**Review**

**Moringa oleifera** Lam.: A Phytochemical and Pharmacological Overview

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**Abstract:** *Moringa oleifera* Lam. is a fast-growing and drought-resistant tree of the Moringaceae family. The tree is known with some common names: miracle, ben oil, drumstick, horseradish or simply moringa. The plant grows in a variety of environments including harsh conditions. The plant has a high content of phytoneutrients, being used as natural integrators to cure malnutrition. Phytochemical studies of plant organs showed the plant as a rich source of primary and secondary metabolites belonging to different classes of organic compounds. Pharmacological studies confirmed the use of the plant to cure several diseases and to possess nutraceutical properties. This review is aimed to contribute to the knowledge of *M. oleifera* by providing its plant description, phytochemistry and pharmacology.

**Keywords:** *Moringa oleifera* Lam.; botany; phytochemistry; bioactive metabolites; nutrition; pharmacology

1. **Introduction**

*Moringa oleifera* Lam. is a fast-growing and drought-resistant tree of the Moringaceae family. It is commonly known with several names including moringa, drumstick tree (for the long and slender seedpods), horseradish tree (for the roots taste resembling horseradish), ben oil tree (being rich in behenic acid), and miracle tree (for the medicinal properties) [1]. *M. oleifera* is the most popular of the thirteen species in the genus *Moringa*, the unique of the family Moringaceae which includes: *M. concanensis*, *M. drouhardii*, *M. arborea*, *M. borziana*, *M. hildebrandtii*, *M. longituba*, *M. pygmaea*, *M. rivae*, *M. ruspoliana*, *M. ovalifolia*, *M. peregrine*, and *M. stenopetala*. The species are native to Africa, Arabia, India, Southeast Asia, South America, the Pacific Islands, and the Caribbean [2]. It is distributed in Ethiopia, Florida, Philippines, and Sudan [2].

*M. oleifera* is one of the most popular plants that can grow in a variety of environments due to its ability to resist to harsh conditions such as high temperatures and limited water availability [3]. Therefore, it grows in a variety of soils including semi-dry, desert, or tropical soils and rainfall conditions [3]. The plant tolerates also different pH levels, ranging from 5.0 to 9.0. However, it prefers neutral pH and well-drained soils. It thrives in temperatures ranging from 25 to 40 °C, though it can withstand temperature swings of −1 to 3 °C and 38 to 48 °C during the coldest and warmest months, respectively [3]. The plant yield related to cutting period and planting density was studied and the data indicated that the highest planting density of 0.2 m × 0.2 m in combination with the intermediate cutting height of 30 cm produced the highest fresh matter and dry matter yield throughout the whole evaluation period [4].

*M. oleifera* is among the food plants richest in nutrients [5]. It has a high content in essential amino acids, proteins, minerals, vitamins, and polyphenols. It is a rich source
of phytochemicals including flavonoids, anthocyanins, isothiocyanates, anthraquinone, alkaloids, essential oils, tannic acid, saponins, steroids, terpenoids, cardiac glycosides [6]. In addition, it is used to treat individuals with extreme malnutrition as well as for its pharmacological (hepatoprotective, antihypertensive, cholesterol-lowering, anti-urolithiasis, antifertility, antidiabetic, and antioxidant activity, nutraceutical properties, and antimicrobial) [6]. Moreover, *M. oleifera* is being used to help to breastfeed mothers improving postpartum milk production [7]. It is also used in Ayurvedic tradition specifically for cancer treatment [8]. *M. oleifera* leaves and buds were used against headache by rubbing them on the temples. Roots and root barks were used as anti-scorbutic [6]. The eye diseases were treated with the juice of the leaves added with honey. Dried seeds of *M. oleifera* were used in ophthalmic preparation, venereal affection anti-inflammatory and purgative and as tonic [6]. The use in ethnomedicine encouraged scientists to study this plant to determine its chemical composition and to investigate its pharmacological potential. This review is aimed to contribute to the knowledge of *M. oleifera* by providing plant description, phytochemistry and pharmacology.

2. Plant Description

*M. oleifera* is a short, slender, deciduous, perennial tree that grows to about 10 m tall, slender with drooping branches; branches and stem are brittle, with corky bark. Leaves are feathery, pale green, compound, tripinnate, (30–60 cm long), with many small leaflets, 1.3–2 cm long, 0.6–0.3 cm wide, lateral ones slightly elliptic, terminal ones obovate, and slightly larger (Figure 1). Flowers are fragrant, white, or creamy-white, 2.5 cm in diameter, and borne in sprays. The stamens are yellow, and the pods are pendulous, brown, triangular, splitting lengthwise into three parts when dry, and containing about 20 seeds embedded in the pith. The pod has nine ribs on both ends and the seeds are dark brown with three papery wings [1].

![Images of M. oleifera tree, leaves, seeds, and flowers.](image)

The feathery leaves of the tripinnate complex have green curved leaflets that are 1–4 cm long (Figure 1). Because of its leaves, the tree is frequently mistaken for a leguminous plant. The alternate twice or thrice pinnate leaves appear at the branch tips in most cases. They have a long petiole with 8–10 pairs of pinnae, each bearing two sets of inverse elliptic leaflets and one at the apex and are 20–70 cm long when young [1].

The seeds have three papery wings and are oval with a tannish semi-permeable seed arrangement (Figure 1). Their arrangements are mostly brown to dark brown but can be
white if portions are of low viability. It almost within a week, viable seeds sprout. The body has three white wings that run at 130 intervals from start to finish [1].

Flowers are prominent, softly fragrant, and are borne on inflorescences 15–25 cm long. They are mostly white to cream in color, 2.5 cm in diameter, and tinged pink in a few varieties (Figure 1). However, the flowers are fragrant and 2.5 cm wide, and they bloom profusely in auxiliary, dropping panicles that are 10–25 cm long. They have a dotted base and are white in color. The direct lanceolate sepals are five-reflexed. The five petals are rumored to be thin. Except for the lowest stamen, they are reflexed and consist of five stamens and five staminodes [1].

3. Phytochemistry

Table 1 lists all the metabolites that have been found in *M. oleifera* extracts. Many studies have focused their efforts in analyzing the polar and non-polar extracts of *M. oleifera* leaves and seeds to characterize the chemical composition of this plant and try to explain the beneficial characteristics demonstrated by lots of works in recent years. Selected characteristic compounds of the plant are reported in Figure 2.

![Figure 2. Bioactive metabolites isolated from *M. oleifera*.](image-url)
Table 1. List of compounds extracted from *Moringa oleifera* leaves and seeds.

| Compounds                  | Molecular Formula | Quantity | Method     | Reference |
|----------------------------|-------------------|----------|------------|-----------|
|                            |                   | Leaves   | Seeds      |           |
| Caproic acid               | C₆H₁₂O₂           | 0.1%     | -          | GC-MS [9] |
| Capric acid                | C₁₀H₂₀O₂          | 0.1%     | -          | GC-MS [9] |
| Lauric acid                | C₁₂H₂₄O₂          | 0.72%    | 0.03%      | GC-MS [9] |
| Myristic acid              | C₁₄H₂₈O₂          | 1.13%    | 0.16%      | GC-MS [9] |
|                            |                   | 23.65%   | 7.92%      | GC-MS [9] |
| Palmitic acid              | C₁₆H₃₂O₂          |          | Madurai: 6.17%<sup>a</sup> | GC-MS [10] |
|                            |                   | 1.08%<sup>a</sup> | - | GC-MS [11] |
|                            |                   | 0.15%    | -          | GC-MS [12] |
|                            |                   | 2.5703% <sup>a</sup> | - | GC-MS [13] |
| Palmitoleic acid           | C₁₆H₃₆O₂          |          | 1.49%      | GC-MS [9] |
| Heptadecanoic acid         | C₁₇H₃₄O₂          | 0.46%    | 0.32%      | GC-MS [9] |
|                            |                   |          | Madurai: 5.12%<sup>a</sup> | GC-MS [10] |
| Stearic acid               | C₁₈H₃₆O₂          | 1.2051%<sup>a</sup> | - | GC-MS [13] |
|                            |                   | 3.93%    | 7.10%      | GC-MS [9] |
| Oleic acid                 | C₁₈H₃₄O₂          | 5.92%    | 74.50%     | GC-MS [9] |
| Linoleic acid              | C₁₈H₃₆O₂          |          | Madurai: 5.12%<sup>a</sup> | GC-MS [10] |
| Arachidic acid             | C₂₀H₄₀O₂          | 6.84%    | 0.62%      | GC-MS [9] |
| Cis-11-Eicosenoic acid     | C₂₀H₃₈O₂          | 54.44%   | -          | GC-MS [9] |
| Linolenic acid             | C₁₈H₃₂O₂          |          | 2.50%      | GC-MS [9] |
| Cis-11, 14-Eicosadienoic acid | C₂₀H₄₀O₂         |          | 0.48%      | GC-MS [9] |
| Lysine                     | C₆H₁₄N₂O₂         | Madurai: 4.10 g/16 gN | Madurai: 4.20 g/16 gN | HPLC [10] |
|                           |                   | Chennai: 3.30 g/16 gN | Chennai: 3.70 g/16 gN |           |
|                           |                   | 129.4 mg/100g dw | - | HPLC [14] |
| Histidine                  | C₄H₇N₃O₂         | Madurai: 2.50 g/16 gN | Madurai: 6.30 g/16 gN | HPLC [10] |
|                           |                   | Chennai: 3.12 g/16 gN | Chennai: 5.70 g/16 gN |           |
|                           |                   | 73.9 mg/100g dw | - | HPLC [14] |
| Valine                     | C₃H₁₁NO₂         | Madurai: 4.55 g/16 gN | Madurai: 3.36 g/16 gN | HPLC [10] |
|                           |                   | Chennai: 4.75 g/16 gN | Chennai: 2.37 g/16 gN |           |
|                           |                   | 111.5 mg/100g dw | - | HPLC [14] |
| Leucine                    | C₄H₁₃NO₂         | Madurai: 4.80 g/16 gN | Madurai: 4.80 g/16 gN | HPLC [10] |
|                           |                   | Chennai: 4.40 g/16 gN | Chennai: 4.70 g/16 gN |           |
|                           |                   | 211.5 mg/100g dw | - | HPLC [14] |
| Isoleucine                 | C₄H₁₃NO₂         | Madurai: 5.90 g/16 gN | Madurai: 8.30 g/16 gN | HPLC [10] |
|                           |                   | Chennai: 9.10 g/16 gN | Chennai: 7.70 g/16 gN |           |
|                           |                   | 81.3 mg/100g dw | - | HPLC [14] |
| Threonine                  | C₄H₇NO₃         | Madurai: 4.10 g/16 gN | Madurai: 3.20 g/16 gN | HPLC [10] |
|                           |                   | Chennai: 4.50 g/16 gN | Chennai: 3.80 g/16 gN |           |
|                           |                   | 91.8 mg/100g dw | - | HPLC [14] |
| Alanine                    | C₃H₇NO₂         | Madurai: 3.23 g/16 gN | Madurai: 4.29 g/16 gN | HPLC [10] |
|                           |                   | Chennai: 3.90 g/16 gN | Chennai: 3.55 g/16 gN |           |
|                           |                   | 121.2 mg/100g dw | - | HPLC [14] |
| Compounds             | Molecular Formula | Quantity                      | Method   | Reference |
|-----------------------|-------------------|-------------------------------|----------|-----------|
| **Leaves**            | **Seeds**         |                               |          |           |
| Aspartic acid         | C$_4$H$_7$NO$_4$  | Madurai: 6.44 g/16 gN         | HPLC     | [10]      |
|                       |                   | Chennai: 6.86 g/16 gN         |          |           |
|                       |                   | Madurai: 6.02 g/16 gN         |          |           |
|                       |                   | Chennai: 6.37 g/16 gN         |          |           |
|                       | 195.8 mg/100 g dw |                               | HPLC     | [14]      |
| Serine                | C$_3$H$_7$NO$_3$  | Madurai: 4.25 g/16 gN         | HPLC     | [10]      |
|                       |                   | Chennai: 4.66 g/16 gN         |          |           |
|                       |                   | Madurai: 4.22 g/16 gN         |          |           |
|                       |                   | Chennai: 4.11 g/16 gN         |          |           |
|                       | 90.2 mg/100 g dw  |                               | HPLC     | [14]      |
| Proline               | C$_3$H$_7$NO$_2$  | Madurai: 6.02 g/16 gN         | HPLC     | [14]      |
|                       |                   | Chennai: 6.37 g/16 gN         |          |           |
|                       |                   | Madurai: 6.44 g/16 gN         |          |           |
|                       |                   | Chennai: 6.86 g/16 gN         |          |           |
|                       | 128.9 mg/100 g dw |                               | HPLC     | [14]      |
| Glutamic acid         | C$_5$H$_9$NO$_4$  | Madurai: 15.86 g/16 gN        | HPLC     | [10]      |
|                       |                   | Chennai: 15.33 g/16 gN        |          |           |
|                       |                   | Madurai: 14.23 g/16 gN        |          |           |
|                       |                   | Chennai: 14.74 g/16 gN        |          |           |
|                       | 268.7 mg/100 g dw |                               | HPLC     | [14]      |
| Glycine               | C$_2$H$_5$NO$_2$  | Madurai: 5.13 g/16 gN         | HPLC     | [10]      |
|                       |                   | Chennai: 5.11 g/16 gN         |          |           |
|                       |                   | Madurai: 5.00 g/16 gN         |          |           |
|                       |                   | Chennai: 4.70 g/16 gN         |          |           |
|                       | 102.4 mg/100 g dw |                               | HPLC     | [14]      |
| Arginine              | C$_6$H$_{14}$N$_2$O$_2$ | Madurai: 4.30 g/16 gN | HPLC     | [10]      |
|                       |                   | Chennai: 3.70 g/16 gN         |          |           |
|                       |                   | Madurai: 9.44 g/16 gN         |          |           |
|                       |                   | Chennai: 11.00 g/16 gN        |          |           |
|                       | 125.4 mg/100 g dw |                                | HPLC     | [14]      |
| Cysteine              | C$_3$H$_7$NO$_2$S | Madurai: 4.22 g/16 gN         | HPLC     | [10]      |
|                       |                   | Chennai: 4.11 g/16 gN         |          |           |
|                       |                   | Madurai: 4.25 g/16 gN         |          |           |
|                       |                   | Chennai: 4.66 g/16 gN         |          |           |
|                       | 29.6 mg/100 g dw  |                                | HPLC     | [14]      |
| Tyrosine              | C$_3$H$_{11}$NO$_3$ | Madurai: 3.50 g/16 gN         | HPLC     | [10]      |
|                       |                   | Chennai: 3.90 g/16 gN         |          |           |
|                       | 79.3 mg/100 g dw  |                                | HPLC     | [14]      |
| Methionine            | C$_3$H$_{11}$NO$_2$S | Madurai: 3.50 g/16 gN        | HPLC     | [10]      |
|                       |                   | Chennai: 3.90 g/16 gN         |          |           |
|                       | 212.4 mg/100 g dw |                                | HPLC     | [14]      |
| Phenylalanine         | C$_3$H$_{11}$NO$_2$ | Madurai: 4.20 g/16 gN         | HPLC     | [10]      |
|                       |                   | Chennai: 4.60 g/16 gN         |          |           |
|                       | 117.2 mg/100 g    |                                | HPLC     | [14]      |
| 1,30-Triacontanediol  | C$_{30}$H$_{62}$O$_2$ | Madurai: 3.06%                 | GC-MS    | [12]      |
|                       |                   | Chennai: 14.98%               |          |           |
| Octacosane            | C$_{28}$H$_{58}$  | Madurai: 8.57%                | GC-MS    | [10]      |
|                       |                   | Chennai: 5.00%                |          |           |
| 2,2-dimethyl-1-oxa-2- | C$_{28}$H$_{58}$  | Madurai: 8.3%                 | GC-MS    | [10]      |
| silacyclotridic         |                       | Chennai: 1.21%                |          |           |
| 13-docosenamide, (Z)-  | C$_{26}$H$_{50}$  | Chennai: 13.62%               | GC-MS    | [10]      |
| 6-octadecenoic acid   | C$_{18}$H$_{32}$O$_2$ | Madurai: 52.24%               | GC-MS    | [10]      |
| Propionamide          | C$_2$H$_5$NO      | Chennai: 5.05%                 | GC-MS    | [10]      |
| Ethyl oleate          | C$_{26}$H$_{40}$O$_2$ | Chennai: 4.33%                | GC-MS    | [10]      |
| Compounds                  | Molecular Formula | Quantity                      | Method        | Reference          |
|---------------------------|-------------------|-------------------------------|---------------|--------------------|
| Glucomoringin             | C_{21}H_{31}NO_{14}S_{2} | 16.14–17.3 μmol/g dry weight | UPLC          | [15]               |
|                           |                   |                               |               |                    |
|                           |                   | Domesticated: 75.29 μmol/g dry weight | HPLC          | [15]               |
|                           |                   | Wild: 18.00 μmol/g dry weight |               |                    |
|                           |                   | 110–112 μmol/g dry weight     | HPLC          | [15]               |
|                           |                   | 21.84–22.56 mg/g dry weight  | HPLC          | [15]               |
|                           |                   | 0.77 mg/g dry weight          | HPLC-ESI-MS/MS| [15]               |
|                           |                   | 0.018 mg/g                   |                |                    |
|                           |                   | 0.189 mg/g                   |                |                    |
| Glucomoringin Isomer I    | C_{21}H_{31}NO_{14}S_{2} | 7.15–8.92 μmol/g dry weight  | UPLC          | [15]               |
| Glucomoringin Isomer II   | C_{21}H_{31}NO_{14}S_{2} | 3.85–4.84 μmol/g dry weight  | UPLC          | [15]               |
| Glucomoringin Isomer III  | C_{21}H_{31}NO_{14}S_{2} | 18.71–22.58 μmol/g dry weight| UPLC          | [15]               |
| Glucosoonjnain            |                   | 3.48 mg/100 g                | HPLC          | [16]               |
| β-carotene                | C_{40}H_{56}       | 27.22 mg/100 g               | N/A           | [16]               |
|                           |                   | 36 mg/100 g                  | AOAC 2004     | [16]               |
| Lutein                    | C_{40}H_{56}O_{2}  | 4.09 mg/g                    | HPLC          | [16]               |
|                           |                   | 5.56 mg/g                    | HPLC          | [16]               |
| Caffeic acid              | C_{9}H_{8}O_{4}    | 0.409 mg/g                   | HPLC          | [16]               |
| Chlorogenic acid          | C_{16}H_{10}O_{9}  | 0.489 mg/g                   | HPLC and MS/MS| [16]               |
| o-Coumaric acid           | C_{9}H_{8}O_{3}    | 6.457 mg/g                   | HPLC          | [16]               |
| Ellagic acid              | C_{14}H_{6}O_{8}   | 0.018 mg/g                   | HPLC and MS/MS| [16]               |
|                           |                   | 0.189 mg/g                   |                |                    |
| Ferulic acid              | C_{10}H_{10}O_{4}  | 0.078 mg/g                   | HPLC and MS/MS| [16]               |
| Gallic acid               | C_{7}H_{6}O_{3}    | 1.034 mg/g                   | HPLC and MS/MS| [16]               |
| Compounds   | Molecular Formula | Quantity                     | Method         | Reference |
|-------------|-------------------|------------------------------|----------------|-----------|
|             |                   | Leaves | Seeds |                     |           |
| Epicatechin | C_{15}H_{14}O_{6} | 5.68 mg/g | - | HPLC | [16]       |
|             |                   | 0.118 mg/g | - | HPLC | [16]       |
|             |                   | 0.52 mg/g | - | HPLC-DAD-ESI-MS | [16]       |
|             |                   | 0.72 mg/g | - | HPLC-DAD-ESI-MS | [16]       |
|             |                   | 0.04 mg/g | - | HPLC | [16]       |
|             |                   | 2.360 mg/g | - | HPLC | [16]       |
|             |                   | 0.198 mg/g | - | HPLC and MS/MS | [16]       |
|             |                   | 1.412 mg/g | - | HPLC | [16]       |
|             |                   | 0.8 mg/g | - | HPLC | [16]       |
|             |                   | 1.23 mg/g | - | LC/MS | [16]       |
|             |                   | 4.59 mg/g | - | HPLC | [16]       |
| Isorhamnetin| C_{16}H_{13}O_{7} | 3.92 mg/g | - | HPLC-DAD-ESI-MS | [16]       |
|             |                   | 2.16 mg/g | - | HPLC-DAD-ESI-MS | [16]       |
|             |                   | 2.25 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 1.27 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 1.05 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 2.9 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 2.3 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 3.5 mg/g | - | LC/MS | [16]       |
|             |                   | 0.3 mg/g | - | HPLC-DAD | [16]       |
| Kaempferol  | C_{15}H_{10}O_{6} | 0.16 mg/g | - | HPLC | [16]       |
|             |                   | 5.804 mg/g | - | HPLC | [16]       |
|             |                   | 0.281 mg/g | - | HPLC | [16]       |
|             |                   | 0.207 mg/g | - | HPLC and MS/MS | [16]       |
|             |                   | 0.207 mg/g | - | HPLC | [16]       |
|             |                   | 0.807 mg/g | - | HPLC and MS/MS | [16]       |
| Myricetin   | C_{15}H_{10}O_{8} | 5.2 mg/g | - | LC/MS | [16]       |
|             |                   | 5.8 mg/g | - | LC/MS | [16]       |
|             |                   | 7.57 mg/g | - | LC/MS | [16]       |
|             |                   | 12.84 mg/g | - | HPLC-DAD-ESI-MS | [16]       |
|             |                   | 16.64 mg/g | - | HPLC-DAD-ESI-MS | [16]       |
|             |                   | 9.26 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 6.34 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 7.70 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 5.47 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 9.1 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 15.2 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 0.58 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 0.46 mg/g | - | HPLC-DAD | [16]       |
| Quercetin   | C_{15}H_{10}O_{7} | 3.529 mg/g | - | HPLC | [16]       |
|             |                   | 5.2 mg/g | - | HPLC | [16]       |
|             |                   | 7.57 mg/g | - | HPLC | [16]       |
|             |                   | 12.84 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 16.64 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 9.26 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 6.34 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 7.70 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 5.47 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 9.1 mg/g | - | HPLC-DAD | [16]       |
|             |                   | 15.2 mg/g | - | LC-MS | [16]       |
|             |                   | 0.58 mg/g | - | LC-MS | [16]       |
|             |                   | 0.46 mg/g | - | LC-MS | [16]       |
Table 1. Cont.

| Compounds | Molecular Formula | Leaves | Seeds | Method | Reference |
|-----------|------------------|--------|-------|--------|-----------|
| Rutin     | C$_{27}$H$_{30}$O$_{16}$ | 1.674 mg/g | - | HPLC | [16] |
| 4-(α-L-rhamnopyranosyl)-benzyl-isothiocyanate (moringin) | C$_{14}$H$_{17}$NO$_{5}$S | 2.256 mg/g | - | HPLC-DAD-ESI-MS | [16] |
| 4-O-(α-Lacetylrhamnopyranosyl)-benzyl-isothiocyanate isomer 1 | C$_{16}$H$_{19}$NO$_{6}$S | 1.80 mg/g | - | HPLC-DAD-ESI-MS | [16] |
| 4-O-(α-Lacetylrhamnopyranosyl)-benzyl-isothiocyanate isomer 2 | C$_{16}$H$_{19}$NO$_{6}$S | 1.52 mg/g | - | HPLC-DAD-ESI-MS | [16] |
| 4-O-(α-Lacetylrhamnopyranosyl)-benzyl isomer 3 | C$_{16}$H$_{19}$NO$_{6}$S | 20.16 mg/g | - | HPLC-DAD-ESI-MS | [16] |
| Toluene   | C$_7$H$_8$         | 1.27%  | -     | GC-MS  | [12] |
| 5-tert-Butyl-1,3-cyclopentadiene | C$_5$H$_{14}$ | 0.07%  | -     | GC-MS  | [11] |
| Benzaldehyde | C$_4$H$_{13}$CHO | 0.55%  | -     | GC-MS  | [11] |
| 5-Methyl-2-furaldehyde | C$_6$H$_{16}$O | 0.27%  | -     | GC-MS  | [11] |
| Benzeneacetaldehyde | C$_6$H$_{14}$O | 2.16%  | -     | GC-MS  | [11] |
| 2-Ethyl-3,6-dimethylpyrazine | C$_8$H$_{12}$N$_2$ | 0.12%  | -     | GC-MS  | [11] |
| Undecane  | C$_{11}$H$_{24}$  | 0.12%  | -     | GC-MS  | [11] |
| Isophorone | C$_9$H$_{14}$O    | 0.10%  | -     | GC-MS  | [11] |
| Benzyl nitrite | C$_7$H$_7$N | 1.10%  | -     | GC-MS  | [11] |
| 2,6,6-Trimethyl-2-cyclohexane-1,4-dione | C$_9$H$_{12}$O$_2$ | 0.05%  | -     | GC-MS  | [11] |
| 2,2,4-Trimethyl-pentadiol | C$_{12}$H$_{24}$O$_3$ | 0.09%  | -     | GC-MS  | [11] |
| 2,3-Epoxy carbane | C$_{10}$H$_{16}$O | 0.16%  | -     | GC-MS  | [11] |
| p-Menth-1-en-8-ol | C$_{10}$H$_{16}$O | 0.08%  | -     | GC-MS  | [11] |
| 2,6,6-Trimethylcyclohexa-1,3-diene carbaldehyde | C$_9$H$_{14}$ | 0.23%  | -     | GC-MS  | [11] |
| Indole    | C$_5$H$_{9}$N     | 1.20%  | -     | GC-MS  | [11] |
| Tridecane | C$_{13}$H$_{28}$  | 0.16%  | -     | GC-MS  | [11] |
| Ionone    | C$_{13}$H$_{26}$O | 0.03%  | -     | GC-MS  | [11] |
| 1,1,6-Trimethyl-1,2-dihydronaphthalene | C$_{13}$H$_{16}$ | 0.41%  | -     | GC-MS  | [11] |
| Ionene    | C$_{13}$H$_{18}$  | 0.09%  | -     | GC-MS  | [11] |
| Damascenone | C$_{13}$H$_{16}$O | 0.28%  | -     | GC-MS  | [11] |
| Ledene oxide | C$_{15}$H$_{24}$O | 0.60%  | -     | GC-MS  | [11] |
| Compounds                        | Molecular Formula | Quantity  | Method    | Reference |
|---------------------------------|------------------|-----------|-----------|-----------|
| 2-tert-Butyl-1,4-dimethoxybenzene | C_{12}H_{18}O_{2} | 0.39% a   | -         | GC-MS     | [11]      |
| (E)-6,10-dimethylnonadeca-5,9- dien-2-one | C_{13}H_{25}O | 0.26% a   | -         | GC-MS     | [11]      |
| 4,6-Dimethyldecane             | C_{14}H_{30}O   | 0.29% a   | -         | GC-MS     | [11]      |
| 3,3,5,6-Tetramethyl-1-indanone  | C_{13}H_{26}O   | 0.23% a   | -         | GC-MS     | [11]      |
| Dihydro-actiridioide           | -                | 1.21% a   | -         | GC-MS     | [11]      |
| 2,3,6-Trimethyl-naphthalene     | C_{13}H_{14}O   | 0.37% a   | -         | GC-MS     | [11]      |
| Megastigmatrienone             | C_{13}H_{28}O   | 0.57% a   | -         | GC-MS     | [11]      |
| 1-[2,3,6-Trimethyl-phenyl]-2- butanone | C_{13}H_{26}O | 3.44% a   | -         | GC-MS     | [11]      |
| 1-[2,3,6-Trimethyl-phenyl]-3- buten-2-one | C_{13}H_{26}O | 0.75% a   | -         | GC-MS     | [11]      |
| Isolongifolene                 | C_{13}H_{24}O   | 0.56% a   | -         | GC-MS     | [11]      |
| Hexahydrofarnesylactone        | C_{14}H_{26}O   | 1.30% a   | -         | GC-MS     | [11]      |
| Farnesylacetone               | C_{14}H_{28}O   | 0.08% a   | -         | GC-MS     | [11]      |
| Methyl palmitate               | C_{15}H_{30}O_{2} | 0.08% a | -         | GC-MS     | [11]      |
| 6E,10E]-7,11,15-trimethyl- methylene-1,6,10,14-hexadeca-tetraene | C_{20}H_{32} | 0.11% a   | -         | GC-MS     | [11]      |
| Phytol                         | C_{20}H_{40}O   | 0.9664% a | -         | GC-MS     | [13]      |
| Docosane                       | C_{22}H_{46}O   | 7.66% a   | -         | GC-MS     | [11]      |
| 1-Docosene                     | C_{22}H_{44}O   | 0.41% a   | -         | GC-MS     | [11]      |
| Tetracosane                    | C_{24}H_{50}O   | 1.45% a   | -         | GC-MS     | [11]      |
| Pentacosane                    | C_{25}H_{52}O   | 17.41% a  | -         | GC-MS     | [11]      |
| Hexacosane                     | C_{26}H_{54}O   | 11.20% a  | -         | GC-MS     | [11]      |
| Pyridine                       | C_{5}H_{5}N     | 0.78%     | -         | GC-MS     | [12]      |
| Hepane,2-methyl-               | C_{6}H_{12}O    | 0.68%     | -         | GC-MS     | [12]      |
| Heptane,3-methyl-              | C_{7}H_{14}O    | 0.50%     | -         | GC-MS     | [12]      |
| Cyclohexane,1,3-dimethyl-,cis- | C_{8}H_{16}O    | 0.78%     | -         | GC-MS     | [12]      |
| Octane                         | C_{9}H_{18}O    | 1.75%     | -         | GC-MS     | [12]      |
| Cyclohexane,1,4-dimethyl-,cis- | C_{9}H_{18}O    | 0.14%     | -         | GC-MS     | [12]      |
| Octane,2-methyl-               | C_{10}H_{20}O   | 0.17%     | -         | GC-MS     | [12]      |
| Cyclohexane,ethyl-             | C_{9}H_{18}O    | 0.43%     | -         | GC-MS     | [12]      |
| Ethylbenzene                   | C_{9}H_{18}O    | 0.83%     | -         | GC-MS     | [12]      |
| Benzene,1,3-dimethyl-o-Xylene  | C_{9}H_{18}O    | 4.63%     | -         | GC-MS     | [12]      |
| Nonane                         | C_{9}H_{20}O    | 2.82%     | -         | GC-MS     | [12]      |
| Benzene,(1-methylethyl)-       | C_{9}H_{12}O    | 0.16%     | -         | GC-MS     | [12]      |
| Benzene,propyl-                | C_{9}H_{12}O    | 1.04%     | -         | GC-MS     | [12]      |
| Benzene,1-ethyl-3-methyl-      | C_{9}H_{12}O    | 5.95%     | -         | GC-MS     | [12]      |
| Benzene,1-ethyl-4-methyl-      | C_{9}H_{12}O    | 2.39%     | -         | GC-MS     | [12]      |
| Benzene,1,3,5-trimethyl-       | C_{9}H_{12}O    | 4.31%     | -         | GC-MS     | [12]      |
| Benzene,1-ethyl-2-methyl-      | C_{9}H_{12}O    | 2.55%     | -         | GC-MS     | [12]      |
Table 1. Cont.

| Compounds                                      | Molecular Formula | Quantity  | Method  | Reference |
|------------------------------------------------|------------------|-----------|---------|-----------|
| Leaves Seeds                                   |                  |           |         |           |
| Benzene,1,2,4-trimethyl-                       | C₉H₁₂             | 16.96%    | -       | GC-MS     | [12]      |
| Benzene,1,2,3-trimethyl-                       | C₉H₁₂             | 4.14%     | -       | GC-MS     | [12]      |
| Indane                                         | C₁₀H₁₈            | 0.39%     | -       | GC-MS     | [12]      |
| Benzene,1-ethyl-2,4-dimethyl-                  | C₁₀H₁₄            | 0.64%     | -       | GC-MS     | [12]      |
| Benzene,1-ethyl-2,3-dimethyl-                  | C₁₀H₁₄            | 0.99%     | -       | GC-MS     | [12]      |
| Benzene,1,2,3,5-tetramethyl-                   | C₁₀H₁₄            | 1.59%     | -       | GC-MS     | [12]      |
| Benzene,1,2,4,5-tetramethyl-                   | C₁₀H₁₄            | 2.12%     | -       | GC-MS     | [12]      |
| Benzene,1,2,3,4-tetramethyl-                   | C₁₀H₁₄            | 0.58%     | -       | GC-MS     | [12]      |
| Naphthalene                                    | C₁₀H₈             | 1.21%     | -       | GC-MS     | [12]      |
| Dimethoate                                     | C₅H₁₂NO₃PS₂       | 0.52%     | -       | GC-MS     | [12]      |
| Nonadecane                                     | C₁₉H₃₈            | 0.22%     | -       | GC-MS     | [12]      |
| Heptacosane                                     | C₂₀H₄₀            | 0.31%     | -       | GC-MS     | [12]      |
| γ-Tocopherol                                    | C₂₀H₄₀O₂           | 0.75%     | -       | GC-MS     | [12]      |
| Sulfurous acid, hexyl pentadecyl ester        | C₂₁H₄₄O₃S         | 0.34%     | -       | GC-MS     | [12]      |
| dl-α-Tocopherol                                 | C₂₆H₅₀O₂           | 1.05%     | -       | GC-MS     | [12]      |
| β-Amyrin                                        | C₃₀H₆₀            | 4.60%     | -       | GC-MS     | [12]      |
| 1,3-Dihydroxyacetone dimer                     | C₆H₁₂O₆           | 3.8551%   | -       | GC-MS     | [13]      |
| Acetic acid, [(amino carboxylamino)oxo]-       | C₃H₂N₂O₄          | 3.2996% a | -       | GC-MS     | [13]      |
| 4(1H)-Pyrimidinone, 2,6-diamino-               | C₄H₆N₄O          | 2.2433% a | -       | GC-MS     | [13]      |
| 4H-Pyran-4-one, 2,3-dihydro-3,5-dihydroxy-6-methyl- | C₄H₆O₄         | 4.1801%   | -       | GC-MS     | [13]      |
| 2-Hexynoic acid                                | C₄H₈O₂            | 1.1214% a | -       | GC-MS     | [13]      |
| Butanedioic acid, 2-hydroxy-2-methyl, (S)-     | C₇H₁₂O₃           | 3.1422% a | -       | GC-MS     | [13]      |
| 3,3-Iminobispropylamine                        | C₆H₁₂N₄          | 1.9275% a | -       | GC-MS     | [13]      |
| 1-Hexanamine                                   | C₆H₁₄N           | 0.0774% a | -       | GC-MS     | [13]      |
| 1,3-Dioxolan-2-one, 4,5-dimethyl-             | C₆H₁₄O₃           | 6.1627% a | -       | GC-MS     | [13]      |
| 2-Butenethioic acid, 3-(ethylthio), S-(1-methylethyl) ester | C₉H₁₆OS₂ | 1.3083% a | -       | GC-MS     | [13]      |
| Propanamide, N,N-dimethyl-                     | C₄H₁₁NO          | 3.0349% a | -       | GC-MS     | [13]      |
| 2-Isopropoxyethyl propionate                   | C₄H₁₆O₃          | 16.8738% a | -     | GC-MS     | [13]      |
| D-Mannohexulose                                | C₇H₁₂O₇           | 2.5622% a | -       | GC-MS     | [13]      |
| Azetidin-2-one, 3,3-dimethyl-4-(1-aminoethyl)- | C₇H₁₄N₂O         | 4.6738% a | -       | GC-MS     | [13]      |
| Carbonic acid, butyl 2-pentyl ester            | C₁₀H₁₆O₃         | 20.6431% a | -       | GC-MS     | [13]      |
| Tetra acetyl-d-xylonic nitrile                 | C₁₄H₁₇NO₃      | 5.0379% a | -       | GC-MS     | [13]      |
| α-D-Glucose                                    | C₆H₁₂O₆         | 3.445% a  | -       | GC-MS     | [13]      |
| 1H-Cyclopenta[c][furan-3(3xH)-one, 6,6α-dihydro-1-(1,3-dioxolan-2-yl)- (3aR,1-trans,6α-cis)- | C₁₀H₁₃O₄ | 1.2587% a | -       | GC-MS     | [13]      |
| Compounds                                                   | Molecular Formula | Leaves Quantity | Seeds Quantity | Method     | Reference |
|-------------------------------------------------------------|------------------|----------------|----------------|------------|-----------|
| 3-(1-(4-Cyano-1,2,3,4-tetrahydronaphthyl)propanenitrile    | C_{14}H_{14}N_{2} | 1.5253%        | -              | GC-MS      | [13]      |
| Quinolinium, 1-ethyl-, iodide                              | C_{11}H_{13}IN   | 1.4067%        | -              | GC-MS      | [13]      |
| N-Isopropyl-3-phenylpropanamide                            | C_{12}H_{17}NO   | 0.9462%        | -              | GC-MS      | [13]      |
| 1,2-Ethanediamine, N-(2-aminooethyl)-                       | C_{4}H_{13}N_{3} | 0.8805%        | -              | GC-MS      | [13]      |
| 1,4-Benzenediol, 2-methyl-                                 | C_{6}H_{12}O_{2} | 4.3169%        | -              | GC-MS      | [13]      |
| Ethene, ethoxy-                                            | C_{4}H_{6}O      | 0.5981%        | -              | GC-MS      | [13]      |
| Dihydroxyacetonine                                         | C_{4}H_{6}O_{3}  | 2.4651%        | -              | GC-MS      | [13]      |
| Glycerine                                                  | C_{3}H_{8}O_{3}  | 1.8656%        | -              | GC-MS      | [13]      |
| Erythritol                                                 | C_{4}H_{8}O_{4}  | 0.5327%        | -              | GC-MS      | [13]      |
| Monomethyl malonate                                         | C_{3}H_{6}O_{4}  | 2.5684%        | -              | GC-MS      | [13]      |
| 4,5-Diamino-6-hydroxypryridimine                           | C_{3}H_{12}N_{4}O_{2} | 0.6434%   | -              | GC-MS      | [13]      |
| Furan, 2,3-dihydro-4-methyl-                               | C_{4}H_{6}O      | 0.2105%        | -              | GC-MS      | [13]      |
| Catecholborane                                             | C_{6}H_{8}BO_{2} | 0.6806%        | -              | GC-MS      | [13]      |
| 2-Fluoropyridine                                           | C_{3}H_{4}FN     | 0.8121%        | -              | GC-MS      | [13]      |
| 1,2,3-Propanetriol, 1-acetate                              | C_{6}H_{10}O_{4} | 1.4375%        | -              | GC-MS      | [13]      |
| 3,4-Furandiol, tetrahydro-, trans-                         | C_{4}H_{6}O      | 0.1746%        | -              | GC-MS      | [13]      |
| 1-Nitro-β-d-arabino furanone, tetraacetate                  | C_{15}H_{17}NO_{11} | 0.5114%    | -              | GC-MS      | [13]      |
| 1,8-Diamino-3,6-dioxoanate                                 | C_{4}H_{12}N_{2}O_{2} | 0.1172%   | -              | GC-MS      | [13]      |
| 1,7-Diaminoheptane                                         | C_{7}H_{10}N_{2} | 1.7997%        | -              | GC-MS      | [13]      |
| N,N-Dimethylacetamide                                       | C_{4}H_{4}ON     | 0.454%         | -              | GC-MS      | [13]      |
| 2-Oxoglutaric acid                                         | C_{4}H_{6}O_{5}  | 0.5466%        | -              | GC-MS      | [13]      |
| Oxazolidine, 2-ethyl-2-methyl-                              | C_{4}H_{11}NO    | 0.9008%        | -              | GC-MS      | [13]      |
| Heptanal                                                   | C_{7}H_{14}O     | 0.7112%        | -              | GC-MS      | [13]      |
| 6-Methoxy-3-pyridazinethio                                 | C_{5}H_{12}N_{2}O_{3} | 0.4293%   | -              | GC-MS      | [13]      |
| 3-Piperidinol                                              | C_{5}H_{11}NO    | 0.5971%        | -              | GC-MS      | [13]      |
| 1,3-Propanediol, 2-ethyl-2-(hydroxymethyl)-                 | C_{6}H_{12}O_{3} | 21.1909%       | -              | GC-MS      | [13]      |
| Benzeneacetoinitrile, 4-hydroxy-                           | C_{4}H_{7}NO     | 3.4763%        | -              | GC-MS      | [13]      |
| Benzenebutanal, γ, 4-dimethyl-                             | C_{12}H_{16}O    | 0.395%         | -              | GC-MS      | [13]      |
| 2(4H)-Benzofuranone, 5,6,7,α-tetrahydro-4,4,7,α-trimethyl- | C_{11}H_{16}O_{2} | 0.4491%      | -              | GC-MS      | [13]      |
| Ethanamine, N-ethyl-N-nitroso-                              | C_{4}H_{10}N_{2}O | 5.2161%       | -              | GC-MS      | [13]      |
| Propanoic acid, 2-methyl-, octyl ester                     | C_{12}H_{24}O    | 15.0279%       | -              | GC-MS      | [13]      |
| 3-Deoxy-d-mannonoic lactone                                | C_{7}H_{13}O_{2} | 3.2947%        | -              | GC-MS      | [13]      |
| d-Glycerol-dido-heptose                                     | C_{7}H_{13}O_{7} | 0.3814%        | -              | GC-MS      | [13]      |
| D-erythro-Pentose, 2-deoxy-                                 | C_{5}H_{10}O     | 0.3314%        | -              | GC-MS      | [13]      |
| N-Methoxy-1-ribofuranosyl-4-imidazolocarboxylic amide      | C_{10}H_{15}N_{3}O_{6} | 0.5345%    | -              | GC-MS      | [13]      |
| Formamide, N,N-dimethyl-                                   | C_{3}H_{7}NO     | 0.5847%        | -              | GC-MS      | [13]      |
### Table 1. Cont.

| Compounds                                      | Molecular Formula | Quantity | Method | Reference |
|------------------------------------------------|-------------------|----------|--------|-----------|
| d-Talonic acid lactone                         | C_{6}H_{10}O_{6}  | 0.5651%  | GC-MS  | [13]      |
| Sorbitol                                        | C_{6}H_{14}O_{6}  | 0.482%   | GC-MS  | [13]      |
| Allo-Inositol                                    | C_{6}H_{12}O      | 0.8749%  | GC-MS  | [13]      |
| D-chiro-Inositol, 3-O-(2-amino-4-((carboxyiminomethyl)amino)-2,3,4,6-tetradeoxy-α-D-arabino-hexopyranosyl)- | C_{14}H_{25}N_{3}O_{9} | 1.595%  | GC-MS  | [13]      |
| Muco-Inositol                                   | C_{6}H_{12}O      | 2.0264%  | GC-MS  | [13]      |
| Inositol                                        | C_{6}H_{12}O      | 2.0545%  | GC-MS  | [13]      |
| Cyclohexane, 1-methyl-4-(2-hydroxyethyl)-       | C_{6}H_{10}O_{1}  | 0.4822%  | GC-MS  | [13]      |
| Hexadecanoic acid, methyl ester                | C_{12}H_{26}O_{2} | 0.8519%  | GC-MS  | [13]      |
| Phenol, 2-methyl-                               | C_{7}H_{6}O       | 0.3737%  | GC-MS  | [13]      |
| (1S)-Propanol, (2S)-[(tert.butyloxycarbonyl)amino]-1-phenyl- | C_{14}H_{21}NO_{3} | 0.967%  | -       | [13]      |
| 9-Octadecenoic acid (Z)-, methyl ester         | C_{18}H_{36}O_{2} | 1.0307%  | GC-MS  | [13]      |
| 9,12,15-Octadecatrienoic acid, (Z,Z,Z)-         | C_{18}H_{30}O_{2} | 5.0063%  | GC-MS  | [13]      |
| 4-Allyl-3-(dimethylhydrazono)-2-methylhexane-2,5-diol | C_{12}H_{24}N_{2}O_{2} | 0.4257%  | GC-MS  | [13]      |
| Benzyl-β-d-glucoside                            | C_{13}H_{26}O_{7} | 0.6056%  | GC-MS  | [13]      |
| 4,6-dimethyl-2-propyl-1,3,5-dithiazinane        | C_{8}H_{17}NS_{2} | 0.1698%  | GC-MS  | [13]      |
| 1,2-Benzenediol, 2-methyl-                      | C_{6}H_{12}O_{2}  | 0.545%   | GC-MS  | [13]      |
| 9-Octadecenamide, (Z)-                         | C_{18}H_{35}NO    | 1.4694%  | GC-MS  | [13]      |

| Abbreviations: a: calculated as peak area; b: mean value of different samples; c: mature/old leaves; d: tender/young leaves; e: vegetative plants; f: obtained considering a moisture of 75%; g: flowering plants. |

#### 3.1. Non-Polar Compounds

Fatty acids are present in very high quantities. In their work, Lee et al. [9] performed an extraction on *M. oleifera* leaves powder using 70% methanol and then partitioning the outcome with hexane, dichloromethane, ethyl acetate, butanol, and water. Finally, the last part of extraction was conducted by supercritical fluid extraction (SFE). The most abundant fatty acid found in the resulting extracts were cis-11-Eicosenoic acid (also called gondoic acid), a monounsaturated acid contained in many oils extracted from plants, followed by palmitic acid (23.65%), and then linoleic acid and oleic acid (6.84% and 5.92%), and apart from them, the other ones are present in quantities lower than 5%. In seeds extracts, the highest content of fatty acid was registered for oleic acid (74.5%), then palmitic acid (7.92%), stearic acid (7.10%), and then minor fatty acids present in smaller quantities [9]. These fatty acids are important because they take part in metabolic processes that lead to the synthesis of ω-3 fatty acid, which have important roles in regulation processes of cellular development and immune system [14]. As showed by the same group, cis-11-eicosenoic acid, linoleic acid, and oleic acid exhibited an antimicrobial effect on *Staphylococcus aureus*, avoiding biofilm formation by inhibition of cell growth [9,10]. The oily fraction of *M. oleifera* leaves is also very important for cosmetic usage, as they are used as a fundamental part of skin and hair products [10].
Moreover, alkanes and alkanes derivatives are another abundant class of molecules in moringa extracts. Chelliah et al. [10] conducted a complete analysis on *M. oleifera*, performing different extractions with hexane, methanol and water. The extracts were then analyzed with GC-MS, and several compounds were found, the most abundant of which were 1,30-triacontanediol (14.98%) and octacosane (8.57%), in leaves from Madurai variety, and nonacosane (15.55%) and γ-Sitosterol (9.56%) for Chennai variety. Conversely, in seeds they found 6-octadecenoic acid (52.24%), n-hexadecanoic acid (palmitic acid) (6.17%), oleic acid (5.12%) as major compounds in Madurai variety, and (Z)-13-docosenamide (13.62%), propionamide (4.48%), ethyl oleate (4.33%) in Chennai variety [10]. Most of them are alkanes or their derivatives, and they are synthesized by plants to build up an external waxy layer (also called epicuticular wax) to protect leaves from surface wetting and water evaporation. 6-octadecenoic acid, also called petroselinic acid, and palmitic acid are two fatty acids regularly present in plant oils. γ-Sitosterol belongs to the class of the phytosterols, and it is an isomer of β-sitosterol, well known because there is scientific evidence that proved the antidiabetic properties of γ-Sitosterol in vivo against type II diabetes [17,18]. Similar compounds were found by Zhao & Zhang [12] that carried on a Soxhlet extraction with hexane. The procedure was conducted for 8 h, and when the dried extract was analyzed by GC-MS analysis, they stated that the most abundant compounds in the extract were nonacosane (18.65%), 1, 2, 4-trimethylbenzene (16.96%), heptacosane (7.45%) [12].

Essential oils are mixtures of dozens or hundreds of molecules, mainly volatiles compounds belonging to the classes of terpenes, alcohols, aldehydes, ketones, esters, phenols and others, often extracted with hydrodistillation, solvent-free microwave extraction (SFME) or supercritical fluid extraction using supercritical carbon dioxide. Despite a wide utilization of essential oils in alternative medicine and aromatherapy, they possess some important properties, such as: antibacterial activity against gram-positive species, probably because of external membrane disruption due to phenolic compounds activity; antioxidant activity, probably due to secondary metabolites with conjugated double bonds; anti-inflammatory activity, triggered by a wide variety of compounds, including antioxidants. An interesting analysis on essential oils was conducted by Chuang et al. [11] that extracted essential oils from *M. oleifera* leaves using hydrodistillation and tested the extract on fungi cultures finding out that both essential oils and seeds extracts had a positive effect against fungi development, probably exerting anti-fungal activity by disrupting the cellular membrane. The non-polar extract obtained were run through GC-MS instrument, and the main compounds found were mostly alkanes and alkenes, including hexacosane, pentacosane, tetracosane making up almost 30% of total extracts. Moreover, phytol was identified (7.66% based on peak area) [11]. Phytol is a diterpene alcohol very abundant in plants because it is one of the two components of chlorophyll. Phytol is used as a food additive, and a group of scientists have proposed that it can be useful to treat schistosomiasis, a disease caused by a parasite [19]. Lastly, a small quantity of terpenes and terpene derivative were found, with hexahydrofarnesylacetone being the most abundant one (1.30%) and then smaller quantities of linalool oxide, farnsylacetone, isolongifolene α-ionene, and α- and β-ionone, barely present in traces [11].

3.2. Polar Compounds

*M. oleifera* leaves are very high in protein content, and thus, they are highly rich in amino acids. All the essential amino acids can be found in *M. oleifera* leaves and they contribute to more than 50% of total amino acids content. Amino acids presence in leaves was represented by high quantities of glutamic acid. Chelliah et al. [10] found 15.86 g over 16 g of nitrogen (g/16 gN) of glutamic acid in Madurai variety and 15.33 g/16 gN in Chennai variety, while Lalas et al. [14] reported 268.7 mg/100 g of glutamic acid. According to Chelliah et al. [10], methionine was the least abundant in Madurai variety, which was found in quantity of 1.90 g/16 gN, and histidine was the lowest in Chennai variety, present in 3.12 g/16 gN. Conversely, according to Lalas et al. [14] the least abundant was cysteine.
which was present in 29.6 mg/100 g of dried leaves. In seeds, the most present amino acid was glutamic acid (14.23 g/16 gN in Madurai, 14.74 g/16 gN in Chennai), whereas the least abundant was threonine for Madurai variety (3.20 g/16 gN) and valine for Chennai variety (2.37 g/16 gN) [10,14]. Regarding glucosinolates, Lopez-Rodriguez et al. [15] have collected few interesting studies that performed extractions with 70% and 80% methanol, and by using HPLC and UPLC they found different isomers of glucoromin (4-[(α-L-rhamnosyloxy)-benzyl]-glucosinolate) (Figure 2(1)) in M. oleifera leaves and seeds [15]. These compounds belong to the class of the glucosinolates, natural compounds synthesized by plants responsible for pungent flavor and smell. Their basic structure is formed by a sugar (thioglucose group), bound to a central carbon atom, and the carbon is then bound to a side chain, a nitrogen and a sulphate group, derived from an amino acid. The side chain varies depending on the glucosinolate, determining their biological activity. The main role of glucosinolates is plant defense against infections and diseases. There can be three types of glucosinolates depending on the amino acid that participates to the synthesis of them: aliphatic, aromatic or indolic glucosinolates. Aromatic glucosinolates seems to be the main ones that can be found in M. oleifera leaf extracts [15]. Moreover, glucosinolates are precursors of isothiocyanates, molecules containing the -N=C=S group that have showed biological activity in vitro, in particular, they showed hypoglycemic effects, antioxidant and anti-inflammatory effects, anticancer and chemopreventive effects, and it seems that isothiocyanates from M. oleifera are more stable than the ones found in other crucifers, because they contain rhamnose, which makes the structure more stable [15]. The review written by Leone et al. [16] shows a series of tables with many phenolic acids, molecules that belong to the class of polyphenols. They are formed by a monohydroxybenzoic acid unit and different side groups bound the phenolic ring depending on the compound and their importance is due to the documented antioxidant, anti-inflammatory, antimutagenic and anticancer activities [16]. Phenolic acids were extracted with polar solvents, using different percentages of methanol, water, ethanol, and analyzed via HPLC and LC-MS. The major compounds belonging to this category were caffeic acid, chlorogenic acid (Figure 2(2)), o-coumaric acid (Figure 2(3)), ellagic acid, ferulic acid and gallic acid, all of them present in quantities between 0.018 mg/g of dried leaves (ellagic acid) and 6.457 mg/g of dried leaves (coumaric acid). More polyphenols are listed in the following tables of the same study. The main polyphenols found are epicatechin (Figure 2(4)), kaempferol (Figure 2(5)), myricetin, quercetin (Figure 2(6)). Myricetin, quercetin and kaempferol are three natural polyphenols that belong to the class of flavonoids, molecules that have a common 15-carbon backbone made by two phenyl rings and a heterocyclic ring between them. The difference between these three molecules lies in the different numbers of hydroxyl groups bound to the basic structure, and thus they exert similar effects on human health. Previous studies listed in the review of Salvamani et al. have shown that quercetin could have several beneficial effects on human health, such as vasorelaxation to lower blood pressure, positive effects on dyslipidemia, on obesity by reducing accumulation of fats and promoting beta oxidation, anti-inflammatory, anti-atherogenic and anti-atherosclerotic effects in in vivo experiments [20]. Antimicrobial activity of quercetin was also reported. In a recent work, Montone et al. [21] demonstrated the importance of quercetin combined with lactoferrin and hydroxyapatite (LF-HA)—whose beneficial biological effects were widely investigated [22–24]—as antimicrobial agent in food industry. Kaempferol shows a similar impact on health, being a powerful agent against cardiovascular diseases endothelial damage, oxidative stress and against arteriosclerosis. Myricetin effects are explained by the same group, who listed several beneficial effects of this flavonoid, such as antioxidant, antiviral, anticarcinogenic, antiplatelet, hypoglycemic, and cytoprotective properties. Similarly to quercetin, it also showed positive effects on hypertension and accumulation of fats in the body of laboratory rats [20]. Quercetin was the most abundant flavonoid in M. oleifera extracts. The highest quantity of quercetin found among the studies is 16.64 mg/g of dried weight (dw), whereas for myricetin is 5.804 mg/g dw, and for kaempferol is 4.59 mg/g dw. The last relevant molecules considered in Leone et al. [16] are the glucosinolates, in particular
they listed 4-(α-L-acetylrhamnopyranosyloxy)-benzyl-isothiocyanate (commonly known as moringin) (Figure 2(7)) and three isomers of 4-O-(α-Lacetylrhamnopyranosyloxy)-benzyl-isothiocyanate. Totally, they ranged from 21.84 mg/g dw to 59.4 mg/g dw among the two works considered in the above-mentioned review [16]. Moreover, some studies have reported cancer chemoprotective activity, allelopathic activity and repellent/insecticidal activity [25].

Finally, Bhalla et al. [13] conducted identification on two different extracts of M. oleifera. They found out that carbonic acid, butyl 2-pentyl ester (20.64%), 2-Isopropoxyethyl propionate (16.87%), 4H-Pyran-4-one, 2,3-dihydro-3,5-dihydroxy-6-methyl- (8.99%), 1,3-Dioxolan-2-one, 4,5-dimethyl- (6.16%) were the compounds present in highest peak area in the aqueous extract. A methanolic extraction was performed as well, and because of the different polarity, more compounds were identified in the methanolic extraction compared to the aqueous. In methanolic extracts, 1,3-propanediol, 2-ethyl-2-(hydroxymethyl)- (21.19%), propionic acid, 2-methyl-, octyl ester (15.02%), ethanamine, N-ethyl-N-nitroso- (5.21%), and 9,12,15-octadecatrienoic acid were the most abundant. These extracts were also tested to examine their antioxidant activity on free radicals, with the methanolic extract showing a higher capacity of scavenging free radicals than the aqueous extracts [13]. As stated by the authors, this is probably due to higher polyphenolic molecules found in methanolic extract. A high antioxidant activity can be useful because of a high request both by food and cosmetic industry, as they are used for skin treatment and for food enrichment.

4. Pharmacology

The pharmacological studies on M. oleifera are listed in Table 2. Different plant organs were extracted with a series of organic solvents and the obtained extracts were tested to determine and confirm the bioactivity known by the ethnomedicine. Pure compounds were also tested and showed interesting bioactivities (Table 2 and Figure 2).
Table 2. Pharmacological activity of organic compounds or extracts of *Moringa oleifera*.

| Material Tested                        | Plant Organ | Disease Model Used | Disease/Effect | Active Principle | Quantity | References |
|----------------------------------------|-------------|--------------------|----------------|------------------|----------|------------|
| Aqueous, ethanolic and methanolic extracts | Leaves     | Anti-microbial     | Enteropathogenic and orthopedics wounds bacteria and fungi | Inhibitory properties for wounds and certain fungal infections | n.i.     | [26]       |
| Ethanol (80%), methanol (70%), hydroalcoholic extracts | Leaves | Metabolic disease and diabetes | Beta-carotene-linoleic acid system, Liposome peroxidation, and liver microsomes | Antioxidant | quercetin and kaempferol | 1 g 80% methanol, and 70% ethanol hydroalcoholic extracts have DPPH equal to 20, 57, 35 mg of pure quercetin | [27] |
| Ethanol solution (90%)                  | Leaves     | Antioxidant        | HPLC, MS-MS, chemometrics, DPPH, ABTS, FRAP | Correlation between phytochemical profile and antioxidant potential | kaempferol 3-O-rutinoside, quercetin 3-O-(6′′-malonyl-glucoside), kaempferol 3-O-glucoside, and quercetin derivative | n.r. | [28] |
| Ethanolic and methanolic extract       | Leaves     | Antioxidant        | DPPH           | Higher antioxidant potential observed for ethanolic and methanolic extract | n.i.     | [29]       |
| Ethyl acetate extract                  | Leaves     | Cardiovascular     | RAW Macrophages | Decreased expression of inflammatory markers | phenolics | LC50, 212.73 µg/mL, LC10, 57.53 µg/mL | [30] |
| Hot water and ethanol extracts         | Leaves     | Cancer             | Cancer breast cells | Inhibition of NF-kB signaling | n.i.     | IC50 0.31 and 0.26 µg/mL, hot water and ethanolic extract, respectively | [31] |
| Hydroalcoholic extract (80% ethanol)   | Leaves     | NAFLD              | Mice           | Protected HFD-induced liver dam | n.i.     | 150 mg/kg body weight, administered by gavage | [32] |
| Leaf powder                            | Leaves     | Anti-inflammatory  | Guinea pigs   | Lower cholesterol and triglycerides and reduced inflammation in the liver | n.i.     | 10% or 15% Moringa enriched diet, 6 weeks | [33] |
| Leaf powder                            | Leaves     | Cardiovascular     | High-fat fed rats | Decrease cholesterol absorption | β-sitosterol | Bread fortified with extract at 10% & 15% for 45 days | [34] |
| Leaf powder                            | Leaves     | NAFLD              | Guinea pigs   | Decrease expression of CD68, SERBP1c | n.i.     | 10% or 15% Moringa diets with 0.25% dietary cholesterol to induce hepatic steatosis, 6 weeks | [35] |
| Leaf powder                            | Leaves     | Protein deficiency | Fish          | Protein supply | n.i.     | 10%, 20%, and 30% of Moringa leaf meal of total dietary protein. The 10% gave the best response | [36] |
Table 2. Cont.

| Material Tested         | Plant Organ | Disease                        | Model Used                                      | Observed Effect                                           | Active Principle                                    | Quantity                                      | References         |
|-------------------------|-------------|--------------------------------|------------------------------------------------|----------------------------------------------------------|-------------------------------------------------------|-----------------------------------------------|--------------------|
| Methanolic extract      | Leaves      | Obesity                        | High-fat induced obesity rats                   | Anti-obesity properties                                   | n.i.                                                  | 200 mg and 400 mg/kg reduced atherogenic index (1.7 ± 0.6 and 0.87 ± 0.76, respectively) | [37]               |
| N, α-L-Rhamnosyl vincosamide | Leaves | Cardiovascular | Isoproterenol (ISO)-induced cardiac toxicity in rats | Increasing the levels of troponin-T, creatine kinase-MB, lactate dehydrogenase and glutamate pyruvate transaminase | N, α-L-Rhamnosyl vincosamide | 40 mg/kg for 7 days, oral administration n | [38]               |
| Total Alkaloids         | Leaves      | Antihypertensive               | Frog heart                                     | Negative inotropic effect on the heart                    | Total Alkaloids                                      | 3 to 48 ng                                   | [39]               |
| Water extract           | Leaves      | Antioxidant                    | DPPH                                           | Gold metal ions reduction (Au+ to Au°) to form nanoparticles | n.i.                                                  | 5 mL extract + 100 mL 1 mM aqueous gold chloride soln | [40]               |
| Hydroalcoholic extract  | Leaves      | Cytotoxic and anticancer       | Male Wistar rats                               | Decreasing tumor markers (α-fetoprotein and carcinoembryonic antigen) and increasing apoptosis | Male Wistar rats, 7–14 days                         | 150 mg/kg–250 mg/kg Male Wistar rats, 7–14 days |                    |
| Water extract           | Leaves      | Antioxidant                    | DPPH                                           | Improvement extraction for antioxidant tests              | tannins                                              | T 80.54°C for 12.19 min. Protein 17.4%, anti-oxidant activity 10.3 µg/mL, tannin 7.8% | [41]               |
| Leaves                  | Antioxidant | DPPH                           | Tannins were determined as the active principle for the activity | tannins                                                  | n.r.                                                 | [25]                           |                    |
| Leaves                  | Antioxidant | DPPH                           | Tannins were determined as the active principle for the activity | tannins                                                  | n.r.                                                 | [25]                           |                    |
| Leaves                  | Cardiovascular and Diabetes | Starch and phosphate buffer solution | Inhibition of α-glucosidase                       | quercetin                                                | IC50 0.78 ± 0.21 mg/mL (inhibitor sucrose esterase) | At 10 mg/mL inhibited cholesterol micelle formation of 40.22 ± 2.64% | [42]               |
Table 2. Cont.

| Material Tested | Plant Organ | Disease Model Used | Observed Effect | Active Principle | Quantity | References |
|-----------------|-------------|--------------------|-----------------|------------------|----------|------------|
| Aqueous extracts (AE) | Seeds | Anti-allergic, and Immunosuppressive properties | Human peripheral blood mononuclear cells | Cytotoxic AE and cMOL, not cytotoxic wsMOL | | |
| | | | NCI-H292, HT-29 and HEp-2 cancer cell lines | Weekly/moderately cytotoxic AE, cMOL, wsMOL | | |
| | | | Murine erythrocytes | Not hemolytic AE, cMOL, wsMOL | | |
| | | | Lipopolysaccharide-stimulated murine macrophages | AE, cMOL, and wsMOL regulated the production of nitric oxide, TNF-α and IL-1β | | |
| | | | Mouse model of carrageenan-induced pleurisy | AE reduced leukocyte migration | | |
| Ethanol extracts | Seeds | Anti-inflammatory, and anti-diabetic | Human peripheral blood mononuclear cells | Bio-accessibility of 1, using the TNO Intestinal Model (TIM-1), determined as 61% and 62% in fasted and fed states, respectively | Isothiocyanate 1 | 50 mg/kg intravenously with pure 1 or orally gavage with seed extract |
| Methanolic extract | Seeds | Anti-inflammatory | Influenza A virus (H1N1) | Inhibitory effects against the H1N1 virus | Moringa A, Glucomoringin, and Vitexin | IC50 0.26 ± 0.03, 0.98 ± 0.17, and 3.42 ± 0.37 µg/mL for Moringa A, Glucomoringin and Vitexin, respectively |
| Phenolic extract | Seeds | Anti-microbial | Raw264.7 cells | Decreasing the levels of TNF-α, IL-6, and IL-1β | Moringa A, Glucomoringin, and Vitexin | IC50 0.9 ± 0.05 |
| Stem powder | Stems | Anti-inflammatory | Keratinocytes and macrophages induced by PGN, TNF-α, or LPS | Suppression of inflammatory factors. Reduction of NLRP3 expression, and inhibition of caspase-1 activation | β-sitosterol | dose range: 7.5 to 30 µM |
| Material Tested | Plant Organ          | Disease         | Model Used                               | Observed Effect                                                                 | Active Principle              | Quantity                          | References |
|-----------------|----------------------|-----------------|------------------------------------------|--------------------------------------------------------------------------------|------------------------------|----------------------------------|------------|
| Ethanolic extracts | Leaves, seeds, pods | Anti-allergic   | Rat basophilic leukemia (RBL-2H3) cells | Extracts and compounds inhibited mast cell degranulation by inhibiting beta-hexosaminidase and histamine release from RBL-2H3 cells, and IL-4 and TNF-α release | β-Sitosterol-3-O-glucoside, Glucomoringin and Quercitin | IC50 of 1.20 ± 0.29, 2.52 ± 0.35 and 2.52 ± 0.15 µg/mL, respectively | [47]       |

n.i. = not identified; n.r. = not reported. Abbreviations: ABTS = 2,2′-Azino-bis(3-ethylbenzothiazoline-6-sulfonic acid; cMOL: coagulant Moringa oleifera lectins; DPPH = 2,2-difenil-1-picrylhydrazyli); FRAP = fluorescence recovery after photobleaching; HDF = high-fat diet; NAFLD = non-alcoholic fatty liver disease; NF-κB = nuclear factor kappa-light-chain-enhancer of activated B cells; RAW = Ralph, rAschke, Watson.
A recent review by Vergara-Jimenez et al. [25] reports the major studies dealing with the use of *M. oleifera* in prevention and alleviation of several chronic conditions, including hypercholesterolemia, high blood pressure, diabetes, insulin resistance, non-alcoholic liver disease, cancer, and overall inflammation.

### 4.1. Leaves: Cardiovascular Activity

Going into details, plant leaves were extracted mainly with hydroalcoholic solvents by different research groups and the extract were tested to evaluate their potential in cardiovascular diseases. Panda et al. [38] isolated from the extracts the alkaloid N, α-L-rhamnopyrinosyl vincosamide (Figure 2(8)) and studied the protective effect in isoproterenol (ISO)-induced cardiac toxicity in rats. Oral administration of the alkaloid at 40 mg/kg for seven days markedly reduced the ISO-induced increase in the levels of serum cardiac markers such as troponin-T, creatine kinase-MB, lactate dehydrogenase, and glutamate pyruvate transaminase, as well as cardiac lipid peroxidation. A parallel increase in the cellular antioxidants suggested its cardio-protective and free radical scavenging potential, which was later confirmed by in vitro study [38]. Rats treated with test compound also improved the ISO-induced abnormal changes in ECG, as well as in cardiac histology. A reduction in myocardial necrosis was further evidenced by the tri-phenyl tetrazolium chloride (TTC) stain in isolated test drug pre-treated rats. The work of Halaby et al. [34] aimed to study the potential effect of fortified bread with *M. oleifera* leaves powder under 5%, 10%, and 15% concentrations to give more protection against hyperlipidemia. Thirty-two male albino rats were used in this experiment. Results indicated that bread fortified with the extract at 10% and even better at 15% when given to rats for 45 days caused reduction of total cholesterol, triglycerides, low-density lipoprotein, and very-low-density lipoprotein. Kidney function was improved and there was significant reduction in uric acid serum, urea, and creatinine compared to the positive control group [34]. The study of Adisakwattana et al. [42] investigated the effect of leaf water extract of *M. oleifera* on inhibition of alpha-glucosidase and pancreatic alpha-amylase related to diabetes mellitus. Moreover, the study also determined in vitro bile acid binding capacity as well as inhibition of cholesterol micellization, pancreatic lipase, and cholesterol esterase activity. Specific inhibition to intestinal sucrase was found (IC50 value of $0.78 \pm 0.21$ mg/mL) along with a markedly inhibition (40.22% \( \pm \) 2.64%) of cholesterol micelle formation [42].

### 4.2. Leaves: Anti-Inflammatory Activity

Kooltheat et al. [30] prepared an ethyl acetate fraction of *M. oleifera* from fresh leaves that showed high levels of phenolic and antioxidant activities. Since macrophages, TNF and related cytokines play an important pathophysiologic role in lung damage induced by cigarette smoke, the authors examined the effects of MOEF on cigarette smoke extract-induced cytokine production by human macrophages. Human monocyte-derived macrophages (MDM) pre-treated with varying concentrations of *M. oleifera* showed decreased production of TNF, IL-6, and IL-8 in response to LPS and cigarette smoke extract. The decrease was evident at both cytokine protein and mRNA levels. Furthermore, the extract inhibited the expression of RelA, a gene implicated in the NF-κB p65 signaling in inflammation. The findings highlight the ability of *M. oleifera* to inhibit cytokines (IL-8) which promote the infiltration of neutrophils into the lungs and others (TNF, IL-6) which mediate tissue disease and damage [30].

### 4.3. Leaves: Antihypertensive Activity

Dangi et al. [39] studied the antihypertensive activity of the leaves extract. Preliminary studies indicated that a leaves water extract is efficacious in reducing the chronotropic and inotropic effects on the isolated frog heart. The alkaloids obtained by extract fractionation converted into their salt form, were tested for their activity on the isolated frog heart. The total alkaloidal salts were found to have a negative inotropic effect on the frog heart [39]. This activity was further characterized by test on the isolated guinea pig ileum. In their
study Bais et al. [37] evaluated the anti-obesity activity of methanolic leaves extract in rats. Chronic administration of high fat diet (HFD) in rats produced hypercholesterolemia leading to an increase in the body weight total cholesterol, triglycerides, and attenuation in the levels of HDL. Treatment of obese rats with the extract for 49 days resulted in a significant decrease in the level of liver biomarkers, organ weight, and blood glucose level [37].

4.4. Leaves: Radical Scavenging and Antioxidant Activity

Siddhuraju et al. [27] studied the radical scavenging capacities and antioxidant activities. All leaf extracts were capable of scavenging peroxyl and superoxyl radicals. Among the three different moringa samples, both methanol and ethanol extracts of Indian origins showed the highest antioxidant activities, 65.1% and 66.8%, respectively, in the beta-carotene-linoleic acid system. Nonetheless, increasing concentration of all the extracts had significantly \( p < 0.05 \) increased reducing power, which may in part be responsible for their antioxidant activity. The major bioactive compounds of phenolics were found to be flavonoid groups such as quercetin and kaempferol [39]. After this paper several other authors reported studied on the antioxidant activity of the leaves extract [25,29]. Xu et al. [28] correlated the antioxidant activity to some flavonoid glycosides (Figure 2(9–11)) while Wahyuni et al. [41] to the tannin content [28,41]. This last paper reported a new approach based on HPLC, tandem MS spectrometry (MS-MS), and chemometrics allowing to correlate specific flavonoid glycosides in the extract with the antioxidant activity tested. Tiloke et al. [40] studied the antioxidant potential of the extract by the synthesis of phytonanoparticles and the fundamental role as a potential antiproliferative agents against cancer. The resultant gold phytonanoparticles are useful in cancer therapies with improved survival rates and quality of life.

4.5. Leaves: Anticancer Activity

Khalafalla et al. [31] tested different leaves extracts against leukemia and hepatocarcinoma cells in vitro. The extracts could kill majority (70–86%) of the abnormal cells among primary cells harvested from 10 patients with acute lymphoblastic leukemia and 15 with acute myeloid leukemia, as well as a culture of hepatocarcinoma cells (75% death), but most significantly by the hot water and ethanol extracts.

4.6. Leaves: Hepatoprotective and Nutraceutical Activity

Consumption of high-fat diet (HFD) induces nonalcoholic fatty liver disease) and may lead to multiple complications affecting human health. In the study of Das et al. [32] the preventive as also curative hepatoprotective activity effect of M. oleifera leaf extract in alleviating HFD induced liver injury in mice has been reported. Results suggested that M. oleifera leaf extract treatment protected HFD-induced liver damage as indicated by histopathology and liver enzyme activity compared to only-HFD fed group \( p < 0.05 \). Interestingly, early signs of HFD-induced fatty liver were also alleviated by M. oleifera leaf extract. Moreover, significant increase in endogenous antioxidant parameters and lower lipid peroxidation were found in liver of all M. oleifera leaf extract treated groups [32].

Almatrafi et al. [33,35] studied the mechanisms by which M. oleifera leaves modulate hepatic lipids on guinea pigs. Low moringa or 15% high moringa diets with 0.25% dietary cholesterol to induce hepatic steatosis. This study demonstrates that M. oleifera leaves may prevent hepatic steatosis by affecting gene expression related to hepatic lipids synthesis resulting in lower concentrations of cholesterol and triglycerides and reduced inflammation in the liver.

Richter et al. [36] evaluated the suitability of freeze-dried moringa leaf meal as alternative protein source for Nile tilapia. Three experimental diets were formulated substituting 10%, 20%, and 30% of the total dietary protein of fish with plant leaves. Diets with higher % of moringa leaves significantly depressed growth performance of the fish compared the other diets. Total phenolics, non-hemolytic saponins, phytic acid, neutral detergent
fiber, and acid detergent fiber in the 30% diet may have contributed to the poorer growth performance in these groups thus suggesting the 10% substituting the best option.

4.7. Leaves: Antimicrobial Activity

Oluduro et al. [26] investigated the antimicrobial activities of the leaf extract of on a series of enteropathogenic and orthopaedics’ wounds bacteria and fungi using paper disc diffusion method. All the leaf extracts showed little inhibitory effect on the enteropathogens, whereas aqueous and methanolic extracts showed appreciable inhibitory effects on the orthopaedic’s wounds bacteria at 30 mg/mL. Ethanolic extract did not show any zone of growth inhibition on the wound bacteria. Minimum inhibitory concentration was 20 mg/mL on all the enteropathogens and ranged from 3.75 to 30 µg/mL on the orthopaedics’ wounds organisms. The study showed that leaves possess inhibitory properties thus can serve as an alternative therapy for wounds and certain fungal infections.

4.8. Leaves, Stems, Pods: Anti-Allergic Activity

The anti-allergic activity of the extracts of leaves, seeds, and pods and of the isolated compounds (β-sitosterol-3-O-glucoside (Figure 2(12)), glucomoringin and quercitin) was evaluated by Abd Rani et al. [47] using rat basophilic leukaemia (RBL-2H3) cells for early and late phases of allergic reactions using as positive control the drug, ketotifen fumarate. The early phase was determined based on the inhibition of beta-hexosaminidase and histamine release, while the late phase was based on the inhibition of interleukin (IL-4) and tumor necrosis factor (TNF-α) release. Both extract and pure compounds showed inhibitions (Table 2) thus indicating a potential use as anti-allergic drug candidate.

4.9. Seeds: Anti-Inflammatory Activity

Wolff et al. [48] reported from the ethanolic seed extract high concentrations of a glucosinolate with isothiocyanate functional group (MIC-1, 1) a compound possessing anti-inflammatory and antidiabetic properties. Because it was not characterized metabolically, the authors studied its bioaccessibility using a human intestinal model and bioavailability using serum from treated rats. The results suggest that the compound remains largely unmodified during uptake, unlike other isothiocyanates, and has favorable bioaccessibility and bioavailability characteristics for a potential therapeutic agent.

The extracts of moringa seeds are used in rural areas of developing countries to treat drinking water because the seeds contain lectins, carbohydrate-binding proteins able to reduce water turbidity because of their coagulant activity. The study by Araújo et al. [43] evaluated the cytotoxic and anti-inflammatory properties of the aqueous seed extract on NCI-H292, HT-29, and HEp-2 cancer cell lines, and on murine erythrocytes (Table 2). In particular, the extracts exhibited anti-inflammatory activity on lipopolysaccharide-stimulated murine macrophages by regulating the production of nitric oxide, TNF-α and IL-1β. The aqueous seed extract reduced leukocyte migration in a mouse model of carrageenan-induced pleurisy; the myeloperoxidase activity and nitric oxide, TNF-α and IL-1β levels were similarly reduced. Histological analysis of the lungs showed that the extract reduced the number of leukocytes.

4.10. Seeds: Antiviral Activity

Xiong et al. [44] isolated eleven compounds from methanol seeds extract, including two previously unknown and nine known compounds. These compounds were authenticated as a carbamate, three phenylglycosides, four phenol glycosides, two nucleosides, and one flavonoid. Antivirus activity analyses revealed that Moringa A, glucomoringin, and Vitexin (Figure 2(13)) possessed strong inhibitory effects against the H1N1 virus, having IC50 values in the range of IC50 = 0.26 ± 0.03, 0.98 ± 0.17, and 3.42 ± 0.37 µg/mL, respectively. Furthermore, these three compounds could decrease the levels of TNF-α, IL-6, and IL-1β, which occur in hosts because of H1N1 infections.
4.11. Seeds: Anticancer Activity

Five-Fluorouracil (5-FU) is a strong anticancer agent commonly used for the treatment of various malignancies. Famurewa et al. [49] explored whether moringa seed oil could protect against 5-FU-induced nephrotoxicity and its mechanism of action in Wistar rats. Rats were subjected to prophylactic oral treatment of moringa seed oil (Table 2). The data obtained suggested that 5-FU-induced nephrotoxicity by oxidative stress, exacerbation of pro-inflammation and apoptosis. The inhibition of the alterations by moringa seed oil is relevant in the clinical management of 5-FU nephrotoxicity in cancer patients.

4.12. Seeds: Antioxidant Activity

Phenolic compounds, antioxidant and antibacterial activities of defatted seed flour were investigated by Singh et al. [45]. The results showed that extractability of phenolic compounds was significantly higher ($p < 0.05$) in bound phenolic extract (4173.00 ± 32.22 mg gallic acid equivalents (GAE)/100 g) than in free phenolic extract (780.00 ± 14.2 mg GAE/100 g) and it showed higher antioxidant and antimicrobial activities. The IC50 value for DPPH radical scavenging activity was 0.9 ± 0.05 and 14.9 ± 0.07 mg/mL for bound phenolic and free phenolic extracts, respectively. Bound phenolic extract was more effective (minimum inhibitory concentration (MIC), 0.06–0.157%) than free phenolic extract (MIC, 0.117–0.191%) against tested bacteria. The data indicated that moringa seeds could be a good source of antioxidants and antibacterials for food and pharmaceutical industries.

Finally, Liao et al. [46] identified β-sitosterol isolated from *M. oleifera* stems as an anti-inflammatory compound on two cell lines, keratinocytes and macrophages induces by PGN, TNF-α, or LPS. β-sitosterol over a dose range of 7.5 to 30 µM, dispersed in the medium of the well as nanoparticles with diameters of 50 ± 5 nm, was able to suppress the secretion of the above-mentioned inflammatory factors.

5. Conclusions

*M. oleifera* is an ancient medicinal plant growing in a variety of environments and climates, able to resist harsh environments that recently received an increased interest in the food industry and as phytopharmaceuticals.

The phytochemical analysis of the plant’s organs identified metabolites belonging to different classes, including flavonoids, anthocyanins, isothiocyanates, anthraquinones, alkaloids, essential oils, tannic acid, saponins, steroids, terpenoids, and cardiac glycosides. Pharmacological studies confirmed the use of the plant as traditional medicine, showing bioactivity including hepatoprotective, antihypertensive, cholesterol-lowering, anti-ureolithiasis, antifertility, antidiabetic, and antioxidant activity, nutraceutical and antimicrobial properties. In addition, most published papers reported biological studies on the plant extracts while only few studies tested pure compounds. Interestingly, these compounds belong to the class of phenolics, including phenylpropanoids, flavonoids, flavonoid O-glycosides, flavonoid C-glycosides, and glucosinolates. Thus, the reported paper reviewing the literature on the phytochemistry and pharmacology of the plant indicates that more research is needed to investigate the chemical and biological properties on a larger scale and to clarify the mechanism of action. This aspect will be crucial for its use in phytopharmaceutical, nutraceutical, and food industries.

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30. Kooltheat, N.; Pankla Sranujit, R.; Chumark, P.; Potup, P.; Laytragoon-Lewin, N.; Usuwanthim, K. An ethyl acetate fraction of Moringa oleifera Lam. inhibits human macrophage cytokine production induced by cigarette smoke. *Nutrients* **2014**, *6*, 697–710. [CrossRef]

31. Khalafalla, M.M.; Abdellatif, E.; Daffalla, H.M.; Nassrallah, A.A.; Aboul-Enein, K.M.; Lightfoot, D.A.; El-Deeb, F.E.; El-Shemy, H.A. Active principle from Moringa oleifera Lam leaves effective against two leukemias and a hepatocarcinoma. *Afr. J. Biotechnol.* **2010**, *9*, 8467–8471. [CrossRef]

32. Das, N.; Sikder, K.; Ghosh, S.; Fromenty, B.; Dey, S. Moringa oleifera lam. leaf extract prevents early liver injury and restores antioxidant status in mice fed high fat diet. *Indian J. Exp. Biol.* **2012**, *50*, 404–412.

33. Almatrafi, M.M.; Vergara-Jimenez, M.; Smyth, J.A.; Medina-Vera, I.; Fernandez, M.L. Moringa oleifera leaves do not alter adipose tissue cholesterol accumulation or inflammation in guinea pigs fed a hypercholesterolemic diet. *EC Nutr.* **2017**, *18*, 1330.

34. Halaby, M.S.; Elmetwally, E.M.; Omar, A.A.A. Effect of Moringa oleifera on serum lipids and kidney function of hyperlipidemic rats. *J. Appl. Sci. Res.* **2013**, *9*, 5189–5198.

35. Almatrafi, M.M.; Vergara-Jimenez, M.; Murillo, A.G.; Norris, G.H.; Blesso, C.N.; Fernandez, M.L. Moringa leaves prevent hepatic lipid accumulation and inflammation in guinea pigs by reducing the expression of genes involved in lipid metabolism. *Int. J. Mol. Sci.* **2017**, *18*, 1330. [CrossRef] [PubMed]

36. Richter, N.; Siddhuraju, P.; Becker, K. Evaluation of nutritional quality of moringa (Moringa oleifera Lam.) leaves as an alternative protein source for Nile tilapia (Oreochromis niloticus L.). *Aquaculture* **2003**, *217*, 599–611. [CrossRef]

37. Bais, S.; Singh, G.S.; Sharma, R. Antiobesity and Hypolipidemic Activity of Moringa oleifera Leaves against High Fat Diet-Induced Obesity in Rats. *Adv. Biol. 2014*, *9*, 457–466. [CrossRef]

38. Panda, S.; Kar, A.; Sharma, P.; Sharma, A. Cardioprotective potential of N-α-l-rhamnopyranosyl vincosamide, an indole alkaloid, isolated from the leaves of Moringa oleifera in isoproterenol induced cardiotoxic rats: In Vivo and in vitro studies. *Bioorg. Med. Chem. Lett. 2013*, *23*, 959–962. [CrossRef] [PubMed]

39. Dangi, S.Y.; Jolly, C.I.; Narayanan, S. Antihypertensive activity of the total alkaloids from the leaves of Moringa oleifera. *Pharm. Biol.* **2002**, *40*, 144–148. [CrossRef] [PubMed]

40. Tiloke, C.; Anand, K.; Gengan, R.M.; Chuturgeroon, A.A. Moringa oleifera and their phytonanoparticles: Potential antiproliferative agents against cancer. *Biomol. Pharmaother. 2014*, *9*, 457–466. [CrossRef]

41. Wahyuni, R.; Wignyanto, W.; Wijana, S.; Sucipto, S. Optimization of protein and tannin extraction in Moringa oleifera leaf as protein source for Nile tilapia (Oreochromis niloticus L.). *J. Funct. Foods** *2013*, *5*, 1883–1891. [CrossRef]

42. Adisakwattana, S.; Chanathong, B. A-Glucosidase Inhibitory Activity and Lipid-Lowering. *Eur. Rev. Med. Pharmacol. Sci. 2011*, *15*, 803–808.

43. Arauto, I.C.C.; Aguiar, J.S.; Napoleão, T.H.; Mota, F.V.B.; Barros, A.L.S.; Moura, M.C.; Coriolano, M.C.; Coelho, L.C.B.B.; Silva, T.G.; Paiva, P.M.G. Evaluation of cytotoxic and anti-inflammatory activities of extracts and lectins from Moringa oleifera seeds. *PLoS ONE* **2013**, *8*, e81973. [CrossRef] [PubMed]

44. Xiong, Y.; Riaz Rajoka, M.S.; Zhang, M.X.; He, Z. Isolation and identification of two new compounds from the seeds of Moringa oleifera and their antiviral and anti-inflammatory activities. *Nat. Prod. Res. 2020*, *1–9*. [CrossRef]

45. Govardhan Singh, R.S.; Negi, P.S.; Radha, C. Phenolic composition, antioxidant and antimicrobial activities of free and bound phenolic extracts of Moringa oleifera seed flour. *J. Funct. Foods** *2013*, *5*, 1883–1891. [CrossRef]

46. Liao, P.C.; Lai, M.H.; Hsu, K.P.; Kuo, Y.H.; Chen, J.; Tsai, M.C.; Li, C.X.; Yin, X.J.; Jeyashoke, N.; Chao, L.K.P. Identification of β-Sitosterol as in Vitro Anti-Inflammatory Constituent in Moringa oleifera. *J. Agric. Food Chem. 2020*, *68*, 10748–10759. [CrossRef] [PubMed]

47. Abd Rani, N.Z.; Kumolosasi, E.; Jasamai, M.; Jamal, J.A.; Lam, K.W.; Blesso, C.N.; Fernandez, M.L. Moringa oleifera leaves do not alter adipose tissue cholesterol accumulation or inflammation in guinea pigs fed a hypercholesterolemic diet. *EC Nutr.* **2017**, *18*, 1330.

48. Wolff, K.; Jaja-Chimedza, A.; Kim, Y.; Waterman, C.; Pouley, A.; Raskin, I.; Ribnicky, D. Moringa isothiocyanate-1 is bioaccessible and bioavailable as a stable unmodified compound. *Phytochem. Lett.* **2020**, *38*, 33–38. [CrossRef]

49. Almatrafi, M.M.; Vergara-Jimenez, M.; Murillo, A.G.; Norris, G.H.; Blesso, C.N.; Fernandez, M.L. Moringa leaves prevent hepatic lipid accumulation and inflammation in guinea pigs by reducing the expression of genes involved in lipid metabolism. *Int. J. Mol. Sci.* **2017**, *18*, 1330. [CrossRef] [PubMed]

50. Almatrafi, M.M.; Vergara-Jimenez, M.; Murillo, A.G.; Norris, G.H.; Blesso, C.N.; Fernandez, M.L. Moringa oleifera leaves do not alter adipose tissue cholesterol accumulation or inflammation in guinea pigs fed a hypercholesterolemic diet. *EC Nutr.* **2017**, *18*, 1330.