

Review

# The Chemistry and Pharmacology of Fungal Genus *Periconia*: A Review

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**Abstract:** *Periconia* is filamentous fungi belonging to the Periconiaceae family, and over the last 50 years, the genus has shown interest in natural product exploration for pharmacological purposes. Therefore, this study aims to analyze the different species of *Periconia* containing natural products such as terpenoids, polyketides, cytochalasan, macroshelides, cyclopentenes, aromatic compounds, and carbohydrates carbasugar derivates. The isolated compound of this kind, which was reported in 1969, consisted of polyketide derivatives and their structures and was determined by chemical reaction and spectroscopic methods. After some years, 77 compounds isolated from endophytic fungus *Periconia* were associated with eight plant species, 28 compounds from sea hare *Aplysia kurodai*, and ten from endolichenic fungi *Parmelia* sp. The potent pharmacological agents from this genus are periconicin A, which acts as an antimicrobial, pericochlorosin B as an anti-human immunodeficiency virus (HIV), peribysin D, and pericosine A as cytotoxic agents, and periconianone A as an anti-inflammatory agent. Furthermore, information about taxol and piperine from *Periconia* producing species was also provided. Therefore, this study supports discovering new drugs produced by the *Periconia* species and compares them for future drug development.



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## 1. Introduction

*Periconia* is filamentous fungi belonging to the Periconiaceae Family (Ascomycetes), and according to Index Fungorum [1], there are 202 epithets with 12 varieties. However, only 40 species of the *Periconia* genus are currently recognized [2–4]. This genus has spread as endophytes and saprobes in plants with multiple habitats and has been shown to be plant pathogens. Leukel [5] reported that Milo disease, found in crops, can be caused by one species of *Periconia*, *P. circinata* (*L. mangin*). Interestingly, these fungi can live in the gastrointestinal tract of animals [6] and as an endolichenic fungus [7–9].

Over the past 50 years, *Periconia* has become an exciting resource for natural product research because it produces many bioactive secondary metabolites. This genus is reported to have tremendous activities as antimicrobial, anti-human immunodeficiency virus (HIV), potent cytotoxicity, and anti-inflammatory agents from many groups of secondary metabolites, which include terpenoid, meroterpene, polyketide, cytochalasan, macrolide, macroshelide, aromatic compound, and carbohydrate derivates [6–14].

The study has been reported in at least two main areas: (1) chemical and bioactivities studies of the isolated secondary metabolites from different species of fungi and various host origins. The species produce secondary metabolites with multiple biosynthesis pathways. (2) Molecular and genomic studies for species identification and revised genus annotation. A comprehensive review of the *Periconia* fungal genus for chemical content and its pharmacological aspects between these studies has not been conducted.

Since 1969, more than a hundred bioactive secondary metabolites of the genus *Periconia* have been discovered. Interestingly, study communities have gained more attention for *Periconia* since the fungus reportedly produces Taxol (anticancer agent) [15] and also Piperine (antimycobacterial agent) [12]. For this significant interest, the potential of *Periconia* species producing secondary metabolites was explored. This highlights chemical contents from different strains classified by the natural products group and with particular emphasis on their pharmacological activities. Furthermore, the parts and origin, function, and biological roles were detailed. All databases with the keyword “periconia” from search engines Scopus, Scifinder, PubMed, and Google Scholar were obtained from May 1969 to October 2020. This study should be helpful in the discovery of new drugs in the future.

## 2. Phylogeny of the Genera *Periconia*

The genus *Periconia* was first introduced in 1791 by H. J. Tode, namely *P. lichenoides* Tode ex Mérat [2]. *Periconia* is classified as a member of the Family Periconiaceae and separated from Massariaceae as a sister taxon because of distinct phylogeny [16]. In addition, the phylogenetic tree is Kingdom Fungi, Phylum Ascomycota, Subphylum Pezizomycotina, Class Dothideomycetes, Subclass Pleosporomycetidae, Order Pleosporales, Family Periconiaceae, and Genus *Periconia* [1,16–18].

*Periconia* has macronematous conidiophores with pale to dark brown, branched or unbranched, smooth or verruculose. Conidia cells are spherical, aseptate, catenate or solitary, smooth or verruculose, pale to dark brown. The conidiogenous cells are monoblastic or polyblastic, integrated or discrete, ovoid to clavate formed on the terminal or intercalary of the stipe [2]. This genus is registered on Index Fungorum and Mycobank with Registration Identifier (ID): 9263 and The National Center for Biotechnology Information (NCBI) taxonomy browser with ID: 97971 [1,16–18].

## 3. Natural Product Diversity

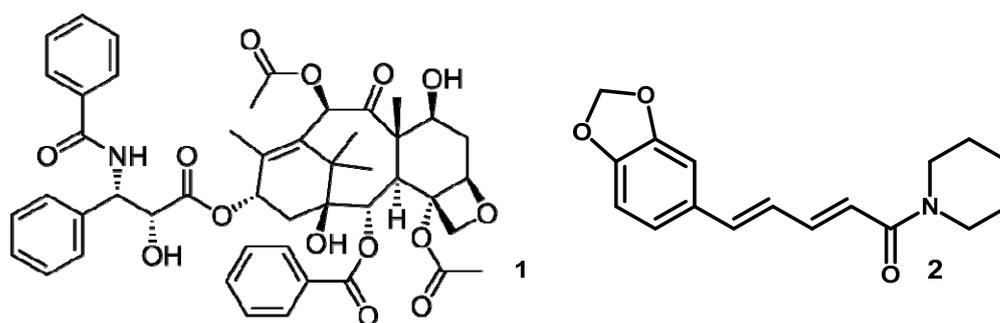
### 3.1. Overview of Compounds Derived from the *Periconia* Fungus

From 1969 to 2020, a total of 104 compounds were isolated from *Periconia*. Terpenes are the major classes of naturally occurring secondary metabolites from the *Periconia* genus comprising a total of 43 terpenoids (41.3%). Furthermore, there are 40 polyketides (38.4%): 11 cytochalasans, 1 macrolide, 10 macrophelides, 4 cyclopentes, and 14 other polyketides. There are 14 aromatic compounds (13.4%) and 5 carbohydrates (4.8%) which were isolated from the *Periconia* genus. The other compounds, taxol and piperine, which its plant host originally produced, this fungus also reportedly made.

*Periconia* compound endophytic fungi in eight plant species: *Annona muricata* (42 compounds), *Taxus cuspidata* (2 compounds), *Xylopi aromaticum* (2 compounds), *Thysanoleana latifolia* (2 compounds), *Sorghum bicolor* (5 compounds), *Rosa chinensis* (1 compound), *Piper longum* (1 compound), and terrestrial herbaceous plants (4 compounds). Endosymbionts gastrointestinal sea hare *A. kurodai* (28 compounds), and endolichenic fungi *Parmelia* sp (10 compounds) also form part of *Periconia*. These compounds are summarized in Table 1 by type of compounds, molecular weight (MW) and mass spectrometry (MS) data, strain type, various host and origins.

### 3.2. Taxol and Piperine Producing *Periconia* Species

Taxol or paclitaxel (1) are well-known anticancer drugs with a tetracyclic diterpenoid structure (Figure 1). Initially, its small quantity was obtained from ethanol extract of the bark of *Taxus brevifolia* [19]. The amount was only 0.01–0.03% dry weight and this is a significant factor contributing to its high price. This problem is emerging in the scientific community to investigate another source of the compound.



**Figure 1.** The chemical structure of taxol (1) and piperine (2) from the *Periconia* genus.

The discoveries continued with this development, and Stierle et al. [20] found that the fungal endophytes of *Taxomyces andreanae* in the Pacific region can produce Taxol like the host. On the contrary, Li et al. [15] studied fungal endophytes of *Torreya grandifolia*, when taxol is not synthesized.

Furthermore, *Periconia* from this plant was demonstrated unequivocally for taxol production by a semi-synthetic medium. The fungi were fermented with a special medium through an activator of metabolism. Several compounds studied include serinol, *p*-hydroxybenzoic acid, and a mixture of phenolic acids. Benzoic acid, known as not a taxol precursor, activates taxol production metabolism in *Periconia*. This was conducted at 0.01 mM with the best yield concentration  $831 \pm 27 \text{ ng L}^{-1}$  through spectroscopic and immunological methods.

According to Verma et al. [12], *Periconia* can also produce compounds similar to its host. For example, Piperine (2), an alkaloid (Figure 1) with many pharmacological activities obtained from the ethanol extract of fruits *Piper longum* (only consist 3–5% dry weight basis), also being produced by *Periconia*. The 2 were obtained from ethyl acetate extract after four weeks of fermentation on a potato dextrose broth. The isolated 2 then crystallized, elucidated by single-crystal X-ray crystallography, and studied extensively as an antimycobacterial agent. These piperidine derivatives (2) were treated for multidrug-resistant *Mycobacterium tuberculosis* and an atypical mycobacterium *M. smegmetis* with a Minimum Inhibitory Concentration (MIC) 1.74 and 2.62  $\mu\text{g/mL}$  (strong activity), respectively, (control positive or negative are not included). In this context, the scientific community increased interest in natural product exploration from endophytic fungi and other fungal sources, especially for the *Periconia* genus.

### 3.3. Terpenoids

Some species of *Periconia* appear to produce terpenoids and have engaging pharmacological activities. According to Bach and Rohmer [21], the starting unit of terpenoid is from the five-carbon precursor unit, isopentenyl diphosphate (IPP), and dimethylallyl diphosphate (DMAPP). This is continued with a head-to-tail condensation repetitive reaction of IPP and DMAPP with prenyltransferases. These produce prenyl diphosphates of increasing length: geranyl-, farnesyl-, geranylgeranyl-diphosphate that lead to synthesizing the different isoprenoids and products. There are thirty-nine terpenoid compounds isolated from this genus. This includes six monoterpenoids (carene and menthene-types), two diterpenoids (fusicoccane-type), twenty-five sesquiterpenoids (cadinane, and eremophilane-types), one norsesquiterpenoid, one meroterpene (diterpenoid alkaloid), and one steroid.

Six monoterpenoids were obtained from the *Periconia* sp. F-31, and they live inside the leaves of *A. muricata* as an endophyte. Furthermore, broth extract of this fungus led to isolation of three new 2-carene derivatives: 2-carene-5,8-diol (3), 2-carene-8,10-diol (4), 2-carene-8-acetamide (5), one new menthene-type: 8-hydroxy-1,7-epoxy-2-menthene (6). Other derivatives include one known monoterpenes, anethofuran (7) [22], and a new carene-type: (-)-(1*R*,4*R*,6*S*,7*S*)-2-carene-4,8-olide (8) [23].

Carene-types monoterpenoid (3) was determined using a combination of 1D, 2D Nuclear Magnetic Resonance (NMR) with  $^1\text{H}$ - $^1\text{H}$  Homonuclear Correlation Spectroscopy

(COSY), Heteronuclear Multiple Bond Correlation (HMBC), and Nuclear Overhauser Effect Spectroscopy (NOESY). Additionally, the absolute configuration was calculated with the Time Dependent-Discrete Fourier Transform (TD-DFT) method at the B3LYP/6–31G(d) level. There are two stereoisomers of relative configuration (1*R*,5*R*,6*S*,7*S*) and (1*S*,5*S*,6*R*,7*R*), which were determined using experimental electronic circular dichroism (ECD). Another two carene-type monoterpenoids (4–5) were determined by 1D NMR spectra. This was because of the nearly identical structure feature of 3, confirmed with HMBC correlation, and The Nuclear Overhauser Effect Difference (NOEDIFF) experiments. Meanwhile, 6 was determined by menthene-type monoterpenoids using the same method. The configuration of 8 was assigned by the NOE difference and circular dichroism (CD) spectroscopic data. CD spectrum confirmed exhibited positive and negative Cotton effect (203 nm) and (228 nm) supporting the 1*R*,4*R*,6*S*,7*S* configuration.

These five monoterpenoids (3–6, 8), biogenetically share usual intermediate cation (methyl/ $\alpha$ -terpinyl), then cyclization by monoterpene synthases (non specific) to produce dissimilar structural types (carene and menthene) [22]. Furthermore, 2-carene was transformed to 8 through a series of oxidation and lactonization [23]. Monoterpenoids play an essential role in plants' resistance to high temperatures and increase heat stress (photosynthetic device) [24].

Diterpenoid from *Periconia* was reported in 2004, and it isolated periconicins A (9) and B (10) from *Periconia* sp. This was obtained from the small inner branches of *T. cuspidata* Kangwon Region of Korea [25]. The chemical structure of both compounds was determined by 1D, 2D NMR (<sup>1</sup>H-<sup>1</sup>H COSY, Heteronuclear Multiple Quantum Coherence HMQC, HMBC, and NOESY), Infra-Red (IR) Spectra and High-Resolution Electron Ionization Mass Spectrometry (HREIMS) techniques. In addition, Periconicin B (10) was also isolated from *P. atropurpurea*, the endophytic fungi from *X. aromatica* leaves [11]. The structure of both compounds has the same carbon structure as the fusicoccins [25]. These fusicoccins are the first member of a group called fusicoccanes with a dicyclopenta[a,d]cyclooctane ring system and isolated from the phytopathogenic fungi *Fusicoccum amygdali* [26]. The compounds are reportedly found in fungi, insects, and plants. Fusicoccanes are a plant growth regulator for the elongation mechanism, promoting the opening of the leaf stomata, root inducing formation, and fast seed germination [27].

Further studies should be conducted with *Periconia* to produce sesquiterpenes, a 15-carbon-length natural product that is synthesized by sesquiterpene synthase from farnesyl pyrophosphate [27]. The reaction continues with the cyclic carbocation undergoing further cyclization and rearrangement until the active site is extinguished [28,29]. In addition, eremophilane and cadinane-types are isolated from strains of *Periconia*. Eremophilane-types possessed contradiction with isoprene rules head-to-tail linkages with a biological role for phytotoxic, phytohormonal and mycotoxic for protection [30].

Yamada and his co-worker isolated eremophilane terpenes called peribysin from the strain *P. byssoides* OUPS-N133, which lives in gastrointestinal sea hare *A. kurodai*. This strain was fermented in 90 L medium of malt extract 1%, glucose 1%, and peptone 0.5% with artificial seawater with pH adjusted to 7.5 in 4 weeks. Peribysins A–J (11–20) were isolated from ethyl acetate extract of culture filtrate [13,31–33].

The peribysins were examined extensively with NMR technique such as COSY, HMBC, Distortionless Enhancement by Polarization Transfer (DEPT), and NOESY and mass spectrum for relative configuration. The absolute stereochemistry of these compounds was studied years later. In 2020, Athawale et al., conducted a stereochemical revision of peribysins A (11), B (12), C (13), F (16), and G (17), guided by enantiospecific total synthesis starting from (+)-nootkatone. The synthesis highlight of enone transposition and kinetic iodination results in diastereomers separation [34]. The structure of peribysin D (14) was revised to 6*b*,13-epoxy-7(11)-eremophilene-8*a*,12-diol (14) with the Canonical-representation of Stereochemistry (CAST)/CNMR system, and NOE data [35]. Meanwhile, the absolute configuration of peribysin E (15) was revised by Angeles et al., [36]. This was conducted with natural and non-natural enantiomers obtained from (*R*)- or (*S*)-carvone, respectively,

and was supported with reinvestigation by Inose et al. [37] through time-dependent DFT-based ECD spectral calculations. Peribysins H (18), I (19), J (20) have an absolute stereo structure base upon spectroscopic analyses using 1D and 2D NMR techniques. This was conducted through some chemical transformations, including the modified Mosher's method [32,33].

Further isolation of peribysins from *Periconia* was conducted by Inose et al., [37], where it was shown that peribysins O–Q (21–23) were obtained from the *P. macropinosa* terrestrial herbaceous plant Kanagawa in 2018. The relative structures elucidate with a quantitative NOE experiment. The absolute structures of 21 and 22 were obtained with the modified Mosher's method and 23 with the ECD exciton coupling theory. Additionally, theoretical ECD calculations were conducted for 21–23 [37]. Reconfirmation of peribysin Q absolute structures were performed by Athawale et al., [34] through enantiospecific total synthesis from (+)-nootkatone.

Another eremophilane sesquiterpene, namely periconianones and endophytic fungi, is isolated from *Periconia* sp. F-31 and leaves of *A. muricata*. Zhang et al. [38] isolated periconianone A (24) and B (25), which are polyoxygenated sesquiterpenoid with a new 6/6/6 tricyclic structure. Both structures and absolute configurations were elucidated with extensive spectroscopic analyses, calculated ECD, and single-crystal X-ray diffraction (Cu K $\alpha$ ). Two years later, Liu et al. [39] isolated periconianones C–K (26–34), where C reportedly contained a linkage of -8/C-11 and a 7,12-epoxy moiety. All compounds were elucidated through extensive spectroscopic data analysis, electronic circular dichroism, Mo<sub>2</sub>(AcO)<sub>4</sub>-induced circular dichroism, and single-crystal X-ray diffraction (Cu sK $\alpha$ ). From the same fungus, Ge et al., [23] isolated (+)-(3*S*,6*S*,7*R*,8*S*)-periconone A (35), and its structure was determined based on extensive spectroscopic data and circular dichroism (CD) exciton chirality for absolute configuration. Furthermore, norsesquiterpenoid, dihydronaphthalene-2,6-dione (36) were isolated from the endophytic fungus *Botryosphaeria rhodina* [38].

Cadinane-type sesquiterpenoid from *Periconia* that was firstly reported by Wu et al., [8] is pericoterpenoid A (37), and is an isolate of *Periconia* sp. (No. 19-4-2-1) obtained from a lichen *Parmelia* sp. The relative structure determines by the spectroscopic data (IR, MS, 1D- and 2D-NMR) is known as 14-hydroxycadalen-15-oic acid (24). Cadinane-types sesquiterpenes have 1,7-dimethyl-4-isopropyldecaline structure, presume has biogenesis cyclization pathway of bisabolane or germacrane cation or a 2-*cis*-6-*cis*-farnesyl precursor [40]. This type of compound has a biological role in defence responses after being attacked by herbivores, fungal and bacterial infections [41]. For this exciting defence mechanism, cadinane-type sesquiterpene was developed for fungicides in the future [8].

Ergosterol such as periconiastone A (38) was also obtained from *Periconia* sp. which was collected from the leaves of *R. chinensis* [42]. This steroid compound had an unprecedented pentacyclo-[8.7.0.01,5.02,14.010,15]-heptadecane system. With the ECD spectrum, it allows an explicit assignment of the absolute configuration as 4*R*,5*R*,8*R*,9*R*,10*R*,13*R*,14*S*,17*R*,20*R*,24*R*. From the biosynthetic study, this compound showed a vinylogous  $\alpha$ -ketol rearrangement and an aldol condensation reaction as crucial steps for its formation [42].

Several compounds from *Periconia* have hybrid structural molecules as meroterpenes. This chemical investigation showed a class of diterpenoid alkaloids, where pericolactines A–C (36–38), the first nitrogen-containing fusicoccane diterpenoids are isolated from *Periconia* sp. (No. 19-4-2-1) endolichenic fungus *Parmelia* sp. Furthermore, the structures of 39–41 and their absolute configurations were determined by spectroscopic analyses and quantum chemical ECD calculation. These compounds represent an unusual 5-5-8-5 tetracyclic ring system. The systems were derived from geranylgeranyl diphosphate (GGDP) and serine conjugated biosynthesis. They belong to the atypical diterpenoid alkaloids [7]. Meanwhile, another hybrid molecule meroterpenes namely periconones B–E (42–45) were isolated from EtOAc extract *Periconia* sp. F-31 through semi-preparative HPLC. Their structures and absolute configurations were established by extensive spectroscopic data analysis and electronic circular dichroism (ECD). It was suggested that these hybrids are derived

from typical polyketide-terpenoid pathways formed from one acetyl-CoA starter and five malonyl-CoA extenders coupled with one C5 unit [43]. A summary of the number and types of terpenoids obtained is demonstrated in Table 1. Meanwhile, Figures 2 and 3 showed the structure of the terpenoids compounds.

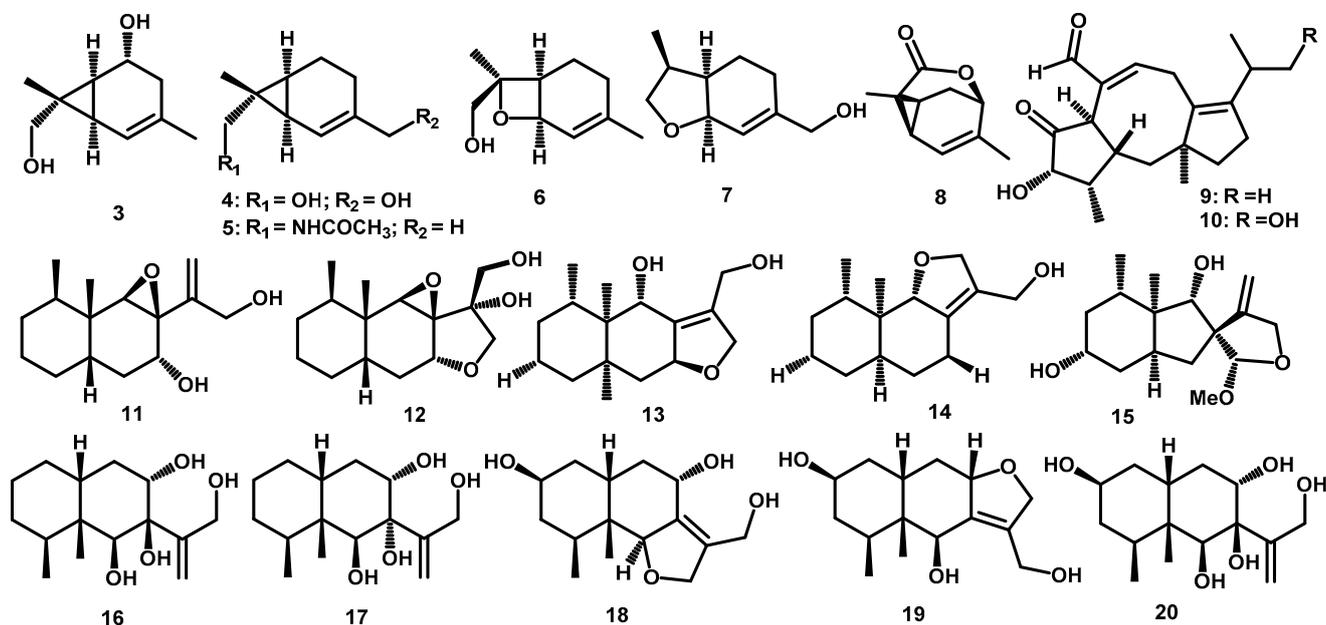
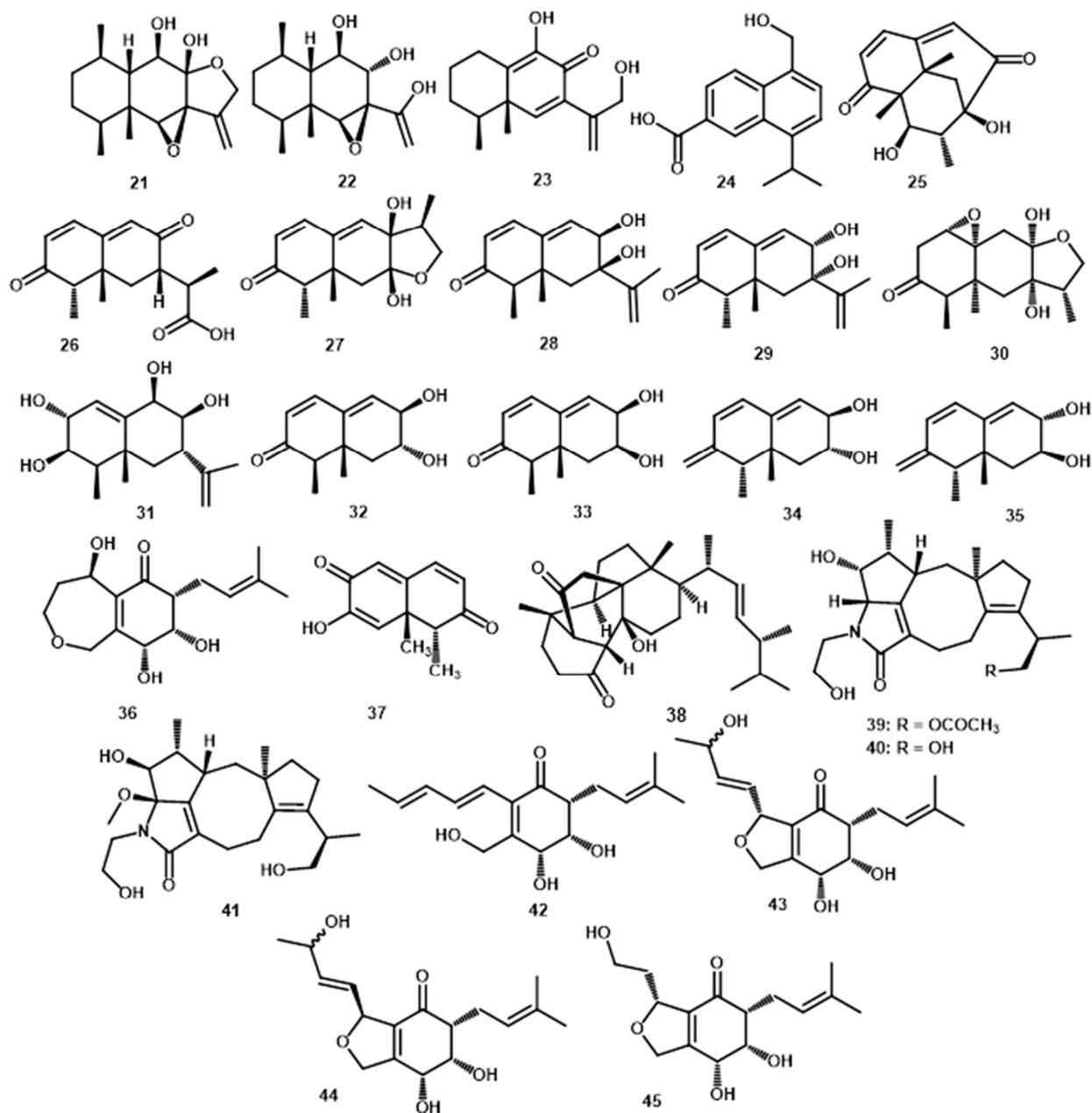


Figure 2. The chemical structures of terpenoids 3–20 from the *Periconia* genus.

### 3.4. Polyketides

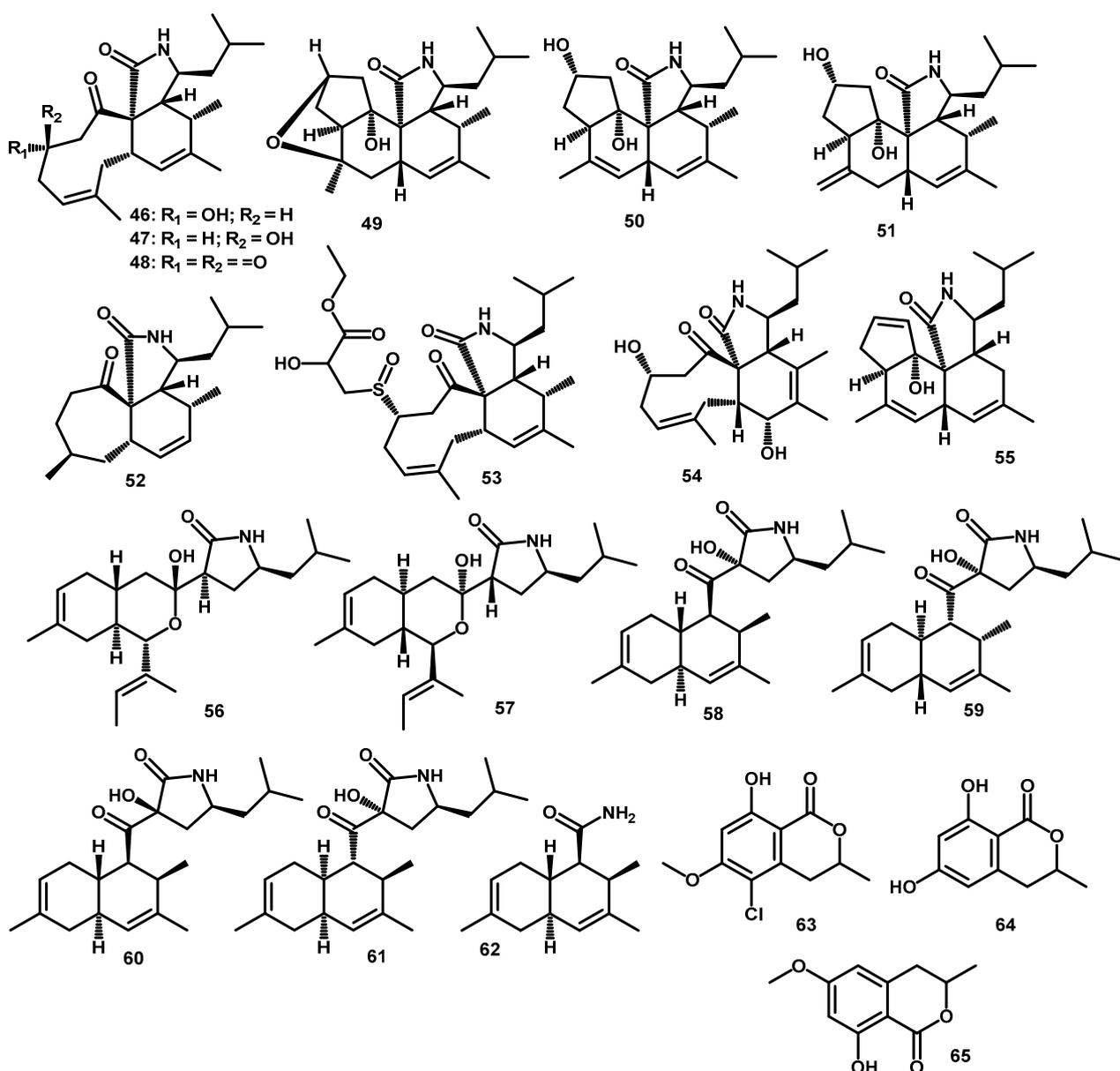
Polyketides are a secondary metabolites group with a large diverse structure derived from the poly- $\beta$ -keto chain [44,45]. They are biosynthesized from acyl-CoA thioesters and undergo repetitive Claisen condensation reactions catalyzed by polyketide synthases (PKSs), which are homologous to fatty acid synthases (FASs) [46]. These compounds are usually used as pharmaceutical agents due to their antibacterial, antifungal, immunosuppressive and antitumor properties [44]. Moreover, polyketides can be produced from diverse microorganisms such as *Periconia*. Twenty twenty-one polyketide compounds were isolated from *Periconia*, including ten cytochalasans and eleven other polyketide derivatives.

Polyketides from *Periconia* genus are isolated from the same natural source known as *A. muricata*. The first were periconiasin compounds, which as a cytochalasan group play a role in fungi and plants to compete and act as an inhibitor of spore germination, hyphal growth, and sporulation of *B. cinerea* [47]. These compounds were also known as a group of polyketide synthase-nonribosomal peptide synthetase (PKS-NRPS) hybrid metabolites. Periconiasins A–C (46–48) (Figure 4) were first isolated with the 9/6/5 tricyclic ring system [48], and their structures were determined using  $^1\text{H-NMR}$ ,  $^{13}\text{C-NMR}$ , DEPT, HMBC,  $^1\text{H-}^1\text{H COSY}$ , HSQC, and IR spectroscopic data. Furthermore, the absolute configuration of 46 was determined using single-crystal X-ray diffraction data, which yielded 3*S*, 4*R*, 5*S*, 8*S*, 9*S*, and 17*S* configurations. In addition, the circular dichroism (CD) spectrum confirmed the negative Cotton effect at 301.5 nm for 39, 296 nm for 47, and 294 nm for 48, where the  $n \rightarrow \pi^*$  transition indicates the 9*S* configuration for 46–48. A comprehensible explanation for the unprecedented ring system was supported by the proposed biosynthesis, which occurs from the attachment of an unusual seven acetate/malonate polyketide backbone to one leucine moiety. This was conducted by PKS-NRPS, followed by Diels–Alder and other reactions [48].



**Figure 3.** The chemical structures of terpenoids 21–45 from the *Periconia* genus.

Further periconiasin compounds sharing the same 9/6/5 tricyclic ring system were obtained, and to this context, periconiasins D–F (49–51) was isolated [49]. Their structures were elucidated using <sup>1</sup>H-NMR, <sup>13</sup>C-NMR, DEPT, HMBC, <sup>1</sup>H-<sup>1</sup>H COSY, and HSQC spectroscopic data. The absolute configuration of 49 was established by ECD calculations and single-crystal X-ray diffraction analyses assigned as 3*S*,4*R*,5*S*,8*S*,9*S*,14*R*,15*S*,17*R*,19*S* configuration (50). Meanwhile, periconiasins G–J (52–55) were separated in 2016 [14,50,51]. Their structures were determined using several spectroscopic data analyses, including IR spectra, <sup>1</sup>H-NMR, <sup>13</sup>C-NMR, DEPT spectra, <sup>1</sup>H-<sup>1</sup>H COSY, HMBC, and NOESY. The structure of 52 possesses a rare sulfoxide group based on strong absorption at 1028 cm<sup>-1</sup> in its IR spectra [14]. In addition, the calculated and experimental ECD curves, which resulted in the absolute configuration of 53 was 3*S*,4*R*,7*R*,8*R*,9*R*,17*S* [50]. Two periconiasins with another unprecedented ring system were reported, and their structures were established as a 7/6/5 tricyclic ring system of periconiasin G (54) and 5/6/6/5 tetracyclic ring system of periconiasin J (55) [14,51].



**Figure 4.** The chemical structures of polyketides 46–65 from the *Periconia* genus.

Many PKS-NRPS compounds that represent the new carbon skeleton have been reported. Additionally, pericoannosin A (56), bearing an unusual hexahydro-1*H*-isochromen-5-isobutylpyrrolidin-2-one structure was isolated from endophytic fungi *Periconia* sp. F-31 [49]. Elucidation of the structure was conducted using <sup>1</sup>H-NMR, <sup>13</sup>C-NMR, DEPT, HMBC, <sup>1</sup>H-<sup>1</sup>H COSY, and HSQC spectroscopic data. The absolute configuration was established by ECD calculations and single-crystal X-ray diffraction analyses assigned as 2*S*,3*R*,5*R*,10*S*,11*S*,16*S* [49]. This compound has a similar biosynthetic precursor with periconiasin, which shares the same aminoaldehyde intermediate. However, this pericoannosin has a different route, which underwent the Claisen and hemiacetal reaction to subsequently yield the pericoannosin A [49,51].

Meanwhile, pericoannosin B (57) was isolated in 2016 from the same source and it is known to be the stereoisomer of pericoannosin A (56) [52]. According to Zhang et al., [52], the non-stereoselective Claisen reaction and hemiacetal reaction led to the yield of stereoisomer pairs. The structure of 57 was determined with extensive spectroscopic analysis of hexahydro-1*H*-isochromen-5-isobutylpyrrolidin-2-one structure skeleton. Meanwhile, the absolute configuration was determined using CD spectra analysis to yield 2*R*,3*S*,5*S*,10*R*,11*R*,16*S* [52].

Subsequently, the investigation for the pericoannosins that was guided by a biosynthetic hypothesis led to the isolation of pericoannosins C–F (58–61) [51]. In this case, Fan et al. [51] proposed the possibility of another Diels–Alder reaction which was conducted by different pairs of conjugated diene and double bond from the same precursor of pericosins and pericoannosins. Their structures were elucidated using  $^1\text{H-NMR}$ ,  $^{13}\text{C-NMR}$ , DEPT, HSQC, HMBC,  $^1\text{H-}^1\text{H COSY}$ , and NOESY spectroscopic data. The absolute configurations of 58 were determined as 2*R*,4*R*,5*S*,10*S*,13*R*,16*S* using ECD spectra and TD-DFT comparison method at the B3LYP/6–31G(d) level. By analogy, the absolute configurations of 59–61 were also determined by calculating the ECD [51].

Recently, Liu et al. [10] reported pericoannosin G (62) to be the product of the oxidative cleavage of pericoannosins C (58) and E (60) in its biosynthetic pathway. The structure was determined using  $^1\text{H-NMR}$ ,  $^{13}\text{C-NMR}$ , DEPT, HSQC, HMBC,  $^1\text{H-}^1\text{H COSY}$ , and NOESY spectroscopic data. The absolute configuration of 62 was determined as 2*R*,3*S*,8*S*,11*R* and was calculated using the TD-DFT method at the B3LYP/6–31G(d) level (10).

Several polyketide pathway compounds were obtained from the *Periconia* genus. There are chlorinated coumarin derivatives of 5-chloro-3,4-dihydro-8-hydroxy-6-methoxy-3-methylisocoumarin (63) obtained from *P. macrospinosa* with medium Czapeck–Dox by NMR varian A60, and HA100 [53]. Henderson et al., [54] isolated 3,4-dihydro-6,8-hydroxy-6-methoxy-3-methylisocoumarin (64) and 3,4-dihydro-8-hydroxy-6-methoxy-3-methylisocoumarin (65) when chlorine are depleted from the medium.

Some polyketide-peptide hybrid compounds pathways were obtained from *P. circinata* (Mangin) Sacc with the part origin and crown root of the grain sorghum, *Sorghum bicolor* (L.). The first compound was circinatin (66), which has a biogenetic relationship with PC-toxins. It is the important pathogenicity factor from endophytic fungi *P. circinata* causing disease symptoms on its host plant, which is the cultivars of sorghum [55]. After two years, Macko et al., isolated the suspected toxins such as peritoxin A–B (67–68), along with periconin A–B (69–70) [56]. From the structural comparison studied, circinatin (66) was reported to be the precursor for the formation of peritoxin A–B (67–68) [56].

Furthermore, macrophelides, macrolide, and cyclopentenone compounds were also isolated from the *Periconia* genus. These compounds were synthesized through the polyketide pathway [57–59]. Modiolide A (71) is a macrolide compound isolated from *P. siamensis* and obtained from the leaves of *T. laffolia* [60,61]. The relative structures were determined by IR, NMR, and Mass Spectra [60]. Meanwhile, crystal structures were examined by Fun et al., [61], and the absolute configurations at C4, C7, and C9 were assigned and determined by the exciton chirality method using the *p*-methoxycinnamoyl chloride treatment. For the first time, 71 was obtained as a colorless oil from fungus *Paraphaeosphaeria* sp. (strain N-119) derived from a marine horse mussel *Modiolus auriculatus* [62].

Macrophelides is another macrolide member isolated from *P. byssoides* symbionts of sea hare *A. kurodai* and during the chemical investigation, a total of ten were isolated. Numata et al. [6] reported the macrophelides E, F–H (74, 76–78) along with C (73), and their structures were established based on  $^1\text{H-NMR}$ ,  $^{13}\text{C-NMR}$ , DEPT,  $^1\text{H-}^1\text{H COSY}$ ,  $^1\text{H-}^{13}\text{C COSY}$ , and HMBC spectral analysis. Furthermore, these compounds were re-isolated along with macrophelide A (72) and I (79) by Yamada et al. [63]. In this report, the determination of the absolute stereochemistry of 74, 76, 77, and 79 were conducted using 1D and 2D NMR spectral analysis and some chemical transformations. Furthermore, the absolute configuration of macrophelide C (73) was determined using X-ray analysis and application of the modified Mosher method [63]. The next macrophelide is seco-macrophelide E (75), which has a similar configuration as macrophelide E, followed by H (78) and L (79) along with their absolute stereochemistry based on spectroscopic analysis [64,65]. The recent investigation of macrophelide compounds from *Periconia* was conducted in 2007, where macrophelide M (81) was isolated along with the determination of its absolute stereochemistry. This was conducted using 1D and 2D NMR techniques and some chemical transformations including the modified Mosher's method [33].

Moreover, cyclopentenones were also isolated from *P. macrospinosa* by Giles and Turner [53]. There are 2-allyl-3,5-dichloro-1,4-dihydroxycyclopent-2-enoate methyl ester (82), Cryptosporiopsinol (84), and Cryptosporiopsin (85) isolate. Their fermentation was conducted using a Czapeck-Dox medium, and their structures were determined by NMR Varian A60 and HA100. Inose et al. [57] isolated another cyclopentenone, from *P. macrospinosa* cyclopericodiol (83), and the biosynthetic relation between melleins and the keto-form derivative was also proposed to yield 83. A summary of the number and types of polyketides obtained from the genus *Periconia* is demonstrated in Table 1, while Figures 4 and 5 showed the structure of the polyketide compounds.

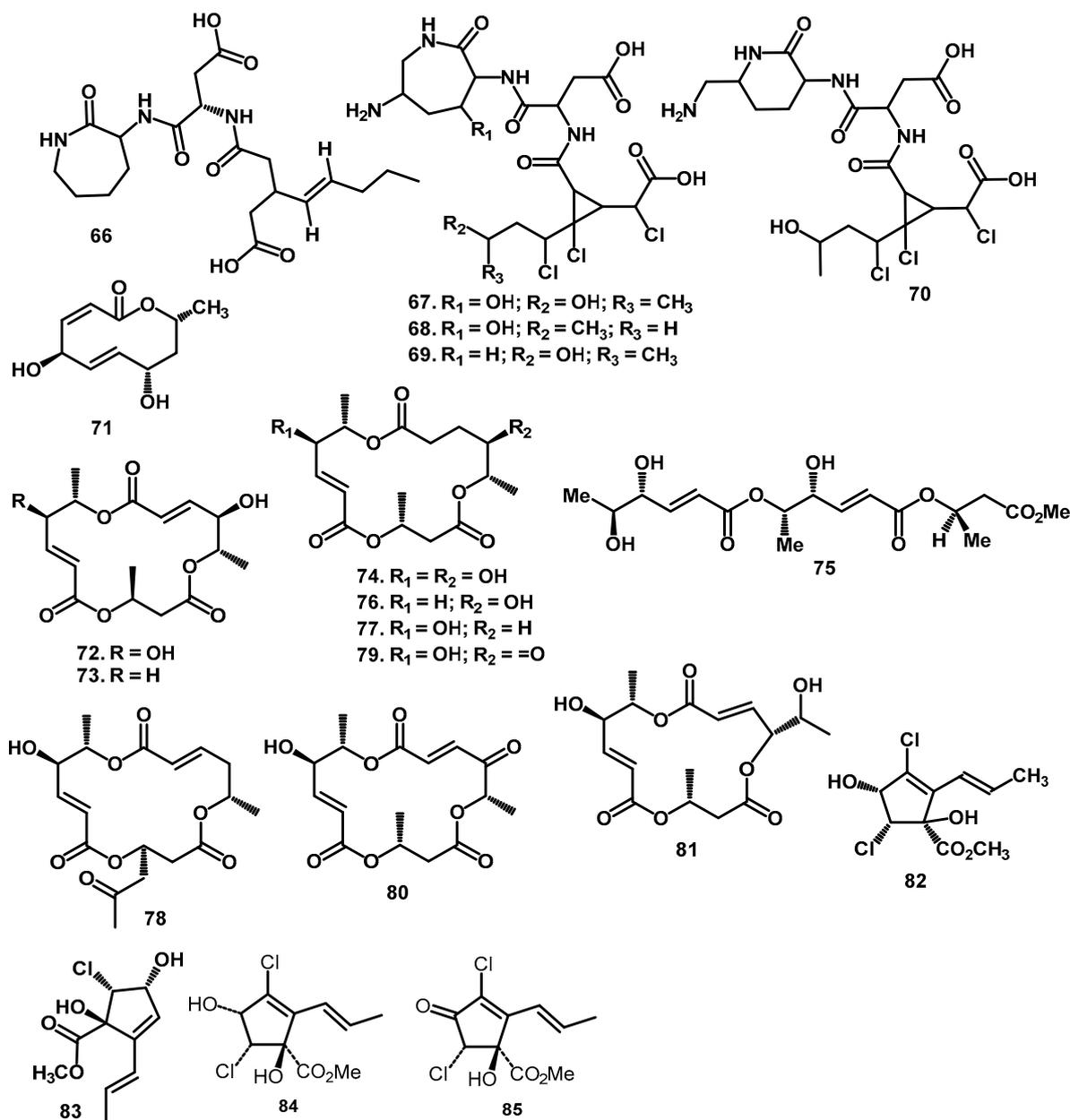


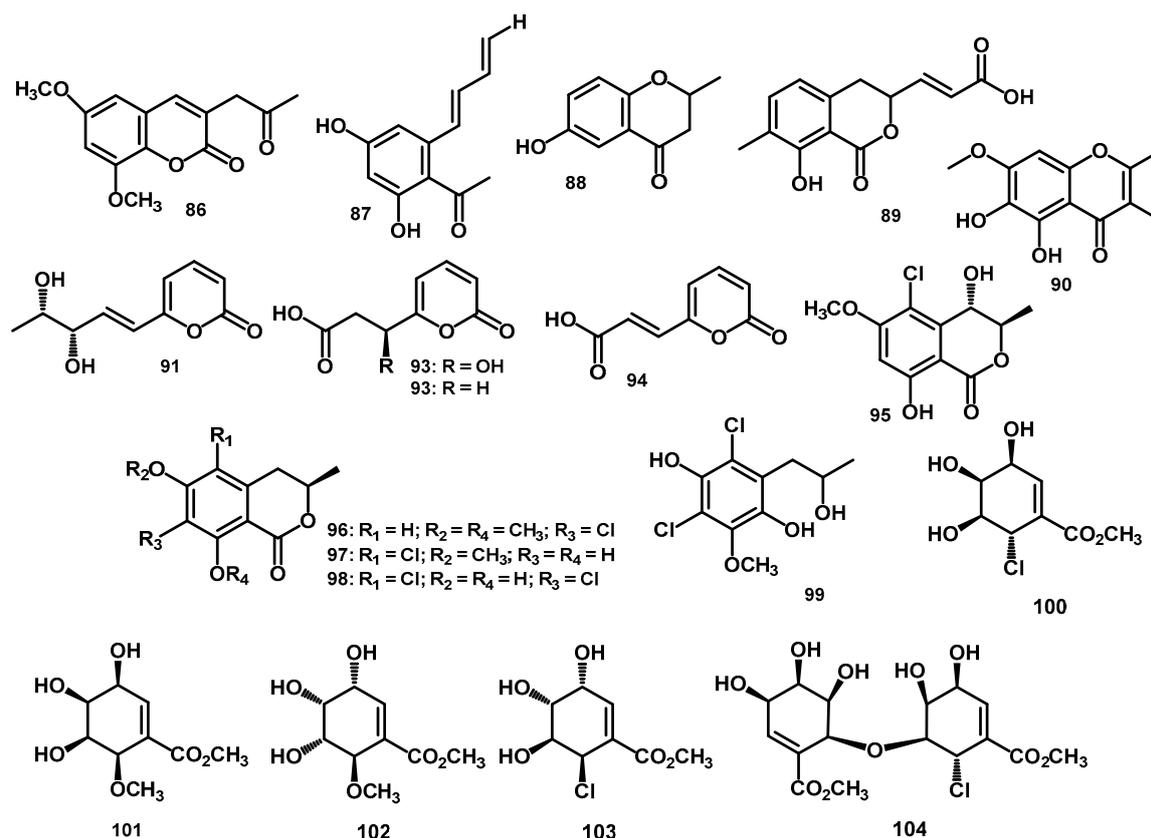
Figure 5. The chemical structures of polyketides 66–85 from the *Periconia* genus.

### 3.5. Aromatic Compounds and Carbohydrates Derivates

Fourteen aromatic compounds were isolated from this genus. These include 6,8-dimethoxy-3-(20-oxo-propyl)-coumarin (86) and 2,4-dihydroxy-6-[(10*E*,30*E*)-penta-1',3'-dienyl]-benzaldehyde (87), which were isolated from *P. atropurpurea* in the leaves of *Xylopi* *aromatica* [11]. Others are 4-chromanone, 6-hydroxy-2-methyl-(5Cl) (88), isolated from *P.*

*siamensis* in the leaves of *T. latifolia* [60]. Subsequently, many aromatic compounds derived from the polyketone biosynthetic pathway were isolated from *Periconia* sp., *Parmelia* sp., pericosins A–D (89–92), 3-(2-oxo-2H-pyran-6-yl)propanoic acid (93), and (*E*)-3-(2-oxo-2H-pyran-6-yl)acrylic acid (94) [9]. The other aromatic compounds were obtained from three melleins (3*R*,4*S*)-5-chloro-4-hydroxy-6-methoxymellein (95), (*R*)-7-chloro-6-methoxy-8-*O*-methyl-mellein (96), and (*R*)-5-chloro-6-methoxy-mellein (97) from *P. macrospinosa* KT3863 [57]. In addition, the same endophytic fungi *Periconia* sp., which was isolated from the different host plant, *A. muricata* was the source of chlorine-containing dihydroisocoumarin perichlorosins A (98) and a chlorinated phenol pericochlorosin B (99) [10]. The structures of all the aromatic compounds were determined by extensive 1D and 2D NMR for relative configurations.

The carbohydrate carbasugar derivatives pericosine A (100) and B (101) were isolated from *P. byssoides* in the sea hare *A. kurodai* through reverse-phase HPLC [6]. In addition, these compounds were re-isolated in 2007 from the same natural source along with pericosine C–E (102–104), which was separated as enantiomeric mixtures [66]. Spectroscopic analyses including NMR, MS, and IR led to the elucidation of their structures. Additionally, the structure of pericosine D (103) was revised by (–)-quinic acid precursor synthesis to methyl (3*R*,4*R*,5*S*,6*R*)-6-chloro-3,4,5-trihydroxy-1-cyclohexene-1-carboxylate [67]. Table 1 shows the summary of the number and types of aromatic compounds as well as carbohydrate derivatives obtained from the genus *Periconia*, while Figure 6 shows the structure of compounds.



**Figure 6.** The chemical structures of aromatic compounds 86–99 and carbohydrate derivatives 100–104 from the *Periconia* genus.

#### 4. Pharmacological Activities

Some metabolites of the genus *Periconia* have shown pharmacological activities for a long time. This genus has tremendous activities such as acting as antimicrobial (antibacterial, antifungal and anti-HIV), potent cytotoxicity, and anti-inflammatory agents [6–9,11–13,43,53,62].

#### 4.1. Antimicrobial Activity

The compounds from the *Periconia* genus have shown antimicrobial activities for numerous bacterial, fungi, and also human immunodeficiency virus. Bhilabutra et al. [60] accessed antibacterial properties from *P. siamensis* CMUGE015 associated with *T. latifolia* in modiolide A (71) and 4-chromanone as well as 6-hydroxy-2-methyl-(5CI) (88). The metabolites 71 inhibit more Gram-positive bacteria rather than Gram-negative. The results showed that the third compound could inhibit *Bacillus cereus*, *Listeria monocytogenes*, Methicillin-resistant *Staphylococcus aureus* (MRSA), *Pseudomonas aeruginosa*, and *Escherichia coli* with Minimum Inhibitory Concentration (MIC) value of 3.12, 6.25, 25, 12.5 and 50 µg/mL, respectively. As a comparison, penicillin G can be used for standard control with MIC values 6.25, 6.25, 25, 12.5, and 50 µg/mL, respectively. These bacteria are common human pathogen agents, which can cause foodborne disease, listeriosis, skin infection, and lung disease [60]. Furthermore, the second compound from this fungi, 88 has antibacterial activity with MIC values of 6.25, 12.5, 50, 12.5 and 100 µg/mL, respectively [62]. The similar structure of this lactone compound, tuckolide, synthesized by Andrus and Shih, acts as an inhibitor for cholesterol in liver cells [68].

An ergosterol, periconiastone A (38) isolated from *Periconia* sp. TJ403-rc01 also exhibits significant antibacterial activity against *S. aureus* with MIC 4 µg/mL and *E. faecalis* with MIC 32 µg/mL [42]. Furthermore, fussiococanne diterpenes, periconicin A (9), and B (10) have antibacterial effects on *Escherichia coli* (ATCC 25922), *Bacillus subtilis* (ATCC 6633), *Staphylococcus aureus* (ATCC 6538p), *Staphylococcus epidermis* (ATCC 12228), *Salmonella typhimurium* (ATCC 14028) *Micrococcus leuteus* (IFO 12708), *Proteus vulgaris* (ATCC 3851). *Klebsiella pneumoniae* (IFO 13541) has several MIC values. For comparison, gentamicin used with a MIC value 1.56, 3.12, 3.12, 3.12, 3.12, 6.12, 6.12, 12.5 µg/mL, respectively. Periconicin A (9) exhibited significant antibacterial activity with MIC range of 3.12–12.5 µg/mL and B (10) in the range of 25–50 µg/mL [25]. Additionally, periconicin A (7), also tested for antifungal activity, and has potent activity against human mycoses *Candida albicans*, *Trichophyton mentagrophytes*, *T. rubrum* with MIC in the range of 6.25, 6.25, 3.12 µg/mL with a control antifungal nystatin MIC range of 3.12, 1.56, 0.78 µg/mL, respectively [69].

Meanwhile, Teles et al. [11] isolated periconicin B (10) from *P. atropurpurea* and tested for anti fungi against *Cladosporium sphaerospermum* and *C. cladosporioides* through the direct bioautography method with 25.0 µg limit detection (weak activity). From the same species, 2,4-dihydroxy-6-[(10E,30E)-penta-10,30-dienyl]-benzaldehyde (87) has vigorous activity against both fungal with 1 µg limit detection compared to nystatin as a positive control (no information about limit detection).

Pericocins A–D (89–92) were obtained from endolichenic fungi, *Periconia* sp. that has a moderate and weak activity to *Aspergillus niger* (MIC value of 31 µg/mL) and *C. albicans* (MIC value of 500 µg/mL) [9] respectively. Additionally, pericoterpenoid A (24) showed moderate and weak activity to *A. niger* with MIC 31 µg/mL, and *C. albicans* with MIC 500 µg/mL [8].

HIV is a virus that attacks the human immune system, and if it is not treated, can lead to AIDS (Acquired Immunodeficiency Syndrome). Liu et al. [10] accessed the isolated compound Pericochlorosin B (99) for anti-HIV activity. The IC<sub>50</sub> value of 99 is 2.2 µM compared to positive control efavirenz with an IC<sub>50</sub> value of 1.4 nM by Firefly Luciferase Assay System. The other compounds, pericochlorosin A (98) and pericoannosin G (62), were also tested for the same assay, but they have weak activity with an IC<sub>50</sub> value of >100 µM [10]. A meroterpene, periconone B (42) has weak anti-HIV activity with an IC<sub>50</sub> value of 18 mM [43]. Additionally, cytochalasans from *Periconia*, periconiasin F (51) [49], G (52) and J (55) [51] have weak activity with an IC<sub>50</sub> value of 29.2, 67.0, 25.0 µM, respectively. Other polyketides, Pericoannosin B (57) [52], C (58), D (59) and F (61), also have weak activity with IC<sub>50</sub> with a value > 100, 15.5, 13.5 and 81.5 µM, respectively. From this genus, periconicin A (9) has antimicrobial potential, and pericochlorosin B (99) can be used as an anti-HIV agent in future developments.

#### 4.2. Cytotoxic Activity

The cytotoxic effects of compounds isolated from the genus of *Periconia* have shown promising activities against some of the cancer cells. Carbasugars, pericosine A (**100**), and B (**101**), were significant growth inhibitors of murine P-388 leukemia cell line with ED<sub>50</sub> 0.1 and 4.0 µg/mL [6]. Pericosine A (**100**) also exhibited significant in vivo tumor-inhibitory activity against P-388 in mice and inhibited protein kinase EGFR (at a concentration of 100 µg/mL by 40–70%) and topoisomerase II (with IC<sub>50</sub> value of 100–300 mM) [66]. These two compounds were studied in synthetic ways for lead cancer drugs [70–84].

Eremophilane-type sesquiterpenoid, peribysins A–D (**11–14**) from *P. byssoides* symbion of sea hare *A. kurodai*, showed promising inhibition of the adhesion of Human Leukimia HL-60 cells line and Human Umbilical Vein Endothelial Cell (HUVEC) with IC<sub>50</sub> values of 0.3, 2.7, 2.7, and 0.1 µM, respectively [13]. Compounds from this same species and host peribysins E–G (**15–17**) [31] as well as H–J (**16–18**) [32] were tested. They were successfully inhibited by similar cell lines with IC<sub>50</sub> values of 11.5, 20.7, 15.6, 15.2, 20.1, and 11.8 µM compared with herbimycin A with IC<sub>50</sub> value of 38 µM. Peribysins D (**14**) showed promising inhibition activity of HL-60 cells to HUVEC, more potent 380 times than herbimycin A [13].

Macrosphelide, a potent inhibitor of adhesion HL-60 cells to HUVEC was initially isolated from *Microsphaeropsis* sp. FO-5050 [85]. These compounds have also been produced by *P. byssoides* symbion of sea hare *A. kurodai*. Macrosphelides A (**72**), C (**73**), E (**74**), F (**76**), G (**77**), H (**78**) [63], and L (**80**) [65] were also isolated from this species. They have anti-adherent activity against HL-60 and HUVEC with IC<sub>50</sub> values of 18.7, 36.5, 19.5, 27.2, 22.5, 8.6, and 5.6 µM with the positive control, herbimycin A 38 µM. Numata et al., [6] tested macrosphelides E (**74**), F (**76**), G (**77**) and H (**78**) to P388 lymphocytic leukemia cell culture with cytotoxicity ED<sub>50</sub> > 100 µg/mL.

The HST -8 (Human Ileocecal Adenocarcinoma) and BGC-823 (Human Gastric Cancer) cell lines were successfully inhibited by periconiasin A (**46**) and B (**47**) cytochalasans from *Periconia* sp. F-31. Both cell lines were inhibited by periconiasin A (**46**) with IC<sub>50</sub> values of 0.9 and 2.1 µM and periconiasin B (**47**), with IC<sub>50</sub> values of 0.8, and 9.4 µM, respectively. Furthermore, periconiasin B (**47**) also tested for Bel-7402, a human hepatoma cell line with IC<sub>50</sub> values 5.1 µM [49]. Another cytochalasan, periconiasin I (**54**) and meroterpenes periconone E (**45**), showed in vitro cytotoxicity against human MCF-7 tumour cell line with an IC<sub>50</sub> value of 4.8 and 4.2 µM [50].

Periconicin B (**10**), a fusicoccane-type diterpenoid is isolated from *Periconia* sp. small branches of *T. cuspidata* [25] and *P. atropurpurea* from leaves *Xylopiya aromatica* [11]. It has decreased levels of HeLa and CHO cell viability, with an IC<sub>50</sub> value of 8.0 µM, and the constituent showed similar potency known as antiplastic agent Cisplatin (IC<sub>50</sub> value of 5.0 µM) [11]. Meanwhile, other terpenoids (+)-(3*S*,6*S*,7*R*,8*S*)-periconone A (**36**) and (-)-(1*R*,4*R*,6*S*,7*S*)-2-caren-4,8-olide (**8**) have weak cytotoxic activities against six human tumor cell lines (HCT-8, Bel-7402, BGC-823, A549, A2780 and MCF-7) with IC<sub>50</sub> > 10<sup>-5</sup> mM [23].

Therefore, Pericosine A (**100**) is the best growth inhibitor for the P-388 leukemia cell line with ED<sub>50</sub> 0.1 µg/mL, peribysin D (**14**) for HL-60 cells and HUVEC inhibitors with IC<sub>50</sub> 0.1 µM. Periconiasin B (**46**) is good for HST -8 and BGC-823 inhibitors with IC<sub>50</sub> values of 0.8 µM, Periconone E (**45**) is good for the MCF-7 tumor cell line with IC<sub>50</sub> 4.2 µM, and Periconicin B (**10**) is good for HeLa and CHO cells with IC<sub>50</sub> 8.0 µM. These previously mentioned compounds may be potential for future cancer treatments.

#### 4.3. Anti-Inflammatory Activity

Sesquiterpenoid, periconianone A (**25**), B (**26**), and norsesquiterpenoid dihydronaphthalene-2,6-dione (**37**) from *Periconia* sp. and host *A. muricata* were tested using lipopolysaccharide (LPS)-induced NO production in mouse microglia BV2 G13 cells for neural anti-inflammatory activity. These compounds exhibited significant inhibition with IC<sub>50</sub> values of 0.15, 0.38, and 0.23 µM compared with curcumin as a positive control with IC<sub>50</sub>

3.9  $\mu\text{M}$  [38]. Similarly, periconianone D (28), G (31) and K (35) also have inhibition effects on lipopolysaccharide-induced NO production in BV2 cells 10.2%, 18.3%, and 16.1%, respectively, compared with curcumin 12.9% inhibition at dose 1.0  $\mu\text{M}$  [39]. Furthermore, microglia are essential immune cells in the central nervous system, that play a critical role in Parkinson's and Alzheimer's Disease [86,87]. Therefore, periconianone A (25), B (26), and dihydronaphthalene-2,6-dione (37), may be potential as a lead compound for Parkinson's and Alzheimer's Disease drug candidates [38].

## 5. Conclusions and Future Perspectives

In summary, after being researched for almost 50 years, *Periconia* fungal genus was found to live in a different ecological niche such as plants, lichens, and animals. Moreover, this fungus can produce plenty of secondary metabolites with various natural products such as terpenes, polyketides, aromatic compounds, and carbohydrate derivatives. Owing to these exciting finds, eventually, the fungus has excellent potential to be a rich source for natural product exploration and can be studied for pharmacological agents.

In this review, several *Periconia* species were studied for natural product exploration, such as *Periconia* sp., *Periconia atropurpurea*, *Periconia byssoides*, *Periconia macrospinosa*, *Periconia circinata*, and *Periconia siamensis*. All these species were found from eight plants (*Annona muricata*, *Taxus cuspidata*, *Xylopiya aromatica*, *Thysanoleana latifolia*, *Sorghum bicolor*, *Rosa chinensis*, *Piper longum*, and terrestrial herbaceous plants). Additionally, there was one lichen (*Parmelia* sp.) and one sea animal (*Aplysia kurodai*). From all *Periconia* species that were already isolated, there are 104 total compounds that were studied from July 1969 to October 2020.

All these compounds have a diverse pharmacological activity that makes research of this genus more interesting. One exciting fact is that *Periconia* was studied for taxol and piperine production. This fungus can produce these host compounds (high-level plants) and if it lives as endophytes. In addition, it turns out that this *Periconia* fungus can produce new secondary metabolites that differ from the characteristics of the metabolites of the host plant. This tremendous discovery makes fungus a gold mine for natural product exploration, and it is fascinating to study their pharmacological activity.

Several potential compounds from the *Periconia* which have pharmacological activity are periconicin A (9), which acts as an antimicrobial, pericochlorosin B (99) possesses anti-HIV properties, peribysin D (14), and pericosine A (100), has vigorous cytotoxic activity, periconianone A (23) can be used as an anti-inflammatory. However, further studies are needed for the action mechanism of apoptosis study. For example, the cytotoxic mechanism of the carbasugar derivative (100). There are intrinsic mechanism studies of mitochondria-dependent cytochrome C, caspase activation, and extrinsic mechanisms of apoptosis (activation of tumor necrotic factor). The mechanism of how this compound can induce apoptosis may lead to new perspectives. On the other hand, the study of the mechanism of anti-inflammatory sesquiterpenoid compounds (25, 26, 37) in BV2 G13 microglia cells needs to be continued for further testing, such as the production of nitrite, iNOS (inducible nitric oxide synthase) mRNA, and protein expression.

Further intensive investigations are recommended concerning how the *Periconia* fungus may produce other new compounds by genetic manipulation. There are mutasynthesis, heterologous expression, and metabolic engineering at genomic level, transcriptome, and proteome (enzyme inhibition, direct precursor biosynthesis, substrate feeding) level treatments. Furthermore, the modification of fungal media can also be carried out using the principle of OSMAC (one strain many compounds), epigenetic modification, semi-synthetic media, and even by Co-Culture with other microorganisms. This treatment may induce the silent gene of the *Periconia* fungus to produce another interesting new compound. Furthermore, omics level studies (metabolomics) using LC-MS or GC-MS tools can identify the new compounds and compare all the treatments. Lastly, the discoveries of new compounds from fungi such as *Periconia* are always interesting for pharmacological studies and future lead compounds.

**Table 1.** Compounds from the *Periconia* fungal genus: type of compounds, MW and MS data, strain type, various host and origins.

Type of Compounds	Name of Compounds	Molecular Formula	Molecular Weights (g/mol)	MS (m/z)	Fungal Strains	Host Origin	Part of Host Origin	Ref.
<b>Terpenes</b>								
Monoterpene	2-carene-5,8-diol (3)	C <sub>10</sub> H <sub>16</sub> O <sub>2</sub>	168.1150	HR-ESI-MS: 151.1115 [M+H-H <sub>2</sub> O] <sup>+</sup> , 133.1011 [M+H-2xH <sub>2</sub> O] <sup>+</sup> ; GC-EI-MS: 168 [M] <sup>+</sup> , 150 [M-H <sub>2</sub> O] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[19]
	2-carene-8,10-diol (4)	C <sub>10</sub> H <sub>16</sub> O <sub>2</sub>	168.1150	HR-ESI-MS: 151.1112 [M+H-H <sub>2</sub> O] <sup>+</sup> , 133.1008 [M+H-2xH <sub>2</sub> O] <sup>+</sup> ; GC-EI-MS: 168 [M] <sup>+</sup> , 150 [M-H <sub>2</sub> O] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[19]
	2-carene-8-acetamide (5)	C <sub>12</sub> H <sub>19</sub> NO	193.1467	HR-ESI-MS: 194.1509 [M+H] <sup>+</sup> , 387.2942 [2M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[19]
	8-hydroxy-1,7-epoxy-2-menthene (6)	C <sub>10</sub> H <sub>16</sub> O <sub>2</sub>	168.1150	HR-ESI-MS: 151.1112 [M+H-H <sub>2</sub> O] <sup>+</sup> ; GC-EI-MS: 168 [M] <sup>+</sup> , 153 [M-CH <sub>3</sub> ] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[19]
	anethofuran (7) (-)-(1R,4R,6S,7S)-2-carene-4,8-olide (8)	C <sub>10</sub> H <sub>16</sub> O <sub>2</sub> C <sub>10</sub> H <sub>12</sub> O <sub>2</sub>	168.1150 164.0837	* HR-ESI-MS: 169.1229 [M+H] <sup>+</sup> HR-ESI-MS: 165.0914 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31 <i>Periconia</i> sp. F-31	<i>Annona muricata</i> <i>Annona muricata</i>	Leaves Leaves	[19] [23]
Diterpene	periconicin A (9)	C <sub>20</sub> H <sub>28</sub> O <sub>3</sub>	316.2038	HR-EI-MS: 316.2046 [M+H] <sup>+</sup>	<i>Periconia</i> sp.	<i>Taxus cuspidata</i>	Small Branches	[25]
	periconicin B (10)	C <sub>20</sub> H <sub>28</sub> O <sub>4</sub>	332.1988	HR-EI-MS: 332.2008 [M+H] <sup>+</sup>	<i>Periconia atropurpurea</i>	<i>Xylopia aromatica</i>	Leaves	[11]
Sesquiterpene	peribysin A (11)	C <sub>15</sub> H <sub>25</sub> O <sub>3</sub>	253.1803	HR-EI-MS: 253.1802 [M+H] <sup>+</sup>	<i>Periconia byssoides</i>	<i>Aplysia kurodai</i>	gastrointestinal tract	[13,34]
	peribysin B (12)	C <sub>15</sub> H <sub>24</sub> O <sub>4</sub>	268.1673	HR-EI-MS: 268.1665 [M] <sup>+</sup>	<i>Periconia byssoides</i>	<i>Aplysia kurodai</i>	gastrointestinal tract	[13,34]
	peribysin C (13)	C <sub>15</sub> H <sub>22</sub> O <sub>2</sub>	234.1619	HR-EI-MS: 234.1619 [M] <sup>+</sup>	<i>Periconia byssoides</i>	<i>Aplysia kurodai</i>	gastrointestinal tract	[13,34]
	peribysin D (14)	C <sub>15</sub> H <sub>22</sub> O <sub>2</sub>	234.1619	HR-EI-MS: 234.1618 [M] <sup>+</sup>	<i>Periconia byssoides</i>	<i>Aplysia kurodai</i>	gastrointestinal tract	[13]
	peribysin E (15)	C <sub>20</sub> H <sub>30</sub> O <sub>6</sub>	366.2042	HR-EI-MS: 366.0948 [M] <sup>+</sup>	<i>Periconia byssoides</i>	<i>Aplysia kurodai</i>	gastrointestinal tract	[31]
	peribysin F (16)	C <sub>21</sub> H <sub>32</sub> O <sub>7</sub>	396.2148	HR-EI-MS: 396.1677 [M] <sup>+</sup>	<i>Periconia byssoides</i>	<i>Aplysia kurodai</i>	gastrointestinal tract	[13,34]
	peribysin G (17)	C <sub>21</sub> H <sub>32</sub> O <sub>7</sub>	396.2148	HR-EI-MS: 396.1677 [M] <sup>+</sup>	<i>Periconia byssoides</i>	<i>Aplysia kurodai</i>	gastrointestinal tract	[13,34]
	peribysin H (18)	C <sub>15</sub> H <sub>24</sub> O <sub>4</sub>	268.1673	HR-EI-MS: 268.1673 [M] <sup>+</sup>	<i>Periconia byssoides</i>	<i>Aplysia kurodai</i>	gastrointestinal tract	[32]
	peribysin I (19)	C <sub>15</sub> H <sub>24</sub> O <sub>4</sub>	268.1673	HR-EI-MS: 268.1673 [M] <sup>+</sup>	<i>Periconia byssoides</i>	<i>Aplysia kurodai</i>	gastrointestinal tract	[32]
	peribysin J (20)	C <sub>15</sub> H <sub>26</sub> O <sub>5</sub>	286.1773	HR-EI-MS: 268.1775 [M] <sup>+</sup>	<i>Periconia byssoides</i>	<i>Aplysia kurodai</i>	gastrointestinal tract	[33]
	peribysin O (21)	C <sub>15</sub> H <sub>22</sub> O <sub>4</sub>	266.1518	ESI-TOF-MS: 289.1416 [M+Na] <sup>+</sup> ; 555.2919 [2M+Na] <sup>+</sup>	<i>P. macrospinosa</i> KT3863	terrestrial herbaceous plant at Kanagawa prefecture in 2018	-	[37]
peribysin P (22)	C <sub>15</sub> H <sub>24</sub> O <sub>4</sub>	268.1675	ESI-TOF-MS: 291.1566 [M+Na] <sup>+</sup> ; 559.3230 [2M+Na] <sup>+</sup>	<i>P. macrospinosa</i> KT3863	terrestrial herbaceous plant at Kanagawa prefecture in 2018	-	[37]	
peribysin Q (23)	C <sub>15</sub> H <sub>20</sub> O <sub>3</sub>	248.1412	ESI-TOF-MS: 249.1496 [M+H] <sup>+</sup> ; 231.1388 [M+H-H <sub>2</sub> O] <sup>+</sup> ; 271.1311 [M+Na] <sup>+</sup>	<i>P. macrospinosa</i> KT3863	terrestrial herbaceous plant at Kanagawa-prefecture in 2018	-	[37]	

Table 1. Cont.

Type of Compounds	Name of Compounds	Molecular Formula	Molecular Weights (g/mol)	MS (m/z)	Fungal Strains	Host Origin	Part of Host Origin	Ref.
	pericoterpenoid A (24)	C <sub>15</sub> H <sub>16</sub> O <sub>3</sub>	244.1099	HR-ESI-MS: 267.1002 [M+Na] <sup>+</sup>	<i>Periconia</i> sp.	<i>Parmelia</i> sp.	-	[8]
	periconianone A (25)	C <sub>15</sub> H <sub>18</sub> O <sub>4</sub>	262.1205	HR-ESI-MS: 263.1271 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[38]
	periconianone B (26)	C <sub>15</sub> H <sub>18</sub> O <sub>4</sub>	262.1205	HR-ESI-MS: 263.1280 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[38]
	periconianone C (27)	C <sub>15</sub> H <sub>20</sub> O <sub>4</sub>	264.1362	HR-ESI-MS: 265.1434 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[39]
	periconianone D (28)	C <sub>15</sub> H <sub>20</sub> O <sub>3</sub>	248.1412	HR-ESI-MS: 249.1458 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[39]
	periconianone E (29)	C <sub>15</sub> H <sub>20</sub> O <sub>3</sub>	248.1412	HR-ESI-MS: 249.1478 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[39]
	periconianone F (30)	C <sub>15</sub> H <sub>22</sub> O <sub>5</sub>	282.1467	HR-ESI-MS: 283.1540 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[39]
	periconianone G (31)	C <sub>15</sub> H <sub>24</sub> O <sub>4</sub>	268.1675	HR-ESI-MS: 291.1567 [M+Na] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[39]
	periconianone H (32)	C <sub>12</sub> H <sub>16</sub> O <sub>3</sub>	208.1099	HR-ESI-MS: 209.1172 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[39]
	periconianone I (33)	C <sub>12</sub> H <sub>16</sub> O <sub>3</sub>	208.1099	HR-ESI-MS: 209.1172 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[39]
	periconianone J (34)	C <sub>12</sub> H <sub>16</sub> O <sub>3</sub>	208.1099	HR-ESI-MS: 209.1172 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[39]
	periconianone K (35)	C <sub>12</sub> H <sub>16</sub> O <sub>3</sub>	208.1099	HR-ESI-MS: 209.1172 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[39]
	(+)-(3S,6S,7R,8S)-periconone A (36)	C <sub>15</sub> H <sub>22</sub> O <sub>5</sub>	282.1467	HR-ESI-MS: 283.1549 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[23]
Norsesqui-terpene	Dihydronaphthalene-2,6-dione (37)	C <sub>12</sub> H <sub>12</sub> O <sub>3</sub>	204.0786	* HR-ESI-MS: 205.0865 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[38]
Steroid	periconiastone A (38)	C <sub>28</sub> H <sub>42</sub> O <sub>3</sub>	426.3134	HR-ESI-MS: 449.3026 [M+Na] <sup>+</sup>	<i>Periconia</i> sp. TJ403-rc01	<i>Rosa chinensis</i>	Leaves	[43]
Meroterpenes	pericolactine A (39)	C <sub>24</sub> H <sub>35</sub> NO <sub>5</sub>	417.2515	HR-ESI-MS: 418.2595 [M+H] <sup>+</sup>	<i>Periconia</i> sp.	<i>Parmelia</i> sp.	-	[7]
	pericolactine B (40)	C <sub>22</sub> H <sub>33</sub> NO <sub>4</sub>	375.2410	HR-ESI-MS: 376.2491 [M+H] <sup>+</sup>	<i>Periconia</i> sp.	<i>Parmelia</i> sp.	-	[7]
	pericolactine C (41)	C <sub>23</sub> H <sub>35</sub> NO <sub>5</sub>	405.2515	HR-ESI-MS: 428.2418 [M+Na] <sup>+</sup>	<i>Periconia</i> sp.	<i>Parmelia</i> sp.	-	[7]
	periconone B (42)	C <sub>17</sub> H <sub>24</sub> O <sub>4</sub>	292.1675	HR-ESI-MS: 293.1741 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[43]
	periconone C (43)	C <sub>17</sub> H <sub>24</sub> O <sub>5</sub>	308.1624	HR-ESI-MS: 331.1516 [M+Na] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[43]
	periconone D (44)	C <sub>17</sub> H <sub>24</sub> O <sub>5</sub>	308.1624	HR-ESI-MS: 331.1516 [M+Na] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[43]
	periconone E (45)	C <sub>15</sub> H <sub>22</sub> O <sub>5</sub>	282.1467	HR-ESI-MS: 283.1540 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[43]
<b>Polyketides</b>								
Cytochalasan	periconiasin A (46)	C <sub>22</sub> H <sub>33</sub> NO <sub>3</sub>	359.2460	HR-ESI-MS: 360.2528 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[49]
	periconiasin B (47)	C <sub>22</sub> H <sub>33</sub> NO <sub>3</sub>	359.2460	HR-ESI-MS: 360.2525 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[48]
	periconiasin C (48)	C <sub>22</sub> H <sub>31</sub> NO <sub>3</sub>	357.2304	HR-ESI-MS: 358.2377 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[48]
	periconiasin D (49)	C <sub>22</sub> H <sub>33</sub> NO <sub>3</sub>	359.2460	HR-ESI-MS: 360.2533 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[49]
	periconiasin E (50)	C <sub>22</sub> H <sub>33</sub> NO <sub>3</sub>	359.2460	HR-ESI-MS: 360.2533 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[49]
	periconiasin F (51)	C <sub>22</sub> H <sub>33</sub> NO <sub>3</sub>	359.2460	HR-ESI-MS: 360.2533 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[49]

Table 1. Cont.

Type of Compounds	Name of Compounds	Molecular Formula	Molecular Weights (g/mol)	MS (m/z)	Fungal Strains	Host Origin	Part of Host Origin	Ref.
	periconiasin G (52)	C <sub>20</sub> H <sub>31</sub> NO <sub>2</sub>	317.2355	HR-ESI-MS: 318.2427 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[14]
	periconiasin H (53)	C <sub>27</sub> H <sub>41</sub> NO <sub>6</sub> S	507.2655	HR-ESI-MS: 508.2703 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[14]
	periconiasin I (54)	C <sub>22</sub> H <sub>33</sub> NO <sub>4</sub>	375.2410	HR-ESI-MS: 376.2474 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[50]
	periconiasin J (55)	C <sub>22</sub> H <sub>31</sub> NO <sub>2</sub>	341.2355	HR-ESI-MS: 342.2422 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[50]
	pericoannosin A (56)	C <sub>22</sub> H <sub>35</sub> NO <sub>3</sub>	361.2617	HR-ESI-MS: 384.2505 [M+Na] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[50]
Other Polyketides	pericoannosin B (57)	C <sub>22</sub> H <sub>35</sub> NO <sub>3</sub>	361.2617	FABMS: 362.2 [M+H] <sup>+</sup> ; HR-ESI-MS: 384.2495 [M+Na] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[52]
	pericoannosin C (58)	C <sub>22</sub> H <sub>33</sub> NO <sub>3</sub>	359.2460	HR-ESI-MS: 360.2527 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[52]
	pericoannosin D (59)	C <sub>22</sub> H <sub>33</sub> NO <sub>3</sub>	359.2460	HR-ESI-MS: 360.2527 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[52]
	pericoannosin E (60)	C <sub>22</sub> H <sub>33</sub> NO <sub>3</sub>	359.2460	HR-ESI-MS: 360.2527 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[52]
	pericoannosin F (61)	C <sub>22</sub> H <sub>33</sub> NO <sub>3</sub>	359.2460	HR-ESI-MS: 360.2527 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[52]
	pericoannosin G (62)	C <sub>14</sub> H <sub>21</sub> NO	219.1623	HR-ESI-MS: 220.1696 [M+H] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[10]
	5-chloro-3,4-dihydro-8-hydroxy-6-methoxy-3-methylisocoumarin (63)	C <sub>11</sub> H <sub>11</sub> ClO <sub>4</sub>	242.0346	* HR-ESI-MS: 243.0414 [M+H] <sup>+</sup>	<i>Periconia macrospinosa</i>	-	-	[55]
	3,4-dihydro-6,8-dihydroxy-3-methylisocoumarin (64)	C <sub>10</sub> H <sub>10</sub> O <sub>4</sub>	194.0579	* HR-ESI-MS: 195.0657 [M+H] <sup>+</sup>	<i>Periconia macrospinosa</i>	-	-	[54]
	3,4-dihydro-8-hydroxy-6-methoxy-3-methylisocoumarin (65)	C <sub>11</sub> H <sub>12</sub> O <sub>4</sub>	208.0736	* HR-ESI-MS: 209.0814 [M+H] <sup>+</sup>	<i>Periconia macrospinosa</i>	-	-	[54]
	Circinatin (66)	C <sub>20</sub> H <sub>31</sub> O <sub>7</sub> N <sub>3</sub>	425	FABMS: 426 [M+H] <sup>+</sup>	<i>Periconia circinata</i> (Mangin) Sacc.	<i>Sorghum bicolor</i> (L.)	root and crown root	[57]
peritoxin A (67)	C <sub>20</sub> H <sub>29</sub> O <sub>9</sub> N <sub>4</sub> Cl <sub>3</sub>	574	FABMS: 575 [M+H] <sup>+</sup>	<i>Periconia circinata</i> (Mangin) Sacc.	<i>Sorghum bicolor</i> (L.)	root and crown root	[56]	
peritoxin B (68)	C <sub>20</sub> H <sub>29</sub> O <sub>8</sub> N <sub>4</sub> Cl <sub>3</sub>	558	FABMS: 559 [M+H] <sup>+</sup>	<i>Periconia circinata</i> (Mangin) Sacc.	<i>Sorghum bicolor</i> (L.)	root and crown root	[56]	
periconin A (69)	C <sub>20</sub> H <sub>29</sub> O <sub>8</sub> N <sub>4</sub> Cl <sub>3</sub>	558	FABMS: 559 [M+H] <sup>+</sup>	<i>Periconia circinata</i> (Mangin) Sacc.	<i>Sorghum bicolor</i> (L.)	root and crown root	[56]	
periconin B (70)	C <sub>20</sub> H <sub>29</sub> O <sub>8</sub> N <sub>4</sub> Cl <sub>3</sub>	558	FABMS: 559 [M+H] <sup>+</sup>	<i>Periconia circinata</i> (Mangin) Sacc.	<i>Sorghum bicolor</i> (L.)	root and crown root	[56]	
Macrolide	modiolide A (71)	C <sub>10</sub> H <sub>14</sub> O <sub>4</sub>	198.21	* HR-EI-MS: 199.0970 [M+H] <sup>+</sup>	<i>Periconia siamensis</i> CMUGE015	<i>Thysanoleana laffolia</i>	Leaves	[60,61]

Table 1. Cont.

Type of Compounds	Name of Compounds	Molecular Formula	Molecular Weights (g/mol)	MS (m/z)	Fungal Strains	Host Origin	Part of Host Origin	Ref.
Macrosphelide	macrosphelide A (72)	C <sub>16</sub> H <sub>22</sub> O <sub>8</sub>	342.1315	HR-EI-MS: 343.1392 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[65]
	macrosphelide C (73)	C <sub>16</sub> H <sub>22</sub> O <sub>7</sub>	326.1366	HR-EI-MS: 327.1442 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[6,63]
	macrosphelide E (74)	C <sub>16</sub> H <sub>22</sub> O <sub>8</sub>	342.1315	HR-EI-MS: 343.1392 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[6,63]
	seco-Macrosphelide E (75)	C <sub>16</sub> H <sub>23</sub> O <sub>8</sub>	342.1315	HR-EI-MS: 342.1315 [M] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[66]
	macrosphelide F (76)	C <sub>16</sub> H <sub>22</sub> O <sub>7</sub>	326.1366	HR-EI-MS: 327.1442 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[6,63]
	macrosphelide G (77)	C <sub>16</sub> H <sub>22</sub> O <sub>8</sub>	342.1315	HR-EI-MS: 343.1392 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[6,63]
	macrosphelide H (78)	C <sub>16</sub> H <sub>22</sub> O <sub>8</sub>	342.1315	HR-EI-MS: 343.1392 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[6,63]
	macrosphelide I (79)	C <sub>16</sub> H <sub>22</sub> O <sub>8</sub>	342.1315	HR-EI-MS: 343.1392 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[65]
	macrosphelide L (80)	C <sub>16</sub> H <sub>22</sub> O <sub>8</sub>	342.1315	HR-EI-MS: 342.1313 [M] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[67]
	macrosphelide M (81)	C <sub>16</sub> H <sub>22</sub> O <sub>8</sub>	342.1315	HR-EI-MS: 343.1392 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[33]
Cyclopentene	methyl 2-allyl-3,5-dichloro-1,4-dihydroxycyclopent-2-enoate methyl ester (82)	C <sub>10</sub> H <sub>12</sub> Cl <sub>2</sub> O <sub>4</sub>	374.0113	* HR-EI-MS: 375.0191 [M+H] <sup>+</sup>	<i>Periconia macrospinosa</i>	-	-	[55]
	cyclopericodiol (83)	C <sub>12</sub> H <sub>12</sub> O <sub>5</sub>	236.0685	ESI-TOF-MS: 255.0399 [M+Na] <sup>+</sup> ; 487.0910 [2M+Na] <sup>+</sup> ; 215.0475 [M+H-H <sub>2</sub> O] <sup>+</sup>	<i>Periconia macrospinosa</i> KT3863	terrestrial herbaceous plant at Kanagawa prefecture in 2018	-	[57]
	Cryptosporiopsinol (84)	C <sub>10</sub> H <sub>12</sub> Cl <sub>2</sub> O <sub>4</sub>	266.0113	* HR-EI-MS: 267.0191 [M+H] <sup>+</sup>	<i>Periconia macrospinosa</i>	-	-	[53]
	Cryptosporiopsin (85)	C <sub>10</sub> H <sub>10</sub> Cl <sub>2</sub> O <sub>4</sub>	263.9956	* HR-EI-MS: 265.0034 [M+H] <sup>+</sup>	<i>Periconia macrospinosa</i>	-	-	[53]

Table 1. Cont.

Type of Compounds	Name of Compounds	Molecular Formula	Molecular Weights (g/mol)	MS (m/z)	Fungal Strains	Host Origin	Part of Host Origin	Ref.
<b>Aromatic Compounds</b>								
	6,8-Dimethoxy-3-(20-oxo-propyl)-coumarin (86)	C <sub>14</sub> H <sub>14</sub> O <sub>5</sub>	262.0841	HR-ESI-MS: 285.0739 [M+Na] <sup>+</sup>	<i>Periconia atropurpurea</i>	<i>Xylopia aromatica</i>	Leaves	[11]
	2,4-dihydroxy-6-[(10E,30E)-penta-1',3'-dienyl]-benzaldehyde (87)	C <sub>12</sub> H <sub>12</sub> O <sub>3</sub>	204.0786	HR-ESI-MS: 203.0708 [M-H] <sup>-</sup>	<i>Periconia atropurpurea</i>	<i>Xylopia aromatica</i>	Leaves	[11]
	4-chromanone, 6-hydroxy-2-methyl-(5CI) (88)	C <sub>10</sub> H <sub>10</sub> O <sub>3</sub>	178.0630	* HR-EI-MS: 179.0708 [M+H] <sup>+</sup>	<i>Periconia siamensis</i> <i>CMUGE015</i>	<i>Thysanoleana latifolia</i>	Leaves	[62]
	pericocin A (89)	C <sub>13</sub> H <sub>12</sub> O <sub>5</sub>	248.0685	HR-ESI-MS: 271.0587 [M+Na] <sup>+</sup>	<i>Periconia</i> sp.	<i>Parmelia</i> sp.	-	[9]
	pericocin B (90)	C <sub>12</sub> H <sub>12</sub> O <sub>5</sub>	236.0685	HR-ESI-MS: 237.0769 [M+H] <sup>+</sup>	<i>Periconia</i> sp.	<i>Parmelia</i> sp.	-	[9]
	pericocin C (91)	C <sub>10</sub> H <sub>12</sub> O <sub>4</sub>	196.0736	HR-ESI-MS: 197.0814 [M+H] <sup>+</sup>	<i>Periconia</i> sp.	<i>Parmelia</i> sp.	-	[9]
	pericocin D (92)	C <sub>8</sub> H <sub>8</sub> O <sub>5</sub>	184.0372	HR-ESI-MS: 185.0450 [M+H] <sup>+</sup>	<i>Periconia</i> sp.	<i>Parmelia</i> sp.	-	[9]
	3-(2-oxo-2H-pyran-6-yl)propanoic acid (93)	C <sub>8</sub> H <sub>8</sub> O <sub>4</sub>	168.0423	* HR-ESI-MS: 169.0501 [M+H] <sup>+</sup>	<i>Periconia</i> sp.	<i>Parmelia</i> sp.	-	[9]
	(E)-3-(2-oxo-2H-pyran-6-yl)acrylic acid (94)	C <sub>8</sub> H <sub>6</sub> O <sub>4</sub>	166.0266	* HR-ESI-MS: 167.0344 [M+H] <sup>+</sup>	<i>Periconia</i> sp.	<i>Parmelia</i> sp.	-	[9]
	(3R,4S)-5-chloro-4-hydroxy-6-methoxymellein (95)	C <sub>11</sub> H <sub>11</sub> ClO <sub>5</sub>	258.0295	ESI-TOF-MS: 259.0368 [M+H] <sup>+</sup> ; 281.0187 [M+Na] <sup>+</sup> ; 296.9927 [M+K] <sup>+</sup> ; 539.0482 [2M+Na] <sup>+</sup>	<i>Periconia macrospinoso</i> KT3863	terrestrial herbaceous plant at Kanagawa prefecture in 2018	-	[57]
	(R)-7-chloro-6-methoxy-8-O-methyl-mellein (96)	C <sub>12</sub> H <sub>13</sub> ClO <sub>4</sub>	256.0502	ESI-TOF-MS: 257.0575 [M+H] <sup>+</sup> ; 279.0395 [M+Na] <sup>+</sup> ; 535.0897 [2M+Na] <sup>+</sup>	<i>Periconia macrospinoso</i> KT3863	terrestrial herbaceous plant at Kanagawa prefecture in 2018	-	[57]
	(R)-5-chloro-6-methoxy-mellein (97)	C <sub>11</sub> H <sub>11</sub> ClO <sub>4</sub>	242.0346	ESI-TOF-MS: 243.0419 [M+H] <sup>+</sup> ; 265.0238 [M+Na] <sup>+</sup> ; 280.9977 [M+K] <sup>+</sup> ; 507.0584 [2M+Na] <sup>+</sup>	<i>Periconia macrospinoso</i> KT3863	terrestrial herbaceous plant at Kanagawa prefecture in 2018	-	[57]
	pericochlorosin A (98)	C <sub>10</sub> H <sub>7</sub> Cl <sub>2</sub> O <sub>4</sub>	260.9721	ESI-MS m/z 263.3 [M+H] <sup>+</sup> , 265.2 [M+2-H] <sup>+</sup> , 267.2 [M+4-H] <sup>+</sup> ; HR-ESI-MS m/z 260.9723 [M-H] <sup>-</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[10]
	pericochlorosin B (99)	C <sub>10</sub> H <sub>11</sub> Cl <sub>2</sub> O <sub>3</sub>	249.0085	ESI-MS m/z 265.6 [M-H] <sup>-</sup> , 267.2 [M+2-H] <sup>-</sup> , 269.3 [M+4-H] <sup>-</sup> ; HR-ESI-MS m/z 249.0080 [M+H-H <sub>2</sub> O] <sup>+</sup>	<i>Periconia</i> sp. F-31	<i>Annona muricata</i>	Leaves	[10]

Table 1. Cont.

CarbohydrateDerivates								
Type of Compounds	Name of Compounds	Molecular Formula	Molecular Weights (g/mol)	MS (m/z)	Fungal Strains	Host Origin	Part of Host Origin	Ref.
	pericosine A (100)	C <sub>8</sub> H <sub>11</sub> ClO <sub>5</sub>	222.0295	HR-EI-MS: 223.0373 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[6,66]
	pericosine B (101)	C <sub>9</sub> H <sub>13</sub> O <sub>6</sub>	217.0712	HR-EI-MS: 218.0790 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[6,66]
	pericosine C (102)	C <sub>9</sub> H <sub>14</sub> O <sub>6</sub>	218.0790	HR-EI-MS: 219.0854 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[66]
	pericosine D (103)	C <sub>8</sub> H <sub>11</sub> ClO <sub>5</sub>	222.0295	HR-EI-MS: 223.0363 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[66]
	pericosine E (104)	C <sub>16</sub> H <sub>21</sub> ClO <sub>10</sub>	408.0832	HR-EI-MS: 409.0900 [M+H] <sup>+</sup>	<i>Periconia byssoides</i> OUPS-N133	<i>Aplysia kurodai</i>	gastrointestinal tract	[66]

Notes: \* Several compounds with MS predictio.

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