



Three Selected Edible Crops of the Genus *Momordica* as Potential Sources of Phytochemicals: Biochemical, Nutritional, and Medicinal Values

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Momordica species (Family Cucurbitaceae) are cultivated throughout the world for their edible fruits, leaves, shoots and seeds. Among the species of the genus *Momordica*, there are three selected species that are used as vegetable, and for medicinal purposes, *Momordica charantia* L (Bitter melon), *Momordica foetida* Schumacher (Bitter cucumber) and *Momordica balsamina* L (African pumpkin). The fruits and leaves of these *Momordica* species are rich in primary and secondary metabolites such as proteins, fibers, minerals (calcium, iron, magnesium, zinc), β -carotene, foliate, ascorbic acid, among others. The extracts from *Momordica* species are used for the treatment of a variety of diseases and ailments in traditional medicine. *Momordica* species extracts are reputed to possess anti-diabetic, anti-microbial, anthelmintic bioactivity, abortifacient, anti-bacterial, anti-viral, and play chemo-preventive functions. In this review we summarize the biochemical, nutritional, and medicinal values of three *Momordica* species (*M. charantia*, *M. foetida* and *M. balsamina*) as promising and innovative sources of natural bioactive compounds for future pharmaceutical usage.

Keywords: momordica, anti-microbial, photochemical, secondary metabolites, inter alia

INTRODUCTION

Momordica charantia L. (Bitter melon), *Momordica foetida* Schumach. (Bitter cucumber), *Momordica balsamina* L. (African pumpkin) are widely cultivated and they belong to the family Cucurbitaceae (Daniel et al., 2014). In some African countries; fruits, leafy shoots and the ripe seeds from these three species are harvested for consumption as vegetable, spices and leaves are mixed with water and consumed as beverages (Cantwell et al., 1996). *Momordica* species (spp.) are annual crops but can be regarded as perennial crop due to their performance during the season (Krawinkel and Keding, 2006; Tan et al., 2008; Costa et al., 2010; Chen et al., 2011; Abegunde et al., 2018). These crop species are mostly adapted to areas of minimum annual average rainfall such as those that receive an average of 400 mm as well as those with mild and frost-free winters (Tokhtar and Doan, 2014). *Momordica* spp. require a minimum temperature of 18°C during emergence and germination and during vegetative stage and flowering and an optimal temperature ranging between 24 and 27°C (Tan et al., 2008). Some researchers have stated that the crops adapt and perform well in deep and well-drained sandy loam and slit loam soils which are rich in organic matter and adapt well in soils with optimum soil pH ranging from 4.3 to 8.0 (Tan C. et al., 2015). However, they can adapt to alkaline soils with up to pH of 8.0 (Grover and Yadav, 2004; Harinantenaina et al., 2006; Tan et al., 2008). *Momordica* spp. are open-pollinated and consist of separate male and female flowers on the same plant (Palada and Chang, 2003). Flowering of *Momordica* species begins a month after planting until reaching dormancy. The fruits are orange in color and soft when ripe with black seeds (Bakare et al., 2010). The local South African indigenous traditional people have been using *Momordica* spp. as culinary species or enhance bitterness of the meat and of vegetables (Madala et al., 2014). However, studies on the trend of food consumption for rural residents in developing countries have often focused relatively on the economic factors such as prices, income, and market development Jia et al. (2017), and have rarely covered social and psychological influences.

Normally, *Momordica* spp. are consumed as vegetables and as nutrient supplements in Asia, Africa and South America and they are also used as medicinal herbs (Tan et al., 2008; Madala et al., 2014; Jia et al., 2017; Islam and Jalaluddin, 2019). Different parts of *Momordica* spp. are rich in both primary and secondary metabolites. The primary metabolites that are derived from *Momordica* spp. include sugars, proteins and chlorophyll (Hassan and Umar, 2006; Daniel et al., 2014), while secondary metabolites are alkaloids, flavonoids, tannins (Roškar and Lušin, 2012; Mada et al., 2013; Madala et al., 2016; Mostafa et al., 2018). *Momordica* spp. are good a source of phenolic compounds, gallic acid, gentisic acid (2,5-dihydroxyl benzoic acid), catechins, chlorogenic acid and epicatechin (Abegunde et al., 2018).

Generally, extracts from *Momordica* spp. are useful for the treatment of various diseases and many other ailments due to the several biological activities exerted such as anti-diabetic Tan et al. (2008), Madala et al. (2014), Jia et al. (2017), Islam and Jalaluddin (2019), Perera et al. (2021), Venugopal and Dhanasekaran (2021), anti-oxidant Acquaviva et al. (2013), Ozusaglam and Karakoca

(2013), Huang et al. (2020), Perumal et al. (2021), anti-microbial (Kumar et al. (2009), Patel et al. (2010), Gupta et al. (2011), anti-bacterial (Koneri et al. (2006), Orhan et al. (2010), Leelaprakash et al. (2011), Puškárová et al. (2017), Chen et al. (2019), Islam and Jalaluddin (2019), Khadijeh et al. (2019), anti-viral Orhan et al. (2010), Santhosh Kumar et al. (2020), anti-fungal Santos et al. (2012), anthelmintic Beloin et al. (2005), Jia et al. (2017), insecticidal Pari et al. (2020), antiglycation Chen et al. (2011), anti-inflammatory Chen et al. (2019), Khadijeh et al. (2019), Perera et al. (2021), Venugopal and Dhanasekaran (2021), anti-thrombotic Patil and Rathod (2020), anti-allergic Asna et al. (2020), hepatoprotective Ramalhete et al. (2011), estrogenic Badhwar et al. (2020), hypolipidemic effect Aeri and Raj (2020), anti-carcinogenic Kesari et al. (2020), anti-proliferative Yue et al. (2020), anti-mutagenic Rolnik and Olas (2020), and chemo-preventive activities (Gupta, 2020).

Momordica spp. have been reported to have inhibitory effects against diabetes, cancer, and cardiovascular diseases in animal and humans (Trakoon-osot et al., 2013; Fongod et al., 2014; Ingle et al., 2017; Jabeen and Khanum, 2017; Perera et al., 2021). Some studies shows that the bioactivity effectiveness in lowering of blood sugar (Huang et al., 2008; Rahmatullah et al., 2012; Balouiri et al., 2016; Anjamma and Bhavani, 2018), in controlling eye disorders and enhancing eyesight due to the presence of β -carotene (Ludidi et al., 2019). In addition, some studies have suggested that *Momordica* spp. are used to treat gastrointestinal parasitic disease (Grover and Yadav, 2004), diarrhea, bleeding of gums Abidemi (2013), piles and hemorrhoids Lii et al. (2009), respiratory problems Leung et al. (2009) and skin infections (Kumar and Bhowmik, 2010). The anti-viral and anti-leukemic activities of extract from *M. balsamina* against liver cancer, leukemia, melanoma and solid sarcomas have also been reported (Lu et al., 2011; Aparicio et al., 2021).

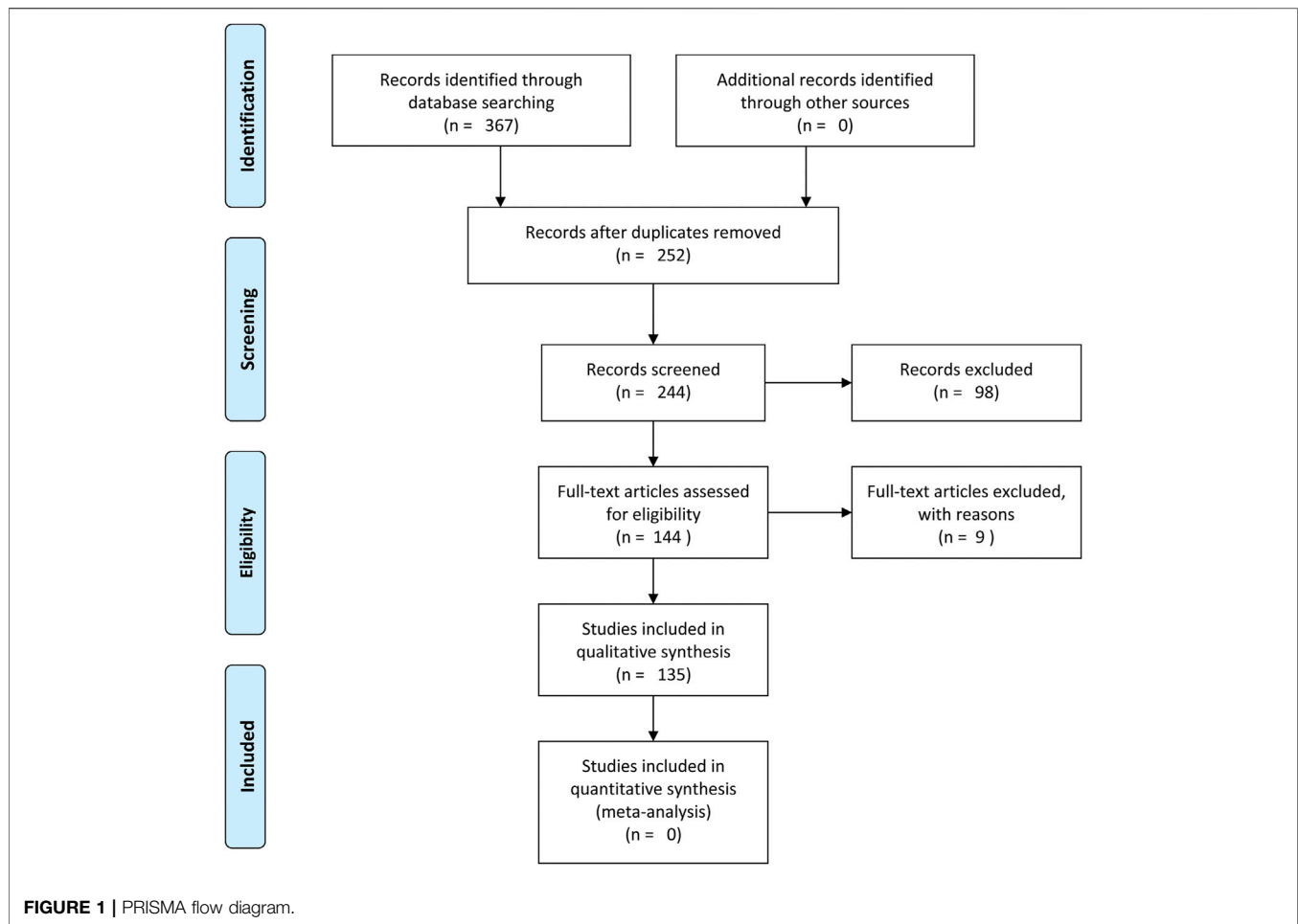
Momordica spp. have flavonoids with anti-microbial activities (Orhan et al., 2010) and *Momordica* leaves extracts have shown anti-bacterial activities against *Staphylococcus aureus*, *Salmonella typhii*, *Escherichia coli*, *Klebsiella pneumoniae*, *Bacillus subtilis*, *Pseudomonas* sp. and *Entamoeba histolytica* (Tansey et al., 2004; Froelich et al., 2007; Ghosh, 2014; Balouiri et al., 2016) and the fruit extract possesses anti-microbial activity against *Helicobacter pylori* (Mwambete, 2009).

The use of the plant extracts of *Momordica* spp. to fully explored their effectiveness, may lead to new drug discovery or advance the use of indigenous herbal medicines for orthodox treatment of certain diseases (Rahmatullah et al., 2012). The aim of this review was to discuss the biochemical, nutritional, and medicinal values of three *Momordica* spp. (*M. charantia*, *M. foetida* and *M. balsamina*) as promising and innovative sources of natural bioactive compounds for future pharmaceutical usage.

METHODOLOGY

Type of Review

The systematic review method involved six steps: 1) scoping; 2) planning; 3) search process and identification of articles; 4) Screening of articles; 5) assessment of relevancy and 6)



Presentation and interpretation of the results. As a rigorous and unbiased method of evaluating and screening the literature attributed to its replicability and exhaustiveness, it is a method of choice suitable for summarizing the current literature on the three *Momordica* spp (*M. charantia*, *M. foetida* and *M. balsamina*).

Literature Sources and Searching Procedure

The search was conducted from six electronic database (Web of Science, Science.gov, Science Direct, Google Scholar, Scopus and Ebscohost) using the following guidelines: 1) “*Momordica charantia*” OR “Bitter melon”, 2) “*Momordica. Foetida*” or “Bitter cucumber”, and 3) “*Momordica. Balsamina*” OR “African pumpkin”. The common search for peer reviewed article used included the biochemical, nutritional, and medicinal values or composition of *Momordica* spp. The same search was also conducted using the common names of the species. The search was filtered based on the year of publications (Year = 1990–2020) to obtain most recent literature however, it did not exclude the old publications (Year < 1990) which form the basic knowledge of the associated

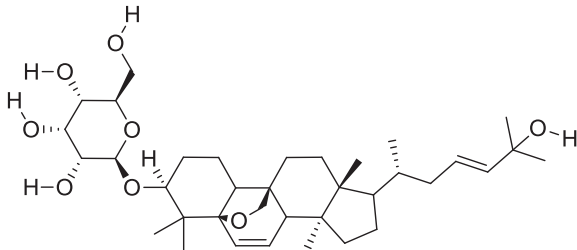
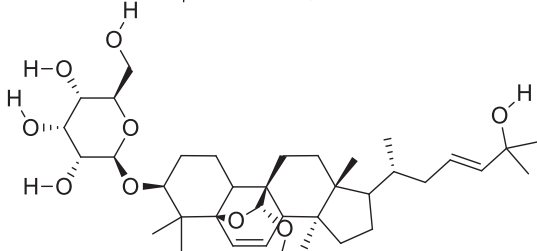
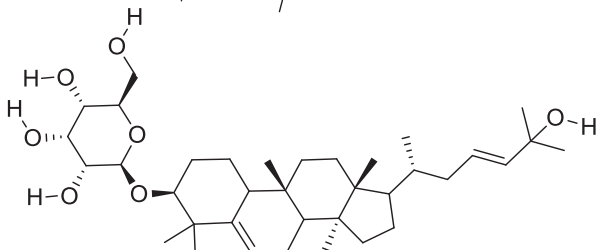
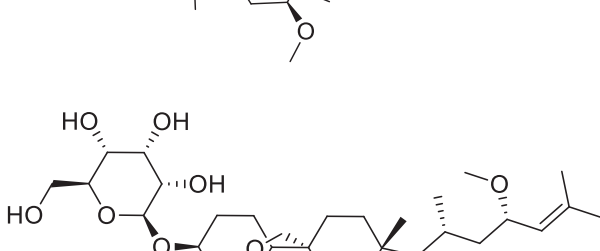
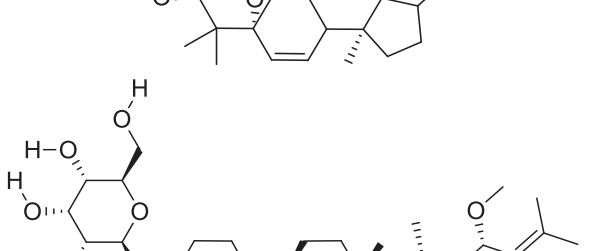
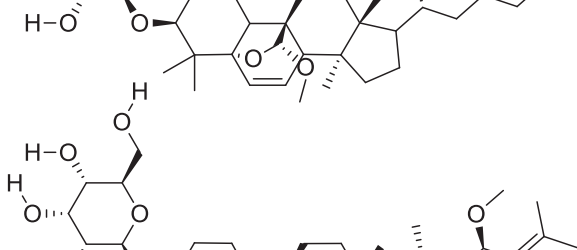
literature. Since, the study was limited to qualitative synthesis, the PRISMA 2009 Flow Diagram (Moher et al., 2009) was to modified to exclude the meta-analysis aspects (Figure 1).

BIOCHEMICAL COMPOSITION AND ACTIVITY OF THE THREE SELECTED *MOMORDICA* SPECIES

Momordica charantia L.

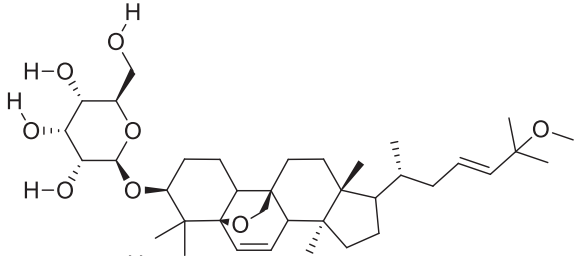
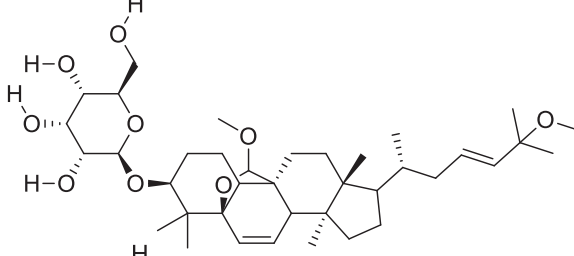
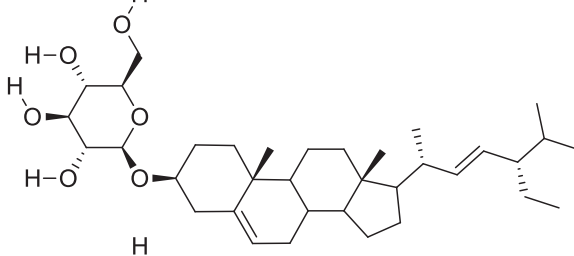
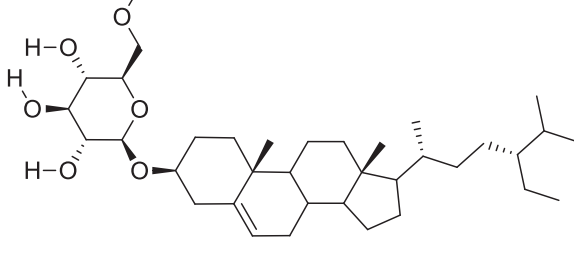
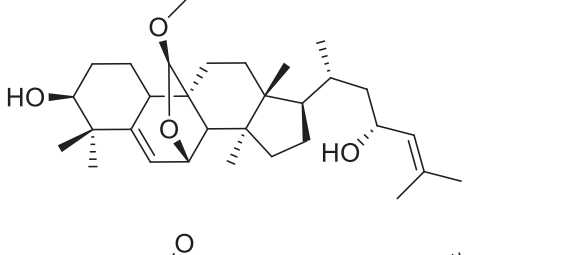
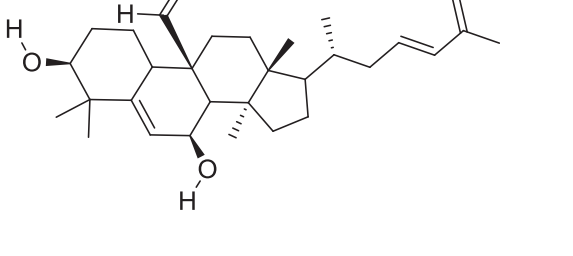
M. charantia is a good source of primary metabolites such as carbohydrates, fibers, proteins, minerals, and vitamins. The most important chemical constituents from *M. charantia* include heteropolysaccharides (e.g., arabinose, galactose, glucose, mannose and rhamnose); proteins and peptides (e.g., momordins, momorcharins, etc); terpenoids and saponins (e.g., cucurbitanes and cucurbitacines); flavonoids and phenolic compounds (Chen et al., 2003; Harinantenaina et al., 2006; Schrot et al., 2015; Dandawate et al., 2016; Lee et al., 2021). *M. charantia* also contains triterpenes, proteins, steroids, alkaloids, lipids (lauric, myriaatic, palmitic, stearic, linoleic, and eleostearic acids), phenolic compounds and minerals (Cu,

TABLE 1 | Compounds extracted from *Momordica charantia* L. and their health benefits.

Compound	Structure	Health benefits	References
Momordicoside F ₂		Anti-inflammatory and anti-cancer	Ju et al. (2004), Cao et al. (2018)
Goyaglycoside B		Anti-cancer	Raina et al. (2016), Cao et al. (2018)
Karaviloside III		Anti-aging	Cao et al. (2018)
Charantoside VI		Anti-aging	Cao et al. (2018)
Charantagenin E		Anti-aging	Cao et al. (2018)
Charantoside II		Anti-aging	Cao et al. (2018)

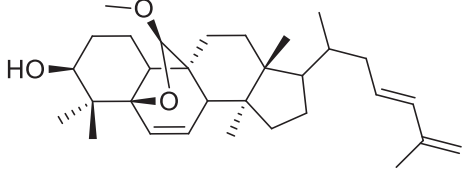
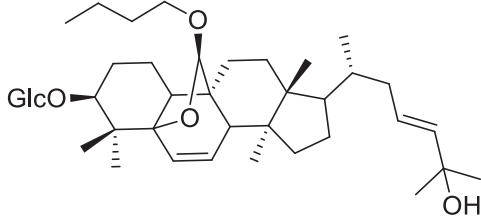
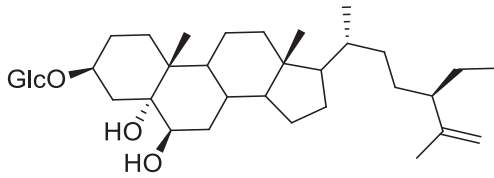
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TABLE 1 | (Continued) Compounds extracted from *Momordica charantia* L. and their health benefits.

Compound	Structure	Health benefits	References
Momordicoside G		Anti-aging	Cao et al. (2018)
Goyaglycoside D		Anti-cancer	Wang et al. (2012), Cao et al. (2018)
Stigmasterol glucoside		Anti-angiogenic and anti-cancer	Desai and Tatke, (2015), Kangsamaksin et al. (2017)
β -sitosterol glucoside		Cholesterol reduction and immune system modulation	Kaur et al. (2011), Desai and Tatke, (2015)
(19R)-7 β ,19-epoxy-19-methoxycucurbita-5,24-dien-3 β ,23-diol		Anti-aging	Jiang et al. (2016), Cao et al. (2018)
Kuguacin J		Anti-cancer	Limtrakul et al. (2013), Chen et al. (2015)

(Continued on following page)

TABLE 1 | (Continued) Compounds extracted from *Momordica charantia* L. and their health benefits.

Compound	Structure	Health benefits	References
Cucurbitane triterpenoid		Prevention and therapy of human breast cancer	Liu et al. (2009), Bishayee et al. (2011)
Cucurbitane triterpenoid		Anti-cancer	Liu et al. (2009), Bishayee et al. (2011)
Steroidal glycoside		Reduce blood sugar level	Liu et al. (2009), Tan et al. (2015b), Lee et al. (2021)

Fe, Mg, Zn, and Ca) (Yuwai et al., 1991; Kumar and Bhowmik, 2010; Joseph and Jini, 2013; Tokhtar and Doan, 2014; Tan E. S. et al., 2015; Yoshime et al., 2016). The leaves and flowers of *M. charantia* contains triterpenoids (momordicine and charantin), carotenoids (antheraxanthin, lutein, violaxanthin, α -carotene, and β -carotene), and phenylpropanoids (caffeic acid, chlorogenic acid, epicatechin, gallic acid, p-coumaric acid, rutin, and *trans*-cinnamic acid) (Sathasivam et al., 2021). The water content of the fruits of *M. charantia* is about 93.2%, while protein and lipids content ranges between 18.02 and 0.76% (Saad et al., 2017). The fruit pulp of *M. charantia* consists of soluble pectin and galactouronic acid. Moreover, the fruits consist of glycosides, saponins, alkaloids, reducing sugars, resins, phenolic constituents, fixed oil and free acids (Bakare et al., 2010). Green fruits of *M. charantia* contain Vitamin A, Vitamin C, and Vitamin *p*, thiamine, riboflavin, niacin, and minerals (Gupta et al., 2011). The essential oil, obtained from water stressed seeds also contains sesquiterpenes, phenylpropanoids and monoterpenes, however seed oil contain tocopherols and polyphenols (Nyam et al., 2009). The phenolic compounds that are found on all parts of the crop protect the plant from oxidative damage (Yoshime et al., 2016; Wardani et al., 2021).

Among the bioactive compounds that have been isolated from the fruits of *M. charantia* and elucidated by nuclear magnetic resonance (NMR) and mass spectrometry (MS) spectroscopic techniques include Monordicophenoide A (4-hydroxyl-benzoic acid 4-*O*- β -D-apiofuranosyl (1 \rightarrow 2)-*O*- β -D-glucopyranoside;

dihydrophaseic acid 3-*O*- β -D-glucopyranoside; momordicolide (10*E*)-3-hydroxyl-dodeca-10-en-9-olide; 6,9-dihydroxy-megastigman-4,7-dien-3-one (blumenol); adenosine; guanosine; uracil; and cytosine (Li et al., 2009). Those isolated from the leaves of *M. charantia* include Momordicin I, Momordicin IV, aglycone of Momordicoside, aglycone of Momordicoside L; and Karavilagenin D were identified (Li et al., 2015). The essential oil such as *trans*-nerolidol, apiole, *cis*-dihydrocarveol, and germacrene D were isolated from the seeds of *M. charantia* and identified by gas chromatography mass spectrometry (GC/MS) (Braca et al., 2008).

Main phenolic acids found in *M. charantia* are gallic acid, chlorogenic acid, catechin, caffeic acid, p-coumaric acid, and ferulic acid (Lee et al., 2017). Compounds such as capxanthin, lutein, zeaxanthin, β -cryptoxanthin, lycopene, α -carotene, and β -carotene have also detected from the fruit of *M. charantia* (Lee et al., 2017). The variation in phenolic acids and carotenoids contents at different maturity stages create a strong relationship with the anti-oxidant activities of *M. charantia* fruit. The crop consists of non-essential amino acids such as arginine, alanine, aspartic acid, glycine, glutamic acid, proline, histidine and serine. Such non-essential amino acids are inherently higher in concentration, as compared to essential amino acids such as leucine, cysteine, phenylalanine, isoleucine, methionine, lysine, threonine, tyrosine and valine. From the leaves of *M. charantia* leucine and aromatic (tyrosine and phenylalanine) were the predominant amino acids, while aspartic acid and glutamic

acid are major non-essential amino acids of this plant (Khanna, 2004).

Other reported phytochemical compounds from *M. charantia* plant were obtained after being extracted and their chemical structures identified using NMR. These include β -sitosterol; (23E)-5 β ,19-epoxycucurbita-6,23-diene-3 β ,25-diol; daucosterol; uracil; and allantoin using spectroscopic analysis including MS and 1H- and C-NMR (Kim et al., 2018). The structures of momordicosides K were identified as 7-O- β -D-glucopyranosides of 3 β ,7 β -dihydroxy-25-methoxy-cucurbita-5; 23-dien-19-al; and momordicosides L as 3 β ,7 β ,25-trihydroxy-cucurbita-5,23-dien-19-al (Okabe et al., 1982).

***Momordica foetida* Schumach**

M. foetida have been reported to contain secondary metabolites such as sitosterylglucoside, 5,25-stigmastadien-3 β -ylglucoside, and 1 β -hydroxyfriedel-6-en-3-one, and a lot of cucurbitane-type triterpenoid derivatives (Molehin and Adefegha, 2014). Phenolic glycosides of *M. foetida* include eriodictyol-, kaempferol- 5,7,4'-trihydroxyflavanone, and 5,7-dihydroxychromone-7-O- β -D-glucopyranoside (Froelich et al., 2007).

The crop species of *M. foetida* contain 25-trihydroxycucurbita-5,23-dien-19-al and 3 β ,7 β -dihydroxy-25 methoxycucurbita-5,23-dien-19-al; 3 β , 7 β , 23 ξ -trihydroxycucurbita-5,24-dien-19-al, 3 β , 7 β . Others include 5 β ,19-epoxy-25-methoxycucurbita-6,23-diene-3 β ,19-diol,5 β ,19-epoxycucurbita-6,23-diene-3 β ,19,25-triol,5 β ,19-epoxy-19 methoxycucurbita-6, 23-diene-3 β ,25-diol,5 β ,19-epoxy-19,25-dimethoxycucurbita-6,23-dien-3 β -ol and 5 β ,19-epoxy-25-methoxy-cucurbita-6,23-dien-3 β -ol (Mulholland et al., 1997). Secondary metabolites detected in *M. foetida* are steroids, alkaloids, cardiac glycosides, phenolics, tannins, flavonoids and saponin (Ndam et al., 2014).

The chemical structures of the isolated compounds include 3 β ,7 β -dihydroxyl-cucurbita-5,23,25-trien-19-al followed by Kaempferol-3-O- β -D-glucopyranoside have anti-microbial activity (Odeleye et al., 2009). The fruits and leaves of *M. foetida* crop contain Momordicines and Foetidins (identical to Charantin). Phytochemical analysis of *M. foetida* revealed phenolic glycosides such as eriodictyol-, 5,7,4'-trihydroxyflavanone-, kaempferol- and 5,7-dihydroxychromone-7-O- β -D-glucopyranoside (Froelich et al., 2007). Other phytochemical analysis of *M. foetida* extract led to the presence of alkaloids, saponins and cardenolides (Oduote and Awaraka, 2004).

***Momordica balsamina* L.**

The leaves, fruits, seeds, and bark of the *M. balsamina* contains secondary metabolites such as alkaloids, flavonoids, glycosides, steroids, terpenes, cardiac glycoside, saponins, tannins and lectins (Talukdar and Hossain, 2014). The matured seeds are the main source of plant lectins (Abegunde et al., 2018). The primary metabolites include crude lipid (2.66%), common sugars, proteins (11.29%), crude fiber (29%), energy, amino acid, carbohydrates (39%) and chlorophyll (Hassan and Umar, 2006; Thakur et al., 2009; Orhan et al., 2010; Ibrahim et al., 2014; Birla, 2016; Abegunde et al., 2018).

The leaves are an important source of nutrients, in addition they contain 17 amino acids and an adequate mineral

composition (Hassan and Umar, 2006; Talukdar and Hossain, 2014; Wardhani et al., 2015; Wang et al., 2016; Nitu and Patidar, 2017). *M. balsamina* is rich potassium, calcium, magnesium, sodium, phosphorus, manganese, zinc, and iron which contribute to prevent micronutrient deficiencies in human.

NUTRITIONAL VALUE OF THE THREE SELECTED *MOMORDICA* SPECIES

***Momordica charantia* L.**

M. charantia fruit is commonly eaten unripe as a vegetable and is also used as herbal plant that is useful in metabolic and physiological processes of the human body. It is advisable to add peeled fruits to minimize bitter taste before preparation which involves soaking the fruits and shoots and then boiled, fried or pickled (Kumar and Bhowmik, 2010). *M. charantia* fruits are known to have high levels of nutritional value (protein, vitamins, thiamine, riboflavin, calcium and iron) compared to other cucurbits it has higher in folate and Vitamin C and the vine tips as an excellent source of Vitamin A. *M. charantia* leaves can be harvested, cooked and consumed alone or mixed with other indigenous vegetables to limit the bitterness (Hassan and Umar, 2006). The leaves and fruits of *M. charantia* have been used to make teas and beer or season soups in the Western world (Kumar and Bhowmik, 2010).

M. charantia contains iron, β -carotene, calcium potassium, vitamins, phosphorus and good dietary fiber (Daniel et al., 2014). About 93.2% of the fruit of *M. charantia* is composed of water, while protein and lipids account for 18.02 and 0.76%, respectively, (Li et al., 2009). Leaves powder from *M. charantia* contained 38% water, 38.2% dietary fiber, 67% minerals, 45% crude protein and 27% crude fat (Anilakumar et al., 2015). Protein lectin that is found on the seed of *M. charantia* is responsible for the inhibition of protein synthesis in the intestinal walls of an animal without producing any gastrointestinal symptoms in humans.

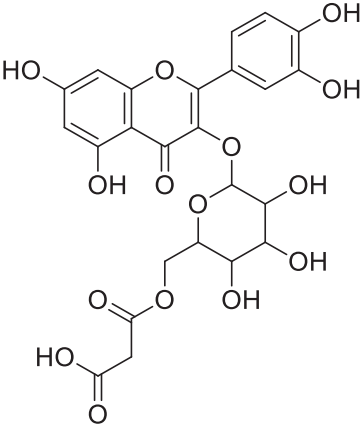
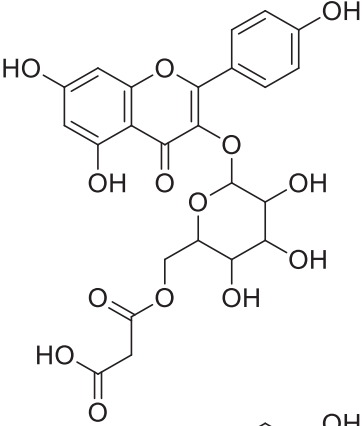
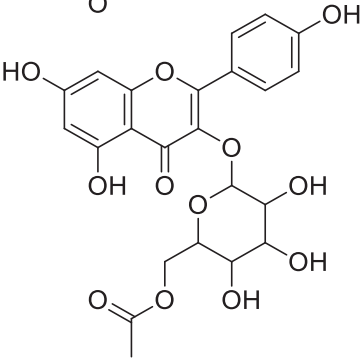
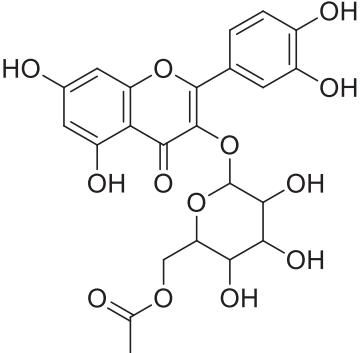
M. charantia seeds may induce favism in humans with glucose-6 phosphate dehydrogenase deficiency (Harinantenaina et al., 2006; Krawinkel and Keding, 2006; Ludidi et al., 2019). Some studies reported that *M. charantia* leaf powder inhibited adipocyte hypertrophy in diet-induced obese rats due to the process of decreasing lipogenic genes including fatty acid synthase, acetyl CoA carboxylase, lipoprotein lipase and adipocyte fatty acid and protein in epididymis fat (Tansey et al., 2004; Nerurkar et al., 2010; Anilakumar et al., 2015).

M. charantia reduced the accumulation of visceral fat that was fed to rats with high fat diet it and thus help in reducing insulin sensitizing and glucose due to its anti-adiposity effect (Chen et al., 2003; Huang et al., 2008; Dellavalle et al., 2011). Circulating levels of catecholamine and non-esterified fatty acids, lipid oxidation in the liver and muscle are reported to be as a result of *M. charantia* supplemented rat (Cohen et al., 1998; Waako et al., 2005; Harinantenaina et al., 2006; Bulbul, 2016; Devyani et al., 2016).

***Momordica foetida* Schumach**

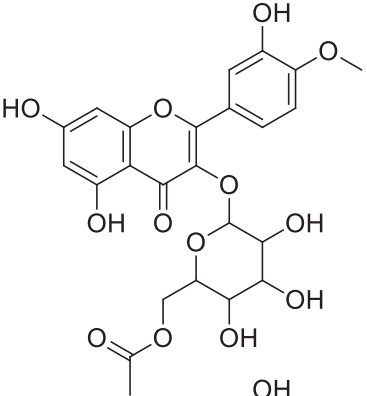
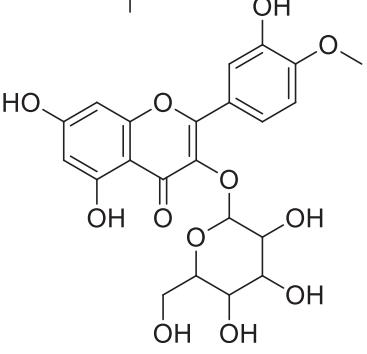
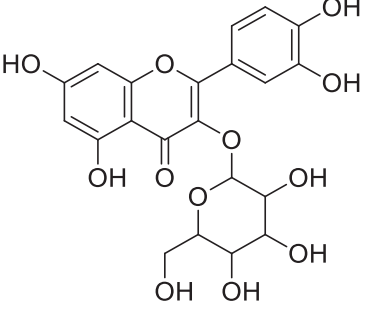
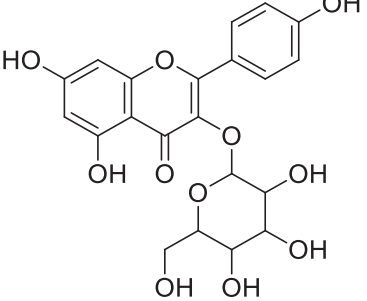
The crop is commercially grown for their nutritional value and they are also used for medicinal purposes (Waako et al., 2005).

TABLE 2 | Compounds extracted from *Momordica foetida* Schumach. and their health benefits.

Compound	Structure	Health benefits	References
Quercetin malonyl-glycoside		Lowers blood pressure and inflammation	Khoza et al. (2016), Dabeek and Marra, (2019)
Kaempferol malonyl-diglycoside		Anti-tumor, anti-oxidant and anti-inflammatory	Khoza et al. (2016), Abegunde et al. (2018)
Kaempferol Acetyl-glycoside		Anti-tumor, anti-oxidant and anti-inflammatory	Khoza et al. (2016), Abegunde et al. (2018)
Quercetin acetyl-glycoside		Lowers blood pressure and inflammation	Khoza et al. (2016), Dabeek and Marra, (2019)

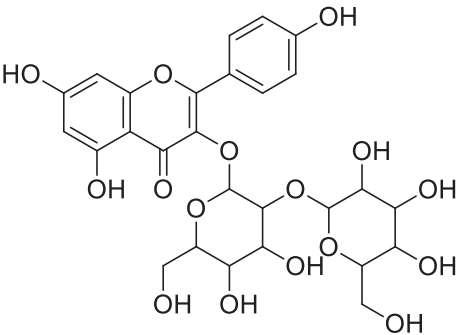
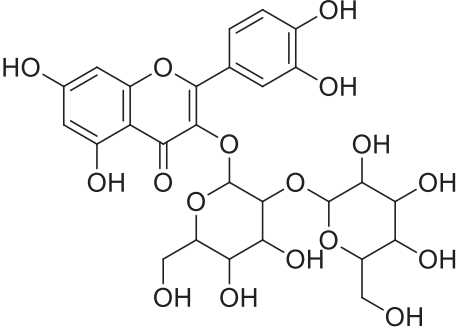
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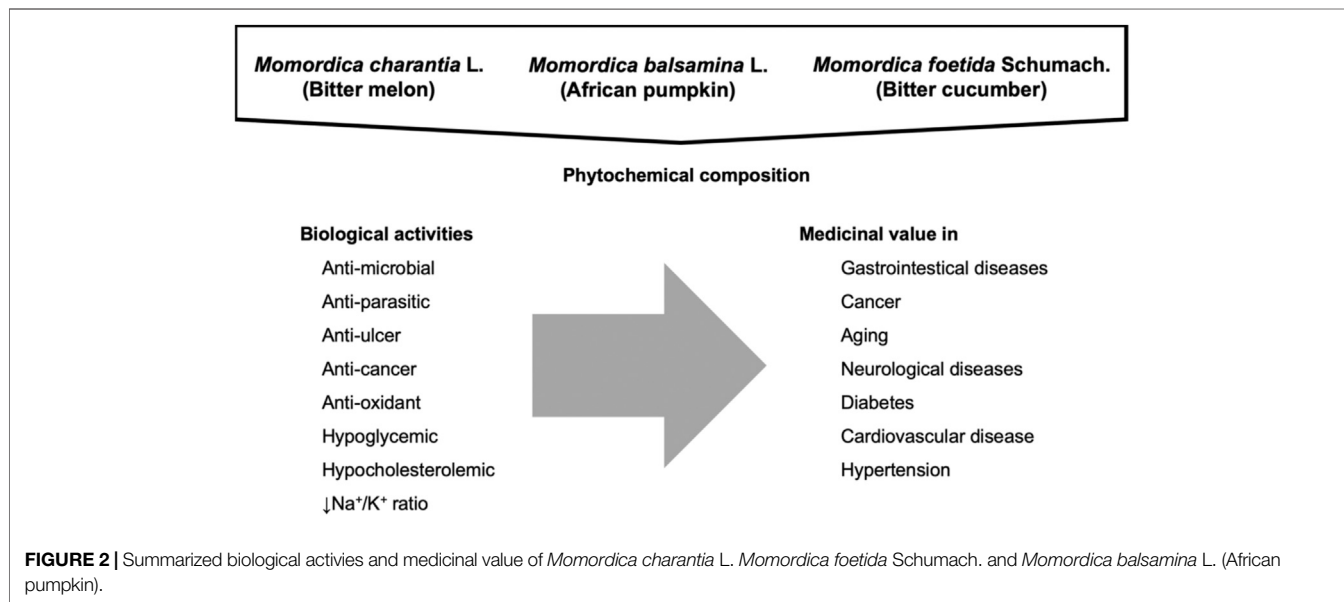
TABLE 2 | (Continued) Compounds extracted from *Momordica foetida* Schumach. and their health benefits.

Compound	Structure	Health benefits	References
Isorhamnetin acetyl-glycoside		Anti-oxidant, anti-viral, anticancer, anti-microbial, and anti-inflammatory	Khoza et al. (2016), Kandakumar and Manju, (2017)
Isorhamnetin glycoside		Anti-oxidant, anti-viral, anticancer, anti-microbial, and anti-inflammatory	Khoza et al. (2016), Kandakumar and Manju, (2017)
Quercetin glycoside		Lowers blood pressure and inflammation	Khoza et al., (2016), Dabeek and Marra, (2019)
Kaempferol glycoside		Anti-tumor, anti-oxidant and anti-inflammatory	Khoza et al. (2016), Abegunde et al. (2018)

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TABLE 2 | (Continued) Compounds extracted from *Momordica foetida* Schumach. and their health benefits.

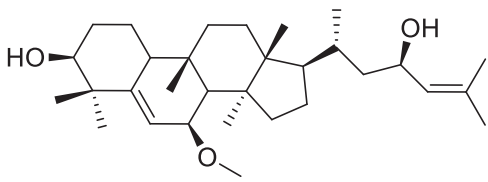
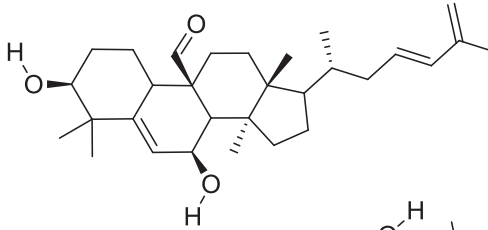
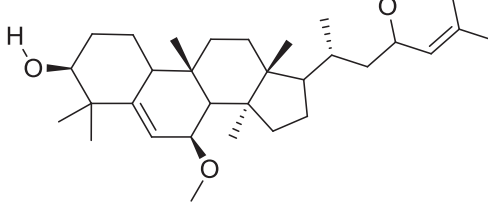
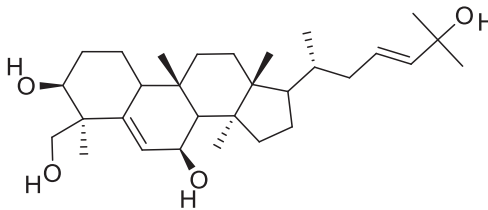
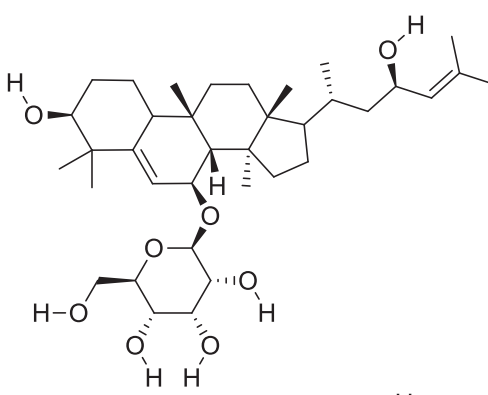
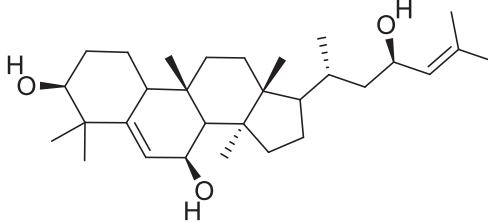
Compound	Structure	Health benefits	References
Kaempferol diglycoside		Anti-tumor, anti-oxidant and anti-inflammatory	Khoza et al. (2016), Abegunde et al. (2018)
Quercetin diglycoside		Lowers blood pressure and inflammation	Khoza et al. (2016), Dabeek and Marra, (2019)



The leaves of *M. foetida* are harvested from the wild and consumed when they are prepared as a vegetable in Gabon, Sudan, Uganda, Tanzania, South Africa and Malawi and some other African countries. The leaves of *M. foetida* are harvested, boiled and consumed with beans or peas together with a staple food. It is utilized in small amounts as a famine food and in emergency situations.

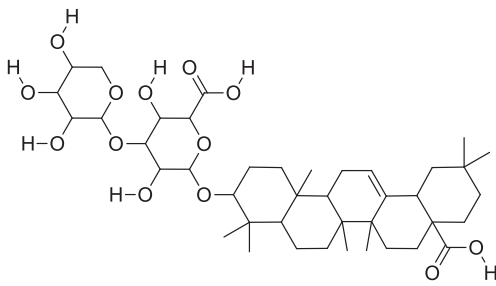
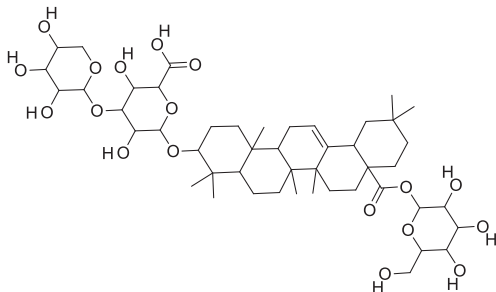
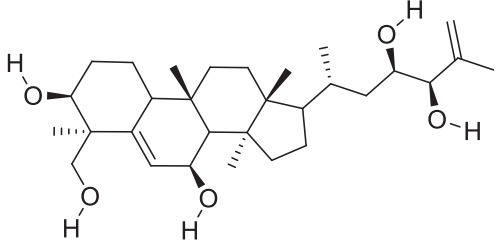
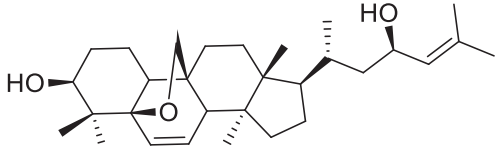
Traditional people prefer to consume *M. foetida* pulp of ripe fruits even if it has bitter taste (Mada et al., 2013; Mostafa et al., 2018; Ludidi et al., 2019). *M. foetida* leaves can be mixed with other vegetables to make a stew. It is also sauced with shrimp, meat, pork, and chicken and served with gravy. The leaves are mixed with ground peanuts and honey as a sauce in meat (Thakur et al., 2009; Dellavalle et al., 2011; Oranje et al., 2012; Sen and

TABLE 3 | Compounds extracted from *Momordica balsamina* L. and their health benefits.

Compound	Structure	Health benefits	References
Cucurbitane glycoside		Anti-diabetic	Spengler et al. (2009), Farooq et al. (2019)
Kuguacin J		Anti-cancer	Spengler et al. (2009), Santos et al. (2012)
Karavilagenin C		Anti-cancer	Ramalhete et al. (2009), Ma et al. (2012)
Balsaminagenin A		Anti-cancer	Ramalhete et al. (2009), Ma et al. (2012)
Balsaminoside B		Anti-cancer	Ramalhete et al. (2009), Ma et al. (2012)
Balsaminol F		Schistosomicidal activity against <i>Schistosoma mansoni</i> adult worms	Ramalhete et al. (2011), Ramalhete et al. (2016)

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TABLE 3 | (Continued) Compounds extracted from *Momordica balsamina* L. and their health benefits.

Compound	Structure	Health benefits	References
Momordin I		Anti-viral	Thakur et al. (2009)
Momordin II		Anti-viral	Thakur et al. (2009)
Balsaminapentaol		Anti-malarial	Ramalhete et al. (2009), Ma et al. (2012)
Cucurbalsaminol		Anti-parasitic	Ma et al. (2012), Shah et al. (2014)

Batra, 2012; Fongod et al., 2014; Talukdar and Hossain, 2014; Bulbul, 2016; Poyraz and Derdovski, 2016; Nitu and Patidar, 2017; Mostafa et al., 2018).

M. foetida leaves are used as fodder and protein supplement in some countries and are suitable for increasing rabbits weight (Jabeen and Khanum, 2017). However, reports from Kenya state that cattle avoid *M. foetida* because of its unusual smell and bitter taste.

Nutritional composition of *M. foetida* leaf is energy, protein, fiber, calcium, iron, magnesium, zinc, β -carotene, foliate and ascorbic acid (Oloyede and Aluko, 2012; Puškárová et al., 2017; Abegunde et al., 2018). The leaves of *M. foetida* contain considerable amount of anti-oxidants and are capable of inhibiting lipid peroxidation (Oloyede and Aluko, 2012; Acquaviva et al., 2013; Anilakumar et al., 2015; Dzotam

et al., 2016; Puškárová et al., 2017; Anjamma and Bhavani, 2018). Triterpenes are found in both *M. charantia* and *M. foetida*, particularly in the fruits and seeds, and are potentially cytotoxic.

***Momordica balsamina* L.**

M. balsamina is important in producing nutrients supplements due its high protein and fat values with low fiber content (Bakare et al., 2010; Costa et al., 2010; Roškar and Lušin, 2012; Ghosh, 2014; Madala et al., 2014; Talukdar and Hossain, 2014; Abegunde et al., 2018). The leaves of *M. balsamina* are an important source of nutrients, including 17 amino acids (Cohen et al., 1998; Hassan and Umar, 2006; Costa et al., 2010; Semenya and Potgieter, 2015). *M. balsamina* leaves are mixed with peanuts and honey and used as a meat sauce (Dellavalle et al., 2011; Oronje et al., 2012; Sen and

Batra, 2012; Fongod et al., 2014; Bulbul, 2016; Poyraz and Derdovski, 2016; Mostafa et al., 2018).

The fruit of *M. balsamina* contains the red, soft flesh that surround the edible seed (Tan C. et al., 2015; Birla, 2016; Devyani et al., 2016; Souda et al., 2018). The tender fruits and shoots of *M. balsamina* are usually boiled with meat and both can be decided to be added to the soup. Young fruits of *M. balsamina* are cooked and eaten, after peeling it is advisable to put on the salt water to minimize the bitterness before cooking, additionally the South Africans and Nigerians appreciate the fruit for its better taste (Acquaviva et al., 2013; Ghosh, 2014; Souda et al., 2018). The fruits of *M. balsamina* are frequent ingredients used in Indo Pakistan pickles and they are often used in curries and meat dishes.

Tender shoots of *M. balsamina* are usually utilized with Okra soup by the Kanuris of Borno State where the plant is locally known as the vegetable (Jabeen and Khanum, 2017). *M. balsamina* is preferred as an ingredient in aphrodisiac preparations however in Senegal they prefer to use the fruits as purgative agents, poultice and vermifuge (Leung et al., 2009; Souda et al., 2018).

MEDICINAL VALUE OF THE THREE MOMORDICA SPECIES

Momordica charantia L.

M. charantia has medicinal value and is also used as a vegetable. The compounds present in *M. charantia* exert anti-diabetic, anti-cancerous and anti-tumorous, anti-microbial, anti-viral, anthelmintic, antimalarial, anti-ulcerative and immunomodulatory activities (Gupta et al., 2011; Kulkarni et al., 2021; Lur et al., 2021; Malik et al., 2021). Some of the structures of compounds isolated from *M. charantia* are listed in **Table 1**.

Various medicinal properties have been claimed to be associated with *M. charantia* and they include anti-diabetic, abortifacient, anthelmintic, contraceptive, anti-malarial and laxative are used for various treatment such as dysmenorrhea, eczema, emmenagogue, galactagogue, gout, jaundice, kidney (stone), leprosy, leucorrhoea, pneumonia, psoriasis, rheumatism, scabies, and piles (Grover and Yadav, 2004; Kumar and Bhowmik, 2010; Chen et al., 2011; Acquaviva et al., 2013; Ghosh, 2014; Bulbul, 2016; Perera et al., 2021).

The fruit juice and a leaf tea extracted from *M. charantia* can be used for the treatment of diabetes, malaria, colic, sores and wounds, infections, worms and parasites, as an emmenagogue, and for measles, hepatitis, and fevers (Kumar and Bhowmik, 2010). The anti-oxidants and chemo-protective are mostly available on *M. charantia* fruit extracts drug that helps with reducing risk of cancer (Dzotam et al., 2016; Khalid et al., 2021). The leaves of *M. charantia* are used for treatment of wounds as well against worms and parasites. *M. charantia* has been used as a traditional medicine in China, India, Africa, and the south-eastern United States (Grover and Yadav, 2004). In the 1980s, the seeds of *M. charantia* were investigated in China as a potential contraceptive (Kumar and Bhowmik, 2010).

Natural products from *M. charantia* have been reported to significantly decrease prostaglandin E₂, interleukin and tumor necrosis factor and increases transforming growth factor and IL-10 secretion in RAW 264.7 macrophages, Caco-2 cells and THP-1 cells (Tan et al., 2008). The *M. charantia* fruit effectively enhanced T helper 2 hormonal responses and T helper 1 cellular immunity. A mixture of steroidal saponins contribute to the hypoglycemic and anti-hyperglycemic activity of *M. charantia* (Nitu and Patidar, 2017). Gentisic acid found as an active metabolite of salicylic acid and relation of the anti-inflammatory property of salicylic acid by inhibiting cyclooxygenase-2 (COX-2) mRNA expression and activity as well as prostaglandin E₂ (Lii et al., 2009; Semanya and Potgieter, 2015; Wardhani et al., 2015; Madala et al., 2016; Wang et al., 2016).

M. charantia have been reported to exhibit anti-diabetic, anthelmintic, abortifacient, anti-bacterial, anti-viral, hypoglycemic agents and chemo-preventive functions (Grover and Yadav, 2004; Costa et al., 2010; Ghosh, 2014; Bulbul, 2016; Dzotam et al., 2016; Ludidi et al., 2019; Kulkarni et al., 2021). The fruits and leaves of *M. charantia* display various biological activities including anti-diabetic, anti-rheumatic, and anti-ulcer, anti-inflammatory and anti-tumor (Dzotam et al., 2016). *M. charantia* contains various chemicals that have a hypoglycemic activity which reduces the amount of sugar in the blood, furthermore it stimulates appetite and helps in the entire digestion process. *M. charantia* is used for the treatment of digestive problems (Ludidi et al., 2019).

The leaves of *M. charantia* are used to treat diabetes, intestinal gas, promote menstruation, and as anti-viral agent against measles and hepatitis viruses (Tansey et al., 2004; Leung et al., 2009; Costa et al., 2010; Nerurkar et al., 2010; Nitu and Patidar, 2017; Perera et al., 2021; Liu et al., 2021). *M. charantia* is used by most of the culture for minimizing blood glucose and treating diabetes and as the other admired herbal resource (Leung et al., 2009; Birla, 2016). Hypoglycemic potential of *M. charantia* has been demonstrated in patients with type 2 diabetes. Emerging evidence also suggest that hypolipidemic action of *M. charantia* lowers serum cholesterol, hepatic total cholesterol and triglyceride and elevated cholesterol (Chen et al., 2003; Tan et al., 2008). The crop juice is believed significantly to reduce lipid accumulation and increase lipolysis in primary human adipocytes (Tan E. S. et al., 2015; Ingle et al., 2017). Animal studies indicate that *M. charantia* juice reduces body weight by inducing a reduced adipose hypertrophy, inhibition of lipogenic genes and enlarged plasma catecholamines (Nerurkar et al., 2010).

The ripe fruit of *M. charantia* fruits had been used as a remedy for tumors, asthma, skin infections and hypertension (Semanya and Potgieter, 2015). The seeds are surrounded by sweet red fleshy pulp that is edible tasting like watermelon (Poyraz and Derdovski, 2016). In some parts of West Africa, the leaves of *M. charantia* are cooked as part of green vegetable soup for post-natal mothers to prevent loss of blood during labor and purify breast milk. However, *M. charantia* can be recommended to use for livestock feeding as supplement for protein (Kuetee et al., 2010).

***Momordica foetida* Schumach**

Traditional medicinal uses are numerous, and many are shared with other *Momordica* spp. In East and Central Africa, *M. foetida* extracts are used to treat hypertension, peptic ulcers, and diabetes. Foetidin isolated from *M. foetida* is capable of reducing blood-glucose level in fasting albino rats (Marquis et al., 1977). Likewise, *M. foetida* is used for treatment of digestive problems due to its hypoglycemic activity that reduces the amount of sugar in the blood and also stimulate appetite and helps in the entire digestion process (Leung et al., 2009; Kumar and Bhowmik, 2010; Chen et al., 2011; Daniel et al., 2014; Birla, 2016).

Additionally, *M. foetida* has slight anti-spasmodic and anticholinergic effects (Hassan and Umar, 2006; Acquaviva et al., 2013; Ingle et al., 2017; Puškárová et al., 2017). Some of the structures of compounds extracted from *M. foetida* are listed in **Table 2**.

Aqueous extract of *M. foetida* has anti-oxidant activity due to high content in phenolic and flavonoid compounds (Acquaviva et al., 2013). Oxidative stress is closely related with Alzheimer and Parkinson diseases, diabetes, rheumatoid arthritis, and several diseases. The use of anti-oxidants in pharmacology has been instrumental in treatments of stroke and neurodegenerative diseases (Jabeen and Khanum, 2017).

The juice of *M. foetida*, dried and crushed leaves are used to relieve cough, intestinal disorders, headache, earache, toothache and as an cure for snake bites (Jabeen and Khanum, 2017). Leaves can be dried or boiled and used to treat skin problems, spitting cobra poison and malaria. The roots of *M. foetida* may be stated to be poisonous, and the crushed seeds are used in East Africa to cure of constipation. *M. foetida* helps to reduce weight in obese people (Nerurkar et al., 2010; Talukdar and Hossain, 2014).

The fruit pulp of *M. foetida* is said to be poisonous to common pests such as weevils, moths and ants, making it to be utilized as potential insect repellent in Tanzania. Momordocin isolated from *M. foetida* has insecticidal ability (Olaniyi and Marquis, 1975; Pinar et al., 1983; Pirillo et al., 1995). In Uganda the *M. foetida* whole plant is use on their cattle as an ox pecker repellent. In Gabon, *M. foetida* soaked and dried leaves are used to stuff cushions for commercial purpose. *M. foetida* is preferred to be grazed by cattle (Kuete et al., 2010; Mada et al., 2013) and is said to be good supplement of protein (Talukdar and Hossain, 2014). The fruits have the essential oils they have been used as insecticides, food additives and aromatherapy (Dandawate et al., 2016).

***Momordica balsamina* L.**

M. balsamina is regarded as the most important medicinal plants popularly used as a source of life saving drugs that is most important to the world's population (Hassan and Umar, 2006; Thakur et al., 2009; Kumar and Bhowmik, 2010; Tan C. et al., 2015; Souda et al., 2018). The crop can be consumed as vegetable in order to supply protein and potassium supplement for diets in poor rural communities, however its high potassium content is known for the executive of hypertension and other cardiovascular conditions (Rahmatullah et al., 2012; Semenya and Potgieter, 2015; Jabeen and Khanum, 2017; Souda et al., 2018). Some of the structures of compounds extracted from *M. balsamina* are listed in **Table 3**.

In some parts of Africa and Europe, soaked leaves of *M. balsamina* are used to treat wounds. Furthermore the fruits and leaves are used for treatment of wounds in Nigeria and Syria as hemostatic anti-septic (Mada et al., 2013; Semenya and Potgieter, 2015; Madala et al., 2016). The fruit of *M. balsamina* is used as liniment because of its strong smell and mixed with olive oil and almond oil to treat chapped hands, band and hemorrhoids (Mada et al., 2013). In West Africa *M. balsamina* fruit pulp is mixed with oil as an anti-phlogistic dressing. Mashed fruit of *M. balsamina* is utilized as poultice and bitter tonic (Semenya and Potgieter, 2015). *M. balsamina* plant is used as a medicine for treatment of fevers and yaws. However, Tsonga and Zulus have used leaves as tea for blood-liver deficiencies, stomach and intestinal ailments (Kumar and Bhowmik, 2010).

The Portuguese recommended *M. balsamina* leaves for blood, stomach, and liver deficiencies, furthermore they use leaves for herbal medicine and culinary herb (Thakur et al., 2009; Madala et al., 2014). *M. balsamina* leaves are used for diabetes, digestion disorder, fevers, ulcers, and mild form of malaria. Additionally, in West Africa *M. balsamina* is used as medicine in human and animals particular for fever, yaws and purgative (Oloyede and Aluko, 2012; Ingle et al., 2017; Jabeen and Khanum, 2017; Mostafa et al., 2018). *M. balsamina* roots are used as ingredient in an aphrodisiac preparation and in treatment of urethral discharge on both human and animals. It is advisable to use the whole *M. balsamina* parts of the plant for used in treating skin disease such as scabies (Waako et al., 2005; Ozusaglam and Karakoca, 2013; Pauliuc and Botau, 2013; Wikaningtyas and Sukandar, 2016). In South Africa, mostly the Pedi people use *M. balsamina* leaves and tendrils as potherb and anti-emetic, while the Venda people used the leaves infusions as anti-emetic and Congo as colic. *M. balsamina* leaves and fruit are widely used in Okavango delta for medicinal and spiritual purpose and also as skin ailments in China (Molehin and Adefegha, 2014). In some countries seed can be used as poison in arrow for hunting. The whole plant extract has insecticidal properties (Trakoon-osot et al., 2013; Wardhani et al., 2015; Wang et al., 2016).

The aqueous leaf extract of *M. balsamina* has also been used to minimize and relieve period pain in young girls and postnatal women used it to stimulate milk production (Tan et al., 2008). In Hausa land of Nigeria and Republic of Niger, the leaves are cooked as part of green vegetables soup for post-natal women, where it is believed to help the mother to restore her lost blood during labor and to cleanse her breast milk (Molehin and Adefegha, 2014).

Anti-oxidants are recognized as ingredient and supplement of good human health because they maintain health and prevent diseases such as cancer, coronary heart disease and even altitude sickness in human and animals (Tan et al., 2008; Ingle et al., 2017; Puškárová et al., 2017). *M. balsamina* is used for treatment of diabetes, malaria, colic, sores and wounds, infections, worms and parasites, as an emmenagogue, measles, and hepatitis, fevers, diarrheal, cancer, ulcer, HIV, bacterial infection, and constipation (Semenya and Potgieter, 2015; Abegunde et al., 2018; Aryanti and Lamdayani, 2021). In addition, *M. balsamina* is believed to treat mental illness (Anjamma and Bhavani, 2018). The anti-HIV properties are

found at the fruit pulp of *M. balsamina* (Abidemi, 2013). The leaves and fruit extracts of *M. balsamina* contain anti-plasmodial activity against malaria (Balouiri et al., 2016). The crude extract of protein on African pumpkin seeds contains a hemagglutinating activity attributed by the presence of sugar D-galactose and lactose (Bulbul, 2016). Moreover, several previous studies did test the hemagglutinating activity on human and animals of the different blood type and proved that the hemagglutinating was greater toward O blood type compared to other blood cell type (Scorzoni et al., 2007; Souda et al., 2018).

The fruit juice of *M. balsamina* exerted hypoglycemic activity, stimulates appetite and helps in the entire digestion process (Kumar and Bhowmik, 2010; Chen et al., 2011). Hypoglycemic potential of *M. balsamina* has been established in normal and diabetic rats and in patients with type 2 diabetes (Waako et al., 2005; Scorzoni et al., 2007; Leung et al., 2009; Kumar and Bhowmik, 2010; Rahmatullah et al., 2012; Joseph and Jini, 2013; Trakoon-osot et al., 2013; Ludidi et al., 2019). Extract of P-Insulin, polypeptide from the fruits and seeds of *M. balsamina* rapidly reduces and normalized the blood sugar level in rats.

The leaves have been found to be highly hemolytic and hepatotoxic in rats while the fruits are regarded as toxic to various organs and tissues of rats in very high dose (Semenya and Potgieter, 2015). Thus, it is important to determine the correct dose to optimize its potential use as a medicinal plant.

CONCLUSION AND FUTURE PERSPECTIVE

The biochemical composition of *M. charantia*, *M. foetida* and *M. balsamina* gives them a great and interesting nutritional and medicinal value. Many of its components have singular biological activity and the synergy of them may exert interesting pharmacological properties. Several studies demonstrated that these important plants have the potential to be utilized for medicinal purposes as they exhibit anti-diabetic, anti-microbial, anthelmintic, abortifacient, anti-bacterial, anti-viral, and chemo-preventive activities. These species possess a promising and innovative source of natural bioactive agents such as resins, alkaloids, flavonoids, glycosides, steroids, terpenes, cardiac glycoside, saponins, pectin, carbohydrates, amino acids, proteins, fats, fiber, chlorophyll, phosphorus, calcium, potassium, magnesium,

sodium, zinc, manganese, and iron. The main biological activities of *Momordica* spp. has been summarized in **Figure 2**.

The anti-microbial, anti-parasitic and anti-ulcer activities of *Momordica* spp. are the mechanisms related with their inhibitory effects against gastrointestinal diseases. In addition, anti-cancer effects of several biochemicals isolated from *Momordica* spp. has been described. Anti-oxidant activities of *Momordica* spp. also prevent cancer diseases avoiding oxidative stress and oxidative damage such as lipid peroxidation. Moreover, anti-oxidants exerts anti-aging and neuroprotection effects. Additionally, the hypoglycemic activities of *Momordica* spp. exert an anti-diabetes effect, the hypocholesterolemic activities exerts a cardioprotective effect, and their low ratio of Na^+/K^+ exerts beneficial effects against hypertension. In addition to being able to be consumed as food or supplement, clinical evidence suggests their effectiveness in managing various ailments such as dysmenorrhea, eczema, emmenagogue, galactagogue, gout, jaundice, kidney (stone), leprosy, leucorrhea, pneumonia, psoriasis, rheumatism, scabies, piles, cancer, coronary heart disease diabetes, digestion disorder, fevers, ulcers, malaria, tumors, asthma, skin infections, and hypertension.

Thus, the review suggests that the three selected crops have medicinal and herbal properties that can be used for various ailments. A better biochemical characterization is needed to better understand of medicinal properties of these plants. This review also assists to understand the commercial properties of the crop since the cultivation as commercial vegetable is currently underway. A better understand of pharmacological activities of *Momordica* spp. will give added value to these plants and benefit the producers.

AUTHOR CONTRIBUTIONS

All authors contributed equally to the manuscript. Conceptualization, MM, CQ, PT, TM, FM, MM, US, and RMK; validation investigation–data curation writing—all authors; review and editing, BS, AFAR, and JS-R. All the authors read and approved the final manuscript.

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Conflict of Interest: The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

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