

***Brachylaena elliptica* and *B. ilicifolia* (Asteraceae): A Comparative Analysis of their Ethnomedicinal Uses, Phytochemistry and Biological Activities**

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Abstract: *Brachylaena elliptica* and *B. ilicifolia* are shrubs or small trees widely used as traditional medicines in southern Africa. There is need to evaluate the existence of any correlation between the medicinal uses, phytochemistry and pharmacological properties of the two species. Therefore, in this review, analyses of the ethnomedicinal uses, phytochemistry and biological activities of *B. elliptica* and *B. ilicifolia* are presented. Results of the current study are based on data derived from several online databases such as Scopus, Google Scholar, PubMed and Science Direct, and pre-electronic sources such as scientific publications, books, dissertations, book chapters and journal articles. The articles published between 1941 and 2020 were used in this study. The leaves and roots of *B. elliptica* and *B. ilicifolia* are mainly used as a mouthwash and ethnoveterinary medicines, and traditional medicines for backache, hysteria, ulcers of the mouth, diabetes, gastro-intestinal and respiratory problems. This study showed that sesquiterpene lactones, alkaloids, essential oils, flavonoids, flavonols, phenols, proanthocyanidins, saponins and tannins have been identified from aerial parts and leaves of *B. elliptica* and *B. ilicifolia*. The leaf extracts and compounds isolated from the species exhibited antibacterial, antidiabetic, antioxidant and cytotoxicity activities. There is a need for extensive phytochemical, pharmacological and toxicological studies of crude extracts and compounds isolated from *B. elliptica* and *B. ilicifolia*.

Keywords: Asteraceae, *Brachylaena elliptica*, *Brachylaena ilicifolia*, indigenous pharmacopeia, traditional medicine.

INTRODUCTION

The genus *Brachylaena* R. Br. is one of the most important sources of traditional medicines among the Asteraceae genera. The family Asteraceae is characterized by phytochemical compounds such as polyacetyles, acetophenones, chalcone, caffeoylquinic acids, diterpenoids, phloroglucinols, polyacetyles, flavonoids, pyrrolizidine alkaloids and polyphenols [1-5]. Biological activities associated with Asteraceae species include analgesic, anthelmintic, anti-allergic, antidiabetic, antimigraine, anti-inflammatory, antiproliferative, antimicrobial, cardiogenic, antioxidant, neuroprotective, antipyretic, antiulcer, antinociceptive, antimalarial, antiprotozoal, antileishmanial, cytotoxicity and hepatoprotective properties [2,3,5-9]. *Brachylaena elliptica* (Thunb.) DC. and *B. ilicifolia* (Lam.) E. Phillips and Scheweick. are widely used as herbal medicines in southern Africa [10]. The roots of *B. elliptica* are an ingredient of "inembe" herbal concoction, mixed with the roots of *Cyphostemma natalitium* (Szyszl.) J.J.M. van der Merwe, *Gunnera perpensa* L., *Rhoicissus tridentata* subsp. *cuneifolia* (Eckl. & Zeyh.) Urton and *Triumfetta rhomboidea* Jacq. [10,11]. The "inembe" herbal concoction is used to induce or augment labour,

postnatal medication to expel afterbirth, abortifacient, and also administered to animals to expel placenta and to treat and manage endometritis [10,12-15]. The leaves of *B. elliptica* and *B. ilicifolia* are traded as traditional medicines in the informal herbal medicine markets in the Eastern Cape province in South Africa [16-18]. Apart from being used as herbal medicines for similar medical conditions, *B. elliptica* and *B. ilicifolia* have been recorded in overlapping geographical areas in southern Africa (Figure 1). It is, therefore, within this context that the current review was undertaken aimed at providing a comparative analysis of the ethnomedicinal uses, phytochemistry and biological activities of *B. elliptica* and *B. ilicifolia*.

Botanical Description of *Brachylaena elliptica* and *B. ilicifolia*

Brachylaena elliptica is a shrub or a small tree growing up to 7 metres in height [19]. The bark of *B. elliptica* is light grey to brown in colour, vertically fissured with young branches that are grooved with very fine hairs. The leaves are lanceolate to ovate in shape, dark green above and white-felted below. The leaf margins are entire to irregularly toothed with a broadly tapering to rounded apex. The flower head is terminal and axillary branched cluster that is creamy white in colour with male florets in small heads while female florets are in large heads. The fruit is a small nutlet with an apical tuft of bristles. *Brachylaena elliptica* has been recorded in coastal areas, river bush,

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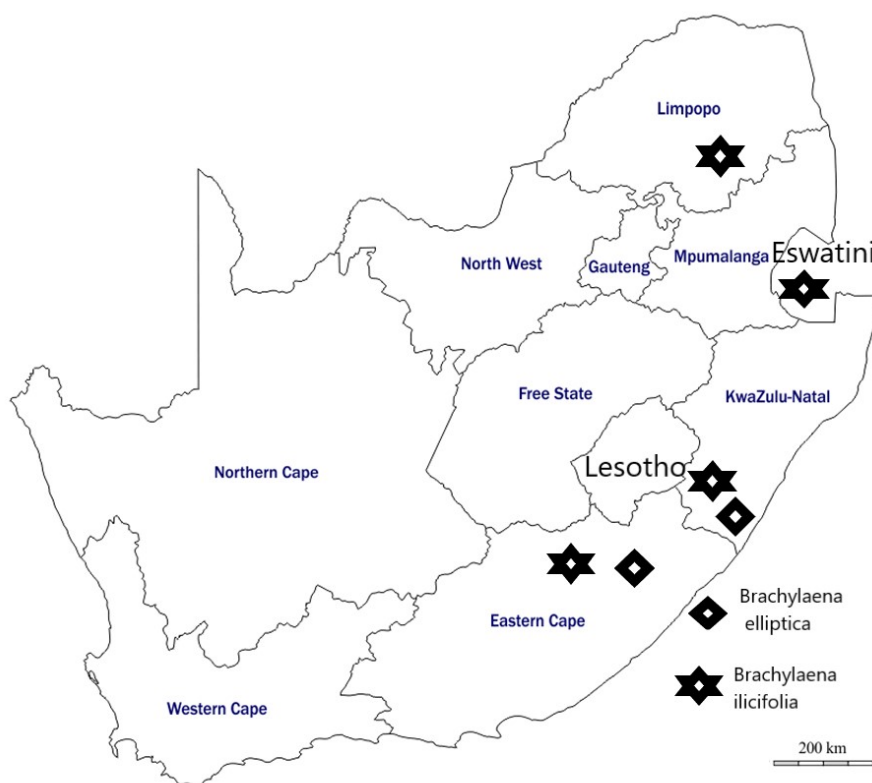


Figure 1: Geographical distribution of *Brachylaena elliptica* and *B. ilicifolia*.

valley bushveld, scrub, grassland, on rocky outcrops and along evergreen forest margins at sea level to 1200 m above sea level [19,20,21]. *Brachylaena ilicifolia* is an evergreen shrub or a small tree growing up to 5 metres in height [19,22]. The main stem is multi-stemmed with dark grey-brown bark that is rough and fissured in older trees. The leaves are crowded at the ends of the branches, alternate, elliptic to ovate in shape, shiny green and smooth above and below are usually covered with white hairs and prominent midrib. The leaf base is rounded to broadly tapering, with entire to finely toothed leaf margins. The flower heads of *B. ilicifolia* are grouped into axillary panicles with creamy-white coloured flowers characterized by a faint, sweet scent. The fruits of *B. ilicifolia* are small achenes characterized by an apical tuft of creamy brown bristles [21,23]. *Brachylaena ilicifolia* has been recorded in bush, scrub forest, on rocky hillsides in seasonally dry open woodland and in dry thickets along watercourses at an altitude ranging from 20 m to 1350 m above sea level [19,20].

Medicinal Uses of *Brachylaena elliptica* and *B. ilicifolia*

The leaves and roots of both *B. elliptica* and *B. ilicifolia* are widely used as traditional medicines in southern Africa. A total of 15 human and livestock diseases and ailments are treated with herbal

concoctions prepared from *B. elliptica* and *B. ilicifolia* (Table 1). The leaves and roots of *B. elliptica* and *B. ilicifolia* are mainly used as a mouthwash and ethnoveterinary medicines, and traditional medicines for backache, hysteria, ulcers of the mouth, diabetes, gastro-intestinal and respiratory problems (Table 1, Figure 2). In South Africa, the roots of *B. elliptica* are used as substitutes for those of *Gymnanthemum corymbosum* (Thunb.) H. Rob. as an abortifacient and ethnoveterinary medicine, and a traditional medicine for stomach ache and hysteria [10,24-29]. In South Africa, the leaves of *B. ilicifolia* are mixed with those of *Leucas capensis* (Benth.) Engl. and sap of *Aloe ferox* Mill. and used as ethnoveterinary medicine [28,30,31].

Phytochemical and Biological Activities of *Brachylaena elliptica* and *B. ilicifolia*

Zdero and Bohlmann [50] identified sesquiterpene lactones, elemanolide and onopordopicrin from the aerial parts of *B. elliptica*. Sagbo [51] and Sagbo *et al.* [52] identified alkaloids, flavonoids, flavonols, phenols, proanthocyanidins, saponins and tannins from leaves of both *B. elliptica* and *B. ilicifolia*. Furthermore, Sagbo [51] identified essential oils from the leaves of both *B. elliptica* and *B. ilicifolia* with benzoic acid, 2,5-bis(trimethylsiloxy)-, trimethylsilyl ester, butylated hydroxytoluene, caryophyllene oxide and oxalic acid, cyclohexylmethyltridecyl ester reported in both

Table 1: Medicinal Applications of *Brachylaena elliptica* and *B. ilicifolia*

Medicinal use	Parts used	Reference
<i>Brachylaena elliptica</i>		
Abortion	Roots used as substitutes for those of <i>Gymnanthemum corymbosum</i> (Thunb.) H. Rob.	[10,24]
Backache	Leaf infusion taken orally	[10,12,27,29]
Diabetes	Leaf decoction taken orally	[10,23,32-43]
Gastro-intestinal problems (biliousness)	Leaf infusion taken orally	[10,12,44]
Stomach ache	Roots used as substitutes for those of <i>Gymnanthemum corymbosum</i>	[10,24,27,29]
Hysteria	Roots used as substitutes for those of <i>Gymnanthemum corymbosum</i>	[10,24-26]
Mouthwash	Leaf decoction taken orally	[23,32,33,35,36,38,42,45]
Pain	Root infusion taken orally	[10,44]
Respiratory problems (pneumonia and sore throat)	Leaf and root infusion taken orally	[10,12,27,34,38,44]
Ulcers of the mouth	Leaf decoction taken orally	[10,12,38,45]
Ethnoveterinary medicine (diarrhoea in calves)	Roots used as substitutes for those of <i>Gymnanthemum corymbosum</i>	[10,24,28]
<i>B. ilicifolia</i>		
Diabetes	Leaf infusion taken orally	[17,22,23,32,35,36,40,41,43,46,47]
Gastro-intestinal problems (diarrhoea)	Leaf infusion and decoction taken orally	[40,48]
Mouthwash	Leaf decoction taken orally	[35]
Pimples in the mouth	Leaf infusion taken orally	[17]
Respiratory problems (asthma, cough and sore throat)	Leaf infusion taken orally	[17]
Ritual	Whole plant	[49]
Ethnoveterinary medicine (paratyphoid in sheep)	Leaf infusion taken orally	[17]
Diarrhoea in lambs	Leaves mixed with those of <i>Leucas capensis</i> (Benth.) Engl. and sap of <i>Aloe ferox</i> Mill.	[28,30,31]

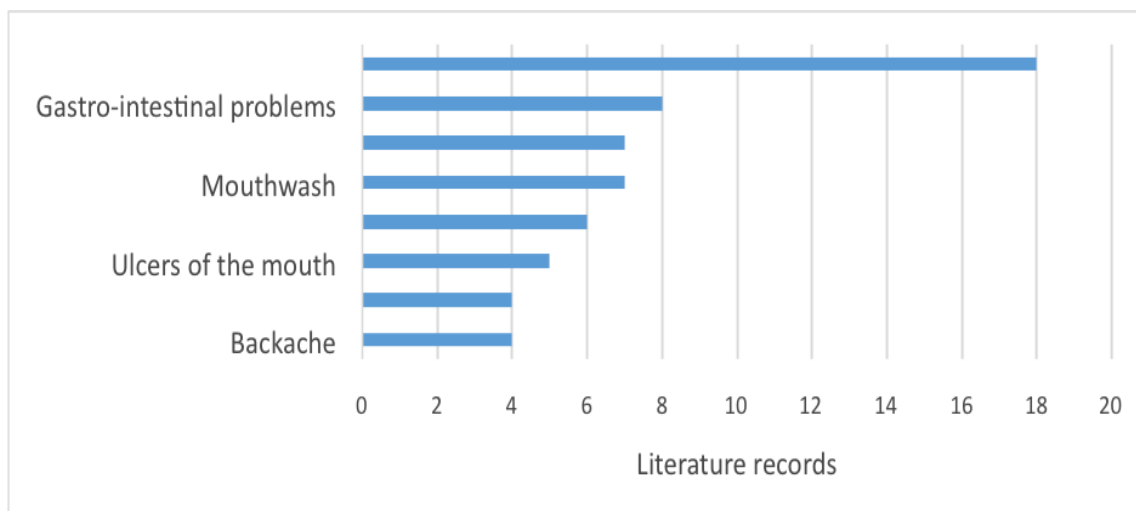


Figure 2: Medicinal applications of *Brachylaena elliptica* and *B. ilicifolia* derived from literature records.

Table 2: Phytochemical Composition of *Brachylaena elliptica* and *B. ilicifolia*

Phytochemical compound	<i>B. elliptica</i>	<i>B. ilicifolia</i>	Plant part	Reference
1R- α -pinene (%)	1.1	Nd*	Leaves	[51]
9-Borabicyclo[3.3.1]nonane,9-hydroxy- (%)	Nd	0.1	Leaves	[51]
Alkaoids (mg/g)	24.3	38.3	Leaves	[52]
Benzoic acid, 2,5-bis(trimethylsiloxy)-,trimethylsilyl ester (%)	0.6	0.1	Leaves	[51]
Butylated hydroxytoluene (%)	4.0	2.2	Leaves	[51]
α -Calacorene (%)	0.7	Nd	Leaves	[51]
δ -Cadinene (%)	1.5	Nd	Leaves	[51]
Carvone oxide, cis- (%)	Nd	0.3	Leaves	[51]
Caryophyllene oxide (%)	2.7	0.5	Leaves	[51]
Damascenone (%)	1.0	Nd	Leaves	[51]
Elemanolide		Nd	Aerial parts	[50]
Eucalyptol (%)	1.2	Nd	Leaves	[51]
Flavanoids (mg/g)	11.5	8.9	Leaves	[52]
Flavonols (mg/g)	44.1	48.8	Leaves	[52]
Hexadecanoic acid, 15-methyl-,methyl ester (%)	0.3	Nd	Leaves	[51]
τ - Muurolol (%)	3.5	Nd	Leaves	[51]
Onopordopicrin		Nd	Aerial parts	[50]
Oxalic acid,cyclohexylmethyltridecyl ester (%)	11.4	2.9	Leaves	[51]
Phenols (mg/g)	75.4	98.6	Leaves	[52]
Phenol,2,5-dichloro-4-methoxy- (%)	0.3	Nd	Leaves	[51]
Proanthocyanidins (mg/g)	156.0	417.0	Leaves	[52]
Pyrazine,methoxy (%)	0.8	Nd	Leaves	[51]
Santolina triene (%)	Nd	0.3	Leaves	[51]
Saponins (mg/g)	34.3	13.3	Leaves	[52]
Squalane (%)	0.8	Nd	Leaves	[51]
Tanins (mg/g)	143.0	211.0	Leaves	[52]

*Nd = not detected.

species (Table 2). Some of these phytochemical compounds may be responsible for the biological activities of the species.

Sagbo *et al.* [52] evaluated the antibacterial activities of ethanol extracts of *B. elliptica* leaves against *Pseudomonas aeruginosa*, *Proteus mirabilis*, *Proteus vulgaris*, *Staphylococcus aureus* and *Streptococcus pyogenes* using the agar dilution method with amoxicillin and ciprofloxacin as positive controls. The extract exhibited activities against all tested pathogens with the exception of *Staphylococcus aureus* with the minimum inhibitory concentration (MIC) values ranging from 2.5 mg/mL to 5.0 mg/mL in comparison to the MIC values of 0.3 mg/mL to 0.6 mg/mL exhibited by the positive controls [52]. Sagbo *et al.* [52] evaluated the antibacterial activities of ethanol

extracts of *B. ilicifolia* leaves against *Pseudomonas aeruginosa*, *Proteus mirabilis*, *Proteus vulgaris*, *Staphylococcus aureus* and *Streptococcus pyogenes* using the agar dilution method with amoxicillin and ciprofloxacin as positive controls. The extract exhibited activities against all tested pathogens with the exception of *Proteus vulgaris* with the MIC values ranging from 2.5 mg/mL to 5.0 mg/mL [52].

Preliminary antidiabetic evaluations carried out by Marles and Farnsworth [53] showed that extracts from the roots and stems of *B. elliptica* lowered blood glucose in experimental animals. Similarly, Sagbo *et al.* [42] evaluated the antidiabetic activities of aqueous leaf extract of *B. elliptica* by assessing the effects of the extract on glucose utilization in HepG2 cells and L6 myoblasts, lipid accumulation in 3T3-L1, glucose

metabolism in INS-1 cells and nitric oxide (NO) production in RAW macrophage cells. The authors also assessed the effects of the extract on alpha-amylase, alpha-glucosidase, pancreatic lipase, dipeptidyl peptidase IV (DPP-IV), collagenase and cytochrome P450 (3A4) enzymes. The extract exhibited activities by causing an increase in the glucose uptake in HepG2 liver cells, producing a concentration-dependent reduction in NO production in RAW macrophages and exhibiting proliferative effect on INS-1 cells at 25.0 µg/ml [42].

Kgopa *et al.* [55] evaluated the antioxidant activities of aqueous extracts of *B. ilicifolia* leaves using the 2,2-azinobis (3-ethyl benzothiazolium-6-sulfonic acid (ABTS), 2,2-diphenyl-1-picrylhydrazyl (DPPH) and ferric reducing antioxidant potential (FRAP) assays. The extract exhibited activities in DPPH showing inhibition of 91.8% and the extract also showed ferric reducing abilities [55]. Sagbo *et al.* [52] evaluated the antioxidant activities of ethanol extracts of *B. elliptica* leaves using the ABTS, DPPH, nitric oxide (NO), hydrogen peroxide (H₂O₂) and ferric reducing power assays with butylatedhydroxytoluene (BHT) and vitamin C as positive controls. The extract exhibited activities with half maximal inhibitory concentration (IC₅₀) values ranging from 0.5 mg/mL to 0.7 mg/mL in ABTS, DPPH, H₂O₂ and NO, and the extract also showed ferric reducing abilities at all tested concentrations when compared with the positive controls [52]. Sagbo *et al.* [52] also evaluated the antioxidant activities of ethanol extracts of *B. ilicifolia* leaves using ABTS, DPPH, nitric oxide, hydrogen peroxide (H₂O₂) and ferric reducing power assays with butylatedhydroxytoluene (BHT) and vitamin C as positive controls. The extract exhibited activities with IC₅₀ values ranging from 0.3 mg/mL to 0.7 mg/mL in ABTS, DPPH, H₂O₂ and NO, and the extract also showed ferric reducing abilities at all tested concentrations when compared with the positive controls [52]. Sagbo *et al.* [42] evaluated the cytotoxicity activities of aqueous leaf extract of *B. elliptica* against HepG2 liver cells using the tetrazolium-based colorimetric 3-(4,5-dimethylthiazol-2-yl)-5-(3,4-diphenyl tetrazolium bromide) (MTT) assay. The extract demonstrated a low level of toxicity with the IC₅₀ value of 250.0 µg/ml [42].

CONCLUSION

The present review summarizes the ethnomedicinal uses, phytochemistry and biological activities of *B. elliptica* and *B. ilicifolia*. Based on the presented

information, these two species are closely related and deemed as potent traditional medicines for treating and managing backache, hysteria, ulcers of the mouth, diabetes, gastro-intestinal and respiratory problems. *Brachylaena elliptica* and *B. ilicifolia* should be subjected to detailed phytochemical, pharmacological and toxicological evaluations aimed at correlating their medicinal uses with their phytochemistry and pharmacological properties.

CONFLICT OF INTEREST

No conflict of interest is associated with this work.

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