

Research Progress on the Chemical Components and Pharmacological Activities of Gesneriaceae

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(Received May 24, 2022; Revised November 30, 2022; Accepted December 02, 2022)

Abstract: The chemical components of the family Gesneriaceae are complex and diverse. The compounds isolated from the plants of this family mainly include flavonoids, phenylethanoid glycosides, terpenoids, anthraquinones, organic carboxylic acids, and steroids. Gesneriaceae have a wide range of pharmacological activities, including antioxidant, antibacterial, anti-tumor, enzyme inhibition and activation, cytotoxicity, and anti-Alzheimer activities. In this study, the chemical components and pharmacological activities of Gesneriaceae in recent years were summarized to provide a reference for future studies.

Keywords: Gesneriaceae; chemical components; flavonoids; phenylethanoid glycosides; pharmacological activity. © 2022 ACG Publications. All rights reserved.

1. Introduction

Gesneriaceae are distributed in eastern and southern Asia, Africa, southern Europe, Oceania, South America, and tropical to temperate areas of Mexico. The plants of the Gesneriaceae family are widely distributed in China [1]. There are 805 known plants (including infraspecies) found in China as of December 31, 2021. They belong to 45 genera and are mainly distributed in southwest to south China, and endemic and narrowly distributed species are abundant [2]. At present, Gesneriaceae plants are widely used for ornamental planting because of their simple reproduction and good flowering. In addition, a variety of chemical constituents and pharmacological activities of the plants from the Gesneriaceae family have been discovered and extensively studied. Liu et al. [3] for the first time isolated two flavonoids, 7-hydroxy-6,8,4'-trimethoxy-5-O- β -D-glucoflavone glycoside (**9**) and 7-hydroxy-6,8,4'-trimethoxy-5-O- $[\beta$ -D-glucose-(1 \rightarrow 6)]- β -D-glucose flavonoid glycoside (**10**), from *Lysionotus pauciflorus* Maxim (*L. pauciflorus*), from the family Gesneriaceae. A survey of the literature has identified more than 200 compounds from Gesneriaceae over the past few decades. However, few reviews have systematically evaluated the chemical constituents and pharmacological

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activities of this family. In this review, we discuss the research progress on Gesneriaceae in terms of chemistry and pharmacology by collecting and reviewing the related literature both in China and abroad. This review provides a comprehensive and in-depth perspective for the further investigation of the Gesneriaceae family.

2. Traditional uses

Plants of the Gesneriaceae family are widely distributed and often used in folk medicine to treat a variety of diseases. In China, some ethnic minorities mostly use the whole plant as medicine and analyze its taste, meridian, and efficacy in combination with the theory of traditional Chinese medicine. For example, *L. pauciflorus* whole plants are used in medicine to clear lung phlegm, stop bleeding, and remove dampness stagnation, and the Miao folk have utilized this for the treatment of lymphatic tuberculosis [4, 5]. *Chirita longgangensis* W. T. Wang var. *hongyao* S. Z. Huang (*C. longgangensis* var. *hongyao*) is sweet, astringent, flat, and has been used to nourish the blood, relieve pain due to swelling, and improve blood circulation, and detoxify and has been commonly used in Zhuang medicine [6, 7]. It is recorded in *Yunnan Chinese Herbal Medicine* and *Commonly Used Herbs in Kunming* as bitter, pungent, and cold. Table 1 summarizes the scientific names, aliases, geographical distribution and traditional uses of eight well-studied genera from the Gesneriaceae family.

Table 1. Traditional uses of eight species of Gesneriaceae

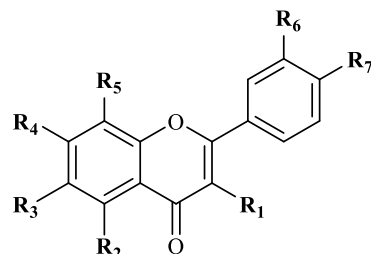
Scientific Name	Common Name	Distribution	Traditional Uses
<i>L. pauciflorus</i>	Cherhnexjenlvieeb, Shi diaolan, Yan gang dou (Chinese)	China Vietnam Japan	Treatment of chronic bronchitis, fall injury, and lung heat cough [8], Miao folk treatment for lymphatic tuberculosis [4, 5]
<i>C. longgangensis</i> var. <i>hongyao</i>	Shao yao, Mo guhua (Chinese)	China Vietnam Japan Mongolia Siberia	Treatment of early cold upper respiratory tract infection and irregular menstruation [7], treatment of physical weakness, anemia, and fracture [9]
<i>Hemiboea subcapitata</i> Clarke (<i>H. subcapitata</i>)	Hu shanye, Si taihua (Chinese)	China	Treatment of poisoning and boils [10]
<i>Corallodiscus kingianus</i> (<i>C. kingianus</i>)	Shi lianhua (Chinese)	China Sikkim Bhutan	Treatment of hot diarrhea [11]
<i>Boea hygrometrica</i> (Bge.) R. Br (<i>B. hygrometrica</i>)	Niu ercao, Mao erduo (Chinese)	China	Treatment of traumatic injuries, low back pain, traumatic hemorrhage, and otitis media [12]
<i>Corallodiscus flabellatus</i> (Craib) Burt (<i>C. flabellatus</i>)	Shi hua, Yan zhijia (Chinese)	Southwest China	Treatment of irregular menstruation, palpitations, sore carbuncle, and intractable tinea [13, 14]
<i>Oreocharis acericula</i> (S.Moore) Clarke (<i>O. acericula</i>)	Yan baicai (Chinese)	Southern China	Treatment of traumatic injuries, carbuncles, boils, and various bleeding [15]
<i>Chirita eburnea</i> Hance (<i>C. eburnea</i>)	Shi sanqi, Shi huer (Chinese)	Southern China	To treat tuberculosis and hypertension in Guangxi folk [16]

3. Chemical Constituents

3.1. Flavonoids and Their Glycosides

Flavonoids and their glycosides have been widely studied in Gesneriaceae plants. The flavonoid skeleton of Gesneriaceae plants mainly includes flavonoids, flavonols, dihydroflavonoids, and dihydroflavonols. Flavonoids often exist in the form of glycosides, aerobic glycosides, and carbon glycosides, and most of the glycosides consist of glucose units.

Flavonoids and their glycosides are derived from 10 species of Gesneriaceae. Apigenin (**6**) and 5,7,4'-trihydroxy-6-methoxy-flavone (**7**) were isolated from *A. superbis* for the first time [17]. Moreover, 5,7,3',4'-tetrahydroxy-6-C- β -D-glucopyranoside (**26**) has been isolated and elucidated from ethyl acetate and n-butanol fractions of *Chirita linearifolia* W. T. Wang (*C. linearifolia*) [18]. Figure 1 shows the structure of flavonoids and their glycosides. Table 2 summarizes the names, sources, and references of flavonoids and their glycosides.



	R ₁	R ₂	R ₃	R ₄	R ₅	R ₆
1	H	OH	OMe	OH	OMe	H
2	H	OH	OH	OMe	OMe	H
3	H	OH	OMe	OGlc	OMe	H
4	H	OH	H	OMe	H	H
5	H	OH	OMe	OH	H	H
6	H	OH	H	OH	H	H
7	H	OH	OMe	OH	H	H
8	H	OH	OMe	Glc(1 \rightarrow 6)GlcO	OMe	H
9	H	OGlc	OMe	OH	OMe	H
10	H	Glc(1 \rightarrow 6)GlcO	OMe	OH	OMe	H
11	OH	OH	OMe	OH	OMe	H
12	H	OH	Glc	OMe	H	H
13	H	OH	OMe	OMe	Glc	H
14	H	OH	OMe	OMe	Xyl(1 \rightarrow 2)Glc	OH
15	H	OH	OMe	OMe	Xyl(1 \rightarrow 2)Glc	H
16	H	OH	OMe	OMe	Api(1 \rightarrow 2)Glc	H
17	OH	OH	OMe	OMe	Glc	H
23	H	OH	H	Rha(1 \rightarrow 6)GlcO	H	H
24	H	OH	OMe	Xyl(1 \rightarrow 2)XylO	H	H
25	OH	OH	H	OH	H	H
27	H	OH	OMe	Rha(1 \rightarrow 6)GlcO	H	H
28	H	OH	Glc	OH	H	OH
35	H	OH	Ara	OH	Gal	H

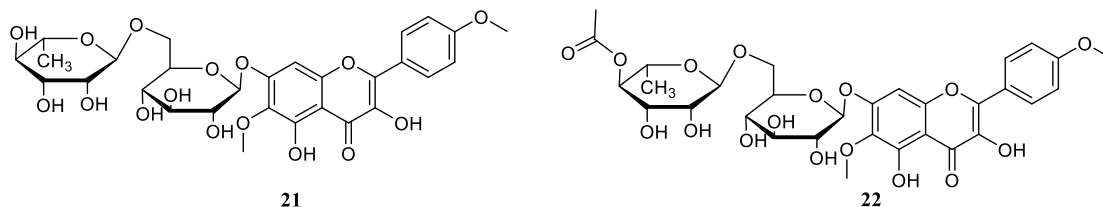
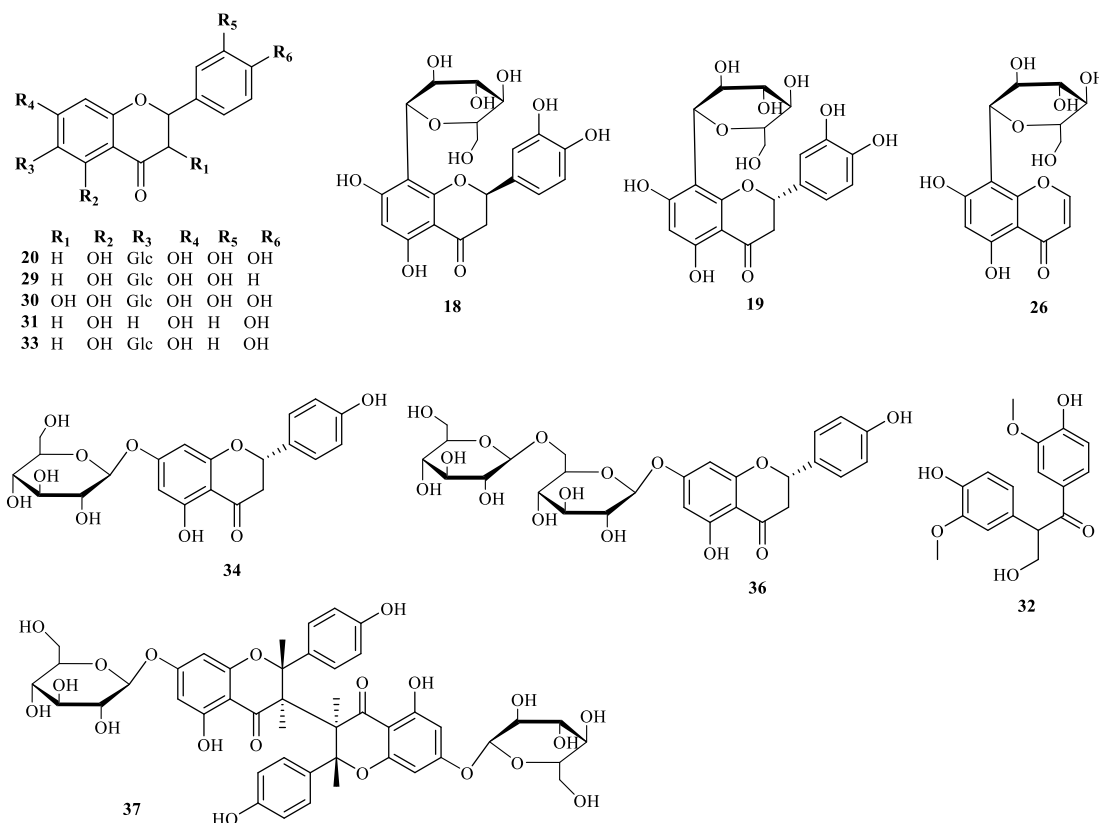


Figure 1. Flavonoids and their glycosides isolated from the genus

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**Figure 1.** Flavonoids and their glycosides isolated from the genus (*continued..*)**Table 2.** Chemical constituents of flavonoids and their glycosides

No	Compound Name	Source	Ref.
1	5,7-dihydroxy-6,8,4'-trimethoxy flavone		
2	5,6,4'-trihydroxy-7,8-dimethoxy flavone	<i>L. pauciflorus</i>	[1, 19-21]
3	5-hydroxy-6,8,4'-trimethoxy-flavone-7-O- β -D-glucoside		
4	5-hydroxy-7,4'-dimethoxy flavone	<i>C. kingianus</i>	[1]
5	5,7-dihydroxy-6,4'-dimethoxy flavone		
6	Apigenin	<i>A. superbis</i>	[17]
7	5,7,4'-trihydroxy-6-methoxy flavone	<i>Chirita fimbrisepala</i> Hand.-Mazz (<i>C. fimbrisepala</i>)	[17, 22] [23]
8	5-hydroxy-6,8,4' -trimethoxy-7-O-[β -D-glucose-(1 \rightarrow 6)]- β -D-glucoseflavonoid glycoside	<i>L. pauciflorus</i>	[19]
9	7-hydroxy-6,8,4'-trimethoxy-5-O- β -D-glucoflavone glycoside		
10	7-hydroxy-6,8,4'-trimethoxy-5-O-[β -D-glucose-(1 \rightarrow 6)]- β -D-glucose flavonoid glycoside	<i>L. pauciflorus</i>	[3, 19, 20]

11	5,7-dihydroxy-6,8,4'-trimethoxy flavonol		
12	4',5-dihydroxy-7-methoxy-6-C- β -D-glucoflavone glycoside	<i>L. pauciflorus</i>	[20]
13	4',5-dihydroxy-6,7-dimethoxy-8-C- β -D-glucoflavone glycoside		
14	5,3',4'-trihydroxy-6,7-dimethoxy-8-C-[β -D-xylose-(1 \rightarrow 2)]- β -D-glucoflavonecarboglycoside		
15	5,4'-dihydroxyl-6,7-dimethoxy-8-C-[β -D-xylose-(1 \rightarrow 2)]- β -D-glucoflavonecarboglycoside	<i>C. flabellatus</i>	[24]
16	5,4'-dihydroxyl-6,7-dimethoxyl-8-C-[β -D-apiofuranosyl (1 \rightarrow 2)]- β -D-glucopyranosyl flavone		
17	5,4'-dihydroxy-6,7-dimethoxy-8-C- β -D-glucose flavonol	<i>B. hygrometrica</i>	[25]
18	2 (R)-Eriodictyol-8-C-B-D-glucopyranoside	<i>C. longgangensis</i> var. <i>hongyao</i>	[26]
19	2 (S)-Eriodictyol-8-C-B-D-glucopyranoside		
20	5,7,3',4'-tetrahydroxy-6-C- β -D-glucodihydroflavone carbon glycoside	<i>C. linearifolia</i>	[18]
21	Pectolinarin		
22	4'''-acetyl-Pectolinarin	<i>C. flabellatus</i>	[27]
23	linarin		
24	mahuangchiside	<i>C. fimbrisepala</i>	[23]
25	kaempferol		
26	5,7-dihydroxy-8-C- β -D-glucochroone carbon glycoside		
27	5-hydroxy-6,4'-dimethoxy-7-[α -L-rhamnose-(1 \rightarrow 6)]-O- β -D-glucochromone glucoside		
28	5,7,3',4'-tetrahydroxy-6-C- β -D-glucoflavonol carbon glycoside	<i>Aeschynanthus moningeriae</i> (<i>A. moningeriae</i>)	[28]
29	5,7-dihydroxy-6-C- β -D-gluco dihydro flavone carbon glycoside		
30	5,7,3',4'-tetrahydroxy-6-C- β -D-gluco dihydro flavonol carbon glycoside		
31	naringenin		
32	evofolin B		
33	hemiphloin		
34	prunin	<i>Aeschynanthus bracteatus</i> Wall. ex A. DC. (<i>A. bracteatus</i>)	[29]
35	corymboside		
36	pyrroside B		
37	ormocarpin		

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3.2. Phenylethanol Glycosides

The number of glyco groups in most phenylethanol glycosides in Gesneriaceae ranges from one to three, and the types of glyco groups are rich and varied. In addition, aromatic acyl substituents are often associated, and coffee acyl substituents are the most common. The phenylethanol glycosides described in this review were derived from eight genera of Gesneriaceae. Figure 2 shows the structures of phenylethanol glycosides. Table 3 summarizes the names, sources, and references of phenylethanol glycosides.

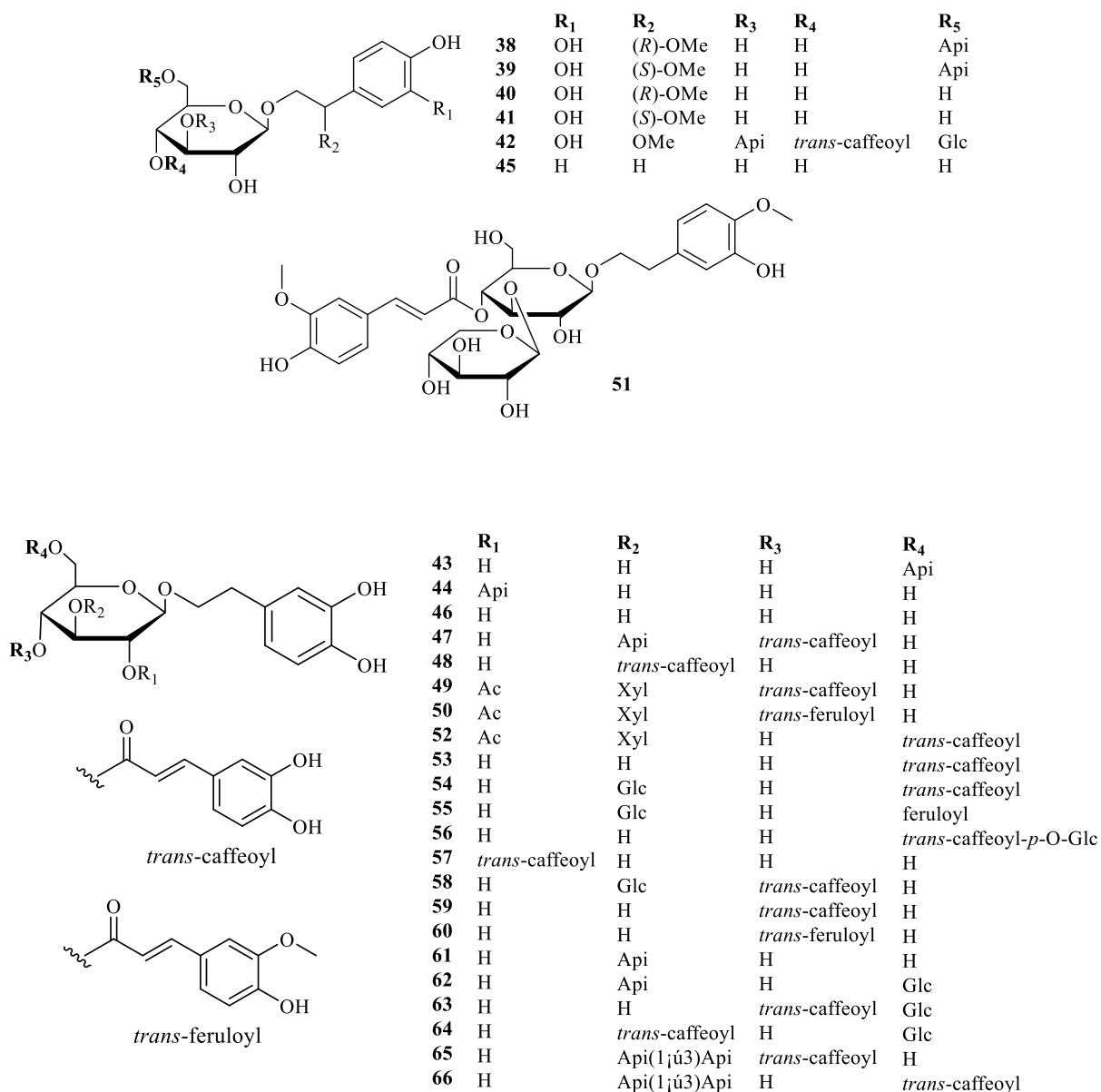


Figure 2. Phenylethanol glycosides isolated from the genus

Table 3. Chemical constituents of phenylethanol glycosides

No	Compound Name	Source	Ref.
38	(7S)-methoxyl-3,4-dihydroxyphenylethanol-8-O- β -D-apiofuranosyl (1 \rightarrow 6)- β -D-glucopyranoside		
39	(7R)-methoxyl-3,4-dihydroxyphenylethanol-8-O- β -D-apiofuranosyl (1 \rightarrow 6)- β -D-glucopyranoside		
40	(7S)-methoxyl-3,4-dihydroxyphenylethanol-8-O- β -D-glucopyranoside		
41	(7R)-methoxyl-3,4-dihydroxyphenylethanol-8-O- β -D-glucopyranoside		
42	(7 ξ) -methoxyl-3,4-dihydroxyphenylethanol-8-O- β -D-apiofuranosyl (1 \rightarrow 3)-[β -D-glucopyranosyl (1 \rightarrow 6)]-4-O-trans-caffeoyl- β -D-glucopyranoside	<i>C. flabellatus</i>	[27]
43	3,4-dihydroxyphenylethanol-8-O- β -D-apiofuranosyl (1 \rightarrow 6)- β -D-glucopyranoside		
44	3,4-dihydroxyphenylethanol-8-O- β -D-apiofuranosyl (1 \rightarrow 2)- β -D-glucopyranoside		
45	4-hydroxyphenylethanol-8-O- β -D-glucopyranoside		
46	3,4-dihydroxyphenylethanol-8-O- β -D-glucopyranoside		
47	isonuomioside A		
48	sanangoside		
49	aeschynanthoside A		
50	aeschynanthoside B	<i>A. bracteatus</i>	[29]
51	aeschynanthoside C		
52	aeschynanthoside D		
53	3,4-dihydroxyphenyl alcohol-6-O-caffeoyl- β -D-glucopyranoside (calceolarioside B)	<i>C. longgangensis</i> var. <i>hongyao</i>	[30] [26, 31]
54	3,4-dihydroxyphenyl alcohol- β -D-glucopyranosyl (1 \rightarrow 3)-6-O-caffeoyl- β -D-glucopyranoside (plantainoside D)	<i>C. longgangensis</i> var. <i>hongyao</i>	[31]
55	3, 4-dihydroxyphenyl alcohol-3-O-cafeoyl- β -D-glucopyranoside (plantainoside A)		
56	3, 4-dihydroxyphenyl alcohol-6-O-glucopyranosylcafeoyl- β -D-glucopyranoside (chiritoside C)	<i>C. eburnea</i>	[32]
57	3, 4-dihydroxyphenyl alcohol-2-O-cafeoyl- β -D-glucopyranoside (plantaninoside B)		
58	3, 4-dihydroxy-phenyl alcohol- β -D-glucopyranosyl (1 \rightarrow 3)-4-O-cafeoyl- β -D-glucopyranoside (plantamajoside)	<i>C. longgangensis</i> var. <i>hongyao</i>	[30] [26] [32]
59	3, 4-dihydroxyphenyl alcohol-4-O-cafeoyl- β -D-glucopyranoside	<i>C. eburnea</i>	[32]

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60	β -D-Glucopyranoside,2-(3,4-dihydroxyphenylethyl,4-[(2E)-3-(4-hydroxy-3-methoxyphenyl)-2-propenoate]	<i>C. linearifolia</i>	[33]
61	1'-O- β -D-(3,4-dihydroxyphenyl)-ethyl- β -D-apiofuranosyl (1 \rightarrow 3')-glucopyranoside	<i>C. flabellatus</i>	[34]
62	1'-O- β -D-(3,4-dihydroxyphenyl)-ethyl- β -D-apiofuranosyl (1 \rightarrow 3')- β -D-glucopyranosyl (1 \rightarrow 6')-glucopyranoside		
63	lugrandoside	<i>Selaginella tarmariscina</i> (Beauv.) Spring	[34]
64	isolugrandoside		
65	paraboside A	<i>Paraboea glutinosa</i> (P. <i>glutinosa</i>)	[35]
66	paraboside B		

3.3. Quinones

There are also many quinones in the plants of Gesneriaceae family, most of which are 9,10-anthraquinones. The substituents are mostly hydroxyl and methyl groups. The quinones in this review were derived from seven members of Gesneriaceae. Figure 3 shows the structures of quinone compounds. Table 4 summarizes the names, sources, and references of quinones.

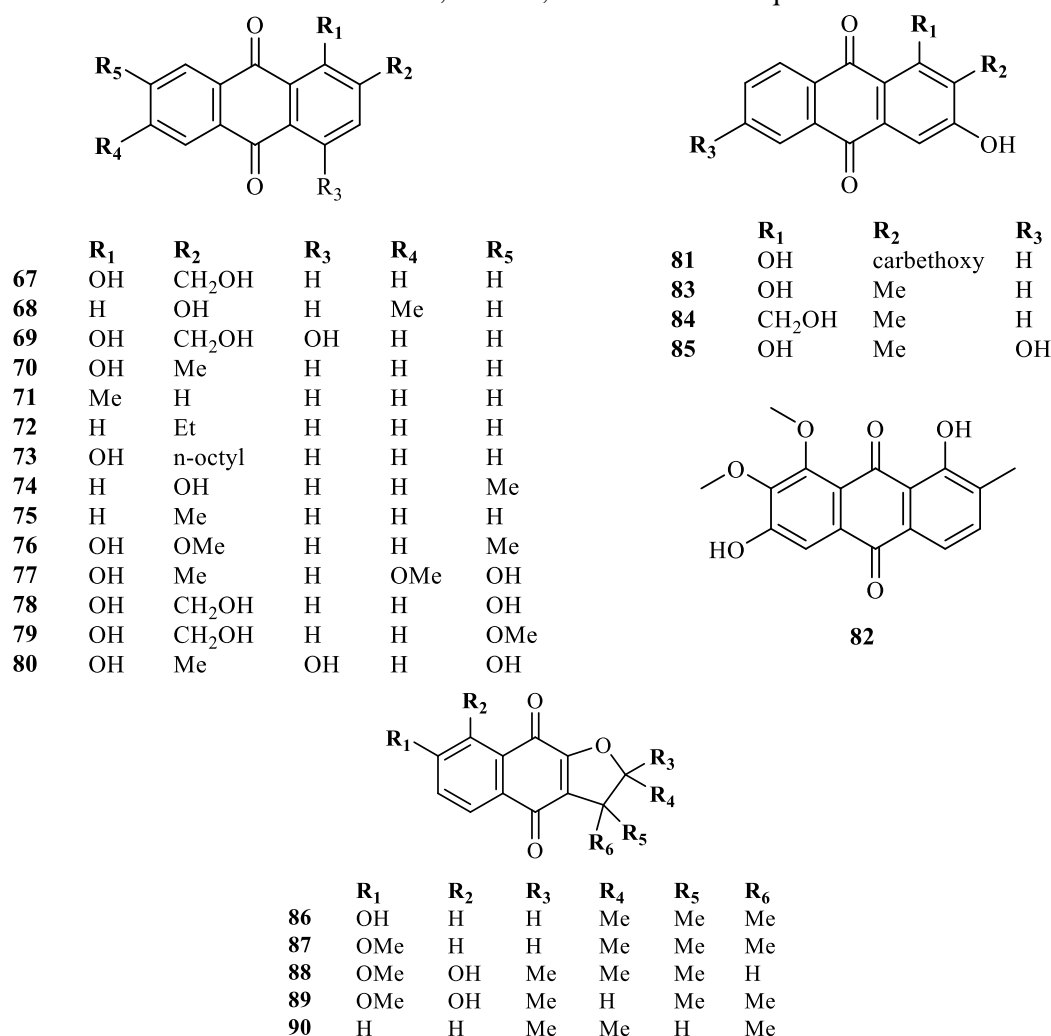


Figure 3. Quinones isolated from the genus

Table 4. Chemical constituents of quinones

No.	Compound Name	Source	Ref.
67	1-hydroxy-2-hydroxymethyl anthraquinone		
68	2-hydroxy-6-methyl anthraquinone	<i>C. linearifolia</i>	[18]
69	1,4-dihydroxy-2-hydroxymethyl anthraquinone		
70	1-hydroxy-2-methyl anthraquinone		
71	1-methyl anthraquinone	<i>C. linearifolia</i>	[33]
72	2-ethyl anthraquinone		
73	2-octyl-1-hydroxy-9,10-anthraquinone		
74	2-hydroxy-7-methyl-9,10-anthraquinone	<i>C. longgangensis</i> var. <i>hongyao</i>	[9]
75	2-methyl-9,10-anthraquinone		
76	1-hydroxy-2-methoxy-7-methy lanthraquinone		
77	1,7-dihydroxy-6-methoxy-2-methy lanthraquinone	<i>C. eburnea</i>	[36]
78	1,7-dihydroxy-2-hydroxymethyl-9,10-anthraquinone		
79	1-hydroxy-7-methoxy-2-hydroxymethyl-9,10-anthraquinone	<i>H. subcapitata</i>	[37]
80	1,4,7-trihydroxy-2-methyl-9,10-anthraquinone		
81	1,3-dihydroxy-2-ethoxycarbonyl-9,10-anthraquinone		
82	rhynchotechol		
83	rubiadin	<i>Rhynchotechum</i> <i>vestitum</i> (<i>R. vestitum</i>)	[38]
84	Rubiadin-1-methylether		
85	6-hydroxy-rubiadin		
86	7-hydroxy-2,3-dihydro-2,3,3-trimethylnaphtho [2,3-b] furan-4,9-dione	<i>C. longgangensis</i> var. <i>hongyao</i>	[9, 26]
87	(3R)-7-methoxy-R-dunnione	<i>C. linearifolia</i>	[18]
88	tenuiflorone		
89	7-metoxy-8-hydroxy- α -dunnione	<i>Paliavana tenuiflora</i> Mansf (<i>P. tenuiflora</i>)	[39]
90	α -isodunnione	<i>Sinningia</i> <i>allagophylla</i> (<i>S.</i> <i>allagophylla</i>)	[40]

3.4. Phenolic Acids

Phenolic acids are the most important organic acid, and there are a variety of phenolic acids in Gesneriaceae. This review describes phenolic acids from 11 species of Gesneriaceae plants. Some polyesters are also found in these compounds. For example, compounds **101**, **103** [20], and **115** [41] were separated from *L. pauciflorus*. Figure 4 shows the structures of phenolic acids. Table 5 summarizes the names, sources, and references of phenolic acids.

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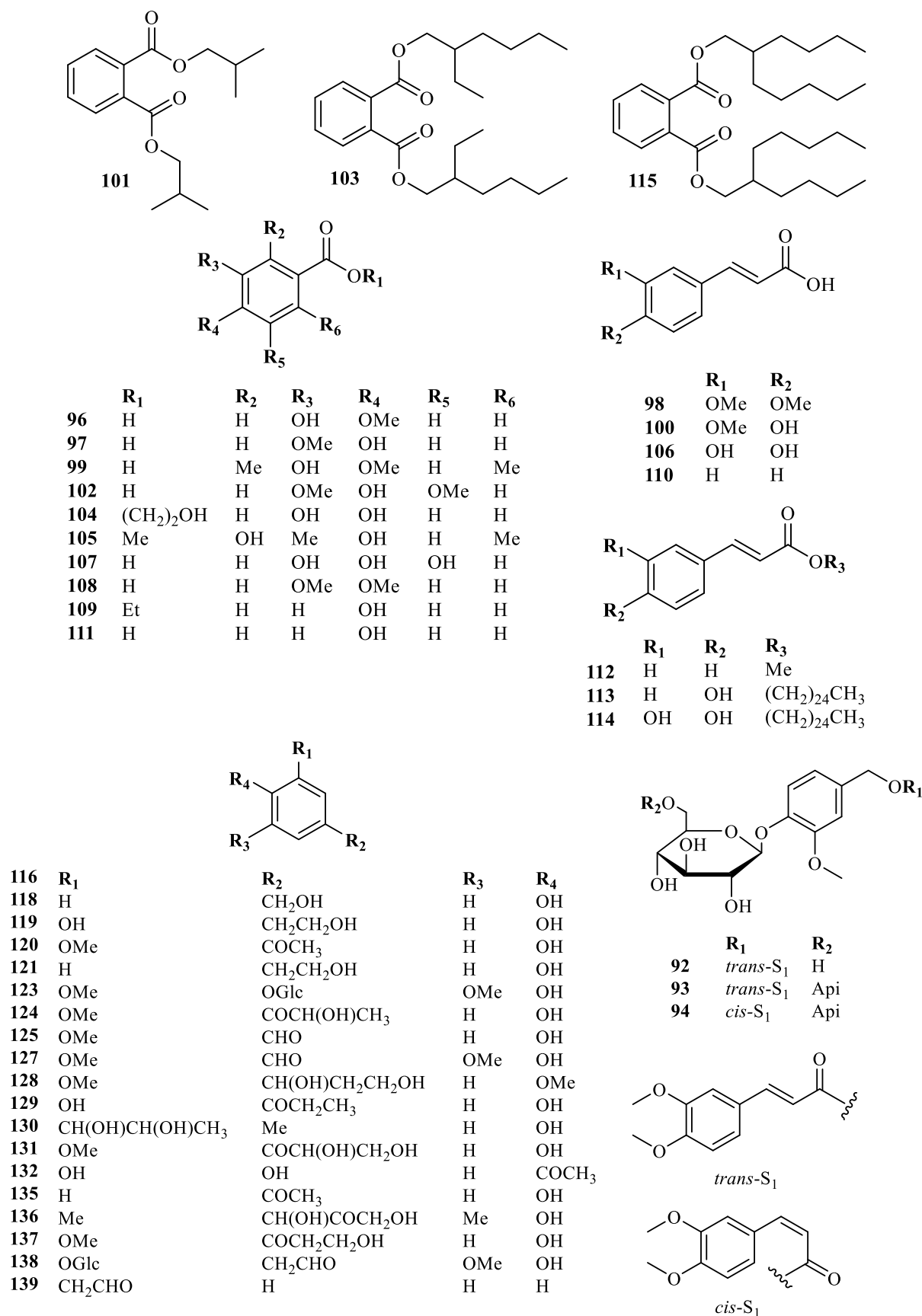


Figure 4. Phenolic acids isolated from the genus

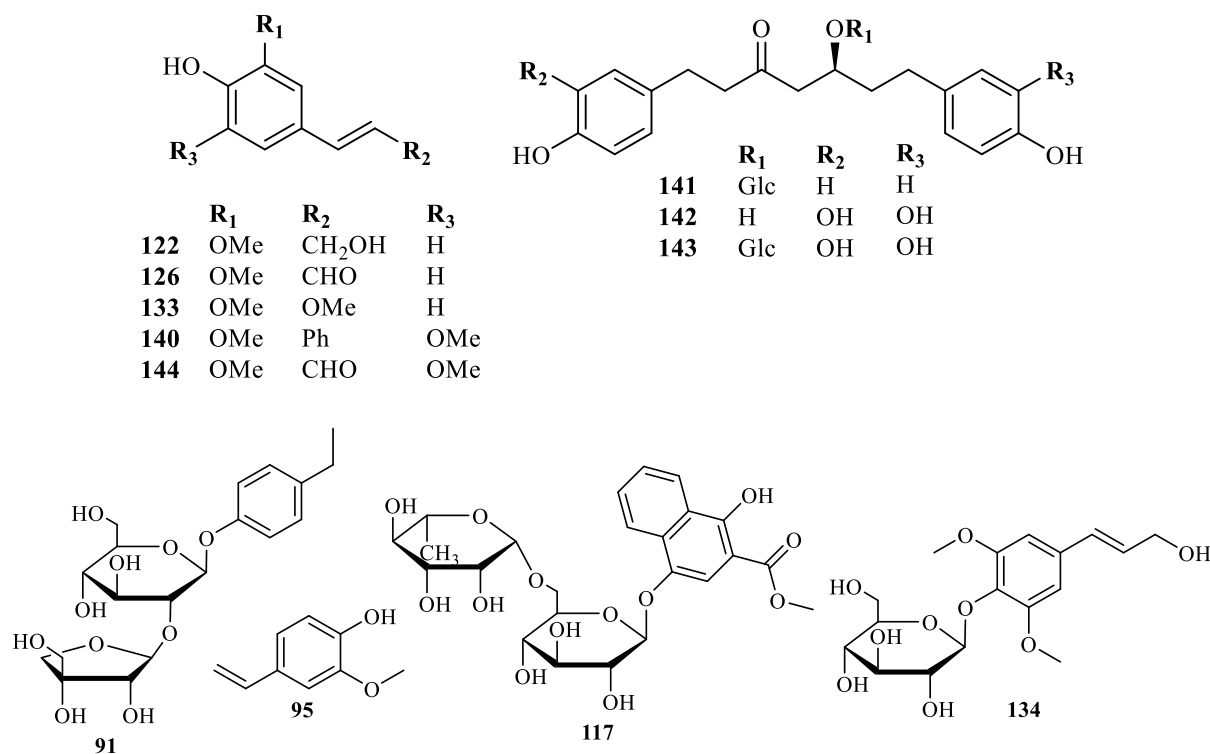


Figure 4. Phenolic acids isolated from the genus (*continued..*)

Table 5. Chemical constituents of phenolic acids

No.	Compound name	Source	Ref.
91	1'-O-β-D-(4-hydroxyphenylethyl)-β-D-apiofuranosyl (1→2')-glucopyranoside	<i>B. hygrometrica</i>	[25]
92	paraboside I		
93	paraboside II	<i>P. glutinosa</i>	[35]
94	paraboside III		
95	4-ethenyl-2-methoxyphenol	<i>O. acericula</i>	[42]
96	isovanillic acid	<i>Aeschynanthus longicaulis</i>	[1,
97	vanillic acid	(<i>A. longicaulis</i>)	43]
98	3,4-dimethoxy cinnamic acid	<i>Puraboeo rutescens</i> (<i>P. rutescens</i>)	[1]
99	2,6-dimethyl-4-methoxy-5-hydroxybenzoic acid	<i>A. superbus</i>	[17]
100	ferulic acid	<i>L. pauciflorus</i> <i>A. bracteatus</i>	[20] [29]
101	diisobutyl phthalate		
102	syringic acid	<i>L. pauciflorus</i>	[20]
103	Phthalic acid-bis-(2-ethylhexyl) ester		
104	3,4-dihydroxy-phenylacetic acid glycol ester	<i>C. flabellatus</i>	[24]
105	2,4-dihydroxy-3,6-dimethylbenzoate	<i>B. hygrometrica</i>	[25]
106	caffeic acid	<i>C. longgangensis</i> var.	[26]
107	gallic acid	<i>hongyao</i>	

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108	veratric acid		
109	ethylparaben		
110	cinnamic acid	<i>A. bracteatus</i>	[29]
111	4-hydroxybenzoic acid		
112	phenyl ethyl 2-acetate	<i>O. acericula</i>	[42]
113	2-(p-hydroxyphenyl) ethyl hexacosanoate		
114	2-(3,4-dihydroxyphenyl) ethyl hexacosanoate	<i>P. tenuiflora</i>	[39]
115	bis (2-butylhexyl) phthalate	<i>L. pauciflorus</i>	[41]
116	P-hydroxyphenyl ethanol	<i>B. hygrometrica</i>	[25]
117	1,4-Dihydroxy-2-naphthalenecarboxylic acid methyl ester-4-O- β -L-Rhamnopyranosyl (1 \rightarrow 6)- β -D-Glucopyranoside	<i>C. longgangensis</i> var. <i>hongyao</i>	[9, 26]
118	hydroxytyrosol		
119	Apocynin	<i>C. flabellatus</i>	[24]
120	4-hydroxyphenylethanol	<i>C. flabellatus</i>	[27]
121	koaburaside		
122	coniferyl alcohol		
123	(R)-2-hydroxy-1 (4-hydroxy-3-methoxyphenyl) propan-1-one		
124	vanillin		
125	syringaldehyde		
126	coniferaldehyde		
127	1-(3,4-dimethoxyphenyl)-1,3-propanediol		
128	3,4-dihydroxypropiophenone		
129	1-(2-hydroxy-5-methoxyphenyl)-1,2-propanediol	<i>A. bracteatus</i>	[29]
130	C-veratroylglycol		
131	2,4-dihydroxyacetophenone		
132	piceol		
133	2-methoxy-4-(3-methoxy-1-propenyl)-phenol		
134	syringenin		
135	1,3-dihydroxy-1-(4-hydroxy-3,5-dimethoxyphenyl) propan-2-one		
136	x-hydroxypropioguaiacone		
137	alasanioside C		
138	Benzene acetaldehyde		
139	Benzene ethanol	<i>O. acericula</i>	[42]
140	3,5-dimethoxy-4-hydroxy- <i>trans</i> -stilbene	<i>L. pauciflorus</i>	[41]
141	platyphylloside		
142	hirsutanonol	<i>C. longgangensis</i> var. <i>hongyao</i>	[44]
143	hirsutanonol-5-O- β -D-glucopyranoside		
144	<i>trans</i> -3,5-dimethoxy-4-hydroxycinnamaldehyde	<i>A. bracteatus</i>	[45]

3.5. Terpenoids

Terpenoids are derived from meglutaric acid, and most of the molecular formula follows the $(C_5H_8)_n$ general formula. Terpenoids are abundant in Gesneriaceae, which include monoterpenes,

sesquiterpenes, diterpenes, triterpenes, and terpenes. Figure 5 shows the structures of terpenoids. Table 6 summarizes the names, sources, and references of terpenoids.

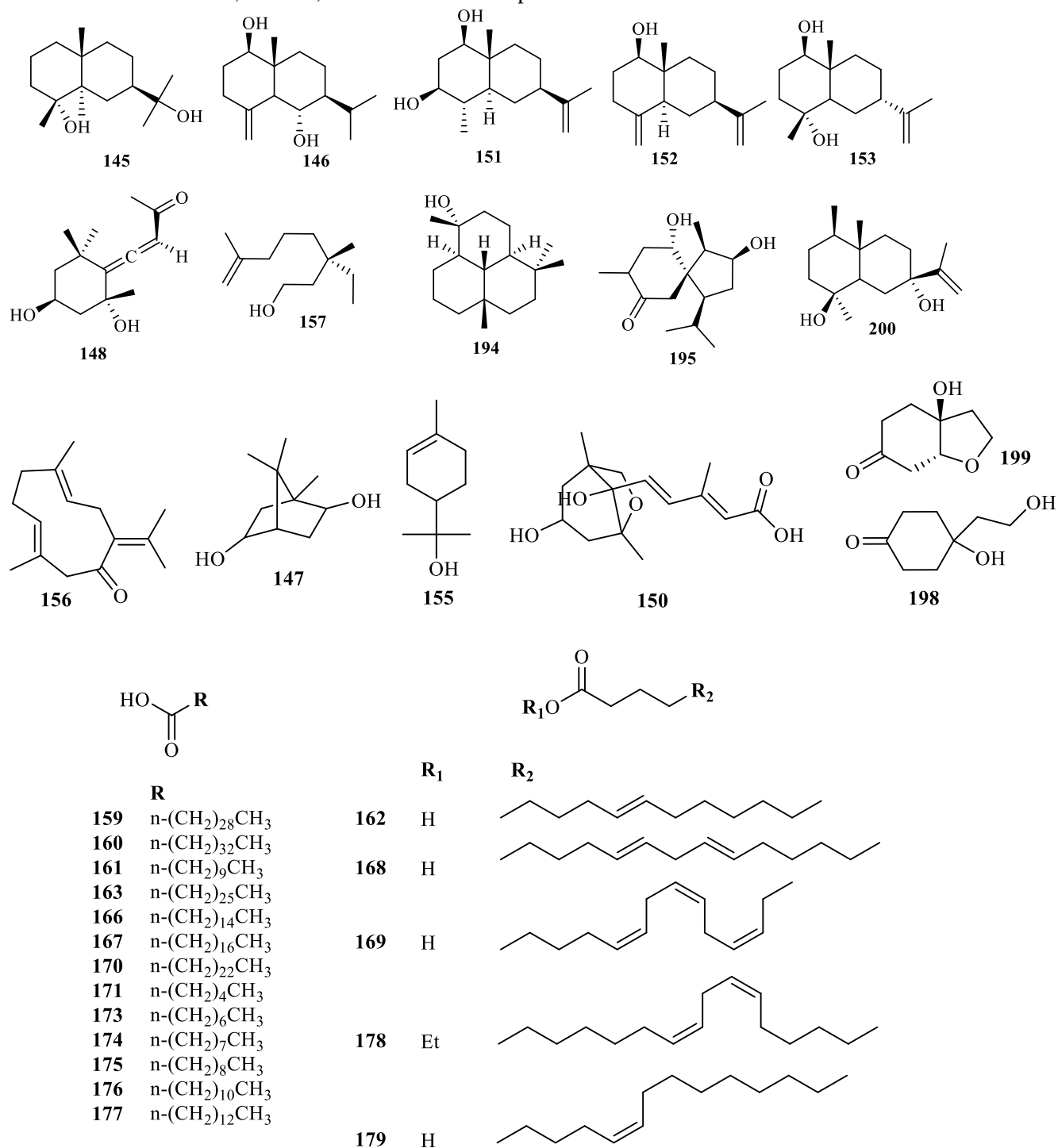


Figure 5. Terpenoids isolated from the genus

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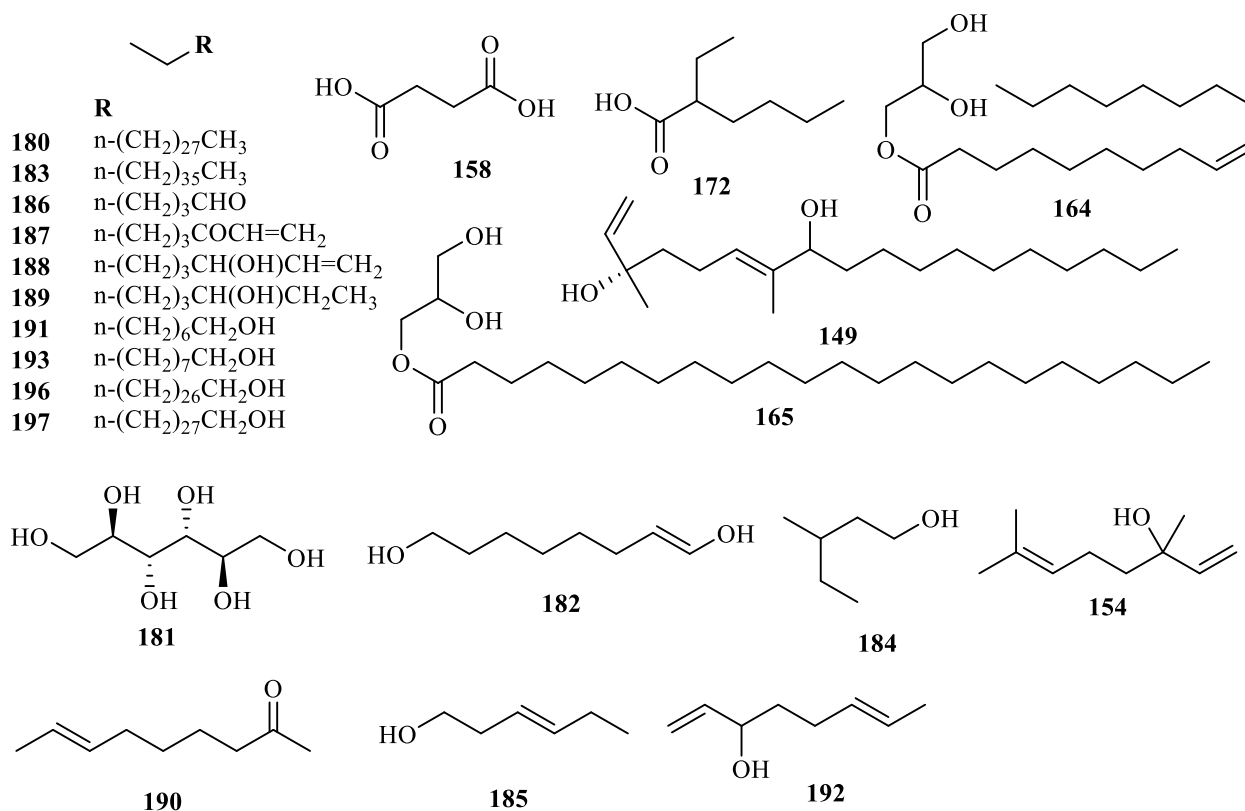


Figure 5. Terpenoids isolated from the genus (continued..)

Table 6. Chemical constituents of terpenoids

No.	Compound name	Source	Ref.
145	cryptomeridiol		
146	4 (15)-eudesmene-1β,6α-diol	<i>A. longicaulis</i>	[1, 43]
147	2,5-bornanediol		
148	grasshopper ketone		
149	9-hydroxylinalool	<i>A. bracteatus</i>	[29]
150	epidihydrophaseic acid		
151	AM-2	<i>Aeschynanthus mengxinensis</i> (<i>A. mengxinensis</i>)	[46]
152	AM-6		
153	AM-8		
154	linalool		
155	A-terpilenol	<i>O. acericula</i>	[42]
156	germacrone		
157	citronellol		
158	succinic acid		
159	N-hexadecanoic acid	<i>C. longgangensis</i> var. <i>hongyao</i>	[26]
160	N-tetradecanoic acid		
161	eleven carbonate	<i>C. linearifolia</i>	[18]
162	palmitic acid		
163	twenty-seven alkanolic acid		
164	glycerol monooleate	<i>C. linearifolia</i>	[33]
165	triglyceride dodicarboxylate		

166	palmitic acid	<i>C. kingianus</i>	[47, 48]
167	stearic acid	<i>Rhynchothecum ellipticum (R. ellipticum)</i>	[49]
168	9,12-octadecadienoic acid	<i>C. kingianus</i>	[47, 48]
169	(Z, Z, Z)-9,12,15-octadecanotrienoic acid		
170	tetracanoic acid	<i>R. vestitum</i>	[38]
171	hexanoic acid	<i>O. acericula</i> <i>A. moningeriae</i>	[42] [50]
172	2-ethyl-hexanoic acid		
173	octylic acid		
174	n-nonanoic acid		
175	decylic acid		
176	dodecanoic acid	<i>O. acericula</i>	[42]
177	myristic acid		
178	11,14-eicosadienoate ethyl ester		
179	oleic acid		
180	triacontane	<i>C. longgangensis var. hongyao</i>	[9, 26]
181	mannitol		
182	7-octene-1,6-diol	<i>A. bracteatus</i>	[29]
183	octatriacontane	<i>C. linearifolia</i>	[18]
184	2-ethyl-4-butanol		
185	3-ene-1-hexanol		
186	hexanal		
187	1-octene-3-ketone		
188	1-ene-3-octanol	<i>O. acericula</i>	[42]
189	3-octanol		
190	<i>trans</i> -2-enonone		
191	nonyl alcohol		
192	1,6-diene-3-octanol		
193	decanol		
194	presilphiperfolan-9-ol	<i>P. tenuiflora</i>	[39]
195	3,10-dihydroxyacoronene	<i>L. pauciflorus</i>	[41]
196	dodecyl alcohol	<i>L. pauciflorus</i>	[51]
197	N-triethanol		
198	cleroindicin B	<i>S. allagophylla</i>	[40]
199	cleroindicin C		
200	AM-15	<i>A. mengxinensis</i>	[46]

3.6. Triterpenoids

Triterpenes are composed of several isoprenoids joined end-to-end with a hydroxyl molecule removed. Triterpenoids in Gesneriaceae are pentacyclic triterpenoids, and their skeletons are of the ursane and oleanane type. Most of the 28 substituents are carboxyl or glycoside with glucose. Figure 6 shows the structures of triterpenes. Table 7 summarizes the names, sources, and references of triterpenes.

Chemical components and pharmacological activities of Gesneriaceae

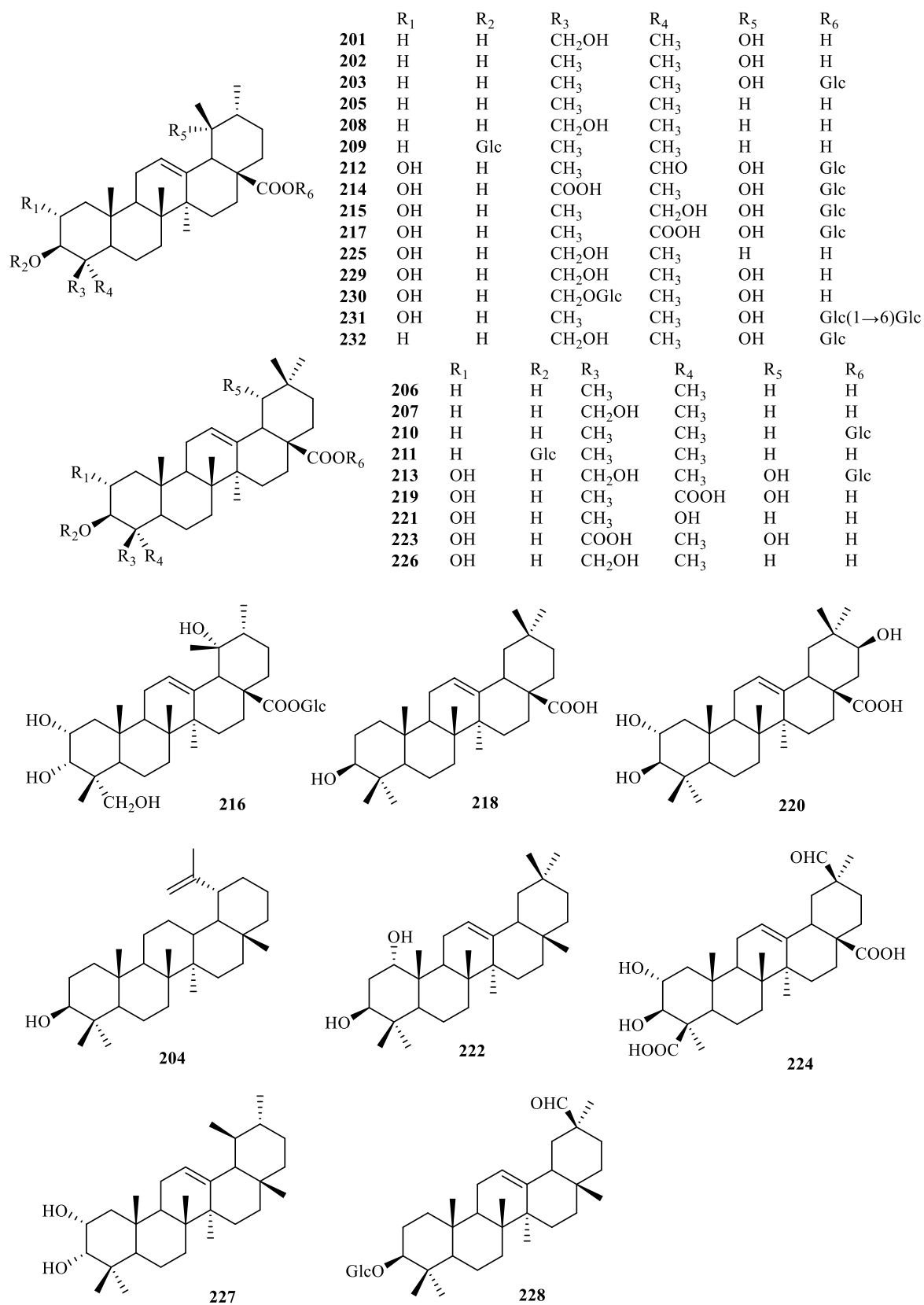


Figure 6. Triterpenoid isolated from the genus

Table 7. Chemical constituents of triterpenoid

No	Compound name	Source	Ref.
201	barbinervic acid	<i>A. moningeriae</i>	[1] [28]
202	3 β ,19 α -dihydroxy-12-ene-28-ursolic acid	<i>P. rutescens</i>	[1]
203	28-O- β -D-glucopyranosyl pomolic acid		
204	lupeol	<i>A. superbus</i> <i>Aeschynanthu maculatus</i> (<i>A. maculatus</i>)	[17, 22] [17]
205	3-hydroxy-20 (29)-lupen-28-oic acid	<i>A. superbus</i> <i>A. maculatus</i> <i>C. longgangensis</i> var. <i>hongyao</i> <i>C. linearifolia</i>	[17, 22] [17] [26] [33]
206	3-hydroxy-5,12-oleanadien-28-oic acid	<i>A. superbus</i>	[17, 22]
207	3,24-dihydroxy-12-oleanen-28-oic acid		
208	3,24-dihydroxy-12-ursen-28-oic acid		
209	12-ursolic acid-3-O- β -D-glucoside	<i>A. maculatus</i>	[17]
210	12-oleanolic acid-28-O- β -D-glucoside		
211	12-oleanolic acid-3-O- β -D-glucoside		
212	4-epipinfaensin		
213	paradrymonoside		
214	trachelosperogenin A1		
215	nigaichigoside F1		
216	nigaichigoside F2		
217	suavissimoside F1		
218	3 β -hydroxy-9,12-oleanadien-28-oic acid	<i>C. kingianus</i>	[46, 47]
219	2 α ,3 β ,19 β -trihydroxy-olean-12-ene-23,28-dioic acid	<i>A. mengxinensis</i>	[52]
220	2 α ,3 β ,21 β -trihydroxyolean-12-ene-28-oic acid		
221	2 α ,3 β ,23-trihydroxyurs-12-ene-28-oic acid	<i>A. mengxinensis</i>	[46, 52]
222	CK01	<i>C. kingianus</i>	[46]
223	AM-1	<i>A. mengxinensis</i>	[46]
224	AM-7		
225	AM-3		
226	AM-9		
227	AM-11		
228	oleanolane-3-O- β -D-glucoside	<i>R. ellipticum</i>	[49]
229	24-hydroxytormentic acid	<i>P. glutinosa</i>	[53]
230	24-hydroxytormentic acid ester glucoside		
231	28-O- β -D-glucopyranosyl (1 \rightarrow 6)- β -D-glucopyranosyl-24-hydroxytormentic acid		
232	2,3,19,24-tetrahydroxy-12-ene-28-ursolate-28-O- β -D-glucoside	<i>A. moningeriae</i>	[28]

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3.7. Steroids

Steroids are a class of thickened tetracyclic aliphatic compounds with a cyclopentane polyhydrophenanthrene nucleus. Steroidal compounds in Gesneriaceae plants contain 3-position hydroxyl groups or glycosides with glucose, and 20-position substituents are chained hydrocarbon groups. β -carotenoside (**235**) was isolated from the aboveground portion of *A. superbis* and *A. maculatus* [17, 22], and the compound was found in five species of Gesneriaceae. In addition, the methanol extract of *A. superbis* contained β -sitosterol (**236**) [17, 22], and β -sitosterol was found in up to 11 species of Gesneriaceae. Figure 7 shows the structures of steroids. Table 8 summarizes the names, sources, and references of steroids.

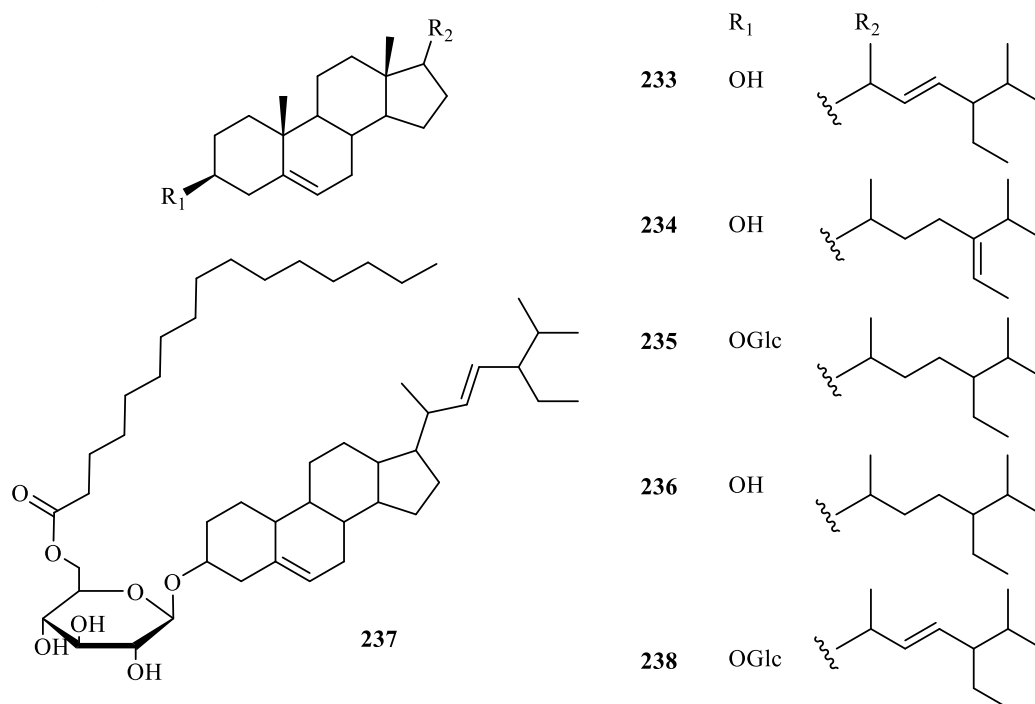


Figure 7. Steroids isolated from the genus

Table 8. Chemical constituents of steroids

No	Compound Name	Source	Ref.
233	stigmasterol	<i>R. ellipticum</i>	[17,
		<i>C. linearifolia</i>	22]
		<i>A. moningeriae</i>	[18]
		<i>A. maculatus</i>	[28]
		<i>P. tenuiflora</i>	[49]
234	Steroidal-5,22 (E)-diene-3- β -alcohol	<i>A. maculatus</i>	[17,
		<i>A. longicaulis</i>	22]
		<i>A. mengxinensis</i>	[43]
			[52]
235	β -daucosterol	<i>C. linearifolia</i>	[18,
		<i>R. ellipticum</i>	33]
		<i>P. glutinosa</i>	[49]
		<i>A. moningeriae</i>	[53]
		<i>C. longgangensis</i> var. <i>hongyao</i>	[28] [9]
236	β -sitosterol	<i>P. rutescens</i>	[1]
		<i>A. longicaulis</i>	[1]

		<i>C. longgangensis</i> var. <i>hongyao</i>	[26]
		<i>A. moningeriae</i>	[28]
		<i>C. linearifolia</i>	[33]
		<i>R. ellipticum</i>	[49]
		<i>P. glutinosa</i>	[53]
		<i>C. eburnea</i>	[36]
		<i>P. tenuiflora</i>	[39]
		<i>L. pauciflorus</i>	[54]
		<i>B. hygrometrica</i>	[12]
237	β -sitosteryl-D-glucoside-6'-palmitate	<i>C. longgangensis</i> var. <i>hongyao</i>	[9, 26]
238	3-O- β -D-glucostigmasterol glycoside	<i>A. moningeriae</i>	[28]

3.8. Volatiles and Fatty Oils

The lipid soluble components of *C. flabellatus* were extracted by the Soxhlet extraction method. The lipid soluble components were methyl esterified, and their chemical constituents were analyzed by GC-MS [55]. A total of 47 compounds were identified, including fatty acids, terpenes, alkanes, alcohols, ketones, and methyl ester, accounting for 84.34% of the liposoluble components.

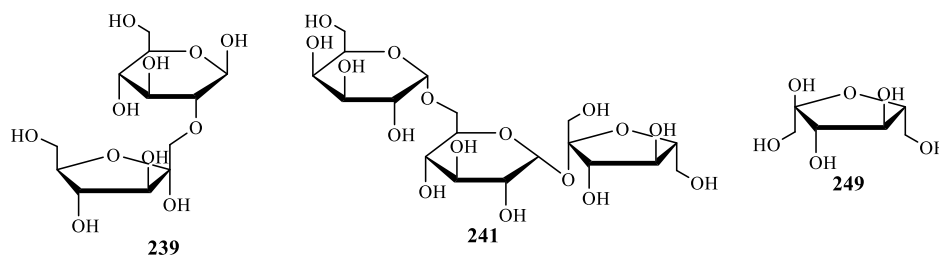
The chemical constituents of volatile oil in *A. moningeriae* were analyzed and extracted by SPME [50]. The components of the volatile oil were separated and identified by GC-MS, and the relative content of each component was determined by the area normalization method. Sixty-three types of components were separated. Among these, 58 components were identified, accounting for approximately 98.06% of the total chemical constituents. The ingredient with the highest content in this study was [2R-(2à,4aà,8aá)]-4a,8-dimethyl-2-(1-methylethenyl)-1,2,3,4,4a,5,6,8a-octahydronaphthalene (12.80%).

3.9. Other Components

In addition to the eight chemical compounds mentioned above, there are many other compounds in the plants of Gesneriaceae family, such as lignans and sugars.

Two naphthol dimer compounds allagophyllidimer D (**250**) and allagophyllidimer E (**251**) were isolated and identified in *S. allagophylla* [50]. Figure 8 shows the structures of other compounds. Table 9 summarizes the names, sources, and references of other compounds.

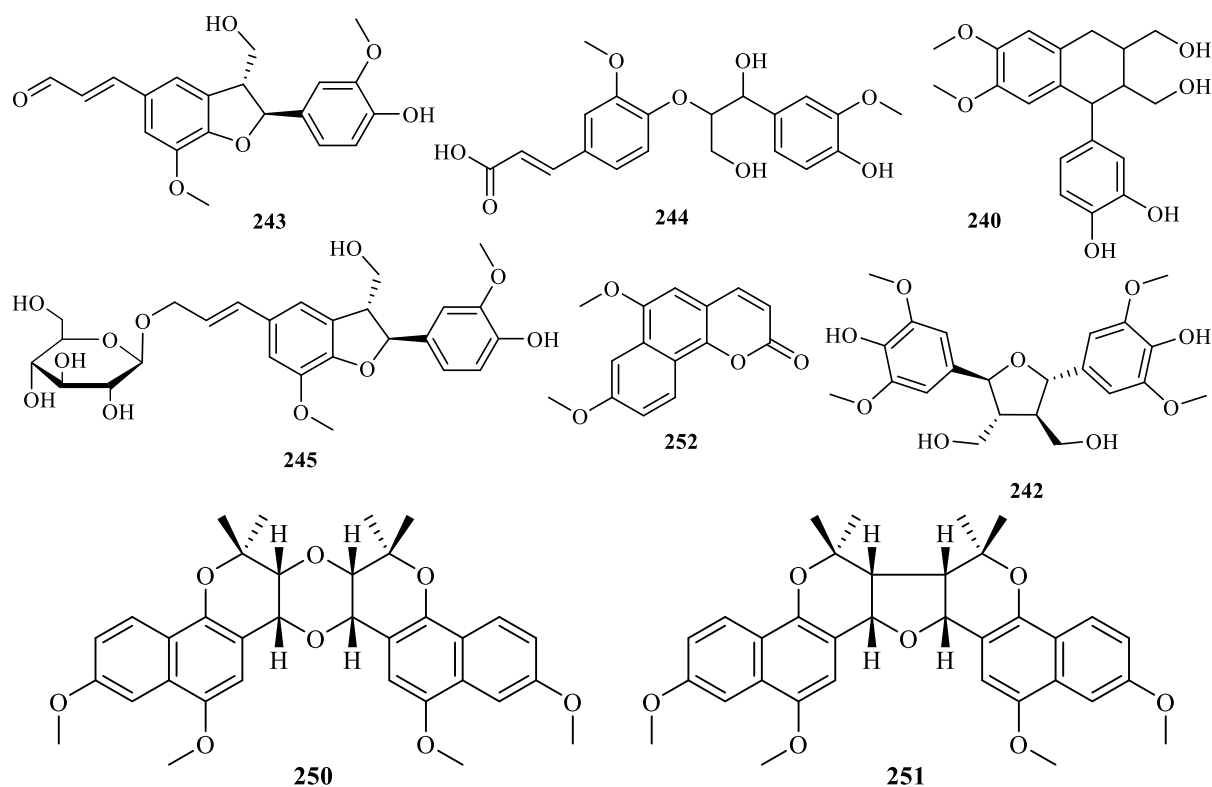
Ultra-high-performance liquid chromatography quadrupole time-of-flight mass spectrometry (UPLC-Q-TOF-MS) was used to analyze the chemical components in *L. wilsonii* [30]. The results showed that 57 components were identified, which included 42 phenylethanoid glycosides, five benzyl alcohol glycosides, six flavonoids, and four other components. Forty-three of these compounds were first identified in Gesneriaceae, and one benzyl alcohol glycoside may be a new compound.



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Table 9. Chemical constituents of other compounds

No	Compound Name	Source	Ref.
239	2-O- α -D-furan fructosyl- α -D-glucose	<i>B. hygrometrica</i>	[25]
240	3',4',9,9-Tetrahydroxy-4,5-dimethoxy-2,7'-cyclolignan	<i>C. longgangensis</i> var. <i>hongyao</i>	[9, 26]
241	D-(+)-raffinose	<i>L. pauciflorus</i>	[54]
242	icariol A2		
243	balanophonin		
244	guaiacylglycerol-b-ferulic acid ether		
245	(7S, 8R) dehydrodiconiferyl alcohol 9'-glucopyranoside	<i>A. bracteatus</i>	[29]
246	dianthoside		
247	6'-O- β -D-apiofuranosyl-dianthoside		
248	caprolactam	<i>C. linearifolia</i>	[18]
249	α -D-furanfructose	<i>B. hygrometrica</i>	[12]
250	allagophyllidimer D		
251	allagophyllidimer E		
252	6,8-dimethoxybenzocoumarin	<i>S. allagophylla</i>	[40]
253	warmingiin A		
254	warmingiin B		

**Figure 8.** Other components isolated from the genus

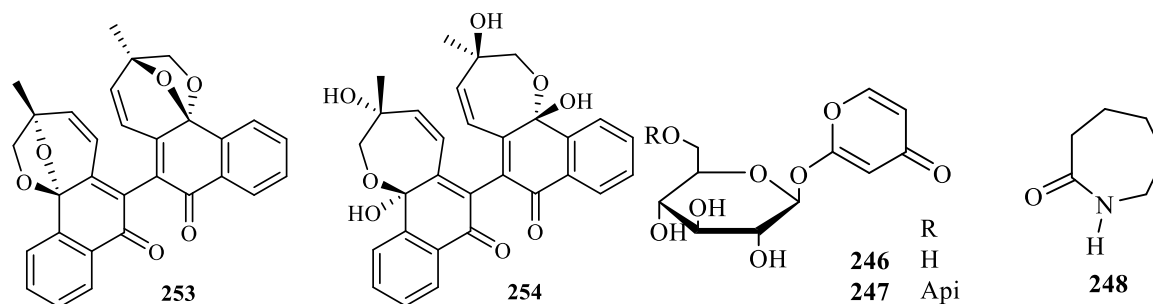


Figure 8. Other components isolated from the genus (*continued.*)

4. Pharmacological Activities

4.1. Antioxidant Activity

Tian [17] used DPPH, ABTS, and FRAP to comprehensively evaluate the antioxidant activities of different solvent extracts of *A. superbus*, *A. maculatus*, and *R. ellipticum* and the isolated compounds *in vitro*. The results showed that ethyl acetate and methanol fractions of *A. superbus* and ethyl acetate and methanol fractions of *A. maculatus* had better antioxidant activities than other fractions. The scavenging ability of the three plant extracts on DPPH free radicals and ABTS free radicals and the reducing ability on Fe^{3+} are all dose-dependent.

Qiao *et al.* [55] used a 96-well plate method to determine the antioxidant capacity of fat-soluble ingredients. The test results showed that the fat-soluble component of *C. flabellatus* had a certain scavenging effect on DPPH. Kang *et al.* [56] used DPPH, ABTS, and FRAP for the first time to study the *in vitro* antioxidant activity of *C. kingianus* and *A. mengxinensis*. It was found that among the four extracts, the methanol extract of *C. kingianus* had a stronger DPPH free radical scavenging ability ($\text{IC}_{50}=4.92 \mu\text{g/mL}$) than the positive control BHT ($\text{IC}_{50}=18.79 \mu\text{g/mL}$). The scavenging ability of the methanol extract of *C. kingianus* to ABTS free radicals ($\text{IC}_{50}=11.10 \mu\text{g/mL}$) was slightly lower than that of BHT ($\text{IC}_{50}=6.04 \mu\text{g/mL}$). In addition, the ability of extract of *C. kingianus* to reduce Fe^{3+} ($\text{FRAP}=2,403.77\pm 38.05 \mu\text{molTE/g}$) was higher than BHT ($\text{FRAP}=1,748.49\pm 3.46 \mu\text{molTE/g}$). Among the four extracts, the methanol extract of *C. kingianus* showed the highest antioxidant capacity.

Liu [57] used two analytical methods, namely, DPPH and FRAP, to determine the antioxidant activity of each fraction of *A. moningeriae* *in vitro*. The results showed that the order of the antioxidant activity of DPPH and FRAP *in vitro* was water fraction > ethyl acetate extraction fraction > 70% crude extract > petroleum ether fraction; 30% methanol enrichment fraction > 50% methanol enrichment fraction > 10% methanol enrichment fraction > 70% methanol enriched fraction. Chen [1] used the DPPH microplate method to determine the antioxidant activity of the compounds isolated from *A. longicaulis* and *L. pauciflorus*. Among these, 5,7-dihydroxy-6,8,4'-trimethoxyflavone (**1**) ($\text{IC}_{50}=17.01 \mu\text{g/mL}$) and vanillic acid (**97**) ($\text{IC}_{50}=13.63 \mu\text{g/mL}$) both exhibited significant antioxidant activity. However, the antioxidant activity of 5,6,4'-trihydroxy-7,8-dimethoxyflavone (**2**) ($\text{IC}_{50}=51.7 \mu\text{g/mL}$) was slightly lower than the positive control BHT.

4.2. Antibacterial Activity

For the first time, Chen [1] studied the antibacterial activities of extracts from three Gesneriaceae plants and their compounds. Analysis of the extracts of *A. longicaulis* and *L. pauciflorus* on SA, MRSA, and ESBLs-SA revealed that the antibacterial effect of *A. longicaulis* was better than that of *L. pauciflorus*, which mainly existed in the parts of petroleum ether and ethyl acetate. Tian [17] studied the *in vitro* antibacterial activity of *A. superbus*, *A. maculatus*, and *R. ellipticum* total methanol extracts. The total methanol fraction of *A. superbus* had an inhibitory effect on SA and MRSA. When the concentration was $250 \mu\text{g}\cdot\text{disc}^{-1}$, the inhibition zone for SA was 9 mm, and the inhibition zone for MRSA was 8 mm. The total methanol part of *A. maculatus* had an inhibitory effect on SA, and the

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inhibition zone was 9 mm; the total methanol part of *R. ellipticum* had an inhibitory effect on SA, and the inhibition zone was 8 mm.

Wei et al. [21] used the paper diffusion method to track the activity of the extracts from *P. rutescens* and *L. pauciflorus* and found that the ethyl acetate fractions of the two plants had the best antibacterial activity against SA, MRSA, and ESBLs-SA. Barbinervic acid isolated from *P. rutescens* had the best inhibition of SA activity ($IC_{50}=0.098$ g/L, **201**) and ESBLs-SA activity ($IC_{50}=0.270$ mg/L). Meanwhile, 3 β ,19 α -dihydroxy-12-ene-28-ursolic acid ($IC_{50}=0.130$ g/L, **202**) had the best inhibitory activity against MRSA.

Xu et al. [5] found that lysionotin (**1**) 200 μ g/mL *in vitro* test has a significant anti-TB effect. Zhang et al. [58] studied the clinical observation test of *L. pauciflorus* tablets in adjuvant treatment of retreatment of pulmonary tuberculosis. The experiment found that the negative conversion rate of sputum bacteria in the treatment group (95.7%) was significantly higher than that in the control group (79.6%) ($P<0.05$), and there were also significant differences between the two groups in drug-resistant cases ($P<0.01$). This may be due to the fact that the phytoplankton enhances the bactericidal activity of sensitive drugs and prevents the sensitive drugs from developing selective resistance.

4.3. Enzyme Activation/Inhibition Activity

For the first time, Chen [1] conducted α -glucosidase inhibitory activity experiments on monomer compounds, extracts, and various parts of *L. pauciflorus* Maxim and *A. longicaulis*, and found that *L. pauciflorus* total acetone water extract ($IC_{50}=415.30$ μ g/mL), ethyl acetate ($IC_{50}=312.67$ μ g/mL), and n-butanol ($IC_{50}=884.10$ μ g/mL) have α -glucosidase inhibitory activity, and the ethyl acetate part has the highest activity. The α -glucosidase inhibitory activity of these three parts is higher than that of all parts of *A. longicaulis*. 5,6,4'-trihydroxy-7,8-dimethoxy flavone (**2**) ($IC_{50}=55.50$ μ g/mL) and vanillic acid (**97**) ($IC_{50}=13.63$ μ g/mL) had α -glucosidase inhibitory activity.

Tian [17, 59] used the 96-well plate method to study the *in vitro* α -glucosidase inhibitory activity of *A. superbus*, *A. maculatus*, and *R. ellipticum* solvent extracts. The results showed that the *in vitro* α -glucosidase inhibitory activity of each part of the three plants was in the order of *R. ellipticum* petroleum ether component ($IC_{50}=112.4$ μ g/mL) > *A. maculatus* petroleum ether component ($IC_{50}=127.3$ μ g/mL) > *A. superbus* petroleum ether component ($IC_{50}=230.2$ μ g/mL) > *R. ellipticum* ethyl acetate component ($IC_{50}=351.4$ μ g/mL) > *A. maculatus* ethyl acetate component ($IC_{50}=750.8$ μ g/mL) > *A. maculatus* the methanol component ($IC_{50}=1,350.6$ μ g/mL). The inhibition rate of the first five components was higher than the positive control Acarbose ($IC_{50}=1,103.01$ μ g/mL), and had a certain concentration dependence. In addition, 5,7,4'-trihydroxy-6-methoxy flavone ($IC_{50}=40.06$ μ g/mL, **7**) and lupeol ($IC_{50}=25.41$ μ g/mL, **204**), and ursolic acid ($IC_{50}=4.42$ μ g/mL, **205**) had good *in vitro* α -glucosidase inhibitory activity.

Yao [27] used the 96-well plate method to study the effects of phenethyl alcohol and glycosides isolated from *C. flabellatus* on mushroom tyrosinase. Using arbutin as positive control, compound **42** showed certain mushroom tyrosinase inhibitory activity, which was concordant to the activity of this type of compound (containing caffeoyl) in the literature. Compounds **43–46** and **120** had no inhibitory activity and showed agonistic activity, which may be related to the caffeoyl compound structure and may be used in the treatment of white disease or vitiligo.

4.4. Anti-Alzheimer's Activity

Wang et al. [60] explored the intervention mechanism of *C. flabellatus* extract for Alzheimer's disease (AD) through UPLC-Q-TOF/MS metabolomics. Statistical analysis showed that the metabolic profiles of the AD model group were significantly separated from the control group, while the CF group was closer to the control and positive groups. A total of 28 potential AD-related biomarkers were detected, and metabolic pathways were established by MetPA. This study elucidated the involvement of metabolic pathways in oxidation, disorders of energy metabolism, and neuromodulator disorders. In addition, this is the first time that a metabolomics study has been used to determine the therapeutic mechanism of CF extract on AD model mice.

Zeng *et al.* [61] studied the effect and mechanism of action of *C. flabellatus* (SDC) extract and isolated isonuomioside A (isA) on A β_{25-35} -induced brain injury. The experiment was completed by inducing brain injury by infusion of A β_{25-35} (200 μ M, 3 μ l/20 g, i.c.v.) and then dosing with SDC extract (155 mg/kg, i.g.) or isA (20 mg/kg, i.g.) over 4 weeks. In addition, the antagonism of MK-801 (NMDA receptor blocker, 10 μ M) in the presence of isA (10 μ M) or SDC (20 μ g/ml) extracts was studied in A β_{25-35} (20 μ M, 24 h)-induced PC-12 and N9 cells to assess whether the effects induced by isA and SDC extracts were mediated through the NMDAR2B/CamK II/PKG pathway. It was demonstrated that isA and SDC extracts ameliorated A β_{25-35} -induced brain injury by inhibiting apoptosis, oxidative stress, and autophagy, possibly through the NMDAR2B/CamK II/PKG pathway.

Zhang *et al.* [62] investigated the intervention effect and mechanism of carbenoside B (CB) on Alzheimer's disease (AD) induced by amyloid β -protein fragment 25-35 (A β_{25-35}). A mouse model of AD was established by injecting A β_{25-35} into the brain. Y maze and new object recognition experiments were used to determine the learning and memory ability of mice. Pathological changes in the hippocampus were observed by HE staining, Nissner staining, and electron microscopy. The levels of A β_{1-42} /A β_{1-40} and p-Tau in the brain were detected by enzyme-linked immunosorbent assay. The content or activity of MDA, SOD, and GSH-Px in serum was detected using biochemical methods. The level of reactive oxygen species in the brain was detected using flow cytometry. Protein western blotting was used to detect the expression levels of LC $_3$ B, Beclin-1, and P62-related autophagy proteins in the brain. In addition, PC-12 cells were cultured *in vitro* and combined with autophagy inhibitor 3-methyladenine (3-MA) to observe whether the effect of CB on A β_{25-35} -induced PC-12 cells is related to autophagy.

The results showed that CB could significantly improve the learning and memory ability of mice with A β_{25-35} -induced mice, brain injury and improve hippocampus injury and neuronal atrophy. In addition, it can also reduce the levels of A β_{1-42} /A β_{1-40} , p-Tau, and oxidative stress, and enhance the level of autophagy. *In vitro* results showed that CB could significantly improve the migration and proliferation ability of PC-12 cells. In summary, CB can alleviate AD model damage induced by A β_{25-35} by enhancing autophagy.

4.5. Other Activities

Zheng *et al.* [63] used guinea pigs to determine that the water decoction of *L. pauciflorus* had antitussive effect, could increase the secretion of the trachea of mice, and had an expectorant effect. It had a certain protective effect on the asthma-like features caused by histamine inhalation in guinea pigs. The animals were quieter and less active after the medication. The drug had a certain sedative effect on the central nervous system. Wang [31] used preliminary pharmacodynamic experiments to observe the effect of the methanol extract of *C. longgangensis* var. *hongyao* on the writhing response induced by acetic acid in mice. The experimental results show that the writhing times of mice using the methanol extract of *C. longgangensis* var. *hongyao* and the n-butanol part of the methanol extract of *C. longgangensis* var. *hongyao* were (3.4 \pm 4.3) and (4.7 \pm 9.8), respectively, which were significantly different from the model group (17.2 \pm 14.0) (P <0.05). Studies have shown that the methanol extract of *C. longgangensis* var. *hongyao* has a significant inhibitory effect on acetic acid-induced pain in mice.

Han *et al.* [64] found that the antihypertensive effect of lysionotin increased with the increase in plasma drug concentration during intravenous infusion, and decreased with the decrease in plasma drug concentration after entering the elimination phase. The maximum antihypertensive effect occurs 5–10 minutes after the maximum blood concentration is reached. Sun *et al.* [65] studied the effect of lysionotin (**1**) on the hemodynamics of anesthetized thoracotomy cats. Intravenous injection of lysionotin at a dose of 2.5 mg/kg did not result in significant changes in HR, dP/dtmax, t-dP/dtmax, and CO. When the dose was increased to 5 mg/kg, blood pressure and TPVR decreased further, while HR, dP/dtmax, t-dP/dtmax, and CO significantly decreased. The results show that the hypotensive effect of low-dose lysionotin is mainly caused by the relaxation of blood vessels, and the inhibitory effect of larger doses of lysionotin on the heart may also be involved in its antihypertensive effects.

Chen *et al.* [36] used the MTT method to assess the cytotoxic activity of the compounds obtained from the *C. eburnea* fraction. Compounds **55–59** had varying degrees of inhibitory effects on the human lung cancer SPC-A1 cells. Compounds **58** and **59** had weak cytotoxic effects on SPC-A1 cells.

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The half inhibitory concentrations (IC₅₀) were 207.23 µmol/L and 180.34 µmol/L. Compounds **55–57** showed no cytotoxicity (IC₅₀> 300 µmol/L). Compounds **55–59** had no cytotoxicity to liver cancer HepG2 cells, gastric cancer SGC-7901 cells and lung large cell carcinoma NCI-H460 cells, and its IC₅₀ was > 300 µmol/L.

Cao et al. [66] took the senescence-accelerated mouse prone 8 (SAMP8) as the animal model, and gave the mice different doses of the extract of *C. flabellatus* (CF) via gavage for one month. The degree of renal senescence was observed by β-galactosidase staining, and the degree of renal fibrosis was evaluated by Masson staining and the expression levels of Col-I, α-SMA and FN. The protein expression levels of the NRF2 pathway and the Wnt/β-catenin/RAS pathway in renal tissues were detected. In addition, β-galactosidase (β-GAL)-induced NRK-52E cells were used as an *in vitro* model to screen the effective components. The separation of 3,4-dihydroxyphenyl ethanol (SDC-0-14, 16) and 3, 4-dihydroxyphenylethanol-8-O-(4-O-trans-caffeoyl-β-D-apiofuranosyl-(1 → 3)-β-D-glucopyranosyl (1→6)-β-D-pyran glycosidase (SDC-1-8) can delay the senescence of NRK52E cells. It has been shown that CF ethanol extract may reduce renal fibrosis in SAMP8 mice through the Wnt/β-catenin/RAS pathway. Hence, SDC-0-14, SDC-0-16 and SDC-1-8 may be the material basis for their anti-renal senescence effects.

5. Conclusions

Gesneriaceae plant resources are abundant, widely distributed, and have broad applications and development potential. In recent years, researchers have extensively studied Gesneriaceae, yet not all pharmacological properties of this family have been investigated, including mechanisms of action and structure-activity relationships of its active components. New techniques for the isolation, purification, and structural identification of chemical components, pharmacological activities, and mechanisms of action of the compounds or/and extracts of the plants from Gesneriaceae family are thus necessary.

Acknowledgments

This work was supported jointly by the National Basic Research Program of China (2019YFC1708802), National Natural Science Foundation of China (82104386), and Scientific and technological key project in Henan Province (212102310345). We thank LetPub (www.letpub.com) for its linguistic assistance during the preparation of this manuscript.

Supporting Information

Supporting information accompanies this paper on <http://www.acgpubs.org/journal/records-of-natural-products>

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References

- [1] L. Chen (2010). Studies on biological activity of four Gesneriaceae plants, Henan University, 37-41.
- [2] F. Wen, L.F. Fu, Z.B. Xin, C. Xiong and Y.G. Wei (2022). Endangered status and biodiversity conservation of China's Gesneriaceae, *Guihaia* 1-78.

- [3] Y. Liu, H. Wagner and R. Bauer (1996). Nevadensin glycosides from *Lysionotus pauciflorus*, *Phytochemistry* **42** (4), 1203-1205.
- [4] Shanghai Institute of Materia Medica (1976). Chinese Herbal Medicine Newslette. Hunan Institute of Pharmaceutical Industry, Hunan.
- [5] G. Xu, Z.B. Hu, S.C. Feng and G.J. Fan (1979). Studies on the anti-tuberculosis principles from *Lysionotus pauciflora* Maxim, *Acta Pharm. Sin. B.* **7**, 447-448.
- [6] M.Y. Wang, Y.J. Fan, J. Zhang and M.X. Gong (2010). Simultaneous determination of plantainoside D and verbascoside from stem of *Chirita longgangensis* var *hongyao* by RP-HPLC, *Chin. J. Chin. Mat. Med.* **35** (23), 3188-3191.
- [7] Y.Z. Wang, Y.Y. Liu, X.K. Zheng, Y.G. Jia and Z.Y. Feng (2011). A reversed-phase high performance liquid chromatography (RP-HPLC) procedure for determination of four main components in *Boea hygrometrica* (Bunge) R. Brown, *Chin. Pharm. J.* **46** (19), 1517-1520.
- [8] H. Cao, S.Y. Wang and X.Y. Li (2006). Research progress in study of *Lysionotus pauciflorus*, *J. Southeast Guizhou Nat. Teacher's Coll.* **24** (6), 44-45.
- [9] M.Y. Wang, L. Yang and Y.Y. Tu (2006). Studies on the chemical constituents from stem of *Chirita longgangensis* var *hongyao*, *Chin. J. Chin. Mat. Med.* **31** (4), 307-308.
- [10] M.Y. Zeng (1994). Traditional Chinese medicine resource flora. Science and Technology Press, Beijing.
- [11] Z.Y. Li and Y.Z. Wang (2005). Plants of Gesneriaceae in China. Henan Sci-Tec Press, Zhengzhou.
- [12] P.F. Lv, X.K. Zheng and W.S. Feng (2006). Studies on the chemical constituents of ethyl ether and ethyl acetate parts from *Boea hygrometrica* (Bge.)R.Br., In:2006 Annual Symposium of Chinese Medicine Chemistry Branch, Chinese Association of Traditional Chinese Medicine, Beijing, pp.13-15.
- [13] Editorial Board of Chinese Materia Medica State Administration of Traditional Chinese Medicine (1999). Chinese materia medica. Shanghai Science and Technology Press, Shanghai.
- [14] W.C. Wang, K.Y. Pan and Z.Y. Li (1990). Flora reipublicae popularis sinicae. Sciences Press, Beijing.
- [15] National Compilation Group of Chinese Herbal Medicine (1978). National compilation of Chinese herbal medicine. People's Medical Publishing House, Beijing.
- [16] Guangxi Institute of Chinese Medicine and Pharmaceutical Science (1986). List of medicinal plants in guangxi. Guangxi People's Publishing House, Nanning.
- [17] P.Y. Tian (2012). Studies on the chemical and biological activity constituents of four plants, Henan University, 67-68.
- [18] Z.Y. Shen (2018). Chemical constituents of ethyl acetate and n-butanol fraction from *Chirita linearifolia* W.T.Wang, *Guangxi Univ. Chin. Med.* 62-63.
- [19] W.S. Feng, Q. Li and X.K. Zheng (2006). Study on the chemical constituents of diethyl ether and n-butanol fractions of *Lysionotus pauciflorus* Maxim, In:2006 China Association for Science and Technology Annual Conference, Beijing, pp.3359-3361.
- [20] W.S. Feng, Q. Li, X.K. Zheng and X.H. Kuang (2006). Chemical constituents from *Lysionotus pauciflorus*, *Nat. Prod. Res. Dev.* **18**, 617-620.
- [21] J.F. Wei, L. Chen, J.M. Wang and W.Y. Kang (2011). Anti-bacteria activity of *Puraboea rutescens* and *Lysionotus pauciflorus*, *Chin. J. Chin. Mat. Med.* **36** (14), 1975-1978.
- [22] P.Y. Tian, J.M. Wang and W.Y. Kang (2011). Chemical constituents of *Aeschynanthus superbus* Clarke, *Chin. Pharm. J.* **46** (23), 1795-1797.
- [23] L.D. Zhou, J.G. Yu, J. Guo and S.L. Yang (2001). Compounds from roots of *Chirita fimbrisejala* Hand.-Mazz, *Chin. J. Chin. Mat. Med.* **26** (2), 114-117.
- [24] W.S. Feng, X.K. Zheng, Y.B. Liu and J. Li (2004). Isolation and structural identification of C-glycosylflavones from *Corallodiscus flabellata*, *Acta Pharm. Sin.* **39** (2), 110-115.
- [25] X.K. Zheng, P.F. Lv and W.S. Feng (2005). Studies on the chemical constituents of *Boea hygrometrica* (Bge) R.Br, In:Proceedings of the 8th National Symposium on Traditional Chinese Medicine and Natural Medicine and the 5th National Symposium on Medicinal Plants and Plant Pharmacy, Wuhan, p.100.
- [26] M.Y. Wang (2005). Preliminary study on chemical constituents and biological activity of *Chirita longgangensis* W. T. Wang var. *hongyao* S. Z. Huang, *China Acad. of TCM*, 30-32.
- [27] Y. Yao (2012). Study on chemical constituents of *Corallodiscus flabellatus*, Kunming University of Science and Technology, 15-43.
- [28] H.X. Liu, H.B. Liao and K. Yuan (2012). Chemical constituents contained in *Aeschynanthus moningeriae*, *Chin. J. Chin. Mat. Med.* **37** (13), 1963-1967.
- [29] S.M. Li, X.W. Yang, Y.H. Shen, L. Feng, Y.H. Wang, H.W. Zeng, X.H. Liu, J.M. Tian, Y.N. Shi, C.L. Long and W.D. Zhang (2008). Chemical constituents of *Aeschynanthus bracteatus* and their weak anti-

Chemical components and pharmacological activities of Gesneriaceae

- inflammatory activities, *Phytochemistry* **69**, 2200-2204.
- [30] J. Hu, Q. Zhang, M.D. Qi, L.P. Kang, T.G. Nan, J. Yang, Y. Yuan, Z.L. Zhan, Y. Liu and L.Q. Huang (2016). Study on chemical constituents in *Lysionotus wilsonii* by UPLC-Q-TOF-MS, *Chin. J. Chin. Mat. Med.* **41** (9) 1658-1669.
- [31] M.Y. Wang, L. Yang and Y.Y. Tu (2005). Phenylethanoid glycosides from stem of *Chirita longgangensis* var *hongyao*, *Chin. J. Chin. Mat. Med.* **30** (24), 1921-1923.
- [32] W.J. Chen, Y.X. Wen, Y.Y. Chen, Y.L. Huang and D.P. Li (2010). Phenylethanoid glycosides and their cytotoxicity of *Chirita eburnea* Hance, *Chin. Tradit. Pat. Med.* **32** (6), 1000-1003.
- [33] J.J. Wang (2017). Chemical constituents of petroleum ether fraction from *Chirita linearifolia* W. T. Wang, *Guangxi Univ. Chin. Medicin.* 63-64
- [34] Y.F. Bi (2002). Research on the chemical constituents of *Corallo-discus flabellate*, *Selaginella tarmariscina* (Beauv.) Spring and pharmacological activity of *Selaginella tarmariscina* (Beauv.) Spring, Beijing University of Chinese Medicine, 9-25.
- [35] X.Q. Wang, L. Li, Z.F. Bai, Y. Peng, P.G. Xiao and Y. Liu (2011). Five new phenylpropanoid glycosides from *Paraboea glutinosa* (Gesneriaceae), *J. Nat. Med.* **65**, 301-306.
- [36] W.J. Chen (2010). Studies on the chemical constituents and bioactivity of *Chirita eburnea* Hance of Gesneriaceae, Guangxi Normal University, 56-58.
- [37] B.L. Qiu, L.X. Xu, X.Y. Wei, M. Kang and L.D. Lin (2014). Anthraquinones from *Hemiboea subcapitata*, *J. Trop. Subtrop. Bot.* **22** (5), 507-510.
- [38] C.Y. Chen, Y. Lu and Z.N. Chen (2001). Studies on the chemical constituents of anthraquinone of *Rhynchotechum Vestitum*, *Acta Univ. Med. Second Shanghai.* **21** (6), 488-491.
- [39] K.A. Sales, E.F.d. Silva, P.T.R.d. Figueiredo, V.C.d.O. Costa, M.T. Scotti, M.d.F. Agra, J.F. Tavares and M.S.d. Silva (2018). Chemical constituents from *Paliavana tenuiflora* Mansf. (Gesneriaceae), *Biochem. Syst. Ecol.* **80**, 76-80.
- [40] M.H. Verdan, V. Winiewski, D.R. Scharf, M.J.e. Salvador, E.L.d. Sa and M.E.A. Stefanello (2021). Dimeric naphthochromenes and other constituents from *Sinningia allagophylla* (Gesneriaceae), *Phytochem. Lett.* **42**, 104-108.
- [41] Q.F. Wen, Y.J. Zhong, X.H. Su, C.Y. Li, J. Liu, Y.F. Li and B. Liang (2013). A new acorane sesquiterpene from *Lysionotus pauciflorus*, *Chin. J. Nat. Med.* **11** (2), 185-187.
- [42] S.Y. Zhu, J. Hou, X.Z. Zheng, H.D. Yu and G.L. Zhou (2004). Chemical composition of essential oil of *Oreocharis acericula*, *Amino Acid. Biotic. Resources.* **26** (4), 8-9.
- [43] L. Chen and W.Y. Kang (2009). Chemical constituents from *Aeschynanthus longicaullis*, *Chin. J. Chin. Mat. Med.* **34** (21), 2758-2760.
- [44] H.J. Huang and L.Y. He (2014). Chemical constituents from roots of *Chirita longgangensis* var. *hongyao*, *Chin. J. Chin. Mat. Med.* **39** (6), 1040-1042.
- [45] Q. Wang and F.H. Xu (2013). Chemical constituents of ethyl acetate fraction from *Aeschynanthus bracteatus*, *J. Chin. Med. Mat.* **36** (3), 414-415.
- [46] X.Y. Zang (2008). Chemical research on *Aeschynanthus mengxingensis* and *Corallo-discus kingianus*, Henan University, 90.
- [47] W.Y. Kang, L. Chen and X.Y. Zang (2009). Triterpenes from *Corallo-discus kingianus*, *Chin. J. Chin. Mat. Med.* **34** (20), 2607-2609.
- [48] W.Y. Kang, Z.Q. Ji and J.M. Wang (2009). Research on fatty acids in *Corallo-discus kingianus* and *Aeschynanthus mengxingensis*, *Nat. Prod. Res. Dev.* **21** (2), 203-207.
- [49] P.Y. Tian, W. Wang, J.M. Wang and W.Y. Kang (2013). Chemical constituents from *Rhynchotechum ellipticum*, *Nat. Prod. Res. Dev.* **25** (7), 919-921.
- [50] G.H. Wang, W.F. Huang and K. Yuan (2010). Analysis of chemical constituents of volatile oil in *Aeschynanthus moningeriae* (Merr.) Chun by SPME-GC/MS, *Chem. World* **51** (12), 725-728.
- [51] F.M. Yang, X.S. Yang, B. Luo and H.Y. Zhu (2003). Studies on the chemical constituents of *Lysionotus pauciflorus* Maxim, *Nat. Prod. Res. Dev.* **15** (6), 508-509.
- [52] W.Y. Kang, X.Y. Zang, J.M. Wang and Q.T. Xu (2008). Terpenoids from leaves of *Aeschynanthus mengxinensis*, *Chin. J. Chin. Mat. Med.* **33** (17), 2118-2121.
- [53] X.Q. Wang, Y. Peng, L.J. Xu, W. Xiao, P.G. Xiao and Y. Liu (2009). Triterpenes and triterpene glycosides from aerial part of *Paraboea glutinosa*, *Chin. J. Chin. Mat. Med.* **34** (10), 1228-1230.
- [54] W.S. Feng, Q. Li and X.K. Zheng (2007). Studies on chemical constituents of *Lysionotus pauciflorus*, *Chin. Pharm. J.* **42** (5), 337-338.
- [55] L. Qiao, L. Li, M.J. Song, Y.M. Hou and X.J. Li (2020). A GC-MS analysis of liposoluble components of herb of fan-shaped corallo-discus and a study of the antioxidative activity, *Chin. Med. Pharm.* **10** (17), 78-81.
- [56] W.Y. Kang, C.F. Li and L. Zhang (2009). Antioxidant activity of *Corallo-discus kingianus* and

- Aeschynanthus mengxingensis*, *Nat. Prod. Res. Dev.* **21** (3), 470-472.
- [57] H.X. Liu (2012). Studies on chemical constituents of *Aeschynanthus moningeriae* and antioxidant activity, Zhejiang Normal University, 43-44.
- [58] Q. Zhang, H.P. Xiao and J.R. Shao (2007). Clinical observation of shidiaolan tablets in adjuvant treatment of retreatment of pulmonary tuberculosis, *Chin. J. Antitubercul.* **6**, 541-542.
- [59] P.Y. Tian and W.Y. Kang (2012). α -glucosidase inhibitory activity of *Aeschynanthus maculatus*, *Chin. J. Chin. Mat. Med.* **37** (19), 2910-2912.
- [60] Y.Y. Wang, N. Zhou, Y.P. Si, Z.Y. Bai, M. Li, W.S. Feng and X.K. Zheng (2021). A UPLC-Q-TOF/MS-based metabolomics study on the effect of *Corallodiscus flabellatus* (Craib) B. L. Burt extract on alzheimer's disease, *Evid. Based Complement. Alternat. Med.* **2021**, doi: 10.1155/2021/8868690.
- [61] M.N. Zeng, A.Z. Feng, M. Li, M. Liu, P.L. Guo, Y.H. Zhang, Q.Q. Zhang, B.B. Zhang, B. Cao, J.F. Jia, R. Wang, J. Lyu and X.K. Zheng (2022). *Corallodiscus flabellata* B. L. Burt extract and isonuoside A ameliorate A β (25-35)-induced brain injury by inhibiting apoptosis, oxidative stress, and autophagy via the NMDAR2B/CamKII/PKG pathway, *Phytomedicine* **101**, doi: 10.1016/j.phymed.2022.154114.
- [62] Y.H. Zhang, M. Li, M.N. Zeng, P.L. Guo, M. Liu, J.B. Liang, B. Cao, X.K. Zheng and W.S. Feng (2022). Carbon glycosides of *Corallodiscus flabellata* B. L. Burt ameliorate the alzheimer's disease in mice, *Chin. J. New Drugs.* **31** (17), 1718-1726.
- [63] X.K. Zheng, J. Li, W.S. Feng and Y.B. Liu (2003). Advances in study of Gesneriaceae, *Chin. J. New Drugs.* **12** (4), 261-263.
- [64] G.Z. Han, C.Y. Su and Y. Zhang (1982). The absorption distribution and elimination of nevadensin in the rat and the relationship between plasma concentration of the drug and its hypotensive effect, *Acta Pharm. Sin.* **17** (8), 572-578.
- [65] A.S. Sun, J.S. Shi, Q. Wu and Z.M. Li (1996). Effects of nevadensin on hemodynamics in anesthetized cats, *J. Zunyi Med. Univ.* **19** (1), 5-7.
- [66] B. Cao, M.N. Zeng, Y.P. Si, B.B. Zhang, Y.Y. Wang, R.Q. Xu, Y.J. Huang, W.S. Feng and X.K. Zheng (2022). Extract of *Corallodiscus flabellata* attenuates renal fibrosis in SAMP8 mice via the Wnt/ β -catenin/RAS signaling pathway, *BMC Complement. Med. Ther.* **22** (1), 52.

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