

**REVIEW ARTICLE** 



# *Helichrysum kraussii* Sch.Bip.: Review on its medicinal uses, phytochemistry and pharmacological properties

# Alfred Maroyi

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

Email: amaroyi@ufh.ac.za

# 

#### **ARTICLE HISTORY**

Received: 01 February 2023 Accepted: 28 May 2023 Available online Version 1.0 : 30 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. *Helichrysum kraussii* Sch.Bip.: Review on its medicinal uses, phytochemistry and pharmacological properties. Plant Science Today (Early Access). https://doi.org/10.14719/ pst.2402

# Abstract

Helichrysum kraussii Sch.Bip. is a woody shrub that naturally occurs in grasslands and woodlands in South-Central Africa. The various parts of H. kraussii serve as components used in traditional medicines within the South Central Africa region. This review provides an overview of the existing literature on the medicinal uses, phytochemical composition, and pharmacological properties of H. kraussii. The study reveals that various parts of the plant, including branches, flowers, leaves, roots, seeds, twigs, and whole plant parts, are used in ritual practices and traditional medicine to treat a range of human ailments. These include venereal diseases, nasal congestion, chest pain, skin infections, tuberculosis, respiratory infections, and cough. Phytochemical analysis of H. kraussii demonstrates the presence of acylated flavonol glucoside, diterpenes, flavonoids, phloroglucinol, and terpenoids. The pharmacological assessments indicate that crude extracts and isolated phytochemical compounds from this species possess antioxidant, antibacterial, antifungal, antiviral, cytotoxic, and antiinflammatory activities. This mini review underscores the traditional uses, phytochemical composition, and pharmacological properties of *H. kraussii*. Based on the findings, it is recommended to conduct comprehensive ethno pharmacological evaluations of H. kraussii, focusing on phytochemistry, pharmacological properties, toxicological assessments, as well as in vivo and clinical research.

#### **Keywords**

Asteraceae, Compositae, Helichrysum kraussii, materia medica, traditional medicine

# Introduction

Helichrysum kraussii Sch.Bip. (Fig. 1) is a well-known plant species in South Central Africa, widely used for its traditional medicinal properties (1-3). It is a perennial shrub belonging to the Asteraceae family. The genus name Helichrysum Mill. originates from the Greek words "helios" meaning "sun" and "chryson" meaning "gold," referring to the characteristic "golden flowers" of this genus (3). The species name "kraussii" is a tribute to Christian Ferdinand Friedrich Krauss, a German traveler, scientist, and plant collector who explored the Cape and Natal regions in South Africa during the 19th century (3). *H. kraussii* is also known by several synonyms, including *H. steetzii* O.Hoffm., *Achyrocline batocana* Oliv. & Hiern., *A. steetzii* (Vatke) O.Hoffm., and *Gnaphalium kraussii* (Sch.Bip.) Sch.Bip. (4, 5). This plant species is naturally found in grasslands and woodlands in South Central Africa (Fig. 2). While considered invasive in most rangelands in Zimbabwe, *H. kraussii* is listed as a weed in South Africa, primarily spread by humans due to

#### 2 MAROYI ET AL.

its ornamental value (6-8). The local population in South Central Africa has acquired indigenous knowledge and skills to utilize the medicinal properties of *H. kraussii*. Medicinal plants, their extracts, and active components derived from them play a crucial role in traditional therapy (9). Therefore, it is essential to understand the phytochemical composition of *H. kraussii*, as these active compounds directly or indirectly contribute to disease prevention, treatment, and overall health maintenance. Hence, this review aims to gather and summarize information on the medicinal uses, phytochemistry, and pharmacological properties of *H. kraussii*.



**Figure 1**. *Helichrysum kraussii* showing the habit of the species (Photo: BT Wursten)





utilized (refer to Table 1). Additionally, traditional sources like books, journal articles, dissertations, book chapters, theses, and scientific articles from the University library were consulted. To facilitate the search, keywords such as "Helichrysum kraussii," "biological activities of Helichrysum kraussii," "pharmacological properties of Helichrysum kraussii," "ethnobotany of Helichrysum kraussii," "medicinal uses of *Helichrysum kraussii*," "phytochemistry of Helichrysum kraussii," and "traditional uses of Helichrysum kraussii" were employed. The chemical structures of phytochemical compounds isolated from H. kraussii were depicted using the ChemSketch program. Only literature sources that evaluated the medicinal uses, phytochemistry, pharmacological, or biological activities of H. kraussii were included in this review. Excluded from this review were articles that were only partially accessed, such as abstracts, as well as published or unpublished ethnopharmacological surveys lacking information on the medicinal uses, phytochemistry, pharmacological, or biological activities of *H. kraussii*.

Table 1. Website links for all databases used in this study

| Name of database | Website link                       |
|------------------|------------------------------------|
| Google Scholar   | http://scholar.google.com          |
| JSTOR            | https://www.jstor.org              |
| PubMed           | http://www.ncbi.nlm.nih.gov/PubMed |
| Science Direct   | http://www.science direct.com      |
| Scopus           | http://scopus.com                  |

#### Habitat and morphology

Helichrysum kraussii, a woody shrub, reaches a maximum height of 1.5 meters (4, 10). Its sturdy stems branch repeatedly (Fig. 1), displaying a brownish color. The stems are finely woolly, becoming less hairy with age, and are adorned with leaves. The leaves of H. kraussii are linear and positioned along the smaller branches. They spread out, resembling narrow needles, and have a smooth underside while being woolly and white on the upper surface. The leaves are sessile, with slightly rolled-under margins, a sharply pointed apex, and occasional fluffy galls on the branchlets. The cylindrical capitula, arranged in a corymbose manner, occur at the tips of the branches. The flowers of H. kraussii form dense terminal heads and exhibit pale yellow to straw-colored hues. This species is documented in Angola, Botswana, Eswatini, Malawi, Mozambigue, South Africa, Zambia, and Zimbabwe (5, 10) within altitudes ranging from 5 to 1850 meters above sea level (5, 10). It is found in submontane grasslands, escarpment woodlands, rocky ridges on the highveld, as well as on poor soils and overgrazed land.

#### Ethnobotanical and ethno pharmacological uses

Helichrysum kraussii is mentioned in the monograph "Medicinal and magical plants of southern Africa: An annotated checklist" (11). It is also sold in informal herbal medicine markets in South Africa's Limpopo province (12-14) and is highly valued as a traditional medicine source. KwaZulu-Natal diviners in South Africa burn *H. kraussii* as incense (15, 16), and religious beliefs have ensured its careful harvesting to preserve the species. Additionally, the woolly stems of *H. kraussii* are used for making twig brooms in South Africa (17). Brooms are widely traded and commonly purchased by rural households instead of being made at home (17).

*H. kraussii* is used for traditional medicines in Eswatini, Malawi, Mozambique, South Africa, and Zimbabwe, accounting for 62.5% of the countries where the species is native (Table 2). The plant parts, including branches, flowers, leaves, roots, seeds, twigs, and whole plants, are used to treat and manage 14 human diseases and ailments in southern Africa. Crude extracts of *H. kraussii* are primarily used to treat venereal diseases, as indicated by a single country record and three literature sources. Other common uses include treating blocked

Table 2. Medicinal uses of Helichrysum kraussii

treating fever, headache, and temporary blindness (18-20). In combination with the leaves of Warburgia salutaris (G. Bertol.) Chiov., the leaves of H. kraussii are used as traditional medicine for asthma when taken orally as a decoction (21). When combined with the branches of Lippia javanica (Burm.f.) Spreng. and the leaves of Trichilia emetica Vahl, the branches of H. kraussii are used as a decoction to alleviate blocked nose, chest pains, and cough (22-26). In Zimbabwe, the roots or whole plant parts of *H. kraussii* are burned, mixed with salt, and taken orally as traditional medicine for coughs (1, 27). While the practice of combining H. kraussii with other medicinal plants is common in south-central Africa, research on Chinese herbal medicines has shown that multi-herbal and/or drug-herbal combinations do not necessarily lead to synergistic effects (28).

| Region                       | Plant part                                      | Mode of preparation   | Pharmacological<br>action or traditional<br>uses | Reference                                  |
|------------------------------|---|---|--|--|
| South Africa                 | Leaves  | Leaves mixed with those of <i>Warburgia salutaris</i> (G. Bertol.)<br>Chiov. and decoction taken orally                             | Asthma   | (21)                                       |
| South Africa                 | Leaves  | Leaf smoke inhaled  | Blocked nose                                     | (24, 29, 30)                               |
| South Africa                 | Branches  | Branches mixed with <i>Lippia javanica</i> (Burm.f.) Spreng. and<br><i>Trichilia emetica</i> Vahl leaves and decoction taken orally | Blocked nose                                     | (23, 26)                                   |
| South Africa                 | Branches  | Branches mixed with <i>Lippia javanica</i> and <i>Trichilia emetica</i><br>leaves and decoction taken orally                        | Chest pains                                      | (23, 26)                                   |
| South Africa                 | Leaves  | Leaf decoction taken orally   | Chest pains                                      | (24, 29, 30)                               |
| South Africa                 | Branches  | Branches mixed with <i>Lippia javanica</i> and <i>Trichilia emetica</i><br>leaves and decoction taken orally                        | Cough  | (22-26)                                    |
| Zimbabwe                     | Roots and whole plant                           | Roots or whole plant burnt and salt added to the ashes and the mixture taken orally   | Cough  | (1, 27)                                    |
| South Africa                 | Flowers,<br>leaves and<br>seeds                 | Flower, leaf and seed infusion taken orally   | Cough  | (22-24, 29-37)                             |
| Mozambique                   | Leaves  | Leaf infusion taken orally  | Fever  | (20)                                       |
| South Africa                 | Whole plant                                     | Whole plant decoction taken orally  | Headache   | (19)                                       |
| Mozambique                   | Leaves  | Leaves burnt and smoke inhaled  | Intoxicant                                       | (20)                                       |
| Eswatini                     | Leaves and<br>twigs                             | Leaves and twigs used   | Pesticide  | (18)                                       |
| South Africa and<br>Zimbabwe | Leaves  | Leaf infusion taken orally  | Respiratory infections                           | (27, 38)                                   |
| South Africa                 | Leaves and<br>twigs                             | Leaves and twigs used   | Ritual and protective charm                      | (11, 16, 33)                               |
| South Africa                 | Leaves  | Leaf decoction applied topically  | Skin infections                                  | (22, 24, 33, 36,<br>37, 39)                |
| Mozambique                   | Leaves  | Leaf decoction applied topically  | Temporary blindness                              | (20)                                       |
| Malawi                       | Leaves and roots                                | Body washed with infusion of leaves and roots   | To drive away bad<br>spirits                     | (1, 22, 33)                                |
| South Africa                 | Flowers,<br>leaves,<br>seeds and<br>whole plant | Flower, leaf, seed and whole plant decoction taken orally   | Tuberculosis                                     | (22, 24, 29, 31-<br>33, 35, 36, 40-<br>46) |
| South Africa                 | Roots   | Root decoction taken orally   | Venereal diseases                                | (36, 47, 48)                               |

nose and chest pains (one country record and five literature records), skin infections (one country record and six literature records), tuberculosis (one country record and 15 literature records), respiratory infections (two country records and two literature records), and as ritual and protective charms (two country records and five literature records). It is also used to alleviate coughs (two country records and 17 literature records) (Fig. 3).

*H. kraussii* has additional medicinal applications, such as being used as an intoxicant and pesticide, and for





# Phytochemical constituents of Helichrysum kraussii

Chemical compounds from the aerial parts, flowers, and leaves of H. kraussii have been identified by various researchers (Fig. 4; Table 3). These compounds include flavonoids, kaurenoic acid, prenyl-butyrylphloroglucinol, and terpenoids (31, 35, 40, 49-51). Terpenoids and essential oils have pharmacological effects on both transmissible and non-transmissible diseases. Their antimicrobial, anti-inflammatory, anticancer. antispasmodic, and antidiabetic properties are welldocumented (52, 53). Similarly, flavonoid compounds have potential medicinal applications due to their anticancer, antibacterial, antioxidant, skin protective, antiinflammatory, cardio-protective, antiviral. and neuroprotective effects (54, 55).

**Figure 4.** Chemical structures of acylated flavonol glucosides, diterpene, flavonoids and terpenoids isolated from *Helichrysum kraussii* 



Table 3. Phytochemical composition of Helichrysum kraussii

# Biological activities of Helichrysum kraussii

# Antibacterial activity

Table 4 presents the antibacterial properties of various extracts and compounds derived from *H. kraussii*. Kaurenoic acid,  $\alpha$ -amyrin,  $\beta$ -amyrin, and essential oils obtained from *H. kraussii* demonstrated antibacterial effects (31, 40, 51, 56). The acetone extracts derived from the aerial parts of *H. kraussii* displayed antibacterial activity (32). Antibacterial activities were observed in the chloroform: methanol extracts of *H. kraussii* leaves and stems (34). The ethanol extracts of *H. kraussii* flowers, leaves, and stems exhibited antibacterial properties (39). Moreover, water extracts from the leaves and stems of *H. kraussii* also showed antibacterial activity (56).

### **Antifungal activities**

.

. .

Table 5 presents the antifungal activities of various extracts and essential oils obtained from *H. kraussii*. The essential oils derived from the leaves and stems of *H. kraussii* demonstrated antifungal effects (56). Antifungal activities were also observed in the acetone extracts of the aerial parts of *H. kraussii* (32), as well as in the dichloromethane:methanol and aqueous extracts from the leaves and stems of the plant species (56).

| Type of compound               | Name of the compound                    | Plant part from<br>where extracted/<br>identified/<br>documented | Process of detection/<br>isolation/extraction | Reference  |
|--------------------------------|---|--|---|------------|
| Flavonoid                      | 3',4',5,7-tetrahydroxy-3-methoxyflavone | Flowers  | GC-MS   | 35, 49, 50 |
| Flavonoid                      | 5,6-dihydroxy-3,7,8-trimethoxyflavone   | Flowers  | GC-MS   | 35         |
| Flavonoid                      | 5,7-dihydroxy-3-methoxyflavone          | Flowers  | GC-MS   | 35         |
| Diterpene                      | Kaurenoic acid                          | Aerial parts   | GC-MS   | 31         |
| Phloroglucinol                 | Prenyl-butyrylphloroglucinol            | Aerial parts   | H-and C-NMR and El-MS                         | 31         |
| Acylated flavonol glucoside    | 3,5-dihydroxy-6,7,8-trimethoxyflavone   | Flowers  | H-and C-NMR and El-MS                         | 49, 50     |
| Acylated flavonol glucoside    | Helichrysoside                          | Flowers  | H-and C-NMR and El-MS                         | 49, 50     |
| Acylated flavonol glucoside    | L(-)-2-O-methylinositol                 | Flowers  | H-and C-NMR and El-MS                         | 49, 50     |
| Acylated flavonol<br>glucoside | Quercetin-3β-⊠-(Þ-coumaroyl)glucoside   | Flowers  | H-and C-NMR and El-MS                         | 49         |
| Monoterpene                    | 1,8-Cineole                             | Aerial parts   | GC and GC-MS                                  | 51         |
| Monoterpene                    | <b>α-</b> Pinene                        | Aerial parts   | GC and GC-MS                                  | 51         |
| Monoterpene                    | α-Terpineol                             | Aerial parts   | GC and GC-MS                                  | 51         |
| Triterpene                     | α- and β-Amyrin                         | Leaves   | H-and C-NMR and GC-MS                         | 40         |
| Bicyclic sesquiterpene         | Trans-α-Bergamotene                     | Aerial parts   | GC and GC-MS                                  | 51         |
| Bicyclic sesquiterpene         | δ- and ⊠-Cadinene                       | Aerial parts   | GC and GC-MS                                  | 51         |
| Bicyclic sesquiterpene         | β-Caryophyllene                         | Aerial parts   | GC and GC-MS                                  | 51         |
| Bicyclic sesquiterpene         | β-Caryophyllene oxide                   | Aerial parts   | GC and GC-MS                                  | 51         |
| Bicyclic sesquiterpene         | α- and β-Selinene                       | Aerial parts   | GC and GC-MS                                  | 51         |
| Monocyclic sesquiterpene       | α-Humulene                              | Aerial parts   | GC and GC-MS                                  | 51         |
| Tricyclic sesquiterpene        | <b>α-</b> Copaene                       | Aerial parts   | GC and GC-MS                                  | 51         |
| Tricyclic sesquiterpene        | ⊠-Gurjunene                             | Aerial parts   | GC and GC-MS                                  | 51         |
| Tricyclic sesquiterpene        | β-Sesquiphellandrene                    | Aerial parts   | GC and GC-MS                                  | 51         |
| Tricyclic sesquiterpene        | Viridiflorol                            | Aerial parts   | GC and GC-MS                                  | 51         |
| Tricyclic sesquiterpene        | Widdrene                                | Aerial parts   | GC and GC-MS                                  | 51         |

Table 4. Summary of antibacterial activities of the extracts and compounds isolated from different parts of Helichrysum kraussii

|                                | <b>,</b>                        |                            |  |           |
|--------------------------------|---------------------------------|----------------------------|--|-----------|
| Extract/<br>compound           | Plant part                      | Model                      | Effect   | Reference |
| Kaurenoic acid                 | Aerial parts                    | Agar diffusion             | Showed activities against <i>Escherichia coli</i> with minimum inhibitory concentration (MIC) value of 1.0 µg/ml) and MIC value of 10.0 µg/ml) against <i>Bacillus cereus, Bacillus subtilis, Staphylococcus aureus</i> and                  | 31        |
| Acetone                        | Aerial parts                    | Agar diffusion             | Showed activities against Bacillus subtilis, Bacillus cereus, Pseudomonas<br>aeruginosa, Bacillus pumilus, Micrococcus kristinae and Staphylococus<br>aureus with MIC value of 1.0 mg/ml   | 32        |
| Essential oils                 | Aerial parts                    | Dilution<br>technique      | Showed activities against Staphylococcus aureus, Escherichia coli,<br>Klebsiella pneumoniae, Enterobacter cloacae, Pseudomonas aeruginosa<br>and Staphylococcus epidermidis with MIC values ranging from 0.073 mg/<br>ml to >20.000 mg/ml    | 51        |
| Terpenoids                     | Leaves                          | TLC<br>bioautography       | Terpenoids α-amyrin and β-amyrin exhibited activities against <i>Bacillus</i><br><i>cereus</i>   | 40        |
| Chloroform :<br>methanol       | Leaves and stems                | Microdilution<br>technique | Showed activities against <i>Staphylococcus aureus, Staphylococcus</i><br>epidermidis, Klebsiella pneumoniae and Bacillus cereus with MIC values<br>ranging from 0.004 mg/ml to 4.0 mg/ml  | 34        |
| Dichlorometha<br>ne : methanol | Leaves and stems                | Micro-dilution             | Exhibited activities against Staphylococcus aureus, Klebsiella pneumoniae,<br>Mycobacterium smegmatis and Moraxella catarrhalis with MIC values<br>ranging from 1.3 mg/mL to 4.0 mg/mL   | 56        |
| Aqueous                        | Leaves and stems                | Micro-dilution             | Exhibited activities against Staphylococcus aureus, Klebsiella pneumoniae,<br>Mycobacterium smegmatis and Moraxella catarrhalis with MIC values<br>ranging from 2.67 mg/mL to 13.33 mg/mL  | 56        |
| Essential oils                 | Leaves and stems                | Micro-dilution             | Exhibited activities against Staphylococcus aureus, Klebsiella pneumoniae<br>and Moraxella catarrhalis with MIC values ranging from 3.0 mg/mL to<br>>16.0 mg/mL  | 56        |
| Ethanol                        | Flowers,<br>leaves and<br>stems | Micro-dilution             | Exhibited activities against <i>Propionibacterium acnes</i> with MIC value of 125.0 μg/ml  | 39        |
| Table 5. Summar                | y of antifungal                 | activities of the ex       | tracts and essential oils isolated from different parts of Helichrysum kraussii  |           |
| Extract/<br>compound           | Plant<br>part                   | Model                      | Effect   | Reference |
| Acetone                        | Aerial<br>parts                 | Agar diffusion             | Showed activities against Cladosporium cladosporioides, Aspergillus<br>niger, Cladosporiums sphaerospermum, Phytophthora capsici, Aspergillus<br>flavus and Cladosporium cucumerinum with MIC values ranging from<br>0.01 mg/ml to 1.0 mg/ml | 32        |
| Dichloro-<br>methane :         | Leaves<br>and                   | Micro-dilution             | Exhibited activities against <i>Cryptococcus neoformans</i> with MIC value of  | 56        |

0.83 mg/ml methanol stems Leaves Exhibited activities against Cryptococcus neoformans with MIC value of Micro-dilution 56 Ageous and 16.0 mg/ml stems Leaves Exhibited activities against Cryptococcus neoformans with MIC value of 56 Essential oils and Micro-dilution 1.0 mg/ml stems

#### **Antiviral activities**

The antiviral activities of H. kraussii flowers, leaves, and stems against the herpes simplex virus type-1 (HSV-1) using the cytopathic effect (CPE) inhibition assay with acyclovir as the positive control have been assessed (48). The extract exhibited potential anti-viral activities at 50.00 µg/ml with 100% viral inhibition when tested at the highest viral dose (100TCID50), which was comparable to the antiviral activities exhibited by the positive control, which showed 100% inhibition at 1.00 µg/ml (48). In another study, the antiviral activities of acetone, dichloromethane, hexane, and methanol: water extracts of H. kraussii leaves using a colorimetric cell-based (HeLa-SXR5) assay have been evaluated (37). The acetone, dichloromethane, and hexane extracts exhibited potential anti-viral activities at 2.5 µg/ml with 97.0% HIV inhibition in the cell-based assay (37).

#### Anti-inflammatory activities

The anti-inflammatory activities of the flavonoid 5,6dihydroxy-3,7,8-trimethoxyflavone isolated from the flowers of *H. kraussii* were assessed using a cyclooxygenase-1 and -2 (COX-1 and COX-2) catalyzed reaction of the arachidonic acid-prostaglandin pathway with indomethacin as the positive control (35). The flavonoid compound exhibited prostaglandin synthesis inhibition of 54.4% and 84.7% against COX-1 and COX-2, respectively (35). Similarly, the anti-inflammatory activities of ethanol extracts of *H. kraussii* flowers, leaves, and stems were assessed using the COX-2 assay with ibuprofen as a positive control (48). The extract exhibited weak activities with COX-2 inhibition at 10.0  $\mu$ g/ml, showing an inhibition of 57.2% compared to the 90.2% exhibited by the positive control (48).

#### **Antioxidant activities**

The flowers of *H. kraussii* were analyzed to determine the antioxidant activities of three flavonoids: 3',4',5,7tetrahydroxy-3-methoxyflavone, 5,6-dihydroxy-3,7,8trimethoxyflavone, and 5,7-dihydroxy-3-methoxyflavone. The assessment was conducted using the  $\alpha,\alpha$ -diphenyl- $\beta$ picrylhydrazyl (DPPH) free radical scavenging assay (35). The results indicated that these flavonoids demonstrated weak antioxidant activities, ranging from 40.7% to 44.4% (35). In another experiment, the antioxidant activities of ethanol and methanol extracts obtained from the flowers, leaves, and stems of H. kraussii were evaluated. The DPPH free radical scavenging assay was employed, with vitamin C serving as the positive control (39). Both extracts exhibited antioxidant activities, with half maximal inhibitory concentrations (IC50) values ranging from 4.0  $\mu$ g/ml to 4.2  $\mu$ g/ml (39). Moreover, the ethanol extracts of H. kraussii flowers, leaves, and stems were further

examined to determine their antioxidant activities using the DPPH radical scavenging and nitric oxide (NO) radical scavenging assays. Ascorbic acid was used as a positive control (48). The ethanol extract showed significant DPPH scavenging activity, with an IC50 value of 4.66  $\mu$ g/ml (36, 48).

# Cytotoxicity activities

Table 6 presents the cytotoxicity activities of various extracts derived from *H. kraussii*. Cytotoxic effects were observed in the chloroform: methanol extract obtained from both the leaves and stems of *H. kraussii* (34). The flowers, leaves, and stems of *H. kraussii* yielded ethanol extracts that displayed cytotoxic activities (39, 48).

The present review aims to provide comprehensive information on the medicinal uses, phytochemistry, biological activities, and botany of H. kraussii, encompassing literature published from 1962 to 2023 (34). H. kraussii is a member of the Helichrysum genus, which is widely utilized as a source of traditional medicines in southern Africa. Other species within this genus, such as H. caespititium (DC.) Harv., H. cymosum (L.) D.Don ex G.Don, H. longifolium DC., H. nudifolium (L.) Less., H. odoratissimum (L.) Sweet, H. pedunculatum Hilliard & B.L. Burtt, and H. petiolare Hilliard & B.L. Burtt, have also been documented for their medicinal properties (57-63). Some of these Helichrysum species, including H. nudifolium and H. odoratissimum, have commercial potential as ritual incense, sedatives, and herbal remedies in southern Africa, particularly for colds and chest pains. Additionally, their essential oils exhibit commercial potential for inhalation and aromatherapy purposes (64-66). These reports align with the findings of our review, which highlight the diverse medicinal uses of H. kraussii, varying across different countries.

The treatment of respiratory infections, such as asthma, blocked nose, chest pains, cough, and tuberculosis (TB), using herbal concoctions prepared from H. kraussii, is common in south central Africa. These reports of H. kraussii's use for similar ailments and diseases in different countries call for detailed ethnopharmacological research focusing on its effectiveness and safety (38, 67). Pharmacological studies on H. kraussii extracts and isolated phytochemical compounds have demonstrated various activities, including antioxidant, antibacterial, antifungal, antiviral, cytotoxicity, and anti-inflammatory effects. However,

there is a lack of information on both in vitro and in vivo studies specifically examining its anti-TB activities or tests against a panel of various Mycobacteria species. Respiratory infections are a significant public health concern in south central Africa, and traditional medicines have shown great potential in their treatment and management (43, 44, 68, 69). Therefore, it is important to understand the ethno pharmacological properties of H. kraussii extracts and phytochemical compounds isolated from the species, as these active ingredients may act directly or indirectly to prevent or treat diseases and maintain health. Evaluating the phytochemical properties of medicinal plants is essential not only to identify the main phytochemical compounds but also to gain a scientific understanding of the medicinal properties of the plant species (70). Medicinal plants like H. kraussii have been used for centuries in the treatment and management of various diseases, and their improved phytochemical profiling and ethno pharmacological research contribute to the development of current therapeutic systems. The pharmacological properties of traditional medicines depend on their primary and secondary phytochemical constituents (71).

Phytochemical compounds identified from various parts of H. kraussii include acylated flavonol glucosides, diterpenes, essential oils, flavonoids, phloroglucinol, and terpenoids (31, 35, 40, 49-51). Some of these secondary phytochemical constituents may be responsible for the documented biological activities of H. kraussii. Therefore, it is necessary to investigate the resulting pharmacological and toxicological properties of these phytochemical constituents in controlled clinical trials to determine their therapeutic potential and their association with specific diseases or indications related to the species. Although several phytochemical compounds have been isolated, purified, and characterized from *H. kraussii*, many of them still require detailed studies. So far, limited ethno pharmacological research has been conducted on the pharmacokinetics studies related to the mechanism of action of individual isolated phytochemical compounds under in vivo conditions. Ethno pharmacological research should also focus on toxicity, preclinical and clinical trials aimed at assessing the safety, effectiveness, and side effects of the phytochemical compounds in H. kraussii formulations used as traditional medicines.

Table 6. Summary of cytotoxicity activities of the extracts isolated from different parts of Helichrysum kraussii

| Extract/<br>compound     | Plant<br>part                      | Model  | Effect  | Reference |
|--------------------------|------------------------------------|--|---|-----------|
| Chloroform :<br>methanol | Leaves<br>and<br>stems             | Sulforhodamine B assay   | Exhibited activities against transformed human kidney<br>epithelial (Graham) cells with inhibition ranging from 9.0%<br>to 45.2%  | 34        |
| Ethanol                  | Flowers,<br>leaves<br>and<br>stems | 2,3-bis[2-methoxy-4-nitro-5-<br>sulfophenyl]-2H-tetrazolium-5<br>-carboxanilide reduction (XTT)<br>assay | Showed activities half maximal inhibitory concentrations $({\rm IC}_{\rm 50})$ value of 51.7 $\mu g/ml$   | 39        |
| Ethanol                  | Flowers,<br>leaves<br>and<br>stems | XTT assay  | Exhibited activities against human melanoma (A375),<br>epidermoid carcinoma (A431), cervical epithelial carcino-<br>ma (HeLa) and human embryonic kidney cells (HEK-293)<br>with IC50 values ranging from 34.9 µg/ml to 151.0 µg/ml | 48        |

# Conclusion

This mini-review summarizes the medicinal use, chemical composition, and pharmacological properties of H. kraussii. Such studies are crucial for plants commonly employed in traditional medicine, as they allow for the assessment of their phytochemistry, pharmacology, and toxicology. Notably, various phytochemical compounds have been isolated from H. kraussii, including acylated flavonol glucosides, diterpenes, essential oils, flavonoids, phloroglucinol, and terpenoids. These compounds have exhibited antibacterial, anti-inflammatory, and antioxidant activities. However, it is important to note that these studies are preliminary, and further ethno pharmacological research is needed. This research should focus on comprehensive evaluations, safety assessments, in vivo mechanisms of action, and clinical investigations to validate the traditional medicinal applications of H. kraussii. Due to the variations in phytochemical constituents depending on the origin and plant parts, it is crucial to establish a standardized phytochemical profiling protocol to isolate and obtain pure compounds from H. kraussii. Additionally, considering the diverse mechanisms of action exhibited by the phytochemical compounds mentioned in this review, future studies should prioritize the assessment of toxicity and safety using animal models.

### Acknowledgements

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

# **Authors' contributions**

SN and OS conceptualized and designed the study, and also drafted the manuscript. SN and CK conducted the research. UC performed statistical analysis, and also prepared the figures and tables. EG, CC and OE participated in the design, coordination of the research, and also helped draft the manuscript. All authors read and approved the final manuscript.

#### **Compliance with ethical standards**

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

Ethical issues: None .

# References

- 1. Gelfand M, Mavi S, Drummond RB, Ndemera B. The traditional medical practitioner in Zimbabwe: His principles of practice and pharmacopoeia. Gweru: Mambo Press; 1985.
- 2. Van Wyk B-E, Van Oudtshoorn B, Gericke N. Medicinal plants of South Africa. Pretoria: Briza Publications; 2017.
- Schmidt E, Lotter M, McCleland W. Trees and shrubs of Mpumalanga and Kruger National Park. Johannesburg: Jacana Media; 2017.
- Hyde MA, Wursten BT, Ballings P, Palgrave CM. Flora of Zimbabwe: Species information: *Helichrysum kraussii* Sch. Bip. [Internet]. 2023 [cited 18 January 2023].: https:// www.zimbabweflora.co.zw/speciesdata/species.php? species\_id=159680.

- POWO. Helichrysum kraussii Sch. Bip. Plants of the World Online. Facilitated by the Royal Botanic Gardens, Kew. 2023 [cited 23 January 2023]. https://powo.science.kew.org/ taxon/urn:lsid:ipni.org;names:212914-1.
- Wells MJ, Balsinhas VM, Joffe H, Engelbrecht VM, Harding G, Stirton CH. A Catalogue of problem plants in southern Africa, incorporating the national weed list of South Africa. Pretoria: Botanical Research Institute; 1986.
- 7. Randall RP. A global compendium of weeds. Department of Agriculture and Food, Western Australia; 2017.
- Gusha J. Invasive plant species of savanna rangelands: Are they a threat or an opportunity? [PhD Thesis]. Harare: University of Zimbabwe; 2018.
- Hedberg I, Staugård F. Traditional medicine in Botswana: Traditional medicinal plants. Gaborone: Ipeleng Publishers; 1989.
- Germishuizen G, Meyer NL. Plants of southern Africa: An annotated checklist. Pretoria: National Botanical Institute; 2003.
- 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. Pretoria: National Botanical Institute; 2002.
- Moeng TE. An investigation into the trade of medicinal plants by muthi shops and street vendors in the Limpopo province, South Africa. [MSc Dissertation]. Sovenga: University of Limpopo; 2010.
- Moeng TE, Potgieter MS. The trade of medicinal plants by muthi shops and street vendors in the Limpopo province, South Africa. J Med Plants Res. 2011;5(4):558–64.
- 14. Dzoyem JP, Tshikalange E, Kuete V. Medicinal plants market and industry in Africa. In: Kuete V, editors. Medicinal plant research in Africa: Pharmacology and chemistry. London: Elsevier; 2013. p. 859-91.
- Cooper K. The conservation status of the indigenous forests of the Transvaal, Natal and O.F.S. Durban: Wildlife Society of Southern Africa; 1983.
- Cunningham AB. African medicinal plants: Setting priorities at the interface between conservation and primary health care. Paris: UNESCO; 1993. People and Plants working paper 1.
- Shackleton SE. The significance of the local trade in natural resource products for livelihoods and poverty alleviation in South Africa. [PhD Thesis]. Grahamstown: Rhodes University; 2005.
- Long C. Swaziland's Flora: SiSwati names and uses. Mbambane: Swaziland National Trust Commission; 2005. [Internet]. Available from: http://www.sntc.org.sz/index.asp [cited 14 January 2023].
- Mashiloane MRR. Medicinal plants used by Mamotintana community in Mankweng, Limpopo province. [BSc Honours Dissertation]. Sovenga: University of Limpopo; 2010.
- Sitoe E. Medicinal ethnobotany of Mozambique: A review and analysis. [MSc Dissertation]. Johannesburg: University of Johannesburg; 2020.
- Ramarumo LJ, Maroyi A, Tshisikhawe MP. Warburgia salutaris (G. Bertol.) Chiov.: An endangered therapeutic plant used by the Vhaven⊠a ethnic group in the Soutpansberg, Vhembe Biosphere Reserve, Limpopo province, South Africa. Res J Pharm Tech. 2019;12(12):5893-8. https:// doi.org/10.5958/0974-360X.2019.01022.9
- 22. Lourens ACU, Viljoen AM, Van Heerden FR. South African Helichrysum species: A review of the traditional uses,

8 MAROYI ET AL.

biological activity and phytochemistry. J Ethnopharmacol. 2008;119:630–52. https://doi.org/10.1016/j.jep.2008.06.011

- York T, de Wet H, van Vuuren SF. Plants used for treating respiratory infections in rural Maputaland, KwaZulu-Natal, South Africa. J Ethnopharmacol. 2011;135:696–710. https:// doi.org/10.1016/j.jep.2011.03.072
- 24. York T. An ethnopharmacological study of plants used for treating respiratory infections in rural Maputaland. MSc Dissertation, University of Zululand, Kwa-Dlangezwa. 2012.
- Maroyi A. Lippia javanica (Burm.f.) Spreng.: Traditional and commercial uses and phytochemical and pharmacological significance in the African and Indian subcontinent. Evidence -Based Compl Alt Med. 2017;2017:6746071. http:// dx.doi.org/10.1155/2017/6746071
- 26. Van Vuuren SF, Motlhatlego KE, Netshia V. Traditionally used polyherbals in a southern African therapeutic context. J Ethnopharmacol. 2022;114977. https://doi.org/10.1016/ j.jep.2022.114977
- Nyagumbo E, Pote W, Shopo B, Nyirenda T, Chagonda I, Mapaya RJ, Maunganidze F, Mavengere WN, Mawere C, Mutasa I, Kademeteme E, Maroyi A, Taderera T, Bhebhe M. Medicinal plants used for the management of respiratory diseases in Zimbabwe: Review and perspectives potential management of COVID-19. Physics Chem Earth. 2022;128:103232. https://doi.org/10.1016/j.pce.2022.103232
- Zhou X, Seto SW, Chang D, Kiat H, Razmovski-Naumovski V, Chan K, Bensoussan A. Synergistic effects of Chinese herbal medicine: A comprehensive review of methodology and current research. Front Pharmacol. 2016;7:201. https:// doi.org/10.3389/fphar.2016.00201.
- 29. Watt JM, Breyer-Brandwijk MG. The medicinal and poisonous plants of southern and eastern Africa. Livingstone, London. 1962.
- Semenya SS, Maroyi A. Data on medicinal plants used to treat respiratory infections and related symptoms in South Africa. Data Brief. 2018;21:419-23. https://doi.org/10.1016/ j.dib.2018.10.012
- Bremner PD, Meyer JJM. Prenyl-butyrylphloroglucinol and kaurenoic acid: Two antibacterial compounds from Helichrysum kraussii. S Afr J Bot. 2000;66:115–17. https:// doi.org/10.1016/S0254-6299(15)31072-3
- 32. Mathekga ADM. Antimicrobial activity of Helichrysum species and the isolation of a new phloroglucinol from Helichrysum caespititium. PhD Thesis, University of Pretoria, Pretoria. 2001.
- Lourens ACU. Structural and synthetic studies of sesquiterpenoids and flavonoids isolated from Helichrysum species. PhD Thesis. University of KwaZulu Natal, Pietermaritzburg. 2008.
- Lourens ACU, Van Vuuren SF, Viljoen AM, Davids H, Van Heerden FR. Antimicrobial activity and in vitro cytotoxicity of selected South African Helichrysum species. S Afr J Bot. 2011;77:229–35. https://doi.org/10.1016/j.sajb.2010.05.006
- Legoale PB, Mashimbye MJ, Van Ree T. Anti-inflammatory and antioxidant flavonoids from Helichrysum kraussii and H. odoratissimum flowers. Nat Prod Comm. 2013;8(10):1403-4.
- Twilley D. Evaluating the anticancer and antioxidant activity of southern African plants and their mechanism of action. MSc Dissertation, University of Pretoria, Pretoria. 2017.
- Yardi SE. Metabolomic analysis on anti-HIV activity of selected Helichrysum species. PhD Thesis, University of Pretoria, Pretoria. 2019.
- 38. 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

- de Canha MN. Antimicrobial and anti-inflammatory effect of southern African plants against Propionibacterium acnes. MSc Dissertation, University of Pretoria, Pretoria. 2014.
- Prinsloo G, Meyer MJJM. In vitro production of phytoalexins by Helichrysum kraussii. S Afr J Bot. 2008;72:482-3. https:// doi.org/10.1016/j.sajb.2006.01.001
- 41. McGaw LJ, Lall N, Meyer JJM, Eloff JN. The potential of South African plants against Mycobacterium infections. J Ethnopharmacol. 2008;119:482–500. doi:10.1016/ j.jep.2008.08.022
- 42. Chinsembu KC. Tuberculosis and nature's pharmacy of putative anti-tuberculosis agents. Acta Trop. 2016;153:46–56. doi:10.1016/j.actatropica.2015.10.004
- 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. doi: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. doi:10.13057/biodiv/d200325
- 45. Cock IE, Van Vuuren SF. The traditional use of southern African medicinal plants for the treatment of bacterial respiratory diseases: A review of the ethnobotany and scientific evaluations. J Ethnopharmacol. 2020;263:113204. doi:10.1016/j.jep.2020.113204
- 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. doi:10.1016/j.hermed.2020.100390
- 47. Mabogo DEN. The ethnobotany of the Vhavenda. MSc Dissertation, University of Pretoria, Pretoria. 1990.
- Twilley D, Langhansová L, Palaniswamy D, Lall N. Evaluation of traditionally used medicinal plants for anticancer, antioxidant, anti-inflammatory and anti-viral (HPV-1) activity. S Afr J Bot. 2017;112:494–500. doi:10.1016/j.sajb.2017.05.021
- Candy HA, Wright W. Helichrysoside: A new acylated flavonoid glycoside from Helichrysum kraussii Sch. Bip. J South Afr Chem Institute. 1975;28:215–19. Available from: https://hdl.handle.net/10520/AJA03794350\_1884
- Candy HA, Laing M, Weeks CM. The crystal and molecular structure of helichrysoside, a new acylated flavonoid glycoside from Helichrysum kraussii. Tetrahedron Lett. 1975;14:1211–14. doi:10.1016/S0040-4039(00)72097-1
- Bougatsos C, Meyer JJM, Magiatis P, Vagias C, Chinou I. Composition and antimicrobial activity of the essential oils of Helichrysum kraussii Sch. Bip. and H. rugulosum Less. from South Africa. Flavour Fragr J. 2003;18:48–51.
- Fokou JBH, Dongmo PMJ, Boyom FF. Essential oil's chemical composition and pharmacological properties. In: El-Shemy H, editor, Essential oils: Oils of nature, Intechopen, London; 2020, pp. 18-32. Available from: http://dx.doi.org/10.5772/ intechopen.86573
- Jugreet BS, Mahomoodally MF. Pharmacological properties of essential oil constituents and their mechanisms of action. In: Swamy M, editor, Plant-derived bioactives. Springer, Singapore; 2020, pp. 387-415. doi:10.1007/978-981-15-2361-8\_18
- 54. Tungmunnithum D, Thongboonyou A, Pholboon A, Yangsabai A. Flavonoids and other phenolic compounds from medicinal

plants for pharmaceutical and medical aspects: An overview. Medicines. 2018;5:93. doi:10.3390/medicines5030093

- 55. Ullah A, Munir S, Badshah SL, Khan N, Ghani L, Poulson BG, Emwas AH, Jaremko M. Important flavonoids and their role as a therapeutic agent. Molecules. 2020;25(22):5243. doi:10.3390/molecules25225243
- York T, Van Vuuren SF, De Wet H. An antimicrobial evaluation of plants used for the treatment of respiratory infections in rural Maputaland, KwaZulu-Natal, South Africa. J Ethnopharmacol. 2012;144:118–27. doi:10.1016/ j.jep.2012.08.038
- 57. Maroyi A. Helichrysum caespititium (DC.) Harv.: Review of its medicinal uses, phytochemistry and biological activities. J Appl Pharmaceut Sci. 2019;9:111-18. doi:10.7324/ JAPS.2019.90616
- Maroyi A. Helichrysum cymosum (L.) D. Don (Asteraceae): medicinal uses, chemistry and biological activities. Asian J Pharmaceut Clinical Res. 2019;12:19-26. doi:10.22159/ ajpcr.2019.v12i7.33771
- Maroyi A. Medicinal uses, biological and phytochemical properties of Helichrysum foetidum (L.) Moench (Asteraceae). Asian J Pharmaceut Clinical Res. 2019;12:13-18. doi:10.22159/ajpcr.2019.v12i7.33607
- Maroyi A. Helichrysum longifolium and H. pedunculatum: a comparative analysis of their medicinal uses, chemistry and biological activities. Asian J Pharmaceut Clinical Res. 2019;12:41-6. doi:10.22159/ajpcr.2019.v12i7.33684
- 61. Maroyi A. Helichrysum nudifolium (L.) Less.: review of its medicinal uses, phytochemistry and biological activities. J Pharm Nutr Sci. 2019;9:189-94. https://doi.org/10.29169/1927 -5951.2019.09.03.8
- 62. Maroyi A. A synthesis and review of medicinal uses, phytochemistry and biological activities of Helichrysum odoratissimum (L.) Sweet. Asian J Pharmaceut Clinical Res. 2019;12:15-23. http://dx.doi.org/10.22159/ ajpcr.2019.v12i8.33508

- Maroyi A. Helichrysum petiolare Hilliard & B.L. Burtt: review of its medicinal uses, phytochemistry and biological activities. Asian J Pharmaceut Clinical Res. 2019;12:32-7. http://dx.doi.org/10.22159/ajpcr.2019.v12i6.33417
- Van Wyk B-E. The potential of South African plants in the development of new medicinal products. S Afr J Bot. 2011;77:812-29. https://doi.org/10.1016/j.sajb.2011.08.011
- 65. Van Wyk B-E. A review of commercially important African medicinal plants. J Ethnopharmacol. 2015;176:118-34. https://doi.org/10.1016/j.jep.2015.10.031
- 66. Van Wyk B-E. A review of African medicinal and aromatic plants. In: Nefati M, Najjaa H, Máthé A, editors. Medicinal and Aromatic Plants of the World Africa. Dordrecht: Springer; 2017. p. 19-60.
- Osman M, Karat AS, Khan M, Meehan S-A, Von Delft A, Brey Z, Charalambous S, Hesseling AC, Naidoo P, Loveday M. Health system determinants of tuberculosis mortality in South Africa: a causal loop model. BMC Health Serv Res. 2021;21:388. https://doi.org/10.1186/s12913-021-06398-0
- Fatima S, Kumari A, Dwivedi VP. Advances in adjunct therapy against tuberculosis: deciphering the emerging role of phytochemicals. Med Comm. 2021;2:494–513. https:// doi.org/10.1002/mco2.82
- Gautam S, Qureshi KA, Jameel Pasha SB, Dhanasekaran S, Aspatwar A, Parkkila S, Alanazi S, Atiya A, Khan MMU, Venugopal D. Medicinal plants as therapeutic alternatives to combat Mycobacterium tuberculosis: a comprehensive review. Antibiotics. 2023;12:541. https://doi.org/10.3390/ antibiotics12030541
- 70. Van Wyk B-E, Wink M. Phytomedicines, Herbal Drugs and Plant Poisons. Pretoria: Briza Publications; 2015.
- Hussein RA, El-Anssary AA. Plants secondary metabolites: the key drivers of the pharmacological actions of medicinal plants. In: Builders PF (Ed), Herbal medicine. London: Intech Open; 2019. pp 11–30. https://doi.org/10.5772/ intechopen.76139