



MINI REVIEW ARTICLE

A systematic review on biological and medicinal properties of *Ehretia rigida* (Thunb.) Druce (Ehretiaceae) in Southern Africa

Alfred Maroyi

Department of Botany, University of Fort Hare, Private Bag X1314, Alice 5700, South Africa

*Email: amaroyi@ufh.ac.za

ARTICLE HISTORY

Received: 14 January 2023 Accepted: 13 July 2023 Available online Version 1.0: 10 September 2023

Check for updates

Additional information

Peer review: Publisher thanks Sectional Editor and the other anonymous reviewers for their contribution to the peer review of this work.

Reprints & permissions information is available at https://horizonepublishing.com/ journals/index.php/PST/open_access_policy

Publisher's Note: Horizon e-Publishing Group remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

Indexing: Plant Science Today, published by Horizon e-Publishing Group, is covered by Scopus, Web of Science, BIOSIS Previews, Clarivate Analytics, NAAS, UGC Care, etc See https://horizonepublishing.com/journals/ index.php/PST/indexing_abstracting

Copyright: © The Author(s). This is an openaccess article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution and reproduction in any medium, provided the original author and source are credited (https://creativecommons.org/licenses/ by/4.0/)

CITE THIS ARTICLE

Maroyi A. A systematic review on biological and medicinal properties of *Ehretia rigida* (Thunb.) Druce (Ehretiaceae) in Southern Africa. Plant Science Today (Early Access). https://doi.org/10.14719/pst.2359

Abstract

Wild plants are widely utilized as important sources of traditional medicines and food. Ehretia rigida (Thunb.) Druce is one such plant species that occurs naturally in southern Africa and different parts of the plant have nutritional, cultural, and pharmaceutical importance. The current study is aimed at providing information on the medicinal and biological properties of Ε. rigida. The current review assessed the existing literature on the biological properties and medicinal uses of the plant. The study revealed that the leaves, twigs, stems, bark and roots of the plant are mainly used for ritual purposes, as traditional medicines for infertility, headache, abdominal pains, chest pains, pain, skin cuts, sprained joints, newborn baby infections, a good-luck charm, fire-making and rain-making ceremonies, warding-off enemies and dangerous animals and as ethnoveterinary medicine. The phytochemical evaluation of the species showed that it is characterized by allantoin, α -amyrin, β -amyrin, flavonoids, phenolics, saponins, tannins, β sitosterol and 1-triacontanol. The pharmacological properties of different crude extracts demonstrated anticholinesterase inhibition, antifungal, antibacterial, antidiabetic, anti-inflammatory, and antioxidant properties. This review highlights the pharmaceutical and health benefits of *E. rigida* in different countries of southern Africa. Therefore, detailed ethnopharmacological evaluations of the species focusing on phytochemistry, pharmacological properties, toxicological evaluations, and in vivo and clinical research investigations are recommended.

Keywords

Boraginaceae; *Ehretia rigida*; Ehretiaceae; indigenous pharmacopeia; toxicological evaluations; traditional medicine

Introduction

The genus name *Ehretia* P.Browne is honoring a German, Georg Dionysius Ehret born on 30 January 1708 and died on 9 September 1770, an entomologist and botanist well-known for his botanical illustrations (1). The specific epithet *rigida* means rigid or stiff branches (2, 3). Synonyms of *E. rigida* include *Capraria rigida* Thunb., *E. eckloniana* H. Buek ex Harv., *E. hottentotica* Burch., *E. violacea* Kunth, *E. zeyheriana* Buek. ex Harv., *Pittosporum commutatum* Krauss and *Freylinia rigida* (Thunb.) G.Don. (4-7). The English common names of *E. rigida* include "Cape lilac", "forest puzzle bush", "puzzle bush" and "stamper wood" (2, 8). *E. rigida* (Thunb.) Druce (Fig. 1) naturally occurs in southern Africa in Botswana, Lesotho, Mozambique, Eswatini, Namibia, Zimbabwe, and South Africa (Fig. 2) (4-6, 9-11). Local people in



Fig. 1. Ehretia rigida A: branch showing leaves and flowers, and B: branch showing flowers and fruits (Photos: MC Palgrave and B Wursten)

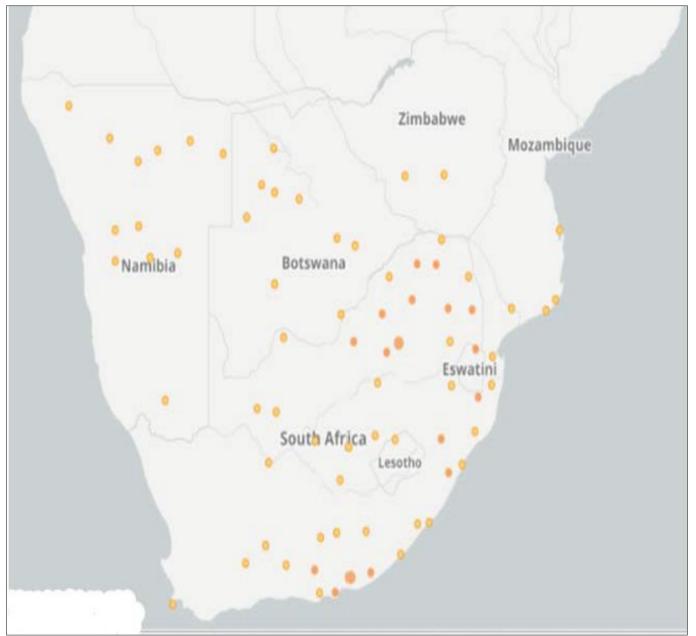


Fig. 2. Distribution of Ehretia rigida in southern Africa)

people in southern Africa have developed indigenous knowledge and skills necessary to provide locally appropriate solutions to local problems, particularly the exploitation of medicinal plants including E. rigida. The use of medicinal plants, crude plant extracts, and their active ingredients, or phytochemicals plays an important role in the primary health care needs of local communities in southern Africa. Medicinal plants are an important aspect of the daily lives of many people and an important part of the Southern African cultural heritage (12). Therefore, understanding the phytochemicals associated with E. rigida is important as these active ingredients act directly or indirectly to prevent or treat diseases and also maintain health. Due to its ability to withstand drought, E. rigida has been identified as an important local species in southern Africa that can be easily integrated into home gardens as an ornamental or hedge plant (2). Therefore, the objective of this review was to collect and summarize information about the medicinal uses, phytochemistry, and biological properties of E. rigida.

Methodology

The literature review of the medicinal and biological properties of *E. rigida* within its geographical ranges in southern Africa was conducted using online databases such as JSTOR, Scopus, PubMed, Science Direct, and Google Scholar. In addition to this, pre-electronic literature sources used which included journal articles, books, theses, dissertations, book chapters, and other scientific articles were obtained from the University of Fort Hare library. Keywords such as *Ehretia rigida*, biological activities, pharmacological properties, ethnobotany, medicinal uses, phytochemistry, and traditional uses of *Ehretia rigida* were used to search for relevant articles as shown in the flow diagram (Fig. 3). Literature sources excluded from this re-

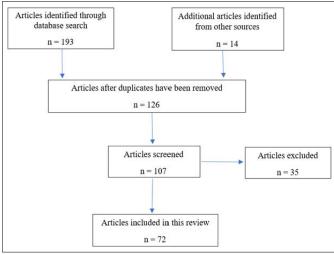


Fig. 3. Flow diagram showing identification and screening of articles used in this review

view are those articles that are partially accessed, that is, accessed as abstracts only, and also published or unpublished ethnopharmacological surveys lacking information on the medicinal uses, phytochemistry, pharmacological or biological activities of *E. rigida*. The chemical structures of phytochemical compounds isolated from *E. rigida* have been drawn using the program ChemSketch.

Results

Morphological description and taxonomy of the species

The genus *Ehretia* consists of about 40 tree and shrub species belonging to the family Ehretiaceae (13). The Ehretia genus has been recorded in the New and Old World tropics with its center of diversity in East Asia and Africa (13). The family Ehretiaceae has been segregated from Boraginaceae sensu lato based on molecular and phylogenetic studies focusing on plastid loci (trnL-trnF, rps16, and trnStrnG) and nuclear (ITS) (13-16). E. rigida is a highly branched shrub or multi-stemmed tree, deciduous, reaching up to 12 meters in height (5). The stems, usually several to many, are often low branching and crooked with grey to pale brown bark with prominent lenticels (4). The inner bark is soft, white, spotted with orange-brown, and turning brown upon exposure (1). The leaves are simple and alternate but often clustered on reduced shoots; obovate, dark green or grey-green above, hairless with prominent venation on lower surfaces (4). The leaf stalks may be absent, short, or sometimes pronounced (1). The flowers are bisexual, regular, pale mauve to dark purple, usually on branched terminal heads, densely congested with peduncles and pedicels (8). The fruit is a berry, subglobose, spherical in shape, yellow to bright orange-red when riped (3, 17). E. rigida is divided into three subspecies distinguished by vegetative and reproductive characteristics, and geographical distribution. E. rigida subsp. rigida is a stunted tree or shrub of up to four meters in height with terminal inflorescences, thick peduncles, and pedicels, and confined to the Albany Centre of Endemism in South Africa (5). Ehretia rigida subsp. silvatica Retief and A.E. Van Wyk and *E. rigida* subsp. *nervifolia* Retief and A.E. Van Wyk are widespread with heights, exceeding five metres (8).

Distribution and growing condition

E. rigida has been recorded in coastal areas, forests, thickets, woodland, and bushland biomes (1). E. rigida naturally occurs in valleys, open flats, stony slopes, granite hills, dunes, Kalahari sand areas and termite mounds at an altitude ranging from sea level to 1700 m above sea level (4, 6-8, 11). In the forest biome, the species loses the shrub characteristic habit and grows upwards to about 12 meters in height with a small tangled crown (1). *E. rigida* is usually in flower from March to December, and fruits are often produced from October to May (3, 4, 8). E. rigida is drought and frost-resistant, easily grown from seeds and cuttings, characterized by fast growth rate of about 600 mm to 700 mm per annum (2, 18). E. rigida can be planted in a garden to attract birds and insects, and also the lilac-to-mauve flowers and orange-to-red fruits make the plant species an interesting and decorative shrub in the garden (2). Subspecies rigida is the most widespread taxon, recorded in Botswana, Mozambique, South Africa, and Zimbabwe (6-11). Subspecies nervifolia has been recorded in Botswana, Eswatini, Lesotho, the Free State, Gauteng, KwaZulu Natal, Limpopo, Mpumalanga, and North-West provinces in South Africa while subspecies sylvatica is confined to the Eastern Cape and KwaZulu Natal provinces in South Africa (6, 8).

Ethnomedicinal significance

In South Africa, *E. rigida* roots are traded as sources of traditional medicines in rural, peri-urban and urban markets in the Gauteng province (19). The fruits are eaten as snacks (1, 18, 20-26), as vegetables and to make non-alcoholic beverages (3). In South Africa, the leaves and twigs are used as cattle feed and eaten by birds and wild animals such as bushbuck (*Tragelaphus scriptus*), grey duiker (*Sylvicapra grimmia*), impala (*Aepyceros melampus petersi*), kudu (*Tragelaphus strepsiceros*), monkey (*Cercopithecus aethiops*) and nyala (*Tragelaphus angasii*) (1, 2, 23). In southern Africa, the herbal concoctions prepared from leaves, bark, roots and twigs are used to treat several animal and human ailments including infertility, headache, abdominal pains, chest pains, pain, skin cuts, sprained joints, newborn baby infections, a good-luck charm, fire-making and rain-making ceremonies, warding off enemies and dangerous animals and as ethnoveterinary medicine (Table 1). In South Africa, the root powder of *E. rigida* is often mixed with the root powder of *Zanthoxylum davyi* (I.Verd.) P.G.Waterman and the bark of *Maerua angolensis* DC. as a remedy against diabetes (27), while the root paste is topically applied with the root paste of *Asparagus falcatus* L. for sprained joints (20, 27).

Region	Plant part used	Mode of preparation	Traditional/ethnomedicinal uses	Reference
South Africa	Roots	Root decoction is taken orally	Abdominal pains	(8, 18, 28, 29)
South Africa	Leaves	Leaf paste topically applied	Burns	(30)
South Africa	Leaves	Leaf paste topically applied	Candidiasis	(31)
Botswana	Whole plant	The tree is used for rain-making ceremonies with the cloud of smoke from twigs being symbolic of rain clouds	Rain-making ceremonies	(1)
Lesotho	Twigs	Hunters smear themselves with the decoction of the twigs	Charm for success when hunting	(32-34)
South Africa	Stems and twigs	People smear themselves with the decoction of stems and twigs	Good luck, hunting and protective charm, warding-off enemies and dangerous animals	(1, 18, 20, 22, 2 29)
South Africa	Roots	Root powder decoction is taken orally	Chest pains	(8, 3, 18, 28, 29)
South Africa	Roots	Root powder mixed with root powder of <i>Zanthoxylum davyi</i> (I.Verd.) P.G.Waterman and the bark of <i>Maerua angolensis</i> DC.	Diabetes	(27)
South Africa	Roots	Root powder infusion is taken orally	Emetic	(35)
South Africa	Roots	Root powder infusion is taken orally	General childhood infections	(36)
South Africa	Roots	Root powder decoction is taken orally	Headache	(27, 37-39)
South Africa	Roots	Root powder decoction is taken orally	Infertility	(20, 27, 40)
South Africa	Roots	Root powder decoction is taken orally	Newborn infections	(27, 41)
Eswatini	Roots	Root powder decoction is taken orally	Pain	(42)
Namibia	Roots	Root powder decoction is taken orally	Pain	(43)
South Africa	Roots	Root powder decoction is taken orally	Pain	(44)
South Africa	Roots	Root powder infusion is taken orally	Respiratory infections	(45)
Eswatini	Roots	Root powder decoction topically applied	Skin cuts	(42)
South Africa	Roots	Root powder decoction topically applied	Skin cuts	(8, 18)
South Africa	Roots	Root paste topically applied and mixed with root paste Asparagus falcatus L	Sprained joints	(20, 27)
South Africa	Roots	Root powder infusion is taken orally	Stomach pains	(3)
South Africa	Roots	Root powder infusion is taken orally	Tonic	(46)
Eswatini	Roots	Root maceration topically applied	Toothache	(42)
South Africa	Leaves	The leaf powder was topically applied	Wounds	(30)
Ethnoveterinary	medicine			
South Africa	Roots	Root powder decoction used	Eating problems in cattle	(47, 48)
South Africa	Roots	Root paste topically applied	Fractures	(48, 49)
South Africa	Roots	Root powder decoction used	Gall sickness in cattle	(2, 8, 18, 29)

Nutritional composition

Various scientists identified elementary, nutritional and chemical compounds from the fruits, bark, roots and leaves of *E. rigida* (Table 2).

Table 2. Nutritional composition of Ehretia rigida .

Nutritional components	Value	Plant part	References
Ash (g/100g)	1.3	Fruits	(50, 51)
Calcium (mg/100g)	30.5	Fruits	(50, 51)
Carbohydrates (g/100g)	10.4	Fruits	(50, 51)
Copper (mg/100g)	0.2	Fruits	(50, 51)
Crude fiber (g/100g)	0.7	Fruits	(50, 51)
Energy (KJ/100g)	215	Fruits	(50, 51)
Fat (g/100g)	0.3	Fruits	(50, 51)
Iron (mg/100g)	0.9	Fruits	(50, 51)
Magnesium (mg/100g)	27.8	Fruits	(50, 51)
Moisture (g/100g)	85.6	Fruits	(50, 51)
Potassium (mg/100g)	547.0	Fruits	(50, 51)
Protein (g/100g)	1.7	Fruits	(50, 51)
Sodium (mg/100g)	2.5	Fruits	(50, 51)
Vitamin C (mg/100g)	6.9	Fruits	(50, 51)
Zinc (mg/100g)	0.2	Fruits	(50, 51)

Phytochemistry

Various researchers identified chemical compounds from the bark, leaves and roots of *E. rigida* (Fig. 4; Table 3). The chemical compounds identified from *E. rigida* include allantoin, α -amyrin, β -amyrin, β -sitosterol, 1-triacontanol, flavonoids, phenolics, saponins and tannins (52-55).

Pharmacological properties

Anticholinesterase inhibition activity

The anticholinesterase inhibition properties of acetone leaf extracts were assessed using an acetylcholinesterase inhibition assay, with eserine as a positive control (54). At a determined concentration of 500.0 μ g/mL, the acetone extracts showed the anticholinesterase inhibitory activity of 50.0% and a dose-dependent half maximum inhibitory concentrations (IC₅₀) average value of 487.4 μ g/mL. The positive control, eserine exhibited anticholinesterase inhibitory activity of 60.0% and an IC₅₀ value of 4.9 μ g/mL (54).

Antimicrobial property

The antibacterial properties of hexane, diethyl ether, chloroform and ethyl acetate leaf extracts were evaluated at 50.0 mg/ml concentration against bacterial pathogens viz. Agrobacterium tumefaciens, Erwinia carotovora subsp. carotovora, Clavibacter michiganense, Xanthomonas campestris and Pseudomonas solanacearum, through an agar diffusion assay using dimethyl dodecyl ammonium chloride (DDAC) as a positive or standard control (56). Ethyl acetate extract demonstrated antibacterial properties against Erwinia carotovora subsp. carotovora exhibiting inhibition of 7.0 mm in comparison with inhibition of 13.0 mm demonstrated by the positive or standard control (56). Similarly, the antibacterial properties of dichloromethane, methanol and ethyl-acetate bark, root and leaf extracts were evaluated against Escherichia coli, Staphylococcus aureus, Micrococcus luteus and Klebsiella pneumoniae through a microdilution assay with neomycin as a positive control (55). The dichloromethane, methanol and ethylacetate extracts demonstrated antibacterial properties against the tested bacterial pathogens with minimum in-

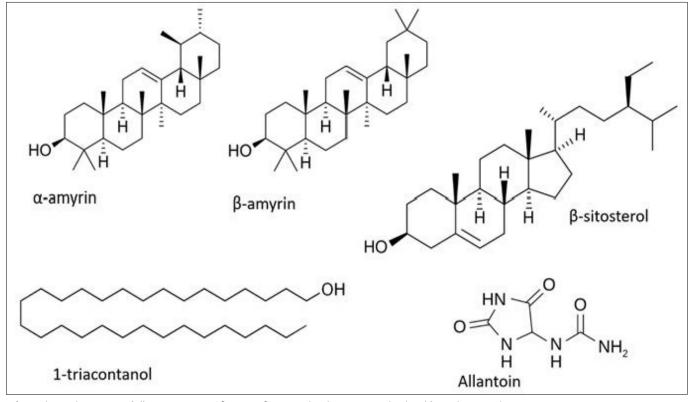


Fig. 4. Chemical structures of allantoin, α -amyrin, β -amyrin, β -sitosterol and 1-triacontanol isolated from Ehretia rigida

Table 3. Phytochemica	l composition	of Ehretia rigida
-----------------------	---------------	-------------------

Type of compound	Name of the compound	Plant part from where extracted/ identified/documented	Process of detection/ isolation/ extraction	References
Glyoxylic acid	Allantoin	Bark	¹ H and ¹³ C NMR	(52)
Triterpene	α-amyrin	Bark	GC	(52)
Triterpene	β-amyrin	Bark	GC	(52)
<u>Phytosterol</u>	β-sitosterol	Bark	¹ H and ¹³ C NMR	(52)
Fatty alcohol	1-triacontanol	Bark	¹ H and ¹³ C NMR	(52)
Flavonoids	-	Bark, leaves, and roots	Calorimetric assay	(54, 55)
Phenolics	-	Bark, leaves, and roots	Folin-Ciocalteau	(53-55)
Saponins	-	Bark, leaves, and roots	Froth test	(55)
Tannins	-	Bark, leaves, and roots	Butanol-HCL assay	(55)

hibitory concentration (MIC) values within the range from 0.78 to >6.25 mg/ml which were higher than the MIC values ranging from 0.01 mg/ml to 0.05 mg/ml demonstrated by the positive control (55).

The antifungal properties of water and acetone leaf extracts were evaluated against Candida parapsilosis, C. albicans, Cryptococcus neoformans, C. glabrata, C. tropicalis and C. krusei, through microdilution method, with clotrimazole as a positive control (30). The plant extracts demonstrated weak antifungal activity in comparison to the positive control with MIC values ranging between 0.04 mg/ml - 0.08 mg/ml (30). The antifungal properties of acetone leaf, root and bark extracts were assessed against C. albicans, Aspergillus fumigatus and Cryptococcus neoformans using a microdilution assay (57). The extracts demonstrated moderate activity with MIC values of 0.02 mg/ml to 0.08 mg/ml (57). Further, the antifungal properties of dichloromethane, methanol and ethyl-acetate plant bark, leaf and root extracts were evaluated against C. albicans through microdilution assay, with an amphotericin B as a positive control (55). The extracts demonstrated considerably significant antifungal activity against C. albicans with MIC values ranging from 0.78 mg/ml to >6.25 mg/ml compared to a positive control (MIC: 0.01 to 0.08 mg/ml (55).

Antidiabetic activity

The antidiabetic properties of leaf, bark and root extracts were evaluated using a microplate serial dilution assay to determine the inhibition of α -glucosidase (55). The α -glucosidase inhibitory activities exhibited by the extracts were dose-dependent ranging from 16.0% to 23.0% at 125.0 µg/ml, 40.0% to 44.0% at 250.0 µg/ml and 59.0% to 65.0% at 500.0 µg/ml (55).

Anti-inflammatory activity

The anti-inflammatory properties of acetone extracts of leaves were assessed using soybean lipoxygenase inhibition, nitric oxide production and the viability of LPS-activated RAW 264.7 macrophage assay, using indomethacin and quercetin as positive controls (54). The acetone extracts demonstrated properties with anti-lipoxygenase activities showing 50.0% inhibition and an IC₅₀ value of 63.0 μ g/mL at 100.0 μ g/mL. Effective dose-dependent inhibition reactions of nitric oxide production (rate of inhibi-

tion: 86.3% to 92.6%; cell viability: 34.3% to 63.9%) were exhibited by acetone extract at 6.3 μ g/mL, 12.5 μ g/mL, 25.0 μ g/mL, and 50.0 μ g/mL concentrations (54).

Antioxidant activities

The antioxidant properties of water extracts of leaves were evaluated against 2,2-diphenyl-1-picrylhydrazyl (DPPH), 2,2-azinobis (3-ethyl benzothiazolium-6-sulfonic acid (ABTS), and ferric reducing antioxidant potential (FRAP) methods (53). The extract showed activities against DPPH radicals only, exhibiting 94.9% inhibition (53). The antioxidant properties of the acetone extracts of E. rigida leaf were assessed using the DPPH, ABTS, and FRAP methods with ascorbic acid and trolox as positive controls (54). The extracts demonstrated significant antioxidant properties with IC₅₀ values of 171.8 μ g/mL for DPPH, 94.1 μ g/mL for FRAP, and 234.8 µg/mL for ABTS (54). The antioxidant properties of acetone and water extracts of leaves were evaluated against DPPH free radicals (30). Both acetone and water extracts exhibited activities with IC₅₀ values ranging from 62.0 µg/ml to 131.0 µg/ml (30). The antioxidant properties of methanol, dichloromethane, and ethylacetate extracts of roots, bark, and leaves were assessed using DPPH free radical scavenging assay with ascorbic acid as a positive control (55). The extracts demonstrated significant dose-dependent antioxidant properties (55).

Discussion

The current review attempted to provide information available on the medicinal uses, phytochemistry and biological activities of *E. rigida* covering literature from 1962 to 2023, a very long period to capture all the relevant research on the ethnopharmacological properties of the species. This species belongs to the *Ehretia* genus which is widely used as a source of herbal medicines in tropical Africa (58, 59). Some of the species belonging to the *Ehretia* genus used as sources of traditional medicines in tropical Africa include *E. amoena* Klotzsch, *E. cymosa* Thonn., *E. obtusifolia* Hochst. ex DC., *E. rigida*, and *E. trachyphylla* C.H. Wright (58-67). The value and importance of *E. rigida* as traditional medicine is demonstrated by the inclusion of the species in the monograph Medicinal and magical plants of Southern Africa: An Annotated Checklist (62) and also its trade as a source of traditional medicines in both informal and formal herbal medicine markets in South Africa (19). Research findings in this review show how indigenous people in southern Africa have adapted and become familiar with the diverse medicinal uses and properties of *E. rigida*. The dependence of rural households on *E. rigida* for non-medicinal and miscellaneous uses for their livelihood needs is also well recognized in southern Africa.

Different classes of phytochemicals of E. rigida including flavonoids, phenolics, saponins, and tannins, have been quantified using different methods (53-55) but several factors such as plant part, season, and environmental factors influence such phytochemical synthesis and quantities (68). Some of these classes of secondary phytochemical constituents may be responsible for the documented biological activities of E. rigida. For example, the chemical compound allantoin identified from E. rigida is known to enhance the smoothness of the skin by stimulating the healing of wounds and soothing irritated skin (69). The chemical compounds α - and β -amyrin identified from E. rigida are characterized by analgesic, anti-inflammatory, anticonvulsant, anti-depressive, gastroprotective, hepatoprotective, anti-pancreatitis, anticholytic, antihyperglycemic and hypolipidemic activities (70). Similarly, the phytosterol, β-sitosterol identified from *E. rigida* is known to have antimicrobial, anti-inflammatory, anticancer, antifertility, angiogenic, antioxidant, immunomodulatory, antidiabetic, and antinociceptive activities (71). This background information highlights the need for detailed phytochemical evaluation of various E. rigida extracts aimed at isolating, purifying, and characterizing phytochemical compounds of this species. Therefore, the little data on the phytochemical compounds of the species highlights a research gap requiring attention from researchers. There is growing interest in phytochemical constituents of plant species in a quest to discover new remedies or to explain the modes of action of these medicinal plants (72).

E. rigida has been subjected to a series of anticholinesterase inhibition, antibacterial, antifungal, antidiabetic, anti-inflammatory, and antioxidant assays as the species is widely used as traditional medicine for abdominal pains, burns, candidiasis, chest pains, childhood infections, diabetes, headache, infertility, pain, respiratory infections, skin cuts, sprained joints, stomach pains, toothache and wounds in humans, eating problems, fractures and gall sickness in cattle (Table 1). The antimicrobial and antioxidant activities exhibited by the crude extracts of E. rigida are directly or indirectly involved in protection against free radicals and the growth of undesirable microbes. There is a need for detailed ethnopharmacological studies focusing on their potential in vitro and in vivo conditions as there is a need to assess the clinical relevance of these different extracts of the species. Therefore, knowledge about the resultant pharmacological and toxicological properties of both crude extracts and the phytochemical constituents isolated from E. rigida should be subjected to controlled clinical trials to determine if such crude extracts and phytochemical compounds are characterized by sufficient therapeutic properties and can be linked to a particular disease or indication associated with

the species. Till now, limited ethnopharmacological research has been made in the pharmacokinetics studies related to the mechanism of action of the individual isolated phytochemical compound under *in vivo* conditions. Also, ethnopharmacological research should focus on toxicity, preclinical, and clinical trials aimed at accounting for the safety, effectiveness, and side effects of the phytochemical compounds in the formulation of *E. rigida* as a source of traditional medicines.

Conclusion

The current study provides a summary of the medicinal, phytochemical and biological activities of *E. rigida*. Such evaluations are important considering that E. rigida is widely used in southern Africa as a food plant and also as an important source of traditional medicines. E. rigida is among the indigenous fruit trees with high medicinal and therapeutic potential that is not fully researched, and have fragmented information in the literature. The presence of carbohydrates, fats, fiber, proteins, vitamins, minerals and biologically active phytochemicals such as flavonoids and phenolic acids makes E. rigida suitable for day-to-day activities as a source of food and medicines. However, there is a lack of alignment between food and ethnomedicinal uses, and existing phytochemical and biological screenings of the species, indicating the need for further ethnopharmacological evaluations. Although the existing research efforts are laudable, several areas of research on E. rigida have been neglected. It is expected that the identified research gaps will serve as a guide for future holistic research focusing on E. rigida. Therefore, detailed ethnopharmacological evaluations of the species focusing on phytochemistry, pharmacological properties, and toxicological evaluations as well as *in vivo* and clinical research aimed at corroborating the traditional medical and food applications of the species are recommended. The outcomes of such research have the potential to accelerate the commercialization of E. rigida health and pharmaceutical products.

Acknowledgements

Funding for this research was provided by the University of Fort Hare, South Africa.

Compliance with ethical standards

Conflict of interest: The author declares no conflict of interest.

Ethical issues: None.

References

- Palmer E, Pitman N. Trees of southern Africa, covering all known indigenous species in the Republic of South Africa, South-West Africa, Botswana, Lesotho and Swaziland. AA Balkema, Cape Town. 1972
- 2. Venter F, Venter J-A. Making the most of indigenous trees. Briza Publications, Pretoria. 2015
- Schmidt E, Lotter M, Mccleland W. 2007. Trees and shrubs of Mpumalanga and Kruger National Park. Jacana Media, Johannesburg. 2017

- Martins ES. Boraginaceae. In: Launert E, Pope GV, editors. Flora Zambesiaca volume 7, part 4. Flora Zambesiaca Managing Committee, London. 1990; pp. 59–110.
- Retief E, Van Wyk AE. The genus *Ehretia* (Boraginaceae: Ehretioideae) in southern Africa. Bothalia 2001;31:9-23. https:// doi.org/10.4102/abc.v31i1.494
- Germishuizen G, Meyer NL. Plants of Southern Africa: An annotated checklist. Strelitzia 14, National Botanical Institute, Pretoria. 2003
- Hyde MA, Wursten BT, Ballings P, Palgrave CM. Flora of Zimbabwe: Species information: *Ehretia rigida* subsp. *rigida*. 2023. Available from: https://www.zimbabweflora.co.zw/speciesdata/ species.php?species_id=148300, accessed on 6 January 2023
- 8. Palgrave MC. Keith Coates Palgrave trees of southern Africa. Struik Publishers, Cape Town. 2002
- Setshogo MF. Preliminary checklist of the plants of Botswana. Southern African Botanical Diversity Network Report No. 37, National Botanical Institute, Pretoria. 2005
- Setshogo MP, Venter F. Trees of Botswana: Names and distribution. Southern African Botanical Diversity Network Report No. 18, National Botanical Institute, Pretoria. 2005
- Burrows J, Burrows S, Lötter M, Schmidt E. Trees and shrubs of Mozambique. Publishing Print Matters, Noordhoek, Cape Town. 2018
- 12. Van Wyk BE, Van Oudtshoorn B, Gericke N. Medicinal plants of South Africa. Briza Publications, Pretoria. 2013
- Gottschling M, Weigend M, Hilger HH. Ehretiaceae. In: Kadereit JW, Bittrich V, editors., Flowering plants: Eudicots: Aquifoliales, Boraginales, Bruniales, Dipsacales, Escalloniales, Garryales, Paracryphiales, Solanales (except Convolvulaceae), Icacinaceae, Metteniusaceae, Vahliaceae: The families and genera of vascular plants, volume 14. Springer, Cham. 2016; pp. 165–78. https:// doi.org/10.1007/978-3-319-28534-4_12
- Gottschling M, Hilger HH. 2001. Phylogenetic analysis and character evolution of *Ehretia* and *Bourreria* (Ehretiaceae, Boraginales) and their allies based on ITS1 sequences. Bot Jahrb Syst 2001;123:249–68. https://doi.org/10.1127/0006-8152/2002/0124-0149
- Gottschling M, Diane N, Hilger HH, Weigend M. Testing hypotheses on disjunctions present in the primarily woody Boraginales: Ehretiaceae, Cordiaceae and Heliotropiaceae, inferred from ITS1 sequence data. Int J Plant Sci 2004;165(S4):S123–S35. https:// doi.org/10.1086/421069
- Gottschling M, Luebert F, Hilger HH, Miller J.S. 2014. Molecular delimitations in the Ehretiaceae (Boraginales). Mol Phylogen Evol 2014;72:1–6. https://doi.org/10.1016/j.ympev.2013.12.005
- 17. Manning J, Goldblatt P. Plants of the Greater Cape Floristic Region 1: The core Cape Flora. Strelitzia 29, South African National Biodiversity Institute, Pretoria. 2012
- Ndou P. *Ehretia rigida* (Thunb.) Druce. 2003. Available from: http://pza.sanbi.org/ehretia-rigida, accessed on 19 December 2022
- Williams VL, Balkwill K, Witkowski ET: A lexicon of plants traded in the Witwatersrand *umuthi* shops. Bothalia 2001;31:71-98. https:// doi.org/10.4102/abc.v31i1.508
- 20. Mabogo DEN. The ethnobotany of the Vhavenda. MSc Dissertation, University of Pretoria, Pretoria. 1990
- 21. Shava S. The use of indigenous plants as food by a rural community in the Eastern Cape: An educational exploration. MSc Dissertation, Rhodes University, Grahamstown. 1999.
- 22. Cocks ML, Wiersum KF. The significance of plant diversity to rural households in Eastern Cape Province of South Africa. For Trees Livelih 2003;13:39-58. https://

doi.org/10.1080/14728028.2003.9752443

- 23. Botha J, Weiersbye IM. Ethnobotanic and forage uses of plants on mine properties in the Witwatersrand basin gold fields, South Africa. In: Mine Closure. Fourie A, Tibbett M, Wiertz J, editors. Australian Centre for Geomechanics, Perth. 2010; pp. 325-42.
- Rasethe MT, Semenya SS, Potgieter MJ, Maroyi A. The utilization and management of plant resource in rural areas of the Limpopo province, South Africa. J Ethnobiol Ethnomed 2013;9:27. https:// doi.org/10.1186/1746-4269-9-27
- Magwede K, Van Wyk B-E, Van Wyk AE. An inventory of Vhavenda useful plants. S Afr J Bot 2019;22:57–89. https://doi.org/10.1016/ j.sajb.2017.12.013
- Welcome AK, Van Wyk B-E. An inventory and analysis of the food plants of southern Africa. S Afr J Bot. 2019;122:136-79. https:// doi.org/10.1016/j.sajb.2018.11.003.004
- Mudau TE, Olowoyo JO, Amoo SO. 2022. Ethnobotanical assessment of medicinal plants used traditionally for treating diabetes in Vhembe district, Limpopo province, South Africa. S Afr J Bot 2022;146:304-324. https://doi.org/10.1016/j.sajb.2021.10.016
- 28. Watt JM, Breyer-Brandwijk MG. The medicinal and poisonous plants of southern and eastern Africa. Livingstone, London. 1962.
- Hutchings A, Scott AH, Lewis G, Cunningham AB. Zulu medicinal plants: An inventory. University of Natal Press, Pietermaritzburg. 1996.
- Ndhlovu PT, Mooki O, Otang-Mbeng W, Aremu AO. Plant species used for cosmetic and cosmeceutical purposes by the Vhavenda women in Vhembe District Municipality, Limpopo, South Africa. S Afr J Bot 2019;122:422–31. https://doi.org/10.1016/ j.sajb.2019.03.036
- Nkuna KV, Masevhe NA. Antifungal activity of the selected medicinal plants used to treat candidiasis. S Afr J Bot 2015;98:214. https://doi.org/10.1016/j.sajb.2015.03.170
- 32. Jacot Guillarmod A. Flora of Lesotho. Cramer, Lehre. 1971
- 33. Moffett RO. Sesotho plant and animal names and plants used by the Basotho. Sun Press, Stellenbosch. 2016
- Moteetee A, Moffett RO, Seleteng-Kose L. A review of the ethnobotany of the Basotho of Lesotho and the Free State province of South Africa (South Sotho). S Afr J Bot 2019;122:21-56. https:// doi.org/10.1016/j.sajb.2017.12.012
- Philander LA. An ethnobotany of Western Cape Rasta bush medicine. J Ethnopharmacol. 2011;138:578-94. https:// doi.org/10.1016/j.jep.2011.10
- Ndhlovu PT, Omotayo AO, Otang-Mbeng W, Aremu AO. Ethnobotanical review of plants used for the management and treatment of childhood diseases and well-being in South Africa. S Afr J Bot. 2021;137:197-215. https://doi.org/10.1016/j.sajb.2020.10.012
- Semenya SS, Maroyi A. Ethnobotanical survey of plants used by Bapedi traditional healers to treat tuberculosis and its opportunistic infections in the Limpopo province, South Africa. S Afr J Bot 2019;122:401–21. https://doi.org/10.1016/j.sajb.2018.10.010
- Semenya SS, Maroyi A. Source, harvesting, conservation status, threats and management of indigenous plant used for respiratory infections and related symptoms in the Limpopo province,South Africa. Biodiversitas 2019;20(3):790-811. https://doi.org/10.13057/ biodiv/d200325
- Semenya SS, Maroyi A. Ethnobotanical survey of plants used to treat respiratory infections and related symptoms in the Limpopo province, South Africa. J Herbal Med 2020;24:100390. https:// doi.org/10.1016/j.hermed.2020.100390
- Arnold H-J, Gulumian M. Pharmacopoeia of traditional medicine in Venda. J Ethnopharmacol 1984;12:35–74. https:// doi.org/10.1016/0378-8741(84)90086-2
- 41. Mongalo NI, Makhafola TJ. Ethnobotanical knowledge of the

lay people of Blouberg area (Pedi tribe), Limpopo province, South Africa. J Ethnobiol Ethnomed 2018;14:46. https:// doi.org/10.1186/s13002-018-0245-4.

- Long C. Swaziland's Flora: SiSwati names and uses. Swaziland National Trust Commission, Mbambane; 2005. Available from: http:// www.sntc.org.sz/index.asp, accessed on 3 January 2023.
- Dushimemaria F, Mumbengegwi DR, Böck R. Indigenous knowledge of medicinal plants used for the treatment of cancer. In: Chinsembu KC, Cheikhyoussef A, Mumbengegwi DR, Kandawa-Schulz M, Kasanda CD, Kazembe L, editors. Indigenous knowledge of Namibia. University of Namibia Press, Windhoek. 2015; pp. 63-88.
- Khumalo GP, Van Wyk B-E, Feng Y, Cock IE. A review of the traditional use of southern African medicinal plants for the treatment of inflammation and inflammatory pain. J Ethnopharmacol. 2022;283:114436. https://doi.org/10.1016/j.jep.2021.114436
- Semenya SS, Maroyi. A. Source of plants, used by Bapedi traditional healers for respiratory infections and related symptoms in the Limpopo province, South Africa. J Biol Sci 2019;19(2):101-21. https://doi.org/10.3923/jbs.2019.101.121
- Mhlongo LS, Van Wyk B-E. Zulu medicinal ethnobotany: new records from the Amandawe area of KwaZulu-Natal, South Africa. S Afr J Bot 2019;122:266–90. https://doi.org/10.1016/ j.sajb.2019.02.012
- Luseba D, Tshisikhawe MP. Medicinal plants used in the treatment of livestock diseases in Vhembe region, Limpopo province, South Africa. J Med Plants Res 2013;7(10):593-601. https:// doi.org/10.5897/JMPR012.1213
- Selogatwe KM, Asong JA, Struwig M, Ndou RV, Aremu AO. A review of ethnoveterinary knowledge, biological activities and secondary metabolites of medicinal woody plants used for managing animal health in South Africa. Vet Sci 2021;8:228. https://doi.org/10.3390/vetsci8100228
- 49. Van der Merwe D, Swan GE, Botha CJ. Use of ethnoveterinary medicinal plants in cattle by Setswana-speaking people in the Madikwe area of the North West province of South Africa. J S Vet Association 2001;72:189–96. https://doi.org/10.4102/ jsava.v72i4.651
- 50. Arnold TH, Wells MJ, Wehmeyer AS. Khoisan food plants: Taxa with potential for future economic exploitation. In: Wickens GE, Goodin JR, Field DV, editors. Plants for arid lands. Proceedings of the Kew International Conference on Economic Plants for Arid Lands. Allen and Unwin, London: London. 1985; pp. 69-86.
- 51. Wehmeyer AS. Edible wild plants of southern Africa: Data on the nutrient contents of over 300 species. CSIR, Pretoria. 1986
- 52. Steyn T. The chemical constituents of *Ehretia rigida, Apodytes dimidiata* and *Ocotea kenyensis*. MSc Dissertation, University of Natal, Pietermaritzburg, 1998.
- Kgopa AH, Bulani SI, Wilhelmi BS, Brand JM. Antioxidant activity of selected plants of the Great Fish River Reserve, Eastern Cape, South Africa. Afr J Range Forage Sci 2010;27:109-12. https:// doi.org/10.2989/10220119.2010.503581
- Dzoyem JP, Eloff JN. Anti-inflammatory, anticholinesterase and antioxidant activity of leaf extracts of twelve plants used traditionally to alleviate pain and inflammation in South Africa. J Ethnopharmacol 2015;160:194–201. https://doi.org/doi: 10.1016/j.jep.2014.11.034
- 55. Mnikathi MMN. A comparative evaluation of the biological activities and phytochemical properties in *Ehretia obtusifolia* and *Ehretia rigida*. MSc Dissertation, University of KwaZulu-Natal, Pietermaritzburg. 2021
- Pretorius JC, Magama S, Zietsman PC, Van Wyk B-E. Growth inhibition of plant pathogenic bacteria and fungi by extracts from selected South African plant species. S Afr J Bot 2003;69 (2):186-92. https://doi.org/10.1016/S0254-6299(15)30344-6

- 57. Ramavhale TT, Mahlo SM. Antifungal activity of selected medicinal plants used for the treatment of "u wela" in Makhado Local Municipality, Limpopo province. S Afr J Bot 2017;109:366. https://doi.org/doi:10.1016/j.sajb.2017.01.166
- 58. Burkill HRM. The useful plants of West Tropical Africa. Royal Botanic Gardens, Kew, Richmond, London. 1994
- Lemmens RHMJ. *Ehretia cymosa* Thonn. In: Lemmens RHMJ, Louppe D, Oteng-Amoako AA, editors., Plant resources of tropical Africa 7(2): Timbers 2. Prota Foundation, Waganingen. 2012; pp. 327–29.
- Lewis WH, Avioli LV. Leaves of *Ehretia cymosa* (Boraginaceae) used to heal fractures in Ghana increase bone remodeling. Econ Bot 1991;45(2):281–2. https://doi.org/10.1007/BF02862055
- 61. Maundu P, Berger D, Ole Saitabau C, Nasieku J, Kipelian M, Mathenge S, Morimoto Y, Höft R. Ethnobotany of the Loita Maasai: Towards community management of the forest of the lost child: Experiences from the Loita Ethnobotany Project. People and Plants Working Paper 8, UNESCO, Paris. 2001.
- 62. Arnold TH, Prentice CA, Hawker LC, Snyman EE, Tomalin M, Crouch NR, Pottas-Bircher C. Medicinal and magical plants of southern Africa: An annotated checklist. National Botanical Institute, Pretoria. 2002
- Wondimu T, Asfaw Z, Kelbessa E. Ethnobotanical study of medicinal plants around 'Dheeraa' town, Arsi Zone, Ethiopia. J Ethnopharmacol 2007;112:152–61. https://doi.org/10.1016/ j.jep.2007.02.014
- 64. Dharani N. Field guide to common trees and shrubs of East Africa. Struik Nature, Cape Town. 2019.
- Maroyi A. Ehretia amoena Klotzsch (Ehretiaceae): Review of its medicinal uses, phytochemistry and pharmacological properties. Int J Res Pharmaceut Sci 2021;12(2):1292-99. https:// doi.org/10.26452/ijrps.v12i2.4679
- Maroyi A. Evaluation of medicinal uses, phytochemistry and biological activities of *Ehretia cymosa* Thonn. (Ehretiaceae). Int J Res Pharmaceut Sci 2021;12(2):1521-28. https:// doi.org/10.26452/ijrps.v12i2.4731
- Maroyi A. Medicinal uses, phytochemistry and biological activities of *Ehretia obtusifolia* Hochst. ex DC. (Ehretiaceae). Int J Res Pharmaceut Sci 2021;12(2):1345-52. https://doi.org/10.26452/ ijrps.v12i2.4688
- Borges CV, Minatel IO, Gomez-Gomez HA, Lima GPP. Medicinal plants: Influence of environmental factors on the content of secondary metabolites. In: Ghorbanpour M, Varma A, editors., Medicinal plants and environmental challenges. Springer, Cham. 2017; pp. 259-77. https://doi.org/10.1007/978-3-319-68717-9_15
- Araújo LU, Grabe-Guimarães A, Mosqueira VC, Carneiro CM, Silva -Barcellos NM. Profile of wound healing process induced by allantoin. Acta Cir Bras 2010;25(5):460-66. https:// doi.org/10.1590/s0102-86502010000500014
- Nogueira AO, Oliveira YIS, Adjafre BL, de Moraes MEA, Aragão GF. Pharmacological effects of the isomeric mixture of alpha and beta amyrin from *Protium heptaphyllum*: A literature review. Fundam Clin Pharmacol 2019;33(1):4-12. https:// doi.org/10.1111/fcp.12402
- Ambavade SD, Misar AV, Ambavade PD. Pharmacological, nutritional and analytical aspects of β-sitosterol: A review. Orient Pharm Exp Med 2014;14:193–211. https://doi.org/10.1007/ s13596-014-0151-9
- 72. Van Wyk B-E, Wink M. Phytomedicines, herbal drugs and plant poisons. Briza Publications, Pretoria. 2015.