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MINI-REVIEW ARTICLE

Garcinia livingstonei T. Anderson: A Potential Source for Bioactive Constituents

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Abstract: *Garcinia livingstonei* T. Anderson (African mangosteen) is a member of family Clusiaceae and is native to South Africa. It is distributed from Malaysia to the Philippines, Burma and India. The plant is used traditionally in the treatment of diarrhea. It contains various chemical constituents belonging to bi-flavonoids, benzophenones and xanthenes. *Garcinia livingstonei* is reported to possess beneficial pharmacological activities as anti-oxidant, anti-bacterial, anti-viral and cytotoxicity. This review aims at summarizing the available literature on the botanical features, phytochemical, ethnomedicinal and pharmacological activities of *Garcinia livingstonei*.

Keywords: *Garcinia livingstonei*, morphological features, traditional uses, nutritive value, phytoconstituents, pharmacological activity.

1. INTRODUCTION

For thousands of years, natural compounds, mostly from plants, have been the mainstay of traditional medicine. They also have been the source of lead compounds for modern medicine [1]. *Garcinia livingstonei* belongs to the family Clusiaceae. Genus *Garcinia* includes more than 250 species distributed in the tropical and temperate regions of Southeast Asia, Central and South America [2, 3]. It is a dioecious evergreen tree with leathery leaves in opposite pairs or whorls and polygamous flowers. The fruit is edible, juicy with a slightly acidic and sweet apricot-like flavor and can be fermented to produce alcoholic beverages [4-7]. Traditionally, different parts of this plant are used for the treatment of cough, fever, parasitic diseases, toothache [8, 9] in addition to treatment of abdominal pains during pregnancy and aid in childbirth [10, 11]. Several secondary metabolites were reported in *Garcinia* species, including xanthenes, flavonoids, benzophenones, lactones and phenolic acids [12]. *G. livingstonei* T. Anderson is a rich source of biflavonoids, benzophenones and xanthenes [4, 13, 14]. *G. livingstonei* was reported to exhibit several therapeutic benefits, including antioxidant, anti-human immunodeficiency virus (HIV), antibacterial in addition to inhibition of melanin formation (skin whitening) which is beneficial in cosmetic preparation. Isolated compounds from *G. livingstonei* showed melanin inhibitory activity which increased activity of the plant as a cosmetic agent [4, 15, 16]. The aim of this review is to highlight the importance of this plant and its phytoconstituents.

2. MORPHOLOGICAL AND MICROSCOPICAL FEATURES

G. livingstonei is a dioecious evergreen tree. It reaches 10-16 meters in height, glabrous with three lateral branches at the node. Leaves are bluish-green in color, coriaceous and petiolate. The texture is leathery or waxy, and the arrangement is opposite pairs or whorls. Lamina shape varies from lanceolate to oblanceolate, obovate, oblong or orbicular. Margin is shallowly crenate. Apex is emarginate, apiculate or round to acute. Base is crenate to round. Fruits are orange-yellow, orange to reddish-orange in color, obovoid to globose in shape, filled with yellow latex, 1-2 seeded, having thin skin and its diameter is 10-40 mm. Flowers are polygamous, have long pedicels, they consist of fascicles; the axils of the older leaves have from five to fifteen or more, while old wood has from four to twenty, 35 mm in length, with five to eight obovate to orbicular petals that is greenish-white to pale yellow in color having longitudinal glandular lines orange to reddish in color; and four unequal sepals which are arranged in two opposite and decussate pairs. Stamens are free and numerous in male flowers and embedded in a fleshy cushion which is the result of united fascicledodes. Staminodes are few in the female flowers and embedded in a fleshy fasciclodal ring beneath the ovary, which is globose; locules number is two or three enclosed in a fleshy and bi-lobed stigma. Seeds number is one or two, 1.5 to 2 cm in length, and the shape of the seeds is cylinder or plano-ovoid. Outer bark of *G. livingstonei* is grey, but underneath the superficial layer is orange, smooth in younger stems, rough, with prominent vertical cracks in mature stem resembles a mosaic appearance, while the inner bark is yellowish red, relatively smooth, saturated with many small dots (exudates) that are dark reddish in color (Figs. 1 and 2) [5-7].

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Fig. (1). Tree of *G. livingstonei* cultivated in Egypt. (A higher resolution / colour version of this figure is available in the electronic copy of the article).



(a) Leaf (x=0.6)

(b) Flower (x=1.06)



(c) Fruit

Fig. (2). Different parts of *G. livingstonei*. (A higher resolution / colour version of this figure is available in the electronic copy of the article).

2.1. *Garcinia livingstonei* T. Andersson Synonyms

G. angolensis Vesque, *G. ferrandii* var. *affinis* Chiov., *G. ferrandii* Chiov., *G. pendula* Engl., *G. pallidinervia* Engl. [10].

2.2. Common English Names

African Mangosteen, Wild Mangosteen, Imbe, Inbe, Gupenja, Lowveld Mangosteen, Mwausungulu, Pama, Veld Mangosteen, Wild Plum, Livingstone's garcinia [6, 10].

2.3. Habitat and Distribution

African mangosteen originates in Sunda Island and the Moluccas or Spice Islands in Indonesia. *G. livingstonei* was

spreading from Malaysia to the Philippines and India. It was introduced in England in 1789, Sri Lanka in 1800 and Australia in 1854. The tree is widespread in many countries, including Egypt, the warmer parts of Africa, extending from the north of Durban to Somalia and Guinea. In Southern Africa, *G. livingstonei* spreads in Limpopo and Zambezi Valleys [5, 17, 18]. In South Africa, It spreads in different regions as scrub, open woodland and in the forest; while in Zimbabwe, the plant is found along the rivers in the Lowveld and in Riparian zone. In Zambia, it is in termite mounds, Munga areas and Mopane woodland. The plant is found on rocky soil and in open coastal forest (Fig. 3) [19].

2.4. Traditional Uses

The seeds, pulp, leaves and bark of *G. livingstonei* had been implicated traditionally in the treatment of diarrhea and digestive problems [7, 20, 21]. The dried roots were used to treat abdominal pains during pregnancy and aid in childbirth [10, 11]. While the stem bark was used as skin lightening agent [4]. Moreover, the fruits and stems were used in the treatment of cough, fever and parasitic diseases [9]. Bark was used to treat internal parasites [7]. In addition, root infusion was used as an aphrodisiac and for the treatment of impotency, while the decoction for the treatment of toothache. Also, when the stem was boiled and eyes were exposed to the vapors effective relief was achieved [8].

2.5. Nutritive Value and Proximate Analysis

G. livingstonei was reported to have high nutritional value. The fruits and seeds contain carbohydrates with content ranges (37.67 ± 0.02 - $95.02 \pm 0.01\%$), crude fiber (2.93 ± 0.01 - $21.13 \pm 0.02\%$), crude protein (0.65 ± 0.02 - 31.76 ± 0.21) and crude fat (1.23 ± 0.04 - $19.55 \pm 0.02\%$). Minerals were also detected, such as nitrogen (6800 - 12940 mg/kg), calcium (4200 - 7600 mg/kg), potassium (2753 - 10830 mg/kg), magnesium (1440 - 2900 mg/kg), phosphorus (391.72 - 577.79 mg/kg), iron (156.6 - 354.6 mg/kg), manganese (7.7 - 246.8 mg/kg), sulphur (4.48 - 21.64 mg/kg), zinc (14.2 - 40.6 mg/kg) and copper (9.7 - 354.6 mg/kg) [5]. This emphasizes the importance of edible fruits for healthy human growth [22, 23].

2.6. Seed Oil

G. livingstonei seed oil is brown in color and semisolid at room temperature. Its specific gravity was 0.90 ± 0.1 , while viscosity was $45.71 \pm 00 \text{ Nm}^2\text{s}$. The free fatty acid content was approximately 1.88% calculated as oleic acid, major unsaturated fatty acid was oleic acid (54.55%) followed by linoleic acid (5.67%) which was reported to have anti-inflammatory effects, play role in immune system and decrease the risk of coronary heart disease. On the other hand, the main saturated fatty acid was recognized as palmitic acid (32.64%) [5, 24, 25].

2.7. Phytochemical Studies

Phytochemical screening of *G. livingstonei* revealed the presence of phenolics and steroids in roots, stems and fruits [26], while the bark contains alkaloids, terpenoids, saponins and phenolics including flavonoids, anthraquinones and tan-

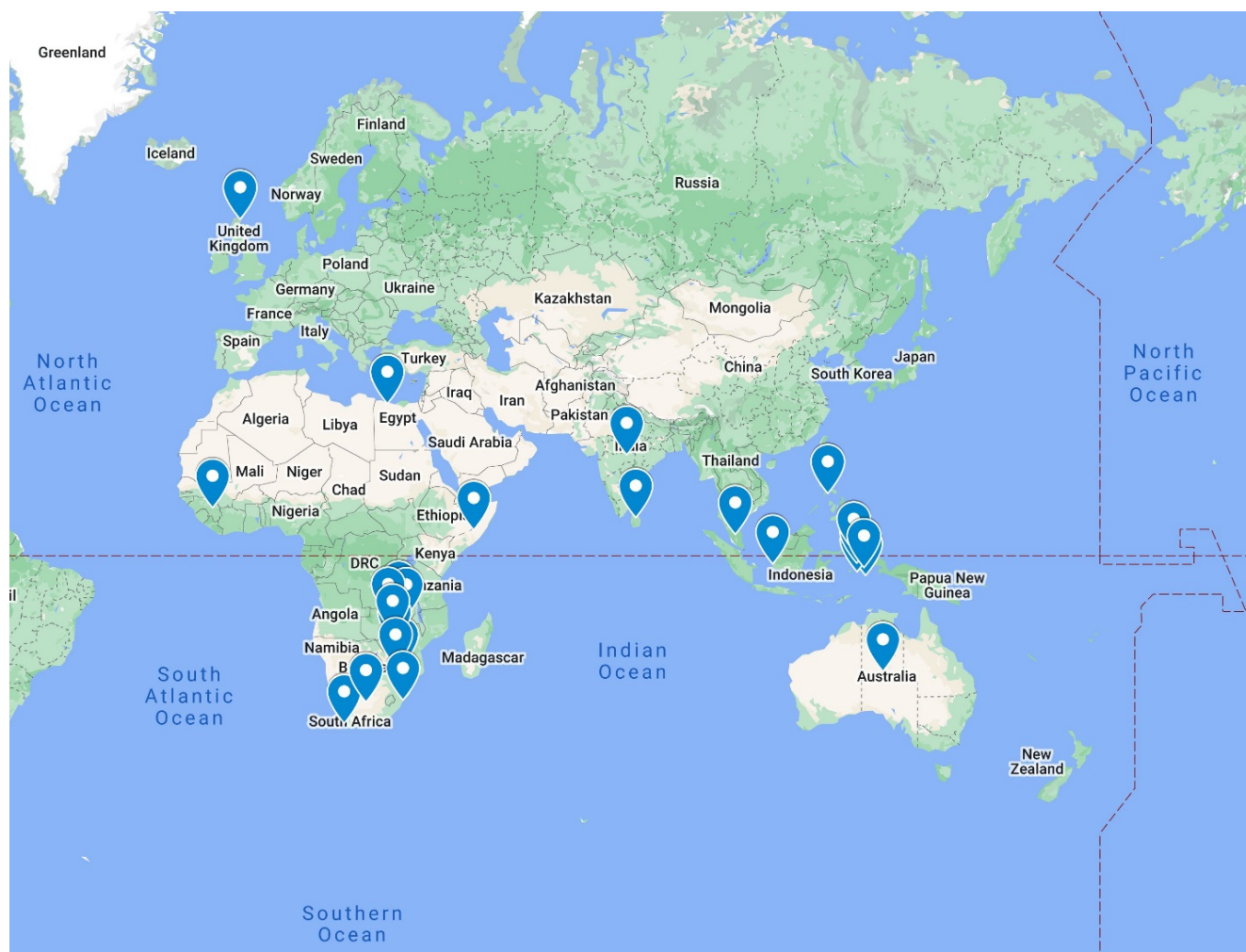


Fig. (3). Distribution of *G. livingstonei*. (A higher resolution / colour version of this figure is available in the electronic copy of the article).

Table 1. Reported total phenolics of different parts of *Garcinia livingstonei* T. Anderson.

| Plant Parts | Phenols (mg GAE/g) | Flavonoids ($\mu\text{g QE/g}$) | Tannin ($\mu\text{g/g}$) | References |
|-------------|--------------------|-----------------------------------|----------------------------|------------|
| Fruit | 115.5 ± 34.1 | - | - | [15] |
| Epicarp | 174.02 ± 0.17 | 49.85 ± 0.06 | 7.96 ± 0.02 | [5] |
| Mesocarp | 83.35 ± 0.08 | 55.32 ± 0.11 | 4.24 ± 0.01 | |
| Seed coat | 10.72 ± 0.00 | 19.25 ± 1.15 | 0.96 ± 0.00 | |
| Seed | 106.41 ± 0.15 | 99.98 ± 0.23 | 1.57 ± 0.00 | |

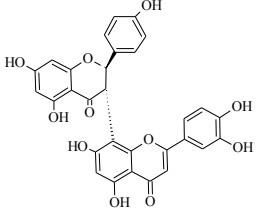
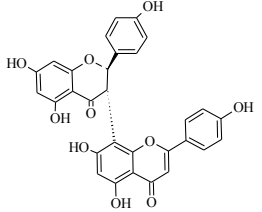
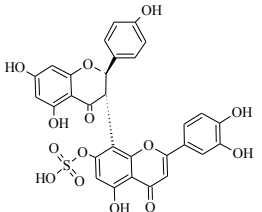
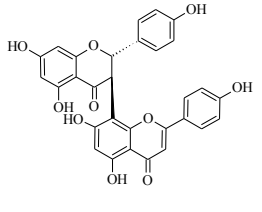
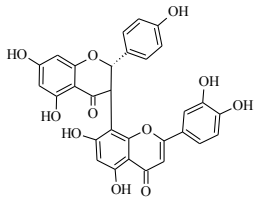
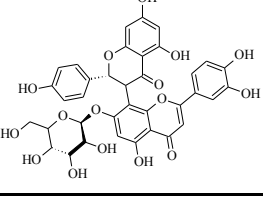
nins [21, 27]. Estimation of phenolic content of different parts of *G. livingstonei* is demonstrated in Table 1 [5, 15]. The total phenolic content of fruits was 115.5 ± 34.1 mg GAE/g of dry fruit; while the epicarp of fruit contains the highest phenolic and tannin contents (174.02 ± 0.17 mg GAE/g and 7.96 ± 0.02 $\mu\text{g/g}$, respectively); the highest flavonoids were reported in the seeds with values 99.98 ± 0.23 $\mu\text{g QE/g}$ [15]. Moreover, the oxalate content in the fruit epicarp was (0.03 ± 0.01 $\mu\text{g/g}$), while the highest amount was found in the seed (0.30 ± 0.19 $\mu\text{g/g}$). On the other hand, mesocarp and seed coat contains nearly the same amount (0.04 $\mu\text{g/g}$) [5].

Different classes of constituents reported in *G. livingstonei* as biflavonoids, benzophenones, xanthenes and triterpenes, are represented in Tables 2-6.

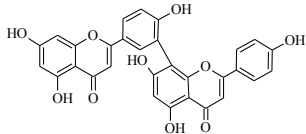
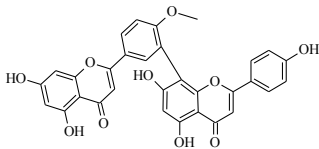
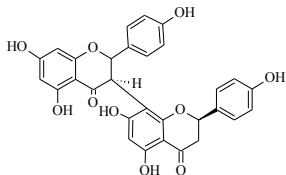
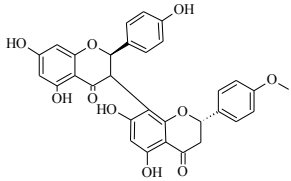
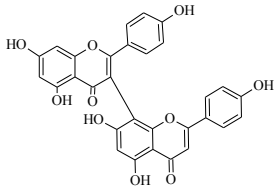
2.8. Biflavonoids

Biflavonoids are flavonoid dimers resulting from the connection of two units of flavone, flavanone, flavanone, flavonol or aurones or mixtures of these, in addition to less common chalcone and isoflavone dimers [28]. Several biological activities were reported for biflavonoids, including

Table 2. Reported biflavonoids from *Garcinia livingstonei* T. Anderson.

| Biflavonoids | | | | | |
|---|----------------------------|------------------------------------|---------------------------|------------|---|
| Compound Name | Plant Part | Extracts/Fractions | Geographical Distribution | References | Chemical Structures |
| Morelloflavone Type | | | | | |
| Morelloflavone | Heartwood, bark and leaves | Acetone extract | India | [32] |  |
| | Stem bark | Ethyl acetate/methanol extract | South Africa | [4] | |
| | Stem wood | Ethyl acetate and acetone extracts | Kasane | [33] | |
| | Fruits | Ethyl acetate fraction | Florida | [14] | |
| | Root bark | Ethyl acetate soluble extract | Tanzania | [13] | |
| BGH-111 (4',4'',5,5'',7,7''-hexahydroxy-3(8''-) flavonyl flavanone | Heartwood, bark and leaves | Acetone extract | India | [32] |  |
| Morelloflavone-7''-sulphate | Stem bark | Ethyl acetate/methanol extract | South Africa | [4] |  |
| Volkensiflavone | Fruits | Methanol extract | Florida | [15] |  |
| | | Ethyl acetate fraction | | [14] | |
| | Stem wood | Ethyl acetate and acetone extracts | Kasane | [33] | |
| | Root bark | Ethyl acetate soluble extract | Tanzania | [13] | |
| Fukugetin | Fruits | Methanol extract | Florida | [15] |  |
| Fukugiside | Fruits | Ethyl acetate fraction | Florida | [15] |  |
| | | Methanol extract | | [14] | |

(Table 2) Contd....

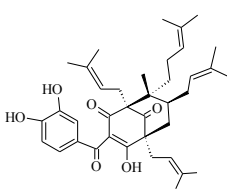
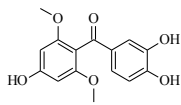
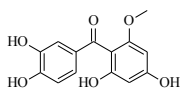
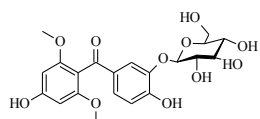
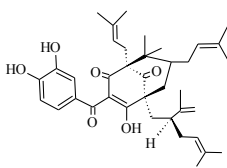
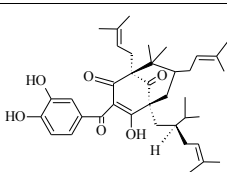
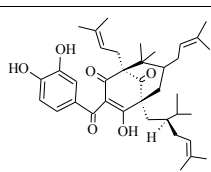
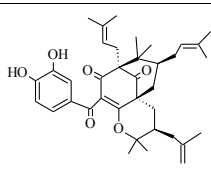
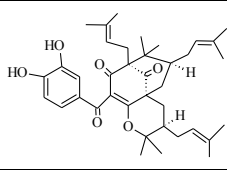
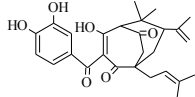
| Biflavonoids | | | | | |
|---|----------------------------|------------------------------------|---------------------------|-------------|---|
| Compound Name | Plant Part | Extracts/Fractions | Geographical Distribution | References | Chemical Structures |
| Amentoflavone Type | | | | | |
| Amentoflavone | Heartwood, bark and leaves | Acetone extract | India | [32] |  |
| | Fruits | Methanol extract | Florida | [15] | |
| | | Ethyl acetate fraction | | [14] | |
| | Leaves | Ethyl acetate fraction | South Africa | [9, 48, 51] | |
| 4'-monomethoxyamentoflavone(Podocarpus flavone A) | Twigs | Ethyl acetate extract | Kasane | [33] |  |
| | leaves | Ethyl acetate fraction | South Africa | [9, 48, 51] | |
| | Heartwood, bark and leaves | Acetone extract | India | [32] | |
| GB-1a Type | | | | | |
| GB-1a | Stem wood | Ethyl acetate and acetone extracts | Kasane | [33] |  |
| Ent-naringeninyl-(1-3 α ,II-8)-4'-O-methylnaringenin | Root bark | Ethyl acetate soluble extract | Tanzania | [13] |  |
| IC3'-IIC8'' Type | | | | | |
| 3,8''-biapigenin | Fruits | Ethyl acetate fraction | Florida | [14] |  |

antioxidant, antimicrobial, antiallergenic, anti-inflammatory, hepatoprotective and antiviral activities [29]. There are many types of biflavonoids depending on position and type of linkage between the two flavonoids [30, 31]. Reported biflavonoids in *G. livingstonei* are shown in Table 2. Morelloflavone type biflavonoids; namely morelloflavone was isolated from the heartwood and bark in India [32], fukugetin and fukugiside were isolated from the fruits in Florida [14, 15]. Amentoflavone type namely, amentoflavone and podocarpusflavone A were isolated from the leaf extract in India [32]. GB-1a type was isolated from the twigs and stem wood in Kasane [33], in addition, ent-naringeninyl-(1-3 α ,II-8)-4'-O-methylnaringenin was extracted from the root bark in Tanzania [13] and 3,8''-biapigenin that belongs to IC3'-IIC8'' type was isolated from the fruits in Florida [14].

2.9. Benzophenones

Most of the isolated benzophenones belong to polyisoprenylated benzophenones that have 13-carbon skeleton forming two rings connected by a carbonyl group. Benzophenones reported in family Clusiaceae are characterized by the presence of two to five isoprene units [34]. Benzophenones have several pharmacological activities as HIV inhibitory activity [35] and were used as skin lightening agents [4]. Benzophenones reported in *G. livingstonei* are demonstrated in Table 3. Guttiferone A was isolated from the fruit in Tanzania as the principal active constituent through bioassay guided fractionation of the HIV inhibitory activity [35] and Guttiferone A and E were also isolated from the fruits in Florida [14]. Moreover, 4,3',4'-trihydroxy-2,6-dimethoxybenzophenone and its glycoside; 3'- β -D-glucosyloxy-4,4'-dihydroxy-2,6-dimethoxybenzophenone were isolated

Table 3. Reported benzophenones from *Garcinia livingstonei* T. Anderson.

| Benzophenones | | | | | | | |
|---|------------|--------------------------------------|---------------------------|------------|---|--|--|
| Compound Name | Plant Part | Extracts/Fractions | Geographical Distribution | References | Chemical Structures | | |
| Guttiferone A | Fruits | Dichloromethane/ methanol extract | Tanzania | [35] |  | | |
| | Fruit peel | Ethyl acetate extract | South Africa | [4] | | | |
| | Fruits | Ethyl acetate fraction | Florida | [14] | | | |
| | Fruits | Methanol extract | | [15] | | | |
| 4,3',4'-trihydroxy-2,6-dimethoxybenzophenone | Twigs | Methanol extract | Kasane | [33] |  | | |
| 3',4,4',6-tetrahydroxy-2-methoxybenzophenone | | | | |  | | |
| 3'-β-D-glucosyloxy-4,4'-dihydroxy-2,6-dimethoxybenzophenone | Stem wood | Ethyl acetate and acetone extracts | | |  | | |
| Guttiferone E | Fruits | Methanol extract | Florida | [15] |  | | |
| Guttiferone K | | Ethyl acetate fraction | | [14] |  | | |
| Xanthochymol | | - | | |  | | |
| Cycloxanthochymol | | | | |  | | |
| Isoxanthochymol | | | | |  | | |
| Gambogenone | | | | |  | | |

together with 4,6,3',4'-tetrahydroxy-2-methoxybenzophenone [33]. Other benzophenones were also isolated, namely; guttiferone E and K, xanthochymol, cycloxanthochymol, isoxanthochymol and gambogone [14].

2.10. Xanthenes

The xanthone skeleton is a conjugated ring system composed of two aromatic rings B fused through a carbonyl group and an oxygen atom; the word "xanthone" is derived from the Greek word xanthos, meaning yellow. Xanthenes have diverse pharmacological properties, including antibacterial, antiviral, antioxidant, anti-inflammatory, anti-coagulant and cytotoxic activity against lung cancer [13, 36]. As shown in Table 4, xanthenes reported in *G. livingstonei* extracted by dichloromethane from the root bark of South Africa afforded 1,4,5-Trihydroxy-3-(3-methylbut-2-enyl)-9H-xanthen-9-one, 6,11-Dihydroxy-2,2-dimethyl-pyrano[3,2-c] xanthen-7(2H)-One, 4-(3',7'-dimethylocta-2', 6'-dienyl)-1,3,5-hydroxy-9H-xanthen-9-One, 6,11-Dihydroxy-3-methyl-3-(4-methylpent-3-enyl)-3H,7H-pyrano[2,3-c]xanthen-7-one and 12b-hydroxy-des-Dgarcigerrin A [37]. While garcilivin A, B and C were isolated from the root bark [38]. Alloathyriol and 6-deoxyisojacareubin, norathyriol and montixanthone were also reported in the twigs and stem wood [14, 33]. In addition, garcilivin A and C, dihydroxy-3-methyl-3-(4-methylpent-3-enyl)pyrano[2,3-c]xanthen-7(3H)-one, 4[(E)-3,7-dimethylocta-2,6-dienyl]-1,3,5-trihydroxy-9H-xanthen-9-one, and 1,4,5-trihydroxy-3-(3-methylbut-2-enyl)-9H-xanthen-9-one were isolated from the root bark [13].

2.11. Flavonoid

Flavonoid was represented by only eriodictyol obtained from the seeds of *G. livingstonei* [39]. Eriodictyol possesses anti-oxidant, anti-diabetic and anti-inflammatory activities [40]. This encourage further studies to reveal other related compounds in Table 5.

2.12. Triterpenes

Betulin, betulin aldehyde and other triterpenes were isolated from the stem of *G. livingstonei*, in addition to sargaol, which was isolated from the hexane fraction of the fruit kernels (Table 6) [4, 33].

2.13. The Pharmacological Activities

Reviewing the available literature, *G. livingstonei* reported benefits from almost all plant parts in addition to its isolated compounds (Tables 7-11). The plant showed wide antimicrobial activity against several micro-organisms such as bacteria, virus and fungi, in addition to anti-oxidant and cytotoxic activities [15, 26, 27, 41].

2.14. Anti-microbial Activity

Diseases caused by bacteria have a great effect on human health [42]. Nowadays, there is great interest in the usage of natural antibacterial remedies extracted from plants [43]. Other *Garcinia* species showed good activity as antibacterial against a broad spectrum of bacteria, for example, the twigs of *G. hombroniana* and the acetone extract and fractions of

G. goudotiana leaves and their isolated compounds exhibited potent antibacterial activity against Gram-positive bacteria [44, 45]; also, *G. kola* seed and some volatile constituents of *G. atroviridis* fruits were effective against Gram-positive and Gram-negative bacteria [46, 47]. Concerning *G. livingstonei*, published data revealed that the acetone extract of leaves showed good antibacterial activity against *Staphylococcus aureus*, *Enterococcus faecalis*, *Escherichia coli* and *Pseudomonas aeruginosa* with MIC (0.07, 0.04, 0.06 and 0.63 mg/ml, respectively). The fractions were also evaluated against the four bacteria and results showed that the most effective fraction against *E. coli* was chloroform fraction with MIC = 0.095 mg/ml; concerning *S. aureus*, the ethyl acetate extract (MIC = 0.195 mg/ml) was the best. The most effective fraction of *E. faecalis* and *P. aeruginosa* was Carbon tetrachloride fraction (MIC = 0.115 mg/ml) [48-50]. MIC values for the isolated 4'-methoxyamentoflavone were 8, 40, 60 and 8 µg/ml against *E. coli*, *S. aureus*, *P. aeruginosa* and *E. faecalis*, respectively. While, for amentoflavone the MIC values were 40, 40, 100 and 60 µg/ml, respectively [9]. MIC for amentoflavone and 4'-monomethoxyamentoflavone against *Mycobacterium smegmatis* were 0.60 ± 0.70 and 1.40 ± 1.56 mg/ml, respectively [51]. Aqueous extract of *G. livingstonei* bark showed MIC of 1.56 mg/mL on all the tested bacteria except for *E. coli*, and *Shigella sonnei* MIC was 0.78 mg/mL, while MIC of the methanol extract was 0.39 mg/mL for all tested bacteria with the exception of *E. coli*, and *S. sonnei* (MIC = 0.78 mg/mL); while MBC of the water and methanol extracts was 12.5 mg/mL for all the tested bacteria except for *S. aureus* MBC of methanol extract was 6.25 mg/mL [21]. The antimicrobial activity of the dichloromethane:methanol (1:1) extract of the bark was evaluated against twelve pathogens and the extract displayed an average broad-spectrum MIC value of 270 µg/mL [27]. Also, the methanol and dichloromethane extracts of the bark were screened against ten different bacteria and the lowest MIC were against *Bacillus cereus* which is equal 0.06 and 0.12 mg/mL, respectively [7].

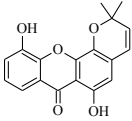
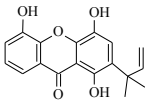
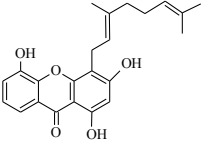
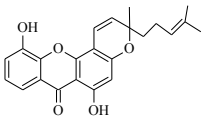
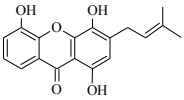
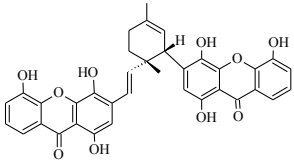
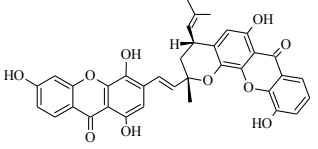
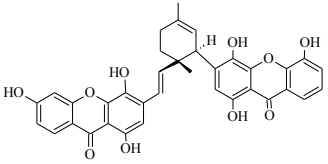
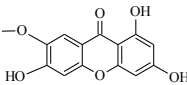
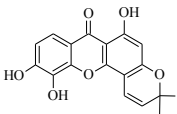
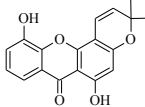
2.15. Therapeutic Index

Therapeutic index (TI) which is the ratio of cytotoxicity to MIC; for the water extract of the bark recorded the highest therapeutic index against *E. coli* and *S. sonnei* with value of 0.98; while for *S. aureus*, *K. oxytoca* and *S. enterica*; it was 0.49. Therapeutic index for methanolic bark extract could not be determined as the value of its cytotoxicity was > 400 µg/ml [21].

2.16. Antioxidant Activity

Antioxidants are our first line of defense against free radical damage that when present at low concentration compared with that of an oxidizable substrate significantly delay or inhibit oxidation of that substrate. The need for antioxidants becomes even more critical with increased exposure to free radicals, pollution, drugs and stress [52, 53]. The approved antioxidant activity of *Garcinia* species was reported, for example, the methanolic extract of *G. penducalata*, the twigs of *Garcinia merguensis*, the isolated biflavonoids from the epicarp of *Garcinia brasiliensis* and chloroform fraction of the methanol extract of *Garcinia morella* fruit, and different

Table 4. Reported xanthenes from *Garcinia livingstonei* T. Anderson.

| Xanthenes | | | | | |
|--|---------------|--|---------------------------|--------------|---|
| Compound Name | Plant Part | Extracts/Fractions | Geographical Distribution | References | Chemical Structures |
| 6,11-Dihydroxy-2,2-dimethyl-pyrano[3,2-c]xanthen-7(2H)-one | Root bark | Dichloromethane extract | South Africa | [37] |  |
| 12b-Hydroxy-des-dgarcigerrin A | | | | |  |
| 4[(E)-3',7'-Dimethylocta-2',6'-dienyl]-1,3,5-trihydroxy-9H-xanthen-9-one | Root bark | Dichloromethane and chloroform extracts | South Africa and Tanzania | [13, 37] |  |
| 6,11-Dihydroxy-3-methyl-3-(4-methylpent-3-enyl)-3H,7H-pyrano[2,3-c]xanthen-7-one | | - | | |  |
| 1,4,5-Trihydroxy-3-(3-methylbut-2-enyl)-9H-xanthen-9-one | Root bark | Dichloromethane and chloroform extracts | South Africa and Tanzania | [13, 37, 38] |  |
| Garcilivin A | | | | [13, 38] |  |
| Garcilivin B | | Dichloromethane extract | South Africa | [38] |  |
| Garcilivin C | | Dichloromethane and ethyl acetate soluble extracts | South Africa and Tanzania | [13, 38] |  |
| Alloathyriol | Fruits | Ethyl acetate fraction | Florida | [14] |  |
| Isojacareubin | Fruit kernels | Dichloromethane extract | South Africa | [4] |  |
| 6-deoxyisojacareubin | | | | |  |
| | Stem wood | Chloroform extract | Kasane | [33] | |

(Table 4) Contd....

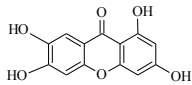
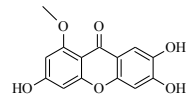
| Xanthenes | | | | | |
|---------------|------------|------------------------------------|---------------------------|------------|---|
| Compound Name | Plant Part | Extracts/Fractions | Geographical Distribution | References | Chemical Structures |
| Norathyriol | Twigs | Methanol extract | - | - |  |
| Montixanthone | Stem wood | Ethyl acetate and acetone extracts | | |  |

Table 5. Reported flavonoid from *Garcinia livingstonei* T. Anderson.

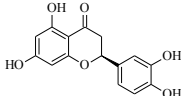
| Compound Name | Plant Part | References | Chemical Structures |
|---------------|------------|------------|---|
| Eriodictyol | Seeds | [39] |  |

Table 6. Reported triterpenes from *Garcinia livingstonei* T. Anderson.

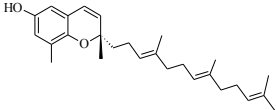
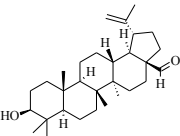
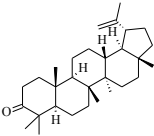
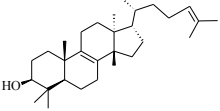
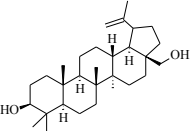
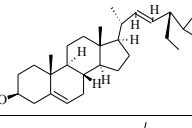
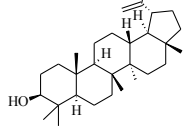
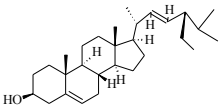
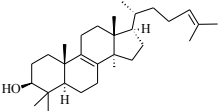
| Compound Name | Plant Part | Extracts/Fractions | Geographical Distribution | References | Chemical Structures |
|--------------------------|---------------|------------------------------------|---------------------------|------------|---|
| Sargaol | Fruit kernels | Hexane fraction | South Africa | [4] |  |
| Betulin aldehyde | Stem bark | Dichloromethane extract | | |  |
| Lupenone | | | | |  |
| Euphol | | - | | |  |
| Betulin | | | | |  |
| | Stem wood | Ethyl acetate and acetone extracts | Kasane | [33] |  |
| | | Chloroform extract | | |  |
| Stigmasterol | Stem bark | Dichloromethane extract | South Africa | [4] |  |
| Lupeol | | | | | Twigs |
| Lanosta-8,24-diene-3b-ol | Stem wood | Chloroform extract | Kasane | [33] |  |

Table 7. Reported anti-microbial activity of leaves acetone extracts and its fractions of *Garcinia livingstonei* T. Anderson.

| Bacteria | MIC of Leaves (mg/mL) | | | | | | | Reference |
|-------------------------------|-----------------------|--------------|---------------|-----------|-----------------------|--------------|--------------------|-----------|
| - | Acetone Extract | Hexane | Ethyl Acetate | Cloroform | Carbon Tetra Chloride | Butanol | 10% Water/Methanol | [48-50] |
| <i>Escherichia coli</i> | 0.06 | 0.23 | 0.175 | 0.095 | 0.115 | 0.78 | 1.875 | |
| <i>Staphylococcus aureus</i> | 0.07 | 0.23 | 0.195 | 0.31 | 0.355 | 1.25 | 1.875 | |
| <i>Pseudomonas aeruginosa</i> | 0.63 | 2.5 | 1.405 | 0.095 | 0.115 | 0.78 | 1.875 | |
| <i>Enterococcus faecalis</i> | 0.04 | Not Detected | 0.99 | 0.195 | 0.115 | Not Detected | Not Detected | |

Table 8. Reported anti-microbial activity of bark extracts of *Garcinia livingstonei* T. Anderson.

| Bacteria | MIC and MBC (mg/mL) of Bark Extracts | | | | |
|---|--------------------------------------|-----------------|---------------------------------|-----------------|------------|
| - | Methanol | Dichloromethane | Dichloromethane: Methanol (1:1) | Water | References |
| <i>Escherichia coli</i> | 0.41 | 2.00 | - | 0.78 | [7] |
| | 0.78 (MBC=12.5) | - | - | 0.78 (MBC=12.5) | [21] |
| <i>Shigella sonnei</i> | 0.83 | 0.50 | - | 0.78 | [7] |
| | 0.78 (MBC=12.5) | - | - | 0.78 (MBC=12.5) | [21] |
| <i>Enterococcus faecalis</i> | 0.18 | 0.75 | - | - | [7] |
| <i>Bacillus cereus</i> | 0.06 | 0.12 | - | - | |
| <i>Salmonella typhimurium</i> | 0.25 | 2.00 | - | - | |
| <i>Klebsiella oxytoca</i> | 0.39 (MBC=12.5) | - | - | 1.56 (MBC=12.5) | [21] |
| <i>Salmonella enterica</i> | 0.39 (MBC=>12.5) | - | - | 1.56 (MBC=12.5) | |
| <i>Staphylococcus aureus</i> | 0.39 (MBC=6.25) | - | - | 1.56 (MBC=12.5) | |
| <i>Brevibacterium agri</i> | - | - | 0.02 | 0.75 | [27] |
| <i>Brevibacterium linens</i> | - | - | 0.06 | 8 | |
| Gentamycin-methicillin resistant <i>S. aureus</i> | - | - | 0.5 | 2 | |
| Methicillin-resistant <i>S. aureus</i> | - | - | 0.25 | 1.5 | |
| <i>Staphylococcus epidermidis</i> | - | - | 0.00006 | 0.004 | |
| <i>Propionibacterium acnes</i> | - | - | 0.25 | >8 | |
| <i>Pseudomonas aeruginosa</i> | - | - | 0.5 | >8 | |
| <i>Candida albicans</i> | - | - | 0.25 | 4 | |
| <i>Microsporium canis</i> | - | - | 0.13 | 0.5 | |
| <i>Trichophyton mentagrophytes</i> | - | - | 0.13 | 6 | |

Table 9. Reported anti-microbial activity of isolated compounds from leaves of *Garcinia livingstonei* T. Anderson.

| Bacteria | MIC of Isolated Compounds (µg/mL) | | References |
|--------------------------------|-----------------------------------|-------------------------------|------------|
| - | Amentoflavone | 4' _ Monomethoxyamentoflavone | - |
| <i>Mycobacterium smegmatis</i> | 0.60 ± 0.70 | 1.40 ± 1.56 | [51] |
| <i>Escherichia coli</i> | 40 | 8 | [9, 48] |
| <i>Staphylococcus aureus</i> | 40 | 40 | |
| <i>Pseudomonas aeruginosa</i> | 100 | 60 | |
| <i>Enterococcus faecalis</i> | 60 | 8 | |

Table 10. Pharmacological activities of extracts of *Garcinia livingstonei* T. Anderson.

| Pharmacological Activities | Plant Parts | Extracts | Values | Geographical Distribution | References |
|--|-------------|---|--|---------------------------|------------|
| Antioxidant activity | Stem wood | Combined ethyl acetate and acetone extracts | IC ₅₀ = 30.3 ± 4.2 µg/ml | Kasane | [33] |
| | | Methanol extract | IC ₅₀ = 53.6 ± 7.4 µg/ml | | |
| | Twigs | Ethyl acetate extract | IC ₅₀ = 100.1 ± 5.9 µg/ml | | |
| | Fruit | Methanol extract | IC ₅₀ = 108.4 ± 12.9 µg/ml | Florida | [15] |
| | Bark | Aqueous extract | IC ₅₀ = 0.35 ± 0.06 µg/ml | South Africa | [21] |
| | | Methanol extract | IC ₅₀ = 0.39 ± 0.04 µg/ml | | |
| Anti-HIV-1 | Fruit | Ethanol extract | EC ₅₀ = 2.25 ± 0.51 µg/ml | Tanzania | [26] |
| Anti-HIV-1 protease activity | Root | | IC ₅₀ = 37.2 µg/ml | - | [16] |
| | Stem | | IC ₅₀ = 32.2 µg/ml | | |
| Anti <i>Cryptococcal meningitis</i> , <i>Herpes zoster</i> , <i>Herpes simplex</i> , skin rashes and tuberculosis chronic diarrhea | Fruit | Not mentioned | - | Zambia | [61] |
| Fungicidal activity | Root bark | Dichloromethane extract | Against <i>C. Cucumerinurn</i> | South Africa | [41] |
| Cytotoxic activity | | | Col15 (ED ₅₀ = 10), SW480 cell lines (ED ₅₀ = 10) | | |
| | | | LC ₅₀ = 17(0.3-43) µg/ml (Brine shrimp toxicity) | | |
| | Fruit | Ethanol extract | A549 (CC ₅₀ = 8 µg/ml), DU145 (CC ₅₀ = 8.2 µg/ml), KB (CC ₅₀ = 5.7 µg/ml), and Kbivin human cell lines (CC ₅₀ = 12 µg/ml). | Tanzania | [26] |
| | Bark | Aqueous extract | (HEK) 293 cells (EC ₅₀ = 769.9 µg/ml ± 36.33) R2 = 0.35 | South Africa | [21] |
| | | Methanol extract | (HEK) 293 cells (EC ₅₀ = > 400 µg/ml for methanol) R2 = 0.89 | | |
| LC ₅₀ = 19.12 after 24 hrs. and 46.85 after 48 hrs | | | | | [7] |

Table 11. Pharmacological activities of isolated compound of *Garcinia livingstonei* T. Anderson.

| Pharmacological Activities | Part Used | Isolated Compound | Values | Geographical Distribution | References |
|----------------------------|---------------------|--|---|---------------------------|------------|
| Antioxidant activity | Twigs and stem wood | 4,3',4'- trihydroxy-2,6-dimethoxybenzophenone | IC ₅₀ = 50.0 µm | Kasane | [33] |
| | | 3'-β-D-glucosy-loxy-4,40-dihydroxy-2,6-dimethoxy-benzophenone | IC ₅₀ = 353.0 µm | | |
| | | 3',4,4',6-tetrahydroxy-2-methoxybenzophenone | IC ₅₀ = 27.2 5 µm | | |
| | | Montixanthone | IC ₅₀ = 59.1 6 µm | | |
| | | 6- deoxyisojacareubin | IC ₅₀ = 1010 µm | | |
| | | Botulin | IC ₅₀ = 743.0 µm | | |
| | | Morelloflavone | IC ₅₀ = 37.0 µm | | |
| | | podocarpusflavone | IC ₅₀ = 498.0 µm | | |
| | | Volkensiflavone | IC ₅₀ = 402.3 µm | | |
| | | GB-1a | IC ₅₀ = 425 µm | | |
| | Leaves | Amentoflavone | TEAC = 0.9 | South Africa | [48] |
| | | 4'-Monomethoxyamentoflavone | TEAC = 2.2 | | |
| HIV inhibitory | Fruit | Guttiferone A | EC = 1-10 µg/ml | Tanzania | [35] |
| Skin lightening activity | Stem bark | Morelloflavone, morelloflavone-7"-sulphate, guttiferone A and sargaol | MeWo cells (MC = < 0.25 at 25 µM/ml) | South Africa | [4] |
| Antiparasitic activities | Root bark | 6,11-Dihydroxy-3-methyl-3-(4-methylpent-3-enyl)-3H,7H-pyrano[2,3-c]xanthen-7-one | <i>T. brucei</i> (IC ₅₀ = 5 ± 3 µm) <i>T. cruzi</i> (IC ₅₀ = 8 ± 3 µm) <i>P. falciparum</i> (IC ₅₀ = 52 ± 17 µm). | Tanzania | [13] |
| | | 4[(E)-3,7-dimethylocta-2,6-dienyl]-1,3,5-trihydroxy-9H-xanthen-9-one | <i>T. brucei</i> (IC ₅₀ = 2 ± 0.1 µm) <i>T. cruzi</i> (IC ₅₀ = 5.7 ± 6.4 µm) <i>P. falciparum</i> (IC ₅₀ = 6 µm) | | |
| | | 1,4,5-trihydroxy-3-(3-methylbut-2-enyl)-9Hxanthen-9-one (27, 10 and 0.78 µm) | <i>L. infantum</i> (IC ₅₀ = 27 µm) <i>P. falciparum</i> (IC ₅₀ = 10 ± 0.1 µm) <i>T. brucei</i> (IC ₅₀ = 0.87 ± 0.23 µm) <i>T. cruzi</i> (IC ₅₀ = 7 ± 0.1 µm) | | |
| | | Garcilivin A | <i>P. falciparum</i> (IC ₅₀ = 6.7 ± 1.5 µm) <i>L. infantum</i> (IC ₅₀ = 32 µm) <i>T. brucei</i> (IC ₅₀ = 0.4 ± 0.1 µm) <i>T. cruzi</i> (IC ₅₀ = 4 ± 2.8 µm) | | |
| | | Garcilivin C | <i>T. brucei</i> (IC ₅₀ = 7.7 ± 1.1 µm) <i>T. cruzi</i> (IC ₅₀ = 39.2 ± 23.2 µm) | | |

(Table 11) Contd....

| Pharmacological Activities | Part Used | Isolated Compound | Values | Geographical Distribution | References |
|---|-----------|--|---|---------------------------|------------|
| - | - | <i>Ent-naringeninyl</i> -(1-3 α ,II-8)-4'- <i>O</i> -methylnaringenin | <i>P. falciparum</i> (IC ₅₀ = 6 μ m) <i>T. brucei</i> (IC ₅₀ = 28.3 \pm 2.9 μ m) <i>T. cruzi</i> (IC ₅₀ = 34.7 \pm 4.6 μ m) <i>P. falciparum</i> (IC ₅₀ = 6 \pm 1.7 μ m) | - | - |
| - | - | Volkensiflavone | <i>T. brucei</i> (IC ₅₀ = 37 \pm 1.0 μ m) <i>T. cruzi</i> (IC ₅₀ = 56 \pm 13.9 μ m) <i>P. falciparum</i> (IC ₅₀ = 48 \pm 14.2 μ m) | - | - |
| Cytotoxicity | Leaves | Amentoflavone | Vero monkey kidney cells (CC ₅₀ = 381 μ g/mL) (A) and > 600 μ g/mL (B) | South Africa | [9, 48] |
| | | 4'-Monomethoxyamentoflavone | Vero monkey kidney cells (CC ₅₀ = 600 μ g/mL) | | |
| | Root bark | Garcilivin A | MRC-5 cells (IC ₅₀ = 2 \pm 0.1 μ m) | Tanzania | [13] |
| | | Garcilivin C | MRC-5 cells (IC ₅₀ = 52.3 \pm 5.5 μ m) | | |
| | | <i>Ent-naringeninyl</i> -(1-3 α ,II-8)-4'- <i>O</i> -methylnaringenin | MRC-5 cells (IC ₅₀ = 38.0 \pm 2.6 μ m) | | |
| | | Volkensiflavone | MRC-5 cells (IC ₅₀ = 40 \pm 4.4 μ m) | | |
| Acetyl cholinesterase inhibition activity | Root bark | 6,11-Dihydroxy-3-methyl-3-(4-methylpent-3-enyl)-3H,7H-pyrano[2,3-c]xanthen-7-one | K _i = 1.9 \pm 1.4 μ M | South Africa | [70] |
| | | 4[(<i>E</i>)-3',7'-Dimethylocta-2',6'-dienyl]-1,3,5-trihydroxy-9 <i>H</i> -xanthen-9-one | K _i = 26.8 \pm 5.4 μ M | | |
| Mono amine oxidase inhibition activity | | 4[(<i>E</i>)-3',7'-Dimethylocta-2',6'-dienyl]-1,3,5-trihydroxy-9 <i>H</i> -xanthen-9-one | MAO-A (IC ₅₀ = 37.4 μ M) MAO-B (IC ₅₀ = 65.5 μ M) | | |

Abbreviations: Col15 = (Colon carcinoma), SW480 = (Colon carcinoma), A549 = (Lung adenocarcinoma), DU145 = (Prostate carcinoma), KB = (Nasopharyngeal carcinoma), Kbivn = (Vincristine-resistant nasopharyngeal), HEK 293 cells = Human Embryonic Kidney, MRC-5 = Human diploid embryonic lung cell line, MeWo = human melanoma cells.

parts of other species all exhibited an interested anti-oxidant activity [54-57]. For *G. livingstonei*, the potent antioxidant activity of 4,3',4'-trihydroxy-2,6-dimethoxybenzophenone, 3',4,4',6-tetrahydroxy-2-methoxybenzophenone, montixanthone and morelloflavone were reported with IC₅₀ values of 50.0, 27.2, 59.1 and 37.0 μ M, respectively, using the DPPH assay isolated from the twigs and stem wood compared to vitamin C (IC₅₀ = 82.4 μ M) [33]. The methanolic extract of the fruit also was reported to exhibit antioxidant activity using the DPPH scavenging assay (IC₅₀ = 108.4 \pm 12.9 μ g/mL) [15]. In addition 4'-monomethoxyamentoflavone had a good antioxidant activity than amentoflavone [48]. Water and methanol extracts of *G. livingstonei* bark showed antioxidant activities with IC₅₀ values 0.35 \pm 0.06 and 0.39 \pm 0.04 μ g/mL, respectively [21].

2.17. Anti-HIV Activity

According to WHO, HIV is considered a major public health problem. In 2020, 1.5 million people were affected by HIV [58]. *Garcinia* exhibited a marked effect against HIV which is evidenced by that the isolated compounds from the fruits of *G. speciosa* and *G. nuntasaenii* for example showed significant inhibitory activities against HIV [59, 60]. The isolated guttiferone A exhibited HIV inhibitory activity [35]. The ethanolic fruit extract from Tanzania showed significant anti-HIV-1 activity with EC₅₀ value of 2.25 \pm 0.51 μ g/ml. The cytotoxicity on the T-cells was CC₅₀ = 6.1 μ g/ml [26]; while the roots and the stems showed anti-HIV-1 protease activity with IC₅₀ values of 37.2 and 32.2 μ g/ml, respectively [16]. The activity of *G. livingstonei* fruits from Zambia

against *Cryptococcal meningitis*, chronic diarrhea, skin rashes, *Herpes simplex*, *Herpes zoster* and tuberculosis was also reported [61].

2.18. Inhibition of Melanoma

Hyperpigmentation is one of the problems that take interest of most population and melanoma is one of the major causes of skin cancer deaths [62, 63]. Xanthoness isolated from *G. mangostana* were effective as anti-melanoma [64], also the hydroalcoholic extract of leaves of *G. gardneriana* was potent in the treatment of hyperpigmentation disorders [65]. Moreover, the aqueous extract of fruits of *G. atroviridis* showed anti tyrosinase activity and inhibited melanin content [66]. Concerning *G. livingstonei*, morelloflavone, morelloflavone-7"-sulphate and sargaol are responsible for its skin lightening activity [4].

2.19. Antiparasitic Activity

The antiparasitic activity of isolated compounds from root bark was evaluated against *Plasmodium falciparum*, *Leishmania infantum*, *Trypanosoma brucei* and *T. cruzi*. 1,4,5-trihydroxy-3-(3-methylbut-2-enyl)-9H-xanthen-9-one showed a marginal activity against *L. infantum*, *P. falciparum* and *T. brucei* with IC_{50} = 27, 10 and 0.78 μ M, respectively, on the other hand garcilivin A showed nonselective cytotoxicity (IC_{50} = 2.0 μ M) against all parasites tested [13].

2.20. Antifungal Activity

The dichloromethane extract of root bark of *G. livingstonei* exhibited fungicidal activity against *Cladosporium cucumerinum* [41]. However, no other reports were traced in the literature concerning this matter.

2.21. Cytotoxic Activity

Cancer is one of the major causes of death worldwide, which needs the use of protocols chemotherapy and radiotherapy protocols in the treatment or further need of surgical intervention [67, 68]. Recently, plant based remedies are preferred for the improvement of health [69]. The cytotoxic activity of root bark extract of *G. livingstonei* was tested against two human colon carcinoma cell lines (Col15 and SW480 cell lines) and the brine shrimp toxicity showed LC_{50} = 17(0.3-43) μ g/ml; the ED_{50} values were 10 and 8 μ g/ml, respectively [41]. Also the ethanolic fruits' extract from Tanzania exhibited cytotoxic activity against other cell lines, A549 (Lung adenocarcinoma), DU145 (Prostate carcinoma), KB (Nasopharyngeal carcinoma) and Kbivin (Vincristine-resistant nasopharyngeal) human cell lines with CC_{50} values of 8, 8.2, 5.7 and 12 μ g/ml, respectively [26]. The mutagenic potential of amentoflavone and 4'-monomethoxyamentoflavone were assessed by the Ames test against *Salmonella* strain TA98. Amentoflavone had some genotoxic effect at the concentration of 100 μ g/plate; however, 4'-monomethoxyamentoflavone was inactive [48]. The safety of the two compounds was assessed with MTT assay using Vero monkey kidney cells. The compounds had low toxicity against the cell line with cytotoxic concentrations of 50% of the cells (LD_{50}) of 386 microg/mL and > 600 microg/mL for amentoflavone and 4"-methoxy amentoflavone, respectively [9]. Moreover, the water extract of bark had the highest EC_{50} value of 769.9 ± 36.33 μ g/mL,

followed by the methanol extract 400 ± 36.33 μ g/mL against HEK cells (Human Embryonic Kidney) [21]. In addition, the methanol extract of bark of *G. livingstonei* showed LC_{50} = 19.12 after 24 hrs and 46.85 after 48 hrs [7].

2.22. Acetylcholinesterase and Monoamine Oxidase Inhibition Activity

The isolated xanthoness, namely; 12b-Hydroxy-desdgarcigerrin A, 6,11-Dihydroxy-3-methyl-3-(4-methylpent-3-enyl)-3H,7H-pyrano[2,3-c]xanthen-7-one, 1,4,5-Trihydroxy-3-(3-methylbut-2-enyl)-9H-xanthen-9-one and 4[(E)-3',7'-Dimethylocta-2',6'-dienyl]-1,3,5-trihydroxy-9H-xanthen-9-one were evaluated for their activity as acetylcholinesterase inhibitors for the treatment of Alzheimer's disease, the most active xanthone was 6,11-Dihydroxy-3-methyl-3-(4-methylpent-3-enyl)-3H,7H-pyrano[2,3-c]xanthen-7-one with K_i value of 1.9 ± 1.4 μ M and 4[(E)-3',7'-Dimethylocta-2',6'-dienyl]-1,3,5-trihydroxy-9H-xanthen-9-one with K_i value of 26.8 ± 5.4 μ M. The later xanthone also showed monoamine oxidase (MAO) inhibition activity, with IC_{50} values for MAO-A and MAO-B equal 37.4 and 65.5 μ M, respectively. In conclusion, 4[(E)-3',7'-Dimethylocta-2',6'-dienyl]-1,3,5-trihydroxy-9H-xanthen-9-one had interesting activity for Alzheimer's disease therapy [70].

CONCLUSION

Garcinia species are members of family Clusiaceae, well known for their edible fruits and bioactive constituents. Several species of *Garcinia* are distributed in different parts of the world such as Malaysia, Indonesia, India, tropical Africa and Egypt. *G. livingstonei* is an economically important fruit crop useful as raw material for plant based products in industrial sector related to food, cosmetic or nutraceuticals. *G. livingstonei* is a rich source of bioactive constituents that are beneficial in several health problems, exemplified by amentoflavone, morelloflavone and volkensiflavone which are effective as anti-oxidant, antibacterial and melanogenesis inhibitors. The presence of benzophenones *viz.*, as guttiferone A with anti-HIV potential and xanthoness as garcilivin C and montixanthone are active anti-parasitic and cytotoxic agents, reflecting their value as potential sources for natural drugs. Moreover, *G. livingstonei* has high nutritive contents including minerals, carbohydrates, crude fat and crude proteins. These elements promote plants as food supplements.

LIST OF ABBREVIATIONS

HIV = Anti-human Immunodeficiency Virus
TI = Therapeutic Index

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The authors declare no conflict of interest, financial or otherwise.

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