

Chapter 11

Flavonoids of *Baccharis*



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Abstract Despite more than 400 species of *Baccharis* occurring worldwide, only less than 20% of the species have chemically been studied. In the *Baccharis* genus, within terpenes and other phenolic compounds (see Chaps. 12 and 13), flavonoids are largely accumulated as aglycone, being apigenin, genkwanin, hispidulin, kaempferol (flavones), quercetin (flavonol), naringenin, sakuranetin (flavanones), and others widely distributed. Additionally, some flavonoid glycosides such as quercitrin, rutin, and others are also found, but in minor frequency. Flavonoids are compounds with a basic skeleton of 15 carbons ($C_6-C_3-C_6$) arranged in two aromatic rings linked through a three-carbon moiety. The oxidation degree of the C_3 moiety is directly related to the classification of flavonoids into flavanones, flavones, isoflavones, flavanonols, and flavonols. With respect to biological activity, flavonoids from *Baccharis* display a significant antioxidant potential, especially for the capacity of suppression of ROS formation, ROS scavenging, and upregulation of antioxidant defenses. In this chapter, the distribution of flavonoids in *Baccharis* is so justified through this antioxidant effect since those species are inserted in areas with direct incidence of sunrays, such as in montane savannas. In addition, flavonoids display hepatoprotective, antimicrobial, anti-inflammatory, antitumoral, antiviral, and other activities. In this chapter, the occurrence and distribution of flavonoids in *Baccharis* species are discussed, as well as their biosynthesis and biological aspects.

Keywords Biological activity · Biosynthesis · Flavonoid distribution · Modulation of redox balance · Structure and composition

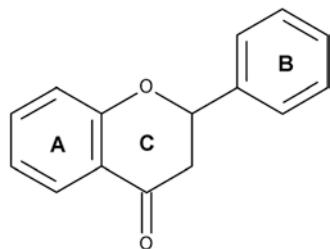
1 Introduction

Although more than 400 *Baccharis* species are distributed worldwide, only a reduced number of plants belonging to this genus were chemically and/or biologically investigated. Among the chemical compounds present in these species, the

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Fig. 11.1 Basic skeleton of a flavonoid



occurrence of flavonoids is reported in about 15% of *Baccharis* species (Campos et al. 2016). Flavonoids are a class of phenolic compounds, widely distributed within the kingdom Plantae, with a basic skeleton of 15 carbons (Fig. 11.1) arranged as two aromatic rings (A and B) connected by a three-carbon moiety ($C_6-C_3-C_6$).

The classification of flavonoids is directly related to the degree of oxidation of the C_3 portion. Flavonoids can be classified into flavanones, flavononols, flavones, isoflavones, flavonols, catechins, and anthocyanidins.

2 Biosynthesis

Secondary metabolites, also known as special metabolites, are substances from natural sources that do not participate in the essential functions of these organisms (reproduction, growth, or development). Most are substances produced in processes of interaction between plants and the environment in which they are inserted. It is important to note that the production of secondary metabolites is also associated with the various conditions to which plant species are subjected. Since these substances are produced to benefit these individuals, depending on the environment surrounding them, it acts as protection against the action of predators, coloring and volatiles agents, attracting pollinators, and also as competition agents among plants in the same habitat. Therefore, the production of these metabolites is fundamental in the maintenance of these species (Dewick 2009).

Besides being responsible for the production of substances that act with different functions in the interaction of these plant species and the environment, which are inserted, the secondary metabolism is also responsible for providing the majority of natural products with pharmacological activity. The secondary metabolism acts as

substrates originating from the metabolic pathways that form the primary metabolism (photosynthesis, glycolysis, and Krebs/ citric acid cycle). These substrates are involved in different metabolic pathways, responsible for the synthesis of several classes of secondary metabolites.

Flavonoids are considered mixed pathway metabolites, being synthesized from precursors of two metabolic routes; shikimic acid (shikimate) and acetate-malonate. From the glycolysis process, the synthesis of phosphoenolpyruvate, together with erythrose-4-phosphate from the pentose-phosphate pathway, is responsible for the synthesis of shikimic acid. In glycolysis, phosphoenolpyruvate, through the enzyme pyruvate kinase, transfers the phosphate ion to a molecule of ADP, generating pyruvate and ATP. This pyruvate is responsible for the synthesis of acetyl-CoA (acetyl coenzyme A). From the shikimic acid path, *p*-coumaric acid (4-coumaric acid) is obtained, resulting from the ammonia elimination from L-phenylalanine side chain (precursors of the C₆-C₃ portions), and precursor of *p*-coumaric alcohol. The elimination of ammonia occurs in the presence of PAL (phenylalanine ammonia lyase) into cinnamic acid, which is converted into *p*-coumaric acid by direct hydroxylation, in the presence of cinnamate-4-hydroxylase (C4H), which is converted to 4-hydroxycinnamoyl-CoA by the 4-coumarate CoA ligase (4CL), by a process known as the general pathway to the formation of phenylpropanoids. The enzyme involved in the condensation process of malonyl-CoA units to form flavonoids is chalcone synthase, through the acetyl-CoA carboxylase (ACC) mediates carboxylation reaction. Therefore, the formation of flavonoids (Fig. 11.2) occurs from 4-hydroxycinnamoyl-CoA units, derived from the pathway of shikimic acid with the addition of malonyl-CoA units (for chain elongation), the route of acetate-malonate (Davies and Schwinn 2006; Dewick 2009).

After the addition of malonyl-CoA units, two pathways could be involved in the formation of different metabolites – flavonoids or stilbenes (e.g., resveratrol). After the formation of the polyketide, different enzymes are involved in aldol or Claisen type condensations to form aromatic ring A. For the formation of flavonoids, the enzyme involved in the condensation process of malonyl-CoA units is chalcone synthase. Chalcones are the precursors of flavonoids (I), followed by enolization (II) of the polyketide (Fig. 11.3).

For the synthesis of flavonoids, chalcones undergo a nucleophilic attack reaction of the OH group to the α,β -unsaturated ketone, forming a heterocyclic ring (C-ring) yielding flavanones. Acidic environments may favor the synthesis of flavanones,

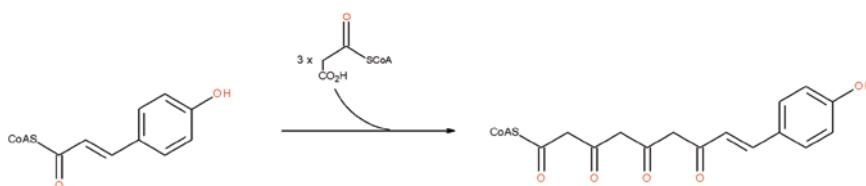


Fig. 11.2 Formation of the polyketide side chain, precursor of flavonoids, by acetate-malonate route

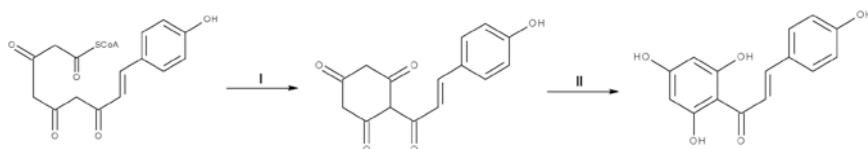


Fig. 11.3 Steps for chalcone biosynthesis from polyketide side chain

while alkaline environments favor the production of chalcones. However, it is worth emphasizing that in nature, these processes occur under very specific conditions, in the presence of stereospecific enzymes that prevent the formation of enantiomers. From the flavanones, a large variety of flavonoids can be synthesized, such as flavones, flavonols, and anthocyanidins (Davies and Schwinn 2006).

According to the basic skeleton of flavonoids, the variations between the different classes of flavonoids are related to the oxygenations and substituents of rings B and C. However, many flavonoids can also lose one or two hydroxyl groups in ring A, a process related to the action of chalcone reductase and chalcone synthase enzymes. The enzymatic complex involved in the biosynthesis of the different classes of flavonoids is broad and with high specificity, through this complex, the alterations of the oxygenation patterns in the aryl moiety are realized. In addition, methylation, glycosylation, and dimethylation processes are responsible for increasing the possibilities of formation of different compounds, increasing the diversity of flavonoids distributed in different plant species (Dewick 2009).

Flavanones, the first group of flavonoids synthesized from chalcones, are the precursors of the other groups of flavonoids. Flavones are synthesized from reactions in the presence of flavone synthase I, oxygen and 2-oxo-glutarate and flavone synthase IIe, oxygen and NADPH. In the presence of oxygen, 2-oxoglutarate, and flavanone 3-hydroxylase enzyme, dihydroflavonols are produced, which are the precursors of flavonols and flavandiols. Finally, flavodiols act as precursors of the catechins and anthocyanidins, through specific enzymatic processes that occur in the presence of oxygen and NADPH, and by water elimination (Davies and Schwinn 2006; Dewick 2009). Figure 11.4 briefly illustrates the biosynthesis of flavonoids.

3 Flavonoid Composition

As described above, the main composition of *Baccharis* are flavonoids, diterpenes, and other phenolic compounds. From the *Baccharis* genus, which is composed of 441 species, only approximately 20% have been investigated in chemical aspects. The occurrence of flavonoids was reported in 86 distinct species of *Baccharis*; these studies lead to the identification of 129 flavonoids separated into 16 flavanones (Table 11.1), 11 flavanonols (Table 11.2), 46 flavones (Table 11.3), and 55 flavonols (Table 11.4).

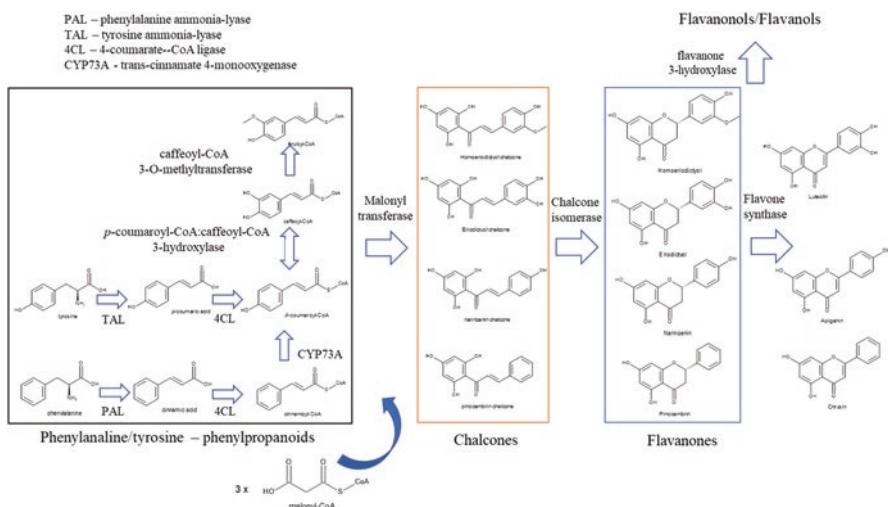


Fig. 11.4 Scheme of flavonoid biosynthesis

4 Flavanones

The main precursor of all flavonoids, naringenin (**1**), was found in aerial parts of *B. alaternoides* (Bohlmann et al. 1979), *B. conferta* (Weimann et al. 2002), *B. ligustrina* (Moreira et al. 2003a, b; Abad and Bermejo 2007), *B. polyccephala* (Davila et al. 2013), *B. retusa* (Grecco et al. 2012a, b; Campos et al. 2016), *B. salzmannii* (Bohlmann et al. 1981a, b), and *B. varians* (Bohlmann et al. 1981a). Compound **1** was also found in the flowers of *B. illinita* (Verdi et al. 2004; Abad and Bermejo 2007; Campos et al. 2016), leaves of *B. dracunculifolia* (Fukuda et al. 2006; Campos et al. 2016) and *B. pseudotenuifolia* (Moreira et al. 2003a, b; Abad and Bermejo 2007). The methylation of position C-7 leads to sakuranetin (**2**) accumulated in the aerial parts of *B. concinna* (Wollenweber et al. 2006), *B. marginalis* (Faine et al. 1987), *B. retusa* (Herz et al. 1977; Grecco et al. 2012a, b, 2014a, b; Rodriguez et al. 2012; Taguchi et al. 2015a, b; Sakoda et al. 2016; Bittencourt-Mernak et al. 2017; Ueno et al. 2018), *B. salicifolia* (del Corral et al. 2012; Campos et al. 2016), *B. serulata* (Bohlmann et al. 1981a), and *B. trinervis* var. *rhexioides* (Bohlmann et al. 1979). This compound was also found in the roots of *B. leptoccephala* (Bohlmann et al. 1981a) and *B. intermixta* (Bohlmann et al. 1981a) and leaves of *B. teindalensis*. The methylation at position C-4' rather than C-7 of naringenin (**1**) affords iso-sakuranetin (**3**) identified in aerial parts of *B. alaternoides* (Bohlmann et al. 1979), *B. conferta* (Weimann et al. 2002), *B. dracunculifolia*, within its roots and vegetative gems (da Silva Filho et al. 2004, 2008; Park et al. 2004; de Alencar et al. 2005; Lemos et al. 2007; Missima et al. 2007; de Sousa et al. 2009; Guimaraes et al. 2012; Figueiredo-Rinhel et al. 2013; Campos et al. 2016) and in *B. polyccephala* (Davila et al. 2013), in the roots of *B. leptoccephala* (Bohlmann et al. 1981b), and in the

Table 11.1 Flavanones identified in parts of species from genus *Baccharis*

	Compounds	Species	Parts	References
1	Naringenin	<i>B. alaternoides</i>	Above ground	Bohlmann et al. (1979)
		<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
		<i>B. ligustrina</i>	Aerial parts	Moreira et al. (2003a, b) and Abad and Bermejo (2007)
		<i>B. polycephala</i>	Aerial parts	Davila et al. (2013)
		<i>B. retusa</i>	Aerial parts	Campos et al. (2016) and Grecco et al. (2012a)
		<i>B. salzmannii</i>	Aerial parts	Bohlmann et al. (1981a) and Campos et al. (2016)
		<i>B. varians</i>	Aerial parts	Bohlmann et al. (1981a)
		<i>B. illinita</i>	Flowers	Verdi et al. (2004), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. dracunculifolia</i>	Leaves	Fukuda et al. (2006) and Campos et al. (2016)
		<i>B. pseudotenuifolia</i>	Leaves, shrub	Moreira et al. (2003a, b), Abad and Bermejo (2007) and Campos et al. (2016)
2	Sakuranetin	<i>B. concinna</i>	Aerial (leaves and stems)	Wollenweber et al. (2006)
		<i>B. marginalis</i>	Aerial parts	Faine et al. (1987)
		<i>B. retusa</i>	Aerial parts	Herz et al. (1977), Grecco et al. (2012b, 2014a, b), Toledo et al. (2013), Taguchi et al. (2015a, b), Sakoda et al. (2016), Bittencourt-Mernak et al. (2017) and Ueno et al. (2018)
		<i>B. salicifolia</i>	Aerial parts	Campos et al. (2016) and del Corral et al. (2012)
		<i>B. serrulata</i>	Aerial parts	Bohlmann et al. (1981a)
		<i>B. trinervis</i> var. <i>rhexioides</i>	Above ground	Bohlmann et al. (1979)
		<i>B. leptocephala</i>	Roots	Bohlmann et al. (1981b)
		<i>B. intermixta</i>	Roots	Bohlmann et al. (1981a)
		<i>B. teindalensis</i>	Leaves	Vidari et al. (2003)
3	Isosakuranetin	<i>B. alaternoides</i>	Above ground	Bohlmann et al. (1979)
		<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
		<i>B. dracunculifolia</i>	Aerial, leaves, roots, and vegetative gems	da Silva Filho et al. (2004, 2008), Park et al. (2004), de Alencar et al. (2005), Lemos et al. (2007), Missima et al. (2007), de Sousa et al. (2009), Guimaraes et al. (2012), Figueiredo-Rinhel et al. (2013) and Campos et al. (2016)

(continued)

Table 11.1 (continued)

	Compounds	Species	Parts	References
		<i>B. polycephala</i>	Aerial parts	Davila et al. (2013)
		<i>B. leptocephala</i>	Roots	Bohlmann et al. (1981b)
		<i>B. leptophylla</i>	Shrub	Almanza et al. (2000) and Mollinedo et al. (2001)
4	5,6,7-trihydroxy-4'-methoxyflavanone	<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
		<i>B. retusa</i>	Aerial parts	Grecco et al. (2010a, b)
		<i>B. teindalensis</i>	Leaves	Vidari et al. (2003)
		<i>B. viminea</i>		Wollenweber et al. (1997)
5	5-hydroxy-4',7-dimethoxyflavanone	<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
		<i>B. polycephala</i>	Aerial parts	Davila et al. (2013)
6	4'-hydroxy-5,7-dimethoxyflavanone	<i>B. alaternoides</i>	Above ground	Bohlmann et al. (1979)
7	Pinocembrin	<i>B. concinna</i>	Roots	Bohlmann et al. (1981b)
		<i>B. oxyodonta</i>	Roots	Bohlmann et al. (1981b) and Abad Martinez et al. (2005)
		<i>B. viminea</i>		Wollenweber et al. (1997)
8	5,7-dihydroxy-3'-methoxyflavanone	<i>B. truncata</i>	Roots	Bohlmann et al. (1981b)
9	Dihydrooroxylin A	<i>B. uncinella</i>	Aerial parts	Campos et al. (2016)
10	Eriodictyol	<i>B. concinna</i>	Aerial parts	Wollenweber et al. (2006)
		<i>B. confertifolia</i>	Aerial parts	Wollenweber et al. (2006)
		<i>B. marginalis</i>	Aerial parts	Faine et al. (1987)
		<i>B. retusa</i>	Aerial parts	Campos et al. (2016) and Grecco et al. (2012a)
		<i>B. pseudotenuifolia</i>	Shrub, leaves	Moreira et al. (2003a, b), Abad and Bermejo (2007) and Campos et al. 2016)
11	Eriodictyol-7-methylether	<i>B. concinna</i>	Aerial parts	Wollenweber et al. (2006)
12	Homoeriodictyol	<i>B. calliprinos</i>	Aerial parts	Gianello et al. (1999)
13	Eriodictyol-3',4'-dimethylether	<i>B. calliprinos</i>	Aerial parts	Gianello et al. (1999)
14	Eriodictyol 7,3',4'-trimethyl ether	<i>B. confertifolia</i>	Aerial parts	Wollenweber et al. (2006)
15	Filifolin	<i>B. concinna</i>	Aerial parts	Wollenweber et al. (2006)
		<i>B. boliviensis</i>	Aerial parts	Campos et al. (2016)
16	8-methoxyeriodictyol	<i>B. concinna</i>	Aerial parts	Wollenweber et al. (2006)

Table 11.2 Flavanonols identified in parts of species from genus *Baccharis*

	Compounds	Species	Parts	References
17	Aromadendrin/ dihydrokaempferol	<i>B. dracunculifolia</i>	Leaves	Guimaraes et al. (2012)
		<i>B. pseudotenuifolia</i>	Shrub	Moreira et al. (2003a, b), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. illinita</i>	Leaves and flowers	Verdi et al. (2004), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. retusa</i>	Aerial parts	Campos et al. (2016) and Grecco et al. (2012a)
18	Aromadendrin-7-methyl ether	<i>B. dracunculifolia</i>	Aerial (leaves and stems)	Campos et al. (2016)
		<i>B. illinita</i>	Aerial (leaves and stems)	Campos et al. (2016) and Pizzolatti et al. (2006)
19	Dihydrokaempferide	<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
		<i>B. dracunculifolia</i>	Leaves, roots, flowers, buds, and stems	Figueiredo-Rinhel et al. (2013), da Silva Filho et al. (2004, 2008), Missima et al. (2007), Lemos et al. (2007); de Sousa et al. (2009, 2011), Resende et al. (2007), Rezende et al. (2014), Cestari et al. (2011), Midorikawa et al. (2003) and Kumazawa et al. (2003)
		<i>B. leptophylla</i>		Almanza et al. (2000) and Mollinedo et al. (2001)
20	3-acetoxy-4',5,7-trihydroxyflavanone	<i>B. varians</i>	Aerial parts	(Bohlmann et al. (1981a) and Abad Martinez et al. (2005))
21	3-acetoxy-4',5-dihydroxy-7-methoxyflavanone	<i>B. dracunculifolia</i>	Leaves	Campos et al. (2016) and Fukuda et al. (2006)
22	Pinobanksin	<i>B. oxyodonta</i>	Roots	Bohlmann et al. (1981b) and Abad Martinez et al. (2005)
		<i>B. dracunculifolia</i>	Leaves and vegetative gems	Park et al. (2004) and de Alencar et al. (2005)
23	Pinobanksin-3-acetate	<i>B. dracunculifolia</i>	Leaf bud	Park et al. (2005)
		<i>B. trinervis</i>		Jakupovic et al. (1986)
24	Taxifolin	<i>B. illinita</i>	Flowers	Verdi et al. (2004), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. retusa</i>	Aerial parts	Campos et al. (2016) and Grecco et al. (2012a)
25	Taxifolin-4'-methyl ether	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
26	8-prenyltaxifolin	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
27	Taxifolin-3-acetate	<i>B. varians</i>	Aerial parts	Bohlmann et al. (1981a) and Abad Martinez et al. (2005)

Table 11.3 Flavones identified in parts of species from genus *Baccharis*

	Compounds	Species	Parts	References
28	Apigenin	<i>B. bigelovii</i>	Aerial parts	Arriaga-Giner et al. (1986)
		<i>B. crispa</i>	Aerial parts	Palacios et al. (1983)
		<i>B. dentata</i>	Aerial parts	Sartor et al. (2013) and Campos et al. (2016)
		<i>B. gaudichaudiana</i>	Aerial parts	Fullas et al. (1994), Guo et al. (2007), Visintini et al. (2013) and Campos et al. (2016)
		<i>B. notosergila</i>	Aerial parts	Palacios et al. (1983)
		<i>B. pedicellata</i>	Aerial parts	Faine et al. (1987)
		<i>B. retusa</i>	Aerial parts	Campos et al. (2016) and Grecco et al. (2012a)
		<i>B. salicifolia</i>	Aerial parts	Bohlmann et al. (1981b), Wollenweber et al. (1986) and Campos et al. (2016)
		<i>B. salzmannii</i>	Aerial parts	Campos et al. (2016)
		<i>B. trimera</i>	Aerial parts and epigaeous parts	Soicke and Leng-Eschlow (1987), Nakasugi and Komai (1998) and Nakasugi (1990)
		<i>B. illinita</i>	Flowers	Verdi et al. (2004), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. pseudotenuifolia</i>	Leaves and shrub	Moreira et al. (2003a, b), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. teindalensis</i>	Leaves	Vidari et al. (2003)
		<i>B. dracunculifolia</i>	Vegetative gems	de Alencar et al. (2005)
		<i>B. genistelloides</i>		Kuroyanagi et al. (1985), Abad Martinez et al. (2005) and Hennig et al. (2011)
		<i>B. heterophylla</i>		Wollenweber et al. (1986)
		<i>B. pteronioides</i>		Wollenweber et al. (1986)
		<i>B. ramosissima</i>		(Bohlmann et al. 1981b)
		<i>B. tola</i>		San Martin et al. (1982, 1983) and Abad Martinez et al. (2005)

(continued)

Table 11.3 (continued)

	Compounds	Species	Parts	References
		<i>B. trinervis</i>		Arriaga et al. (1982)
		<i>B. vaccinioides</i>		Wollenweber et al. (1986)
		<i>B. viminea</i>		Wollenweber et al. (1997)
29	Isoscutellarein	<i>B. pilularis</i>		Wollenweber et al. (1997)
30	Acacetin	<i>B. articulata</i>	Aerial parts	Gianello and Giordano (1984) and Cariddi et al. (2012)
		<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
		<i>B. polyccephala</i>	Aerial parts	Davila et al. (2013)
		<i>B. dracunculifolia</i>	Leaves	Park et al. (2004) and da Silva Filho et al. (2009)
		<i>B. rhomboidalis</i>	Leaves	Silva et al. (1971)
		<i>B. grandicapitulata</i>		Bohlmann et al. (1985)
		<i>B. patagonica</i>		Zdero et al. (1986) and Rivera et al. (1988)
		<i>B. salicifolia</i>		Wollenweber et al. (1986)
		<i>B. trinervis</i>		Arriaga et al. (1982)
		<i>B. vaccinioides</i>		Wollenweber et al. (1986)
		<i>B. viminea</i>		Wollenweber et al. (1997)
31	Genkwanin	<i>B. crispa</i>	Aerial parts	Palacios et al. (1983)
		<i>B. notosergila</i>	Aerial parts	Palacios et al. (1983)
		<i>B. pedicellata</i>	Aerial parts	Faine et al. (1987)
		<i>B. trimera</i>	Aerial parts	Soicke and Leng-Eschlow (1987) and Nakasugi and Komai (1998)
		<i>B. trinervis</i>	Leaves	Herrera et al. (1996)
		<i>B. genistelloides</i>		Abad Martinez et al. (2005) and Hennig et al. (2011)
		<i>B. pilularis</i> var. <i>consanguinea</i>		Wollenweber et al. (1997)
32	Thevetiaflavone	<i>B. gaudichaudiana</i>	Aerial parts	Guo et al. (2007) and Campos et al. (2016)
33	7-hydroxy-5,4'-dimethoxyflavone	<i>B. articulata</i>		de Oliveira et al. (2014)
		<i>B. uesterii</i>	Aerial parts	de Oliveira et al. (2014)

(continued)

Table 11.3 (continued)

	Compounds	Species	Parts	References
34	5-hydroxy-4',7-dimethoxyflavone	<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
		<i>B. crispa</i>	Aerial parts	Bandoni et al. (1978)
		<i>B. illinita</i>	Aerial parts	Arriaga-Giner et al. (1986) and Campos et al. (2016)
		<i>B. rhomboidalis</i>	Leaves	Silva et al. (1971)
		<i>B. teindalensis</i>	Leaves	Vidari et al. (2003)
		<i>B. tola</i>	Leaves	San Martin et al. (1982)
35	Chrysins	<i>B. dracunculifolia</i>	Leaves	Paula et al. (2017) and Park et al. (2004)
		<i>B. viminea</i>		Wollenweber et al. (1997)
36	7-methylchrysins	<i>B. viminea</i>		Wollenweber et al. (1997)
37	2'-methoxychrysins	<i>B. illinita</i>	Flowers	Verdi et al. (2004), Abad and Bermejo (2007) and Campos et al. (2016)
38	Hispidulin	<i>B. flabellata</i>	Aerial parts	Saad et al. (1988)
		<i>B. gaudichaudiana</i>	Aerial parts	Campos et al. (2016), Fullas et al. (1994) and Akaike et al. (2003)
		<i>B. genistelloides</i>	Aerial parts	Daily et al. (1984) and San Martin et al. (2012)
		<i>B. grisebachii</i>	Aerial parts	Abad and Bermejo (2007) and Tapia et al. (2004)
		<i>B. halimifolia</i>	Aerial parts	Joshi et al. (1997) and Jakupovic et al. (1990)
		<i>B. ligustrina</i>	Aerial parts	Abad and Bermejo (2007), Moreira et al. (2003a, b) and Nogueira et al. (2016)
		<i>B. magellanica</i>	Aerial parts	Cordano et al. (1982)
		<i>B. rhomboidalis</i>	Aerial parts	Cordano et al. (1982)
		<i>B. trimera</i>	Aerial and epigeous	Nakasugi and Komai (1998), Soicke and Leng-Eschlow (1987), Nakasugi (1990) and Padua et al. 2014)
	<i>B. uncinella</i>	Aerial parts		Campos et al. (2016) and Grecco et al. (2010a, b, 2014a, b)

(continued)

Table 11.3 (continued)

	Compounds	Species	Parts	References
		<i>B. pseudotenuifolia</i>	Leaves and shrub	Moreira et al. (2003a, b), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. vaccinioides</i>		Wollenweber et al. (1986)
39	Pectolinaringenin	<i>B. concava</i>	Aerial parts	Zamorano et al. (1987)
		<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
		<i>B. decussata</i>	Aerial parts	Morales Mendez et al. (1984) and Rojas and Morales (2000)
		<i>B. grisebachii</i>	Aerial parts	Gianello and Giordano (1987), Abad Martinez et al. (2005) and Feresin et al. (2003)
		<i>B. uncinella</i>	Aerial parts	Grecco et al. (2010a, b, 2014a, b), Passero et al. (2011), Zalewski et al. (2011) and Campos et al. (2016)
		<i>B. pedunculata</i>	Leaf	Rahalison et al. (1995)
		<i>B. trinervis</i>	Branches	Sharp et al. (2000) and Abad Martinez et al. (2005)
		<i>B. macraei</i>	Fresh	Faini et al. (1991)
		<i>B. pilularis var. consanguinea</i>		Wollenweber et al. (1997)
		<i>B. vaccinioides</i>		Wollenweber et al. (1986)
40	Cirsimarin	<i>B. concava</i>	Aerial parts	Zamorano et al. (1987)
		<i>B. concinna</i>	Aerial parts	Wollenweber et al. (2006)
		<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
		<i>B. genistelloides</i>	Aerial parts	Suttisri et al. (1994), Abad Martinez et al. (2005) and Hennig et al. (2011)
		<i>B. rhomboidalis</i>	Aerial parts	Labbe et al. (1986)
		<i>B. rufescens</i> var. <i>rufescens</i>	Aerial parts	Simirgiotis et al. (2003), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. trimera</i>	Aerial parts	Nakasugi (1990) and Nakasugi and Komai (1998)

(continued)

Table 11.3 (continued)

	Compounds	Species	Parts	References
		<i>B. elaeagnoides</i>		Mesquita et al. (1985)
		<i>B. halimifolia</i>		Joshi et al. (1997)
		<i>B. intermedia</i>		Faini et al. (1991)
		<i>B. macraei</i>		Faini et al. (1991)
		<i>B. pilularis</i> var. <i>consanguinea</i>		Wollenweber et al. (1997)
		<i>B. tricuneata</i>		Wagner et al. (1978)
41	Scutellarein-7,4'-dimethyl ether	<i>B. tucumanensis</i>	Aerial parts	Tonn et al. (1982)
42	Salvigenin	<i>B. concava</i>	Aerial parts	Zamorano et al. (1987)
		<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
		<i>B. rhomboidalis</i>	Aerial parts	Labbe et al. (1986)
		<i>B. scandens</i>	Aerial parts	Cabrera et al. (2016)
		<i>B. trinervis</i>	Branches	Sharp et al. (2000) and Abad Martinez et al. (2005)
		<i>B. macraei</i>		Faini et al. (1991)
		<i>B. pilularis</i>		Wollenweber et al. (1997)
43	Desmethoxysudachitin	<i>B. grisebachii</i>	Aerial parts	Gianello and Giordano (1987), Tapia et al. (2004), Abad Martinez et al. (2005) and Campos et al. (2016)
		<i>B. solieri</i>	Aerial parts	Labbe et al. (1986)
44	Nevadensin	<i>B. decussata</i>	Aerial parts	Rojas and Morales (2000)
		<i>B. grisebachii</i>	Aerial parts	Gianello and Giordano (1987), Feresin et al. (2003), Tapia et al. (2004) and Abad Martinez et al. (2005)
		<i>B. nitida</i>	Aerial parts	Chidiak et al. (2007) and Campos et al. (2016)
45	Xantomicrol	<i>B. boliviensis</i>	Aerial parts	Calle et al. (2012) and Campos et al. (2016)
		<i>B. nitida</i>	Aerial parts	Chidiak et al. (2007) and Campos et al. (2016)
		<i>B. patens</i>	Aerial parts	Silva et al. (1985)
		<i>B. scandens</i>	Aerial parts	Cabrera et al. (2016)
		<i>B. tucumanensis</i>	Aerial parts	Tonn et al. (1982)
		<i>B. pentlandii</i>	Leaves	Tarqui et al. (2012)
		<i>B. quitenensis</i>	Roots	Bohlmann et al. (1981b)
46	Gardenin B	<i>B. grisebachii</i>	Aerial parts	Feresin et al. (2003), Tapia et al. (2004) and Campos et al. (2016)

(continued)

Table 11.3 (continued)

	Compounds	Species	Parts	References
		<i>B. scandens</i>	Aerial parts	Cabrera et al. (2016)
47	Tangeretin	<i>B. illinita</i>	Flowers	Verdi et al. (2004), Abad and Bermejo (2007) and Campos et al. (2016)
48	Luteolin	<i>B. articulata</i>	Aerial parts	Cariddi et al. (2012)
		<i>B. bigelovii</i>	Aerial parts	Arriaga-Giner et al. (1986) and Wollenweber et al. (1986)
		<i>B. concinna</i>	Aerial parts	Wollenweber et al. (2006)
		<i>B. confertifolia</i>	Aerial parts	Wollenweber et al. (2006)
		<i>B. gaudichaudiana</i>	Aerial parts	Guo et al. (2007)
		<i>B. genistelloides</i>	Aerial parts	San Martin et al. (2012)
		<i>B. incarum</i>	Aerial parts	Zampini et al. (2009) and Campos et al. (2016)
		<i>B. linearis</i>	Aerial parts	Wollenweber et al. (2006)
		<i>B. lycioides</i>	Aerial parts	Wollenweber et al. (2006)
		<i>B. reticularia</i>	Aerial parts	Bohlmann et al. (1981a)
		<i>B. trimera</i>	Aerial parts	Soicke and Leng-Eschlow (1987), da Silva et al. (2016) and Menezes et al. (2016)
		<i>B. trinervis</i>	Aerial parts	Jaramillo-Garcia et al. (2018)
		<i>B. varians</i>	Aerial parts	Bohlmann et al. (1981a)
		<i>B. illinita</i>	Flowers	Verdi et al. (2004), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. halimifolia</i>		Wollenweber et al. (1997)
		<i>B. microcephala</i>		Bohlmann et al. (1985)
		<i>B. nitida</i>		Bohlmann et al. (1985)
		<i>B. pteronioides</i>		Wollenweber et al. (1986)
49	Chrysoeriol	<i>B. salicifolia</i>	Aerial parts	Warning et al. (1986)
		<i>B. pseudotenuifolia</i>	Leaves and shrub	Moreira et al. (2003a, b), Abad and Bermejo (2007) and Campos et al. (2016)

(continued)

Table 11.3 (continued)

	Compounds	Species	Parts	References
		<i>B. illinita</i>	Flowers	Verdi et al. (2004), Abad and Bermejo (2007) and Campos et al. (2016)
50	Luteolin-4',7-dimethylether	<i>B. trimera</i>	Aerial parts	Padua et al. (2014) and de Araujo et al. (2016)
		<i>B. trinervis</i>	Leaves	Herrera et al. (1996)
51	Luteolin-3',7-dimethylether	<i>B. calliprinos</i>	Aerial parts	Gianello et al. (1999)
		<i>B. rhetinodes</i>	Aerial parts	Gianello et al. (1999)
		<i>B. salicifolia</i>	Aerial parts	Warning et al. (1986)
52	5-hydroxy-7,3',4'-trimethoxyflavone, luteolin-3',4',7-trimethylether	<i>B. crispa</i>	Aerial parts	Bandoni et al. (1978)
		<i>B. latifolia</i>	Aerial parts	Salcedo et al. (2003) and Campos et al. (2016)
		<i>B. trinervis</i>	Leaves	Rivera et al. (1988)
53	6-hydroxyluteolin	<i>B. boliviensis</i>	Aerial parts	Abad and Bermejo (2007)
54	Nepetin; eupafolin	<i>B. concinna</i>	Aerial parts	Verdi et al. (2004)
		<i>B. confertifolia</i>	Aerial parts	Verdi et al. (2004)
		<i>B. flabellata</i>	Aerial parts	Bandoni et al. (1978)
		<i>B. gaudichaudiana</i>	Aerial parts	Salcedo Ortiz et al. (2001) and Abad and Bermejo (2007)
		<i>B. genistelloides</i>	Aerial parts	San Martin et al. (2012)
		<i>B. linearis</i>	Aerial parts	He (1995), He et al. (1996) and Wollenweber et al. (2006)
		<i>B. lycooides</i>	Aerial parts	Wollenweber et al. (2006)
		<i>B. trimera</i>	Aerial parts	Soicke and Leng-Eschlow (1987) and Simões-Pires et al. (2005)
55	Jaceosidin	<i>B. concinna</i>	Aerial parts	Wollenweber et al. (2006)
		<i>B. flabellata</i>	Aerial parts	Saad et al. (1988)
		<i>B. gaudichaudiana</i>	Aerial parts	Akaike et al. (2003) and Campos et al. (2016)
		<i>B. grisebachii</i>	Aerial parts	Tapia et al. (2004)
56	Desmethoxycentaureidin	<i>B. gaudichaudiana</i>	Aerial parts	Akaike et al. (2003) and Campos et al. 2016)
		<i>B. solieri</i>	Stems and leaves	Labbe et al. (1986)
		<i>B. petiolata</i>		Labbé et al. (1990)
		<i>B. salicina</i>		Parodi and Fischer (1988) and Quijano et al. (1998)

(continued)

Table 11.3 (continued)

	Compounds	Species	Parts	References
57	Cirsiliol	<i>B. concinna</i>	Leaves and stems	Wollenweber et al. (2006)
		<i>B. confertifolia</i>	Leaves and stems	Wollenweber et al. (2006)
		<i>B. flabellata</i>	Aerial parts	Saad et al. (1988)
		<i>B. linearis</i>	Leaves and stems	Wollenweber et al. (2006)
		<i>B. rufescens</i> var. <i>rufescens</i>	Aerial parts	Simirgiotis et al. (2003), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. genistelloides</i>		Kuroyanagi et al. (1985), Abad Martinez et al. (2005) and Hennig et al. (2011)
58	Cirsilineol	<i>B. concinna</i>	Aerial parts	Wollenweber et al. (2006)
		<i>B. salicifolia</i>	Aerial parts	Warning et al. (1986)
		<i>B. genistelloides</i>		Abad Martinez et al. (2005) and Hennig et al. (2011)
59	5,6-Dihydroxy-3',4',7-trimethoxyflavone	<i>B. trimera</i>	All plants	Borella et al. (2006)
60	Eupatilin	<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
		<i>B. gaudichaudiana</i>	Aerial parts	Akaike et al. (2003) and Campos et al. (2016)
61	Eupatorin	<i>B. genistelloides</i>	Aerial parts	Suttisri et al. (1994) and Hennig et al. (2011)
		<i>B. trimera</i>	Leaves, stems, flowers, and fruits	Herz et al. (1977), de Mello and Petrovick (2000), Torres et al. (2000), da Silva et al. (2004), Silva et al. (2006), Padua et al. (2014) and de Araujo et al. (2016)
62	5-hydroxy-3',4',6,7-tetramethoxyflavone	<i>B. genistelloides</i>	Aerial parts	Suttisri et al. (1994)
		<i>B. trimera</i>	Leaves	Rendon and Vila (1995) and Silva et al. (2006)
63	Sideritiflavone	<i>B. patens</i>	Aerial parts	Silva et al. (1985)
		<i>B. thymifolia</i>	Aerial parts	Saad et al. (1987)
		<i>B. pentlandii</i>	Leaves	Tarqui et al. (2012) and Campos et al. (2016)

(continued)

Table 11.3 (continued)

	Compounds	Species	Parts	References
64	4',5-dihydroxy-3',6,7,8-tetramethoxyflavone, 7-methylsudachitin, 3'-methoxyxanthomicrol	<i>B. incarum</i>	Aerial parts	Faini et al. (1982a, b)
		<i>B. salicifolia</i>	Aerial parts	Warning et al. (1986)
		<i>B. thymifolia</i>	Aerial parts	Saad et al. (1987)
		<i>B. pentlandii</i>	Leaves	Tarqui et al. (2012) and Campos et al. (2016)
		<i>B. oxyodonta</i>	Roots	Bohlmann et al. (1981b)
65	Gardenin D	<i>B. patens</i>	Aerial parts	Silva et al. (1985)
		<i>B. thymifolia</i>	Aerial parts	Saad et al. (1987)
67	Nobiletin	<i>B. illinita</i>	Flowers	Abad and Bermejo (2007) and Campos et al. (2016)
68	Isoschaftoside	<i>B. gaudichaudiana</i>	Aerial parts	Akaike et al. (2003 and Campos et al. (2016))
69	Vicenin II	<i>B. trimera</i>	Aerial parts	Rabelo et al. (2017)
		<i>B. trimera</i>	Aerial parts	Rabelo et al. (2017)
70	6(8)-C-furanosyl-8(6)-C-hexosyl flavone	<i>B. trimera</i>	Aerial parts	de Araujo et al. (2016)
71	6(8)-C- hexosyl-8(6)-C-furanosyl flavone	<i>B. trimera</i>	Aerial parts	de Araujo et al. (2016) and Rabelo et al. (2017)
72	5,3'-dihydroxy-4'-methoxy-7-O-pyranosyl-furanosyl flavone	<i>B. trimera</i>	Aerial parts	Padua et al. (2014)

shrub *B. leptophylla*. The C-6 hydroxylation of isosakuranetin (**3**) leads to the formation of 5,6,7-trihydroxy-4'-methoxyflavanone (**4**) accumulated in the leaves and aerial parts of *B. conferta* (Weimann et al. 2002), *B. retusa* (Grecco et al. 2010a, b), *B. teindalensis* (Vidari et al. 2003), and *B. viminea* (Wollenweber et al. 1997). The dimethoxy derivative of naringenin (**1**), 5-hydroxy-4',7-dimethoxyflavanone (**5**), was identified in the aerial parts of *B. conferta* (Weimann et al. 2002) and *B. polyccephala* (Davila et al. 2013) and 4'-hydroxy-5,7-dimethoxyflavanone (**6**) in ground parts of *B. alaternoides* (Bohlmann et al. 1979). The 4'-dehydroxylated derivative, pinocembrin (**7**), was identified in *B. viminea* (Wollenweber et al. 1997), in the roots of *B. concinna* (Bohlmann et al. 1981b), and in *B. oxyodonta* (Bohlmann et al. 1981b; Abad Martinez et al. 2005). Less frequently isolated 3'- and 6-monomethoxylated derivatives of pinocembrin (**7**): 5,7-dihydroxy-3'-methoxyflavanone (**8**) and 5,7-dihydroxy-6-methoxyflavanone, known as dihydrooroxylin A (**9**), were identified in the roots of *B. truncata* [27] and aerial parts of *B. uncinella*, respectively (Grecco et al. 2010a, b; Campos et al. 2016).

Table 11.4 Flavanols identified in parts of species from genus *Baccharis*

	Compounds	Species	Parts	References	
73	Kaempferol	<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)	
		<i>B. dentata</i>	Aerial parts	Sartor et al. (2013) and Campos et al. (2016)	
		<i>B. gaudichaudiana</i>	Aerial parts	Campos et al. (2016)	
		<i>B. polyccephala</i>	Aerial parts	Davila et al. (2013)	
		<i>B. retusa</i>	Aerial parts	Grecco et al. (2012a) and Campos et al. (2016)	
		<i>B. illinita</i>	Flowers	Verdi et al. (2004), Abad and Bermejo (2007) and Campos et al. (2016)	
		<i>B. maritima</i>	Flowering tops	Moreira et al. (1975)	
		<i>B. pseudotenuifolia</i>	Shrub, leaves	Moreira et al. (2003a, b), Abad and Bermejo (2007) and Campos et al. (2016)	
		<i>B. trimera</i>	Leaves	da Silva et al. (2016)	
		<i>B. dracunculifolia</i>	Leaves, buds, stems and vegetative gems	Midorikawa et al. (2003), Park et al. (2004, 2005) and de Alencar et al. (2005)	
74	Isokaempferide	<i>B. genistelloides</i>		Daily et al. (1984)	
		<i>B. pilularis</i> var. <i>consanguinea</i>		Wollenweber et al. (1997)	
		<i>B. linearis</i>	Aerial parts	Wollenweber et al. (2006) and Faini et al. (1999)	
		<i>B. lycioides</i>	Aerial parts	Wollenweber et al. (2006)	
		<i>B. pedicellata</i>	Aerial parts	Faine et al. (1987)	

(continued)

Table 11.4 (continued)

	Compounds	Species	Parts	References
75	Kaempferide	<i>B. papillosa</i>	Leaves	Campos et al. (2016) and Escobar et al. (2009)
		<i>B. intermedia</i>		Faini et al. (1991)
		<i>B. linearis</i>		Faini et al. (1991)
		<i>B. macraei</i>		Faini et al. (1991)
		<i>B. pilularis</i> var. <i>consanguinea</i>		Wollenweber et al. (1997)
		<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
75	Kaempferide	<i>B. polycephala</i>	Aerial parts	Davila et al. (2013)
		<i>B. dracunculifolia</i>	Flowers, leaves, buds, stems and vegetative gems	Midorikawa et al. (2003), Park et al. (2004, 2005), de Alencar et al. (2005), Piantino et al. (2008), Campos et al. (2016) and Paula et al. (2017)
		<i>B. leptophylla</i>		Almanza et al. (2000) and Mollinedo et al. (2001)
		<i>B. pilularis</i> var. <i>consanguinea</i>		Wollenweber et al. (1997)
		<i>B. viminea</i>		Wollenweber et al. (1997)
76	Kaempferol-7-methyleter	<i>B. pilularis</i> var. <i>consanguinea</i>		Wollenweber et al. (1997)
77	Ermanin	<i>B. papillosa</i>	Leaves	Campos et al. (2016)
		<i>B. dracunculifolia</i>	Buds; leaves; stems	Midorikawa et al. (2003) and da Silva Filho et al. (2009)
		<i>B. pilularis</i> var. <i>consanguinea</i>		Wollenweber et al. (1997)
		<i>B. viminea</i>		Wollenweber et al. (1997)
78	Kaempferol-7,4'-dimethyl ether	<i>B. crispa</i>	Aerial parts	Bandoni et al. (1978)

(continued)

Table 11.4 (continued)

	Compounds	Species	Parts	References
		<i>B. Illinita</i>	Leaves and stems	Campos et al. (2016) and Pizzolatti et al. (2006)
		<i>B. teindalensis</i>	Leaves	Vidari et al. (2003)
79	Kaempferol-3,7-dimethylether	<i>B. pedicellata</i>	Aerial parts	Faine et al. (1987)
		<i>B. santelicensis</i>	Aerial parts	Zdero et al. (1991)
		<i>B. pilularis var. consanguinea</i>		Wollenweber et al. (1997)
80	Kaempferol-3,4',7-trimethylether	<i>B. illinita</i>	Aerial parts	Campos et al. (2016) and Pizzolatti et al. (2006)
		<i>B. santelicensis</i>	Aerial parts	Zdero et al. (1991)
81	Galangin	<i>B. dracunculifolia</i>	Leaf bud	Park et al. (2005)
		<i>B. viminea</i>		Wollenweber et al. (1997)
82	Galangin-7-methylether	<i>B. viminea</i>		Wollenweber et al. (1997)
83	6-hydroxykaempferol	<i>B. pilularis var. consanguinea</i>		Wollenweber et al. (1997)
84	6-methoxykaempferol	<i>B. dracunculifolia</i>	Leaves	Kumazawa et al. (2003)
85	4',6-dimethoxykaempferol	<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
86	6,7-dimethoxykaempferol	<i>B. pilularis var. consanguinea</i>		Wollenweber et al. (1997)
87	Penduletin	<i>B. pedunculata</i>	Leaves	Rahalison et al. (1995)
		<i>B. trinervis</i>	Branches	Sharp et al. (2000) and Abad Martinez et al. (2005)
		<i>B. salicifolia</i>		Wollenweber et al. (1986)
		<i>B. sarothroides</i>		Wollenweber et al. (1986)
		<i>B. vaccinioides</i>		Wollenweber et al. (1986)
88	Herbacetin-3-methyl ether	<i>B. pilularis var. consanguinea</i>		Wollenweber et al. (1997)
89	Quercetin	<i>B. articulata</i>	Aerial parts	Campos et al. (2016)

(continued)

Table 11.4 (continued)

Compounds	Species	Parts	References
	<i>B. bigelovii</i>	Aerial parts	Wollenweber et al. (1986) and Grecco et al. (2014a, b)
	<i>B. concinna</i>	Aerial parts	Wollenweber et al. (2006)
	<i>B. confertifolia</i>	Aerial parts	Wollenweber et al. (2006)
	<i>B. dentata</i>	Aerial parts	Sartor et al. (2013) and Campos et al. (2016)
	<i>B. genistelloides</i>	Aerial parts	Montes et al. (1971) and Morales Mendez et al. (1984)
	<i>B. grisebachii</i>	Aerial parts	Moreira et al. (1975)
	<i>B. linearis</i>	Aerial parts	Wollenweber et al. (2006)
	<i>B. lycioides</i>	Aerial parts	Wollenweber et al. (2006)
	<i>B. pteronioides</i>	Aerial parts	Wollenweber et al. (1986) and Jakupovic et al. (1990)
	<i>B. retusa</i>	Aerial parts	Grecco et al. (2012a), Ferreira et al. (2015) and Campos et al. (2016)
	<i>B. salicifolia</i>	Aerial parts	Wollenweber et al. (1986)
	<i>B. scandens</i>	Aerial parts	Cabrera et al. (2016)
	<i>B. thesioides</i>	Aerial parts	Liu et al. (1993)
	<i>B. trimera</i>	Aerial parts	Soicke and Leng-Eschlow (1987), Simões-Pires et al. (2005), Padua et al. (2014), da Silva et al. (2016), de Araujo et al. (2016), Menezes et al. (2016) and Sabir et al. (2017)

(continued)

Table 11.4 (continued)

	Compounds	Species	Parts	References
		<i>B. trinervis</i>	Aerial parts	Jaramillo-Garcia et al. (2018)
		<i>B. dracunculifolia</i>	Leaf bud and flowers	Midorikawa et al. (2003) and Park et al. (2005)
		<i>B. pseudotenuifolia</i>	Leaves and shrubs	
		<i>B. spicata</i>	Leaves, stems, and flowers	Moreira et al. (2003a, b), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. Illinita</i>	Flowers	Verdi et al. (2004), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. maritima</i>	Flowering tops	Moreira et al. (1975)
90	Rhamnetin	<i>B. confertifolia</i>	Leaves and stems	Wollenweber et al. (2006)
		<i>B. pilularis var. consanguinea</i>		Wollenweber et al. (1997)
91	Isorhamnetin	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
		<i>B. linearis</i>	Leaves and stems	Wollenweber et al. (2006) and He (1995)
		<i>B. lycioides</i>	Leaves and stems	Wollenweber et al. (2006)
		<i>B. pseudotenuifolia</i>	Leaves and shrub	
		<i>B. viminea</i>		Wollenweber et al. (1997)
92	3-methylquercetin	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
		<i>B. trimera</i>	Aerial parts	de Mello and Petrovick (2000)
		<i>B. linearis</i>	Leaves	Faini et al. (1999)
		<i>B. papillosa</i>	Leaves	Campos et al. (2016)

(continued)

Table 11.4 (continued)

	Compounds	Species	Parts	References
93	Rhamnazin	<i>B. illinita</i>	Flowers	Verdi et al. (2004), Abad and Bermejo (2007) and Campos et al. (2016)
		<i>B. halimifolia</i>		Wollenweber et al. (1997)
		<i>B. intermedia</i>		Faini et al. (1991)
		<i>B. linearis</i>		Faini et al. (1991)
94	Isorhamnetin 3-methyl ether	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
		<i>B. confertifolia</i>	Leaves and stems	Wollenweber et al. (2006)
		<i>B. latifolia</i>		Salcedo Ortiz et al. (2001)
94	Isorhamnetin 3-methyl ether	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
		<i>B. linearis</i>	Leaves and stems	Wollenweber et al. (2006) and Faini et al. (1999)
		<i>B. lycioides</i>	Leaves and stems	Wollenweber et al. (2006)
		<i>B. intermedia</i>		Faini et al. (1991)
		<i>B. linearis</i>		Faini et al. (1991)
95	3,4'-dimethoxy-3',5,7-trihydroxyflavone	<i>B. sarothroides</i>		Abad Martinez et al. (2005)
96	3,7-dimethoxyquercetin	<i>B. triangularis</i>	Aerial parts	Gianello and Giordano (1989)
		<i>B. pilularis var. consanguinea</i>		Wollenweber et al. (1997)
97	3,7-dimethyl-isorhamnetin	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
98	3,7,4'-trimethylquercetin	<i>B. illinita</i>	Leaves and stems	Campos et al. (2016, 46)
		<i>B. grisebachii</i>		Abad and Bermejo (2007)
99	3,5-dihydroxy-7,3',4'-trimethoxyflavone	<i>B. latifolia</i>		Salcedo Ortiz et al. (2001)
100	Retusin	<i>B. illinita</i>	Leaves and stems	Campos et al. (2016, 46)

(continued)

Table 11.4 (continued)

	Compounds	Species	Parts	References
101	3-hydroxy-5,7,3',4'-tetramethoxyflavone	<i>B. latifolia</i>	Aerial parts	Campos et al. (2016, 121)
		<i>B. conferta</i>	Aerial parts	Weimann et al. (2002)
102	Patuletin	<i>B. concinna</i>	Leaves and stems	Wollenweber et al. (2006)
		<i>B. confertifolia</i>	Leaves and stems	Wollenweber et al. (2006)
		<i>B. halimifolia</i>		Wollenweber et al. (1997)
		<i>B. pilularis</i> var. <i>consanguinea</i>		Wollenweber et al. (1997)
103	3'-methylpatuletin	<i>B. halimifolia</i>		Wollenweber et al. (1997)
104	Axillarin	<i>B. incarum</i>	Leaves and stems	Campos et al. (2016)
		<i>B. linearis</i>	Leaves and stems	Wollenweber et al. (2006) and Labbe et al. (1986)
		<i>B. solieri</i>	Leaves and stems	Labbe et al. (1986)
		<i>B. halimifolia</i>		Wollenweber et al. (1997)
105	Eupatolitin	<i>B. confertifolia</i>	Leaves and stems	Wollenweber et al. (2006)
		<i>B. gaudichaudiana</i>	Aerial parts	Akaike et al. (2003)
106	Centaureidin	<i>B. sarothrodes</i>	Leaves and twigs	Abad Martinezt al. (2005) and Montes et al. (1971)
		<i>B. solieri</i>	Leaves and stems	Labbe et al. (1986)
		<i>B. salicina</i>		Parodi and Fischer (1988) and Quijano et al. (1998)
107	5,7-dihydroxy-3,6,3',4'-tetramethoxyflavone	<i>B. salicina</i>		Quijano et al. (1998)
108	3,3'-dimethylgossypetin	<i>B. linearis</i>	Leaves and stems	Wollenweber et al. (2006)
109	3,8-dimethylgossypetin	<i>B. linearis</i>	Leaves and stems	Wollenweber et al. (2006)
110	3,8,3'-trimethylgossypetin	<i>B. linearis</i>	Leaves and stems	Wollenweber et al. (2006)

(continued)

Table 11.4 (continued)

	Compounds	Species	Parts	References
111	3',4',5,7-tetrahydroxy-3,6,8-trimethoxyflavone	<i>B. incarum</i>	Leaves and stems	Campos et al. (2016)
112	4',5,7-trihydroxy-3,3',6,8-tetramethoxyflavone	<i>B. incarum</i>	Leaves and stems	Campos et al. (2016)
		<i>B. solieri</i>	Leaves and stems	Labbe et al. (1986)
113	3',5-dihydroxy-3,4',6,7,8-pentamethoxyflavone	<i>B. boliviensis</i>	Aerial parts	Campos et al. (2016)
114	4',5-dihydroxy3',3,6,7,8-pentamethoxyflavone	<i>B. incarum</i>	Leaves and stems	Faini et al. (1982a, b), Nuño et al. (2012) and Campos et al. (2016)
115	3-O-acetylmyricetin	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
116	Myricetin 7,3'-dimethylether	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
117	Myricetin 7,3',5'-trimethylether	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
118	Myricetin 3',5',7,8-tetramethylether	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
119	6-hydroxy-7,3',5'-trimethylmyricetin	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
120	6-hydroxy-3,7,3',5'-tetramethylmyricetin	<i>B. tola</i>	Aerial parts	Simirgiotis et al. (2016)
121	Apigenin 3-O- β -D-glucopyranoside/ astragalin	<i>B. Dracunculifolia</i>	Aerial parts	Nagatani et al. (2001)
		<i>B. angustifolia</i>		Wagner et al. (1972)
122	Kaempferol-3-O-rutinoside/nicotiflorin	<i>B. antioquensis</i>	Leaves	Mejia-Giraldo et al. (2016)
123	Quercetin 3-O-[β -D-apiofuranosyl (1 \rightarrow 2) α -Lrhamnopyranosyl (1 \rightarrow 6)]- β -D-glucopyranoside	<i>B. thesioides</i>	Aerial parts	Liu et al. (1993)
124	Isoqueretin	<i>B. dracunculifolia</i>	Aerial parts	Nagatani et al. (2001)
		<i>B. thesioides</i>	Aerial parts	Kakehashu et al. (2016)
		<i>B. trimera</i>	Aerial parts	Simões-Pires et al. (2005)
		<i>B. pseudotenuifolia</i>	Leaves and shrub	Moreira et al. (2003a, b), Abad and Bermejo (2007) and Campos et al. (2016)

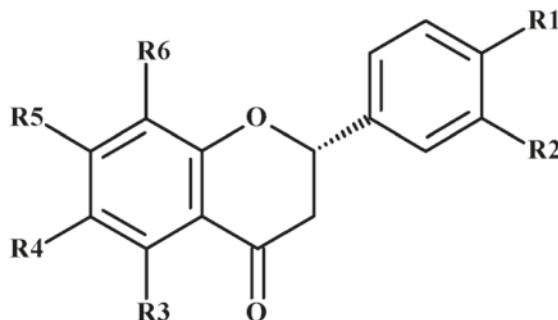
(continued)

Table 11.4 (continued)

	Compounds	Species	Parts	References
		<i>B. angustifolia</i>		Wagner et al. (1972)
		<i>B. ochracea</i>		Schenkel et al. (1997)
125	Hyperoside	<i>B. dracunculifolia</i>	Aerial parts	Nagatani et al. (2001)
		<i>B. thesioides</i>	Aerial parts	Nagatani et al. (2001)
126	Quercitrin	<i>B. gaudichaudiana</i>	Aerial parts	Campos et al. (2016) and Akaike et al. (2003)
		<i>B. microdonta</i>	Leaves	Toyama et al. (2014)
		<i>B. pseudotenuifolia</i>	Leaves and shrubs	Moreira et al. (2003a, b), Abad and Bermejo (2007) and Campos et al. (2016)
127	Rutin	<i>B. gaudichaudiana</i>	Aerial parts	Campos et al. (2016) and Akaike et al. (2003)
		<i>B. thesioides</i>	Aerial parts	Tarqui et al. (2012)
		<i>B. trinervis</i>	Aerial parts	Jaramillo-Garcia et al. (2018)
		<i>B. trimera</i>	Aerial parts and leaves	Gene et al. (1996), Menezes et al. (2016) and Sabir et al. (2017)
		<i>B. antioquensis</i>	Leaves	Mejia-Giraldo et al. (2016)
		<i>B. dentata</i>	Leaves and stems	Campos et al. (2016) and Sartor et al. (2013)
		<i>B. spicata</i>	Leaves, stems, and flowers	Agudelo et al. (2016)
		<i>B. genistelloides</i>		Hennig et al. (2011)
128	Quercetin -3-O-(4"-O-caffeyl)- rhamnopyranosyl-(1 → 6)- galactopyranoside – 174	<i>B. antioquensis</i>	Leaves	Mejia-Giraldo et al. (2016)

Eriodictyol (**10**) was found in aerial parts of *B. concinna* (Bohlmann et al. 1981b), *B. confertifolia* (Wollenweber et al. 2006), *B. marginalis* (Faine et al. 1987), *B. retusa* (Campos et al. 2016; Grecco et al. 2012a, b), and *B. pseudotenuifolia* (Moreira et al. 2003a, b; Abad and Bermejo 2007; Campos et al. 2016). Six eriodictyol derivatives were found in *Baccharis* species: eriodictyol-7-methyl ether (**11**) at *B. concinna* (Wollenweber et al. 2006), homoeriodictyol (**12**) and eriodictyol-3',4'-dimethyl ether (**13**) at *B. calliprinos* (Gianello et al. 1999), eriodictyol 7,3',4'-trimethyl ether (**14**) at *B. confertifolia* (Wollenweber et al. 2006), filifolin (**15**) at *B. concinna* (Wollenweber et al. 2006) and *B. boliviensis* (Campos et al. 2016), and 8-methoxyeriodictyol (**16**) at *B. concinna* (Wollenweber et al. 2006). Flavanones identified at *Baccharis* species are summarized in Table 11.1, followed by their respective structures in Fig. 11.5.

Fig. 11.5 Structures of flavanones **1–16** identified in *Baccharis* species



- 1** – R1 = R3 = R5 = OH; R2 = R4 = R6 = H
- 2** – R1 = R3 = OH; R5 = OMe; R2 = R4 = R6 = H
- 3** – R1 = OMe; R3 = R5 = OH; R2 = R4 = R6 = H
- 4** – R1 = OMe; R3 = R4 = R5 = OH; R2 = R6 = H;
- 5** – R1 = R5 = OMe; R3 = OH; R2 = R4 = R6 = H
- 6** – R1 = OH; R3 = R5 = OMe; R2 = R4 = R6 = H
- 7** – R3 = R5 = OH; R1 = R2 = R4 = R6 = H
- 8** – R2 = OMe; R3 = R5 = OH; R1 = R4 = R6 = H
- 9** – R3 = R5 = OH; R4 = OMe; R1 = R2 = R6 = H
- 10** – R1 = R2 = R3 = R5 = OH; R4 = R6 = H
- 11** – R1 = R2 = R3 = OH; R5 = OMe; R4 = R6 = H
- 12** – R1 = R3 = R5 = OH; R2 = OMe; R4 = R6 = H
- 13** – R1 = R2 = OMe; R3 = R5 = OH; R4 = R6 = H
- 14** – R1 = R2 = R5 = OMe; R3 = OH; R4 = R6 = H
- 15** – R1 = R2 = R3 = R5 = OH; R4 = OMe; R6 = H
- 16** – R1 = R2 = R3 = R5 = OH; R6 = OMe; R4 = H

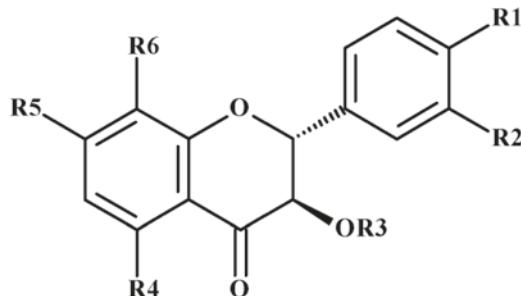
5 Flavanonols

The 3-hydroxylation of flavanones through 3-hydroxylase enzymes leads to the biosynthesis of flavanonol derivatives. Naringenin-3-hydroxylase affords aromadendrin, also known as dihydrokaempferol (**17**), identified in leaves of *B. dracunculifolia* (Guimaraes et al. 2012), shrubs of *B. pseudotenuifolia* (Moreira et al. 2003a, b; Abad and Bermejo 2007; Campos et al. 2016), leaves and flowers of *B. illinita* (Verdi et al. 2004; Abad and Bermejo 2007; Campos et al. 2016), and in aerial parts of *B. retusa* (Campos et al. 2016; Grecco et al. 2012a). Aromadendrin-7-methyl ether (**18**) was isolated from aerial parts of *B. dracunculifolia* (Campos et al. 2016) and *B. illinita* (Pizzolatti et al. 2006). Dihydrokaempferide (**19**) was identified in the aerial parts of *B. conferta* (Weimann et al. 2002), in leaves, roots, flowers, buds, and stems of *B. dracunculifolia* (Figueiredo-Rinhel et al. 2013; da Silva Filho et al. 2004, 2008; Missima et al. 2007; Lemos et al. 2007; de Sousa et al. 2009, 2011; Resende et al. 2007; Rezende et al. 2014; Cestari et al. 2011; Midorikawa et al. 2003; Kumazawa et al. 2003), and from *B. leptophylla* (Mollinedo et al. 2001; Almanza et al. 2000). 3-Acetylated derivative such as 3-acetoxy-4' 5 7-trihydroxyflavanone (**20**) was isolated from aerial parts of *B. varians* (Bohlmann et al. 1981a; Abad Martinez et al. 2005), while 3-acetoxy-4',5-dihydroxy-7-methoxyflavanone (**21**) was obtained from leaves of *B. dracunculifolia*, (Fukuda et al. 2006; Campos et al. 2016). A 4'-dehydroxylated derivative of aromadendrin, pinobanksin (**22**), was isolated from roots of *B. oxydonta* (Bohlmann et al. 1981b; Abad Martinez et al. 2005) as well as from vegetative gems and leaves of *B. dracunculifolia* (Park et al. 2004; de Alencar et al. 2005). Additionally, its acetylated derivative, pinobanksin-3-acetate (**23**), was identified in *B. dracunculifolia* leaf bud (Park et al. 2005) and in *B. trinervis* (Jakupovic et al. 1986). Taxifolin (**24**) was identified in *B. illinita* (Abad and Bermejo 2007), especially in flowers (Verdi et al. 2004; Campos et al. 2016), as well as in aerial parts of *B. retusa* (Grecco et al. 2012a; Campos et al. 2016). 3-hydroxyhesperetin or taxifolin-4'-methyl ether (**25**), together with 8-prenyltaxifolin (**26**), were identified in aerial parts of *B. tola* (Simirgiotis et al. 2016). Furthermore, an acetylated derivative, taxifolin-3-acetate (**27**), was identified in aerial parts of *B. varians* (Bohlmann et al. 1981a; Abad Martinez et al. 2005). Flavanonols identified in *Baccharis* species are summarized in Table 11.2, followed by their respective structures in Fig. 11.6, Flavanonols identified in *Baccharis* species are summarized in Table 11.3, followed by their respective structures in Fig. 11.6.

6 Flavones

The enzymatic dehydration of flavanone naringenin (**1**) through flavone synthase affords apigenin (**28**) identified in the aerial parts of *B. bigelovii* (Arriaga-Giner et al. 1986), *B. crispa* (Palacios et al. 1983), *B. dentata* (Campos et al. 2016), *B. gaudichaudiana* (Fullas et al. 1994; Guo et al. 2007; Visintini et al. 2013; Campos

Fig. 11.6 Structures of flavanono^ls **17–27** identified in *Baccharis* species



17 – R1 = R4 = R5 = OH; R2 = R3 = R6 = H

18 – R1 = R4 = OH; R5 = OMe; R2 = R3 = R6 = H

19 – R1 = OMe; R4 = R5 = OH; R2 = R3 = R6 = H

20 – R1 = R4 = R5 = OH; R3 = Acetyl; R2 = R6 = H

21 – R1 = R4 = OH; R3 = Acetyl; R5 = OMe; R2 = R6 = H

22 – R4 = R5 = OH; R1 = R2 = R3 = R6 = H

23 – R3 = Acetyl; R4 = R5 = OH; R1 = R2 = R6 = H

24 – R1 = R2 = R4 = R5 = OH; R3 = R6 = H

25 – R1 = OMe; R2 = R4 = R5 = OH; R3 = R6 = H

26 – R1 = R2 = R4 = R5 = OH; R6 = Prenyl; R3 = H

27 – R1 = R2 = R4 = R5 = OH; R3 = Acetyl; R6 = H

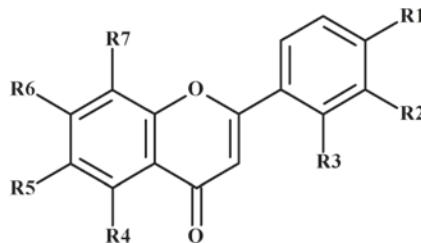
et al. 2016), *B. notosergila* (Palacios et al. 1983), *B. pedicellata* (Faine et al. 1987), *B. retusa* (Grecco et al. 2012a; Campos et al. 2016), *B. salicifolia* (Campos et al. 2016; Bohlmann et al. 1981b), *B. salzmannii* (Campos et al. 2016), as well as in the aerial and epigeous parts of *B. trimera* (Nakasugi 1990, 1998; Fullas et al. 1994). This compound was also identified in the flowers of *B. illinita* (Verdi et al. 2004; Abad and Bermejo 2007; Campos et al. 2016), leaves, and shrubs of *B. pseudotenuifolia* (Moreira et al. 2003a, b; Abad and Bermejo 2007; Campos et al. 2016) and *B. teindalensis* (Vidari et al. 2003), and vegetative gems of *B. dracunculifolia* (de Alencar et al. 2005). The flavone **28** was also found in *B. genistelloides* (Kuroyanagi et al. 1985; Abad Martinez et al. 2005; Hennig et al. 2011), *B. heterophylla* (Wollenweber et al. 1986), *B. pteronioides* (Wollenweber et al. 1986), *B. ramosissima* (Bohlmann et al. 1981b), *B. tola* (San Martin et al. 1982, 1983; Abad Martinez et al. 2005), *B. trinervis* (Arriaga et al. 1982), *B. vaccinioides* (Wollenweber et al. 1986), and *B. viminea* (Wollenweber et al. 1997). Its 8-hydroxylated derivative, isoscutellarein (**29**), was identified only in *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997), while the 4'-methylated derivative, acacetin (**30**), was found in aerial parts of *B. articulata* (Gianello and Giordano 1984; Cariddi et al. 2012), *B. conferta* (Weimann et al. 2002), and *B. polycephala* (Davila et al. 2013), in the leaf parts of *B. dracunculifolia* (Park et al. 2004; da Silva Filho et al. 2009)

and *B. rhomboidalis* (Silva et al. 1971) and from *B. grandicapitulata* (Bohlmann et al. 1985), *B. patagonica* (Zdero et al. 1986; Rivera et al. 1988); *B. salicifolia* (Wollenweber et al. 1986), *B. trinervis* (Arriaga et al. 1982), *B. vaccinioides* (Wollenweber et al. 1986), and *B. viminea* (Wollenweber et al. 1997). Also other two monomethylated derivatives were found: genkwanin (31) in aerial parts of *B. crispa* (Palacios et al. 1983), *B. notosergila* (Palacios et al. 1983), *B. pedicellata* (Faine et al. 1987), *B. trimera* (Nakasugi 1990; Nakasugi and Komai 1998), in leaves of *B. trinervis* (Herrera et al. 1996) as well as from *B. genistelloides* (Abad Martinez et al. 2005; Hennig et al. 2011) and *B. pilularis* (Wollenweber et al. 1997). Thevetiaflavone (32) was identified in aerial parts of *B. gaudichaudiana* (Guo et al. 2007; Campos et al. 2016). 4',5- and 4',7- dimethylated derivatives (33 and 34 derivatives, respectively) were identified in some *Baccharis* species, 7-Hydroxy-5,4'-dimethoxyflavone (33) was isolated from aerial parts of *B. articulata* (de Oliveira et al. 2014) and *B. uesterii* (Salcedo Ortiz et al. 2001), while 5-Hydroxy-4',7-dimethoxyflavone (34), from aerial parts of *B. conferta* (Weimann et al. 2002), *B. crispa* (Bandoni et al. 1978), *B. illinita* (Pizzolatti et al. 2006; Campos et al. 2016), from leaves of *B. rhomboidalis* (Silva et al. 1971), *B. teindalensis* (Vidari et al. 2003), *B. tola* (San Martin et al. 1982), *B. trinervis* (Herrera et al. 1996), and from *B. latifolia* (Salcedo Ortiz et al. 2001). Flavones biosynthesized from phenylalanine-cinnamic acid-pinocembrin were also identified: chrysin (35) and 7-methylchrysin (36) from leaves of *B. dracunculifolia* (Park et al. 2004; Paula et al. 2017) and *B. viminea* (Wollenweber et al. 1997). Its 2'-methoxylated derivative, 2'-methoxychrysin (37), was also identified from flowers of *B. illinita* (Verdi et al. 2004; Abad and Bermejo 2007; Campos et al. 2016). Hispidulin (38) was identified in aerial parts of *B. flabellata* (Saad et al. 1988), *B. gaudichaudiana* (Fullas et al. 1994; Akaike et al. 2003; Campos et al. 2016), *B. genistelloides* (Daily et al. 1984; San Martin et al. 2012), *B. grisebachii* (Tapia et al. 2004; Abad and Bermejo 2007), *B. halimifolia* (Joshi et al. 1997; Jakupovic et al. 1990), *B. ligustrina* (Moreira et al. 2003a, b; Abad and Bermejo 2007; Nogueira Sobrinho et al. 2016), *B. magellanica* (Cordano et al. 1982), *B. rhomboidalis* (Labbe et al. 1986), *B. trimera* (Soicke and Leng-Eschlow 1987; Nakasugi 1990; Nakasugi and Komai 1998; Padua et al. 2014), and *B. uncinella* (Grecco et al. 2010a, b, 2014a, b; Campos et al. 2016). Flavonoid 38 was also identified in leaves and shrub of *B. pseudotenuifolia* (Moreira et al. 2003a, b; Abad and Bermejo 2007; Campos et al. 2016) and nonspecified parts of *B. vaccinioides* (Wollenweber et al. 1986). Pectolinaringenin (39) was identified in aerial parts of *B. concava* (Zamorano et al. 1987), *B. conferta* (Weimann et al. 2002), *B. decussata* (Morales Mendez et al. 1984; Rojas and Morales 2000), *B. grisebachii* (Gianello and Giordano 1987; Feresin et al. 2003; Abad Martinez et al. 2005), and *B. uncinella* (Grecco et al. 2010a, b, 2014a, b; Passero et al. 2011; Zalewski et al. 2011; Campos et al. 2016). Flavonoid (39) was also detected in the leaves of *B. pedunculata* (Rahalison et al. 1995), branches of *B. trinervis* (Sharp et al. 2000; Abad Martinez et al. 2005) [44, 103], and nonspecified parts of fresh *B. macraei* (Faini et al. 1991) [104], *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997) [41] and *B. vaccinioides* (Wollenweber et al. 1986). Cirsimaritin (40) was identified in the aerial parts of *B. concava* (Zamorano et al. 1987), *B. concinna*

(Wollenweber et al. 2006), *B. conferta* (Weimann et al. 2002), *B. genistelloides* (Suttisri et al. 1994; Abad Martinez et al. 2005; Hennig et al. 2011), *B. rhomboidalis* (Labbe et al. 1986), *B. rufescens* var. *rufescens* (Simirgiotis et al. 2003; Abad and Bermejo 2007; Campos et al. 2016), *B. trimera* (Nakasugi 1990; Nakasugi and Komai 1998), *B. elaeagnoides* (Mesquita et al. 1985), *B. halimifolia* (Joshi et al. 1997), *B. intermedia* and *B. macraei* (Faini et al. 1991), *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997), and *B. tricuneata* (Wagner et al. 1978). Scutellarein-7,4'-dimethyl ether (**41**) was identified from aerial parts of *B. tucumanensis* (Tonn et al. 1982), while salvigenin (**42**) was found in the aerial parts of *B. concava* (Zamorano et al. 1987), *B. conferta* (Weimann et al. 2002), *B. rhomboidalis* (Labbe et al. 1986), and *B. scandens* (Cabrera et al. 2016). Flavonoid **42** was identified in the branches of *B. trinervis* (Sharp et al. 2000; Abad Martinez et al. 2005), and in nonspecified parts of *B. macraei* (Faini et al. 1991) and *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997). Desmethoxysudachitin (**43**) was identified in the aerial parts of *B. grisebachii* (Gianello and Giordano 1987; Tapia et al. 2004; Grecco et al. 2010a, b; Campos et al. 2016) and *B. solieri* (Labbe et al. 1986). Its 4'-methylated derivative, nevadensin (**44**), was found in the aerial parts of *B. decussata* (Rojas and Morales 2000), *B. grisebachii* (Gianello and Giordano 1987; Feresin et al. 2003; Tapia et al. 2004; Grecco et al. 2010a, b), and *B. nitida* (Chidiak et al. 2007; Campos et al. 2016). A methylated derivative of **43** in position C-7 afforded xantomicrol (**45**) found in aerial parts of *B. boliviensis* (Calle et al. 2012; Campos et al. 2016), *B. nitida* (Chidiak et al. 2007; Campos et al. 2016), *B. patens* (Silva et al. 1985), *B. scandens* (Cabrera et al. 2016), and *B. tucumanensis* (Tonn et al. 1982). Compound **45** was also found in the leaves of *B. pentlandii* (Tarqui et al. 2012) and roots of *B. quitensis* (Bohlmann et al. 1981b). Gardenin B (**46**) was identified in the aerial parts of *B. grisebachii* (Feresin et al. 2003; Tapia et al. 2004; Campos et al. 2016) and *B. scandens* (Cabrera et al. 2016). A pentamethoxylated derivative, tangeretin (**47**), was found in the flowers of *B. illinita* (Verdi et al. 2004; Abad and Bermejo 2007; Campos et al. 2016). The flavone luteolin (**48**), derived from dehydration of flavanone eriodictyol, was identified in aerial parts of *B. articulata* (Cariddi et al. 2012), *B. bigelovii* (Arriaga-Giner et al. 1986; Wollenweber et al. 1986), *B. concinna* (Wollenweber et al. 2006), *B. confertifolia* (Wollenweber et al. 2006), *B. gaudichaudiana* (Guo et al. 2007), *B. genistelloides* (San Martin et al. 2012), *B. incarum* (Zampini et al. 2009; Campos et al. 2016), *B. linearis* (Wollenweber et al. 2006), *B. lycooides* (Wollenweber et al. 2006), *B. reticularia* (Bohlmann et al. 1981a), *B. trimera* (Soicke and Leng-Eschlow 1987; da Silva et al. 2016; Menezes et al. 2016), *B. trinervis* (Jaramillo-Garcia et al. 2018), and *B. varians* (Bohlmann et al. 1981a). Furthermore, flavonoid **48** was isolated from flowers of *B. illinita* (Verdi et al. 2004; Abad and Bermejo 2007; Campos et al. 2016), and nonspecified parts of *B. halimifolia* (Wollenweber et al. 1997), *B. microcephala* (Bohlmann et al. 1985), *B. nitida* (Bohlmann et al. 1985), and *B. pteronioides* (Wollenweber et al. 1986). Methylated derivatives of luteolin, biosynthesized through luteolin-methyltransferase, were found in several studied *Baccharis* species. A monomethylated (C-3') derivative, chrysoeriol (**49**), was identified in the flowers of *B. illinita* (Verdi et al. 2004; Abad and Bermejo 2007; Campos et al.

2016), shrub of *B. pseudotenuifolia* (Moreira et al. 2003a, b; Abad and Bermejo 2007; Campos et al. 2016), and aerial parts of *B. salicifolia* (Warning et al. 1986). 4',7- and 3',7- Dimethylated derivatives of Luteolin were also found in *Baccharis* genus, luteolin-4',7-dimethylether (**50**) in aerial parts of *B. trimera* (Padua et al. 2014; de Araujo et al. 2016), and leaves of *B. trinervis* (Herrera et al. 1996) and luteolin-3',7-dimethylether (**51**) in aerial parts of *B. calliprinos* (Gianello et al. 1999), *B. rhetinodes* (Gianello et al. 1999), and *B. salicifolia* (Warning et al. 1986). The trimethylated derivative, 5-hydroxy-7,3',4'-trimethoxyflavone/luteolin-3',4',7-trimethylether (**52**), was identified in the aerial parts of *B. crispa* (Bandoni et al. 1978) [80], *B. latifolia* (Salcedo et al. 2003; Campos et al. 2016), and in the leaves of *B. trinervis* (Herrera et al. 1996). 6-Hydroxyluteolin (**53**) was found only in aerial parts of *B. boliviensis* (Campos et al. 2016), while its methylated derivatives were identified in several *Baccharis* species; nepetin, also known as eupafolin (**54**), was identified in aerial parts of *B. concinna* (Wollenweber et al. 2006), *B. confertifolia* (Wollenweber et al. 2006), *B. flabellata* (Saad et al. 1988), *B. gaudichaudiana* (Akaike et al. 2003; Campos et al. 2016), *B. genistelloides* (San Martin et al. 2012), *B. linearis* (He 1995; He et al. 1996; Wollenweber et al. 2006), *B. lycioides* (Wollenweber et al. 2006), and *B. trimera* (Soicke and Leng-Eschlow 1987; Simões-Pires et al. 2005). Jaceosidin (**55**) was isolated from aerial parts of *B. concinna* (Wollenweber et al. 2006), *B. flabellata* (Saad et al. 1988), *B. gaudichaudiana* (Akaike et al. 2003; Campos et al. 2016), and *B. grisebachii* (Tapia et al. 2004). Desmethoxycentaureidin (**56**) was obtained from aerial parts of *B. gaudichaudiana* (Akaike et al. 2003; Campos et al. 2016), *B. solieri* (Labbe et al. 1986), and non-specified parts of *B. petiolata* (Labbé et al. 1990) and *B. salicina* (Parodi and Fischer 1988; Quijano et al. 1998). Cirsiliol (**57**) was identified in the aerial parts of *B. concinna* (Wollenweber et al. 2006), *B. confertifolia* (Wollenweber et al. 2006), *B. flabellata* (Saad et al. 1988), *B. linearis* (Wollenweber et al. 2006), *B. rufescens* var. *rufescens* (Simirgiotis et al. 2003; Abad and Bermejo 2007; Campos et al. 2016), and in nonspecified parts of *B. genistelloides* (Kuroyanagi et al. 1985; Abad Martinez et al. 2005; Hennig et al. 2011) and *B. tricuneata*. Cirsilineol (**58**) was detected in aerial parts of *B. concinna* (Wollenweber et al. 2006), *B. salicifolia* (Warning et al. 1986), and nonspecified parts of *B. genistelloides* (Abad Martinez et al. 2005; Hennig et al. 2011), while 5,6-dihydroxy-3',4',7-trimethoxyflavone (**59**) from young parts of *B. trimera* (Borella et al. 2006). The aerial parts of *B. conferta* (Weimann et al. 2002) and *B. gaudichaudiana* (Akaike et al. 2003; Campos et al. 2016) afforded eupatilin (**60**), while aerial parts of *B. genistelloides* (Suttisri et al. 1994; Hennig et al. 2011) and leaves, stems, flowers, and fruits of *B. trimera* (Herz et al. 1977; de Mello and Petrovick 2000; Torres et al. 2000; da Silva et al. 2004; Silva et al. 2006; Padua et al. 2014; de Araujo et al. 2016) afforded eupatorin (**61**). 5-Hydroxy-3',4',6,7-tetramethoxyflavone (**62**) was identified from aerial parts of *B. genistelloides* (Suttisri et al. 1994) and leaves of *B. trimera* (Rendon and Vila 1995; Silva et al. 2006). Sideritiflavone (**63**) was detected in aerial parts of *B. patens* (Silva et al. 1985) and *B. thymifolia* (Saad et al. 1987) as well as from leaves of *B. pentlandii* (Tarqui et al. 2012; Campos et al. 2016); 4',5-Dihydroxy-3',6,7,8-tetramethoxyflavone (**64**), known as 7-methylsudachitin or 3'-methoxyxanthomictrol, was identified from aerial parts of *B. incarum* (Faini et al. 1982a,b), *B. salicifolia*

(Warning et al. 1986) and *B. thymifolia* (Saad et al. 1987), from leaves of *B. pentlandii* (Tarqui et al. 2012; Campos et al. 2016), and roots of *B. oxyodonta* and *B. quitensis* (Bohlmann et al. 1981b). Gardenin D (**65**) was identified in aerial parts of *B. patens* (Silva et al. 1985). A pentamethoxylated derivative, 5-hydroxy-3',4',6,7,8-pentamethoxytlavone (**66**), was detected only in aerial parts of *B. thymifolia* (Saad et al. 1987). Nobletin was identified only in the flowers of *B. illinita* (**67**). Although in less distribution, flavone glycosides were found in *Baccharis* species: isoschaftoside (**68**) was identified in the aerial parts of *B. gaudichaudiana* (Akaike et al. 2003; Campos et al. 2016), while vicenin II (**69**), 6(8)-C-furanosyl-8(6)-C-hexosyl flavone (**70**), 6(8)-C-hexosyl-8(6)-C-furanosyl flavone (**71**), and 5,3'-dihydroxy-4'-methoxy-7-O-pyranosyl-furanosyl flavone (**72**) were detected in the aerial parts of *B. trimera* (Padua et al. 2014; de Araujo et al. 2016; Rabelo et al. 2017). Flavones identified in *Baccharis* species are summarized in Table 11.3, followed by their respective structures in Fig. 11.7.



28 - R1 = R4 = R6 = OH; R2 = R3 = R5 = R7 = H	29 - R1 = R4 = R6 = R7 = OH; R2 = R3 = R5 = H
30 - R1 = OMe; R4 = R6 = OH; R2 = R3 = R5 = R7 = H	31 - R1 = R4 = OH; R6 = OMe; R2 = R3 = R5 = R7 = H
32 - R1 = R6 = OH; R4 = OMe; R2 = R3 = R5 = R7 = H	33 - R1 = R4 = OMe; R6 = OH; R2 = R3 = R5 = R7 = H
34 - R1 = R6 = OMe; R4 = OH; R2 = R3 = R5 = R7 = H	35 - R4 = R6 = OH; R1 = R2 = R3 = R5 = R7 = H
36 - R4 = OH; R6 = OMe; R1 = R2 = R3 = R5 = R7 = H	37 - R3 = OMe; R4 = R6 = OH; R1 = R2 = R5 = R7 = H
38 - R1 = R4 = R6 = OH; R5 = OMe; R2 = R3 = R7 = H	39 - R1 = R5 = OMe; R4 = R6 = OH; R2 = R3 = R7 = H
40 - R1 = R4 = OH; R5 = R6 = OMe; R2 = R3 = R7 = H	41 - R1 = R6 = OMe; R4 = R5 = OH; R2 = R3 = R7 = H
42 - R1 = R5 = R6 = OMe; R4 = OH; R2 = R3 = R7 = H	43 - R1 = R4 = R6 = OH; R5 = R7 = OMe; R2 = R3 = H
44 - R1 = R5 = R7 = OMe; R4 = R6 = OH; R2 = R3 = H	45 - R1 = R4 = OH; R5 = R6 = R7 = OMe; R2 = R3 = H
46 - R1 = R5 = R6 = R7 = OMe; R4 = OH; R2 = R3 = H	47 - R1 = R4 = R5 = R6 = R7 = OMe; R2 = R3 = H
48 - R1 = R2 = R4 = R6 = OH; R3 = R5 = R7 = H	49 - R1 = R4 = R6 = OH; R2 = OMe; R3 = R5 = R7 = H
50 - R1 = R6 = OMe; R2 = R4 = OH; R3 = R5 = R7 = H	51 - R1 = R4 = OH; R2 = R6 = OMe; R3 = R5 = R7 = H
52 - R1 = R2 = R6 = OMe; R4 = OH; R3 = R5 = R7 = H	53 - R1 = R2 = R4 = R5 = R6 = OH; R3 = R7 = H
54 - R1 = R2 = R4 = R6 = OH; R5 = OMe; R3 = R7 = H	55 - R1 = R4 = R6 = OH; R2 = R5 = OMe; R3 = R7 = H
56 - R1 = R5 = OMe; R2 = R4 = R6 = OH; R3 = R7 = H	57 - R1 = R2 = R4 = OH; R5 = R6 = OMe; R3 = R7 = H
58 - R1 = R6 = OMe; R2 = R4 = R5 = OH; R3 = R7 = H	59 - R1 = R2 = R6 = OMe; R4 = R5 = OH; R3 = R7 = H
60 - R1 = R2 = R5 = OMe; R4 = R6 = OH; R3 = R7 = H	61 - R1 = R5 = R6 = OMe; R2 = R4 = OH; R3 = R7 = H
62 - R1 = R2 = R5 = R6 = OMe; R4 = OH; R3 = R7 = H	63 - R1 = R2 = R4 = OH; R5 = R6 = R7 = OMe; R3 = H
64 - R1 = R4 = OH; R2 = R5 = R6 = R7 = OMe; R3 = H	65 - R1 = R5 = R6 = R7 = OMe; R2 = R4 = OH; R3 = H
66 - R1 = R2 = R5 = R6 = R7 = OMe; R4 = OH; R3 = H	67 - R1 = R2 = R4 = R5 = R6 = R7 = OMe; R3 = H
68 - R1 = R4 = R6 = OH; R5 = Ara; R7 = Glu; R2 = R3 = H	
69 - R1 = R4 = R6 = OH; R5 = R7 = Glu; R2 = R3 = H	
70 - R5 = Furanosyl; R7 = Hexosyl; R1 = R2 = R3 = R4 = R6 = H	
71 - R5 = Hexosyl; R7 = Furanosyl; R1 = R2 = R3 = R4 = R6 = H	
72 - R1 = OMe; R2 = R4 = OH; R6 = O-Pyranosyl-Furanosyl; R3 = R5 = R7 = H	

Fig. 11.7 Structures of flavones **28–72** identified in *Baccharis* species

7 Flavanols

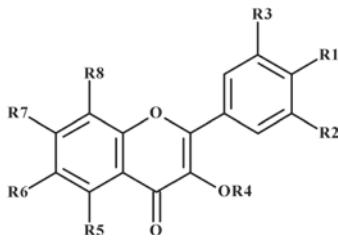
The biosynthesis of flavanols, the main group of flavonoids of genus *Baccharis*, occurs after the formation of flavanonols, through flavanol synthase enzyme (FLS). Thus, naringenin (**1**) is converted to aromadendrin/dihydrokaempferol (**17**) to finally afford the flavanol kaempferol (**73**). Compound **73** was identified in aerial parts of *B. conferta* (Weimann et al. 2002), *B. dentata* (Sartor et al. 2013; Campos et al. 2016), *B. gaudichaudiana* (Campos et al. 2016), *B. polycephala* (Davila et al. 2013), *B. trimera* (da Silva et al. 2016), and *B. retusa* (Campos et al. 2016; Grecco et al. 2012a). This compound was also identified in the flowers of *B. illinita* (Verdi et al. 2004; Abad and Bermejo 2007; Campos et al. 2016), flowering tops of *B. maritima* (Moreira et al. 1975), leaves of *B. pseudotenuifolia* (Moreira et al. 2003a, b; Campos et al. 2016), and buds, stems, and vegetative gems of *B. dracunculifolia* (Palacios et al. 1983; Park et al. 2004, 2005; de Alencar et al. 2005) and nonspecified parts of *B. genistelloides* (Daily et al. 1984), and *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997). The 3-methylation of kaempferol afforded isokaempferide (**74**) identified in the aerial parts of *B. linearis* (Faini et al. 1999; Wollenweber et al. 2006), *B. lycioides* (Wollenweber et al. 2006), and *B. pedicellata* (Faine et al. 1987). Furthermore, this compound was also identified in the leaves of *B. papillosa* (Escobar et al. 2009; Campos et al. 2016), and nonspecified parts of fresh *B. intermedia* (Faini et al. 1991), *B. linearis* (Faini et al. 1991), *B. macraei* (Faini et al. 1991), and *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997). The 4'-methylation affords kaempferide (**75**), found in aerial parts of *B. conferta* (Weimann et al. 2002) and *B. polycephala* (Davila et al. 2013) as well as in the flowers, leaves, buds, stems, and vegetative gems of *B. dracunculifolia* (Midorikawa et al. 2003; Park et al. 2004, 2005; de Alencar et al. 2005; Piantino et al. 2008; Campos et al. 2016; Paula et al. 2017), in no-nspecified parts of *B. leptophylla* (Almanza et al. 2000; Mollinedo et al. 2001) and in *B. pilularis* var. *consanguinea* and *B. viminea* (Wollenweber et al. 1997). Kaempferol-7-methylether (**76**) was identified only in *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997). The leaves of *B. papillosa* (Campos et al. 2016) afforded emanin (**77**), also identified in buds, leaves, and stems of *B. dracunculifolia* (Midorikawa et al. 2003; da Silva Filho et al. 2009) and from nonspecified parts of *B. pilularis* var. *consanguinea* and from *B. viminea* (Wollenweber et al. 1997). Kaempferol-7,4'-dimethyl ether (**78**) was detected in the aerial parts of *B. crispa* (Bandoni et al. 1978) and *B. illinita* (Pizzolatti et al. 2006; Campos et al. 2016) as well as from leaves of *B. teindalensis* (Vidari et al. 2003). Kaempferol-3,7-dimethyl ether (**79**) was identified in the aerial parts of *B. pedicellata* (Faine et al. 1987) and *B. santellicis* (Zdero et al. 1991) and nonspecified parts of *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997). The trimethylated derivative, kaempferol-3,4',7-trimethylether (**80**), was identified in the aerial parts of *B. illinita* (Pizzolatti et al. 2006; Campos et al. 2016) and *B. santellicis* (Zdero et al. 1991). Galangin (**81**) was detected in the leaf bud of *B. dracunculifolia* (Park et al. 2005) and nonspecified parts of *B. viminea* (Wollenweber et al. 1997), that also afforded galangin-7-methylether (**82**). 6-Hydroxylated kaempferol and its derivatives were identified in few *Baccharis* species, 6-hydroxykaempferol (**83**) in *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997); 6-methoxykaempferol (**84**) in leaves

of *B. dracunculifolia* (Kumazawa et al. 2003); 4',6-dimethoxykaempferol (**85**) in aerial parts of *B. conferta* (Weimann et al. 2002); and 6,7-dimethoxykaempferol (**86**) in *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997). Penduletin (**87**) was identified in leaves of *B. pedunculata* (Rahalison et al. 1995), branches of *B. trinervis* (Sharp et al. 2000; Abad Martinez et al. 2005), and nonspecified parts of *B. salicifolia*, *B. sarothroides* and *B. vaccinioides* (Wollenweber et al. 1986), while herbacetin-3-methylether (**88**) was isolated from *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997). Quercetin (**89**) was identified in the aerial parts of *B. articulata* (Campos et al. 2016), *B. bigelovii* (Arriaga-Giner et al. 1986; Wollenweber et al. 1986), *B. concinna* and *B. confertifolia* (Wollenweber et al. 2006), *B. dentata* (Sartor et al. 2013; Campos et al. 2016), *B. genistelloides* (Daily et al. 1984; San Martin et al. 2012), *B. grisebachii* (Tapia et al. 2004), *B. linearis* (Wollenweber et al. 2006), *B. lycioides* (Wollenweber et al. 2006), *B. pteronioides*, *B. retusa*, *B. salicifolia* (Wollenweber et al. 1986), *B. scandens* (Cabrera et al. 2016), *B. thesioides* (Liu et al. 1993), *B. trimera* (Soicke and Leng-Eschlow 1987; Simões-Pires et al. 2005; Padua et al. 2014; da Silva et al. 2016; de Araujo et al. 2016; Menezes et al. 2016; Sabir et al. 2017), *B. trinervis* (Jaramillo-Garcia et al. 2018), in the leaf bud of *B. dracunculifolia* (Park et al. 2005), leaves and shrubs of *B. pseudotenuifolia* (Moreira et al. 2003a, b; Abad and Bermejo 2007; Campos et al. 2016), leaves, stems, and flowers of *B. spicata* (Agudelo et al. 2016) flowers of *B. illinita* (Verdi et al. 2004; Abad and Bermejo 2007; Campos et al. 2016) and *B. maritima* (Moreira et al. 1975) as well as in the nonspecified parts of *B. viminea* (Wollenweber et al. 1997). Rhamnetin (**90**) was detected in leaves and stems of *B. confertifolia* (Wollenweber et al. 2006) and nonspecified parts of *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997), while isorhamnetin (**91**) was detected in the aerial parts of *B. tola*, leaves and stems of *B. linearis* (He 1995; Wollenweber et al. 2006) and *B. lycioides* (Wollenweber et al. 2006), leaves and shrubs of *B. pseudotenuifolia* (Moreira et al. 2003a, b; Abad and Bermejo 2007; Campos et al. 2016), and in nonspecified parts of *B. viminea* (Wollenweber et al. 1997). The 3-methylated derivative of quercetin, 3-methylquercetin (**92**), was identified in the aerial parts of *B. tola* (Simirgiotis et al. 2016) and *B. trimera* (de Mello and Petrovick 2000), in the leaves of *B. linearis* (Faini et al. 1999), flowers of *B. illinita* (Verdi et al. 2004; Abad and Bermejo 2007; Campos et al. 2016), fresh parts of *B. intermedia* and *B. linearis* (Faini et al. 1991) as well as in the nonspecified parts of *B. halimifolia* (Wollenweber et al. 1997). Rhamnazin (**93**) was found in the aerial parts of *B. tola* (Simirgiotis et al. 2016), leaves and stems of *B. confertifolia* (Wollenweber et al. 2006), and nonspecified parts of *B. latifolia* (Salcedo et al. 2001). Isorhamnetin 3-methylether (**94**) was identified in the aerial parts of *B. tola* (Simirgiotis et al. 2016), leaves and stems of *B. linearis* (Faini et al. 1999; Wollenweber et al. 2006) and *B. lycioides* (Wollenweber et al. 2006), and in the nonspecified parts of fresh *B. intermedia* and *B. linearis* (Faini et al. 1991). *B. sarothroides* afforded 3,4'-dimethoxy-3',5,7-trihydroxyflavone (**95**) (Abad Martinez et al. 2005), while 3,7-dimethoxyquercetin (**96**) was found in aerial parts of *B. triangularis* (Gianello and Giordano 1989) and nonspecified parts of *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997). 3,7-Dimethylisorhamnetin (**97**) was isolated from aerial parts of *B. tola* (Simirgiotis et al. 2016), while 3,7,4'-trimethylquercetin (**98**) was obtained from leaves and

stems of *B. illinita* (Campos et al. 2016) and nonspecified parts of *B. grisebachii* (Abad and Bermejo 2007). 3,5-dihydroxy-7,3',4'-trimethoxyflavone (**99**) was detected in *B. latifolia*. Retusin (**100**) was isolated from leaves and stems of *B. illinita* (Pizzolatti et al. 2006; Campos et al. 2016), while 3-hydroxy-5,7,3',4'-tetramethoxyflavone (**101**) was obtained from aerial parts of *B. latifolia* (Salcedo et al. 2003; Campos et al. 2016) and *B. conferta* (Weimann et al. 2002). Patuletin (**102**) was detected in leaves and stems of *B. concinna* and *B. confertifolia* (Wollenweber et al. 2006) as well as in nonspecified parts of *B. halimifolia* and *B. pilularis* var. *consanguinea* (Wollenweber et al. 1997). Flavonoid 3'-methylpatuletin (**103**) was detected in *B. halimifolia* (Wollenweber et al. 1997), while axillarin (**104**) was identified in the leaves and stems of *B. incarum* (Campos et al. 2016), *B. linearis* (Labbe et al. 1986; Wollenweber et al. 2006), and *B. solieri* (Labbe et al. 1986) as well as in the nonspecified parts of *B. halimifolia* (Wollenweber et al. 1997). Eupatolitin (**105**) was isolated from leaves and stems of *B. confertifolia* (Wollenweber et al. 2006) and aerial parts of *B. gaudichaudiana* (Akaike et al. 2003). Centaureidin (**106**) was identified in the leaves and twigs of *B. sarothroides* (Montes et al. 1971; Abad Martinez et al. 2005), leaves and stems of *B. solieri* (Labbe et al. 1986), and nonspecified parts of *B. salicina* (Parodi and Fischer 1988; Quijano et al. 1998). 5,7-Dihydroxy-3,6,3',4'-tetramethoxyflavone (**107**) was detected in *B. salicina* (Quijano et al. 1998), while three gossypetin derivatives (**108–110**) were identified in leaves and stems of *B. linearis* (Wollenweber et al. 2006): 3,3'-dimethyl- (**108**), 3,8-dimethyl- (**109**) and 3,8,3'-trimethyl- (**110**) derivatives. 3',4',5,7-Tetrahydroxy-3,6,8-trimethoxyflavone (**111**) and 4',5,7-trihydroxy-3,3',6,8-tetramethoxyflavone (**112**) were identified in the leaves and stems of *B. incarum* (Campos et al. 2016); compound **112** was also identified in the leaves and stems of *B. solieri* (Labbe et al. 1986). 3',5-Dihydroxy-3,4',6,7,8-pentamethoxyflavone (**113**) was detected in the aerial parts of *B. boliviensis* (Campos et al. 2016) and 4',5-dihydroxy3',3,6,7,8-pentamethoxyflavone (**114**) in the leaves and stems of *B. incarum* (Faini et al. 1982a, b; Nuño et al. 2012; Campos et al. 2016). Six myricetin derivatives (**115–120**) were detected in leaves and stems of *B. tola* (Simirgiotis et al. 2016): 3-O-acetyl- (**115**), 7,3'-dimethyl- (**116**), 7,3',5'-trimethyl- (**117**), 3',5',7,8-tetramethyl- (**118**), 6-hydroxy-7,3',5'-trimethyl- (**119**) and 6-hydroxy-3,7,3',5'-tetramethyl (**120**) derivatives. Additionally, nine flavonol glycosides (**121–129**) were identified in several species of *Baccharis*: apigenin 3-O- β -D-glucopyranoside, also known as astragalin (**121**), from aerial parts of *B. dracunculifolia* (Nagatani et al. 2001) and from *B. angustifolia* (Wagner et al. 1972), kaempferol-3-O-rutinoside, also known as nicotiflorin (**122**), from *B. antioquensis* (Mejia-Giraldo et al. 2016), quercetin 3-O-[β -D-apiofuranosyl-(1 → 2) α -Lrhamnopyranosyl-(1 → 6)]- β -D-glucopyranoside (**123**) from *B. thesioides* (Liu et al. 1993), isoquercetin (**124**) from aerial parts of *B. dracunculifolia* (Nagatani et al. 2001), *B. thesioides* (Kakehashu et al. 2016), *B. trimera* (Simões-Pires et al. 2005), *B. pseudotenuifolia* (Moreira et al. 2003a, b; Abad and Bermejo 2007; Campos et al. 2016), *B. angustifolia* (Wagner et al. 1972), and *B. ochracea* (Schenkel et al. 1997). Hyperoside (**125**) was detected in the aerial parts of *B. dracunculifolia* (Nagatani et al. 2001) and *B. thesioides* (Liu et al. 1993) [144], while quercitrin (**126**) was identified in *B. gaudichaudiana* (Akaike et al. 2003; Campos et al. 2016),

B. microdonta (Toyama et al. 2014), and *B. pseudotenuifolia* (Moreira et al. 2003a, b; Abad and Bermejo 2007; Campos et al. 2016). Rutin (**127**) was identified in the aerial parts of *B. gaudichaudiana* (Akaike et al. 2003; Campos et al. 2016), *B. thesioides* (Liu et al. 1993), *B. trinervis* (Jaramillo-Garcia et al. 2018), and *B. trimera* (Gene et al. 1996; Menezes et al. 2016; Sabir et al. 2017). Furthermore, flavonoid **127** was also isolated from leaves of *B. antioquensis* (Mejia-Giraldo et al. 2016), leaves and stems of *B. dentata* (Sartor et al. 2013; Campos et al. 2016), leaves, stems, and flowers *B. spicata* (Agudelo et al. 2016), and nonspecified parts of *B. genistelloides* (Hennig et al. 2011). Finally, quercetin-3-O-(4"-O-caffeyl)-rhamnopyranosyl-(1 → 6)-galactopyranoside (**128**) was identified in the leaves of *B. antioquensis* (Mejia-Giraldo et al. 2016). Flavanols identified in *Baccharis* species are summarized in Table 11.4, followed by their respective structures in Fig. 11.8.

Flavonoids display a significant role in biological functions, especially those that depend on the modulation of redox balance. Several studies describes the antioxidant and radical scavenger activities of extracts and isolated flavonoids of *Baccharis*, such as the antioxidant activities of isosakuranetin (**3**), dihydrokaempferide (**19**), kaempferol (**75**) (Mollinedo et al. 2001), isokaempferide (**74**), ermanin (**77**), 3-methylquercetin (**92**) (Escobar et al. 2009), luteolin (**48**), axillarin (**104**), 3',4',5,7-tetrahydroxy-3,6,8-trimethoxyflavone (**111**), 4',5,7-trihydroxy-3',6,8-tetramethoxyflavone (**112**), and 4',5-dihydroxy-3',3,6,7,8-pentamethoxyflavone (**114**). Antioxidant activities were also attributed to extracts of *B. spicata*, *B. tola*,



- 89 - R1= R2= R5= R7= OH; R3= R4= R6= R8= H
 91 - R1= R5= R7= OH; R2= OMe; R3= R4= R6= R8= H
 93 - R1= R5= OH; R2= R7= OMe; R3= R4= R6= R8= H
 95 - R1= OMe; R2= R5= R7= OH; R4= Me; R5= R6= R8= H
 97 - R1= R5= OH; R2= R7= OMe; R4= Me; R3= R6= R8= H
 99 - R1= R2= R7= OMe; R5= OH; R3= R4= R6= R8= H
 101 - R1= R2= R7= R5= OMe; R3= R4= R6= R8= H
 103 - R1= R5= R7= OH; R2= R6= OMe; R3= R4= R8= H
 105 - R1= R2= R5= OH; R6= R7= OMe; R3= R4= R8= H
 107 - R1= R2= R6= OMe; R5= R7= OH; R4= Me; R3= R8= H
 109 - R1= R2= R5= R7= OH; R4= Me; R8= OMe; R3= R6= H
 111 - R1= R2= R5= R7= OH; R4= Me; R6= R8= OMe; R3= H
 113 - R1= R6= R7= R8= OMe; R2= R5= OH; R4= Me; R3= H
 115 - R1= R2= R3= R5= R7= OH; R4= Acyl; R6= R8= H
 117 - R1= R5= OH; R2= R3= R7= OMe; R4= R6= R8= H
 119 - R1= R5= R6= OH; R2= R3= R7= OMe; R4= R8= H
 121 - R1= R5= R7= OH; R4= Glu; R2= R3= R6= R8= H
 123 - R1= R2= R5= R7= OH; R4= (Apio)-Rha-Glu; R3= R6= R8= H
 125 - R1= R2= R5= R7= OH; R4= Gal; R3= R6= R8= H
 127 - R1= R2= R5= R7= OH; R4= Rha-Glu; R3= R6= R8= H

- 73 - R1= R5= R7= OH; R2= R3= R4= R6= R8= H
 74 - R1= R5= R7= OH; R4= OMe; R2= R3= R6= R8= H
 75 - R1= OMe; R5 = R7 = OH; R2 = R3 = R4 = R6 = R8 = H
 76 - R1= R5= OH; R7= OMe; R2= R3= R4= R6= R8= H
 77 - R1= OMe; R4= Me; R5= R7= OH; R2= R3= R6= R8= H
 78 - R1= R7= OMe; R5= OH; R2= R3= R4= R6= R8= H
 79 - R4= Me; R7= OMe; R1= R5= OH; R2= R3= R6= R8= H
 80 - R1= R7= OMe; R4= Me; R5= OH; R2= R3= R6= R8= H
 81 - R1= R5= R7= OH; R1= R2= R3= R4= R6= R8= H
 82 - R5= OH; R7= OMe; R1= R2= R3= R4= R6= R8= H
 83 - R1= R5= R6= R7= OH; R2= R3= R4= R8= H
 84 - R1= R5= R7= OH; R6= OMe; R2= R3= R4= R8= H
 85 - R1= R6= OMe; R5= R7= OH; R2= R3= R4= R8= H
 86 - R1= R5= OH; R6= R7= OMe; R2= R3= R4= R8= H
 87 - R1= R5= OH; R4= Me; R6= R7= OMe; R2= R3= R4= R8= H
 88 - R1= R5= R7= R8= OH; R4= Me; R2= R3= R6= H
 90 - R1= R2= R5= R6= OH; R7= OMe; R3= R4= R6= R8= H
 92 - R1= R2= R5= R7= OH; R4= Me; R3= R6= R8= H
 94 - R1= R5= R7= OH; R2= OMe; R4= Me; R3= R6= R8= H
 96 - R1= R2= R5= OH; R4= Me; R7= OMe; R3= R6= R8= H
 98 - R1= R7= OMe; R2= R5= OH; R4= Me; R3= R6= R8= H
 100 - R1= R2= R7= OMe; R4= Me; R5= OH; R3= R6= R8= H
 102 - R1= R2= R5= R7= OH; R6= OMe; R3= R4= R8= H
 104 - R1= R2= R5= R7= OH; R4= Me; R6= OMe; R3= R8= H
 106 - R1= R6= OMe; R2= R5= R7= OH; R4= Me; R3= R8= H
 108 - R1= R5= R7= R8= OH; R2= OMe; R4= Me; R3= R6= H
 110 - R1= R5= R7= OH; R2= R8= OMe; R4= Me; R3= R6= H
 112 - R1= R5= R7= OH; R2= R6= R8= OMe; R4= Me; R3= H
 114 - R1= R5= OH; R2= R6= R7= R8= OMe; R4= Me; R3= H
 116 - R1= R3= R5= OH; R2= R7= OMe; R4= R6= R8= H
 118 - R1= R5= OH; R2= R3= R7= OMe; R8= Me; R4= R6= H
 120 - R1= R5= R6= OH; R2= R3= R7= OMe; R4= Me; R8= H
 122 - R1= R5= R7= OH; R4= Rha-Glu; R2= R3= R6= R8= H
 124 - R1= R2= R5= R7= OH; R4= Glu; R3= R6= R8= H
 126 - R1= R2= R5= R7= OH; R4= Rha; R3= R6= R8= H
 128 - R1= R2= R5= R7= OH; R4= Rha-Glu-caffeyl; R3= R6= R8= H

Fig. 11.8 Structures of flavanols **73–128** identified in *Baccharis* species Biological activities

and *B. trimera* (Agudelo et al. 2016; Simirgiotis et al. 2016; Sabir et al. 2017), while radical scavenger activity (DPPH, ROS) was attributed to *B. dentata* and *B. trimera* extracts (Sartor et al. 2013; Agudelo et al. 2016; Sabir et al. 2017). The last extract also inhibits ROS production through PKC and downregulates p47phox phosphorylation of NADPH oxidase in SK Hep-1 cells. Hispidulin (38), desmethoxyssudachitin (43), jaceosidin (55), and quercetin (89) suppress/scavenge erythrocyte lipoperoxidation, 38 and 89 superoxide anions, and 89 DPPH (Tapia et al. 2004). Eupafolin (54), quercitrin (127), and rutin (128) moderately scavenge DPPH radical (Akaike et al. 2003). Compounds 48, 104, 111, 112, and 114 also displayed antimicrobial activity against methicillin-resistant *Staphylococcus aureus* and *Enterococcus faecalis* (Zampini et al. 2009; Nuño et al. 2012). Antimicrobial activities were also attributed to some flavonoids: penduletin (87) presented antifungal activity against some human pathogenic and phytopathogenic fungi (Rahalison et al. 1995). Isosakuranetin (3) showed antifungal activities against *Neurospora crassa* (Almanza et al. 2000) and *Cryptococcus neoformans* (da Silva Filho et al. 2008); besides the antioxidant activity, previously described, compound 3 also demonstrated strong trypanocidal (da Silva Filho et al. 2004) and antiinflammatory activities (Figueiredo-Rinhel et al. 2013); both activities were also attributed to dihydrokaempferide (19). *B. dentata* extracts presented antibacterial activity against *Staphylococcus aureus* (Sartor et al. 2013), while *B. crispa* and *B. notosergila* ethanolic extracts presented antimicrobial activities against *Bacillus subtilis*, *Micrococcus luteus*, and *Staphylococcus aureus*, due to the presence of genkwanin (31) and apigenin (28). Sakuranetin (2) presented antifungal activity against pathogenic yeast belonging to the genus *Candida* (six species – *C. dubliniensis*, *C. tropicalis*, *C. glabrata*, *C. parapsilosis*, *C. krusei*, and *C. albicans*), *Cryptococcus* (two species/four serotypes – *C. neoformans* – serotypes A and D, *C. gattii* – serotypes B and *C. neoformans*), and *Saccharomyces cerevisiae* (Grecco et al. 2014a, b). Compound 2 demonstrated promising activities to prevent and treat several respiratory disorders, such as acute lung injury (Bittencourt-Mernak et al. 2017), chronic allergic pulmonary inflammation (Sakoda et al. 2016), emphysema, through regulation of NF-κB, oxidative stress, and metalloproteinases (Taguchi et al. 2015a, b), and reverses airway inflammation and remodelling in an asthma murine model (Toledo et al. 2013). Sakuranetin (2) displayed activity against four *Leishmania* species (*L. amazonensis*, *L. brasiliensis*, *L. major*, and *L. chagasi*) and *Trypanosoma cruzi* (Davila et al. 2013) and also presented phytotoxic activities against *Panicum miliaceum* and *Raphanus sativus*, inhibiting its growth and germination (del Corral et al. 2012). Pectolinaringenin (39) and cirsimarinin (40) present spasmolytic activities (Weimann et al. 2002) and antiparasitic potential (antileishmanial and trypanocidal). This was attributed to 5,6,7-trihydroxy-4'-methoxyflavanone (4), hispidulin (38), spectolinaringenin (39), acacetin (30), and ermanin (77) (da Silva Filho et al. 2009; Passero et al. 2011; Grecco et al. 2010a, b, 2014a, b). Luteolin-3',7-dimethylether (51) displayed anti-inflammatory activity (Gianello et al. 1999), which was also attributed to rutin (128), within its analgesic effect (Gene et al. 1996) and pectolinaringenin (39) (Zalewski et al. 2011). *B. gaudichaudiana* and *B. spicata* presented antiviral activities against poliovirus type 2 (PV2) and vesicular stomatitis virus; PV2

attributed to the presence of apigenin (**28**) (Visintini et al. 2013) This compound also enhances the action of nerve growth factor to stimulate neurite outgrowth from PC12D cells, that could be useful in the treatment of neurological disorders (Guo et al. 2007). *B. articulata* extract induced the death of human peripheral blood mononuclear cells (PBMCs) through apoptosis and exerted low mutagenic effects on mice, those could be related to the presence of luteolin (**48**) and acacetin (**30**) and other nonflavonoidic compounds (Cariddi et al. 2012). Antitumoral activities were attributed to 3,4'-dimethoxy-3',5,7-trihydroxyflavone (**95**) and centaureidin (**106**) (Montes et al. 1971), while gardenin B (**46**) demonstrated cytotoxic activities against HL60 and U937 – human leukemia cell lines, leading to activation of extrinsic and the intrinsic apoptotic pathways (Cabrera et al. 2016). Kaempferol and quercetin glycosides (**123**, **128** and **129**) from *B. antioquensis* displayed photoprotective potential, while quercitrin (**127**) displayed antivenom effect (*Crotalus durissus terrificus* snake) through inhibition of sPLA2 enzyme, preventing myotoxicity and edematogenic effect. Genkwanin (**31**), cirsimarinin (**40**), hispidulin (**38**), and apigenin (**28**) showed antimutagenic activity (Nakasugi and Komai 1998), while quercetin (**89**), luteolin (**48**), nepetin (**54**), apigenin (**28**), and hispidulin (**38**) were responsible for antihepatotoxic properties of *B. trimera* (Soicke and Leng-Eschlow 1987), leading to a protective effect against acute hepatic injury induced by acetaminophen (Padua et al. 2014), an effect also detected in *B. dracunculifolia* leaves extract (Rezende et al. 2014). *B. teindalensis* displayed antiulcer and antidiarrhoeic effects (Vidari et al. 2003), also observed for *B. dracunculifolia* extracts, with inhibition of doxorubicin-induced mutagenicity (Resende et al. 2007). Finally, stems of *B. illinita* presented anticoagulant activity, leading to a significant effect on platelet aggregation (Pizzolatti et al. 2006) and *B. trinervis* displayed cytotoxic, genotoxic, and mutagenic activities that could be related to the presence of flavonoids (Jaramillo-Garcia et al. 2018).

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