## CHAPTER II

## HISTORICAL

Meliaceae contains a wide range of chemical compounds. Some of these compounds, such as the triterpenoids, are biosynthesized from the mevalonate pathway. Other compounds are biosynthesized from the shikimate pathyway, i.e. flavonoids, coumarins, lignans and flavaglines.

The only information on alkaloid content in genus Chisocheton was reported by Tzouros in which two spermidine alkaloids, Chisitine 1 [1.1] and Chisitine 2 [1.2], were isolated from the leaves of $C$. weinlandii.


Reviews of the triterpenoid constituents of Chisocheton species, dammaranetype, aromadendrane-type sesquiterpenes and together with a review of flavagline derivatives, are present herein.

## Chemical constituents of Chisocheton spp.

Chemical investigations of a number of Chisocheton spp. have shown them to be a good source of triterpenoids. The literature reviews of triterpenoids from $C$. paniculatus, C. microcarpus and C. macrophyllus are summarized in Table 2.

Table 2 The distribution of triterpenoids in Chisocheton spp.

| Compounds | Source | Part | References |
| :--- | :---: | :---: | :---: |
| Apo-tirucallol <br> triterpenes <br> Compound A [1.3] | Chisocheton <br> paniculatus | Wood | Connolly et al., 1979 |

Table 2 The distribution of triterpenoids in Chisocheton spp. (continued)

| Compounds | Source | Part | References |
| :---: | :---: | :---: | :---: |
| Compound B [1.4] | C. paniculatus | Wood | Connolly et al., 1979 |
| Compound C [1.5] | C. paniculatus | Wood | Connolly et al., 1979 |
| Compound D [1.6] | C. paniculatus | Wood | Connolly et al., 1979 |
| Paniculatin B [1.7] | C. paniculatus | Root wood | $\begin{aligned} & \text { Yadav } \text { et al., 1999a, } \\ & \text { 1999b } \end{aligned}$ |
| Paniculatin C [1.8] | C. paniculatus | Root wood | $\begin{aligned} & \text { Yadav } \text { et al., 1999a, } \\ & \text { 1999b } \end{aligned}$ |
| Paniculatin D [1.9] | C. paniculatus | Root wood | $\begin{aligned} & \text { Yadav } \text { et al., 1999a, } \\ & \text { 1999b } \end{aligned}$ |
| Tirucallane triterpenes <br> Arunachalin [1.10] | C. paniculatus | Root wood | Yadav et al., 1999a, 1999b, 1999c |
| Tetranortriterpenoids 6 $\alpha$-Acetoxyazadirone (Paniculatin) [1.11] | C. paniculatus | Fruits | Saikia et al., 1978; <br> Bhattacharyya et al., <br> 2004 |
|  | C. paniculatus | Seeds | Chatterjee et al., 1989 |
|  | C. paniculatus | Root wood | Yadav et al., 1999a |
| 6 $\alpha$-Acetoxyepoxy azadirone [1.12] | C. paniculatus | Seeds | Chatterjee et al., 1989 |
| 6 $\alpha$-Acetoxygedunin [1.13] | C. paniculatus | Seeds | Connolly et al., 1979; Chatterjee et al., 1989 |
| 6 $\alpha$-Acetoxynimbinin [1.14] | C. paniculatus | Seeds | Connolly et al., 1979 |
| 6 $\alpha$-Acetoxy-16oxoazadirone [1.15] | C. paniculatus | Fruits | Saikia et al., 1978 |

Table 2 The distribution of triterpenoids in Chisocheton spp. (continued)

| Compounds | Source | Part | References |
| :---: | :---: | :---: | :---: |
| $5 \alpha, 7 \alpha, 13 \alpha, 17 \alpha-7-$ <br> Acetyloxy-21,23- $\gamma$ - <br> lactone-4,4,8-trimethyl- <br> 24-norchola-1,14,20,22- <br> tetraene-5-one [1.16] | C. microcarpus | Leaves | Gunning et al., 1994 |
| Compound E [1.17] | C. paniculatus | Seeds | Connolly et al., 1979 |
| Compound F [1.18] | C. paniculatus | Seeds | Connolly et al., 1979 |
| Compound G [1.19] | C. paniculatus | Seeds | Connolly et al., 1979 |
| $5 \alpha, 7 \alpha, 13 \alpha, 17 \alpha-7-$ <br> Deacetyloxy-21,23- $\gamma$ - <br> lactone-4,4,8-trimethyl- <br> 24-norchola-1,14,20,22- <br> tetraene-5-one [1.20] | C. microcarpus | Leaves | Gunning et al., 1994 |
| $\text { 1,2-Dihydro-6 } \alpha-$ <br> acetoxyazadirone [1.21] | C. paniculatus | Fruits | Bordoloi et al., 1993 |
| 6 $\alpha, 7 \alpha$-Dihydroxymeliaca-1,14,20,22-tetraene-3,14dione [1.22] | C. paniculatus | Fruits | Saikia et al., 1978 |
| Gedunin [1.23] | C. paniculaius | Seeds | Connolly et al., 1979 |
| 17 $\beta$-Hydroxy- $6 \alpha$ - | 1-C. paniculatus | Seeds | Connolly et al., 1979 |
| acetoxyazadiradione [1.24] | C. paniculatus | Seeds | Chatterjee et al., 1989 |
| $17 \beta \text {-Hydroxy- } 6 \alpha-$ <br> acetoxynimbinin [1.25] | C. paniculatus | Seeds | Connolly et al., 1979 |
| Vilasinin 1,3-diacetate [1.26] | C. paniculatus | Wood | Connolly et al., 1979 |
| Oleanane triterpenes <br> Moronic acid [1.27] | C. macrophyllus | Leaves | Inada et al., 1993 |
| Lupane triterpenes <br> Betulonic acid [1.28] | C. macrophyllus | Leaves | Inada et al., 1993 |

Table 2 The distribution of triterpenoids in Chisocheton spp. (continued)

| Compounds | Source | Part | References |
| :--- | :---: | :---: | :--- |
| Dammarane triterpenes <br> 24-Hydroxydammara- <br> 20,25-dien-3-one [1.29] | C. macrophyllus | Leaves | Inada et al., 1993 |



Compound $\mathrm{A}[1.3]: \mathrm{R}_{1}=0, \mathrm{R}_{2}=\mathrm{H}, \mathrm{R}_{3}=\mathrm{Ac}$
Compound D [1.6]: $\mathrm{R}_{1}=\mathrm{H}, \alpha-\mathrm{OH}, \mathrm{R}_{2}=\mathrm{H}, \mathrm{R}_{3}=\mathrm{Ac}$
Paniculatin B [1.7]: $\mathrm{R}_{1}=\mathrm{H}, \alpha-\mathrm{OH}, \mathrm{R}_{2}=\mathrm{Ac}, \mathrm{R}_{3}=\mathrm{H}$


Compound $\mathrm{B}[1.4]: \mathrm{R}_{1}=\mathrm{H}, \alpha-\mathrm{OAc}, \mathrm{R}_{2}=\mathrm{H}$
Paniculatin D [1.9] : $\mathrm{R}_{1}=\mathrm{H}, \alpha-\mathrm{OH}, \mathrm{R}_{2}=\mathrm{Ac}$


Compound C [1.5] : $\mathrm{R}_{1}=\mathrm{H}, \alpha-\mathrm{OAc}, \mathrm{R}_{2}=\mathrm{H}, \mathrm{R}_{3}=\mathrm{OH}$
Paniculatin C [1.8] : $\mathrm{R}_{1}=\mathrm{H}, \alpha-\mathrm{OH}, \mathrm{R}_{2}=\mathrm{Ac}, \mathrm{R}_{3}=\mathrm{OH}$
Figure 3. Chemical structures of triterpenoids found in Chisocheton spp.


Arunachalin [1.10]


6 $\alpha$-Acetoxyazadirone (Paniculatin) [1.11]

$$
\begin{aligned}
& : \mathrm{R}_{1}=\mathrm{OAc}, \mathrm{R}_{2}=\mathrm{OAc}, \mathrm{R}_{3}=\mathrm{H}_{2}, \mathrm{R}_{4}=\mathrm{H} \\
& : \mathrm{R}_{1}=\mathrm{OAc}, \mathrm{R}_{2}=\mathrm{OAc}, \mathrm{R}_{3}=\mathrm{O}, \mathrm{R}_{4}=\mathrm{H}
\end{aligned}
$$

$6 \alpha$-Acetoxy-16-oxoazadirone [1.15] $6 \alpha, 7 \alpha$-Dihydroxymeliaca-1,14,20,22-tetraene-3,14-dione [1.22]
$: \mathrm{R}_{1}=\mathrm{OH}, \mathrm{R}_{2}=\mathrm{OH}, \mathrm{R}_{3}=\mathrm{O}, \mathrm{R}_{4}=\mathrm{H}$
$17 \beta$-Hydroxy- $6 \alpha$-acetoxyazadiradione [1.24] : $\mathrm{R}_{1}=\mathrm{OAc}, \mathrm{R}_{2}=\mathrm{OAc}, \mathrm{R}_{3}=\mathrm{O}, \mathrm{R}_{4}=\mathrm{OH}$


6 $\alpha$-Acetoxyepoxyazadirone [1.12]


6 $\alpha$-Acetoxygedunin [1.13] : $\mathrm{R}_{1}=\mathrm{OAc}$
Gedunin [1.23]

$$
: \mathrm{R}_{1}=\mathrm{H}
$$

Figure 3. Chemical structures of triterpenoids found in Chisocheton spp. (continued)

6 $\alpha$-Acetoxynimbinin [1.14] : $\mathrm{R}=\mathrm{H}$
$17 \beta$-Hydroxy- $6 \alpha$-acetoxynimbinin [1.25] : $\mathrm{R}=\mathrm{OH}$

$5 \alpha, 7 \alpha, 13 \alpha, 17 \alpha-7$-Acetyloxy-21,23- $\gamma$-lactone-4,4,8-trimethyl-24-norchola-1,14,20,22-tetraene-5-one [1.16]

$$
: \mathrm{R}=\mathrm{Ac}
$$

5 $\alpha, 7 \alpha, 13 \alpha, 17 \alpha$-7-Deacetyloxy-21,23- $\gamma$-lactone-4,4,8-trimethyl-24-norchola-$1,14,20,22$-tetraene-5-one [1.20] $\quad: \mathrm{R}=\mathrm{H}$




Compound G [1.19] : $\mathrm{R}=$ вон $^{\text {он }}$
Figure 3. Chemical structures of triterpenoids found in Chisocheton spp. (continued)


1,2-Dihydro-6 $\alpha$-acetoxyazadirone [1.21]


Vilasinin 1,3-diacetate [1.26]



Betulonic acid [1.28]


24-Hydroxydammara-20,25-dien-3-one [1.29]

Figure 3. Chemical structures of triterpenoids found in Chisocheton spp. (continued)

## Dammarane - type triterpenoid compounds

Dammarane-type triterpenoid compounds are found in almost every plant family, including Meliaceae. They represent tetracyclic triterpenes as shown below.


The distribution of dammarane-type triterpenes in the Meliaceae is summarized in Table 3 and their chemical structures are shown in Figure 4.

Table 3. Distribution of dammarane-type triterpenoids in the Meliaceae.

| Compounds | Source | Part | References |
| :---: | :---: | :---: | :---: |
| $23(24 \rightarrow 25)$ Abeo- <br> 20R,24-dihydroxy <br> dammaran-3-one [2.1] | Dysoxylum cauliflorum | Fruits | Huang et al., 1999 |
| Aglaiol [2.2] | Aglaia odorata | Leaves | Shiengthong et al., 1965 |
|  | Cabralea polytricha | Fruits | Cascon and Brown, $1972$ |
| Aglaiondiol [2.3] | Aglaia odorata | Leaves | Shiengthong et al., $1974$ |
| Aglaitriol (24R) [2.4] | Aglaia odorata | Leaves | Shiengthong et al., 1974 |
| Aglaitriol (24S) [2.5] | Aglaia odorata | Leaves | Shiengthong et al., 1974 |

Table 3. Distribution of dammarane-type triterpenoids in the Meliaceae. (continued)

| Compounds | Source | Part | References |
| :---: | :---: | :---: | :---: |
| Aglinin A [2.6] | Aglaia lawii | Leaves | Mohamad et al., 1999 |
|  | Amoora yunnanensis | Bark | Luo et al., 2000 |
| Aglinin B [2.7] | Aglaia lawii | Leaves | Mohamad et al., 1999 |
| Aglinin C [2.8] | Aglaia tomentosa | Leaves | Mohamad et al., 1999 |
| Aglinin D [2.9] | Aglaia tomentosa | Leaves | Mohamad et al., 1999 |
| Cabraleadiol [2.10] | Aglaia crassinervia | Bark | Su et al., 2006 |
|  | Aglaia lawii | Stem bark | Qiu et al., 2001 |
|  | Aglaia tomentosa | Leaves | Mohamad et al., 1999 |
|  | Amoora cucullata | Stem bark | Haque et al., 1995 |
|  | Amoora yunnanensis | Leaves | Luo et al., 2000 |
|  | Cabralea eichleriana | Wood | Rao et al., 1975 |
|  | C. polytricha | Fruits | Cascon and Brown, $1972$ |
|  | Dysoxylum malabaricum | Stem bark | Hisham et al., 1996 |
| Cabraleadiol 3-acetate[2.11] | Aglaia lawii | Stem bark | Qiu et al., 2001 |
|  | Aglaia tomentosa | Leaves | Mohamad et al., 1999 |
|  | Cabralea eichteriana | Wood | Rao et al., 1975 |
|  | C. polytricha | Fruits | Cascon and Brown, $1972$ |
| $\begin{aligned} & \text { Cabraleahydroxylactone } \\ & {[2.12]} \end{aligned}$ | Aglaia crassinervia | Bark | Su et al., 2006 |
|  | Cabralea eichleriana | Wood | Rao et al., 1975 |
|  | C. polytricha | Fruits | Cascon and Brown, 1972 |
|  | Amoora yunnanensis | Bark | Luo et al., 2000 |
| Cabralealactone [2.13] | Aglaia leucophylla | Stem bark | Benosman et al., 1994 |
|  | Aglaia tomentosa | Leaves | Mohamad et al., 1999 |
|  | Cabralea eichleriana | Wood | Rao et al., 1975 |
|  | C. polytricha | Fruits | Cascon and Brown, $1972$ |

Table 3. Distribution of dammarane-type triterpenoids in the Meliaceae. (continued)

| Compounds | Source | Part | References |
| :---: | :---: | :---: | :---: |
| Cabralealactone [2.13] | Amoora cucullata | Stem bark | Haque et al., 1996 |
|  | Dysoxylum cauliflorum | Fruits | Huang et al., 1999 |
| Cabralealactone 3acetate [2.14] | Aglaia tomentosa | Leaves | Mohamad et al., 1999 |
| Cabraleone [2.15] | Aglaia lawii | Leaves | Mohamad et al., 1999 |
|  | Aglaia tomentosa | Bark | Mohamad et al., 1999 |
|  | Aglaia leucophylla | Stem bark | Benosman et al., 1994 |
|  | Aglaia silvestris | Fruits | Hwang et al., 2004 |
|  | Aglaia rubiginosa | Leaves | Rivero-Cruz et al., 2004 |
|  | Amoora yunnanensis | Bark | Luo et al., 2000 |
|  | Amoora cucullata | Stem bark | Haque et al., 1995 |
|  | Cabralea eichleriana | Wood | Rao et al., 1975 |
|  | C. polytricha | Fruits | Cascon and Brown, 1972 |
|  | Dysoxylum richii | Fruits | Aalbersberg and Singh, 1991 |
|  | D. muellerii | Wood | Mulholland and Naidoo, 2000 |
| 20S,23R,24R-23-Chloro- 20,24-epoxy- dammarane-3 $\alpha, 24,25$ - triol 3-acetate [2.16] | $\begin{array}{\|l\|} \hline \text { Amoora } \\ \text { yunnanensis } \end{array}$ | Bark | Luo et al., 2000 |
| $\begin{aligned} & \text { 5 } \alpha \text {-Dammar-20-ene-3 } \beta \text { - } \\ & \text { 24,25-triol [2.17] } \end{aligned}$ | Aglaia odorata | Leaves | Boar and Damps, 1977 |
| Dammarenolic acid [2.18] | Cabralea eichleriana | Wood | Rao et al., 1975 |
|  | Aglaia rubiginosa | Leaves | Rivero-Cruz et al., 2004 |

Table 3. Distribution of dammarane-type triterpenoids in the Meliaceae. (continued)

| Compounds | Source | Part | References |
| :---: | :---: | :---: | :---: |
| 11 $\alpha, 20 \xi-$ <br> Dihydroxydammar-24- <br> ene-3-one [2.19] | Astrotrichilia asterotricha | Wood and bark | Mulholland et al., 1994 |
| 24,25-Dihydroxy-dammar-20-en-3-one [2.20] | Aglaia odorata | Leaves | Boar and Damps, 1977 |
|  | Amoora yunnanensis | Bark | Luo et al., 2000 |
| $\begin{aligned} & \hline(20 S, 23 E)-20,25- \\ & \text { Dihydroxy-3,4- } \\ & \text { secodammara-4(28),23- } \\ & \text { dienoic acid }[\mathbf{2 . 2 1 ]} \end{aligned}$ | Aglaia rubiginosa | Leaves | Rivero-Cruz et al., 2004 |
| (20S,23E)-20,25- <br> Dihydroxy-3,4- <br> secodammara-4(28),23- <br> dienoic acid methyl ester [2.22] | Aglaia rubiginosa | Leaves | Rivero-Cruz et al., 2004 |
| Dymalol [2.23] | Dysoxylum <br> malaaricum | Leaves | Govindachari et al., $1994$ |
| Eichlerialactone [2.24] | Amoora yunnanensis | Bark | Luo et al., 2000 |
|  | Cabralea canjerana | Branches | De Campos Braga et al., 2006 |
|  | C. eichleriana | Wood | Rao et al., 1975 |
|  | Dysoxylum cauliflorum | Fruits | Huang et al., 1999 |
|  | D. richii | Leaves | Singh and <br> Aalbersberg, 1992 |
| Eichlerianic acid [2.25] | Aglaia lawii | Leaves | Mohamad et al., 1999 |
|  | Aglaia leucophylla | Stem bark | Benosman et al., 1994 |
|  | Aglaiu. elliptica | Stem | Cui et al., 1997 |
|  | Dysoxylum richii | Fruits | Aalbersberg and Singh, 1991 |

Table 3. Distribution of dammarane-type triterpenoids in the Meliaceae. (continued)

| Compounds | Source | Part | References |
| :---: | :---: | :---: | :---: |
| Eichlerianic acid [2.25] | Cabralea canjerana | Stem and branches | De Campos Braga et $\text { al., } 2006$ |
|  | C. eichleriana | Wood | Rao et al., 1975 |
| 3-Epi-Cabraleahydroxy <br> lactone [2.26] | Aglaia crassinervia | Bark | Su et al., 2006 |
| 3-Epi-ocotillol [2.27] | Aglaia lawii | Leaves | Mohamad et al., 1999 |
|  | Aglaia foveolata | Bark | Roux et al., 1998 |
|  | Aglaia crassinervia | Bark | Su et al., 2006 |
| $24 \xi, 25-\text { Ероху-5 } \alpha-$ <br> dammar-20-en-3-one $[2.28]$ | Aglaia odorata | Leaves | Boar and Damps, 1977 |
| 20S,24-Epoxy-24,25- <br> dihydroxydammar-3-one [2.29] | Amoora yunnanensis | Bark | Luo et al., 2000 |
| $\begin{aligned} & \text { 20S,24S-Epoxy-7 } \beta, 25 \text { - } \\ & \text { dihydroxy-3,4- } \\ & \text { secodammar-4(28)-en-3- } \\ & \text { oic acid }[2.30] \end{aligned}$ | Cabralea <br> canjerana | Stem | $\begin{aligned} & \text { De Campos Braga et } \\ & \text { al., } 2006 \end{aligned}$ |
| 20S,24S-Epoxy-4-hydroxy-3,4-seco-dammar-25(26)-en-3-oic acid [2.31] | Dysoxylum richii | Leaves | Singh and Aalbersberg, 1992 |
| 20S,24S-Epoxy-25- <br> hydroxydammaran-3-one [2.32] | Aglaia <br> elaeagnoidea | Bark | Fuzzati et al., 1996 |
| 20S,24S-Epoxy-25-hydroxymethyldammaran- 3-one [2.33] | Aglaia <br> elaeagnoidea | Bark | Fuzzati et al., 1996 |

Table 3. Distribution of dammarane-type triterpenoids in the Meliaceae. (continued)

| Compounds | Source | Part | References |
| :---: | :---: | :---: | :---: |
| 20S,24S-Epoxy- <br> $7 \beta, 15 \alpha, 25$-trihydroxy- <br> 3,4-secodammar-4(28)- <br> en-3-oic acid [2.34] | Cabralea canjerana | Stem | De Campos Braga et al., 2006 |
| 20S,24S-Epoxy- <br> $7 \beta, 22 \xi, 25$-trihydroxy- <br> 3,4-secodammar-4(28)- <br> en-3-oic acid [2.35] | C. canjerana | Stem | De Campos Braga et $\text { al., } 2006$ |
| 20S,24-Ероху-25,26,27-trisnor-24-oxo-3,4-seco-4(28)-dammaren-3-oic acid [2.36] | Amoora <br> yunnanensis | Bark | Luo et al., 2000 |
|  | Dysoxylum richii | Leaves | Singh and Aalbersberg, 1992 |
| Ethyl eichlerianoate $[2.37]$ | D. cauliflorum | Stem bark | Benosman et al., 2000 |
| Foveolin A [2.38] | Aglaia lawii | Leaves | Mohamad et al., 1999 |
|  | Aglaia foveolata | , Bark | Roux et al., 1998 |
| Foveolin B [2.39] | Aglaia foveolata | Bark | Roux et al., 1998 |
| Methyl richenoate [2.40] | Dysoxylum richii | Fruits | Aalbersberg et al., 1991 |
| Ocotillol [2.41] | Aglaia <br> leucophylla | Stem bark | Benosman et al., 1994 |
|  | Aglaia elliptica | Stem | Cui et al., 1997 |
|  | Cabralea polytricha | Fruits | Cascon and Brown, 1972 |
|  | Dysoxylum cauliflorum | Fruits | Huang et al., 1999 |
| Ocotillone [2.42] | Aglaia <br> leucophylla | Stem bark | Benosman et al., 1994 |
|  | Aglaia silvestris | Fruits | Hwang et al., 2004 |
|  | Aglaia rubiginosa | Leaves | Rivero-Cruz et al., 2004 |

Table 3. Distribution of dammarane-type triterpenoids in the Meliaceae. (continued)

| Compounds | Source | Part | References |
| :---: | :---: | :---: | :---: |
| Ocotillone [2.42] | Amoora yunnanensis | Bark | Luo et al., 2000 |
|  | Cabralea canjerana | Stem and branches | Braga et al., 2006 |
|  | C. eichleriana | Wood | Rao et al., 1975 |
|  | Dysoxylum malaaricum | Leaves | Govindachari et al., $1994$ |
|  | D. richii | Fruits | Aalbersberg and Singh, 1991 |
|  | D. cauliflorum | Fruits | Huang et al., 1999 |
| Richenoic acid [2.43] | Dysoxylum richii | Fruits | Aalbersberg and Singh, 1991 |
| Richenol [2.44] | Dysoxylum richii | Fruits | Aalbersberg and Singh, 1991 |
| Richenone [2.45] | D. richii | Fruits | Aalbersberg and Singh, 1991 |
|  | D. muellerii | Wood | Mulholland and Naidoo, 2000 |
| Shoreic acid [2.46] | Aglaia lawii | Leaves | Mohamad et al., 1999 |
|  | Aglaia elliptica | h Stem | Cui et al., 1997 |
|  | Aglaia rubiginosa | Leaves | $\begin{aligned} & \text { Rivero-Cruz et al., } \\ & 2004 \end{aligned}$ |
|  | Amoora yunnanensis | Bark | Luo et al., 2000 |
|  | Cabralea canjerana | Stem and branches | De Campos Braga et al., 2006 |
|  | C. eichleriana | Wood | Rao et al., 1975 |
|  | Dysoxylum malabaricum | Leaves | Govindachari et al., $1994$ |
|  | D. richii | Fruits | Aalbersberg and Singh, 1991 |

Table 3. Distribution of dammarane-type triterpenoids in the Meliaceae. (continued)

| Compounds | Source | Part | References |
| :--- | :--- | :---: | :--- |
| Shoreic acid [2.46] | D. cauliflorum | Fruits | Huang et al., 1999 |
| $3 \alpha, 11 \alpha, 20 \xi$-Trihydroxy <br> dammar-24-ene [2.47] | Aglaia <br> asterotricha | Wood and | Mulholland et al., 1994 |



$$
\begin{array}{llll}
\mathrm{R}_{1} & \mathrm{R}_{2} & \mathrm{R}_{3} & \mathrm{R}_{4}
\end{array}
$$

| Aglinin C [2.8] | $\beta-\mathrm{H}$ | $\alpha-\mathrm{OH}$ | OH | H |
| :--- | :---: | :---: | :---: | :---: |
| Aglinin D [2.9] |  | $=0$ | OH | H |
| Cabraleadiol [2.10] |  |  | H | $\beta-\mathrm{H}$ |
| Cabraleadiol 3-acetate [2.11] | $\beta-\mathrm{H}$ | $\alpha-\mathrm{OAc}$ | $\beta-\mathrm{H}$ | H |
| Cabraleone [2.15] |  | $=0$ | $\beta-\mathrm{H}$ | H |
| 20S,23R,24R-23-Chloro- | $\beta-\mathrm{H}$ | $\alpha-\mathrm{OAc}$ | $\alpha-\mathrm{OH}$ | $\beta-\mathrm{Cl}$ |

## 20,24-epoxy-dammarane-

$3 \alpha, 24,25$-triol 3-acetate [2.16]
3-Epi-ocotillol [2.27] $\quad \beta$-H $\quad \alpha-\mathrm{OH} \quad \beta$-H $\quad \mathrm{H}$
20S,24-Epoxy-24,25- $\quad=\mathrm{O} \quad \mathrm{OH} \quad \mathrm{H}$
dihydroxydammar-3-one [2.29]
20S,24S-Epoxy-25-hydroxy- $\quad=0 \quad \alpha$-H H
dammaran-3-one [2.32]
20S,24S-Epoxy-25-hydroxy- $\quad=\mathrm{O} \quad \alpha-\mathrm{H} \quad \mathrm{H}$
methyldammaran-3-one [2.33]
$\begin{array}{lcccc}\text { Ocotillol [2.41] } & \alpha-\mathrm{H} & \beta-\mathrm{OH} & \beta-\mathrm{H} & \mathrm{H} \\ \text { Ocotillone [2.42] } & =\mathrm{O} & \beta-\mathrm{H} & \mathrm{H}\end{array}$
$20 S, 23 R, 24 S$-23-Chloro-20,24 $\beta$-H $\quad \alpha$-OAc $\quad \beta$-OH $\quad \beta$-Cl
-epoxy-dammarane-3 $\alpha, 24,25-$
triol 3-acetate [2.36]

Figure 4. Chemical structures of dammarane - type triterpenoids in the Meliaceae

$\begin{array}{ll}\text { Cabraleahydroxylactone [2.12] } & : \mathrm{R}_{1}, \mathrm{R}_{2}=\mathrm{H}, \alpha-\mathrm{OH} \\ \text { Cabralealactone [2.13] } & : \mathrm{R}_{1}, \mathrm{R}_{2}==\mathrm{O} \\ \text { Cabralealactone 3-acetate [2.14] } & : \mathrm{R}_{1}, \mathrm{R}_{2}=\mathrm{H}, \alpha-\mathrm{OAc} \\ \text { 3-Epi-cabraleahydroxylactone [2.26] } & : \mathrm{R}_{1}, \mathrm{R}_{2}=\mathrm{H}, \beta-\mathrm{OH}\end{array}$


Eichlerialactone [2.24]

Figure 4. Chemical structures of dammarane - type triterpenoids in the Meliaceae (continued)

$\begin{array}{lllll}\mathrm{R}_{1} & \mathrm{R}_{2} & \mathrm{R}_{3} & \mathrm{R}_{4} & \mathrm{R}_{5}\end{array}$

| Aglinin A [2.6] | H |  |  | H | H |
| :---: | :---: | :---: | :---: | :---: | :---: |
| Eichlerianic acid, $24 S$ [2.25] |  | H | H | H | H |
| $20 S, 24 S$-Epoxy-7 $\beta, 25$-dihydroxy |  | $\alpha$ - |  | $\beta$-OH | H |
| 3,4-secodammar-4(28)-en- |  |  |  |  |  |
| acid [2.30] |  |  |  |  |  |
| $20 S, 24 S$-Epoxy-7 $\beta, 15 \alpha, 25-$ |  |  | $\alpha$-O | $\beta-\mathrm{OH}$ | H |
| trihydroxy-3,4-secodammar- |  |  |  |  |  |
| 4(28)-en-3-oic acid [2.34] |  |  |  |  |  |
| 20S,24S-Epoxy-7 $\beta, 22 \xi, 25-$ |  |  | H | $\beta-\mathrm{OH}$ | OH |
| trihydroxy-3,4-secodammar- |  |  |  |  |  |
| 4(28)-en-3-oic acid [2.35] |  |  |  |  |  |
| Ethyl eichlerianoate, 24S [2.37] | $\mathrm{CH}_{2}$ |  | H | H | H |
| Shoreic acid, 24R [2.46] | H | H | H | H | H |

Figure 4. Chemical structures of dammarane - type triterpenoids in the Meliaceae (continued)

Aglinin B [2.7]
$: \mathrm{R}_{1}=\mathrm{H}, \mathrm{R}_{2}=\mathrm{OH}$
Dymalol, $24 S$ [2.23]
Foveolin A, 24S [2.38] $: \mathrm{R}_{1}=\mathrm{H}, \mathrm{R}_{2}=\mathrm{H}$
Foveolin B, $24 R[2.39]$
$: \mathrm{R}_{1}=\mathrm{H}, \mathrm{R}_{2}=\mathrm{H}$



Richenol [2.44]
Richenone [2.45]
$: \mathrm{R}_{1}, \mathrm{R}_{2}=\mathrm{H}, \beta-\mathrm{OH}$
$: \mathrm{R}_{1}, \mathrm{R}_{2}==0$

Figure 4. Chemical structures of dammarane - type triterpenoids in the Meliaceae (continued)


Aglaiol [2.2]


$5 \alpha$-Dammar-20-ene-3 $\beta$-24,25-triol [2.17]
24,25-Dihydroxy-dammar-20-en-3-one [2.20]
$: \mathrm{R}_{1}, \mathrm{R}_{2}=\mathrm{H}, \beta-\mathrm{OH}$
$: \mathrm{R}_{1}, \mathrm{R}_{2}==0$

Figure 4. Chemical structures of dammarane - type triterpenoids in the Meliaceae (continued)

$24 \xi, 25$-Epoxy- $5 \alpha$-dammar-20-en-3-one [2.28]



20S,24S-Epoxy-4-hydroxy-3,4-seco-dammar-25(26)-en-3-oic acid [2.31]

Figure 4. Chemical structures of dammarane - type triterpenoids in the Meliaceae (continued)

$23(24 \rightarrow 25)$ Abeo-20R,24-dihydroxydammaran-3-one [2.1]

$11 \alpha, 20 \xi$-Dihydroxydammar-24-ene-3-one [2.19] $: \mathrm{R}==0$
$3 \alpha, 11 \alpha, 20 \xi$-Trihydroxydammar-24-ene [2.47] $: \mathrm{R}=\alpha-\mathrm{OH}, \beta-\mathrm{H}$

(20S,23E)-20,25-Dihydroxy-3,4-secodammara-4(28),23-dienoic acid [2.21]

$$
: \mathrm{R}=\mathrm{H}
$$

(20S,23E)-20,25-Dihydroxy-3,4-secodammara-4(28),23-dienoic acid methyl ester [2.22]

$$
: \mathrm{R}=\mathrm{CH}_{3}
$$

Figure 4. Chemical structures of dammarane - type triterpenoids in the Meliaceae (continued)

## Aromadendrane - type sesquiterpenes

Aromadendranes are a class of sesquiterpene natural products found in a number of plant species (Gijsen et al., 1992), especially in species belonging to Compositae, Leguminosae, and Myrtaceae. They are also found in marine organisms including soft corals, sponges, and liverwort (Staerk et al., 2004). Aromadendranes are tricyclic sesquiterpenes structurally characterized by a dimethyl cyclopropane unit fused to a hydroazulene ring system (Stephen et al., 1996).

Distribution of aromadendrane - type sesquiterpenes in plants is summarized in Table 4 and their chemical structures are shown in Figure 5.
Table 4. Distribution of aromadendrane - type sesquiterpenes in plants.

| Compound | Sources | Family | Part | References |
| :---: | :---: | :---: | :---: | :---: |
| 2'-Acetyl arvoside B [3.1] | Calendula arvensis | Compositae | Aerial parts | $\begin{aligned} & \text { Pizza et al., } \\ & 1988 \end{aligned}$ |
| Alloaromadendran-14 $\beta$ - <br> al [3.2] | Duguetia glabriuscula | Annonaceae | Leaves | De Siqueira et al., 2003 |
| Alloaromadendran-14 $\beta$ oic acid [3.3] | Duguetia glabriuscula | Annonaceae | Leaves | De siqueira et al., 2003 |
| (+)-Aromadendra-1(10),4-dien-15-al-3-one $[3.4]$ | Mandevilla pentlandiana | Apocynaceae | Root | Cabrera et al.,1993 |
| $1(5), 3-$ <br> Aromadendradiene [3.5] | Balsamum tolutanum | Leguminosae | n.i. | Friedel et al., 1987 |
| $1(10), 4-$ <br> Aromadendradiene [3.6] | Balsamum tolutanum | Leguminosae | n.i. | Friedel et al., 1987 |
| (+)-Alloaromadendrane$4 \beta, 10 \alpha$-diol [3.7] | Ambrosia peruviana | Compositae | Stem, <br> leaves | Goldsby and Burke, 1987 |
|  | Phebalium filifolium | Rutaceae | Aerial parts | $\begin{aligned} & \text { Rashid et al., } \\ & 1995 \end{aligned}$ |
| (+)-Alloaromadendran$10,14 \beta$-diol [3.8] <br> (14-hydroxyviridiflorol) | Duguetia <br> glabriuscula | Annonaceae | Leaves | $\begin{aligned} & \text { Matos et al., } \\ & 2006 \end{aligned}$ |
|  | Pulicaria paludosa | Compositae | Aerial parts | Feliciano et al., 1989 |

Table 4. Distribution of aromadendrane - type sesquiterpenes in plants. (continued)

| Compound | Sources | Family | Part | References |
| :---: | :---: | :---: | :---: | :---: |
| Alloaromadendrene [3.9] | Duguetia glabriuscula | Annonaceae | Leaves | De Siqueira et al., 2003 |
| Alloaromadendrane- $4 \beta, 10 \alpha \text {-diol [3.10] }$ | Xylopia <br> brasiliensis | Annonaceae | Leaves | Moreira et al., 2003 |
| Aromadendrane-4 $\alpha, 10 \alpha$ diol [3.11] | X. brasiliensis | Annonaceae | Leaves | Moreira et $\text { al., } 2003$ |
| Aromadendrane-4 $\alpha, 10 \beta$ - <br> diol [3.12] | X. brasiliensis | Annonaceae | Leaves | $\begin{aligned} & \text { Moreira } \text { et } \\ & \text { al., } 2003 \end{aligned}$ |
| Aromadendrane-4 $\beta, 10 \alpha$ - <br> 14-triol [3.13] | X. brasiliensis | Annonaceae | Leaves | $\begin{aligned} & \text { Moreira et } \\ & \text { al., } 2003 \end{aligned}$ |
| Aromadendrane- $4 \beta, 10 \alpha$ diol [3.14] | Brasilia sickii | Compositae | Root | Bohlmann et al., 1983 |
|  | Aglaia grandis | Meliaceae | Leaves | $\begin{aligned} & \text { Inada } \text { et al., } \\ & 2000 \end{aligned}$ |
| Aromadendrane- $4 \beta, 10 \beta$ diol [3.15] | Ambrosia peruviana | Compositae | Stem, leaves | Goldsby and Burke, 1987 |
|  | Aristolochia heterophylla | Aristolc:hiaceae | Root, Stem | Wu, Chan and Leu, 2000 |
| (+)-Aromadendrene [3.16] | Eucalyptus globulus | Myrtaceae | Leaves | $\begin{aligned} & \text { Graham et } \\ & \text { al., } 1960 \end{aligned}$ |
| Arvoside B [3.17] | Calendula arvensis | Compositae | Aerial <br> parts | $\begin{aligned} & \text { Pizza et al., } \\ & 1988 \end{aligned}$ |
| Cyclocolorenone [3.18] | Drimys brasiliensis | Winteraceae | Stem <br> bark, <br> Leaves, <br> Fruits | $\begin{aligned} & \text { Limberger } e t \\ & \text { al., } 2007 \end{aligned}$ |
| Dendroside A [3.19] | Dendrobium nobile | Orchidaceae | Stem | $\begin{aligned} & \text { Zhao et al ., } \\ & 2001 \end{aligned}$ |
| Dendroside D [3.20] | D. nobile | Orchidaceae | Stem | $\begin{aligned} & \text { Ye } \text { et al ., } \\ & 2002 \end{aligned}$ |

Table 4. Distribution of aromadendrane - type sesquiterpenes in plants. (continued)

| Compound | Sources | Family | Part | References |
| :---: | :---: | :---: | :---: | :---: |
| 12,13-Diacetoxy-2-oxo-aromadendr-1(10)-ene [3.21] | Gnephosis arachnoidea | Compositae | Aerial parts | Jakupovic et al., 1988 |
| $\begin{aligned} & \hline(2 S, 4 R, 5 S, 6 R, 7 R, 9 S)- \\ & \text { 2,9-Dihydroxy-1(10)- } \\ & \text { aromadendren-14-oic } \\ & \text { acid 2,14-lactone [3.22] } \end{aligned}$ | Landolphia dulcis | Apocynaceae | Root | $\begin{aligned} & \text { Staerk } \text { et al., } \\ & 2004 \end{aligned}$ |
| (2S,4R,5S,6R,7R,11S)-2,12-Dihydroxy-1(10)-aromadendren-14-oic acid 2,14-lactone [3.23] | L. dulcis | Apocynaceae | Root | $\begin{aligned} & \text { Staerk } \text { et al., } \\ & 2004 \end{aligned}$ |
| $\begin{aligned} & \hline 8 \alpha, 13 \text {-Dihydroxy } \\ & \text { spathulenol [3.24] } \end{aligned}$ | Cineraria fruticulorum | Compositae | Aerial parts | Bohlmann et al., 1982 |
| Epiglobulol [3.25] | Eucalyptus globulus | Myrtaceae | Leaves | $\begin{aligned} & \text { Graham et } \\ & \text { al., } 1960 \end{aligned}$ |
| Flourensadiol [3.26] | Flourensia cernиa | Compositae | Whole plant | $\begin{aligned} & \hline \text { Kingston } \text { et } \\ & \text { al., } 1975 \end{aligned}$ |
| (+)-Globulol [3.27] | Angelica sylvestris | Apiaceae | Root | Vinokurova et al., 1999 |
| (-)-Globulol [3.28] | Eucalyptus <br> globulus | Myrtaceae | Leaves | Graham et $\text { al., } 1960$ |
| (-)- $\alpha$-Gurjunene [3.29] | Lansium anamalayanum | Meliaceae | Wood | Krishnappa and Dev, 1973 |
| (1R,6R,7R,10R,11S)-12- <br> Hydroxy-4(5)-aroma- <br> dendren-3-one $[3.30]$ | Landolphia dulcis | Apocynaceae | Root | $\begin{aligned} & \text { Staerk } \text { et al., } \\ & 2004 \end{aligned}$ |

Table 4. Distribution of aromadendrane - type sesquiterpenes in plants. (continued)

| Compound | Sources | Family | Part | References |
| :---: | :---: | :---: | :---: | :---: |
| (4R,5S,6R,7R,11S)-12-Hydroxy-1(10)-aromadendren-14-al [3.31] | L. dulcis | Apocynaceae | Root | Staerk et al., $2004$ |
| (4R,5S,6R,7R,11S)-12-Hydroxy-1(10)-aromadendren-2-one [3.32] | L. dulcis | Apocynaceae | Root | $\begin{aligned} & \text { Staerk } \text { et al., } \\ & 2004 \end{aligned}$ |
| (4R,5S,6R,7R,11S)- <br> 12-Hydroxy-1(10)- <br> aromadendren-9-one <br> [3.33] | L. dulcis | Apocynaceae | Root | $\begin{aligned} & \text { Staerk } \text { et al., } \\ & 2004 \end{aligned}$ |
| $(2 S, 4 R, 5 S, 6 R, 7 R)-2-$ <br> Hydroxy-1(10)-aromadendren-14oic acid 2,14-lactone [3.34] | L. dulcis | Apocynaceae | Root | $\begin{aligned} & \text { Staerk } \text { et al., } \\ & 2004 \end{aligned}$ |
| $8 \alpha$-Hydroxy-13-oxospathulenol [3.35] | Cineraria <br> fruticulorum | Compositae | Aerial parts | $\begin{aligned} & \text { Bohlmann et } \\ & \text { al., } 1982 \end{aligned}$ |
| $\begin{array}{\|l\|} \hline 8 \alpha \text {-Hydroxy } \\ \text { spathulenol [3.36] } \end{array}$ | C. fruticulorum | Compositae | Aerial parts | $\begin{aligned} & \text { Bohlmann et } \\ & \text { al., } 1982 \end{aligned}$ |
| $\begin{aligned} & (+)-13 \text {-Hydroxy } \\ & \text { spathulenol [3.37] } \end{aligned}$ | Eriostemon brucei | Rutaceae | Aerial parts | $\begin{aligned} & \hline \text { Rashid } e t \\ & \text { al., } 1995 \end{aligned}$ |
| (+)-Ledol [3.38] | Entandrophragm a cylindricum | Meliaceae | Bark | Daniewski et al., 1996 |
| (-)-Ledol [3.39] | Duguetia glabriuscula | Annonaceae | Leaves | De Siqueira et al., 2003 |

Table 4. Distribution of aromadendrane - type sesquiterpenes in plants. (continued)

| Compound | Sources | Family | Part | References |
| :---: | :---: | :---: | :---: | :---: |
| (-)-Ledol [3.39] | Phebalium tuberculosum | Rutaceae | Aerial parts | Rashid et $\text { al., } 1995$ |
|  | Piper clusii | Piperaceae | Fruits | Koul et al., $1993$ |
| $2^{\prime}\left(2^{\prime \prime}-\text { Methyl }-\right.$ <br> butanoyl) arvoside B [3.40] | Calendula arvensis | Compositae | Aerial parts | $\begin{aligned} & \text { Pizza et al., } \\ & 1988 \end{aligned}$ |
| $2^{\prime}\left(2^{\prime \prime}-\right.\text { Methyl-2"- }$ butenoyl) arvoside B [3.41] | C. arvensis | Compositae | Aerial parts | De Tom masi et al., 1990 |
| $\begin{array}{\|l\|} \hline \text { Methyl } \\ (4 R, 5 S, 6 R, 7 R, 11 S)- \\ 2,9 \text {-dioxo-1(10)- } \\ \text { aromadendren-12- } \\ \text { oate [3.42] } \end{array}$ | Landolphia dulcis | Apocynaceae | Root | $\begin{aligned} & \hline \text { Staerk } \text { et al., } \\ & 2004 \end{aligned}$ |
| $2^{\prime}\left(3^{\prime \prime}-\text { Methyl-2" }-\right.$ pentenoyl) arvoside B [3.43] | Calendula arvensis | Compositae | Aerial parts | $\begin{aligned} & \text { Pizza et al., } \\ & 1988 \end{aligned}$ |
| $\begin{aligned} & 2^{\prime}\left(2^{\prime \prime}\right. \text {-Methyl-2"'- } \\ & \text { propanoyl }) \text { arvoside } \\ & \text { B [3.44] } \end{aligned}$ | C. arvensis | Compositae | Aerial <br> parts | De Tom masi et al., 1990 |
| (4R,5S,6R,7R,11S)- 2-Oxo-1(10)- aromadendren-12- oic acid [3.45] | Landolphia dulcis | Apocynaceae | Root | Staerk et al., $2004$ |
| (+)-Spathulenol [3.46] | Duguetia glabriuscula | Annonaceae | Leaves | De Siqueira et al., 2003 |
|  | Guarea <br> guidonia | Meliaceae | Leaves | Brochini and Roque, 2000 |

Table 4. Distribution of aromadendrane - type sesquiterpenes in plants. (continued)

| Compound | Sources | Family | Part | References |
| :---: | :---: | :---: | :---: | :---: |
| (+)-Spathulenol[3.46] | Drummondita basselli | Rutaceae | Aerial parts | Rashid et $\text { al., } 1995$ |
|  | D. calida | Rutaceae | Aerial parts | Rashid et $\text { al., } 1995$ |
|  | Phebalium tuberculosum | Rutaceae | Aerial parts | Rashid et $\text { al., } 1995$ |
|  | P. filifolium | Rutaceae | Aerial parts | $\begin{aligned} & \text { Rashid et } \\ & \text { al., } 1995 \end{aligned}$ |
|  | Xylopia brasiliensis | Annonaceae | Leaves | Moreira et $\text { al., } 2003$ |
| $\beta$-Spathulene [3.47] | Schinus molle | Anacardiaceae | Fruits | Terhune, Hogg and Lawrence, 1974 |
| Squamulosone[3.48] | Hyptis <br> verticillata | Labiatae | n.i. | Collins et al., 2001 |
|  | Phebalium squamulosum | Rutaceae <br> เหาวิทยาลัย | n.i. | Batey, <br> Hellyer and <br> Pinhey, $1971$ |
| (-)-10 $\beta, 13,14-$ <br> Trihydroxy-alloaromadendrane [3.49] | Wyethia arizonica | Compositae | n.i. | Bohlmann et al., 1984 |
| Viridiflorol [3.50] | Duguetia glabriuscula | Annonaceae | Leaves | De Siqueira et al., 2003 |

n.i. $=$ not indicated

(+)-Aromadendra-1(10),4-dien-15-al-3-one [3.4]: $\mathrm{R}_{1}=\mathrm{CHO}, \mathrm{R}_{2}, \mathrm{R}_{3}=\mathrm{O}$

1(10),4-Aromadendradiene [3.6]


1(5),3-Aromadendradiene [3.5]
$: \mathrm{R}_{1}=\mathrm{CHO}, \mathrm{R}_{2}, \mathrm{R}_{3}=\mathrm{O}$

$\beta$-Spathulene [3.47]

$\begin{array}{llll}\mathrm{R}_{1} & \mathrm{R}_{2} & \mathrm{R}_{3} & \mathrm{R}_{4}\end{array}$

Alloaromadendran-14 $\beta$-al [3.2]
Alloaromadendran-14 $\beta$-oic acid [3.3]
(+)-Alloaromadendran-10,14 $\beta$-diol
(14-hydroxyviridiflorol) [3.8]
Flourensadiol [3.26]
H $\quad \beta-\mathrm{OH} \quad \alpha-\mathrm{CH}_{3} \quad \mathrm{OH}$
(-)-Ledol [3.39]
Viridiflorol [3.50]
H $\quad \beta$ - $\mathrm{CHO} \quad \alpha-\mathrm{CH}_{3} \quad \mathrm{H}$

H $\beta-\mathrm{COOH} \quad \alpha-\mathrm{H} \quad \mathrm{H}$
$\mathrm{H} \alpha-\mathrm{CH}_{2} \mathrm{OH} \quad \beta-\mathrm{OH} \quad \mathrm{H}$

Figure 5. Chemical structures of aromadendrane - type sesquiterpenes in plants

$\begin{array}{lllll}\mathrm{R}_{1} & \mathrm{R}_{2} & \mathrm{R}_{3} & \mathrm{R}_{4} & \mathrm{R}_{5}\end{array}$

Alloaromadendrane-4 $\beta, 10 \alpha$-diol
$\mathrm{OH} \quad \alpha-\mathrm{OH} \quad \beta-\mathrm{CH}_{3} \quad \mathrm{CH}_{3} \quad \mathrm{H}$
[3.10]
Dendroside A[3.19]
Dendroside $\mathrm{D}[3.20]$
(-)-10 , 13, 14-Trihydroxy-allo-
H $\alpha-\mathrm{CH}_{2} \mathrm{Oglu} \quad \beta-\mathrm{OH} \quad \mathrm{CH}_{2} \mathrm{OH} \quad \mathrm{H}$
H $\alpha-\mathrm{CH}_{2} \mathrm{Oglu} \beta-\mathrm{OH}$ COOglu H
H $\beta-\mathrm{OH} \quad \alpha-\mathrm{CH}_{2} \mathrm{OH} \quad \mathrm{CH}_{3} \quad \mathrm{OH}$ aromadendrane [3.49]

$\begin{array}{lcccc} & \mathrm{R}_{1} & \mathrm{R}_{2} & \mathrm{R}_{3} & \mathrm{R}_{4} \\ \text { (+)-Alloaromadendrane- } 4 \beta, 10 \alpha \text {-diol [3.7] } & \mathrm{OH} & \alpha-\mathrm{OH} & \beta-\mathrm{CH}_{3} & \mathrm{H} \\ \text { (+)-Globulol [3.27] } & \mathrm{H} & \alpha-\mathrm{OH} & \beta-\mathrm{CH}_{3} & \mathrm{H} \\ \text { (+)-Ledol [3.38] } & \mathrm{H} & \beta-\mathrm{OH} & \alpha-\mathrm{CH}_{3} & \mathrm{H}\end{array}$

Figure 5. Chemical structures of aromadendrane - type sesquiterpenes in plants (continued)


$$
\mathrm{R}_{1} \quad \mathrm{R}_{2}
$$

| Aromadendrane-4 $\alpha, 10 \alpha$-diol [3.11] | $\alpha-\mathrm{OH}$ | $\beta-\mathrm{CH}_{3}$ |
| :--- | :--- | :--- |
| Aromadendrane-4 $\alpha, 10 \beta$-diol [3.12] | $\beta$-OH | $\beta-\mathrm{CH}_{3}$ |



Aromadendrane-4 $\beta, 10 \alpha$-14-triol [3.13] $\mathrm{OH} \quad \beta-\mathrm{CH}_{2} \mathrm{OH} \alpha-\mathrm{OH} \mathrm{CH}_{3} \quad \mathrm{H}$ Aromadendrane- $4 \beta, 10 \alpha$-diol [3.14] GKOR OH $\alpha-\mathrm{OH} \quad \beta-\mathrm{CH}_{3} \quad \mathrm{CH}_{3} \quad \mathrm{H}$
Aromadendrane- $4 \beta, 10 \beta$-diol [3.15] $\quad \mathrm{OH} \quad \beta$ - $\mathrm{OH} \quad \alpha-\mathrm{CH}_{3} \quad \mathrm{CH}_{3} \quad \mathrm{H}$
Epiglobulol [3.25] $\quad \mathrm{H} \quad \beta-\mathrm{OH} \quad \alpha-\mathrm{CH}_{3} \quad \mathrm{CH}_{3} \quad \mathrm{H}$
(-)-Globulol [3.28]
H $\quad \beta-\mathrm{OH} \quad \alpha-\mathrm{CH}_{3} \quad \mathrm{CH}_{3} \quad \mathrm{H}$

Figure 5. Chemical structures of aromadendrane - type sesquiterpenes in plants (continued)

$\begin{array}{lll}\mathrm{R}_{1} & \mathrm{R}_{2} & \mathrm{R}_{3}\end{array}$

Cyclocolorenone [3.18]
$(-)-\alpha$-Gurjunene [3.29]
$\mathrm{H} \quad \mathrm{H}$
( $1 R, 6 R, 7 R, 10 R, 11 S$ )-12-Hydroxy-
$=0 \quad \mathrm{OH}$
4(5)-aromadendren-3-one [3.30]

$\mathrm{R}_{1} \mathrm{R}_{2}$ 等
( $4 R, 5 S, 6 R, 7 R, 11 S$ )-12-Hydroxy-
$=0$
1(10)-aromadendren-2-one [3.32]
( $4 R, 5 S, 6 R, 7 R, 11 S$ )-12-Hydroxy-
H H
1(10)-aromadendren-9-one [3.33]
Squamulosone [3.48]
(4R,5S,6R,7R,11S)-2-Oxo-1(10)-

H
$=0$
$=0$
$\mathrm{CH}_{2} \mathrm{OH}$
H H
$\mathrm{CH}_{2} \mathrm{OH}$ $\mathrm{CH}_{3}$

COOH aromadendren-12-oic acid [3.45]

Figure 5. Chemical structures of aromadendrane - type sesquiterpenes in plants (continued)

$\begin{array}{lllllll}\mathrm{R}_{1} & \mathrm{R}_{2} & \mathrm{R}_{3} & \mathrm{R}_{4} & \mathrm{R}_{5} & \mathrm{R}_{6} & \mathrm{R}_{7}\end{array}$

12,13-Diacetoxy-2-oxo- $\quad=\mathrm{O} \quad \mathrm{H} \quad \mathrm{H} \mathrm{CH} \mathbf{C H A C C H}_{2} \mathrm{OAc}_{\mathrm{OH}}^{3}$ aromadendr-1(10)-ene [3.21]
$(4 R, 5 S, 6 R, 7 R, 11 S)$-12-Hydroxy- $\mathrm{H} \quad \mathrm{H} \quad \mathrm{H} \quad \mathrm{H} \quad \mathrm{CH}_{3} \quad \mathrm{CH}_{2} \mathrm{OH} \mathrm{CHO}$ 1(10)-aromadendren-14-al [3.31]

Methyl (4R,5S,6R,7R,11S)-2,9- $=0 \quad=0 \quad C H_{3} \quad \mathrm{COOCH}_{3} \mathrm{CH}_{3}$ dioxo-1(10)-aromadendren-12oate [3.42]

( $2 S, 4 R, 5 S, 6 R, 7 R$ )-2-Hydroxy-1(10)-aromadendren-14-oic acid 2,14-lactone [3.34]

$$
: \mathrm{R}_{1}=\mathrm{H}, \mathrm{R}_{2}=\mathrm{CH}_{3}
$$

( $2 S, 4 R, 5 S, 6 R, 7 R, 9 S$ )-2,9-Dihydroxy-1(10)-aromadendren-14-oic acid 2,14-lactone [3.22]: $\mathrm{R}_{1}=\mathrm{OH}, \mathrm{R}_{2}=\mathrm{CH}_{3}$
( $2 S, 4 R, 5 S, 6 R, 7 R, 11 S$ )-2,12-Dihydroxy-1(10)-aromadendren-14-oic acid 2,14-lactone [3.23]: $\mathrm{R}_{1}=\mathrm{H}, \mathrm{R}_{2}=\mathrm{CH}_{2}$

Figure 5. Chemical structures of aromadendrane - type sesquiterpenes in plants (continued)


Alloaromadendrene [3.9]

(+)-Spathulenol [3.46]

$$
: \mathrm{R}_{1}=\mathrm{OH}, \mathrm{R}_{2}=\mathrm{H}, \mathrm{R}_{3}=\mathrm{CH}_{3}
$$

(+)-Aromadendrene [3.16]

$$
: \mathrm{R}_{1}=\mathrm{H}, \mathrm{R}_{2}=\mathrm{H}, \mathrm{R}_{3}=\mathrm{CH}_{3}
$$

(+)-13-Hydroxyspathulenol [3.37]
$: \mathrm{R}_{1}=\mathrm{OH}, \mathrm{R}_{2}=\mathrm{H}, \mathrm{R}_{3}=\mathrm{CH}_{2} \mathrm{OH}$
$8 \alpha$-Hydroxyspathulenol [3.36]
$: \mathrm{R}_{1}=\mathrm{OH}, \mathrm{R}_{2}=\mathrm{OH}, \mathrm{R}_{3}=\mathrm{CH}_{3}$
$8 \alpha, 13$-Dihydroxyspathulenol [3.24]
$8 \alpha$-Hydroxy-13-oxo-spathulenol [3.35]

$$
: \mathrm{R}_{1}=\mathrm{OH}, \mathrm{R}_{2}=\mathrm{OH}, \mathrm{R}_{3}=\mathrm{CH}_{2} \mathrm{OH}
$$

$$
: \mathrm{R}_{1}=\mathrm{OH}, \mathrm{R}_{2}=\mathrm{OH}, \mathrm{R}_{3}=\mathrm{CHO}
$$

Figure 5. Chemical structures of aromadendrane - type sesquiterpenes in plants (continued)


2'-Acetyl arvoside B [3.1]
Arvoside B [3.17]
$2^{\prime}\left(2^{\prime \prime}\right.$-Methyl-butanoyl) arvoside B [3.40]

2'(2"-Methyl-2'-butenoyl) arvoside B [3.41]
$2^{\prime}\left(3^{\prime \prime}-\right.$ Methyl-2" -pentenoyl) arvoside B [3.43]
$2^{\prime}\left(2^{\prime \prime}\right.$-Methyl-2"'-propanoyl) aryoside B [3.44]

$$
\begin{aligned}
& : R=A c \\
& : R=H
\end{aligned}
$$






Figure 5. Chemical structures of aromadendrane - type sesquiterpenes in plants (continued)

Aromadendrane-type sesquiterpenes displayed various biological activities. For example dendroside A from the stems of Dendrobium nobile (family Orchidaceae), a plant used as tonic in traditional Chinese medicine, significantly stimulated the proliferation of murine T and B lymphocytes in vitro at concentrations of $1 \times 10^{-7}$ and $1 \times 10^{-5} \mathrm{M}$, respectively (Zhao et al., 2001). Another compound, (+)-alloaromadendran-10,14 $\beta$-diol (14-hydroxyviridiflorol) from Duguetia glabriuscula (family Annonaceae), showed cytotoxic activity by inhibiting the growth of $\mathrm{Hep}_{2}$ (human larynx carcinoma) cell line ( $\mathrm{IC}_{50}<25 \mu \mathrm{~g} / \mathrm{ml}$ ) (Matos et al., 2006).

Cladosporium herbarium and C. cladosporioides are important causes of mould allergies in human. It was found that ( + )-alloaromadendrane- $4 \beta, 10 \alpha$-diol, an aromadendrane sesquiterpene from the stems and leaves of Ambrosia peruviana (family Compositae), together with aromadendrane-4 $\alpha, 10 \beta$-diol and aromadendrane$4 \alpha, 10 \alpha$-diol from the leaves of Xylopia brasiliensis (family Annonaceae), exhibited antifungal activity against these allergic molds. (Goldsby and Buke, 1987; Moreira et al., 2003). Arvoside B, an aromadendrane glycoside from the aerial parts of Calendula arvensis (family Compositae), showed in vitro antiviral activity against vesicular stomatitis virus at MIC of $14 \mu \mathrm{~g} / \mathrm{ml}$ (De Tommasi et al., 1990). Squamulosone, isolated in large quantity from Hyptis verticillata (family Labiatae), a medicinal plant which has been traditionally used in the treatment of eczema, psoriasis, scabies, athlete's foot, rheumatoid arthritis and cold-related problems, displayed insecticidal activity against the sweet potato weevil, Cylas formicarius elegantulus (Collins et al., 2001).


## Flavagline derivatives

Flavagline derivatives are unusual benzofuran derivatives featuring a cyclopenta $[b]$ benzofuran skeleton.


They were initially named rocaglamide derivatives after the parent compound rocaglamide, originally isolated in 1982 (King et al., 1982). These compounds appear to be derived from a flavonoid nucleus linked with a cinnamic acid moiety (Nugroho et al., 1999). The incorporation of nitrogen into rocaglamide represents a late biosynthetic step, therefore, the general use of the term rocaglamides for all derivatives of that basic structure can not be applied to every compound within this group. Regarding the restricted occurrence of this type of compounds to the genus Aglaia of the family Meliaceae and the incorporation of a flavonoid moiety as a central biosynthetic step, the name flavaglines was therefore suggested for this class of compounds (Brader et al., 1998).

Flavaglines can be classified into three groups (Proksch et al., 2001) :

1) Cyclopenta[b]benzofuran derivatives (rocaglamide derivatives)

2) Cyclopenta[bc] benzopyran derivatives (aglain and aglaforbesin derivatives)

aglain derivatives

aglaforbesin derivatives
3) Benzo $[b]$ oxe-pine derivatives (forbagline derivatives)


Many members of the cyclpenta[b]benzofuran groups exhibited biological activities such as insecticidal activity (Ishibashi et al., 1993; Janprasert et al., 1993; Nugroho et al., 1997a, 1997b; Brader et al., 1998; Bacher et al, 1999; Chaidir et al., 1999; Hiort et al., 1999; Nugroho et al., 1999; Schneider et al., 2000; Dreyer et al., 2001; Greger et al., 2001; Bringmann et al., 2003) comparable in potency to azadirachtin the well-known natural insecticide from the neem tree, Azadirachta indica L. These flavaglines also displayed significant inhibitory activity against cancer cell lines at nanomolar concentrations (King et al., 1982; Dumontet et al., 1996; Cui et al., 1997; Lee et al., 1998; Mohamad et al., 1999; Proksch et al., 2001; Baumann et al., 2002; Hausott et al., 2004; Hwang, et al., 2004; Rivero-Cruz, et al., 2004; Chumkaew et al., 2006; Kim et al., 2006; Su et al., 2006; Salim, et al., 2007). However, the cyclopenta[bc]benzopyrans and benzo[b]oxepines evaluated so far were not active (Kim et al., 2006).

The distribution of flavagline derivatives in the family Meliaceae is summarized in Table 5 and their chemical structures are shown in Figure 6.

Table 5. Distribution of flavagline compounds in the family Meliaceae.

| Compounds | Source | part | References |
| :---: | :---: | :---: | :---: |
| Cyclopenta[b]benzofurans 1-O-Acetyl-N-butanoyldidesmethylrocaglamide [4.1] | Aglaia elliptica | Fruits | Nugroho et al., 1997b |
| 1-O-Acetyldemethylrocaglamide [4.2] | A. duperreana | Flowers | Chaidir et al., 1999 |
|  |  | Root | Hiort et al., 1999 |
| 1-O-Acetyl-4'-demethoxy-3',4'-methylenedioxy-methyl rocaglate [4.3] | A. spectabilis | Bark | $\begin{aligned} & \text { Schneider } \text { et al., } \\ & 2000 \end{aligned}$ |
| 1-O-Acetyl-didemethylrocaglamide [4.4] | A. duperreana | Flowers | Chaidir et al., 1999 |
| 1-O-Acetyl-3'hydroxydemethylrocaglamide [4.5] | A. duperreana | Flowers | Chaidir et al., 1999 |
|  |  | Root | Hiort et al., 1999 |
| 1-O-Acetyl-3'-hydroxy methylrocaglate [4.6] | A. duperreana | Flowers | Chaidir et al., 1999 |
|  |  | Root | Hiort et al., 1999 |
| 1-O-Acetyl-3'-hydroxyrocaglamide [4.7] | A. duperreana | Flowers | Chaidir et al., 1999 |
|  |  | Twigs | $\begin{aligned} & \text { Nugroho } \text { et al., } \\ & \text { 1997a } \end{aligned}$ |
|  | งกรณ์มหาวิ | Root | Hiort et al., 1999 |
|  | A. odorata | Twigs | Nugroho et al., 1999 |
|  |  | Leaves | Ishibashi et al., 1993 |
| 1-O-Acetyl-rocaglamide [4.8] | A. duperreana | Root | Hiort et al., 1999 |
| 1-O-Acetylmethyl rocaglate[4.9] | A. duperreana | Root | Hiort et al., 1999 |
|  |  | Flowers | Chaidir et al., 1999 |
|  | A. rubiginosa | Twigs | Rivero-Cruz et al., $2004$ |
| 1-O-Acetylrocaglaol [4.10] | A .rubiginosa | Twigs | Rivero-Cruz et al., 2004 |
| $\begin{aligned} & \text { Aglaiastatin [4.11] } \\ & \text { (aglaroxin D) } \end{aligned}$ | A. duperreana | Twigs | $\begin{aligned} & \text { Nugroho et al., } \\ & \text { 1997a } \end{aligned}$ |

Table 5. Distribution of flavagline compounds in the family Meliaceae. (continued)

| Compounds | Source | part | References |
| :---: | :---: | :---: | :---: |
| Aglaiastatin [4.11] <br> (aglaroxin D) | A. odorata | Leaves | Ohse et al., 1996 |
| N-Butanoyldidesmethylrocaglamide [4.12] | A. elliptica | Fruits | Nugroho et al., 1997b |
| ```Cyclorocaglamide (6- demethoxy-8b,2'-epoxy-3'- methoxy-6,7- methylenedioxy rocaglamide) [4.13]``` | A. oligophylla | Twigs | Bringmann et al., 2003 |
| Dehydroaglaiastatin [4.14] | A. duperreana | Twigs | Nugroho et al., 1997a |
|  |  | Flowers | Chaidir et al., 1999 |
|  |  | Root | Hiort et al., 1999 |
|  | A. odorata | Root | Kokpol et al., 1994 |
|  |  | Leaves | Ohse et al., 1996 |
|  |  | Root | Kokpol et al., 1994 |
|  | A. testicularis | Leaves | Wang et al., 2004 |
| 6-Demethoxy-2'-hydroxy- <br> 3'-methoxy-6,7- <br> methylenedioxy <br> rocaglamide [4.15] | A. oligophylla | Twigs <br> ทยาลัย | Bringmann et al., 2003 |
| 4'-Demethoxy-3', 4'-methylenedioxy-methyl rocaglate [4.16] | A. dasyclada | Leaves | Chaidir et al., 2001 |
|  | A. elliptica | Stem, Fruits | Cui et al., 1997 |
|  |  | Stems | Lee et al., 1998 |
|  | A. spectabilis | Bark | Schneider et al., 2000 |
| 6-Demethoxy-6,7-methylenedioxymethylrocaglate [4.17] (Pannellin) | A.elaeagnoidea | Leaves, <br> Stems, <br> Root , Bark | Brader et al., 1998 |

Table 5. Distribution of flavagline compounds in the family Meliaceae. (continued)

| Compounds | Source | part | References |
| :---: | :---: | :---: | :---: |
| $\begin{aligned} & \hline \text { 6-Demethoxy-6,7- } \\ & \text { methylenedioxymethyl- } \\ & \text { rocaglate [4.17] (Pannellin) } \end{aligned}$ | A. oligophylla | Twigs | Dreyer et al., 2001 |
| $\begin{aligned} & \text { 6-Demethoxy-6,7- } \\ & \text { methylenedioxy rocaglamide } \\ & (\text { aglaroxin A) [4.18] } \end{aligned}$ | A.elaeagnoidea | Stem bark | $\begin{aligned} & \text { Molleyres et al., } \\ & 1999 \end{aligned}$ |
|  | A. oligophylla | Twigs | Dreyer et al., 2001 |
| 4'-Demethoxy-3',4'methylenedioxyrocaglaol[4.19] | A. elliptica | Fruits | Cui et al., 1997 |
|  | A. spectabilis | Bark | Schneider et al., 2000 |
| 1-Oxo-4'-demethoxy-3',4'methylenedioxy rocaglaol [4.20] | A. elliptica | Stem | Cui et al., 1997 |
| Desmethylrocaglamide [4.21] | A, duperreana | Flowers | Chaidir et al., 1999 |
|  | A. odorata | Leaves | Ishibashi et al., 1993 |
| Didesmethylrocaglamide [4.22] | A. argentea | Seeds | $\begin{aligned} & \text { Dumontet et al., } \\ & 1996 \end{aligned}$ |
|  | A. duperreana | Root | Hiort et al., 1999 |
|  | A. elliptica | Fruits | $\begin{aligned} & \text { Nugroho et al., } \\ & \text { 1997b } \end{aligned}$ |
| Episilvestrol [4.23] | A. pyramidata | Twigs | Hwang et al., 2004 |
| 8b-O-Ethyl- <br> demethylrocaglamide [4.24] | A. duperreana | Flowers | Chaidir et al., 1999 |
| 8b-O-Ethyl-3'- <br> hydroxyrocaglamide [4.25] | A. duperreana | Flowers | Chaidir et al., 1999 |
| Ethylrocaglaol [4.26] | A. forbesii | Bark | Dumontet et al., 1996 |
| 1-O-Formylmethyl rocaglate [4.27] | A. dasyclada | Leaves | Chaidir et al., 2001 |
|  | A. spectabilis | Bark | Schneider et al., $2000$ |

Table 5. Distribution of flavagline compounds in the family Meliaceae. (continued)

| Compounds | Source | part | References |
| :---: | :---: | :---: | :---: |
| $\begin{aligned} & \text { 1-O-Formyloxy-4'- } \\ & \text { demethoxy-3',4'- } \\ & \text { methylenedioxy-methyl } \\ & \text { rocaglate }[4.28] \end{aligned}$ | A. elliptica | Stem | $\begin{aligned} & \hline \text { Cui } \text { et al., } 1997 \\ & \text { Lee } \text { et al., } 1998 \end{aligned}$ |
|  | A. spectabilis | Bark | Schneider et al., $2000$ |
| 1-O-Formylrocagloic acid [4.29] | Amoora cucullata | Fruits | Chumkaew et al., $2006$ |
| 3'-Hydroxy- <br> dehydroaglaiastatin [4.30] | A. duperreana | Flowers | Chaidir et al., 1999 |
|  | A. testicularis | Leaves | Wang et al., 2004 |
| 3'-Hydroxydemethylrocaglamide [4.31] | A. odorata | Leaves | Nugroho et al., 1999 |
|  | A. duperreana | Root | Ohse et al., 1996 |
| 3'-Hydroxy-1-O-formyloxymethyl rocaglate [4.32] | A. spectabilis | Bark | $\begin{aligned} & \text { Schneider } \text { et al., } \\ & 2000 \end{aligned}$ |
| 3'-Hydroxydidemethylrocaglamide [4.33] | A. odorata | Leaves | Nugroho et al., 1999 |
| 3'-Hydroxymarikarin [4.34] | A. gracilis | Root and Stem bark | Greger et al., 2001 |
| 3'-Hydroxymethyl rocaglate [4.35] | A. duperreana | Root | Ohse et al., 1996 |
|  | A. odorata | Leaves | Nugroho et al., 1999 |
|  | A. spectabilis | Bark | $\begin{aligned} & \text { Schneider } \text { et al., } \\ & 2000 \end{aligned}$ |
|  | Amoora cucullata | Fruits | Chumkaew et al., $2006$ |
| 3'-Hydroxyrocaglamide[4.36] | A. duperreana | Flowers | Chaidir et al., 1999 |
|  |  | Twigs | $\begin{aligned} & \text { Nugroho et al., } \\ & \text { 1997a } \end{aligned}$ |
|  |  | Root | Hiort et al., 1999 |
|  | A. odorata | Twigs and Leaves | Nugroho et al., 1999 |
| 3'-Hydroxyrocagloic acid [4.37] | Amoora cucullata | Fruits | Chumkaew et al., $2006$ |

Table 5. Distribution of flavagline compounds in the family Meliaceae. (continued)

| Compounds | Source | part | References |
| :---: | :---: | :---: | :---: |
| Marikarin [4.38] | A. gracilis | Root, Stem bark | Greger et al., 2001 |
| 3'-Methoxy-6-demethoxy- <br> 6,7-methylenedioxy <br> rocaglamide [4.39] | A.elaeagnoidea | Stem bark | Molleyres et al., 1999 |
|  | A. oligophylla | Twigs | Dreyer et al., 2001 |
| 3'- Methoxy-methyl rocaglate [4.40] | A. spectabilis | Bark | Schneider et al., 2000 |
| 3'-Methoxypannellin [4.41] | A. elaeagnoidea | Leaves, <br> Stems, <br> Root, Bark | Brader et al., 1998 |
| 3'- Methoxylrocaglaol [4.42] | A. odorata | Twigs | Nugroho et al., 1999 |
|  |  | Leaves | Nugroho et al., 1999 |
| 3'- Methoxyrocaglamide[4.43] | A. duperreana | Twigs | $\begin{aligned} & \text { Nugroho et al., } \\ & \text { 1997a } \end{aligned}$ |
|  | A. odorata | Twigs | Nugroho et al., 1999 |
| 8b-O-methyl-methyl rocaglate [4.44] | A. duperreana | Root | Hiort et al., 1999 |
| Methyl rocaglate [4.45] (Aglafoline) | A. dasyclada | Leaves | Chaidir et al., 2001 |
|  | A. duperreana | Flowers | Chaidir et al., 1999 |
|  | งกรณัมหาวิท | Root | Hiort et al., 1999 |
|  | A.elaeagnoidea | Bark | Fuzzati et al., 1996 |
|  | A. elliptica | Stem and Fruits | Cui et al., 1997 |
|  | A. odorata | Leaves | Ishibashi et al., 1993 |
|  | A. rubiginosa | Twigs | Rivero-Cruz et al., 2004 |
|  | A. spectabilis | Bark | Schneider et al., 2000 |
|  | Amoora cucullata | Fruits | Chumkaew et al., $2006$ |

Table 5. Distribution of flavagline compounds in the family Meliaceae. (continued)

| Compounds | Source | part | References |
| :---: | :---: | :---: | :---: |
| Methyl rocaglate [4.45] (Aglafoline) | A. elliptifolia | n.i. | Ko et al., 1992 |
|  | A. ponapensis | Leaves and twigs | Salim et al., 2007 |
| 8b-O-methylrocaglaol [4.46] | A. duperreana | Root | Hiort et al., 1999 |
| 1-Oxime-3'-methoxymethylrocaglate [4.47] | A. odorata | Leaves | Nugroho et al., 1999 |
| 1-Oxo-2-piriferine-6-demethoxy-6,7methylenedioxy rocaglamide [4.48] | A. oligophylla | Twigs | Dreyer et al., 2001 |
|  | A. spectabilis | Bark | $\begin{aligned} & \text { Schneider } \text { et al., } \\ & 2000 \end{aligned}$ |
| Pannellin 1-O-acetate [4.49] | A. elaeagnoidea | Leaves, Stems and Root, Bark | Brader et al., 1998 |
| Rocaglamide [4.50] | A. duperreana | Twigs | $\begin{aligned} & \text { Nugroho et al., } \\ & \text { 1997a } \end{aligned}$ |
|  |  | Root | Hiort et al., 1999 |
|  | A. elliptica | Fruits | Nugroho et al., $1997 \mathrm{~b}$ |
|  | A. elliptifolia | Root and Stem | King et al., 1982 |
|  | A. odorata UTII | Twigs | Janprasert et al., 1993 |
|  |  | Leaves | Ishibashi et al., 1993 |
| Rocaglaol-3'-rhamnose [4.51] | A. harmsiana | Leaves | $\begin{aligned} & \text { Nugroho et al., } \\ & \text { 1997b } \end{aligned}$ |
| Rocaglaol [4.52] | A. duperreana | Root | Hiort et al., 1999 |
|  | A. elliptifolia | Stem | Cui et al., 1997 |
|  | A. odorata | Leaves | Ishibashi et al., 1993 |
|  |  |  | Ohse et al., 1996 |

Table 5. Distribution of flavagline compounds in the family Meliaceae. (continued)

| Compounds | Source | part | References |
| :---: | :---: | :---: | :---: |
| Rocaglaol [4.52] | A. crassinervia | Bark | Su et al., 2006 |
|  | A. dasyclada | Leaves | Chaidir et al., 2001 |
|  | A. ferruginaea | n.i. | Mulholland and Naidoo, 1998 |
|  | A. forbesii | Bark | Dumontet et al., 1996 |
|  | A. spectabilis | Bark | Schneider et al., 2000 |
|  | A. tomentosa | Bark | Mohamad et al., 1999 |
|  | Amoora cucullata | Fruits | Chumkaew et al., $2006$ |
| Rocagloic acid [4.53] | A. dasyclada | Leaves | Chaidir et al., 2001 |
|  | A. rubiginosa | Twigs | $\begin{aligned} & \text { Rivero-Cruz et al., } \\ & 2004 \end{aligned}$ |
|  | Amoora cucullata | Fruits | Chumkaew et al., $2006$ |
| Silvestrol [4.54] | A. pyramidata | Fruits | Hwang et al., 2004 |
| N -Tetrahydrofuranrocaglamide [4.55] | A. elliptica | Fruits | Nugroho et al., 1997b |
| $\begin{aligned} & \text { Cyclopenta[bc]benzopyrans } \\ & \text { (aglains) } \\ & \text { Aglain A [4.56] } \end{aligned}$ | A. argentea | Leaves | Dumontet et al., 1996 |
|  | A. forbesii | Barks | Dumontet et al., 1996 |
| Aglain B [4.57] | A. argentea | Leaves | Dumontet et al., 1996 |
| Aglain C [4.58] | A. argentea | Leaves | Dumontet et al., 1996 |

Table 5. Distribution of flavagline compounds in the family Meliaceae. (continued)


Table 5. Distribution of flavagline compounds in the family Meliaceae. (continued)

| Compounds | Source | part | References |
| :---: | :---: | :---: | :---: |
| Thapsakon B [4.79] | A. edulis | Roots | Bacher et al., 1999 |
| Cyclopenta[bc]benzopyrans <br> (aglaforbesins) <br> Aglaforbesin A [4.80] | A. forbesii | Bark | Dumontet et al., 1996 |
| Aglaforbesin B [4.81] | A. forbesii | Bark | $\begin{aligned} & \text { Dumontet et al., } \\ & 1996 \end{aligned}$ |
| Aglaxiflorin C [4.82] | A. laxiflora | Leaves | Xu et al., 2000 |
| Aglaforbesin -O-glycoside [4.83] | A. dasyclada | Leaves | Chaidir et al., 2001 |
| 8-Demethoxy-7,8methylenedioxyaglaforbesin [4.84] | A. oligophylla | Twigs | Dreyer et al., 2001 |
| Benzo[b]oxepines <br> (forbaglines) <br> Forbaglin A [4.85] | A. forbesii | Bark | Dumontet et al., $1996$ |
| Forbaglin B [4.86] | A. forbesii | Bark | Dumontet et al., 1996 |
| Forbaglin- $O$-glycoside [4.87] | A. dasyclada ทย | Leaves | Chaidir et al., 2001 |
| Homothapoxepine A [4.88] | A.edulis UNIVE | Roots | Bacher et al., 1999 |
| 4'-Hydroxy-10-acidic-21deglycosyloxy forbaglin [4.89] | A. dasyclada | Leaves | Chaidir et al., 2001 |
| (13R)-Thapoxepine A [4.90] | A. edulis | Roots | Bacher et al., 1999 |
| (13S)-Thapoxepine A [4.91] | A. edulis | Roots | Bacher et al., 1999 |
| (13R)-Thapoxepine B [4.92] | A. edulis | Roots | Bacher et al., 1999 |
| (13S)-Thapoxepine B [4.93] | A.edulis | Roots | Bacher et al., 1999 |


$\begin{array}{lll}R_{1} & R_{2} & R_{3}\end{array}$

1-O-Acetyldemethyl-
$\mathrm{OCOCH}_{3} \mathrm{CONHCH}_{3} \mathrm{H}$ rocaglamide [4.2]
1-O-Acetyl-didemethylrocaglamide $\quad \mathrm{OCOCH}_{3} \mathrm{CONH}_{2} \quad \mathrm{H}$ [4.4]
1-O-Acetyl-3'-hydroxydemethylrocaglamide [4.5]

1-O-Acetyl-3'-hydroxymethyl rocaglate [4.6]

1-O-Acetyl-3'-hydroxy-rocaglamide $\quad \mathrm{OCOCH}_{3} \mathrm{CON}\left(\mathrm{CH}_{3}\right)_{2} \mathrm{OH}$ [4.7]
1-O-Acetyl-rocaglamide [4.8] $\quad \mathrm{OCOCH}_{3} \quad \mathrm{CON}\left(\mathrm{CH}_{3}\right)_{2} \mathrm{H}$
1-O-Acetylmethyl rocaglate [4.9] $\quad \mathrm{OCOCH}_{3} \quad \mathrm{COOH} \quad \mathrm{H}$
1-O-Acetylrocaglaol [4.10] $\quad \mathrm{OCOCH}_{3} \quad \mathrm{H} \quad \mathrm{H}$
Desmethylrocaglamide [4.21] $\quad \mathrm{OH} \quad \mathrm{CONHCH}_{3} \quad \mathrm{H}$
1-O-Formylmethyl rocaglate [4.27] $\quad 0 \mathrm{OCHO} \quad \mathrm{COOCH}_{3} \quad \mathrm{H}$
1-O-Formylrocagloic acid [4.29]
3'-Hydroxy-1-O-formyloxy-
OCHO COOH H
methyl rocaglate [4.32]
3'-Hydroxymethyl rocaglate [4.35]
3'-Hydroxyrocaglamide [4.36]
3'-Hydroxyrocagloic acid [4.37]
3'- Methoxy-methyl rocaglate [4.40]
3'- Methoxylrocaglaol [4.42]
3'- Methoxyrocaglamide [4.43]
$\mathrm{OH} \quad \mathrm{COOCH}_{3} \quad \mathrm{OH}$
$\mathrm{OH} \quad \mathrm{CON}\left(\mathrm{CH}_{3}\right)_{2} \mathrm{OH}$
$\mathrm{OH} \quad \mathrm{COOH} \quad \mathrm{OH}$
$\mathrm{OH} \quad \mathrm{COOCH}_{3} \quad \mathrm{OCH}_{3}$
$\mathrm{OH} \quad \mathrm{H} \quad \mathrm{OCH}_{3}$
$\mathrm{OH} \quad \mathrm{CON}\left(\mathrm{CH}_{3}\right)_{2} \quad \mathrm{OCH}_{3}$

Figurc 6. Chemical structures of flavagline compounds in the family Meliaceae

R

Methyl rocaglate (Aglafoline) [4.45]
Rocaglamide [4.50]
Rocaglaol [4.52]
Rocagloic acid [4.53]
$\mathrm{COOCH}_{3}$
$\mathrm{CON}\left(\mathrm{CH}_{3}\right)_{2}$
H
COOH

$\mathrm{R}_{2}$
$\mathrm{R}_{3}$
1-O-Acetyi-N-butanoyl-didesmethyl- $\quad \mathrm{OCOCH}_{3} \mathrm{CONH}_{\left(\mathrm{CH}_{2}\right)_{4} \mathrm{OH} \quad \mathrm{H}}$ rocaglamide [4.1]
N -butanoyl-didesmethylrocaglamide [4.12]
$\mathrm{OH} \mathrm{CONH}\left(\mathrm{CH}_{2}\right)_{4} \mathrm{OH} \quad \mathrm{H}$
Didesmethylrocaglamide [4.22]
3'-Hydroxydemethyl- rocaglamide $\mathrm{OH} \quad \mathrm{CONHCH}_{3} \quad \mathrm{OH}$ [4.31]
3'-Hydroxydidemethyl-rocaglamide
$\mathrm{OH} \quad \mathrm{CONH}_{2}$ OH [4.33]

Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)


1-Oxime-3'-methoxy- methylrocaglate [4.47]


N -Tetrahydrofuran-rocaglamide [4.55]


1-O-Acetyl-4'-demethoxy-3',4'-methylenedioxy-
$\begin{array}{cc}\mathrm{R}_{1} & \mathrm{R}_{2} \\ \mathrm{OCOCH}_{3} & \mathrm{COOCH}_{3}\end{array}$ methyl rocaglate [4.3]
4'-Demethoxy-3', 4'-methylenedioxy-methyl rocaglate [4.16]
OH
OH
$\mathrm{COOCH}_{3}$
4'-Demethoxy-3',4'-methylenedioxyrocaglaol [4.19]
1-O-Formyloxy-4'-demethoxy-3',4'-methylenedioxyOCHO H methyl rocaglate [4.28]

Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)


1-Oxo-4'-demethoxy-3',4'-methylenedioxy rocaglaol [4.20]


Aglaiastatin (aglaroxin D) [4.11]



Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)


R
6-Demethoxy-6,7-methylenedioxy rocaglamide $\quad \mathrm{H}$
(aglaroxin A) [4.18]
3'-Methoxyl-6-demethoxy-6,7-methylenedioxy $\mathrm{OCH}_{3}$ rocaglamide [4.39]


6-Demethoxyl-6,7-methylenedioxymethylrocaglate $\quad \mathrm{OH} \quad \mathrm{H}$
(Pannellin) [4.17]
3'-Methoxypannellin [4.41] $\mathrm{OH} \quad \mathrm{OCH}_{3}$
Pannellin 1-O-acetate [4.49]
$\mathrm{OCOCH}_{3} \quad \mathrm{H}$


1-Oxo-2-piriferine-6-demethoxy-6,7-methylenedioxy rocaglamide [4.48]
Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)


|  | $\mathrm{R}_{1}$ | $\mathrm{R}_{2}$ |
| :--- | :---: | :---: |
| 8b-O-Ethyl-demethylrocaglamide [4.24] | $\mathrm{CONHCH}_{3}$ | $\mathrm{OC}_{2} \mathrm{H}_{5}$ |
| 8b-O-Ethyl-3'-hydroxyrocaglamide [4.25] | $\mathrm{CON}\left(\mathrm{CH}_{3}\right)_{2}$ | $\mathrm{OC}_{2} \mathrm{H}_{5}$ |
| Ethylrocaglaol [4.26] | H | H |



6-Demethoxy-2'-hydroxy-3'-methoxy-6,7-methylenedioxy rocaglamide [4.15]


Cyclorocaglamide
(6-demethoxy-8b,2'-epoxy-3'-methoxy-6,7-methylenedioxyrocaglamide) [4.13]

Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)


Rocaglaol-3'-rhamnose [4.51]

$\begin{array}{cl}8 \mathrm{~b}-\mathrm{O} \text {-methyl-methyl rocaglate [4.44] } & : \mathrm{R}=\mathrm{COOCH}_{3} \\ \text { 8b-O-methylrocaglaol [4.46] } & : \mathrm{R}=\mathrm{H}\end{array}$


$$
\begin{gathered}
\text { Episilvestrol }[4.23]\left(5^{\prime \prime \prime} S\right) \\
\text { Silvestrol }[4.54]\left(5^{\prime \prime} R\right)
\end{gathered}
$$

Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)


Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)



Elliptifoline [4.68]

Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)


3'-Hydroxyaglain C [4.73]


| Chulalongikorin Universit $\mathrm{R}_{1}$ | $\mathrm{R}_{2}$ |  |
| :--- | :---: | :---: |
| Isothapsakin B [4.74] (H-3 $\beta, \mathrm{H}-4 \alpha)$ | OH | H |
| Thapsakin A 10-O-acetate $[4.75](\mathrm{H}-3 \alpha, \mathrm{H}-4 \beta)$ | H | OAc |
| (13S)-Thapsakin B [4.77] (H-3 $\beta, \mathrm{H}-4 \alpha)$ | H | OH |

Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)

(13R)-Thapsakin B [4.76]


$$
\begin{array}{ll}
\text { CHULALONGIKRNN UNIV } & \mathrm{R}_{1} \\
\text { Thapsakon A [4.78] }(\mathrm{H}-3 \alpha, \mathrm{H}-4 \beta) & =\mathrm{O} \\
\text { Thapsakon B [4.79] (H-3 } \beta, \mathrm{H}-4 \alpha) & =\mathrm{O}
\end{array}
$$

Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)


Grandiamide A [4.69]


Aglain - $O$-glycoside [4.59]

Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)



8-Demethoxy-7,8-methylenedioxyaglaforbesin [4.84]

Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)


Aglaforbesin - $O$-glycoside [4.83]


Forbaglin A [4.85] (13R) (19S)
Forbaglin B [4.86] (13S)
(13R)-Thapoxepine B [4.92] (13R)
(13S)-Thapoxepine B [4.93]

| $\mathrm{R}_{1}$ | $\mathrm{R}_{2}$ | $\mathrm{R}_{3}$ |
| :---: | :---: | :---: |
| $\mathrm{OCH}_{3}$ | H | $\mathrm{C}_{2} \mathrm{H}_{5}$ |
| $\mathrm{OCH}_{3}$ | H | $\mathrm{C}_{2} \mathrm{H}_{5}$ |
| $-\mathrm{OCH}_{2} \mathrm{O}-$ | $\mathrm{CH}_{3}$ |  |
| $-\mathrm{OCH}_{2} \mathrm{O}-$ | $\mathrm{CH}_{3}$ |  |

Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)


Homothapoxepine A [4.88] (13S)
(13R)-Thapoxepine A [4.90] (13R)
(13S)-Thapoxepine A [4.91] (13S)
$\mathrm{R}_{1} \quad \mathrm{R}_{2}$
$\mathrm{R}_{3}$ $\begin{array}{lr}-\mathrm{OCH}_{2} \mathrm{O}- & \mathrm{C}_{2} \mathrm{H}_{5} \\ -\mathrm{OCH}_{2} \mathrm{O}- & \mathrm{CH}_{3} \\ -\mathrm{OCH}_{2} \mathrm{O}- & \mathrm{CH}_{3}\end{array}$

$\begin{array}{lll}\mathrm{R}_{1} & \mathrm{R}_{2} & \mathrm{R}_{3}\end{array}$

Forbaglin- $O$-glycoside [4.87]
Glucose
$\mathrm{CH}_{3}$
$\mathrm{CH}_{3}$
4'-Hydroxy-10-acidic-21-deglycosyloxy H
H
H

## forbaglin [4.89]

Figure 6. Chemical structures of flavagline compounds in the family Meliaceae (continued)

