

## CHAPTER II

### HISTORICAL

#### 1. Bioactive compounds of the Celastraceae

The Celastraceae family is indigenous to tropical and sub-tropical regions of the world , including North Africa , South America , and many parts of East Asia , particularly China.<sup>3,4</sup> The family constitutes approximately 88 genera and 1300 species of plants.<sup>4</sup> They have been valued since antiquity because their extracts have useful medicinal properties.<sup>4,5</sup> Indeed , the variety of properties attributed to crude plant extracts of the Celastraceae in traditional medicine and agriculture is astonishing , and includes stimulant , appetite suppressive , sedative , emetic , purgative , memory-restorative , male contraceptive , anti-tumor , anti-leukemic , anti-bacterial , insecticidal and insect repellent activities.<sup>4</sup> For example , extracts of the root of the thunder-god vine *Tripterygium wilfordii* Hook. f. have been used by the Chinese for centuries to combat life-threatening illness , and have more recently found application in the treatment of leukemia and rheumatoid arthritis.<sup>6</sup> Similarly , the stimulating properties of the leaves of the khat bush *Catha edulis* Forssk. , which grows in certain parts of East Africa and Southern Arabia , were reported in the literature as early as 1237 by the Arabian physician Naguib Ad Din , who proposed the use of khat for alleviating depression. Since khat leaves rapidly lose their effect after picking , the culture of chewing khat leaves for their stimulant/euphoric and appetite/fatigue suppressive effects was for centuries confined to the immediate areas where the plants grows. The habit had become so widespread by the mid-1970s because of the improvements in road communications, a form of addiction known as khatism was beginning to present serious social and economic difficulties in Ethiopia and the Yemen Republic.<sup>4,7</sup> This prompted the World Health Organization to fund an investigation to determine the active constituents of khat. This led to the coincidental discovery of the macrolide maytansine ( 1 ) from *Maytenus ovatus* Loes. in 1972 , which displayed exceptionally high inhibitory activity *in vitro* against certain human carcinomas<sup>8</sup> , that catapulted the natural product chemistry of the Celastraceae into the limelight. The development of maytansine ( 1 ) and related macrolides as anti-tumor lead has halted in the 1980s when they were found to cause serious gastro-intestinal damage in rats<sup>4</sup> , but recent advances in drug

targeting *via* conjugation to cancer-specific humanized antibodies appear to offer new opportunities for clinical application.<sup>4</sup>

Over the last 30 years or so , a large number of secondary metabolites exhibiting a wide range of bioactivity have been extracted from the Celastraceae . The maytansinoids , maytansine ( 1 ) , maytanprine ( 2 ) and maytanbutine ( 3 ) displayed cytotoxic activity.<sup>9-11</sup> The sesquiterpene pyridine alkaloids triptonine A ( 4 ) and triptonine B ( 5 ) isolated from *Tripterygium hypoglaucum* ( Levl. ) Hutch. and *T. wilfordii* showed anti - HIV activity and were considered a new class of potent anti - HIV agents<sup>12</sup> . Ebenifoline E-11 ( 6 ) and congorinine E-1 ( 7 ) , two other compounds isolated from *T. wilfordii* Hook. f. , possessed immunosuppressive activity<sup>13</sup> . Emarginatine A ( 8 )<sup>14</sup> , emarginatine F ( 9 )<sup>15</sup> , and emarginatinine ( 10 )<sup>14</sup> from *Maytenus emarginata* and celahinine A ( 11 )<sup>16</sup> from *Celastrus hindsii* showed cytotoxic activity whereas cathedulin E-5 ( 12 ) from *Catha edulis* exhibited potent insecticidal activity.<sup>4</sup>

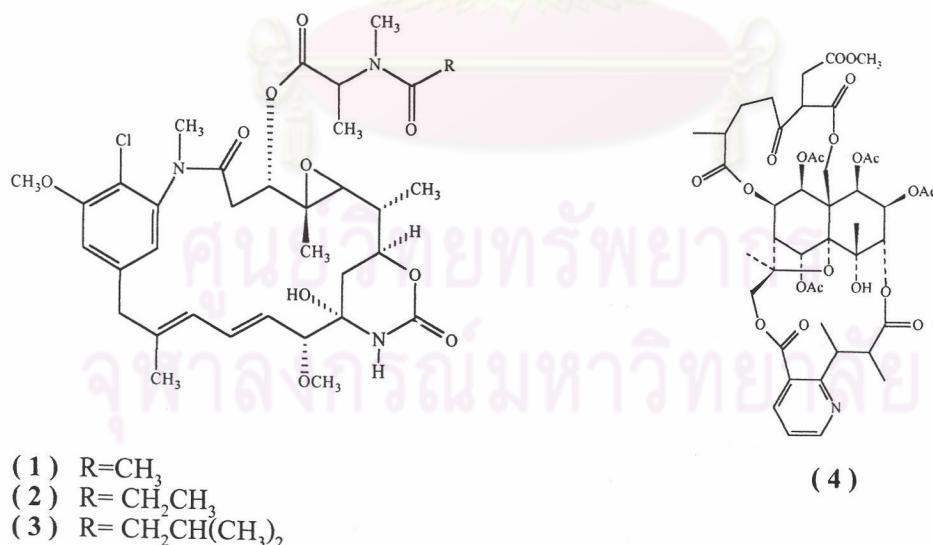
The quinone - methide triterpenes tingenone ( 13 ) , 22 $\beta$  - hydroxy tingenone ( 14 ) , celastrol ( 15 ) , and pristimerin ( 16 ) displayed cytotoxic activity<sup>17,18-19,20</sup> , and several quinone – methide triterpenes have been reported for their antimicrobial activity.<sup>21-26</sup>

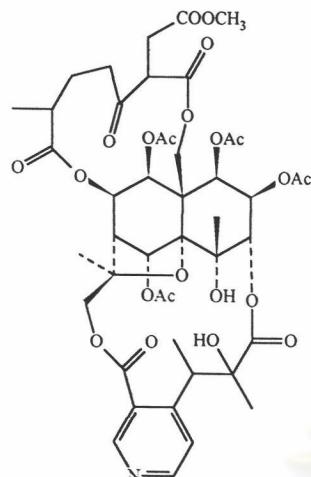
The dihydroagarofuran sesquiterpene , celangulin ( 17 ) , showed insecticidal and anti-feedant activity.<sup>27</sup> Both celafolin D-2 ( 18 ) and celafolin B-2 ( 19 ) exhibited cytotoxic activity.<sup>28</sup> Interestingly , celafolin A-1 ( 20 ) and celorbicol ( 21 ) have also been found to reverse multi-drug resistance ( MDR ) in cancer cells.<sup>29</sup> More recently, a series of sesquiterpenoids based on 4 $\beta$ ,14-dihydroxycelapanol ( 22 ) and 3,13-dideoxyeuonyminol core structure ( 23 and 24 ) have been found to reverse the resistance of parasitic protozoan *Leishmania tropica* to growth inhibition by daunomycin.<sup>30</sup>

The tri-epoxide diterpenoid triptolide ( 25 ) , isolated from the roots of *Tripterygium wilfordii* , has been shown to be responsible for some of the significant anti-leukemic and anti-tumor activity of this plant. The compound also displayed male anti - fertility property and exhibited more potent anti-inflammatory activity than prednisolone ( 26 ) .<sup>31</sup> It is one of a number of diterpene epoxides contained in a commercially available ‘ total multi - glycoside extract’ or ‘T<sub>II</sub> extract’ that has been used clinically for male fertility control , treatment of inflammatory , and autoimmune diseases in China. Triptolide was believed to be the main active principle responsible for the immunosuppressive activity of ‘ T<sub>II</sub> extract ’.<sup>31</sup>

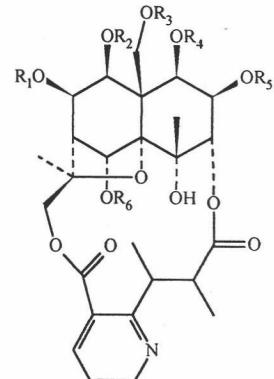
Three triterpenes from different genera of this plant family , 3-oxo-lup-20(29)-en-30,21 $\alpha$ -olide ( 27 ) , elabunin ( 28 ) , and maytenfoliol ( 29 ) were shown to possessed cytotoxic activity.<sup>32-34</sup> Another triterpene , 2 $\alpha$ ,3 $\beta$ -dihydroxy-olean-12-ene-22,29-lactone ( 30 ) , possessed immunosuppressive activity<sup>31</sup>, while kotalagenin 16-acetate ( 31 ) were found to have inhibitory activity on enzyme aldose reductase .<sup>35</sup>

Furthermore , a diverse array of bioactive phenyl-alkylamines , and flavonoids have also been isolated from plants of Celastraceae. In particular , the  $\alpha$ -aminoketone, (-)-cathinone ( 32 ) has now been established as the unstable constituent of fresh khat leaves which is primarily responsible for their stimulant and euphoric effects. The compound is structurally related to , and has the same absolute configuration as , (+)-amphetamine ( 33 ) and (+)-norpseudoephedrine ( 34 ) but is a more potent stimulant.<sup>7</sup> The flavonoid quercetin ( 35 ), which is found in celastraceous plants and a number of other plant families, displays a wide spectrum of anti-oxidant and anti- inflammatory activity as a result of its ability to quench reactive oxygen radicals.<sup>4</sup>





(5)

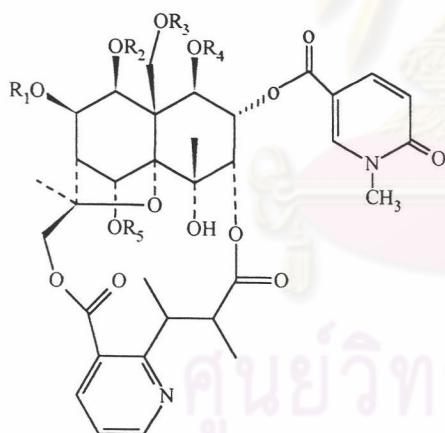


	R <sub>1</sub>	R <sub>2</sub>	R <sub>3</sub>	R <sub>4</sub>	R <sub>5</sub>	R <sub>6</sub>
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(6) Ac Ac Ac Bz Ac Bz

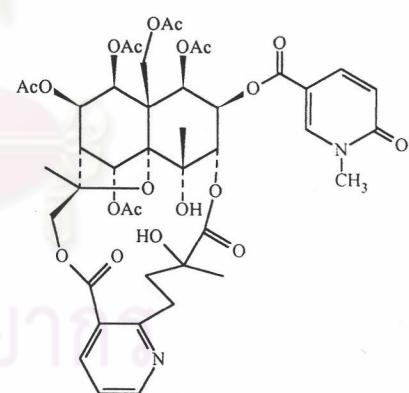
(7) Ac Ac Ac Ac Ac Bz

(11) Ac Bz Ac Bz Ac Bz

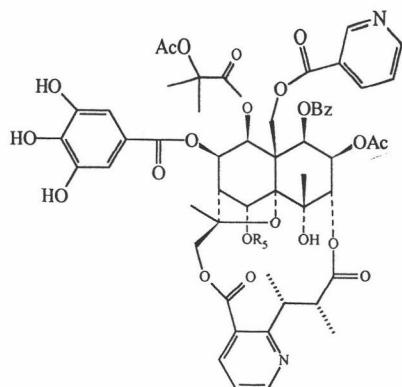


(8) R<sub>1</sub> R<sub>2</sub> R<sub>3</sub> R<sub>4</sub> R<sub>5</sub>

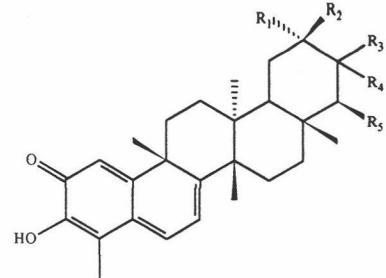
(9) Ac Ac Ac Ac Ac



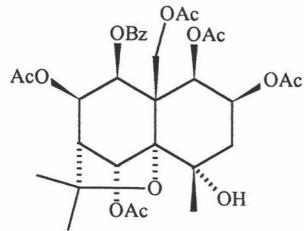
(10)



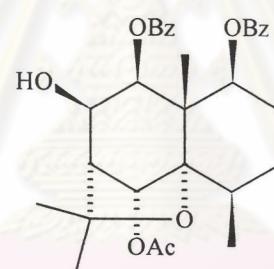
( 12 )


 $R_1 \quad R_2 \quad R_3 \quad R_4 \quad R_5$ 

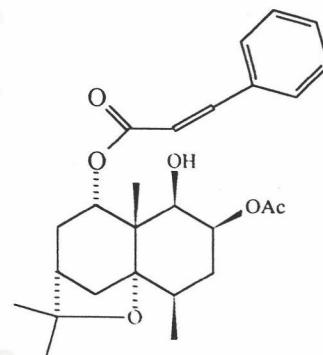
- ( 13 ) H CH<sub>3</sub> =O H  
 ( 14 ) H CH<sub>3</sub> =O OH  
 ( 15 ) COOH CH<sub>3</sub> H H H  
 ( 16 ) COOCH<sub>3</sub> CH<sub>3</sub> H H H



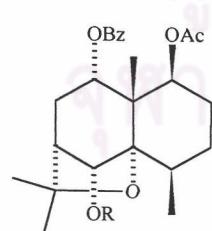
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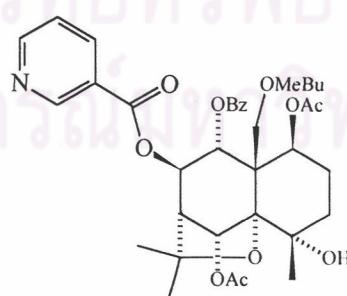
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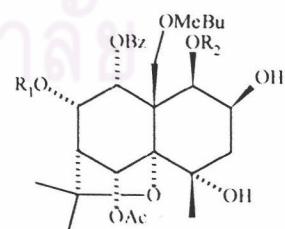
( 19 )



( 20 ) R= Cinnamoyl



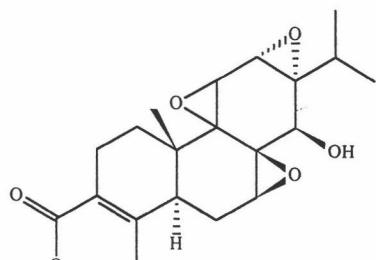
( 22 )


 $R_1 \quad R_2$ 

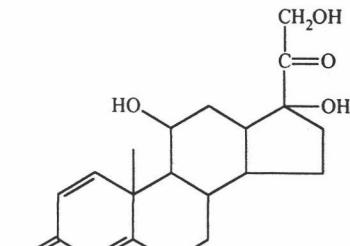
( 21 ) R= Bz

( 23 ) Ac Bz

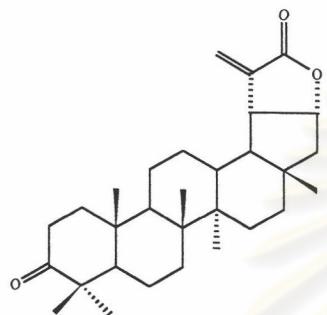
( 24 ) MeBu Ac



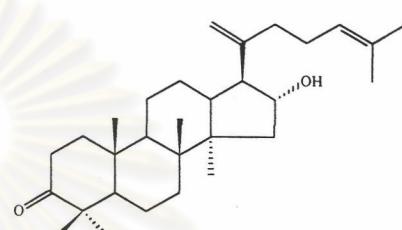
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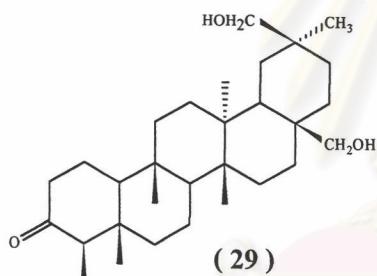
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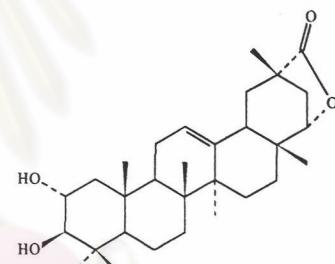
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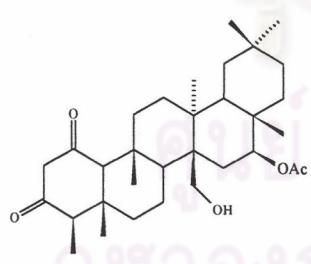
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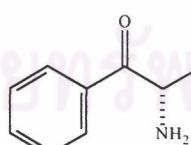
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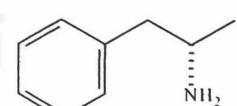
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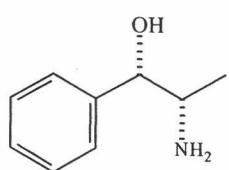
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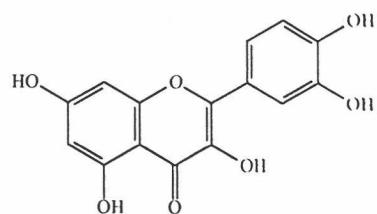
( 32 )



( 33 )



( 34 )



( 35 )

## 2. Quinone-methide triterpenes

Quinone-methide triterpenes are the characteristic orange - red pigments of the Celastraceae and Hippocrateaceae. In 1936 , the first quinone – methide triterpene , tripterin ( 15 ) , was discovered from *Tripterygium wilfordii* Hook. f.<sup>36</sup> and the three year later was again isolated from *Celastrus scandens* Linn., but was called celastrol<sup>37</sup>. In 1951, pristimerin ( 16 ) , the methyl ester of celastrol , was isolated from *Pristimera indica* ( Willd. ) A. C. Smith<sup>38</sup>. It took 30 years for the structure determination of the both compound to be completed.<sup>39-40</sup> Their main skeleton is 24-nor-D:A-friedooleanane triterpene with quinone - methide chromophore on ring A / B and hydroxyl group at C-3 position ( Figure 2 ). Up to the year 2002 , one hundred and six quinone - methide triterpenes have been found from nature. This group of compounds can be called celastroloids after the name of the first isolated quinone – methide , celastrol .<sup>41</sup>

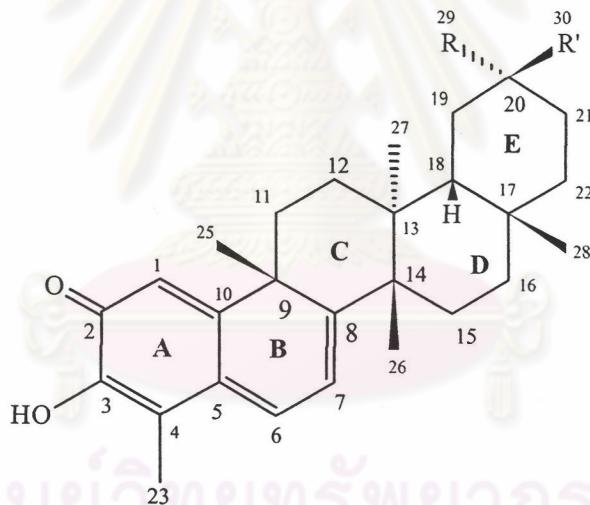


Figure 2. Main skeleton of quinone-methide triterpenes

### 2.1) Classification of quinone-methide triterpenes

Based on the A / B chromophores, quinone-methide triterpenes can be divided into 3 main classes and 5 subclasses as shown in Table1.

Table 1. Classification of quinone – methide triterpenes based on ring A /B chromophores.

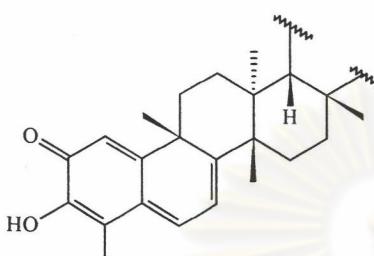
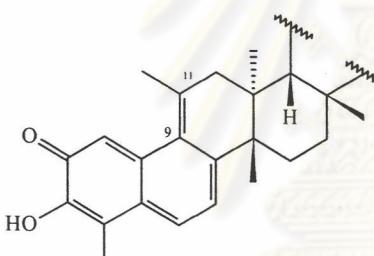
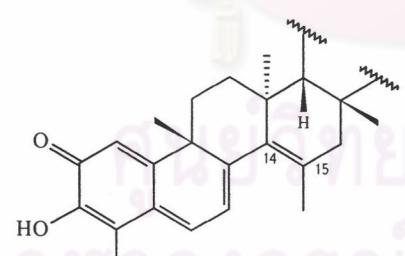
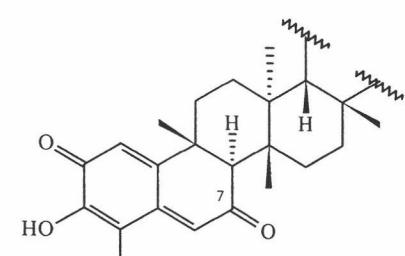
Class/Main Skeleton	Approximated UV $\lambda_{\max}$ ( solvent )	Ref.
<b>Class1 Typical quinone-methide triterpenes</b> Subclass1.1 Simple quinone-methide triterpenes	421 ( MeOH )	17,39
		
<b>Subclass 1.2 Ene-quinone-methide triterpenes</b> 1.2.1) 9(11)-Ene-(9→11)-quinone-methide triterpenes	446 ( MeOH )	39
		
1.2.2) 14(15)-Ene-(14→15) -quinone-methide triterpenes	444 ( EtOH )	40
		
<b>Subclass 1.3 7-Oxo-quinone-methide triterpenes</b>	321, 328, 409 ( EtOH )	41
		

Table1. ( continued )

Class/Main Skeleton	Approximated UV $\lambda_{\text{max}}$ ( solvent )	Ref.
<b>Class 2 Phenolic-D:A-friedo-24-noroleananes</b> Subclass 2.1 Phenolic-(9→8)D:A-friedo-24-noroleananes	305, 376 ( EtOH )	42
<b>Subclass 2.2 6-Oxo-phenolic-D:A-friedo-24-noroleananes</b>	307 ( MeOH )	18
<b>Class 3 Anhydride quinone-methide triterpenes</b>	392 ( EtOH )	43

Natural quinone - methide triterpenes mostly occur as monomers , although dimmers can be found. Nine type of monomeric quinone – methides can be categorized according to the substitution pattern of their E ring and the chromophore character. There are pristimerin, netzahualcoyone, excelsine, tingenone, 21-desoxotingenone, iguesterin, balaenol ,*Salacia* quinone-methide, and celastranhydride types. The dimeric quinone – methides are produced from the linkage of the pristimerin, tingenone or netzahualcoyone type. The linkage can be either one

or two ether-linkages. These naturally occurring quinone-methide triterpenes are summarized in Table 2 and 3.

Table 2. Naturally occurring monomeric quinone – methide triterpenes.

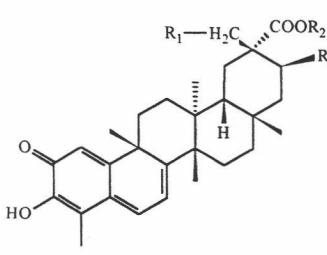
Compound / Structure	Plant source	Reference
<b>1.Pristimerin type</b>		
Pristimerin	<i>Acanthothamnus aphyllus</i> T. S. Brandegee	44
		
( R <sub>1</sub> =R <sub>3</sub> =H, R <sub>2</sub> =CH <sub>3</sub> )	<i>Austroplenckia populnea</i> ( Reiss. ) Lundell	45
	<i>Cassine balae</i> Kostermans	outer root- bark 40 , 46
	<i>Catha edulis</i> Forssk.	root bark 47-48
	<i>Celastrus paniculatus</i> Willd.	root bark 49 outer root-bark 43
	<i>Crossopetalum uragoga</i> O. Ktze.	root 50
	<i>Gymnosporia emarginata</i> ( Willd. ) Hook. f. ex Thw.	root 51
	<i>Gymnosporia montana</i> ( Roth ) Benth.	root bark and stem bark 52
	<i>Hippocratea excelsa</i> H. B. et K.	stem bark and root bark 53
	<i>Kokoona reflexa</i> Thw.	outer root-bark 43
	<i>Kokoona zeylanica</i> Thw.	outer stem-bark root bark 54 - 56 55
	<i>Maytenus boaria</i> Molina	root 57
	<i>Maytenus canariensis</i> ( Loes. ) Kunkel et Sunding	root bark root bark 56 , 58 25 , 57
	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	stem bark 18 , 59
	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	bark 60
	<i>Maytanus disperma</i> ( F.Muell. ) Loes.	outer root-bark 61
	<i>Maytenus horrida</i> Reiss	
	<i>Maytenus ilicifolia</i> Mart. ex. Reiss.	root bark 42 root cortex 62

Table 2. ( continued )

Compound / Structure	Plant source	Reference
Pristimerin ( continued )		
	<i>Maytenus obtusifolia</i> Mart.	root
	<i>Maytenus scutoides</i> ( Griseb. ) Lourteig et O'Donell	root bark
	<i>Maytenus umbellata</i> ( R. Br. ) Mabberley	root
	<i>Pachystigma canbyi</i> A. Gray	root bark
	<i>Plenckia polpunea</i> Reiss.	root
	<i>Pleurostylia opposita</i> ( Wall. ex Carey ) Alston	stem bark
	<i>Prionostemma aspera</i> Miers	root bark
	<i>Reissantia indica</i> ( Halle ) Ding Hou	root root bark
	<i>Rzedowskia tolantoguensis</i> F. Gonzalez -Medrano	root
	<i>Salacia beddomei</i> Gamble	stem bark
	<i>Salacia crassifolia</i> G. Don	
	<i>Salacia macrosperma</i> Wight	root bark
	<i>Salacia reticulata</i> Wight var. $\beta$ <i>diandra</i>	bark outer stem bark outer root bark
	<i>Salacia</i> sp.	root
	<i>Schaefferia cuneifolia</i> A. Gray	root bark root
	<i>Schaefferia cuneifolia</i> Standley	
	<i>Zinowiewia costaricensis</i> Lundell	root bark
	<i>Zinowiewia integerrima</i> Turcs.	root bark
Celastrol ( tripterin )	<i>Catha edulis</i> Forssk.	root bark
( R <sub>1</sub> =R <sub>2</sub> =R <sub>3</sub> =H )	<i>Celastrus paniculatus</i> Willd.	outer root - bark fresh aril
	<i>Celastrus scandens</i> Linn.	root bark
	<i>Celastrus strigulosus</i> Nakai	root
	<i>Hippocratea excelsa</i> H. B. et K.	stem bark and root bark
	<i>Kokoona ochracea</i> ( Elm. ) Mirrill	stem bark
		19

Table 2. ( continued )

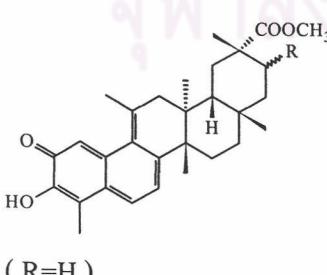
Compound / Structure	Plant source	Reference
Celastrol ( continued )		
	<i>Kokoona zeylanica</i> Thw.	
	outer stem-bark	55 - 56
	root bark	55
	soap cake	56
	<i>Maytenus canariensis</i> ( Loes. ) Kunkel et Sunding	root bark
		57
	<i>Maytenus horrida</i> Reiss.	22
	<i>Maytenus scutoides</i> ( Griseb. ) Lourteig et O' Donell	root bark
		26
	<i>Maytenus umbellata</i> ( R. Br. ) Mabberley	root
		63
	<i>Mortonia greggi</i> A. Gray	root
		83
	<i>Orthosphenia mexicana</i> Standley	root bark
		84
	<i>Salacia reticulata</i> Wight var. $\beta$ <i>diandra</i>	outer root-bark
		76
	<i>Schaefferia cuneifolia</i> Standley	
		22
	<i>Tripterygium wilfordii</i> Hook. f.	
	root	36 , 85
	tissue culture	86 - 89
	crude drug	90
	root bark	91
	<i>Tripterygium hypoglaucum</i> Hutchinson	root
		90
	<i>Tripterygium regelii</i> Sprague et Takeda	root
		82 , 92
21-Hydroxypristimerin ( R <sub>1</sub> = H , R <sub>2</sub> = CH <sub>3</sub> , R <sub>3</sub> = OH )	<i>Salacia</i> sp.	root bark
30-Hydroxy-pristimerin ( R <sub>1</sub> = OH , R <sub>2</sub> = CH <sub>3</sub> , R <sub>3</sub> = H )	<i>Salacia reticulata</i> Wight var. $\beta$ <i>diandra</i>	outer root bark
Pristimerinene	<i>Prionostemma aspera</i> Miers	root bark
		39

Table 2. ( continued )

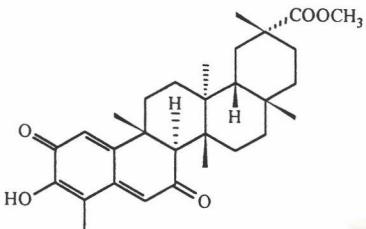
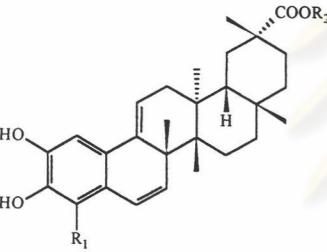
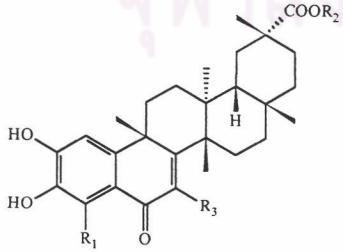
Compound/ Structure	Plant source	Reference
21-Hydroxypristimerinene ( R = OH )	<i>Salacia</i> sp.	root bark
Dispermoquinone	<i>Austroplenckia populnea</i> ( Reiss ). Lundell	root bark
	<i>Austroplenckia populnea</i> ( Reiss ). Lundell var. <i>ovata</i>	bark wood
	<i>Maytenus disperma</i> ( F. Muell. ) Loes.	outer root- bark
Isopristimerin III	<i>Maytenus ebenifolia</i> Reiss.	95
	<i>Maytenus ilicifolia</i> Mart.ex Reiss.	root bark
( R <sub>1</sub> = R <sub>2</sub> = CH <sub>3</sub> )		42
23-Oxoisopristimerin III ( R <sub>1</sub> = CHO, R <sub>2</sub> = CH <sub>3</sub> )	<i>Kokoona zeylanica</i> Thw.	outer stem-bark
		root bark
		inner root bark
		soap cake
Wilforol B ( R <sub>1</sub> = CH <sub>3</sub> , R <sub>2</sub> = H )	<i>Tripterygium wilfordii</i> Hook. f.	root bark
Zeylasterone	<i>Celastrus paniculatus</i> Willd.	outer root- bark
	<i>Kokoona reflexa</i> Thw.	outer root- bark
( R <sub>1</sub> = COOH , R <sub>2</sub> = CH <sub>3</sub> , R <sub>3</sub> = H )	<i>Kokoona zeylanica</i> Thw.	outer stem-bark
		root bark
		inner bark

Table 2. ( continued )

Compound / Structure	Plant source		Reference
Zelasterol ( R <sub>1</sub> = CHO, R <sub>2</sub> = CH <sub>3</sub> , R <sub>3</sub> = H )	<i>Celastrus paniculatus</i> Willd.	outer root- bark	43
	<i>Kokoona reflexa</i> Thw.	outer root- bark	43
	<i>Kokoona zeylanica</i> Thw.	outer stem-bark root bark soap cake	55-56 , 98 55 56
Desmethylzeylasterone ( R <sub>1</sub> = COOH , R <sub>2</sub> = H , R <sub>3</sub> = H )	<i>Kokoona zeylanica</i> Thw.	outer stem-bark root bark	55-56 , 98 55
	<i>Tripterygium wilfordii</i> Hook. f.	root bark	91
Desmetylzeylasterol ( R <sub>1</sub> = CHO, R <sub>2</sub> = H , R <sub>3</sub> = H )	<i>Kokoona zeylanica</i> Thw.	outer stem-bark root bark outer root-bark	55 55 56 , 99
	<i>Tripterygium wilfordii</i> Hook. f.	root bark	91
Wilforol A ( R <sub>1</sub> = CH <sub>3</sub> , R <sub>2</sub> =H , R <sub>3</sub> = H )	<i>Tripterygium wilfordii</i> Hook. f.	root bark	91
6-Oxopristimerol ( R <sub>1</sub> = R <sub>2</sub> = CH <sub>3</sub> , R <sub>3</sub> = H )	<i>Maytenus canariensis</i> ( Loes. ) Kunkel et Sunding	root bark	25
	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	stem bark	18
23-Nor-6-oxopristimerol ( R <sub>1</sub> = H , R <sub>2</sub> = CH <sub>3</sub> , R <sub>3</sub> = H )	<i>Kokoona zeylanica</i> Thw.	outer root-bark	99
23-Nor-6-oxodesmethyl- pristimerol	<i>Kokoona zeylanica</i> Thw.	outer stem-bark root bark outer root-bark	55 55 99
	<i>Tripterygium wilfordii</i> Hook. f.	root bark	91

Table 2. ( continued )

Compound / Structure	Plant source	Reference
2,3,7-Trihydroxy-6-oxo-1,3,5(10),7-tetraene-24-nor-friedelane-29-oic acid methylester  ( R <sub>1</sub> =CH <sub>3</sub> , R <sub>2</sub> =CH <sub>3</sub> , R <sub>3</sub> =OH )	<i>Crossopetalum gaumeri</i> ( Loes.) root	100
<b>2. Netzahualcoyone type</b>		
Netzahualcoyone  	<i>Maytenus horrida</i> Reiss.  <i>Orthosphenia mexicana</i> Standley  root bark	21 - 22  101  102
( R <sub>1</sub> =OH, R <sub>2</sub> R <sub>3</sub> =O , R <sub>4</sub> = CH <sub>3</sub> )	<i>Rzedowskia tolantoguensis</i> F. Gonzalez -Medrano  <i>Schaefferia cuneifolia</i> Standley	69 21  22
Netzahualcoyondiol  ( R <sub>1</sub> =OH , R <sub>2</sub> =H , R <sub>3</sub> =OH , R <sub>4</sub> =CH <sub>3</sub> )	<i>Maytenus horrida</i> Reiss.  <i>Orthosphenia mexicana</i> Standley  <i>Rzedowskia tolantoguensis</i> F. Gonzalez -Medrano	21  101  69 21
Netzahualcoyonol  ( R <sub>1</sub> =OH , R <sub>2</sub> =R <sub>3</sub> =H, R <sub>4</sub> =CH <sub>3</sub> )	<i>Maytenus horrida</i> Reiss  <i>Orthosphenia mexicana</i> Standley  <i>Rzedowskia tolantoguensis</i> F. Gonzalez -Medrano  <i>Schaefferia cuneifolia</i> Standley	21-22  101  69 21  22
Netzahualcoyene ( R <sub>1</sub> =R <sub>2</sub> =R <sub>3</sub> =H , R <sub>4</sub> =CH <sub>3</sub> )	<i>Maytenus horrida</i> Reiss  <i>Maytenus scutoides</i> ( Griseb. ) Lourteig et O'Donell  <i>Rzedowskia tolantoguensis</i> F. Gonzalez -Medrano  <i>Salacia reticulata</i> Wight var. $\beta$ -diandra  <i>Schaefferia cuneifolia</i> Standley	21-22, 101  root bark  26  21  outer root- bark  22

Table 2. ( continued )

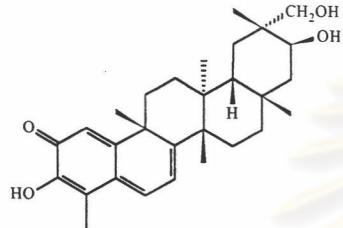
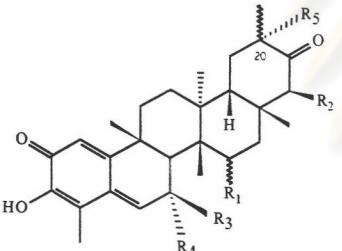
Compound / Structure	Plant source	Reference
Netzahualcoyol ( R <sub>1</sub> = OH , R <sub>2</sub> = R <sub>3</sub> = H , R <sub>4</sub> = COOCH <sub>3</sub> )	<i>Orthosphenia mexicana</i> Standley	101
<b>3. Excelsine type</b> Excelsine	<i>Hippocratea excelsa</i> H. B. et K.	53
	stem bark and root bark	
<b>4. Tingenone type</b> Tingenone ( tingenin A , maitenin )		
		
<i>Acanthothamnus aphyllus</i> T.S.Brandegee	root	44
<i>Cassine balae</i> Kostermans	outer root bark	40 , 46
<i>Cassine papillosa</i> ( Hochst. ) Kuntze	stem bark	103
<i>Maytenus canariensis</i> ( Loes. ) Kunkel et Sunding	root bark	57
<i>Catha edulis</i> Forssk.	root	47
	root bark	48
( 20 $\beta$ -CH <sub>3</sub> , R <sub>1</sub> = R <sub>2</sub> = H , 7,8-dehydro , R <sub>5</sub> = H )	<i>Crossopetalum uragoga</i> O. Ktze.	root bark and root medulla
		50
<i>Euonymus tingens</i> Wall.	stem bark	104
	stem bark	105
<i>Gymnosporia emarginata</i> ( Willd.) Hook. f. ex Thw.	root	51
<i>Gymnosporia montana</i> ( Roth ) Benth.	stem bark and root bark	52 , 106
<i>Hippocratea excelsa</i> H. B. et K.	stem bark and root bark	53
<i>Kokoona ochracea</i> ( Elm.) Mirill	stem bark	19
<i>Maytenus buchananii</i> (Loes.) R. Wilczek	tissue culture	20
<i>Maytenus canariensis</i> ( Loes. ) Kunkel et Sunding	root bark	25 , 58
<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	stem bark	18 , 59

Table 2. ( continued )

Compound / Structure	Plant source	Reference
Tingenone ( continued )		
	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas bark	60
	<i>Maytenus horrida</i> Reiss.	21 , 22
	<i>Maytenus ilicifolia</i> Mart. ex Reiss. root cortex	107
	bark	62
		108
	<i>Maytenus laevis</i> Reiss. root bark	109
	<i>Maytenus obtusifolia</i> Mart. root	57
	<i>Maytenus scutoides</i> ( Griseb. ) Lourteig et O'Donell root bark	26
	<i>Maytenus</i> sp.	110
	<i>Maytenus umbellata</i> ( R. Br. ) Mabberley root	63
	<i>Maytenus wallichiana</i> ( Spreng ex Wight et Arn. ) Raju et Babu tissue culture	111 - 112
	<i>Plenckia polpunea</i> Reiss. root	65
	<i>Prionostemma aspera</i> Miers root bark	39
	<i>Reissantia indica</i> ( Halle ) Ding Hou root bark	68
	<i>Rzedowskia tolantoguensis</i> F. Gonzalez - Medrano root	69
	<i>Rzedowskia tolantoguensis</i> F. Gonzalez - Medrano	21
	<i>Salacia macrosperma</i> Wight root bark	72 - 73
	<i>Salacia reticulata</i> Wight var. $\beta$ <i>diandra</i> outer root bark	76
	<i>Salacia</i> sp. root bark	39
	root	77
	<i>Schaefferia cuneifolia</i> A. Gray root	78
	<i>Schaefferia cuneifolia</i> Standley	22
	<i>Tripterygium wilfordii</i> Hook. f. tissue culture	87 - 89
	<i>Zinowiewia costarricensis</i> Lundell. root bark	79

Table 2. ( continued )

Compound / Structure	Plant source	Reference	
3,20 $\alpha$ -Dihydroxy-24,29-dinor-1(10),3,5,7-friedelatetraene-2,21-dione ( 20 $\beta$ - CH <sub>3</sub> , R <sub>1</sub> =R <sub>2</sub> =H , 7,8-dehydro, R <sub>5</sub> =OH )	<i>Glyptopetalum sclerocarpum</i> Laws.	stem bark	113
3,20 $\alpha$ -,22 $\beta$ -Trihydroxy-24,29-dinor-1(10),3,5,7-friedelatetraene-2,21-dione ( 20 $\beta$ - CH <sub>3</sub> , R <sub>1</sub> =H , R <sub>2</sub> =OH, 7,8-dehydro, R <sub>5</sub> =OH )	<i>Glyptopetalum sclerocarpum</i> Laws.	stem bark	113
3,20 $\beta$ ,22 $\beta$ -Trihydroxy-24,30-dinor-1(10),3,5,7-friedelatetraene-2,21-dione ( 20 $\beta$ - OH, R <sub>1</sub> =H , R <sub>2</sub> =OH , 7,8-dehydro, R <sub>5</sub> =CH <sub>3</sub> )	<i>Glyptopetalum sclerocarpum</i> Laws.	stem bark	113
3,20 $\alpha$ -Dihydroxy-24,29-dinor-1(10),3,5,7- friedelatetraene-2,21,22-trione ( 20 $\beta$ - CH <sub>3</sub> , R <sub>1</sub> =H , R <sub>2</sub> =O, 7,8-dehydro , R <sub>5</sub> =OH )	<i>Glyptopetalum sclerocarpum</i> Laws.	stem bark	113
3,20 $\beta$ -Dihydroxy-24,30-dinor-1(10),3,5,7- friedelatetraene-2,21,22-trione ( 20 $\beta$ - OH , R <sub>1</sub> =H , R <sub>2</sub> =O, 7,8-dehydro , R <sub>5</sub> =CH <sub>3</sub> )	<i>Glyptopetalum sclerocarpum</i> Laws.	stem bark	113
7,8-Dihydro-7-oxo-22- $\beta$ -hydroxytingenone ( 20 $\beta$ - CH <sub>3</sub> , R <sub>1</sub> =H, R <sub>2</sub> =OH , R <sub>3</sub> =R <sub>4</sub> =O, R <sub>5</sub> =H )	<i>Maytenus amazonica</i> C. Martius	root bark	114
7,8-Dihydro-22- $\beta$ -hydroxy-tingenone ( 20 $\beta$ - CH <sub>3</sub> , R <sub>1</sub> =H , R <sub>2</sub> =OH , R <sub>3</sub> =R <sub>4</sub> =R <sub>5</sub> =H )	<i>Maytenus amazonica</i> C. Martius	root bark	114
(8S)-7,8-Dihydro-7-oxo-tingenone ( 20 $\beta$ - CH <sub>3</sub> , R <sub>1</sub> =R <sub>2</sub> =H , R <sub>3</sub> =R <sub>4</sub> =O , R <sub>5</sub> =H )	<i>Maytenus amazonica</i> C. Martius	root bark	115
(7S,8S)-7-Hydroxy-7,8-dihydro-tingenone ( 20 $\beta$ - CH <sub>3</sub> , R <sub>1</sub> =R <sub>2</sub> =R <sub>3</sub> =H , R <sub>4</sub> =OH , R <sub>5</sub> =H )	<i>Maytenus amazonica</i> C. Martius	root bark	115

Table 2. ( continued )

Compound / Structure	Plant source	Reference	
Tingenin B ( 22 $\beta$ -hydroxytingenone ) ( 20 $\beta$ -CH <sub>3</sub> , R <sub>1</sub> =H , R <sub>2</sub> =OH , 7,8 - dehydro , R <sub>5</sub> = H )	<i>Acanthothamnus aphyllus</i> T. S. Brandegee <i>Cassine balae</i> Kostermans <i>Cassine papillosa</i> ( Hochst. ) Kuntze <i>Catha edulis</i> Forssk. <i>Euonymus tingens</i> Wall. <i>Glyptopetalum sclerocarpum</i> Laws. <i>Maytenus buchananii</i> ( Loes. ) R.Wilczek <i>Maytenus canariensis</i> ( Loes. ) Kunkel et Sunding <i>Maytenus chuchuhuasca</i> R. Hamet et Colas <i>Maytenus laevis</i> Reiss. <i>Maytenus obtusifolia</i> Mart. <i>Maytenus</i> sp. <i>Salacia reticulata</i> Wight var. $\beta$ - <i>diandra</i> <i>Tripterygium wilfordii</i> Hook. f.	root outer root- bark stem bark root bark stem bark stem bark tissue culture root bark stem bark root bark root root outer root bark tissue culture	44 40 , 46 103 47 - 48 104 17 20 25 18 60 109 57 110 76 89 46 45 40 - 46 105 116 19 69 72 - 73
15 $\alpha$ ,22 $\beta$ -Dihydroxytingenone ( 20 $\beta$ -CH <sub>3</sub> , R <sub>1</sub> =R <sub>2</sub> = OH , 7,8 - dehydro , R <sub>5</sub> = H )	<i>Cassine balae</i> Kostermans	outer root - bark	46
20-Hydroxy-20-epi-tingenone ( 20 $\beta$ -OH , R <sub>1</sub> =R <sub>2</sub> =H , 7,8- dehydro , R <sub>5</sub> =CH <sub>3</sub> )	<i>Austroplenckia populnea</i> ( Reiss. ) Lundell <i>Cassine balae</i> Kostermans <i>Euonymus tingens</i> Wall. <i>Glyptopetalum sclerocarpum</i> Laws <i>Kokoona ochracea</i> ( Elm. ) Mirrill <i>Rzedowskia tolantoguensis</i> F. Gonzalez-Medrano <i>Salacia macrosperma</i> Wight	root bark outer root - bark bark stem bark stem bark stem bark root root bark	40 - 46 105 116 19 69 72 - 73

Table 2. ( continued )

Compound / Structure	Plant source	Reference
Isotingenone III ( R <sub>1</sub> =CH <sub>3</sub> , R <sub>2</sub> =R <sub>3</sub> =H )	<i>Maytenus ebenifolia</i> Reiss. <i>Maytenus ilicifolia</i> Mart. ex Reiss. root bark	95 42
23-Oxo-isotingenone ( R <sub>1</sub> =CHO , R <sub>2</sub> =R <sub>3</sub> =H )	<i>Maytenus amazonica</i> C. Martius	root bark
2,3,22 $\beta$ -trihydroxy-24,29-dinor-25(9 $\rightarrow$ 8)-1,3,5(10),7-friedelatetraene-21-one-23-al ( R <sub>1</sub> =CHO , R <sub>2</sub> =OH , R <sub>3</sub> =H )	<i>Maytenus amazonica</i> C. Martius	root bark
6-Oxotingenol	<i>Maytenus canariensis</i> ( Loes.) Kunkel et Sunding <i>Maytenus ilicifolia</i> Mart. ex Reiss.	root bark
( R <sub>1</sub> =R <sub>2</sub> =H , R <sub>3</sub> =CH <sub>3</sub> )		
(8S)-7,8-Dihydro-6-oxo-tingenol ( R <sub>1</sub> =R <sub>2</sub> =H , R <sub>3</sub> =CH <sub>3</sub> )	<i>Maytenus amazonica</i> C. Martius	root bark
23-Nor-6-oxo-tingenol ( R <sub>1</sub> =R <sub>2</sub> =H , R <sub>3</sub> =H, 7,8-dehydro )	<i>Maytenus amazonica</i> C. Martius	root bark
2,3,22 $\beta$ -Trihydroxy-24,29-dinor-1,3,5(10),7-friedelatetraene-6,21-dione-23-al ( R <sub>1</sub> =H , R <sub>2</sub> =OH , R <sub>3</sub> =CHO , 7,8-dehydro )	<i>Maytenus amazonica</i> C. Martius	root bark
22 $\beta$ -Hydroxy-6-oxo-tingenol ( R <sub>1</sub> =H , R <sub>2</sub> =OH , R <sub>3</sub> =CH <sub>3</sub> , 7,8 - dehydro )	<i>Maytenus amazonica</i> C. Martius	root bark

Table 2. ( continued )

Compound / Structure	Plant source		Reference
23-Nor-22-hydroxy-6-oxo-tingenol ( R <sub>1</sub> = H , R <sub>2</sub> = OH , R <sub>3</sub> = H , 7,8-dehydro )	<i>Maytenus amazonica</i> C. Martius	root bark	114
3-Methoxy-22 $\beta$ -hydroxy-6-oxo-tingenol ( R <sub>1</sub> = CH <sub>3</sub> , R <sub>2</sub> = OH , R <sub>3</sub> = CH <sub>3</sub> , 7,8-dehydro )	<i>Maytenus amazonica</i> C. Martius	root bark	114
22 $\beta$ -Hydroxy-7,8-dihydro-6-oxo-tingenol ( R <sub>1</sub> = H , R <sub>2</sub> = OH , R <sub>3</sub> = CH <sub>3</sub> )	<i>Maytenus amazonica</i> C. Martius	root bark	114
3-Methyl-6-oxotingenol ( R <sub>1</sub> = CH <sub>3</sub> , R <sub>2</sub> = H , R <sub>3</sub> = CH <sub>3</sub> )	<i>Maytenus canariensis</i> ( Loes. ) Kunkel et Sunding	root bark	25
	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	stem bark	95 18
3-Methyl-22 $\beta$ ,23-dihydroxy-6-oxotingenol ( R <sub>1</sub> = CH <sub>3</sub> , R <sub>2</sub> = OH , R <sub>3</sub> = CH <sub>3</sub> )	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	stem bark	18
<b>5. 21-desoxotingenone type</b>			
3,20 $\alpha$ ,21 $\alpha$ -Trihydroxy-24,29-dinor-1(10),3,5,7-friedelatetraene-2,22-dione	<i>Glyptopetalum sclerocarpum</i> Laws.	stem bark	113
30(20→21)abeo -3,21 $\alpha$ -Dihydroxy-24,29-dinor-1(10),3,5,7-friedelatetraene-2,20,22-trione	<i>Glyptopetalum sclerocarpum</i> Laws.	stem bark	113

Table 2. ( continued )

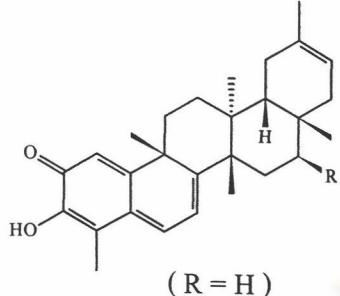
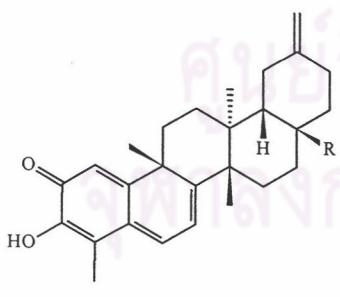
Compound / Structure	Plant source	Reference
<b>6. Igesterin type</b>		
Igesterin	<i>Catha edulis</i> Forsk	root bark
	<i>Gymnosporia emarginata</i> ( Willd. ) Hook. f. ex Thw.	root
( R = H )	<i>Gymnosporia montana</i> ( Roth ) Benth.	stem bark and root bark
	<i>Maytenus canariensis</i> ( Loes. ) Kunkel et Sunding	root bark
	<i>Maytenus horrida</i> Reiss.	
	<i>Maytenus umbellata</i> ( R. Br. ) Mabberly	root
	<i>Rzedowskia tolantoguensis</i> F. Gonzalez -Medrano	
	<i>Salacia reticulata</i> Wight var. $\beta$ -diandra	bark
16 $\beta$ -Hydroxyigesterin ( R = OH )	<i>Maytenus canariensis</i> ( Loes. ) Kunkel et Sunding	root bark
Isoigesterin	<i>Salacia madagascariensis</i> DC.	root
	<i>Salacia reticulata</i> Wight var. $\beta$ -diandra	root bark
( R = CH <sub>3</sub> )		outer root - bark
28-Nor-isoigesterin-17-carbaldehyde ( R = CHO )	<i>Salacia kraussii</i> ( Harv. )	root
		119

Table 2. ( continued )

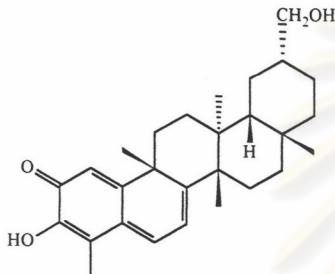
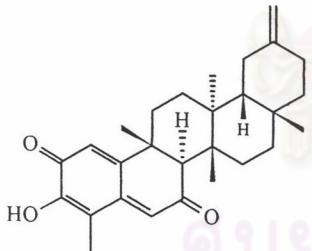
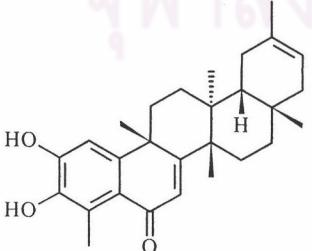
Compound/ Structure	Plant source	Reference
17-(Methoxycarbonyl)-28-nor- isoiguesterin ( R = COOMe )	<i>Salacia kraussii</i> ( Harv. ) root	119
28-Hydroxyisoiguesterin ( R = CH <sub>2</sub> OH )	<i>Salacia kraussii</i> ( Harv. ) root	119
Isoiguesterol	<i>Salacia reticulata</i> Wight var. $\beta$ - <i>diandra</i>	outer root - bark
		76
( R = CH <sub>2</sub> OH )		
Salacquinone	<i>Salacia reticulata</i> Wight var. $\beta$ - <i>diandra</i>	root bark outer root - bark
		41 76
6-Oxo-iguesterol	<i>Maytenus canariensis</i> ( Loes. ) Kunkel et Sunding	root bark
		25

Table 2. ( continued )

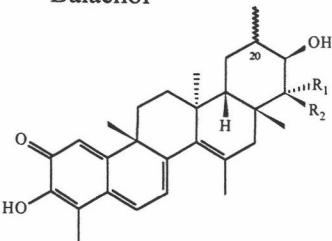
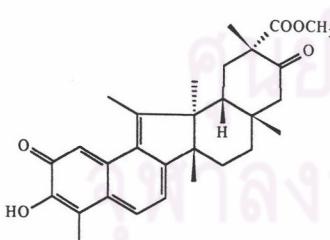
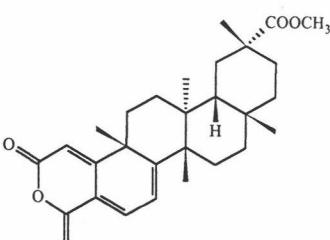
Compound / Structure	Plant source	Reference
<b>7. Balaenol type</b>		
Balaenol 	<i>Cassine balaee</i> Kostermans	outer root-bark      40 , 46 , 120
( 20 $\beta$ - CH <sub>3</sub> , R <sub>1</sub> = R <sub>2</sub> = H )		
Balaenonol ( 20 $\beta$ - CH <sub>3</sub> , R <sub>1</sub> R <sub>2</sub> = O )	<i>Cassine balaee</i> Kostermans	outer root-bark      40 , 46 , 120
Isobalaenol ( 20 $\alpha$ - CH <sub>3</sub> , R <sub>1</sub> = R <sub>2</sub> = H )	<i>Cassine balaee</i> Kostermans	outer root-bark      46
Isobalaendiol ( 20 $\alpha$ - CH <sub>3</sub> , R <sub>1</sub> = H , R <sub>2</sub> = OH )	<i>Cassine balaee</i> Kostermans	outer root-bark      40 , 46
<b>8. Salacia quinonemethide type</b>		
	<i>Salacia macrosperma</i> Wight	root bark      72 - 73
		
<b>9. Celastrananhydride type</b>		
Celastrananhydride 	<i>Cassine balaee</i> Kostermans	43
	<i>Kokoona reflexa</i> Thw.	43
	<i>Kokoona zeylanica</i> Thw.	43
	<i>Reissantia indica</i> ( Halle ) Ding Hou	43

Table 3. Naturally occurring dimeric quinone – methide triterpenes.

Compound/Structure	Plant source	Reference
<b>1. One - ether linkage dimers</b>		
<b>1.1) Pristimerin-pristimerin type</b> Magellanin	<i>Maytenus magellanica</i> Hook.f.	121
Rzedowskia bistriterpenoid ( 4 $\alpha$ -CH <sub>3</sub> )	<i>Rzedowskia tolantoguensis</i> F. Gonzalez - Medrano	122
4-Epimeric Rzedowskia bistriterpenoid ( 4 $\beta$ -CH <sub>3</sub> )	<i>Rzedowskia tolantoguensis</i> F. Gonzalez - Medrano	122

Table3. ( continued )

Compound/Structure	Plant source	Reference
<b>1.2) Tingenone - tingenone type</b>		
Umbellatin $\alpha$	<i>Maytenus umbellata</i> ( R. Br. ) root Mabberley	24
( 4 $\alpha$ -CH <sub>3</sub> )		
Umbellatin $\beta$ ( 4 $\beta$ -CH <sub>3</sub> )	<i>Maytenus umbellata</i> ( R. Br. ) root Mabberley	24
D:A-Friedo-24,30-dinor-oleana-1(10),5,7-triene-2,21-dione,3,4-epoxy-3-[(8 $\beta$ ,20 $\beta$ )-3-hydroxy-6,21-dioxo-D:A-friedo-24,30-dinoroleana-1,3,5(10)-trien-2-yl]oxy](4 $\xi$ ,20 $\beta$ )	<i>Maytenus chuchuhuasca</i> R. Hamel et Colas	95
( R <sub>1</sub> =R <sub>2</sub> =H )		

Table3. ( continued )

Compound/Structure	Plant source	Reference
D:A-Friedo-24,30-dinor-oleana-1,3,5(10),7-tetraene-6,21-dione ,2 [[4ξ,20β]-3,4-epoxy-2,21-dioxo- D:A-friedo-24,30-dinoroleana-1(10),5,7-trien-3-yl]oxy]-3-hydroxy (20β) ( R <sub>1</sub> = R <sub>2</sub> = H, 7,8 - dehydro )	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	95
D:A-Friedo-24,30-dinor-oleana-1,3,5(10),7-tetraene-6,21-dione,2[[4ξ,20β,22β]-3,4-epoxy-22-hydroxy-2,21-dioxo-D:A-friedo-24,30-dinoroleana-1(10),5,7-trien-3-yl]oxy]-3,22-dihydroxy ( 20β,22 β) ( R <sub>1</sub> = H , R <sub>2</sub> = OH, 7,8 - dehydro )	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	95
D:A-Friedo-24,30-dinor-oleana-1,3,5(10),7-tetraene-6,21-dione,2[[4ξ,20β,22β]-3,4-epoxy-22-hydroxy-2,21-dioxo-D:A-friedo-24,30-dinoroleana-1(10),5,7-trien-3-yl]oxy]-3-hydroxy (20β) ( R <sub>1</sub> = R <sub>2</sub> = OH , 7,8 - dehydro )	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	95

## 2. Two - ether linkage dimers

### 2.1) Pristimerin-pristimerin type

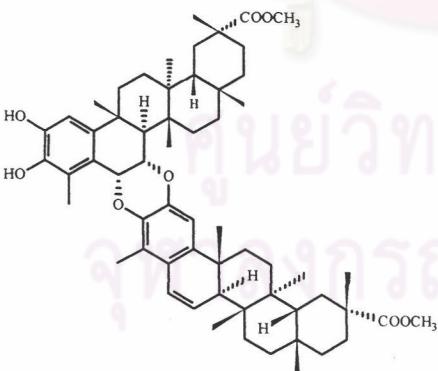
Cangorosin A	<i>Maytenus ilicifolia</i> Mart. ex Reiss.	root	95 , 123 - 124
			

Table3. ( continued )

Compound/Structure	Plant source	Reference
<b>Isocangorosin A</b>	<i>Maytenus ilicifolia</i> Mart. ex Reiss.	root 95 , 123 - 124
<b>6',7'- Dihydroisocangorosin A ( 6',7' - dihydro )</b>	<i>Maytenus ilicifolia</i> Mart. ex Reiss.	root 95 , 123 - 124
<b>Scutionin αA</b>	<i>Maytenus scutiooides</i> ( Griseb. ) Lourteig et O' Donell	root - bark 26
<b>( 3β - OH , 4β - CH<sub>3</sub> )</b>		
<b>7,8-Dihydro-scutionin αA ( 3β - OH , 4β - CH<sub>3</sub> , 7,8 - dihydro )</b>	<i>Maytenus scutiooides</i> ( Griseb. ) Lourteig et O'Donell	root - bark 26

Table3. ( continued )

Compound/Structure	Plant source	Reference
7,8-Dihydro-scutionin $\beta$ A ( 3 $\alpha$ - OH, 4 $\alpha$ - CH <sub>3</sub> , 7,8 - dihydro )	<i>Maytenus scutoides</i> ( Griseb.) Lourteig et O'Donell	root - bark 26
7,8-Dihydro-scutionin $\alpha$ B ( 3 $\beta$ - OH , 4 $\beta$ - CH <sub>3</sub> )	<i>Maytenus scutoides</i> ( Griseb.) Lourteig et O'Donell	root - bark 26
7,8-Dihydro-scutionin $\beta$ B ( 3 $\alpha$ - OH , 4 $\alpha$ - CH <sub>3</sub> )	<i>Maytenus scutoides</i> ( Griseb.) Lourteig et O'Donell	root - bark 26
Scutionin $\alpha$ B	<i>Maytenus blepharodes</i> Lundell	root 125
	<i>Maytenus magellanica</i> Lam.	root - bark 125
6',7'-Dihydroscutionin $\alpha$ B ( 6',7'- Dihydro )	<i>Maytenus blepharodes</i> Lundell	root 125
	<i>Maytenus magellanica</i> Lam.	root - bark 125
6' $\beta$ -methoxy-dihydro-scutionin $\alpha$ B ( 6',7'- Dihydro , 6' $\beta$ - methoxy )	<i>Maytenus blepharodes</i> Lundell	root 125
	<i>Maytenus magellanica</i> Lam.	root - bark 125

Table3. ( continued )

Compound/Structure	Plant source	Reference
Scutidin $\alpha$ A	<i>Maytenus scutoides</i> ( Griseb.) Lourteig et O'Donell	root - bark 26
7,8-dihydroisoxuxuarine E $\alpha$ ( R = H , 7,8-dihydro )	<i>Maytenus chuchuhuasca</i> Raymond - Hamet et Colas	bark 126
7,8-Dihydro-scutidin $\alpha$ B	<i>Maytenus scutoides</i> ( Griseb.) Lourteig et O'Donell	root bark 26
Xuxurine E $\alpha$	<i>Maytenus blepharodes</i> Lundell	root 127

Table3. ( continued )

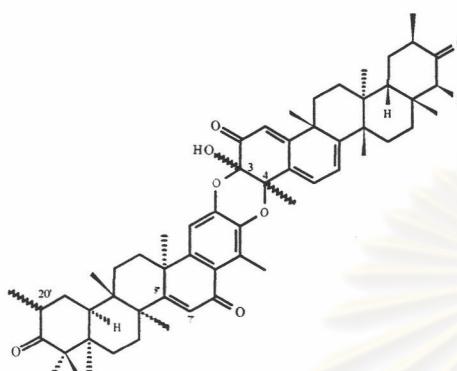
Compound/Structure	Plant source	Reference
<b>2.2 Tingenone - tingenone Type</b>		
Xuxuarine A $\alpha$	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	bark 60
( 3 $\beta$ - OH , 4 $\beta$ - CH <sub>3</sub> , 20' $\alpha$ - CH <sub>3</sub> , R <sub>1</sub> = R <sub>2</sub> = R <sub>3</sub> = H )		
Xuxuarine A $\beta$ ( 3 $\alpha$ - OH , 4 $\alpha$ - CH <sub>3</sub> , 20' $\alpha$ - CH <sub>3</sub> , R <sub>1</sub> = R <sub>2</sub> = R <sub>3</sub> = H )	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	bark 60
Xuxuarine B $\alpha$ ( 3 $\beta$ - OH , 4 $\beta$ - CH <sub>3</sub> , 20' $\alpha$ - CH <sub>3</sub> , R <sub>1</sub> = R <sub>3</sub> = OH , R <sub>2</sub> = H )	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	bark 60
Xuxuarine B $\beta$ ( 3 $\alpha$ - OH , 4 $\alpha$ - CH <sub>3</sub> , 20' $\alpha$ - CH <sub>3</sub> , R <sub>1</sub> = R <sub>3</sub> = OH , R <sub>2</sub> = H )	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	bark 60
Xuxuarine C $\alpha$ ( 3 $\beta$ - OH , 4 $\beta$ - CH <sub>3</sub> , 20' $\alpha$ - CH <sub>3</sub> , R <sub>1</sub> = OH , R <sub>2</sub> = R <sub>3</sub> = H )	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	bark 60
Xuxuarine C $\beta$ ( 3 $\alpha$ - OH , 4 $\alpha$ - CH <sub>3</sub> , 20' $\alpha$ - CH <sub>3</sub> , R <sub>1</sub> = OH , R <sub>2</sub> = R <sub>3</sub> = H )	<i>Maytenus chuchuhuasca</i> R.Hamet et Colas	bark 60
Xuxuarine D $\alpha$ ( 3 $\beta$ - OH , 4 $\beta$ - CH <sub>3</sub> , 20' $\alpha$ - CH <sub>3</sub> , R <sub>1</sub> = R <sub>3</sub> = H , R <sub>2</sub> = OH )	<i>Maytenus chuchuhuasca</i> R.Hamet et Colas	bark 60
Xuxuarine D $\beta$ ( 3 $\alpha$ - OH , 4 $\alpha$ - CH <sub>3</sub> , 20' $\alpha$ - CH <sub>3</sub> , R <sub>1</sub> = R <sub>3</sub> = H , R <sub>2</sub> = OH )	<i>Maytenus chuchuhuasca</i> R.Hamet et Colas	bark 60

Table3. ( continued )

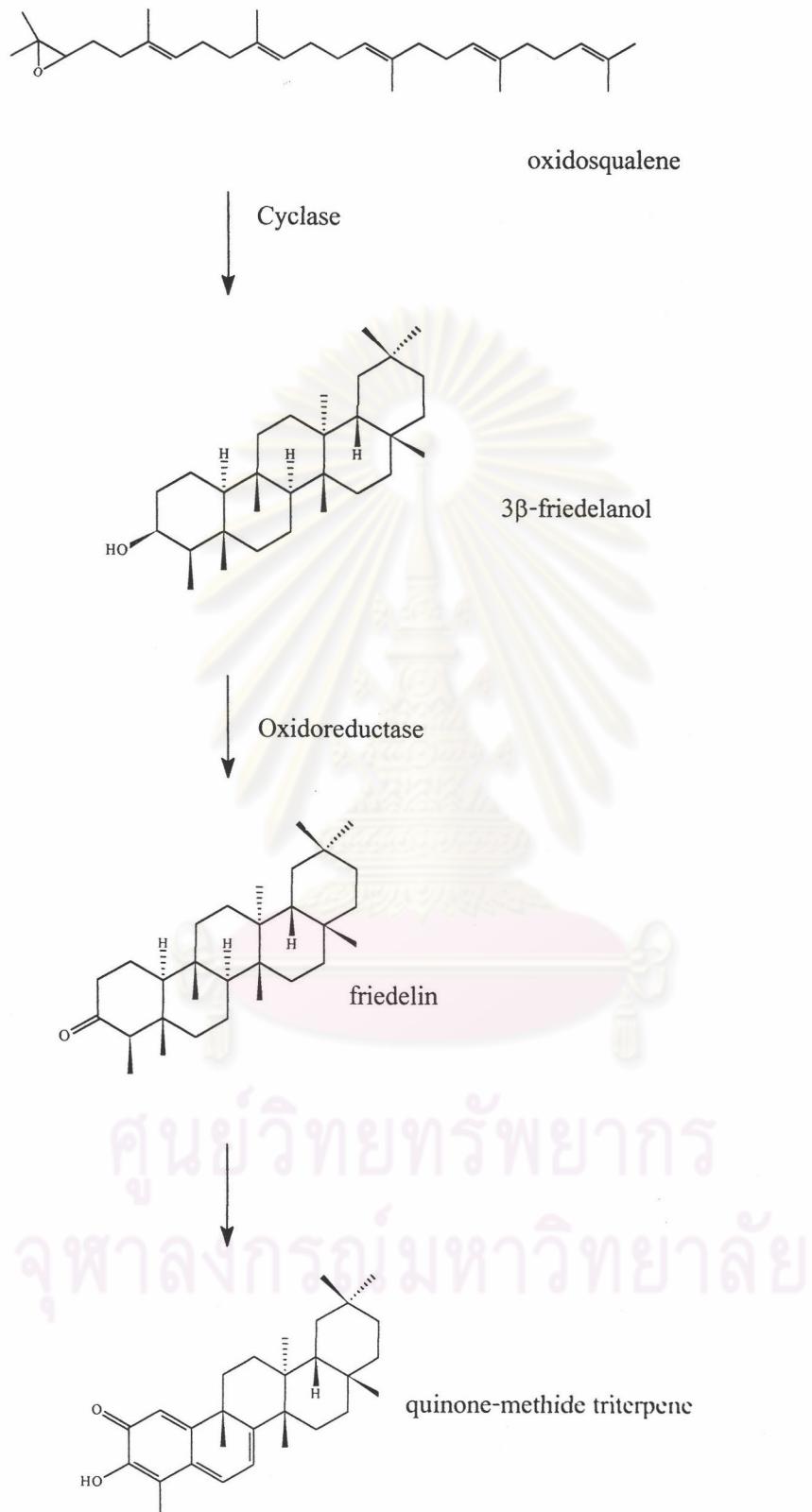
Compound/Structure	Plant source	Reference
7',8'-Dihydroxuxuarine A $\beta$ ( 3 $\alpha$ -OH , 4 $\alpha$ -CH <sub>3</sub> , 20' $\alpha$ -CH <sub>3</sub> , 7',8'dihydro, R <sub>1</sub> =R <sub>2</sub> =R <sub>3</sub> =H )	<i>Maytenus chuchuhuasca</i> R.Hamet et Colas	bark 60
<b>2.3 Pristimerin – tingenone type</b>		
Cangorosin B	<i>Maytenus ilicifolia</i> Mart. ex Reiss.	95 , 124
Xuxuarine F $\beta$ ( R = H )	<i>Maytenus chuchuhuasca</i> R.Hamet et Colas	stem bark 126

Table3. ( continued )

Compound/Structure	Plant source	Reference
Xuxuarine G $\alpha$ ( R = H, cis 3,4 - dioxy bond = $\alpha$ )	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	stem bark
		126
Xuxuarine G $\beta$ ( R = H, cis 3,4 - dioxy bond = $\beta$ )	<i>Maytenus chuchuhuasca</i> R. Hamet et Colas	stem bark
		126
<b>2.4 Pristimerin - netzahualcoyone type Netzascutionin <math>\alpha</math> A</b>	<i>Maytenus scutoides</i> ( Griseb.) Lourteig et O'Donell	root bark
		26

## 2.2) Biosynthesis of quinone-methide triterpenes

The entry point for biosynthesis of quinone-methides requires oxidosqualene as a central intermediate, which by the action of a cyclase would give rise to the first cyclic intermediate, 3 $\beta$ -friedelanol and then, by the action of an oxidoreductase, the conversion to friedelin should occur between leaves and root bark. The transformation/translocation steps for the quinone-methide triterpenes should take place in the root bark.<sup>128</sup> ( Scheme 1. )



Scheme1. Conversion of oxidosqualene to  $3\beta$ -friedelanol and friedelin and their involvement as the biosynthetic precursor to the quinone - methide triterpene.

### 3. Oleanane Triterpenes in Celastraceae

Naturally occurring oleanane triterpenes found in the family Celastraceae up to the year 2002 are summarized in Figure 3 and Table 4.

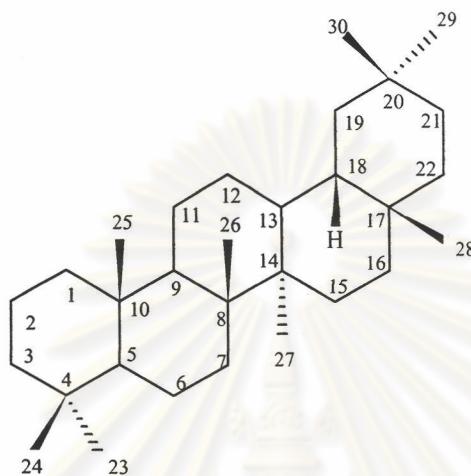


Figure 3. Basic structure of oleanane triterpene

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จุฬาลงกรณ์มหาวิทยาลัย

Table 4. Naturally occurring oleanane triterpenes in Celastraceae.

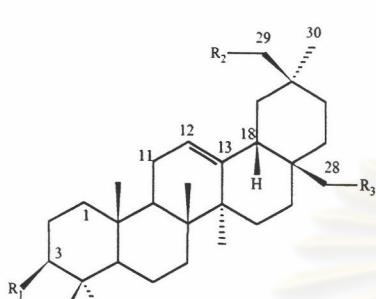
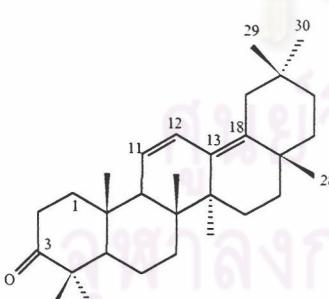
Compound / Structure	Plant source	Reference
$\beta$ -amyrin ( R <sub>1</sub> = OH , R <sub>2</sub> = R <sub>3</sub> = H )	<i>Acanthothamus aphyllus</i> T. S. Brandegree	root 44
	<i>Celastrus paniculatus</i> Willd. <i>Celastrus scandens</i> Linn. <i>Lophopetalum wightianum</i> Wight ex Arn.	3 3 3
	<i>Maytenus arbutifolia</i> ( A. Rich. ) Wilezek	leaves 129
	<i>Maytenus canariensis</i> ( Loes. ) Kunkel et Sunding	3
	<i>Maytenus heterophylla</i> ( Eckl. & Zeyh. )	stem bark 129
Paniculatadiol ( R <sub>1</sub> = OH , R <sub>2</sub> = OH , R <sub>3</sub> = H )	<i>Celastrus paniculatus</i> Willd	3
Erythrodiol ( R <sub>1</sub> = OH , R <sub>2</sub> = H , R <sub>3</sub> = OH )	<i>Pristimera grahamii</i> Miers	3
3-oxo-olean-11, 13(18)-ene	<i>Maytenus disperma</i> ( F. Muell. ) Loes.	3
		

Table 4. ( continued )

Compound / Structure	Plant source	Reference
Crataegolic acid	<i>Euonymus latifolius</i> L.	3
3 $\alpha$ ,23-Dihydroxy-olean-12-en-28-oic acid ( R <sub>1</sub> = OH , R <sub>2</sub> = CH <sub>3</sub> , R <sub>3</sub> = CH <sub>2</sub> OH )	<i>Tripterygium wilfordii</i> Hook.f.	root bark 130
3 $\alpha$ ,24-Dihydroxy-olean-12-en-28-oic acid ( R <sub>1</sub> = OH , R <sub>2</sub> = CH <sub>2</sub> OH , R <sub>3</sub> = CH <sub>3</sub> )	<i>Tripterygium wilfordii</i> Hook.f.	root bark 130
3 $\beta$ ,22 $\alpha$ -Dihydroxyolean-12-en-29-oic acid ( R = $\alpha$ - OH )	<i>Salacia oblonga</i> Wall.	root 35
3 $\beta$ ,22 $\beta$ -Dihydroxyolean-12-en-29-oic acid ( R = $\beta$ - OH )	<i>Tripterygium wilfordii</i> Hook. f.	131

Table 4. ( continued )

Compound / Structure	Plant source	Reference
Germacinone	<i>Acanthothamnus aphyllus</i> T.S. Brandegee	root 44
Hypodiol	<i>Tripterygium hypoglaucum</i> (Levl.) Hutch	outer root bark 132
3-Oxo-olean-12-en-29-oic acid ( R <sub>1</sub> =O , R <sub>2</sub> =COOH , R <sub>3</sub> =H , R <sub>4</sub> =CH <sub>3</sub> , , R <sub>5</sub> =CH <sub>3</sub> )	<i>Austroplenckia populnea</i> ( Reiss ) Lundell	bark wood 131
Mupinensisone ( R <sub>1</sub> =O , R <sub>2</sub> =OH , R <sub>3</sub> =H , R <sub>4</sub> =CH <sub>3</sub> , R <sub>5</sub> =CH <sub>3</sub> )	<i>Euonymus mupinensis</i> L.	131
29-Hydroxy-3-oxo-olean-12-en-28-oic acid ( R <sub>1</sub> =O , R <sub>2</sub> =CH <sub>2</sub> OH , R <sub>3</sub> =H , R <sub>4</sub> = COOH , R <sub>5</sub> =CH <sub>3</sub> )	<i>Tripterygium wilfordii</i> Hook.f.	root 5
22 $\alpha$ -Hydroxy-3-oxo-olean-12-en-29-oic acid ( R <sub>1</sub> =O , R <sub>2</sub> =COOH , R <sub>3</sub> = $\alpha$ -OH , R <sub>4</sub> = R <sub>5</sub> =CH <sub>3</sub> )	<i>Tripterygium wilfordii</i> Hook.f.	root 5

Table 4. ( continued )

Compound / Structure	Plant source	Reference
22 $\beta$ -Hydroxy-3-oxo-olean-12-en-29-oic acid ( R <sub>1</sub> = O , R <sub>2</sub> = COOH , R <sub>3</sub> = $\beta$ - OH , R <sub>4</sub> = R <sub>5</sub> = CH <sub>3</sub> )	<i>Tripterygium wilfordii</i> Hook.f.	root 5
24-Hydroxy-3-oxo-olean-12-en-28-oic acid ( R <sub>1</sub> = O , R <sub>2</sub> = CH <sub>3</sub> , R <sub>3</sub> = H , R <sub>4</sub> = COOH , R <sub>5</sub> = $\beta$ CH <sub>2</sub> OH )	<i>Tripterygium wilfordii</i> Hook.f.	root 5
23-Hydroxy-3-oxo-olean-12-en-28-oic acid ( R <sub>1</sub> = O , R <sub>2</sub> = CH <sub>3</sub> , R <sub>3</sub> = H , R <sub>4</sub> = COOH , R <sub>5</sub> = $\alpha$ CH <sub>2</sub> OH )	<i>Tripterygium wilfordii</i> Hook.f.	root 5
Methyl katonate ( R <sub>1</sub> = O , R <sub>2</sub> = COOCH <sub>3</sub> , R <sub>3</sub> = H , R <sub>4</sub> = CH <sub>3</sub> , R <sub>5</sub> = CH <sub>3</sub> )	<i>Tripterygium wilfordii</i> var. <i>regelii</i> Makino	stem 133
Maytenfolic acid	<i>Acanthothamnus aphyllus</i> T.S. Brandegee	root 44
	<i>Maytenus heterophylla</i> ( Eckl. & Zeyh. )	stem bark 129
	<i>Salacia oblonga</i> Wall.	root 35
Mesembryanthemoidigenic acid	<i>Tripterygium hypoglaucum</i> ( Levl. ) Hutch	outer root bark 132

Table 4. ( continued )

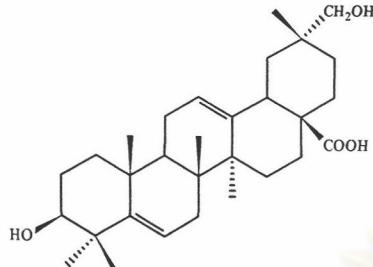
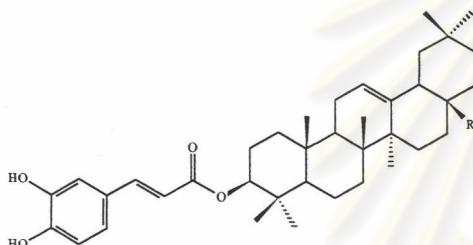
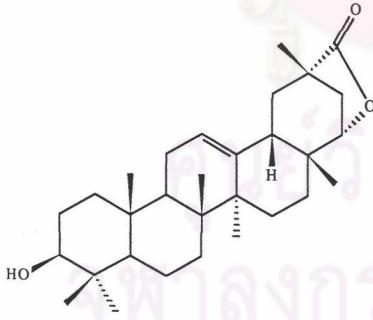
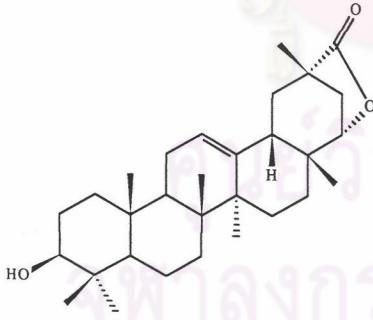
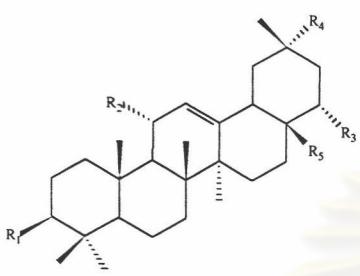
Compound / Structure	Plant source	Reference
Hypoglauterpenic acid	<i>Tripterygium hypoglauicum</i> Hutchinson	131
		
Olean-28-al-3 $\beta$ -yl caffeoate ( R = CHO )	<i>Celastrus stephanofolius</i> L.	stem 134
		
Olean-28-oic-3 $\beta$ -yl caffeoate ( R = COOH )	<i>Celastrus stephanofolius</i> L.	stem 134
		
Regelide	<i>Tripterygium regeli</i> Hook. f.	131
		
	<i>Tripterygium wilfordii</i> Hook. f.	root 133
	<i>Tripterygium wilfordii</i> var. <i>regelii</i> Makino	dry stalk 133

Table 4. ( continued )

Compound / Structure	Plant source	Reference
Triptohypol F ( R <sub>1</sub> = OH, R <sub>2</sub> = OCH <sub>3</sub> , R <sub>3</sub> = H , R <sub>4</sub> = R <sub>5</sub> = CH <sub>3</sub> )	<i>Tripterygium hypoglauicum</i> ( Levl.)	root bark      135
		
Oleanic acid 3-O-acetate ( R <sub>1</sub> = OCOCH <sub>3</sub> , R <sub>2</sub> = R <sub>3</sub> = H , R <sub>4</sub> = CH <sub>3</sub> , R <sub>5</sub> = COOH )	<i>Tripterygium hypoglauicum</i> ( Levl. )	root bark      135
Triptocallic acid D ( R <sub>1</sub> = α - OH , R <sub>2</sub> = H , R <sub>3</sub> = OH , R <sub>4</sub> = COOH , R <sub>5</sub> = CH <sub>3</sub> )	<i>Tripterygium hypoglauicum</i> ( Levl.)	root bark      135
	<i>Tripterygium wilfordii</i> var. <i>regelii</i> Makino	callus- culture      136
3-Epikatonic acid ( R <sub>1</sub> = OH , R <sub>2</sub> = R <sub>3</sub> = H , R <sub>4</sub> = COOH , R <sub>5</sub> = CH <sub>3</sub> )	<i>Tripterygium hypoglauicum</i> ( Levl.)	root bark      135
3β,15α- Dihydroxy-olean-12-ene	<i>Schaefferia cuneifolia</i> Jacq.	131
