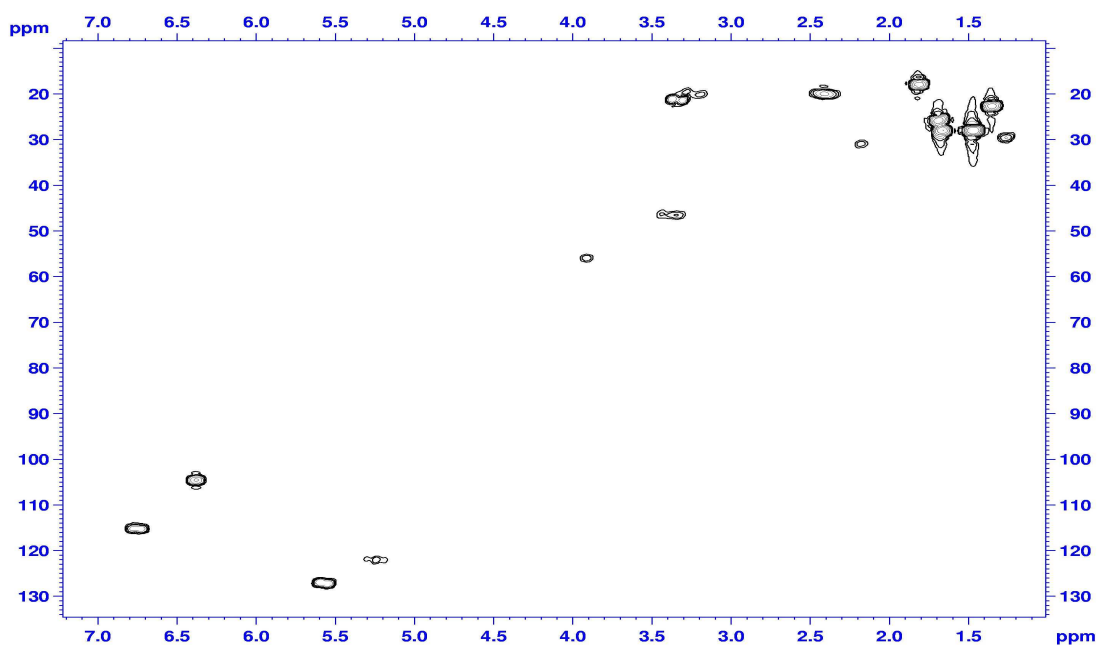


Figure 68 2D HMQC spectrum of AE8

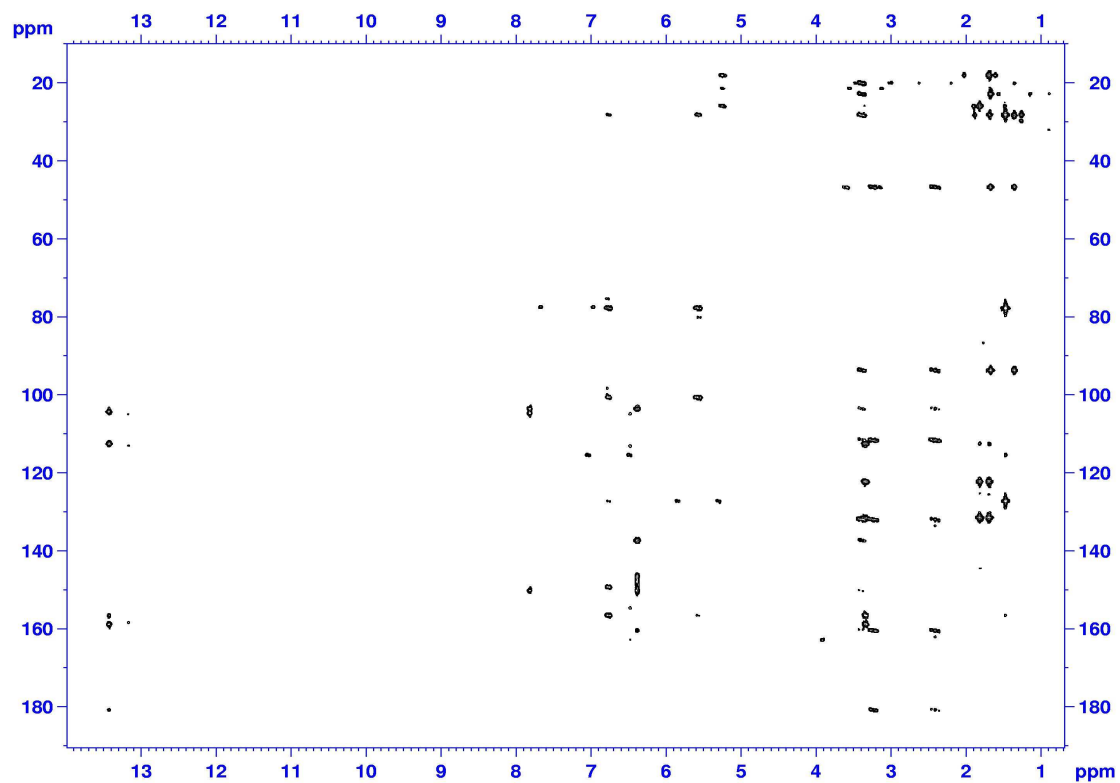


Figure 69 2D HMBC spectrum of AE8

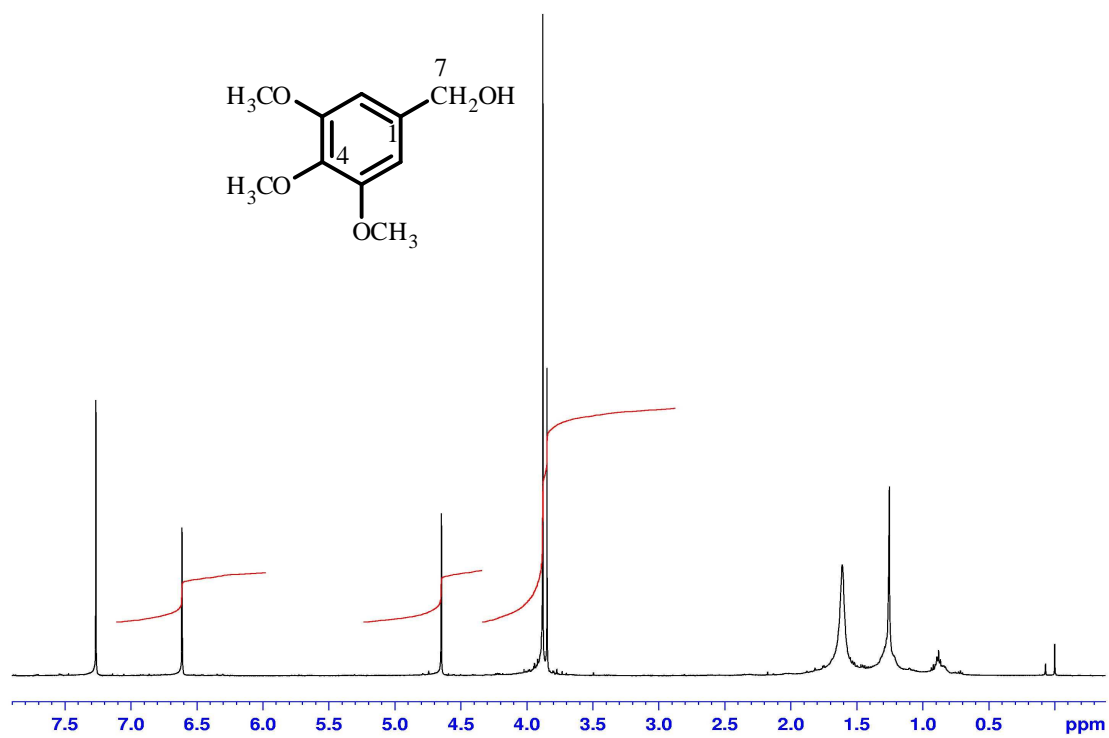


Figure 70 ^1H NMR (500 MHz) (CDCl_3) spectrum of **AE9**

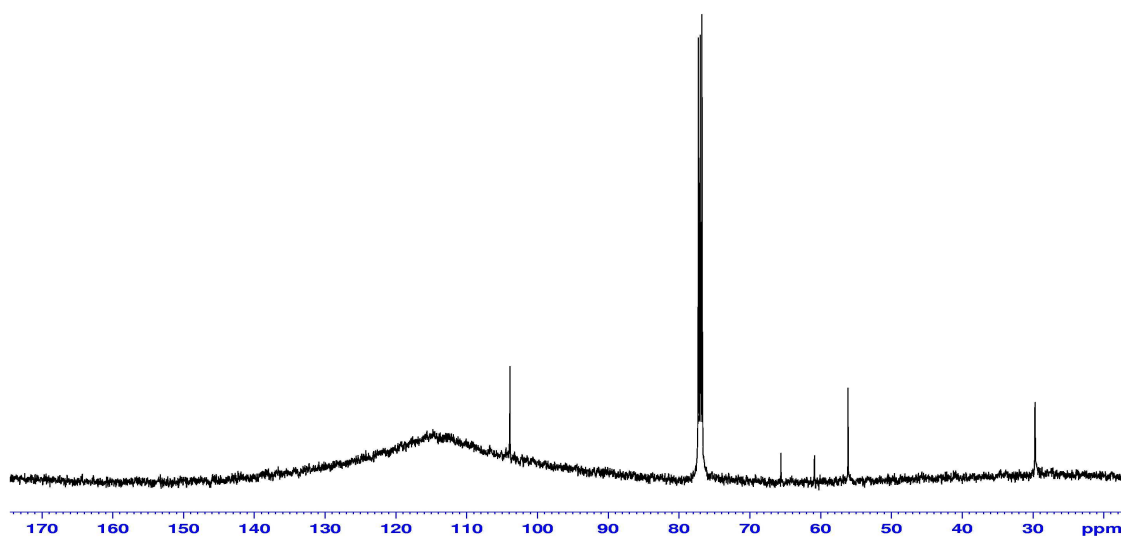


Figure 71 ^{13}C NMR (125 MHz) (CDCl_3) spectrum of AE9

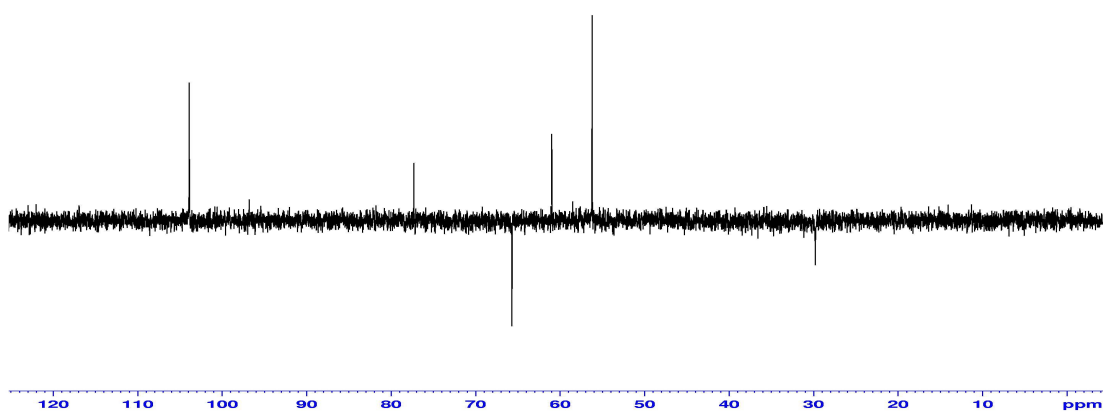


Figure 72 DEPT 135° (CDCl_3) spectrum of AE9

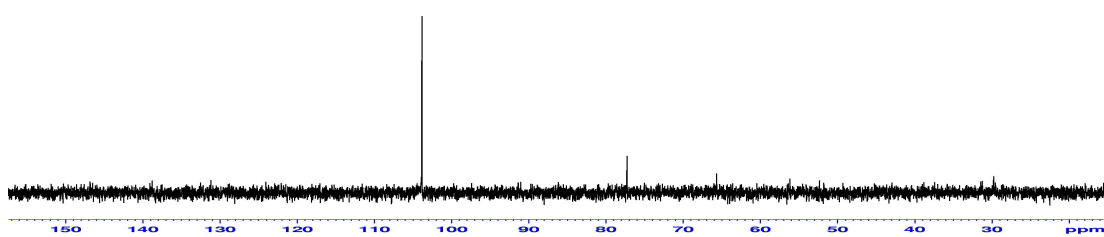


Figure 73 DEPT 90° (CDCl_3) spectrum of AE9

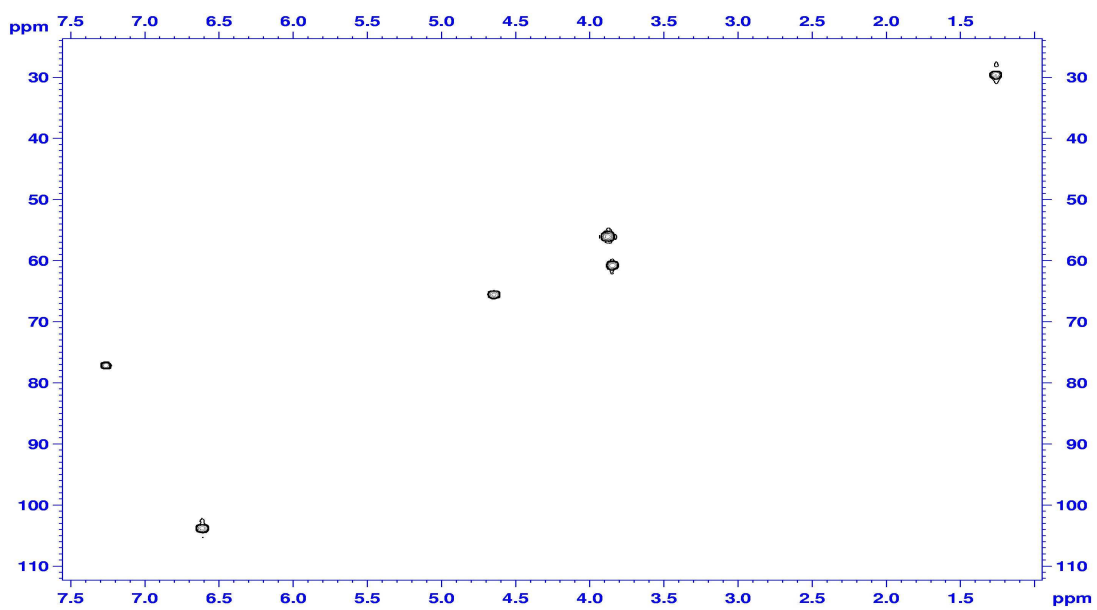


Figure 74 2D HMQC spectrum of AE9

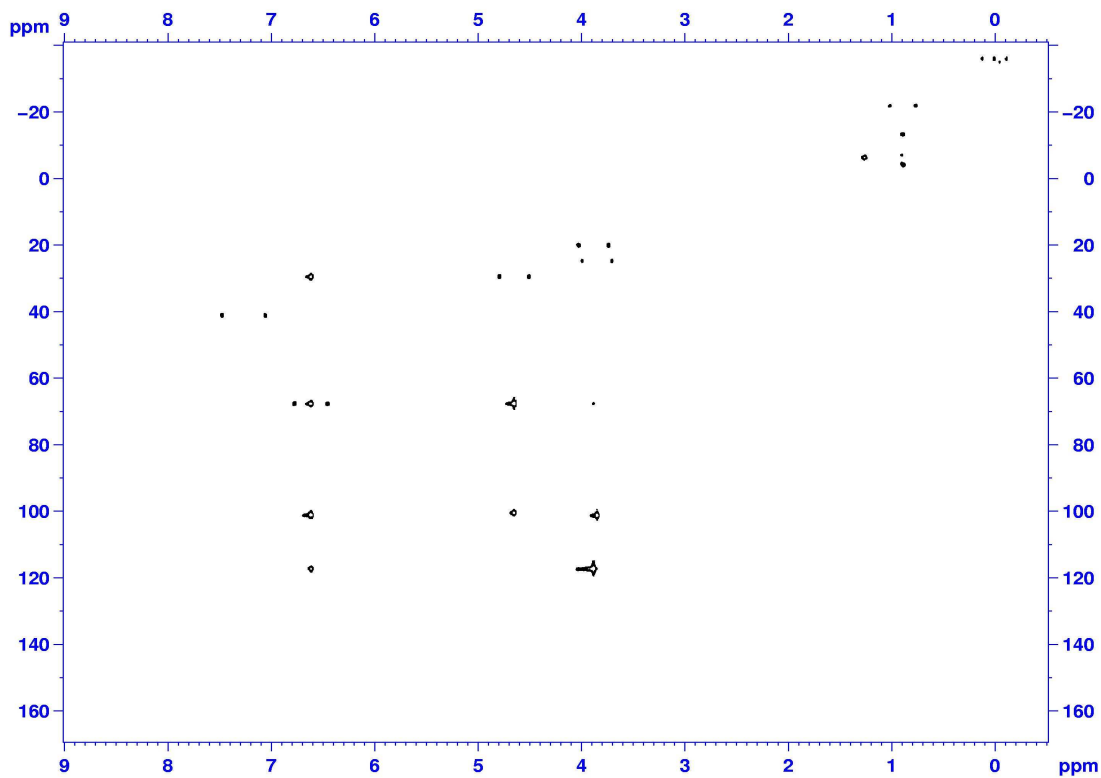


Figure 75 2D HMBC spectrum of AE9

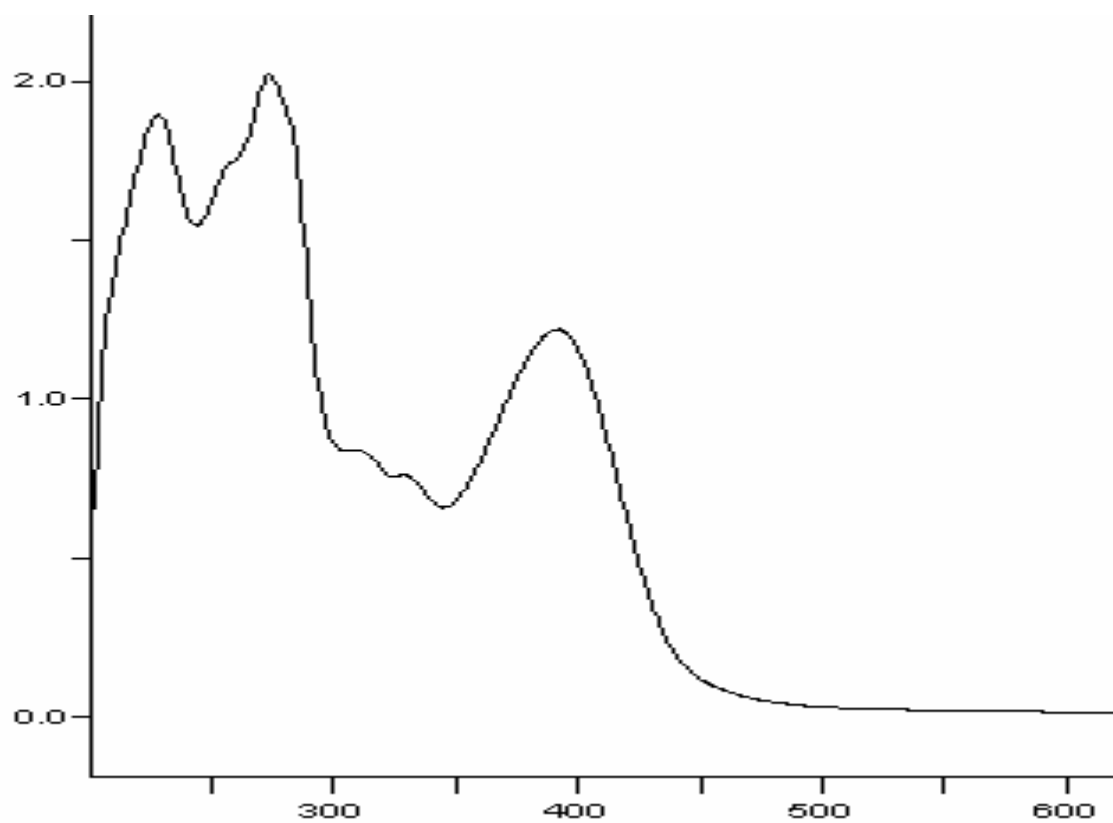


Figure 76 UV (EtOH) spectrum of AE10

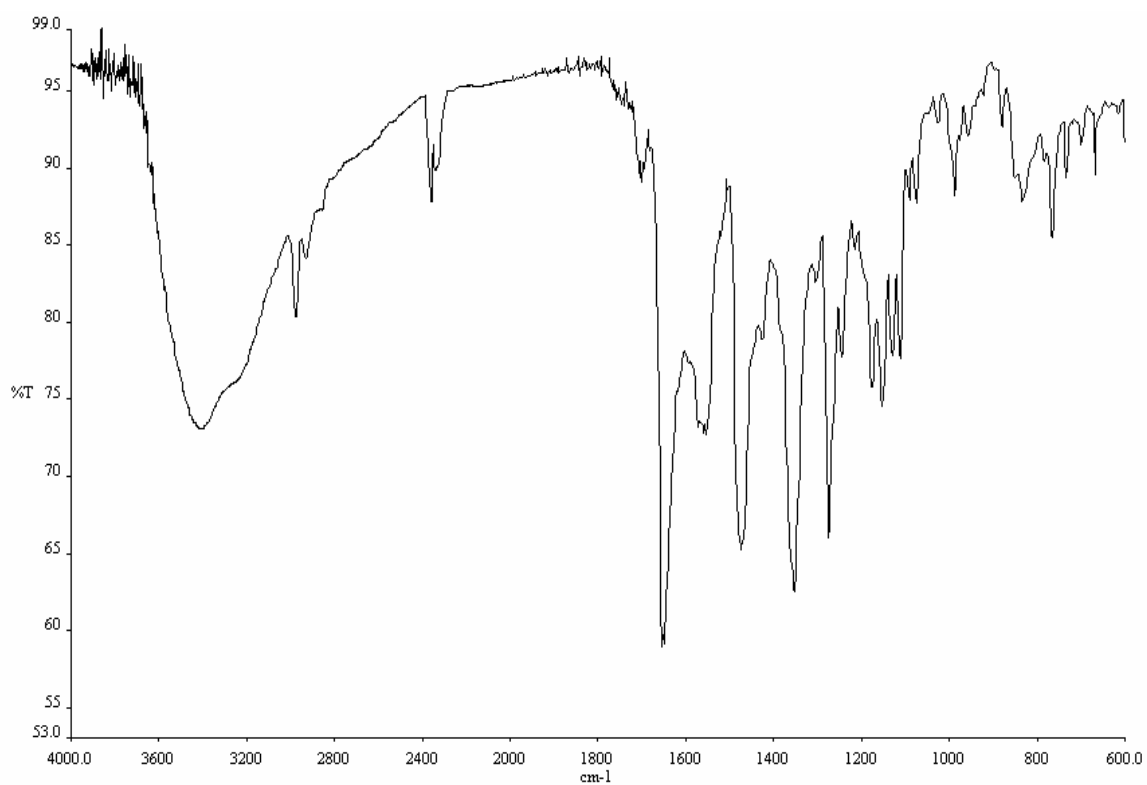


Figure 77 FT-IR (Neat) spectrum of AE10

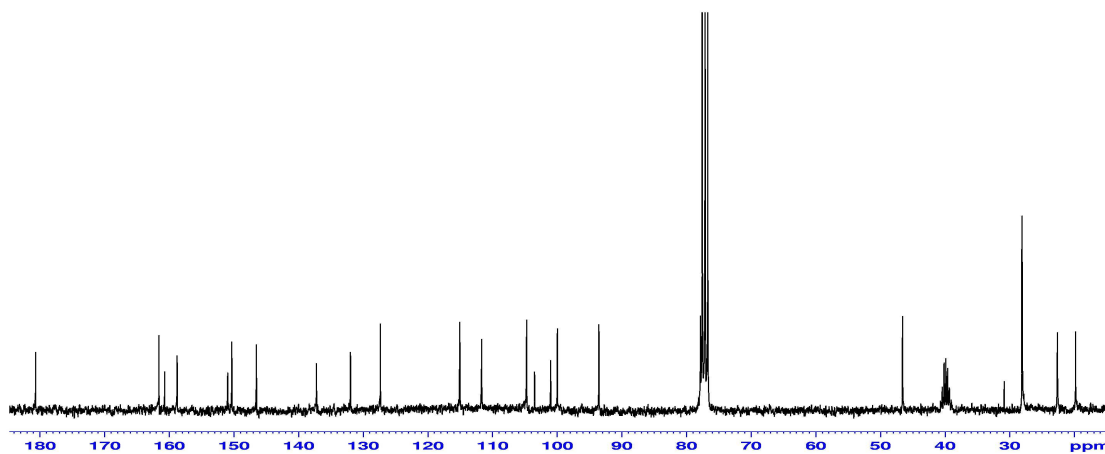


Figure 80 ^{13}C NMR (75 MHz) ($\text{CDCl}_3+\text{DMSO}-d_6$) spectrum of **AE10**

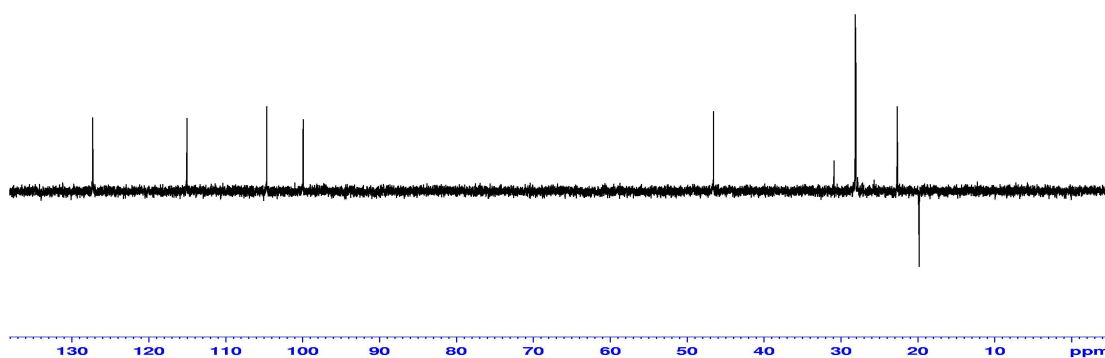


Figure 81 DEPT 135° ($\text{CDCl}_3+\text{DMSO}-d_6$) spectrum of **AE10**

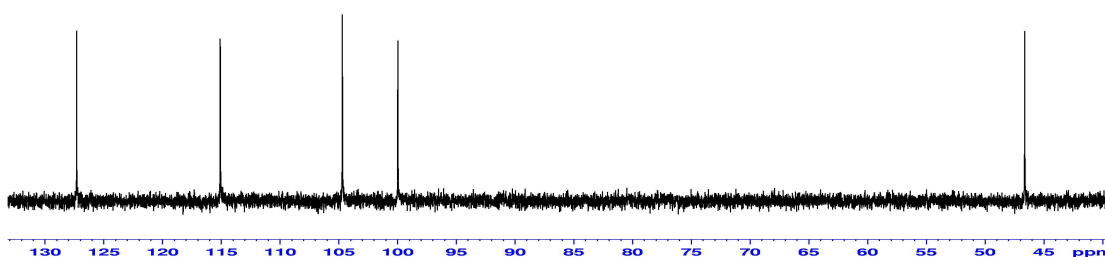


Figure 82 DEPT 90° ($\text{CDCl}_3+\text{DMSO}-d_6$) spectrum of **AE10**

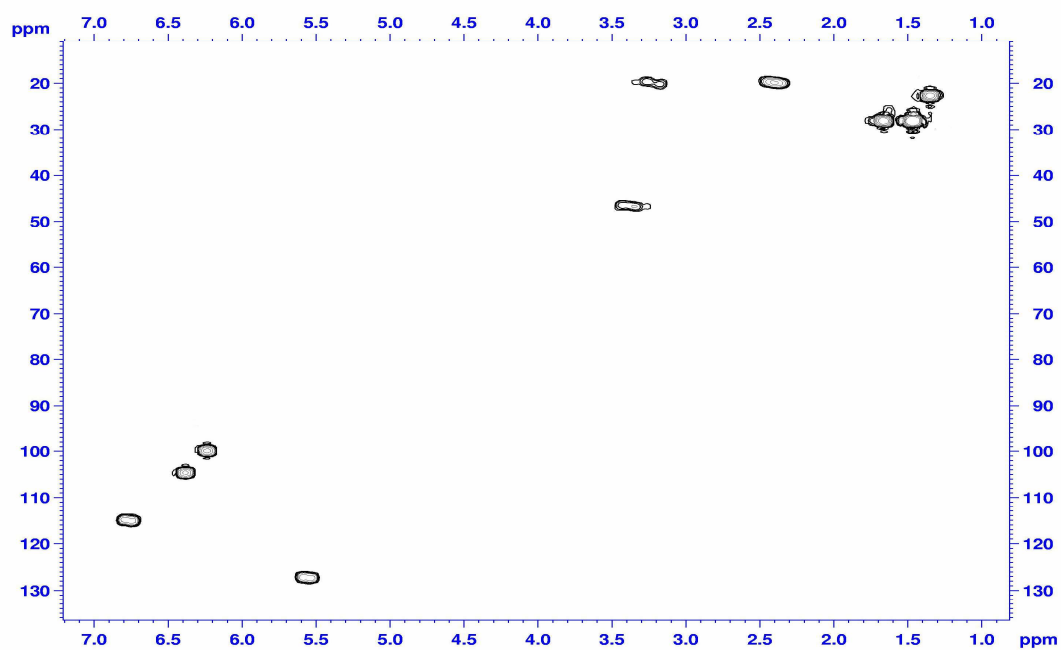


Figure 83 2D HMQC spectrum of AE10

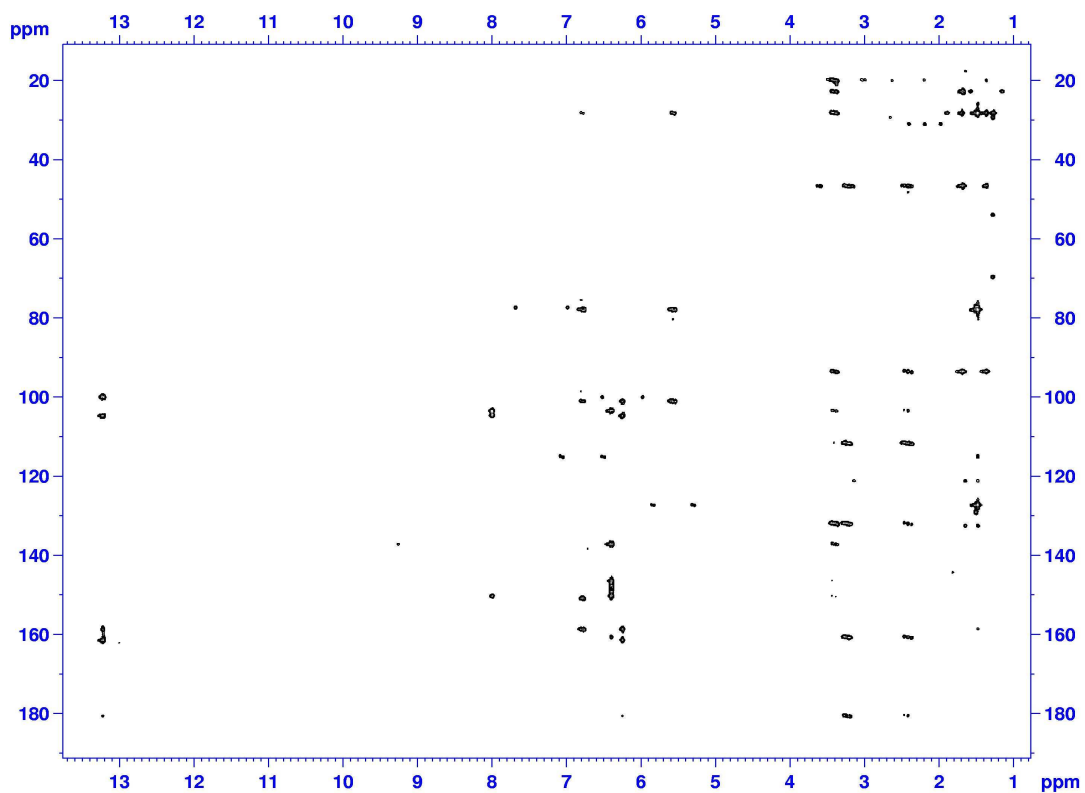


Figure 84 2D HMBC spectrum of AE10

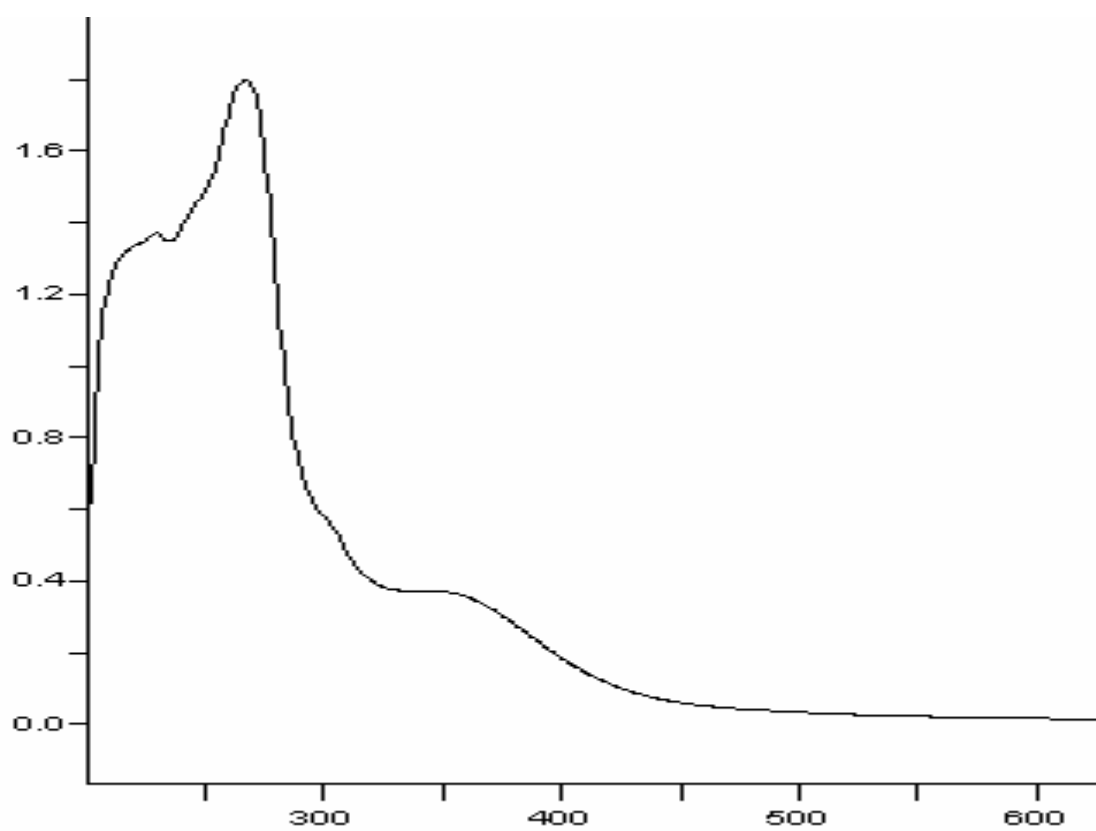


Figure 85 UV (EtOH) spectrum of **AE11**

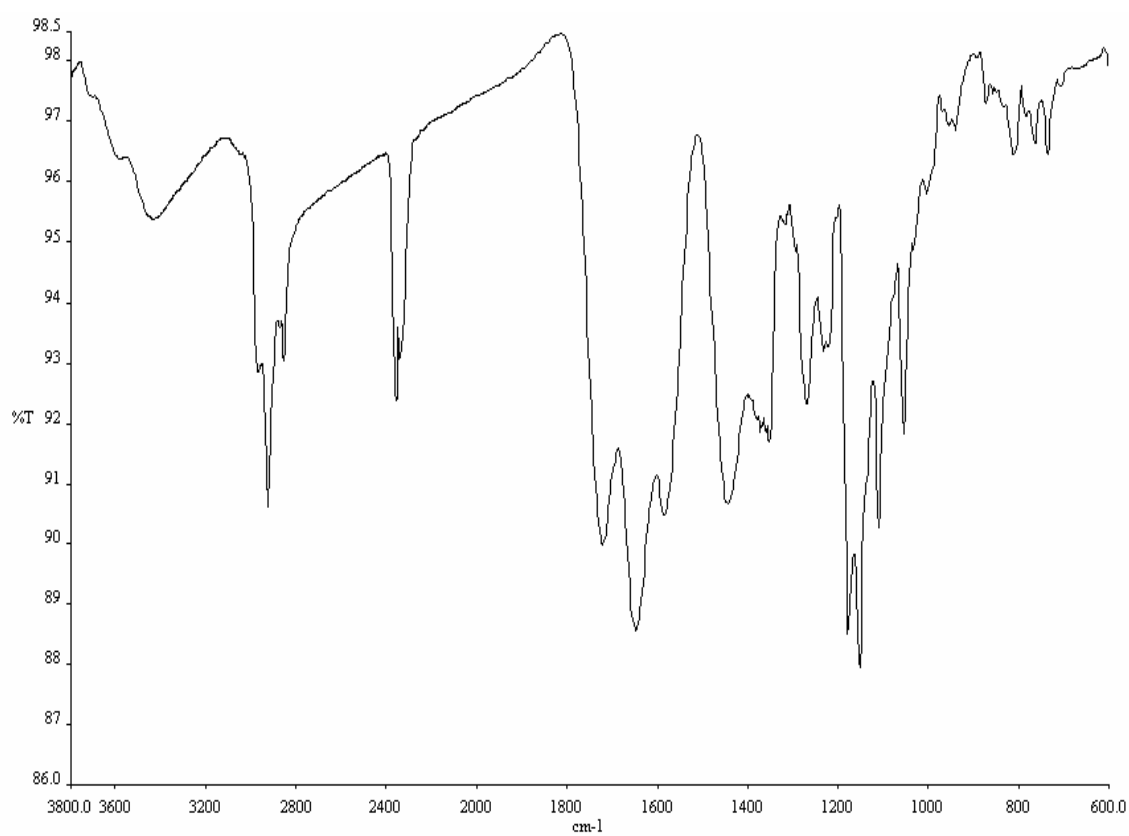
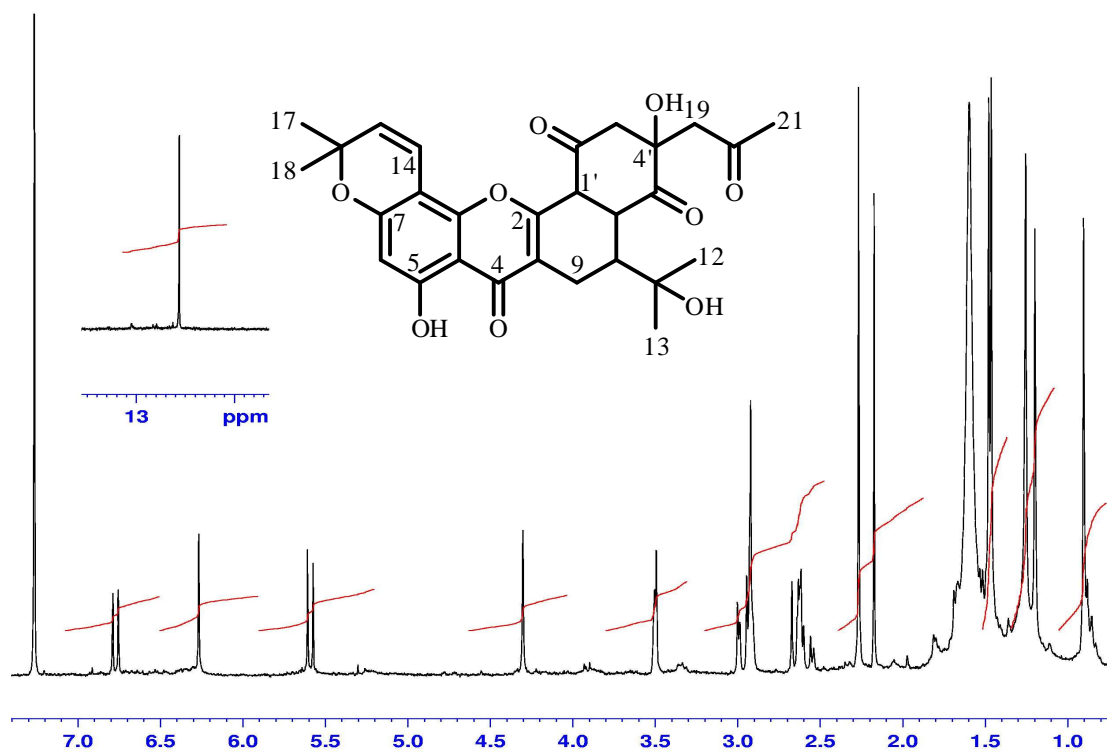
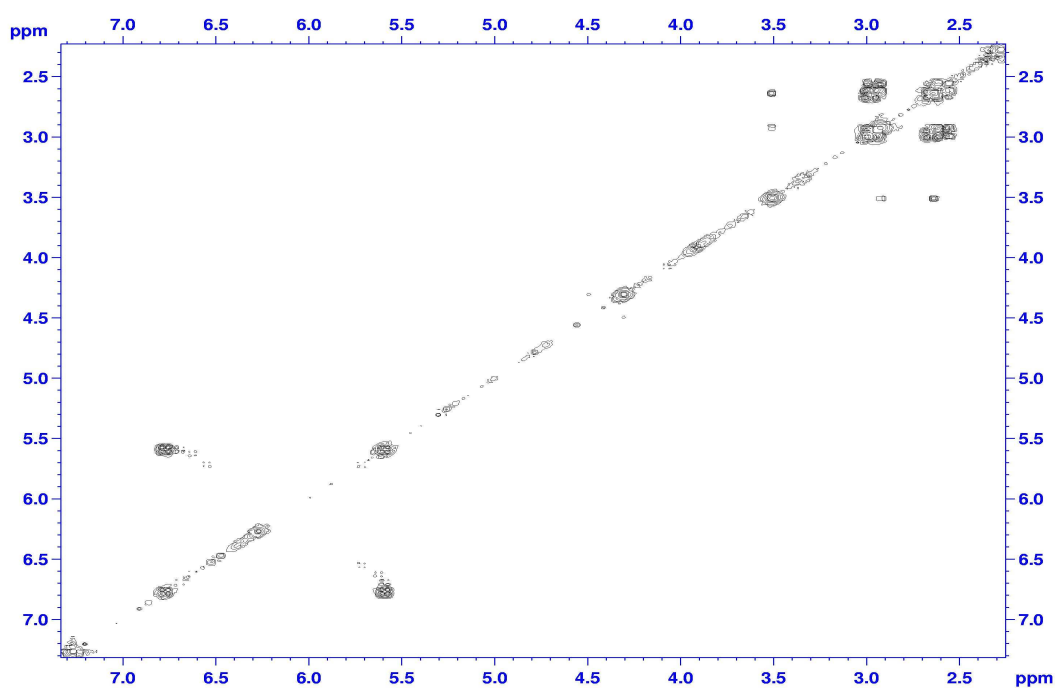


Figure 86 FT-IR (Neat) spectrum of **AE11****Figure 87** ¹H NMR (300 MHz) (CDCl₃) spectrum of **AE11****Figure 88** ¹H-¹H COSY spectrum of **AE11**

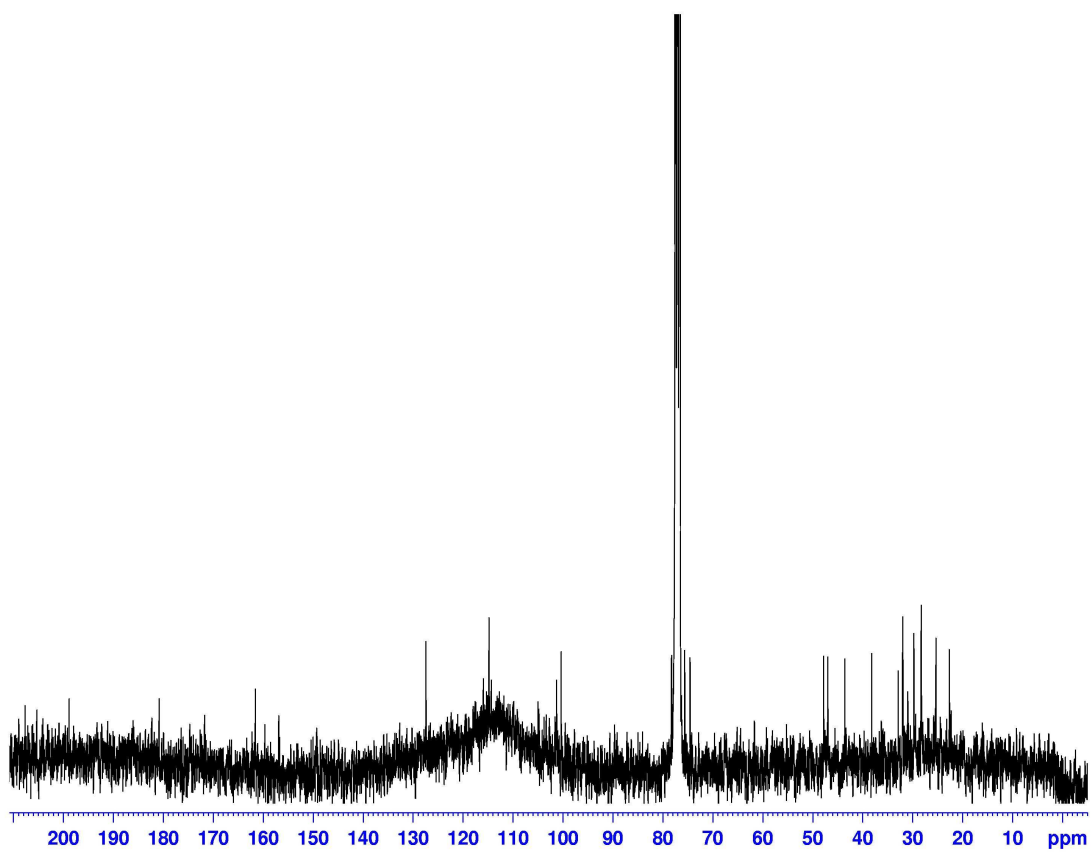


Figure 89 ^{13}C NMR (75 MHz) (CDCl_3) spectrum of AE11

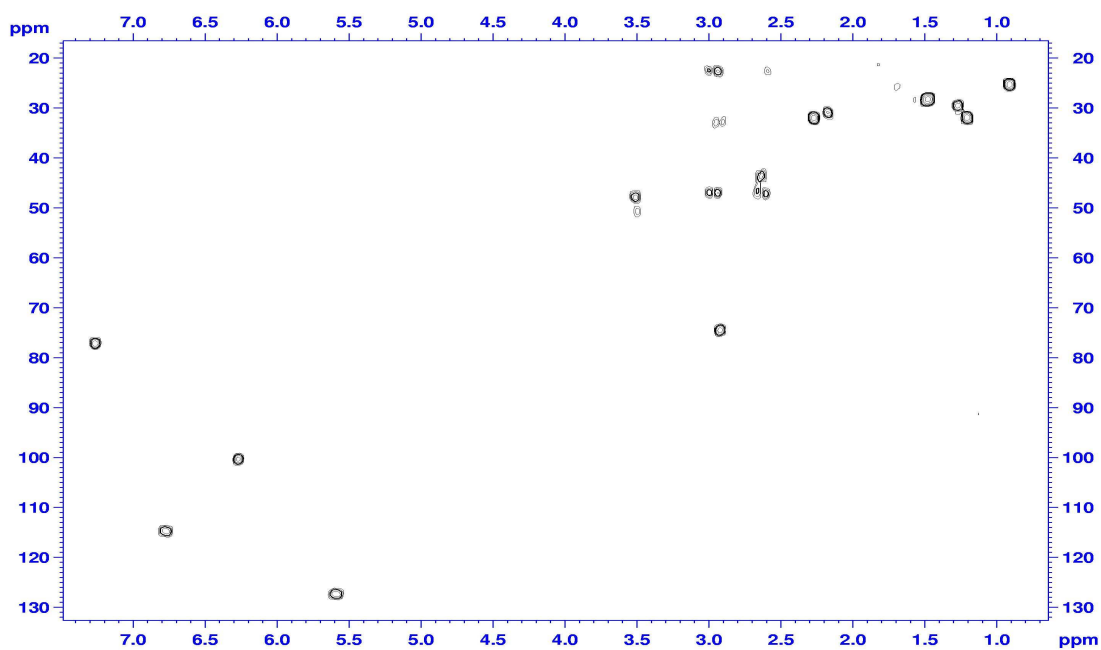


Figure 90 2D HMQC spectrum of AE11

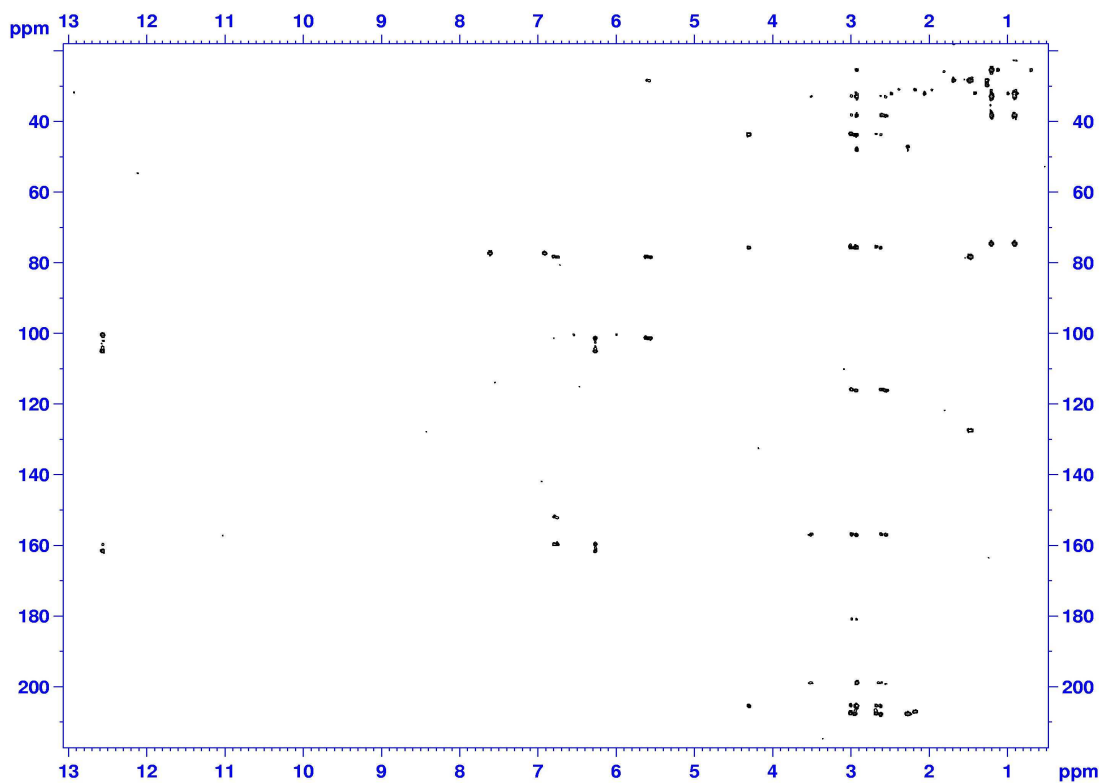


Figure 91 2D HMBC spectrum of AE11

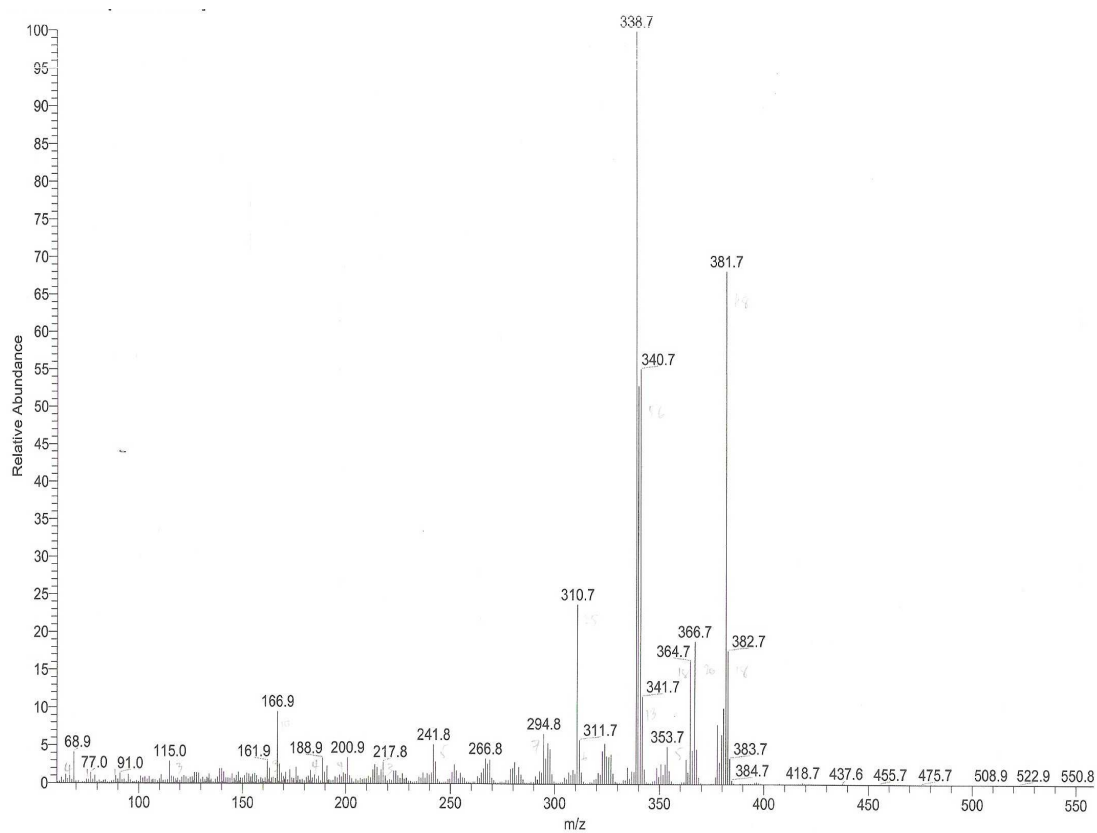


Figure 92 EI-MS spectrum of AE12

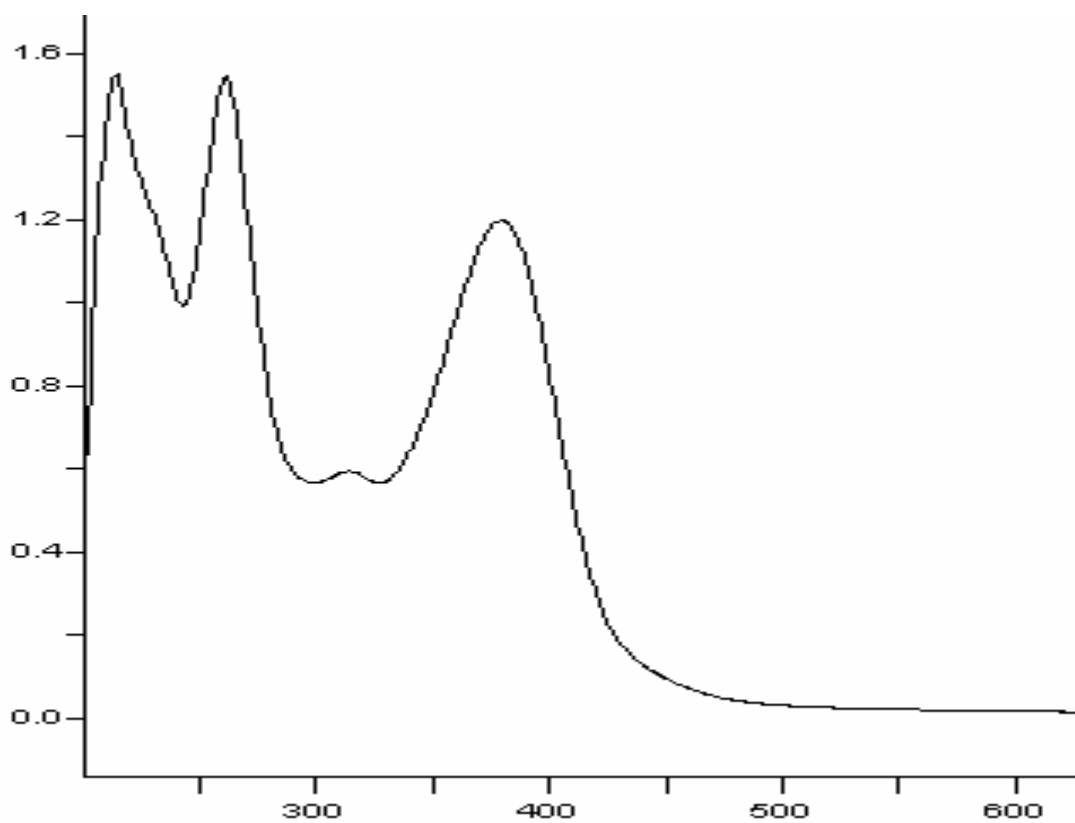


Figure 93 UV (EtOH) spectrum of AE12

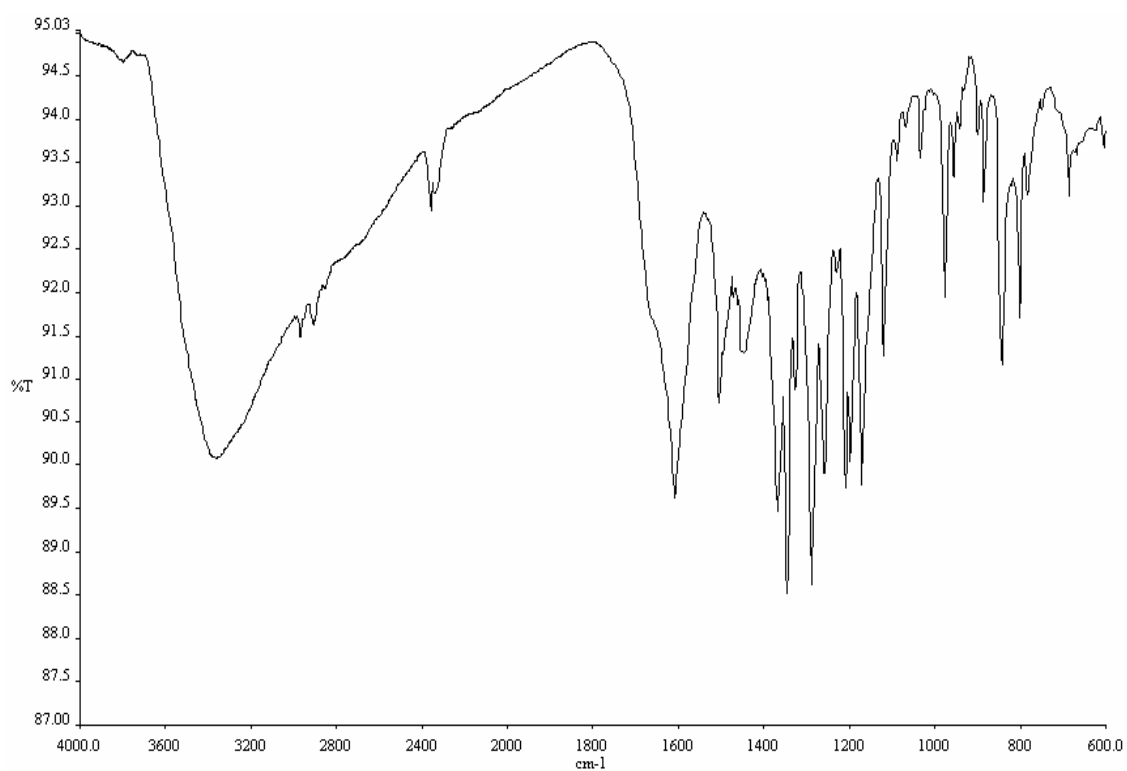


Figure 94 FT-IR (Neat) spectrum of AE12

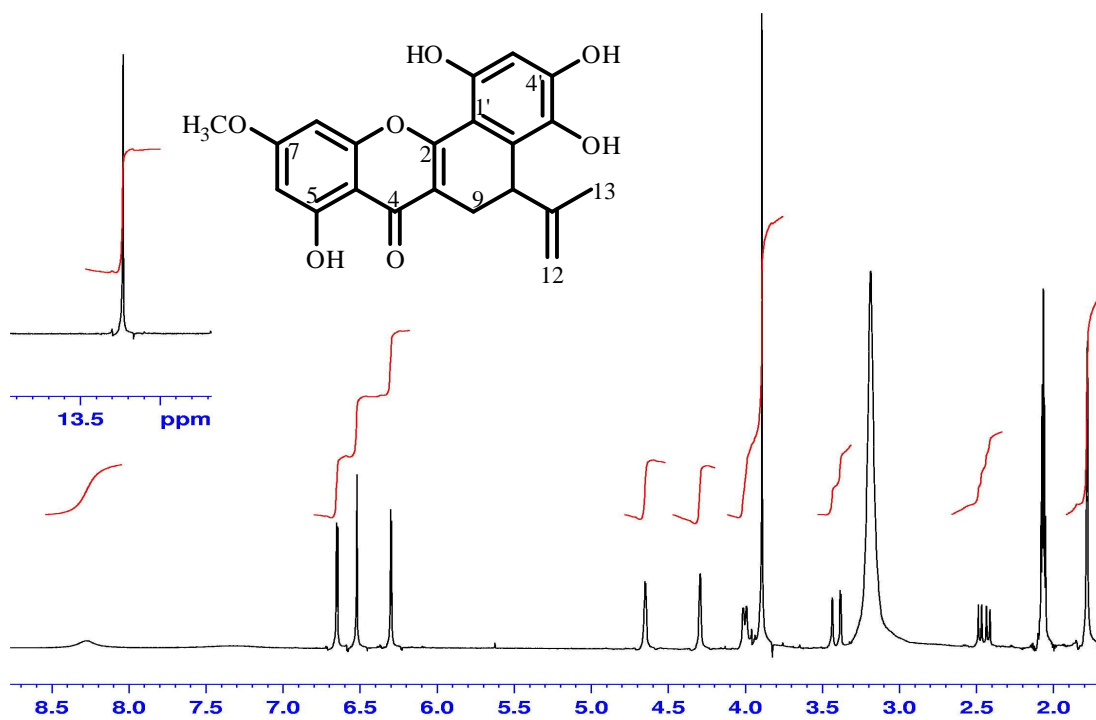


Figure 95 ^1H NMR (300 MHz) ($\text{Acetone-}d_6$) spectrum of AE12

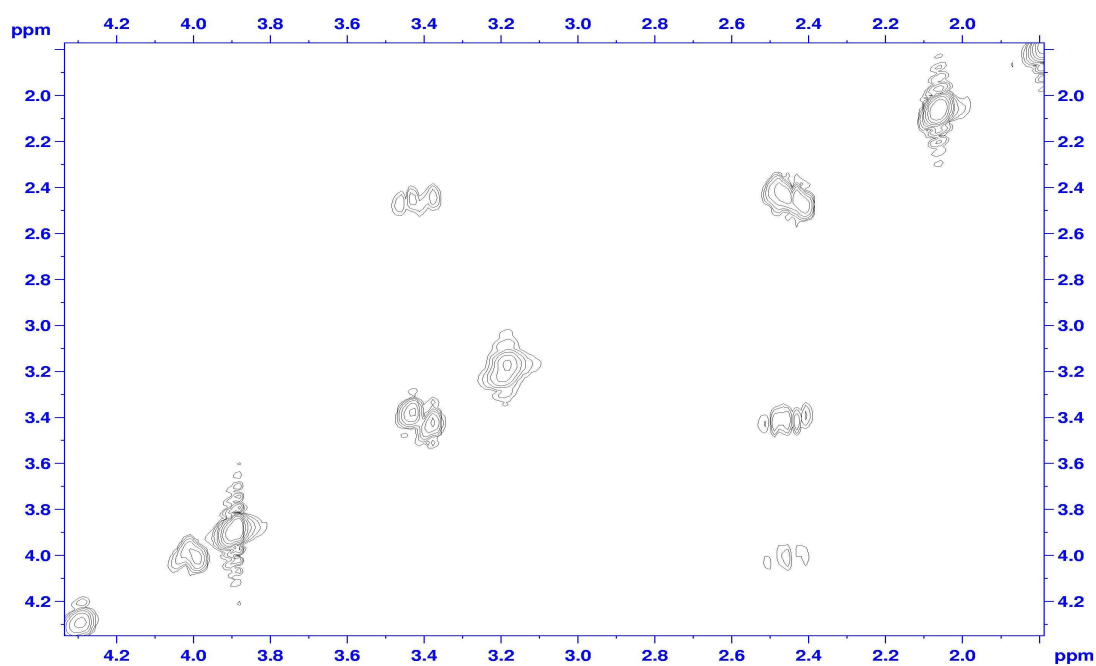


Figure 96 ^1H - ^1H COSY spectrum of AE12

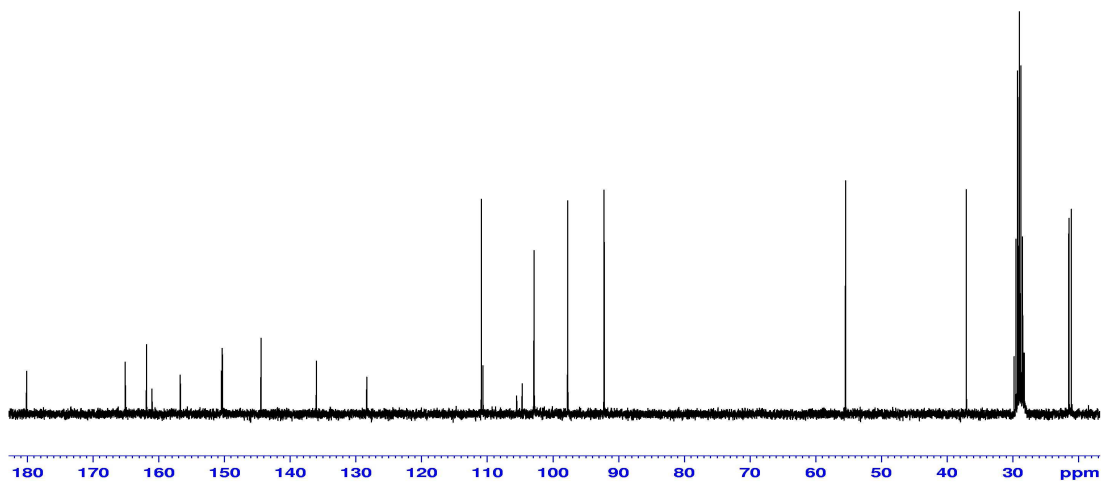


Figure 97 ^{13}C NMR (75 MHz) ($\text{Acetone-}d_6$) spectrum of AE12

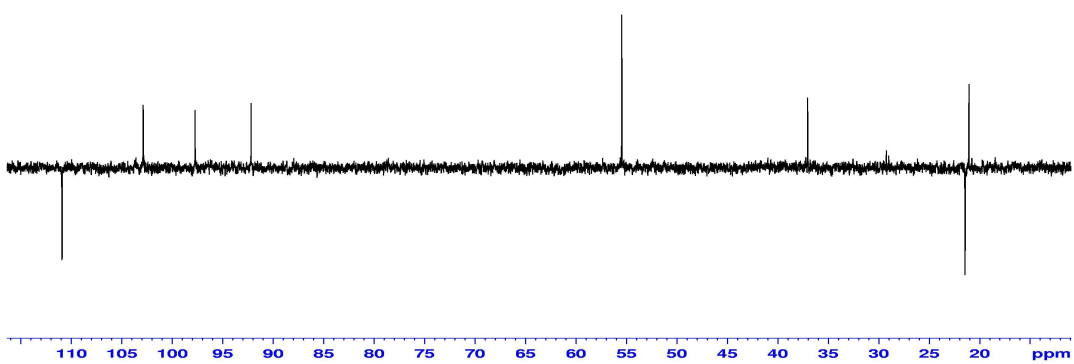


Figure 98 DEPT 135° ($\text{Acetone-}d_6$) spectrum of AE12

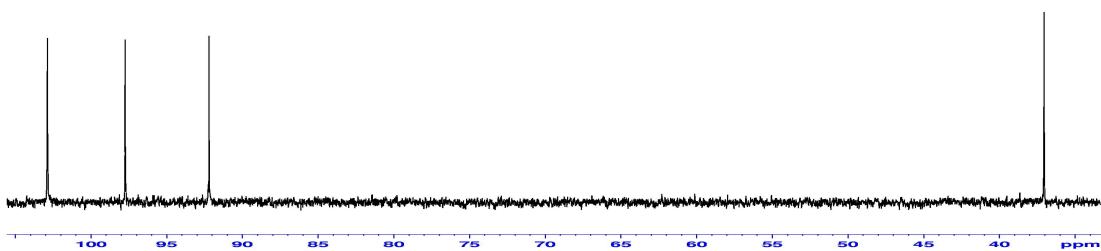


Figure 99 DEPT 90° ($\text{Acetone-}d_6$) spectrum of AE12

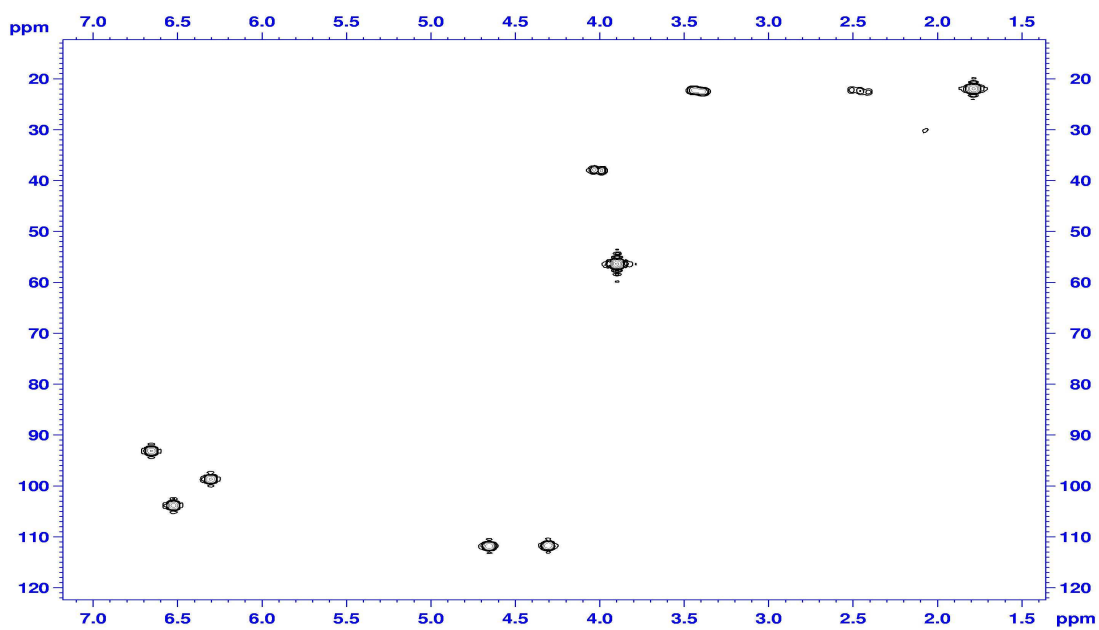


Figure 100 2D HMQC spectrum of AE12

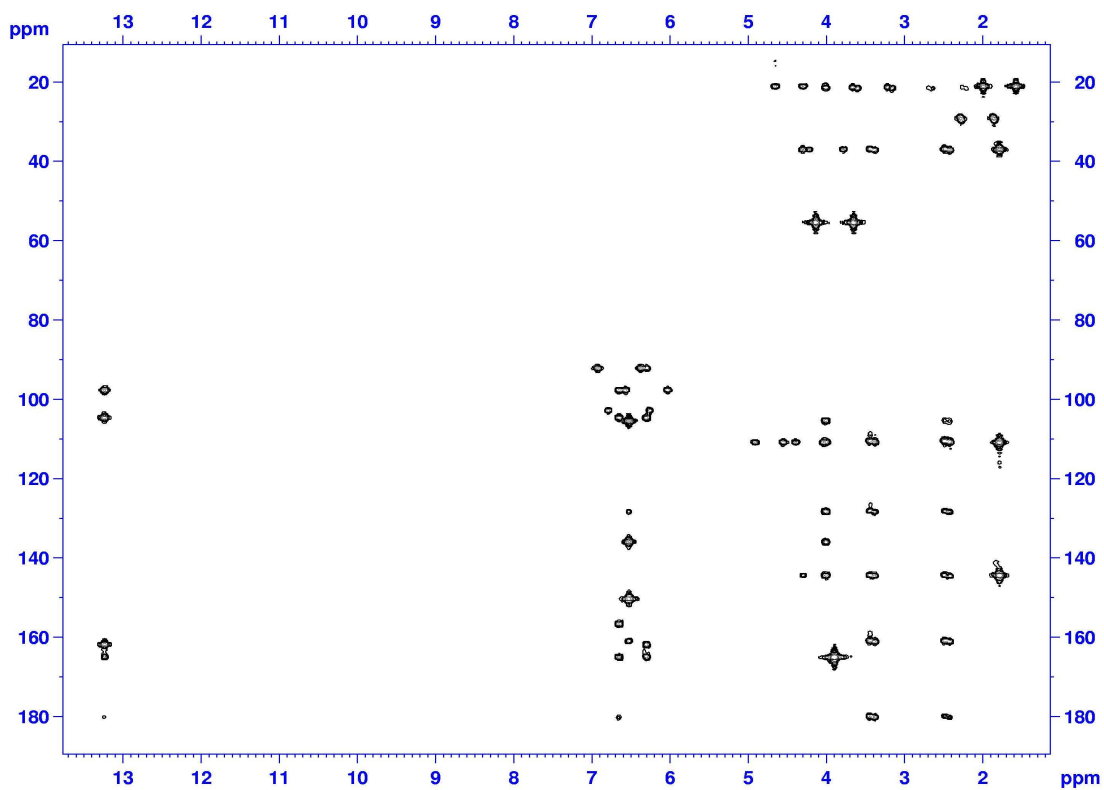


Figure 101 2D HMBC spectrum of AE12

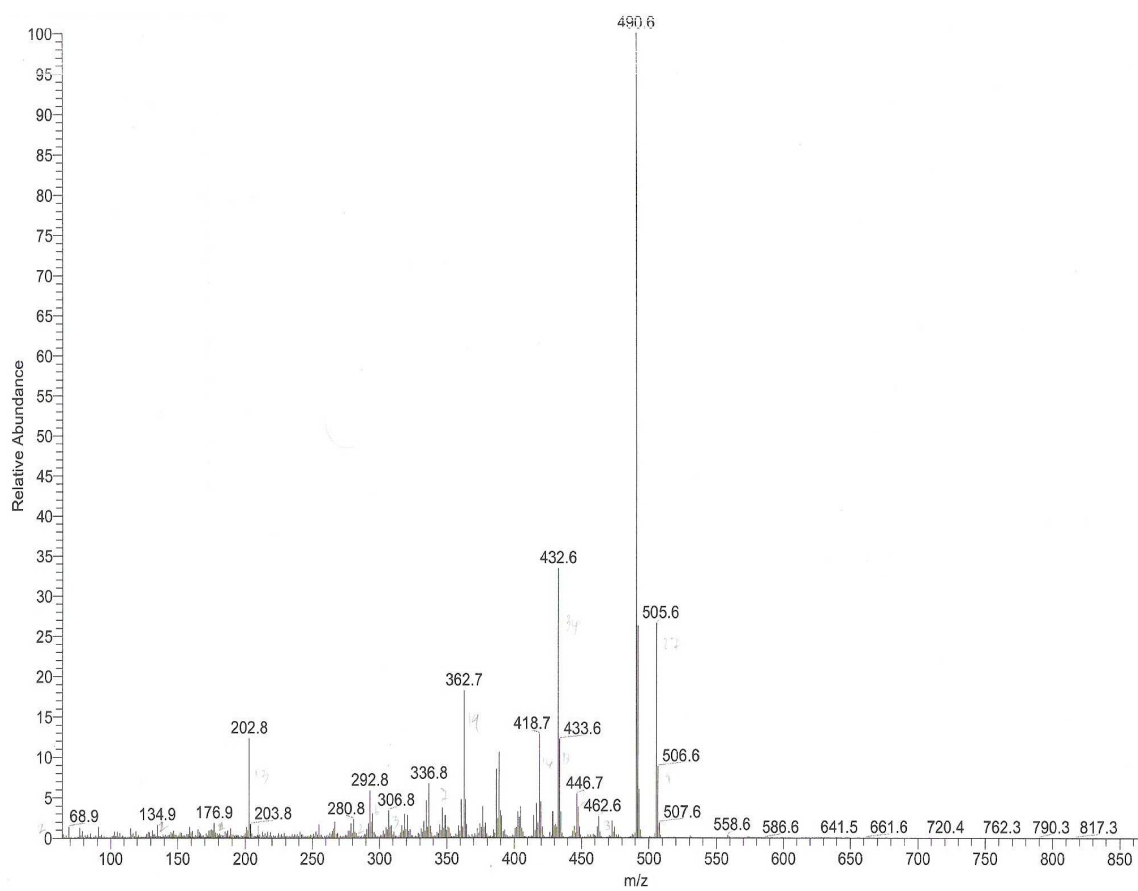


Figure 102 EI-MS spectrum of **AE13**

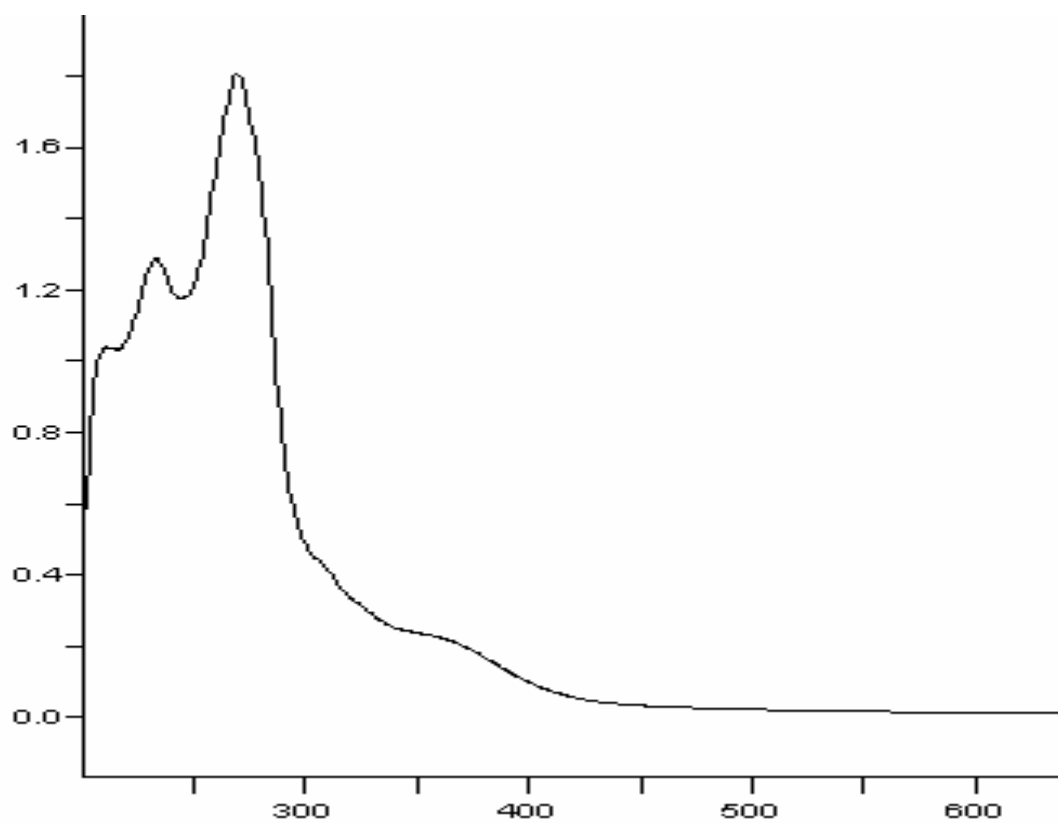


Figure 103 UV (EtOH) spectrum of AE13

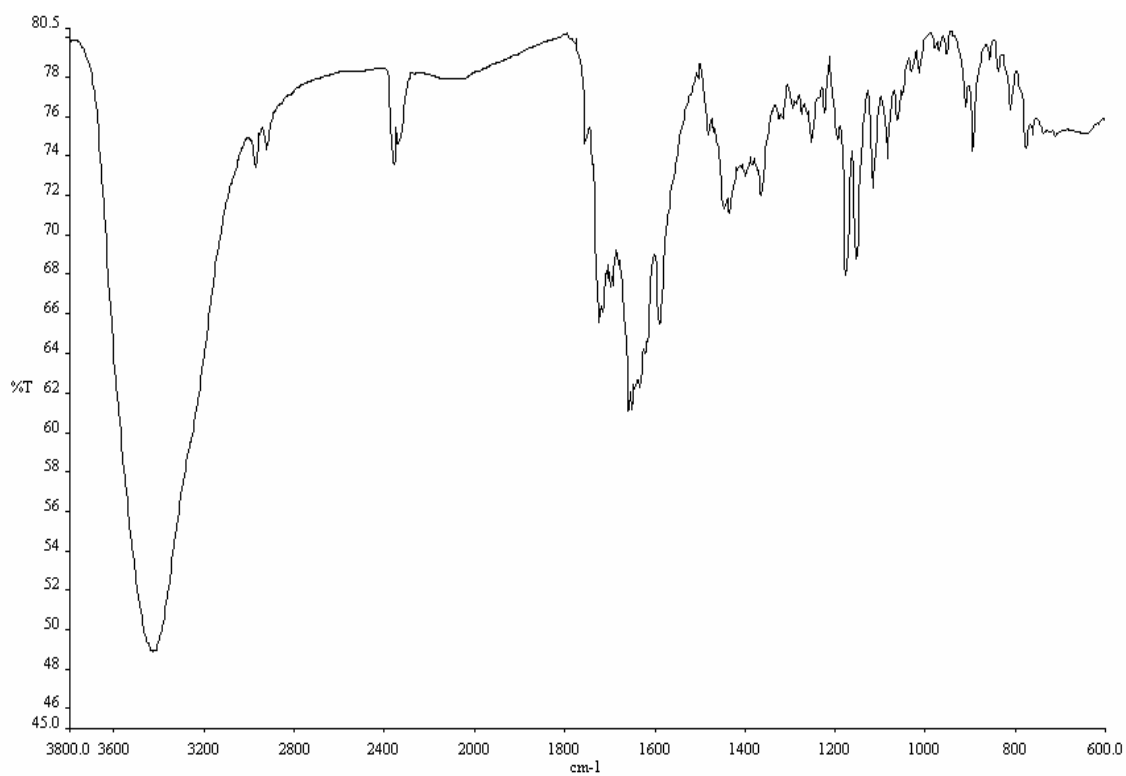


Figure 104 FT-IR (Neat) spectrum of AE13

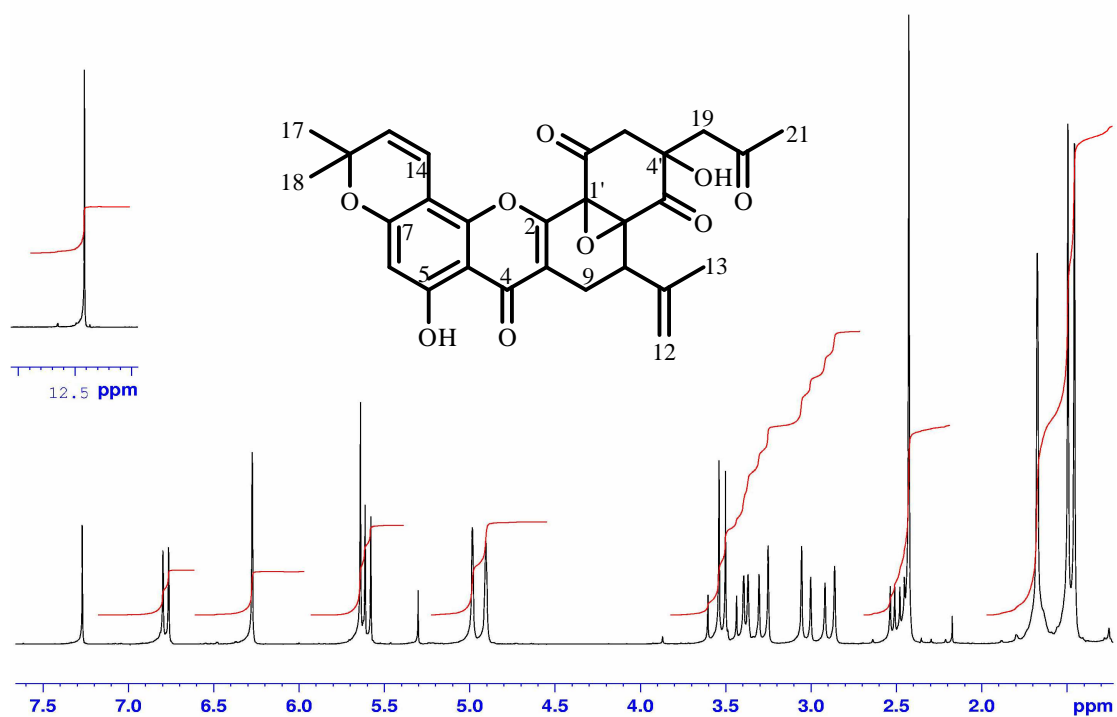


Figure 105 ^1H NMR (300 MHz) (CDCl_3) spectrum of AE13

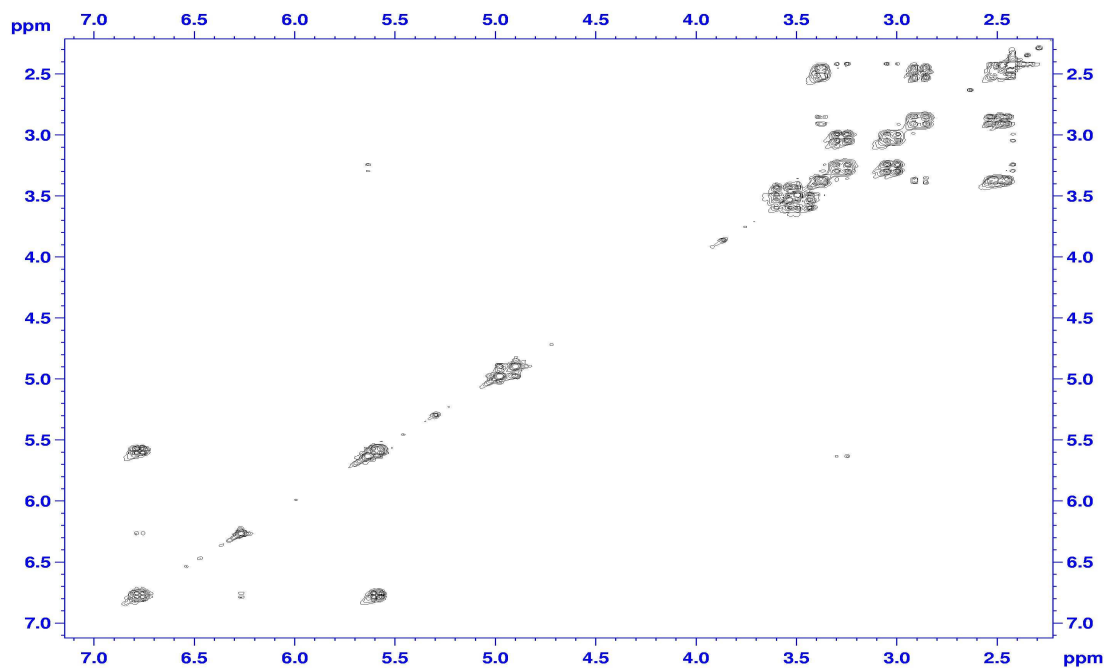


Figure 106 ^1H - ^1H COSY spectrum of AE13

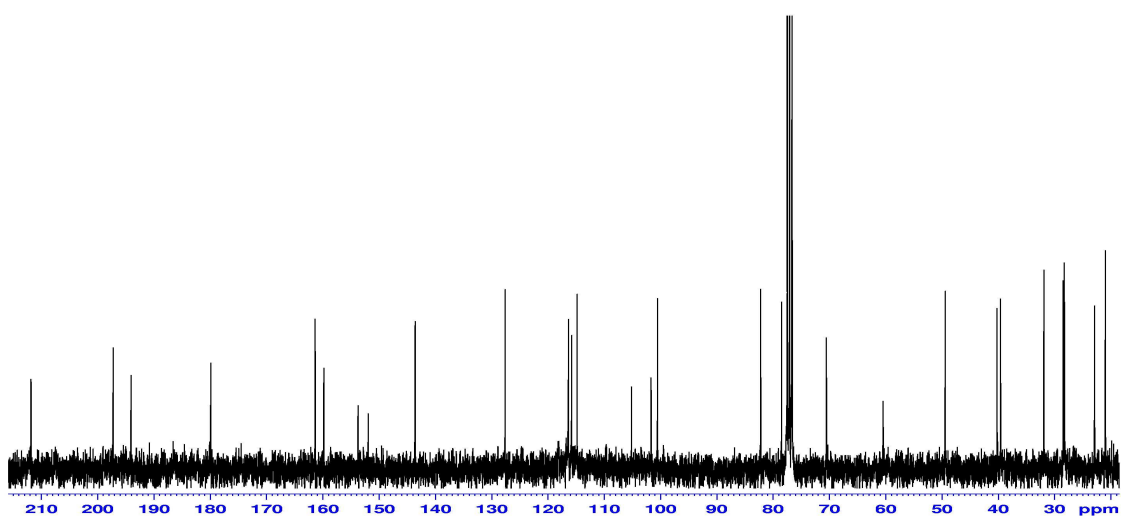


Figure 107 ^{13}C NMR (75 MHz) (CDCl_3) spectrum of AE13

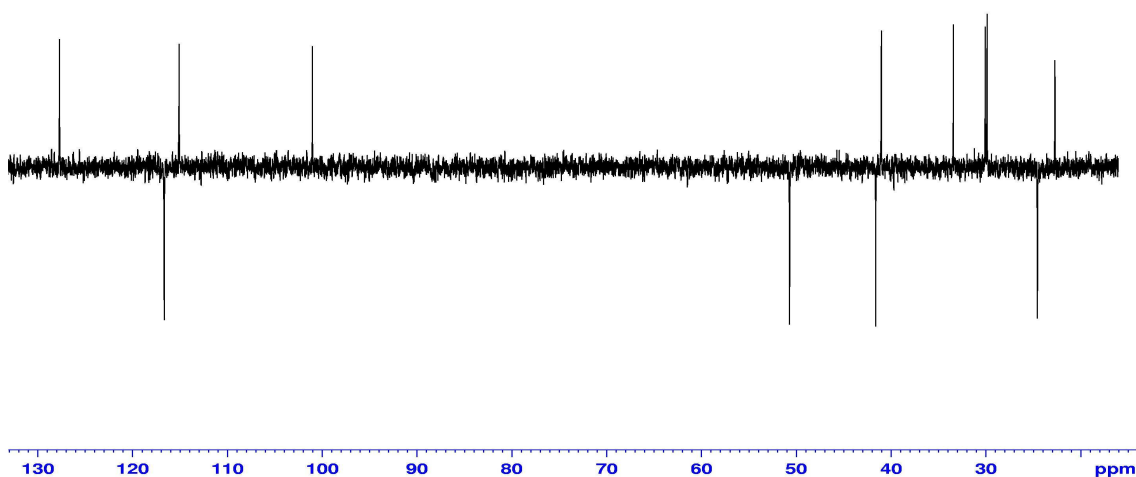


Figure 108 DEPT 135° (CDCl_3) spectrum of AE13

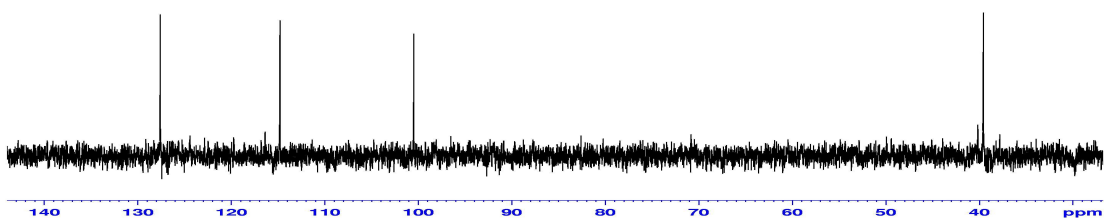


Figure 109 DEPT 90° (CDCl_3) spectrum of AE13

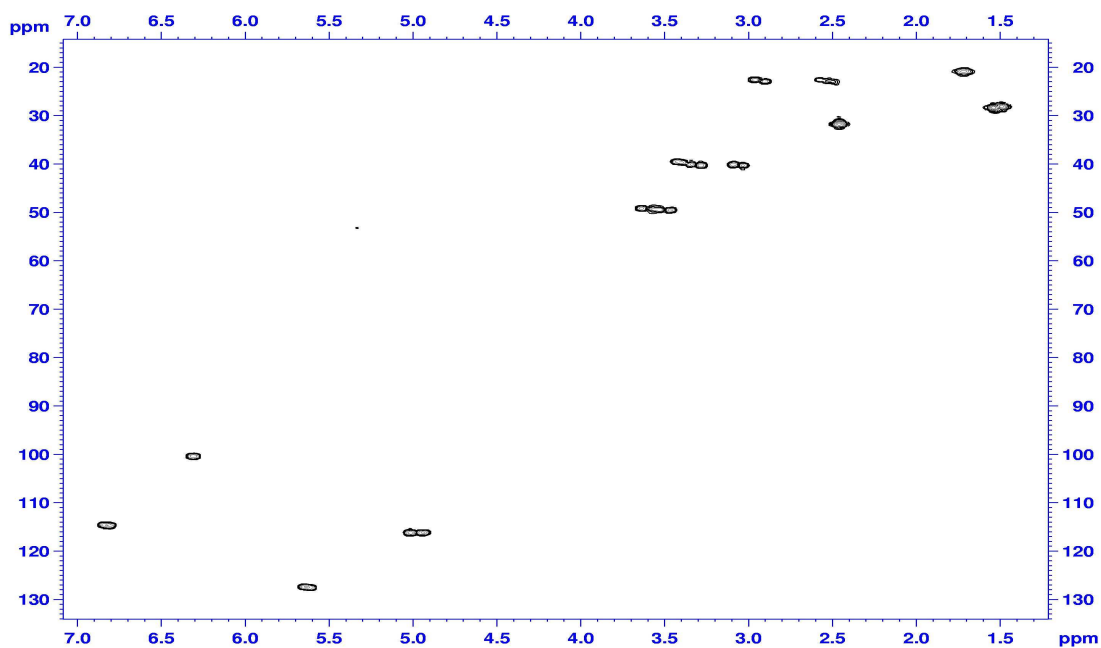


Figure 110 2D HMQC spectrum of AE13

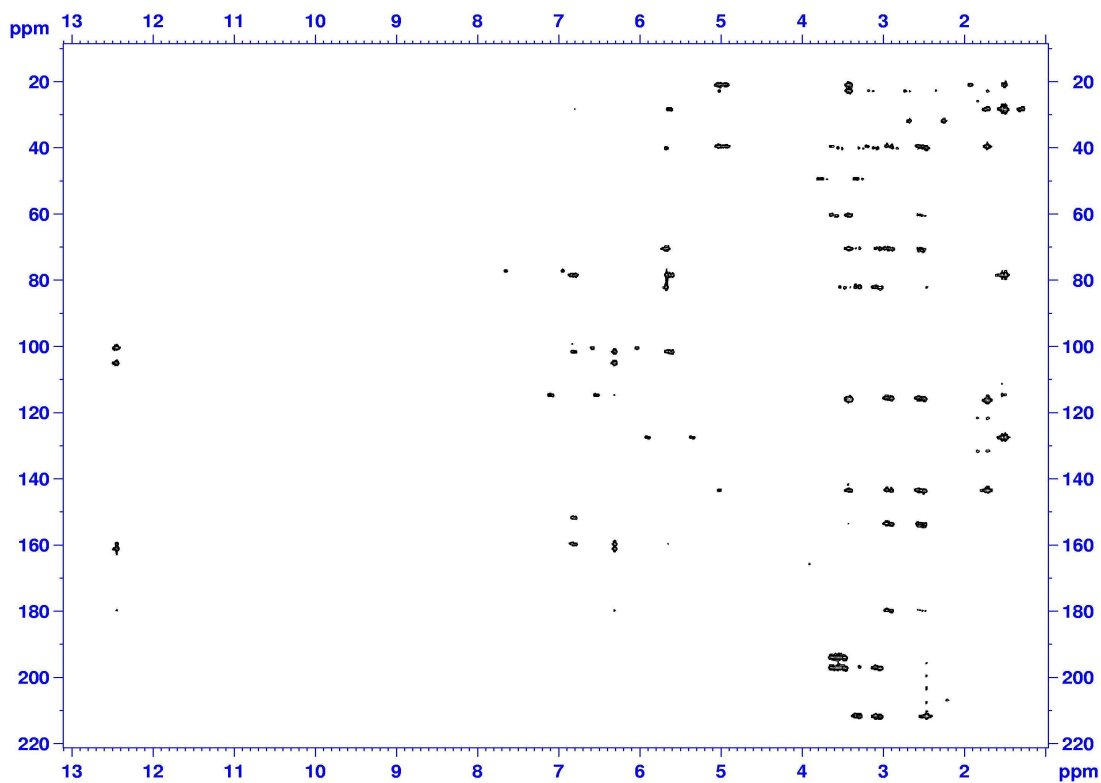


Figure 111 2D HMBC spectrum of AE13

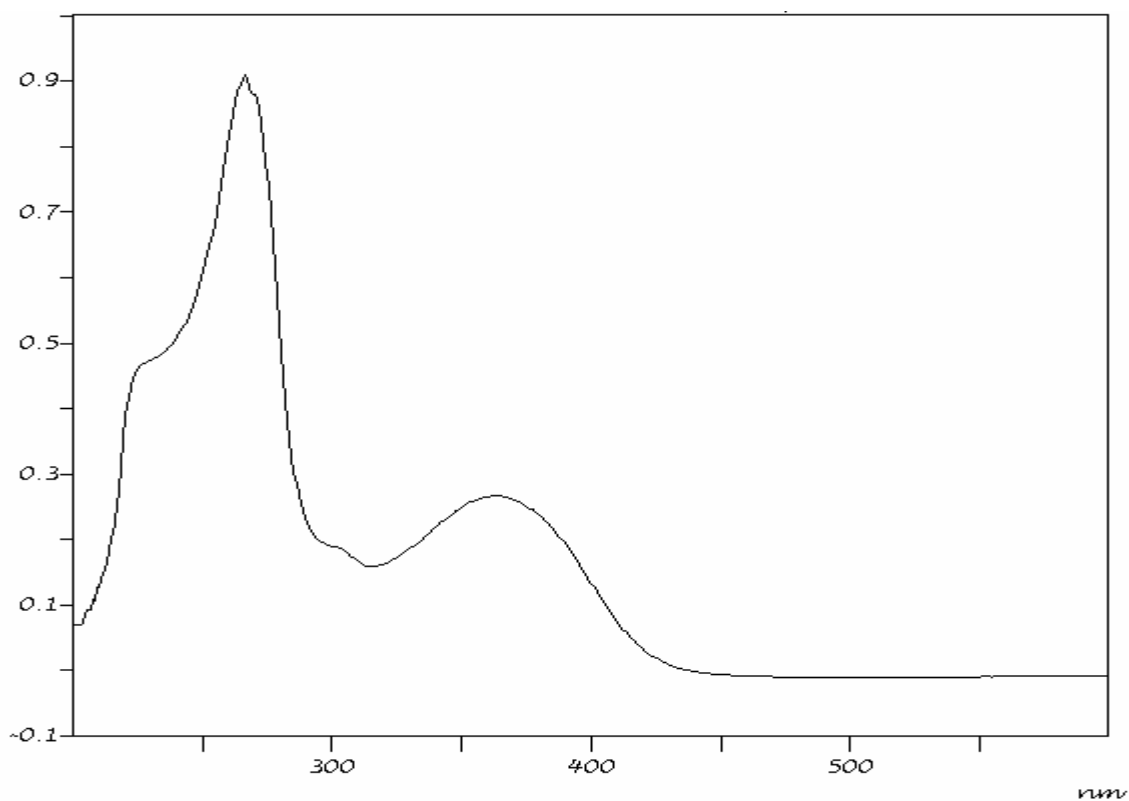


Figure 112 UV (CH₃OH) spectrum of AE14

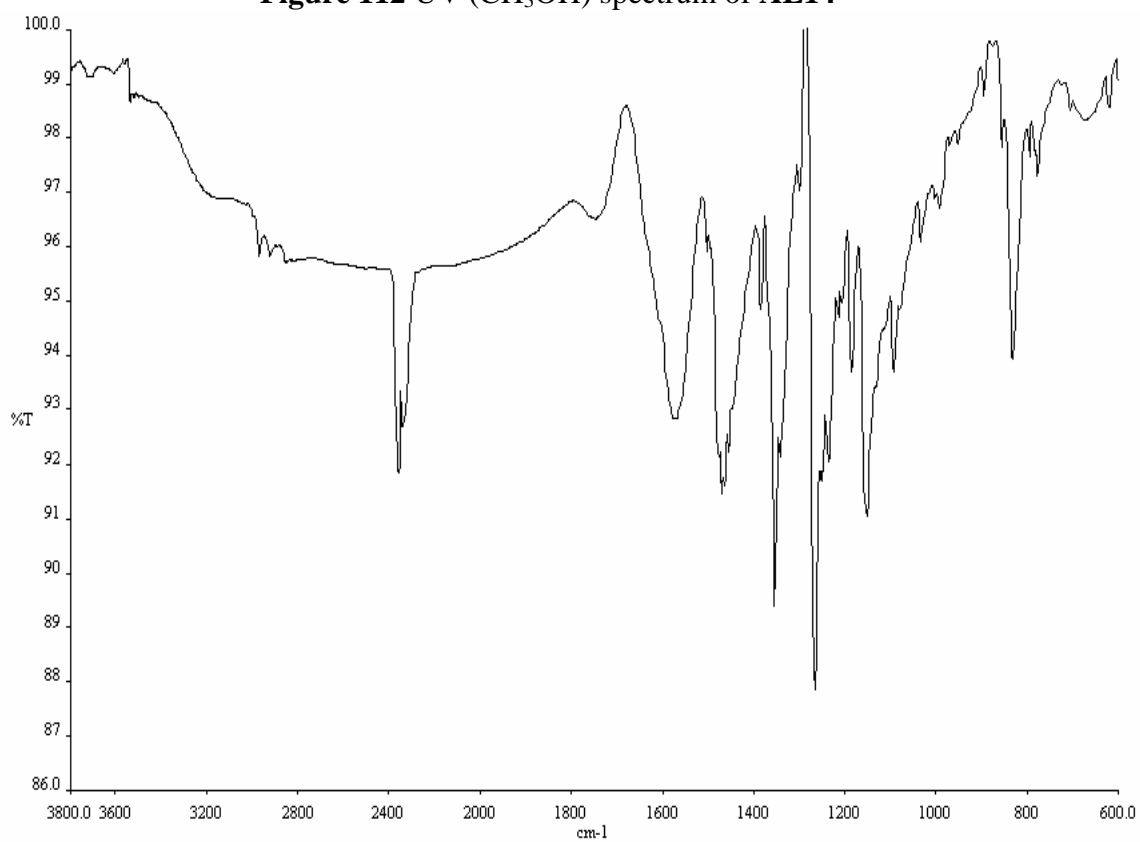


Figure 113 FT-IR (Neat) spectrum of AE14

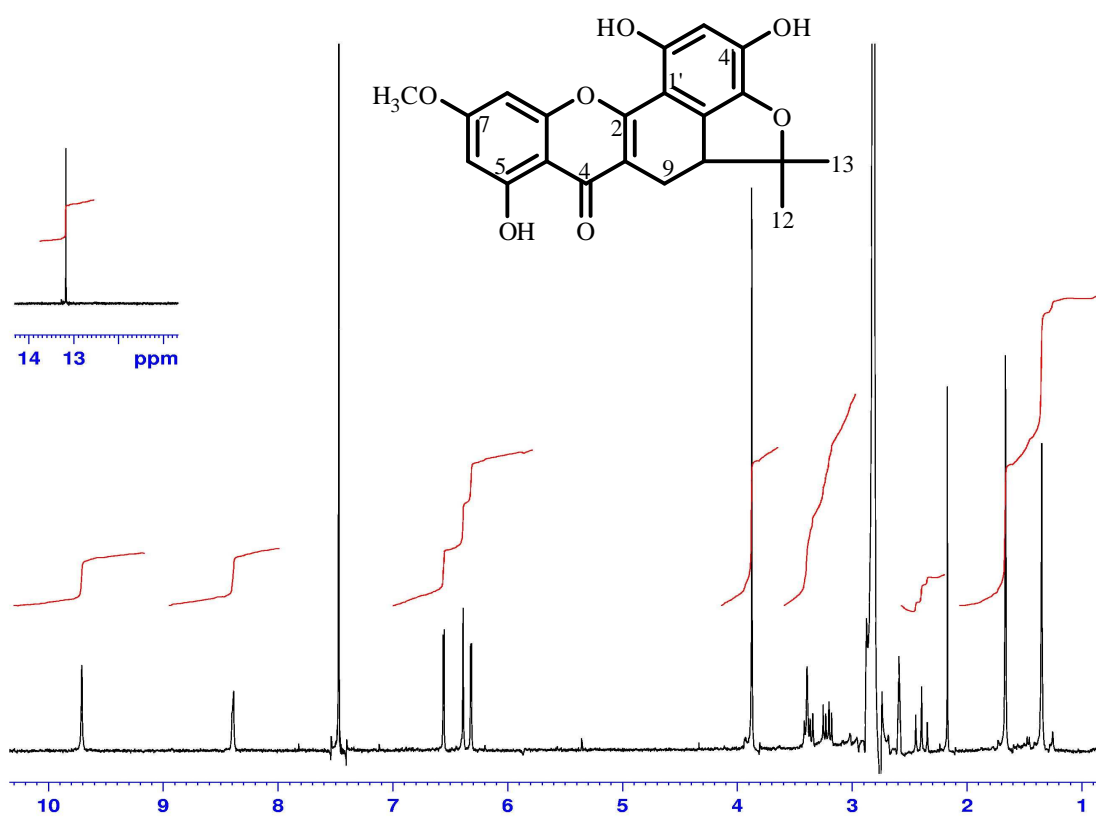


Figure 114 ^1H NMR (300 MHz) ($\text{CDCl}_3+\text{DMSO}-d_6$) spectrum of AE14

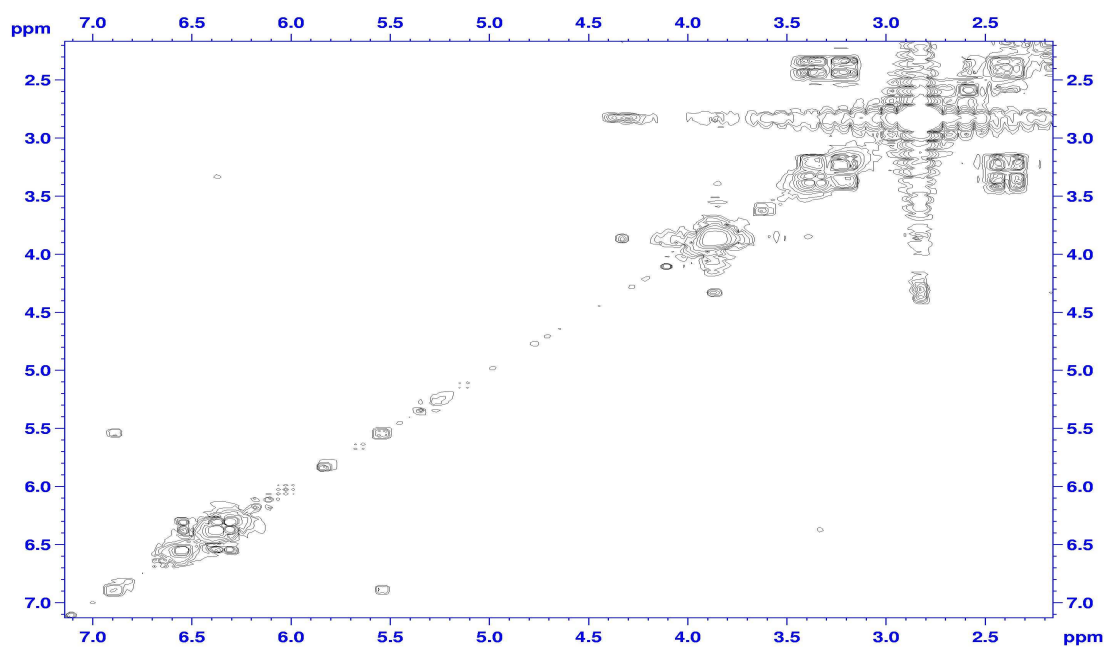


Figure 115 ^1H - ^1H COSY spectrum of AE14

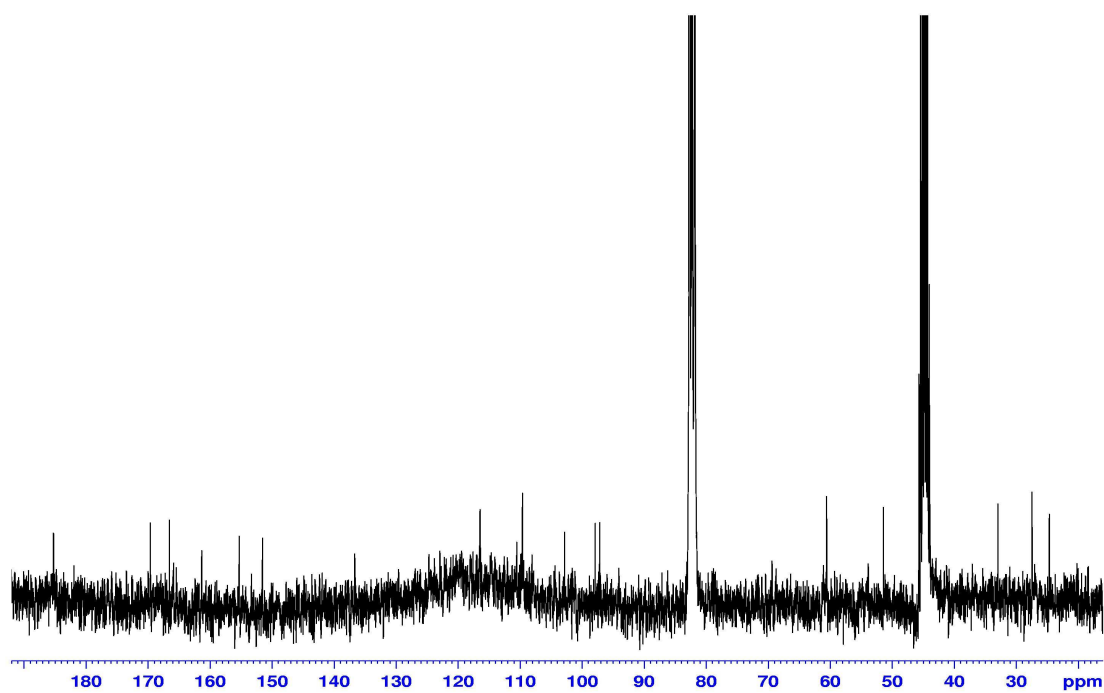


Figure 116 ^{13}C NMR (75 MHz) ($\text{CDCl}_3+\text{DMSO}-d_6$) spectrum of AE14

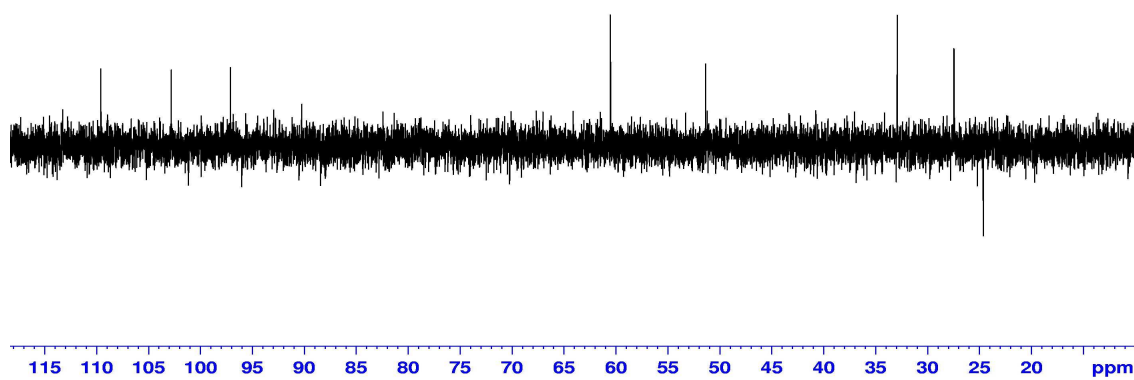


Figure 117 DEPT 135° ($\text{CDCl}_3+\text{DMSO}-d_6$) spectrum of AE14

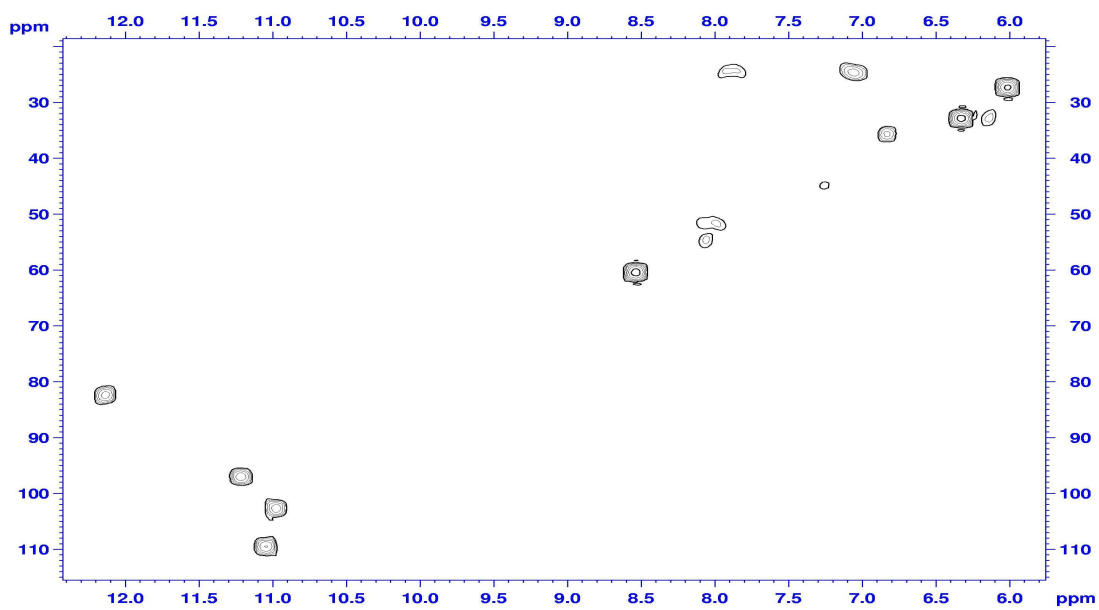


Figure 118 2D HMQC spectrum of AE14

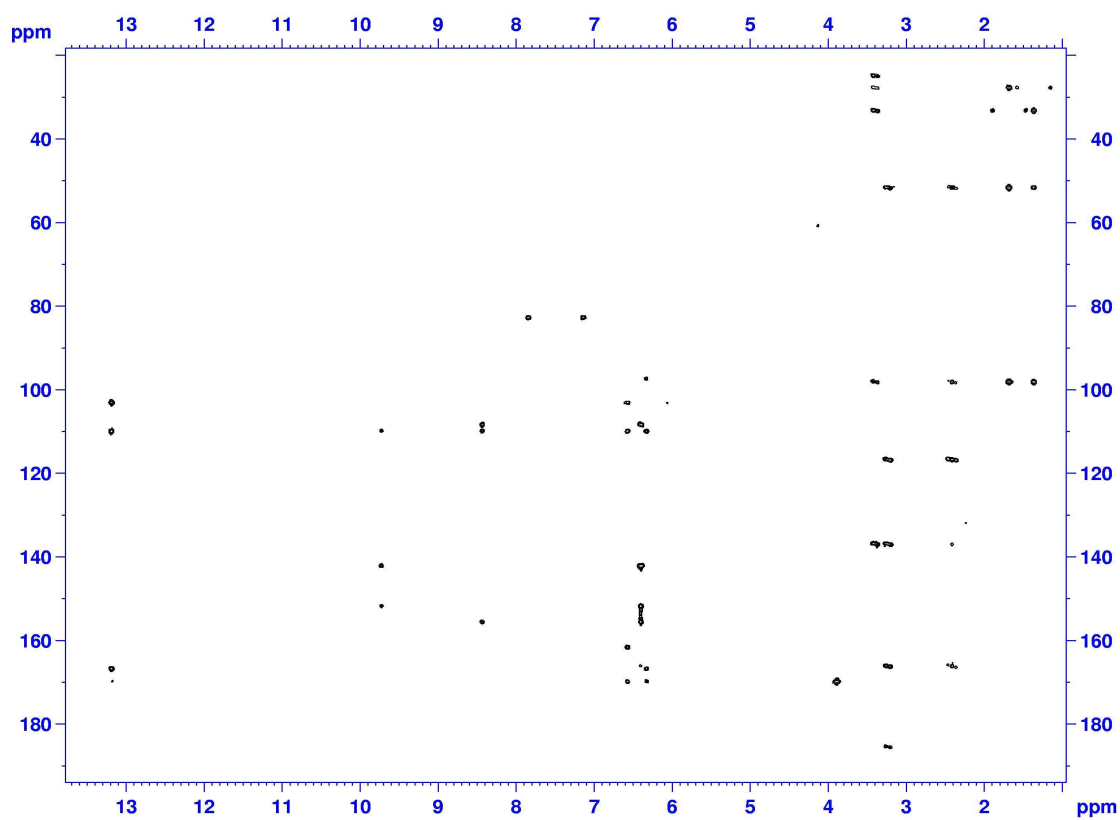


Figure 119 2D HMBC spectrum of AE14

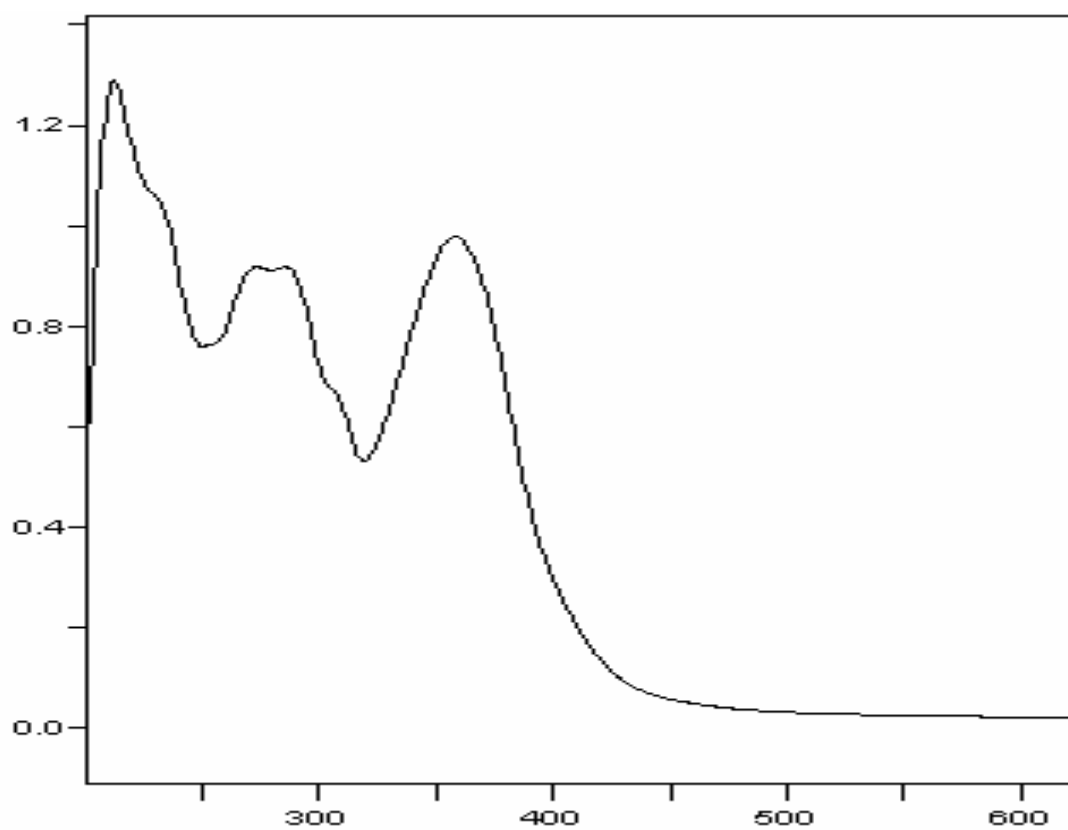


Figure 120 UV (EtOH) spectrum of AE15

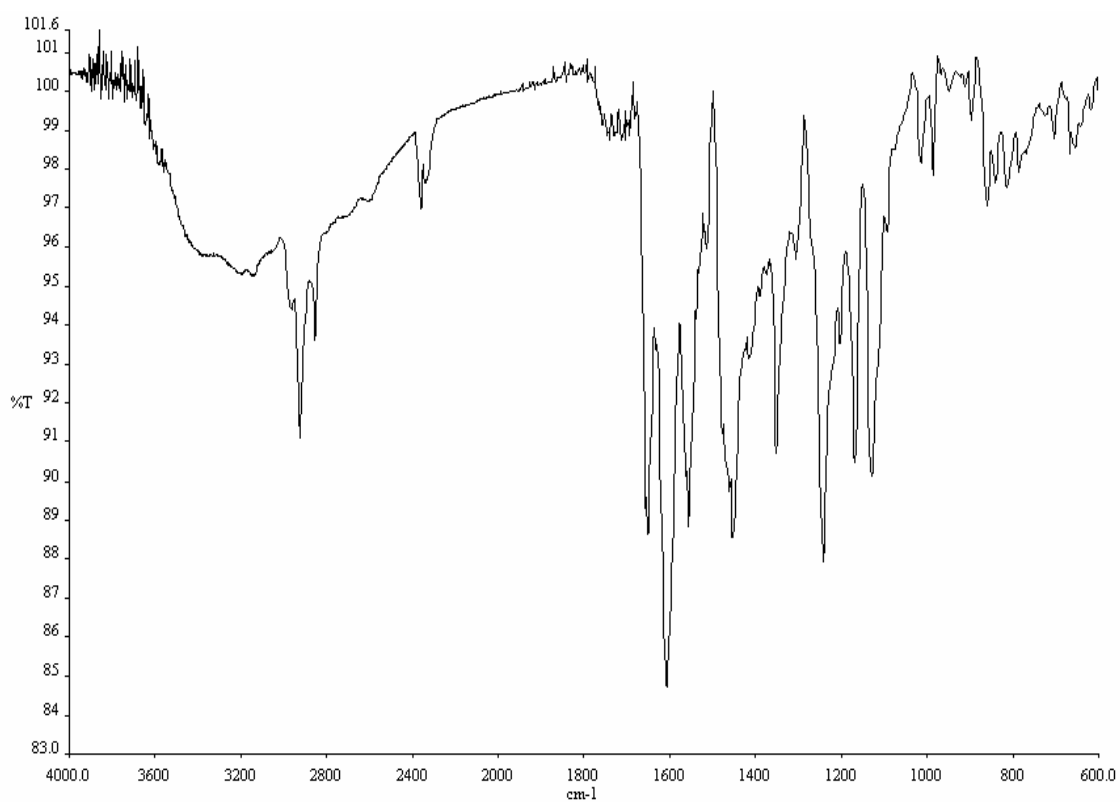


Figure 121 FT-IR (Neat) spectrum of AE15

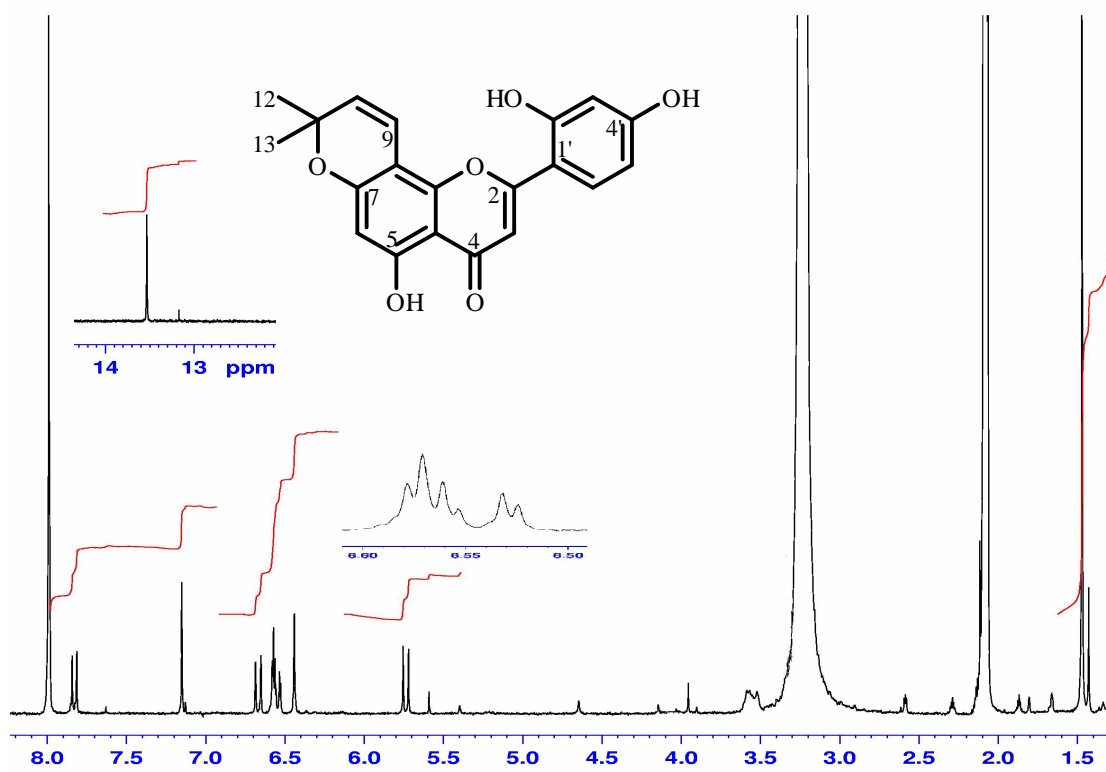


Figure 122 ^1H NMR (300 MHz) (Acetone- d_6) spectrum of AE15

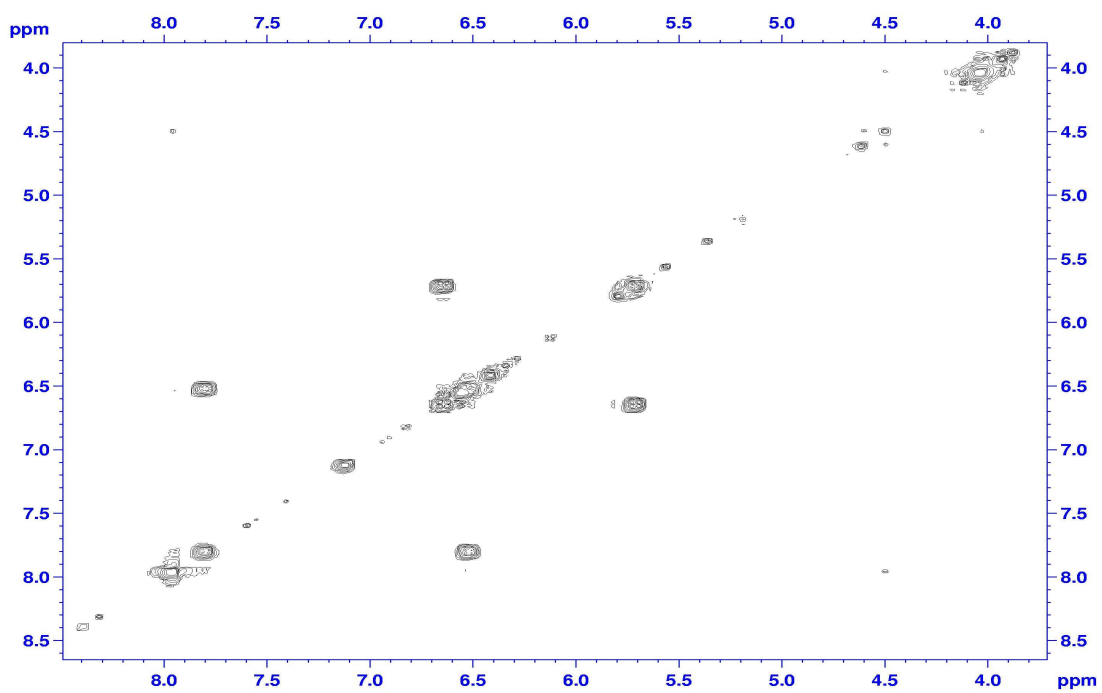


Figure 123 ^1H - ^1H COSY spectrum of AE15

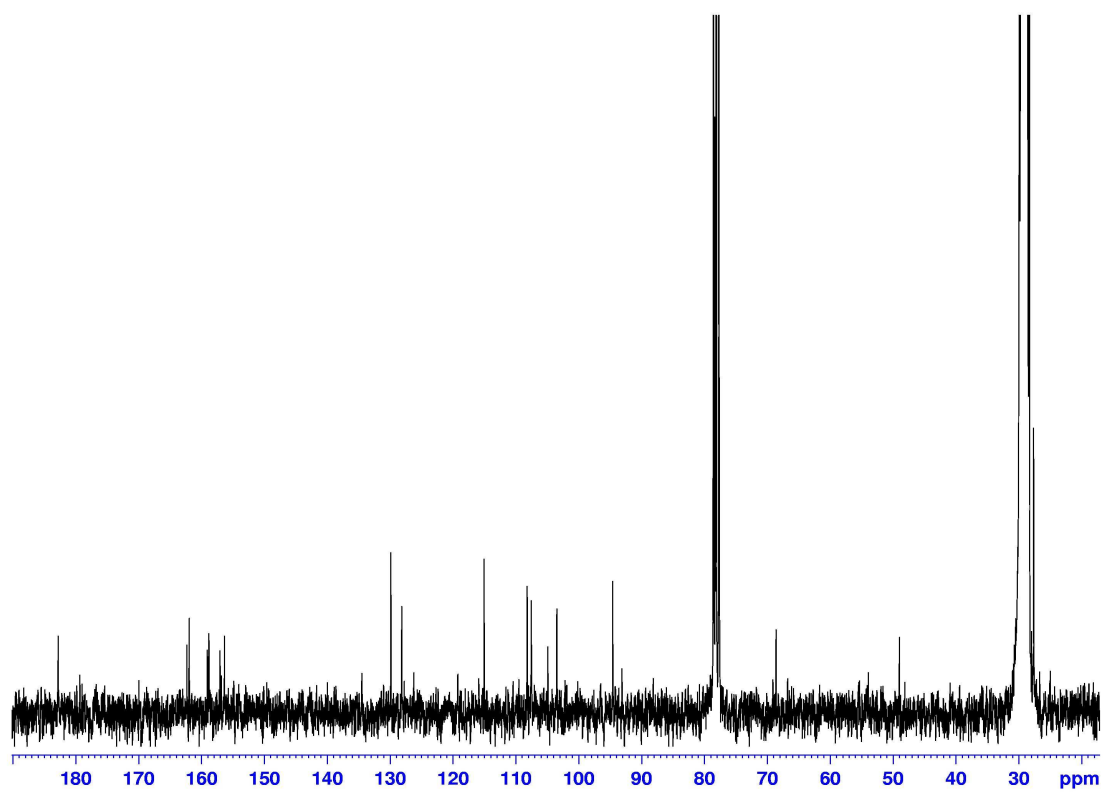


Figure 124 ^{13}C NMR (75 MHz) ($\text{Acetone-}d_6$) spectrum of AE15

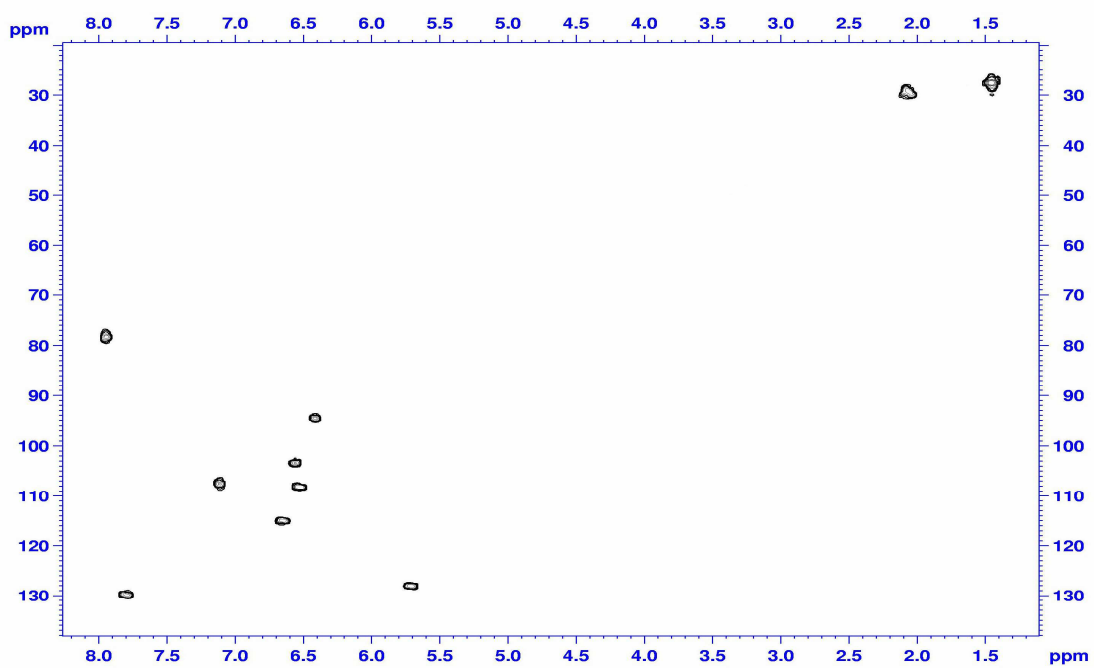


Figure 125 2D HMQC spectrum of AE15

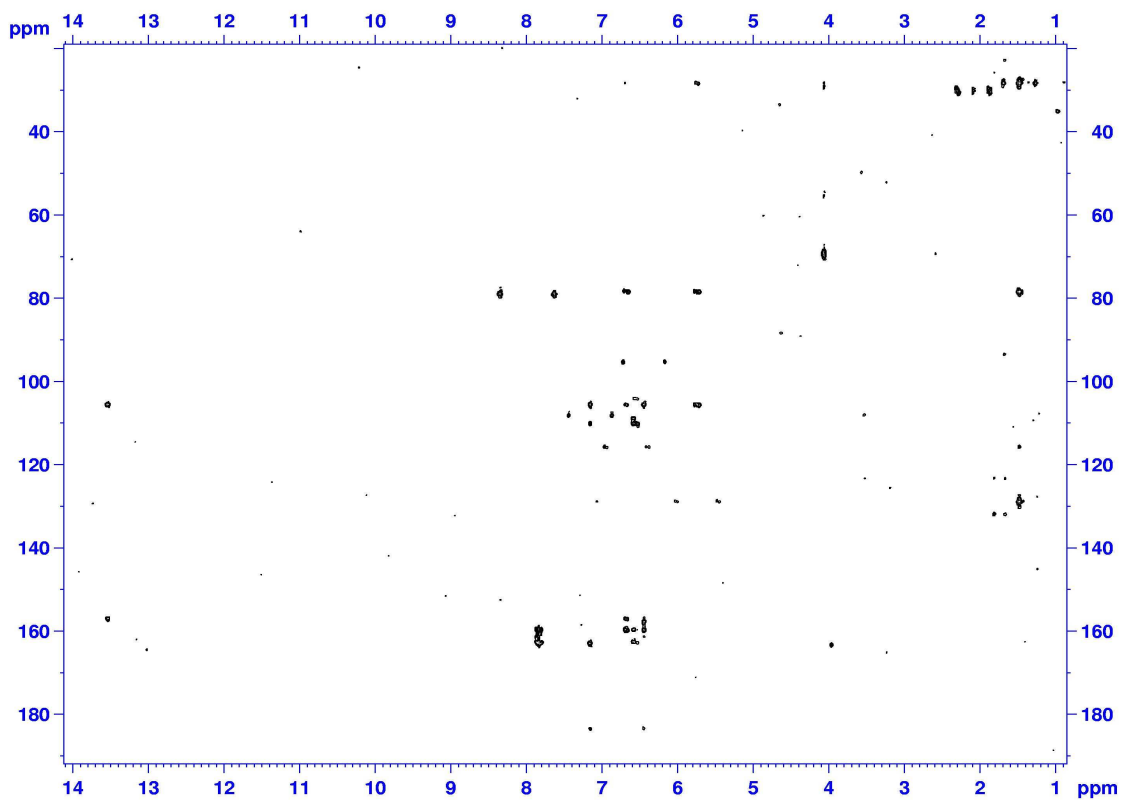


Figure 126 2D HMBC spectrum of AE15

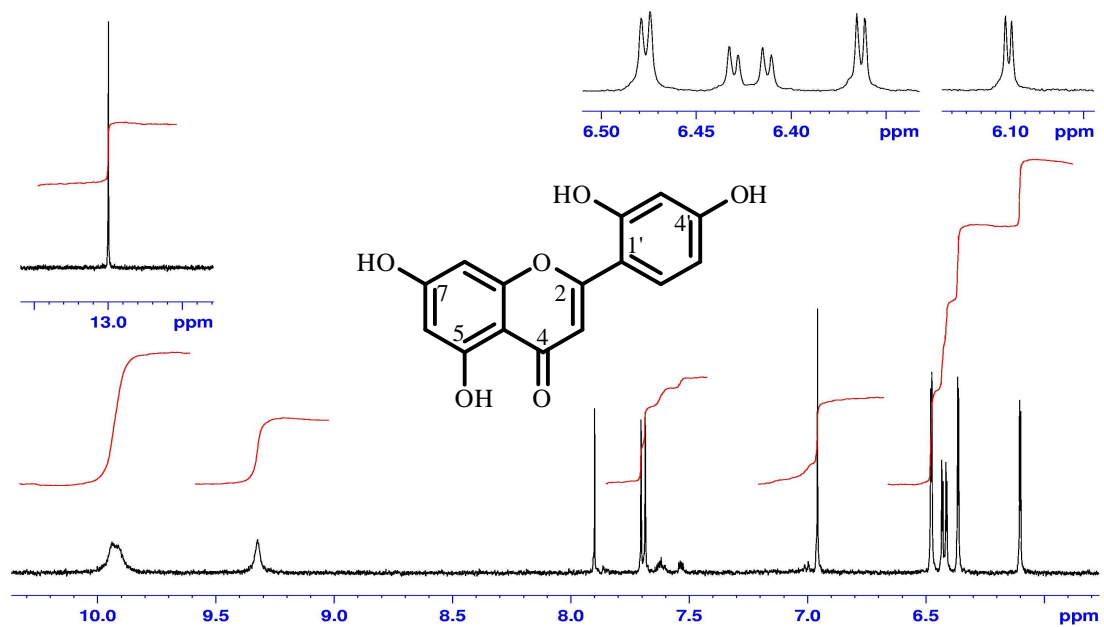


Figure 127 ^1H NMR (500 MHz) (CDCl_3) spectrum of AE16

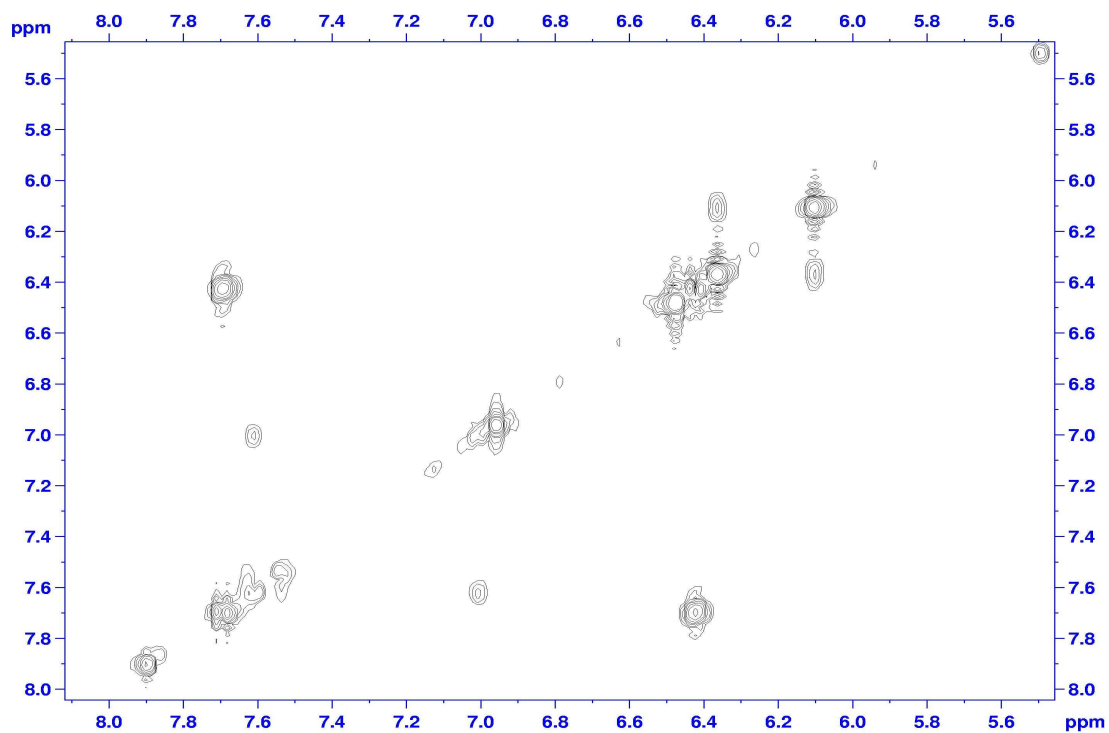


Figure 128 ^1H - ^1H COSY spectrum of AE16

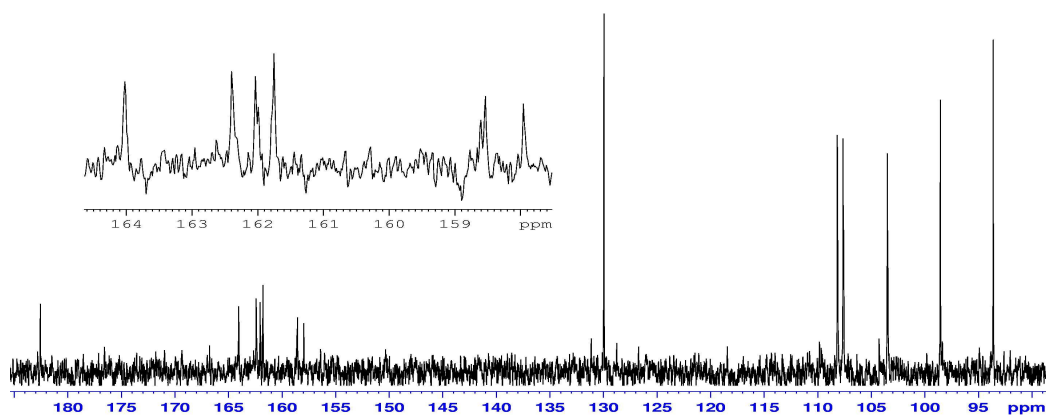


Figure 129 ^{13}C NMR (125 MHz) ($\text{CDCl}_3+\text{DMSO}-d_6$) spectrum of AE16

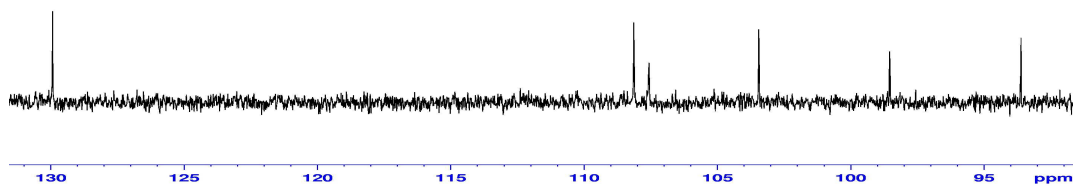


Figure 130 DEPT 135° ($\text{CDCl}_3+\text{DMSO}-d_6$) spectrum of AE16

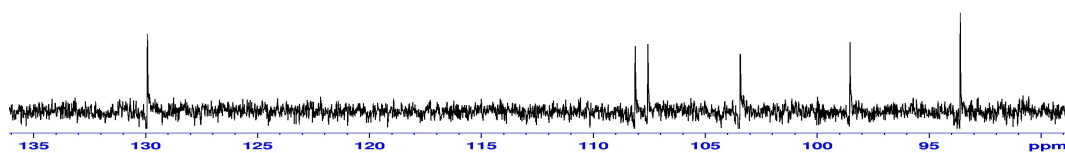


Figure 131 DEPT 90° ($\text{CDCl}_3+\text{DMSO}-d_6$) spectrum of AE16

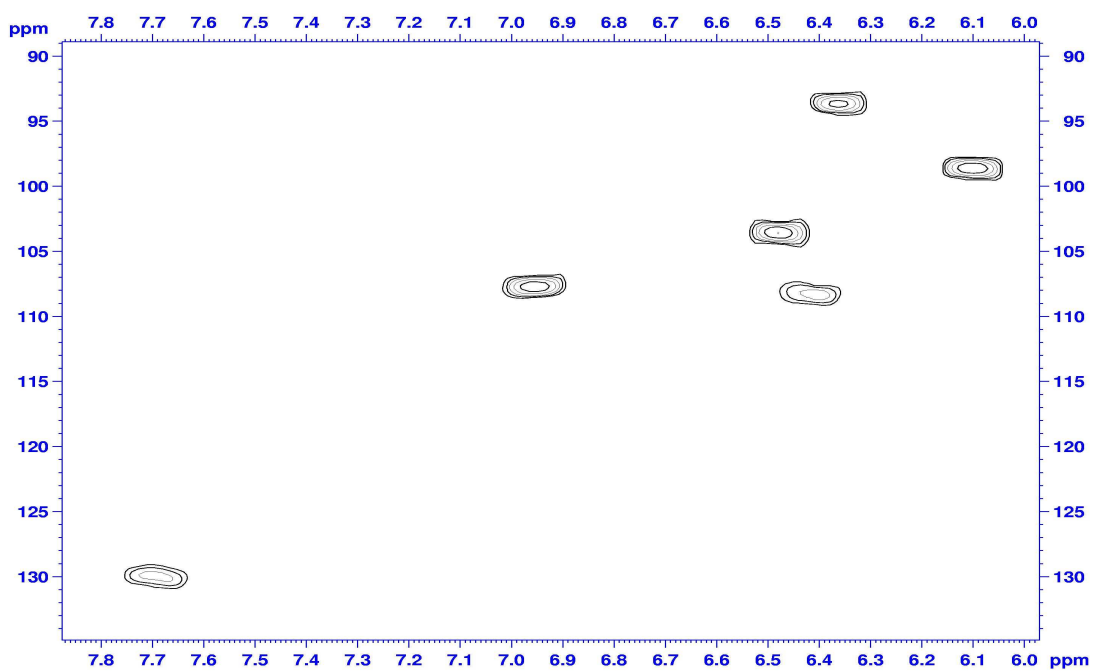


Figure 132 2D HMQC spectrum of AE16

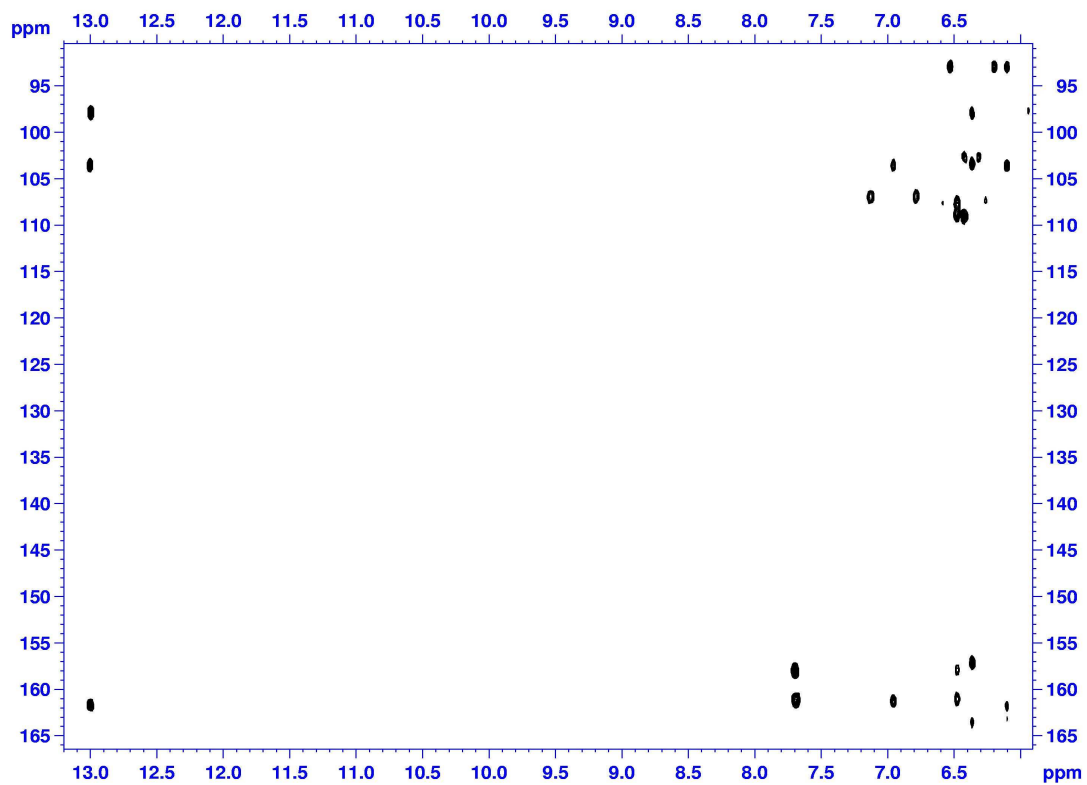


Figure 133 2D HMBC spectrum of AE16

VITAE

Name Miss Aeesoh Yanya

Student ID 5010220176

Educational Attainment

Degree	Name of Institution	Year of Graduation
B.Sc. (Chemistry)	Prince of Songkla University	2006

Scholarship Awards during Enrolment

Center of Excellence for Innovation in Chemistry (PERCH-CIC),
Commission on Higher Education, Ministry of Education

List of Publications and Proceedings

1. A. Yanya and W. Mahabusarakam. "Prenylated Flavonoids from the Bark of *Artocarpus elasticus*". The 6th IMT-GT UNINET CONFERENCE 2008, The Gurney Resort Hotel & Residences Penang, Penang, Malaysia, 28-30 August 2008. (Poster presentation)
2. Aeesoh Yanya and Wilawan Mahabusarakam. "Prenylated Flavones from the Bark of *Artocarpus elasticus*". 4th National Grade Research Conference, Burapha University, Chonburi, Thailand, 13 March 2009. (Poster presentation)