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# A Phytochemical and Biological Review on Plants of the Family Aizoaceae <sup>2</sup>

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# **ABSTRACT 8**

**Objectives:** This study is aimed to be a comprehensive review of the phytochemical constituents and biological activities 9 of Aizoaceae family plants (Mesembryanthemaceae). **Methods:** This study is covering articles between 1969 and 2018, reviewed from internationally accepted databases and scientific data from scientific Journals. **Results:** Phytochemically studied plants of family Aizoaceae have shown the presence of various classes of compounds including; alkaloids, triterpenes, sterols, lignans, phenolic compounds, betacyanins, and essential oils. Biological studies on plants of family Aizoaceae have indicated various bioactive potentials including antioxidant, antidiabetic, antimicrobial, antitumor, hepatoprotective, anti-inflammatory and other effects. The reported medicinal plants of family Aizoaceae were selected and summarized on the basis of their; phytochemical constituents and biological activities. **Conclusion:** The results of this study may inspire further ethno-botanical and ethno-pharmacological research and investigations toward drug discovery.

**Keywords:** Aizoaceae; Biological activities; Phytochemical constituents 10

# **INTRODUCTION 11**

The Aizoaceae (Mesembryanthemaceae) 12 family is mostly composed of succulent plants of which 99% are found in South or Southwest of Africa. The other one-percent are found in coastal areas of Australia, New Zealand, Mediterranean area, Canary Islands, and the western coasts of Chile and California<sup>1</sup>.

Family Aizoaceae constitutes a major part of 13 the Southern African succulent flora. With approximately 1860 species in 127 genera, and in India the family is represented by three genera, namely, *Sesuvium*, *Trianthema* and *Zaleya* with seven species distributed in the coastal and lowland areas of

peninsular India and the Gangetic plains<sup>2</sup>. It is also 14 South Africa's second largest plant family<sup>3</sup>. Based on the division given by Bittrich & Hartmann<sup>4</sup>, five subfamilies are recognized: Aizooideae, Mesembryanthmoideae, Rushioideae, Sesuvioideae and Tetragonioideae<sup>5</sup>. In Egypt, Boulos<sup>6</sup> recognized five genera, but Hosny<sup>7</sup> listed only four genera without considering *Sesuvium*. According to Boulos<sup>6</sup> and Hosny<sup>7</sup>, Trianthema is represented in Egypt by two species: *T. portulacastrum* L. and either *T. triquetra* Willd. Ex Spreng. (according to Boulos<sup>6</sup>) or *T. Crystalline* (Forssk.) Vahl (according to Hosny<sup>7</sup>). *Zaleya* is represented by two species: *Z. pentandra* (L.) C. Jeffrey and *Z. decandra* (L.) Burm. f.<sup>6</sup>, or by only

one species: *Z. pentandra* (L.) C. Jeffrey<sup>7</sup>. This family 1 is represented in Egypt by 6 genera and 10 species<sup>8</sup>. It typically inhabits dry subtropical deserts and wet tropical coasts<sup>5-9</sup>. Native species of Aizoaceae are distributed through Mediterranean coastal habitats and the Nile Valley as weeds of cultivated land, and extend

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to deserts.

Many members of the family are of economic 2 importance as ornamentals and in cultivation worldwide10 as some of them are used to stabilize sand dunes in coastal regions 11 while others are important as medicinal plants<sup>12</sup> e.g. Tetragonia tetragonioides is used in treatment of enteritis and stomach ache as well as stomach cancer and ulcer12. Extracts from these plants are used as preservatives, as a remedy against throat infections and in soap making<sup>13</sup>. Although Aizoaceae is considered as one of the most diverse and abundant families, it was found that it is the least studied in terms of medicinal potential<sup>14</sup>. It would, therefore, be important to extensively investigate other plants of family Aizoaceae for future drug discovery. Thus, we aimed to make a comprehensive review of the chemical constituents and biological activities of family Aizoaceae covering the articles between 1969 and 2018.

#### MATERIAL AND METHODS 3

The research strategy employed in the review 4 of family Aizoaceae includes internationally accepted databases like Science Direct, Scopus and Web of Science as well as scientific data collected from scientific Journals.

# RESULTS AND DISCUSSION 5 Phytochemical constituents

A variety of chemicals have been isolated from 6 different species of family Aizoaceae including; alkaloids, triterpenes, sterols, lignans, flavonoids, phenolic compounds, essential oils, and some miscellaneous compounds<sup>5-15-16</sup>.

# Alkaloids 7

Mesembrine alkaloids are considered to be the 8 primary active constituents of the South African medicinal plant *Sceletium tortuosum* Aizoaceae), and it is used as the dried or fermented aerial material from the plant, which is known as Kanna (Aka, Channa, Kougoed). Traditional regional use ranged from relieving thirst, mild analgesia, and alteration of mood<sup>19</sup>. Only two genera of family Aizoaceae were reported to be rich with alkaloids which are genus *Sceletium* and genus *Trianthema* as shown in **Table 1**. *Sceletium spp*. is currently the only known genus having species with relatively high levels of mesembrine<sup>19</sup>, which belong to the crinane class of compounds based on the alkaloid skeleton<sup>17</sup>. Jeffs et al.

(1982) categorized the various Sceletium alkaloids into 9 groups: (1) the four structural 3a-aryl-cisoctahydroindole class (e.g. mesembrine and the new isolated alkaloid channaine)<sup>17-18</sup>, (2) C-seco mesembrine alkaloids (e.g. joubertiamine), (3) alkaloids containing a 2,3 disubstituted pyridine moiety and 2 nitrogen atoms (e.g. Sceletium alkaloid A4) and (4) a ring C-seco Sceletium alkaloid A4 group (e.g. tortuosamine)<sup>20</sup>. The mesembrine alkaloid is the most studied of the mesembrine alkaloids due to its relative abundance in S. tortuosum and biological activity, it was initially partially isolated, characterized, and named by Zwicky, and it was purified and crystallized as picrate 19 Mesembranol and its acetylated derivative, 4-Oacetylmesembrenol was also isolated together with 4-Odemethylmesembrenol and 4-O-demethylmesembranol from Sceletium spp in addition to mesembrenone. Joubertiamine and its derivatives represented a new structural class different from the known mesembranes. Joubertiamine, dehydrojoubertiamine, and dihydrojoubertiamine were all referred to as the secomesembranes and were isolated from Sceletium joubertii<sup>17</sup>. The preponderance of research on Sceletium alkaloids has revolved around isolation and structural elucidation. Very little is known about the distribution and chemotaxonomic patterns of these alkaloids within the genus<sup>17</sup>.

### Di-, Tri-, tetra-terpenes, and sterols:10

Phytochemical screening and analysis of some 11 species of family Aizoaceae showed presence of di-, tri, and tetraterpenes. β- amyrin, oleanolic acid, Uvaol were isolated from the leaves' methanolic extract of *C. edulis* <sup>16</sup>, while ecdysterone, were isolated from aqueous and organic extracts of *S. portulacastrum*—and *T. portulacastrum* <sup>15-24</sup> Pentandradione and pentandraone were isolated only from the methanolic extract of *Z. pentandra* <sup>50-51</sup>.

Sterols and their glycosides e.g. (Stigmasterol, 12  $\beta$ -Sitosterol, Stigmasterol 3-O- $\beta$ -D-glucoside and  $\beta$ -Sitosterol 3-O- $\beta$ -D-glucoside and the novel and recently isolated sterol 17-(5-ethyl-6-methylheptan-2-yl)-4,4,10,13-tetramethyl-hexadecahydro-1H-cyclopenta ( $\alpha$ ) phenanthren-3-ol ) have been reported to be isolated from T.  $portulacastrum^{15-16-21-22-24-25-26-27-28}$  as shown in **Table 2.** 

# Lignans and neolignans: 13

Aptenia cordifolia and *T. turgidifolium* leaves 14 are the only reported member of family Aizoaceae to show the presence of lignans and neolignans e.g. Pinorsinol, syringaresinol, di-erythro-syringylglycrol-β-*O*-4,4`-syringarsinol ether and apteniol A-G were isolated from the methanolic extract of the leaves of *A. cordifolia*<sup>22-29-30-31</sup>. Isoamericanin A was isolated from *T. turgidifolium* <sup>22</sup> as shown in **Table 3.** 

Table 1. Alkaloids isolated from plants of family Aizoaceae

Source	Used part/extract	Chemical constituents	Compound no
Sceletium spp. Sceletium tortuosum (L.) N. E. Br. Sceletium joubertii L. Bolus	Acid-base extraction of the entire plant	<ul> <li>Δ<sup>7</sup> mesembrenone<sup>17</sup></li> <li>4`-O-demethylmesemranol<sup>17</sup></li> <li>Mesembrenol<sup>17</sup></li> <li>Mesembranol<sup>17</sup></li> <li>4`-O-demethylmesemrenone<sup>17</sup></li> <li>Sceletenone<sup>17</sup></li> <li>N-demethyl-N-formyl mesembrenone<sup>17</sup></li> <li>O-acetylmesembrenol<sup>17</sup></li> <li>Mesembrane<sup>17</sup></li> <li>N-demethylmesembrenol<sup>17</sup></li> <li>N-demethylmesembrenol<sup>17</sup></li> <li>N-demethylmesembranol<sup>17</sup></li> <li>Hordenine<sup>17</sup></li> </ul>	1 12
Sceletium spp. Sceletium tortuosum (L.) N. E. Sceletium joubertii L. Bolus	Acid-base extraction of the entire plant	<ul> <li>Dehyrojoubertiamine<sup>17</sup></li> <li>Dihyrojoubertiamine<sup>17</sup></li> <li>Joubertiamine<sup>17</sup></li> <li>Joubertinamine<sup>17</sup></li> <li>O-methyldehydro-joubertiamine<sup>17</sup></li> <li>O-methyljoubertiamine<sup>17</sup></li> <li>O-methyldihydro-joubertiamine<sup>17</sup></li> <li>3`-methoxy-4`- O-methyljoubertiamine<sup>17</sup></li> <li>4-(3,4-dimethoxyphenyl)-4-[2-acetyl methylamino) ethyl]cyclohexanone<sup>17</sup></li> <li>3`-methoxy-4`- O-methyljoubertiaminol<sup>17</sup></li> <li>4-(3-methox-4-hydroxyphenyl)-4-[2-(acetylmethylamino) ethyl]cyclohexadienone<sup>17</sup></li> <li>3a-(3,4-dimethoxyphenyl)-1-methyl-1,2,3,3a,4,5,6,8,9,9b-decahydro-7H-pyrrolo[2,3-f] quinolin-7-one<sup>17</sup></li> <li>Sceletium alkaloid A4<sup>17</sup></li> <li>Touruosamine<sup>17</sup></li> <li>N-formyltortuosamine<sup>17</sup></li> <li>N-acetyltortuosamine<sup>17</sup></li> <li>Mesembrenone<sup>17</sup></li> <li>Channaine<sup>18</sup></li> <li>Mesembrine<sup>17</sup></li> <li>4`- O-demethylmesemrenol<sup>17</sup></li> </ul>	13
T. decandra L.	Organic extracts of leaves, fruits and seeds	-Trianthemine <sup>22</sup>	
T. portulacastrum L.	Organic extracts of the entire plant	- Punarnavine <sup>23</sup> - Trianthemine <sup>23</sup>	33
Sesuvium portulacastrum L.	Aqueous, methanolic and organic extracts of the whole dried plant	- Capsaicin <sup>21-25</sup>	34

Table 2. Reported plants of family Aizoaceae showing sterols, di-, tri-, and tetra-terpenes: 1

Reported Source	Used part/extract	Chemical constituents	Compound no.
Carpobrotus edulis L. Bolus	Methanolic extraxct of the leaves	- β- amyrin <sup>16</sup> - Oleanolic acid <sup>16</sup> - Uvaol <sup>16</sup>	35
Sesuvium portulacastrum L.	Aqueous, ethanolic and dichloromethane extracts of leaves and stems	<ul> <li>Ecdysterone<sup>15</sup></li> <li>Ecdysone<sup>15</sup></li> <li>22,23-Dihydrostigmasterol<sup>21-25</sup></li> <li>Ethyl iso-allocholate<sup>26</sup></li> <li>Squalene<sup>26</sup></li> <li>Phytol<sup>26</sup></li> </ul>	38
- Trianthema decandra L.	Leaves, fruits and seeds organic extracts	<ul> <li>3-Acetyl aleuritolic acid<sup>22</sup></li> <li>Stigmasterol 3- <i>O</i>-β-D-glucoside<sup>22</sup></li> <li>β-Sitosterol 3- <i>O</i>-β-D-glucoside<sup>22</sup></li> <li>Trianthenol<sup>22</sup></li> </ul>	44
Trianthema portulacastrum L.	Organic extracts of the aerial parts	- Ecdysterone <sup>24</sup> - 3-Acetyl aleuritolic acid <sup>24</sup> - Stigmasterol <sup>24</sup> - β-Sitosterol <sup>24</sup> - Stigmasterol 3- <i>O</i> -β-D-glucoside <sup>24</sup> - β-Sitosterol 3- <i>O</i> -β-D-glucoside <sup>24</sup> - Trianthenol <sup>22-24</sup> - 17-(5-ethyl-6-methylheptan -2-yl)-4, 4, 10, 13- vetramethyl-hexadecahydro-1H cyclopenta(α) phenanthren-3-ol] <sup>27</sup>	44> 50
Zaleya pentandra L.	Methanolic extract of the aerial parts	- Pentandradione <sup>28</sup> - Pentandraone <sup>28</sup>	51, 52

# Fatty acids, and fatty alcohols: 3

Sesuvium portulacastrum is the only member 4 of family Aizoaceae with data reported on isolates of fatty acids and fatty alcohols as shown in **Table 4**. It showed the presence of; Linolenic acid, oleic acid, eicosyl ester, 9,12,15-Octadecatrienoic acid, 2,3-dihydroxypropyl ester, (Z,Z,Z)-, hexadecanoic acid ethyl ester, lauric acid, tridecanoic acid, myristic acid, pentadecanoic acid, palmitic acid, heptadecanoic acid, stearic acid, oleic acid, linoleic acid, nonadecanoic acid, arachidic acid, heneicosanoic acid, behenic acid, octadecanoic acid, 1-Docosanol and rhodopsin, which were isolated from dichloromethane extract of its leaves and stems<sup>11-26-32-33</sup>.

#### Catechins and phenolic acids 5

Reported data showed that catechins and 6 phenolic acids were isolated from six members only of family Aizoaceae which are; *A. cordifolia*, *C. edulis*, *M. crystallinum*, *S. portulacastrum*, *T. decandra*, and *T. portulacastrum*<sup>14-21-22-25-26-29-33-34-35-36</sup>. It has been reported, that epicatechin was isolated from the organic extracts of the leaves of two different plant species *C. edulis* and *S. portulacastrum*<sup>14-21-22-25-26-29-33-34-35-36</sup>. On the other hand, we found, that 2,6-bis(1,1-dimethylethyl)-4-methylphenol is the only phenolic

compound isolated from *M. crystallinum*. Other isolated 7 catechins and phenolic compounds are shown in **Table 5.** 

#### Flavonoids 8

Several flavonoids were isolated from different 9 species of family Aizoaceae. Rutin was isolated from the leaves ethyl acetate extract of *C. edulis* and *Mesembryanthemum forsskaolii*. Moreover, the new compound C-methyl flavone was isolated from the organic extracts of the whole plant of the two species *T. portulacastrum* and *T. decandra*<sup>11-22-24-34-37-38-39-40</sup>. Moreover, 2-(3',4'dihydroxyphenyl) 3,5,7- trihydroxychromen-4 one was also isolated from *T. decandra*<sup>40</sup>. Other isolated compounds are shown in **Table 6.** 

#### Betacyanin 10

The genus *Lampranthus* of family Aizoaceae 11 was observed according to the reported data in **Table** (7) to show the presence of betacyanins. This includes *Lampranhus peersu* and *Lampranthus sociorum*. Betanin was isolated from the two plant species *Drosanthemum floribundum* and *Lampranthus spp*. Also, dopaxanthin was the only betacyanin isolated from *Glottiphyllum longum*, while  $\beta$ -Cyanin was isolated from *T. portulacastrum*<sup>24-41-42-43-44</sup>.

Table 3. Lignans, neolignans isolated from plants of family Aizoaceae 1

Species	Used part/extract	Chemical constituents	Compound no.
Aptenia cordifolia L.F.	Methanolic extract of the leaves	<ul> <li>Pinorsinol<sup>29</sup></li> <li>Syringaresinol<sup>29</sup></li> <li>Di-erythro-syringylglycrol-β-o-4,4`-syringarsinol ether<sup>29</sup></li> <li>Apteniol A, B, C, D, E, F</li> <li>Apteniol G<sup>30</sup></li> </ul>	53> 59
Trianthema turgidifolium F. Muell	Methanolic extract of the leaves	– Isoamericanin A <sup>22</sup>	

Table 4. Fatty acids and fatty alcohols isolated from plants of family Aizoaceae 3

Species	Used part/extract	Chemical constituents	Compound no.
Sesuvium portulacastrum L.	Dichloromethane extracts of leaves and stems	- Linolenic acid <sup>11-32</sup> - Oleic acid eicosyl ester <sup>26</sup> - 9,12,15-Octadecatrienoic acid, 2,3-dihydroxypropyl ester, (Z,Z,Z)- <sup>26</sup> -Hexadecanoic acid ethylester <sup>26</sup>	60
Sesuvium portulacastrum L.	Dichloromethane extracts of leaves and stems	- Lauric acid <sup>32</sup> - Tridecanoic acid <sup>32</sup> - Myristic acid <sup>11,32</sup> - Pentadecanoic acid <sup>32</sup> - Palmitic acid <sup>11,32</sup> - Heptadecanoic acid <sup>32</sup> - Stearic acid <sup>32</sup> - Oleic acid <sup>11,26,32</sup> - Linoleic acid <sup>11,26,32</sup> - Linoleic acid <sup>11,32</sup> - Nonadecanoic acid <sup>32</sup> - Arachidic acid <sup>32</sup> - Heneicosanoic acid <sup>32</sup> - Behenic acid <sup>11,32</sup> - Octdecanoic acid <sup>33</sup> - 1 - Docosanol <sup>26</sup> - Rhodopin <sup>26</sup>	64> 79

# Nitrogen containing compounds 5

The reported data showed that some nitrogen-6 containing compounds like 2-(dimethylamino)-1phenylethanol, 3-(1H-inol-3-yl) propionic acid, 3-(1Hinol-3-yl)propionic acid methyl ester, (2S,E)-N-[2-Hydroxy-2-(4-hydroxyphenyl)ethyl] Ferulamide, (E)-N--[2-Hydroxy-2-(4-hydroxy-3-methoxyphenyl)-ethyl] ferulamide, and (E)-N-[2-(4-Hydroxyphenyl)-2propoxyethyl] ferulamide has been isolated from the methanolic extract of A. cordifolia, in addition to many compounds e.g. Trans-4-hydroxyprolinebtaine, and Pvrrolo [1,2-A]Pyrazin-1,4-dione,hexahydro-3-(phenylmethyl) were isolated from aqueous and organic extracts of S. porulacastrum. Moreover, nicotinic acid is the only isolated nitrogen-containing compound isolated from T. portulacastrum<sup>15-21-23-25-29-31-33</sup>as shown in Table 8.

# Essential oils 7

The available data reported that only the leaves 8 of *S. porulacastrum* yielded oil via hydro-distillation which upon analysis using GC-MS showed the following composition: o-cymene,  $\alpha$ -pinene, trans-caryophyllene, 1,8- cineole, limonene,  $\alpha$ -terpinene,  $\alpha$ -terpinene, camphene, (-)-bornylacetate, tridecane and  $\alpha$ -humulene<sup>21-45</sup> as shown in **Table 9.** 

# Miscellaneous compounds: 9

It has been reported, that different 10 miscellaneous compounds have been isolated from five different plants of family Aizoaceae including; *A. cordifolia*, *C. edulis*, *T. portulacastrum* and *S. portulacastrum* <sup>16-23-24-26-29-33-46-47</sup> as shown in **Table 10**.

Table 5. Catechins and phenolic acids and their esters isolated from plants of family Aizoaceae 1

Species	Used part/extract	Chemical constituents	Compound no.
Aptenia cordifolia L.F.	Methanolic extract of the leaves	<ul> <li>- 4-hydroxybenzoic acid<sup>29</sup></li> <li>- Dihyrocinnamic acid<sup>29</sup></li> <li>- 4-hydroxy-dihydrocinnamic acid<sup>29</sup></li> <li>- Dihydrofrulic acid<sup>29</sup></li> <li>- 3,4-dimethoxy-dihyrocinnamic acid<sup>29</sup></li> <li>- 3,4-dimethoxy-dihyro cinnamic acid methyl ester<sup>29</sup></li> <li>- 3,4-dimethoxy-dihyrocinnamic acid ethyl ester<sup>29</sup></li> <li>- Methylfrulate<sup>29</sup></li> <li>- Sinapic acid<sup>29</sup></li> <li>- Methyl 2,5-dihyroxybenzoate<sup>29</sup></li> <li>- 3,4,5-Trimethoxyphenol<sup>29</sup></li> <li>- 3-Hydroxy-7,8-dihydro-β-ionone<sup>29</sup></li> <li>- 3,4-dimethoxycinnamic acid<sup>29</sup></li> </ul>	80
Carpobrotus edulis L. Bolus	Leaves methanolic and ethylacetate extract	- Ferulic acid <sup>34</sup> - Catechin <sup>14</sup> - Epicatechin <sup>14</sup> - Procyanidin (B5) <sup>14</sup>	86 —— 88
Mesembryanthemum crystallinum L.	Methanolic extract of the leaves	2,6-bis(1,1-dimethylethyl)-4-methylphenol <sup>35,36</sup>	89
Sesuvium portulacastrum L.	Aqueous, ethanolic and dichloromethane extracts of leaves and stems	<ul> <li>Epicatechin<sup>21,25</sup></li> <li>Phenol,2,4-bis(1,1-dimethylethyl)<sup>33</sup></li> <li>Gallic acid<sup>21,25</sup></li> <li>Benzoic acid<sup>21</sup></li> <li>Benzoic acid, 4-ethoxy-, ethyl ester<sup>26</sup></li> </ul>	90 9
Trianthema decandra L.	Leaves, fruits and seeds organic extracts	<ul> <li>5- Hydroxy-2-methoxy benzaldehyde<sup>22</sup></li> <li>- P-Methoxybenzoic acid<sup>22</sup></li> <li>- P-Propoxybenzoic acid<sup>22</sup></li> <li>- Leptorumol<sup>22</sup></li> </ul>	94> 9
Trianthema portulacastrum L.	Organic extracts of the aerial parts	<ul> <li>4-hydroxybenzoic acid<sup>24</sup></li> <li>3,4-Dimethoxycinnamic acid<sup>24,35</sup></li> <li>Pyrogallol<sup>24</sup></li> <li>5- Hydroxy-2-methoxy benzaldehyde<sup>35</sup></li> <li>Protocatechuic acid<sup>24</sup></li> <li>Vanillic acid<sup>24</sup></li> <li>O-Coumaric acid<sup>24</sup></li> <li>Caffeic acid<sup>24</sup></li> <li>P-Methoxybenzoic acid<sup>24,35</sup></li> <li>P-Propoxybenzoic acid<sup>24,35</sup></li> <li>Leptorumol<sup>24</sup></li> </ul>	94

# **Biological activities: 3**

Members of family Aizoaceae are known to 4 have diverse biological activities including antihyperlipidemic, antipyretic, diuretic, antioxidant, anticancer, larvicidal, analgesic, anti-rheumatic, anticholera, emetic, laxative, anti-inflammatory andantimicrobial. As well as being used in the treatment of; skin diseases, specific blood diseases, jaundice, cataract, night blindness, heart diseases, joint pain, dropsy, ascites, edema, and others as shown in **Table 11**.

# **Antioxidant activity 5**

Oxidative stress can cause damage to tissues 6 and cells. Free radicals, such as nitric oxide, superoxide anions, and hydroxyl radicals, can result in oxidative stress and may inflict damage in almost every organ. Furthermore, cancers also can arise from excess reactive oxygen species (ROS) that can damage cellular DNA. The antioxidant potential of the following plant organic extracts: *T. portulacastrum*, *Mesembryanthemum forsskaolii*, *Aizoon canariense*,

Table 6. Flavonoids isolated from plants of family Aizoaceae 1

Species	Used part/extract	Chemical constituents	Figure no.
Carpobrotus edulis L. Bolus	Ethyl acetate extract of	- Rutin <sup>34</sup>	103, 104
	leaves	- Hyperoside <sup>34</sup>	
		- Neohesperidin <sup>34</sup>	
Mesembryanthemum	Methanolic extract of	- Rutin <sup>37</sup>	103
forsskaolii Hochst. ex Bioss	the entire herb	- Apigenin <sup>37</sup>	105
		- Apigenin-7- <i>O</i> -glucoside <sup>37</sup>	
		- Kaempferol-3- <i>O</i> -glucoside <sup>37</sup>	
		- Isorhamnetin-3- <i>O</i> -β-glucopyranoside <sup>37</sup>	
Sesuvium portulacastrum L.	Aqueous, ethanolic	- 3,5,4`-trihyroxy-6,7-dimethoxyflavone <sup>11</sup>	103, 109
	and dichloromethane	- 3,5-dihydroxy-6,3 ',4' -trimethoxy-flavone-7-0- $[\alpha$	
	extracts of the leaves	L-rhamnopyranosyl-1 (1—6)- β-D-	
	and stems	Glucopyranoside <sup>11</sup>	
Trianthema portulacastrum	Organic extracts of the	- Quercetin <sup>24</sup>	103, 110
L.	dried plant	- C-Methyl Flavone <sup>24,38</sup>	
		- 5, 2'-dihydroxy-7-methoxy-6, 8-dimethylflavone <sup>39</sup>	
Trianthema decandra L.	Leaves, fruits and	- C-Methyl Flavone <sup>22</sup>	103, 111
	seeds organic extracts	- 2 - (3', 4' dihydroxyphenyl) 3, 5, 7 - trihydroxy-	
		chromen-4 one <sup>40</sup>	

Table 7. Betacyanins isolated from plants of family Aizoaceae 3

Species	Used part/extract	Chemical constituents	Compound no.
Drosanthemum floribundum	Methanolic extract of	- Betanidine <sup>41</sup>	112
(Haw.) N.E.Br.	the flowers	- Isobetanidine <sup>41</sup>	
		- Betanin <sup>41</sup>	
		- Isobtanin <sup>41</sup>	
Glottiphyllum longum	Methanolic extract of	Dopaxanthin <sup>42</sup>	116
(Haw.) N.E.Br.	the flowers		
Lampranthus spp	Aqueous extract of the	- Betanin <sup>43</sup>	114, 115
	flowers	- Isobtanin <sup>43</sup>	
Lampranthus sociorum	Methanolic extract of	Betanidin 5-0-[2"- <i>O</i> -(E)-feruloyl- β -(1",2')-	117
N.E.Br.	the flowers	glucuronosyl-β-glucoside] <sup>44</sup>	
Lampranhus peersii N.E.Br.	Methanolic extract of	Betanidin- 5-0-[6'- O-(E)-feruloyl- β-glucoside] <sup>44</sup>	118
	the flowers		
Trianthema portulacastrum	Organic extracts of the	β-Cyanin <sup>24</sup>	119
L.	dried plant		

Mesembryanthemum Nodiflorum, Mesembryanthemum 5 crystallinum, Mesembryanthemum forsskaolii, S. portulacastrum Mesembryanthemum nodiflorum, Carpobrotus edulis, Mesembryanthemum edule Mesembryanthemum crystallinum and T. decandra were investigated by 1,1-diphenyl-2-picryl hydrazyl (DPPH) and hydrogen peroxide assays. The results indicated that the organic extracts possessed a concentration-dependent free radical-scavenging activity against DPPH and hydrogen peroxide radicals, which was comparable with standard ascorbic acid 15-22-24-36-37-48-49-50.

# Anticancer activity: 6

The organic extracts of the following plants: *T.7* portulacastrum, *S. portulacastrum*, Carpobrotus edulis and *T. decandra* showed anticancer activity against mouse lymphoma cells and hepatic carcinoma using MTT assay<sup>16-22-24-33</sup>.

The protective role of *T. portulacastrum* 8 against diethylnitrosoamine–induced experimental hepatocarcinogenesis was evaluated. Morphometric evaluation of focal lesions showed a reduction of altered liver cell foci/cm<sup>2</sup> and a reduction of the average focal area. A decrease in the percentage of liver

Table 8. Nitrogen containing compounds isolated from plants of family Aizoaceae 1

Species	Used part/extract	Chemical constituents	Compound no.
Aptenia cordifolia L.F.	Leaves methanolic extract	- 2-(dimethylamino)-1-phenylethanol <sup>29</sup> - 3-(1H-inol-3-yl) propionic acid <sup>29</sup> - 3-(1H-inol-3-yl) propionic acid methyl ester <sup>29</sup> - (2S, E)-N-[2-Hydroxy-2-(4-hydroxyphenyl) ethyl] Ferulamide <sup>31</sup> - (E)-N-[2-Hydroxy-2-(4-hydroxy-3-methoxyphenyl)-ethyl] ferulamide <sup>31</sup> - (E)-N-[2-(4-Hydroxyphenyl)-2-propoxyethyl] ferulamide <sup>31</sup> - (E, E)-N,N-Dityramin-4,40 -dihydroxy-3,50 -dimethoxy-b,30 -bicinnamamide <sup>31</sup> - 7-Hydroxy-1-(4-hydroxy-3-methoxyphenyl)-N2, N3-bis(4-hydroxyphenethyl)-6-methoxy-1,2-dihydronaphthalene-2,3-dicarboxamide <sup>31</sup>	120 125
Sesuvium portulacastrum L.	Aqueous, methanolic and organic extracts of the whole dried plant	- Trans-4-hydroxyprolinebtaine <sup>15</sup> - Pyrrolo[1,2-A] Pyrazin-1,4-dione, hexahydro-3-(phenylmethyl) <sup>33</sup> - Pyrrolo[1,2-A] Pyrazin-1,4-dione, hexahydro-3-(2-methylpropyl) <sup>33</sup> - Butanoic acid, pyrolidide <sup>33</sup> - L-proline, N-Valeryl-, hexadecyl ester <sup>33</sup>	126 → 130
Trianthema portulacastrum L.	Organic extracts of the whole dried plant	- Nicotinic acid <sup>23</sup>	131

Table 9. Essential oils isolated from plants of family Aizoaceae 3

Species	Used part/extract	Chemical constituents	Compound no.
Sesuvium porulacastrum L.	Leaves hydrodistillation	- O-cymene <sup>21,45</sup>	132
		- α-Pinene <sup>21,45</sup>	
		<b>-</b> 2-β-Pinene <sup>45</sup>	
		- Trans-Caryophyllene <sup>21,45</sup>	
		- 1,8- Cineole <sup>21,45</sup>	
		- Limonene <sup>21,45</sup>	
		- α-Terpinene <sup>21-45</sup>	
		- α-Terpinolene <sup>21,45</sup>	
		- Camphene <sup>21,45</sup>	
		- (-)-Bornylacetate <sup>21,45</sup>	
		- Tridecane <sup>21,45</sup>	
		- α- Humulene <sup>21,45</sup>	

Table 10. Miscellaneous compounds isolated from plants of family Aizoaceae 1

Species	Used part/extract	Chemical constituents	Compound no.
Aptenia cordifolia L.F.	Methanolic extract of the leaves	- 4-(hydroxymethyl) phenol <sup>29</sup> - 4-(hydroxymethyl)-2,6 dimethoxyphenol <sup>29</sup> - Dehydrololiolide <sup>29</sup> - (9R)-9-hyroxymegastigm-4-ene-3-one <sup>29</sup> - Mgastigm-4-ene-3,9-dione <sup>29</sup> - 4-oxo-7,8-dihydro-β-ionone <sup>29</sup> - (3R,9R)-3,9-dihyroxymegastigm-5-en-4-one <sup>29</sup> - 3- <i>O</i> -methyl-chiro-inositol <sup>29</sup>	144> 149
Carpobrotus edulis L. Bolus	Methanolic extract of the leaves	Monogalactosyl diacylglycerol <sup>16</sup>	150
Trianthema portulacastrum L.	Organic extracts of the dried whole plant	- β-Carotene <sup>24</sup> - Ascorbic acid <sup>23</sup>	151, 152
Trianthema decandra L.	Methanolic extract of the leaves	Bis (2-ethyl hexyl) phthalate <sup>46</sup>	153
Sesuvium portulacastrum L.	Aqueous, methanolic and organic extracts of the dried plant	<ul> <li>Hentriacontane<sup>33</sup></li> <li>L-(+)-ascorbic acid,2-6-dihexaecanoate<sup>33</sup></li> <li>Vitamin E<sup>26</sup></li> <li>1-monolinoleoylglycerol-trimethylsilyl ether<sup>26</sup></li> <li>Dibutylphthalate<sup>47</sup></li> <li>Diisooctyl phthalate<sup>47</sup></li> </ul>	154

parenchyma occupied by foci seems to suggest the 3 anticarcinogenic potential of the plant extract in DENA-induced hepatocarcinogenesis<sup>22</sup>.

# Analgesic, antinociceptive, antihyperglycemic and 4 hepatoprotective activity

The methanol extract of the leaves of T.5 portulacastrum and T. decandra have remarkable analgesic, antinociceptive activity, antihyperglycemic and hepatoprotective activity<sup>22-24</sup>.

The analgesic activity of the leaf extract of T. 6 decandra was detected by hot plate and acetic acidinduced writhing response method. The results indicate that the administration of leaf extract of T. decandra exhibit central analgesic properties since it exerted a significant and dose-dependent effect on the chemical (acetic acid-induced) and thermic (heat) painful stimuli from the respective doses of 100 and 200 mg/kg<sup>22</sup>.

Many natural plant extracts have been 7 investigated with respect to the suppression of glucose production from carbohydrates in the gut or glucose absorption from the intestine.

 $\alpha$ - amylase catalyzes the hydrolysis of  $\alpha$  -1,4-8 glucosidic linkages of starch, glycogen and various

oligosaccharides and  $\alpha$ -glucosidase further breaks down 9 the disaccharides into simpler sugars, readily available for the intestinal absorption. The inhibition of their activity, in the digestive tract of humans, is considered to be effective to control diabetes by diminishing the absorption of glucose decomposed from starch by these enzymes. Therefore, effective and nontoxic inhibitors of  $\alpha$  - amylase and  $\alpha$ -glucosidase have long been sought, and for the first time, studies have revealed the anti-diabetic potential of T. decandra and these studies could be helpful to develop medicinal preparations for diabetes<sup>22</sup>.

The probable mechanism by which *T.* 10 *portulacastrum* exerts its hepatoprotective action against paracetamol or thioacetamide-induced hepatocellular metabolic alterations could be by the stimulation of hepatic regeneration through an improved synthesis of protein or accelerated detoxification and excretion<sup>22</sup>.

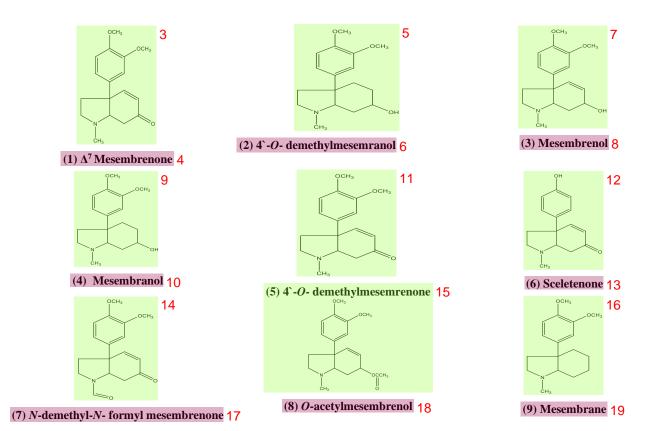
#### Anti-inflammatory activity 11

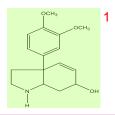
The anti-inflammatory activity of methanolic 12 and organic extracts of the following plants: *T. portulacastrum, Aizoon canariense, Aizoon hispanicum,* 

Plant name	Biological activities	References
Aptenia cordifolia L.F.	- Algicide activity	29
	- Antifungal	
Aizoon canariense L.	- Antioxidant	48,51,52
	- Antibacterial	
	- Skin diseases	
	- Emetic	
	- Antiiflammatory	
	- Antifungal	
	- Wound healing	
Aizoon hispanicum L.	Antiiflammatory	51
Carpobrotus acinaciformis L.	Antidiarrhea and dysentery	50,53
	- Antioxidant	16,34,35,53,54
Carpobrotus edulis L. Bolus		10,54,55,55,5
	- Anticancer	
	- Antibacterial	
	- Antidiarrhea and dysentery	
C	- Anti HFFIV/AIDS	52
Carpobrotus muirii L.	Antidiarrhea and dysentery	53
Calania Africana I	- Antioxidant - Antituberculosis	5.5
Galenia Africana L.		55
G: 1: 1 :1 I	- Asthma and obstructive pulmonary diseases	50
Gisekia phernaceoides L.	Antibacterial	56
Lampranthus francisci L. Bolus	Anticandidal	57
Mesembryanthemum anatomicum Haw.	- Mood altering	58
	- CNS stimulant	
Mesembryanthemum crystallinum L.	- Antioxidant	35,36,48,50
	- Antifungal	
	- Antibacterial	
Mesembryanthemum forsskaolii Hochst.	- Antioxidant	35,37
ex Bioss	- Antibacterial	
	- Antifungal	
Mesembryanthemum nodiflorum L.	- Antibacterial	35,49
	- Antioxidant	
	- Antifungal	
	- Cytotoxic	
Sceletium spp.	- Antiinflammatory	17,58
	- Asthma and obstructive pulmonary diseases	
	- Cytotoxic	
	- Psoriasis	
	- Cocaine like activity	
	- Antidepressant	
	- Psychiatric conditions	
	- Antianxiety	
Sceletium tortuosum (L.) N. E. Br.	- Antidote in alcohol poisoning	17,58,59
	- Antidepressant	
	- Psychiatric conditions	
	- Antianxiety	
	- Hypnotic and sedative	
Sesuvium portulacastrum L.	- Antioxidant	15,33
,	- Anticancer	
	- Antifungal	
	- Antibacterial	
	- Antiulcerogenic	
	- Kidney disorders	

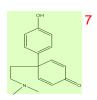
# Table 11. Cont. 1

Sesuvium verrucosum Raf.	Cytotoxic	21	2
Trianthema decandra L.	- Antioxidant	22,40	
	- Anticancer		
	- Analgesic and antinociceptive		
	- Antihyperglycemic		
	- Hepatoprotective		
	- Antiinflammatory		
	- Antibacterial		
	- Antifungal		
	- Antiulcerogenic		
	- Wound healing		
Zaleya pentandra L.	- Antiacetylcholinesterase	28,60,61	
	- Antibutyrylcholinesterase		
	- Antifungal		
	- Stomach diseases		
	- Respiratory tract infection and cough		
	- Asthma and obstructive pulmonary diseases		
	- Laxative		
	- Blood diseases		
	- Jaundince		
	- Larvicidal		
	- Cataract and night blindness		

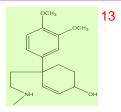




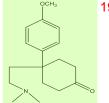
(10) N-demethylmesembrenol 2



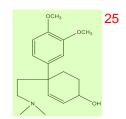
(13) Dehyrojoubertiamine 8



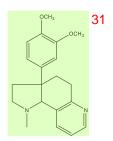
(16) Joubertinamine 15



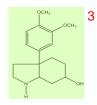
(19) O-methyl dihydrojoubertiamin 20



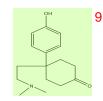
(22) 3`-methoxy-4`- *O*-methyl 26 joubertiaminol



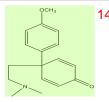
(25) Sceletium alkaloid A4 35



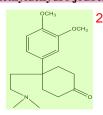
(11) N- demethylmesembranol 4



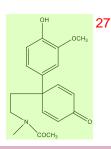
(14) Dihyrojoubertiamine 10



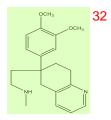
(17) O-methyldehydro joubertiamine 17



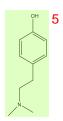
(20) 3`-methoxy-4`-o- methyl 22 joubertiamine



(23) 4-(3-methox-4-hydroxy phenyl)-4- [2 29 (acetyl methyl) amino] ethyl cyclohexanone



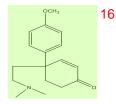
(26) Touruosamine 34



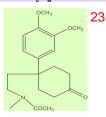
(12) Hordinine 6



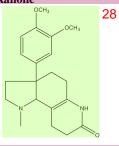
(15) Joubertiamine 12



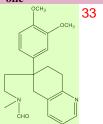
(18) *O*-methyl joubertiamine 18



(21) 4-(3,4dimethoxyphenyl)-4 – [2-24 (acetyl methyl) amino] ethyl cyclohexanone

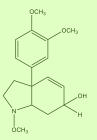


(24) 3a-(3,4-dimethoxy phenyl)-1-30 methyl1,2,3,3a,4,5,6,8,9,9b decahydro-7H-pyrrolo[2,3-f] quinolin-7-one

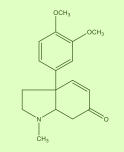


(27) N-formyl tortuosamine 36

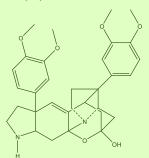
# (28) N-acetyl tortuosamine



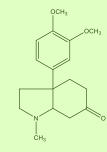
# (31) 4`-O-demethylmesemrenol



(29) Mesembrenone



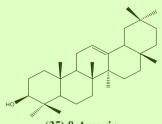
(32) Channaine



(30) Mesembrine

(33) Punarnavine

(34) Capsaicin



(37) Uvaol

(38) Ecdysterone

(39) Ecdysone

(40) 22, 23-Dihydrostigmasterol

(41) Ethyl iso-allocholate

(42) Squalene

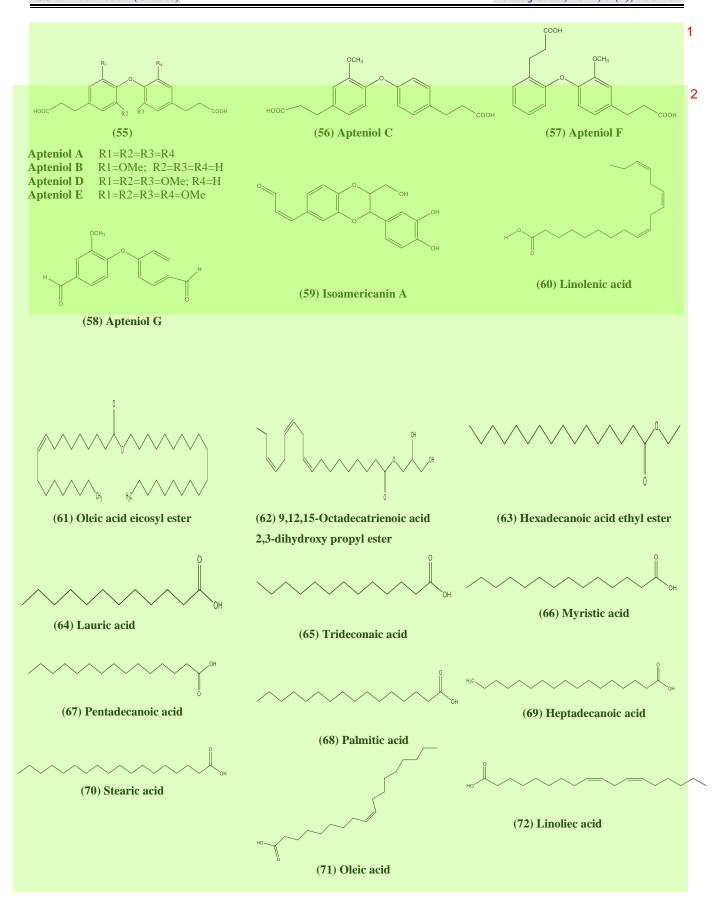
(52) Pentandraone

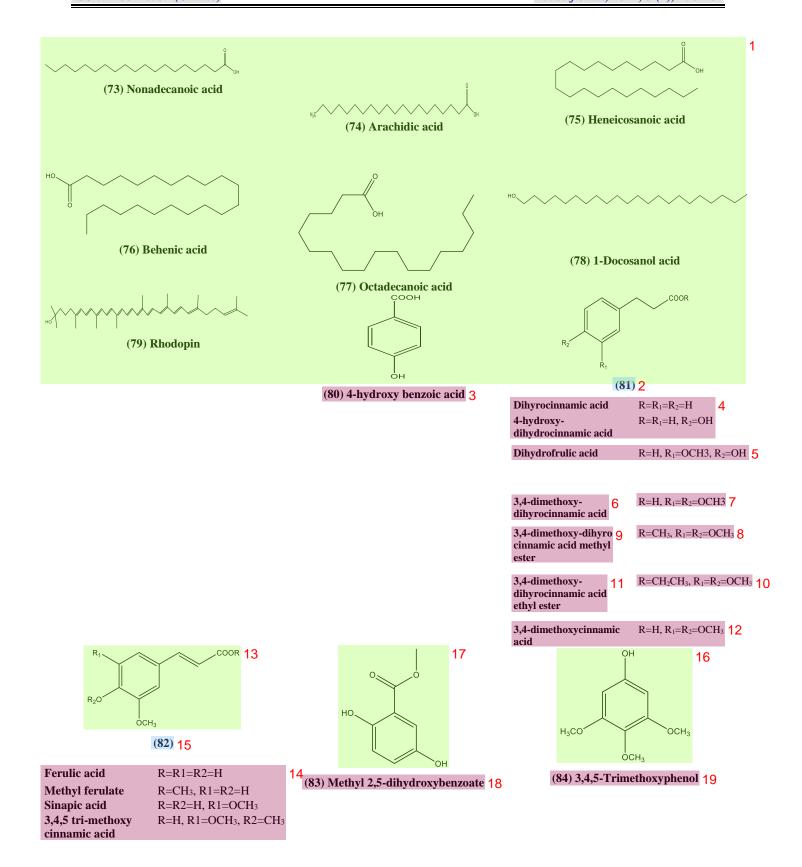
1

(43) Phytol (44) 3-Acetyl aleuritolic acid (45) Stigmasterol 3- *O*-β-Dglucoside (46) β-Sitosterol 3- O- β-D-glucoside (47) Trianthenol (48) Stigmasterol (49) β-Sitosterol  $(50)\ 17\hbox{-}(5\hbox{-ethyl-}6\hbox{-methylheptan -}2\hbox{-yl})\hbox{-}4,$ 4, 10, 13- vetramethyl-hexadecahydro-1H (51) Pentandradione cyclopenta(α) phenanthren-3-ol]

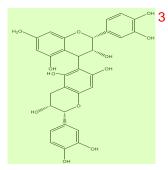
> Pinorsinol R=R1=H Syringaresinol R=OCH<sub>3</sub>, R1=H

(54) Di-erythro-syringylglycrol- $\beta$ -o- 3 4,4 $\hat{}$ - syringaresinol ether

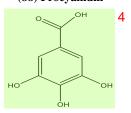




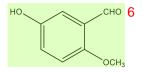
(85) 3-Hydroxy-7,8-dihydro-β-ionone 2



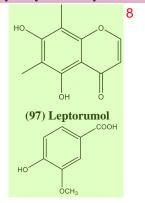
(88) Procyanidin



(91) Gallic acid 5

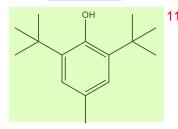


(94) 5-Hydroxy-2-methoxy benzaldehyde 7

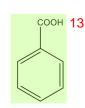


(100) Vanillic acid 23

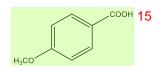
(86) Catechin 10



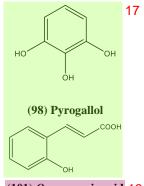
(89) 2,6-bis(1,1-dimethylethyl)-4- 12 methylphenol



(92) Benzoic acid 14



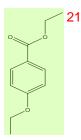
(95)  $\rho$ -methoxy benzoic acid 16



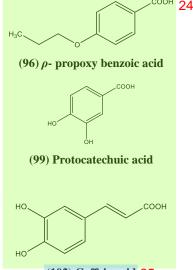
(101) O- coumaric acid 18



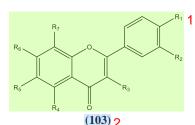
(90) Phenol 2,4-bis(1,1-dimethylethyl) 20



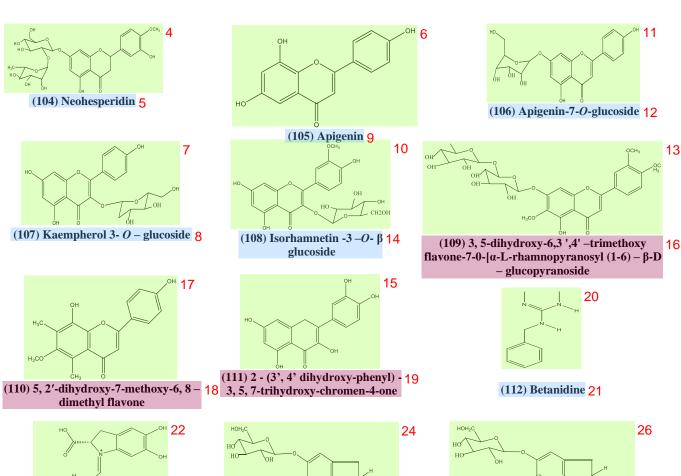
(93) Benzoic acid, 4-ethoxy ethyl ester 22

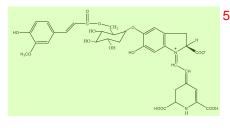


(102) Caffeic acid 25



(103)									
	Compound	R1	R2	R3	R4	R5	R6	<b>R7</b>	3
	Rutin	OH	OH	O-Rutinose	OH	H	OH	Н	
	Quercetin	OH	OH	OH	OH	Н	OH	Н	
	3,5,4`-trihyroxy-6,7-	OH	Н	OH	OH	OCH3	OCH3	Н	
	dimethoxyflavone								
	C-Methyl flavone	Н	Н	Н	OCH3	CH3	OCH3	CH3	
	Hyperoside	OH	OH	O-Galactose	OH	Н	OH	Н	

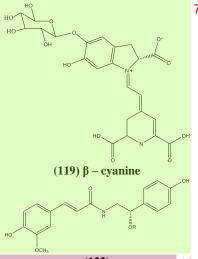


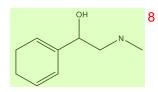


(116) Dopaxanthin 2

(117) Betanidin 5-O-[2"-O-(E)-feruloyl-  $\beta$  - 4 (1",2')- glucuronosyl- $\beta$ -glucoside]

(118) Betanidin- 5-O-[6'- O-(E)-feruloyl-  $\beta$ - 6 glucoside]





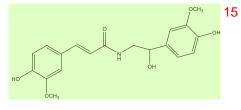
COOR 10

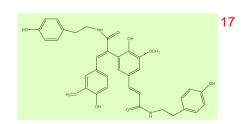
N
H
(121) 11

(120) 2-(dimethylamino)- 9 1-phenylethanol

3-(1*H*-inol-3-yl) propionic acid R=H 12 3-(1*H*-inol-3-yl) propionic acid methyl ester R=CH<sub>3</sub>

(122) 13 (2S, E)-N-[2-Hydroxy-2-(4hydroxyphenyl) ethyl] ferulamide R=H

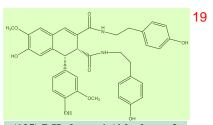


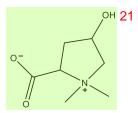


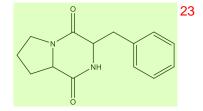
(E)-N-[2-(4-Hydroxyphenyl)-2- propoxyethyl] ferulamide R= n-Propyl

(123) (E)-N-[2-Hydroxy-2-(4-hydroxy-3methoxyphenyl)ethyl] ferulamide

(124) (E, E)-N, N-Dityramin-4,40 -dihydroxy- 18 3,50 - dimethoxy-b,30 -bicinnamamide





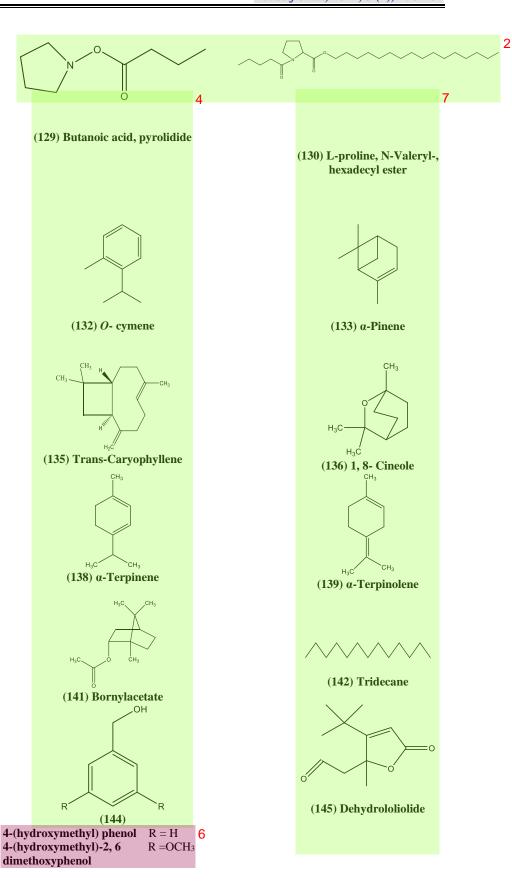


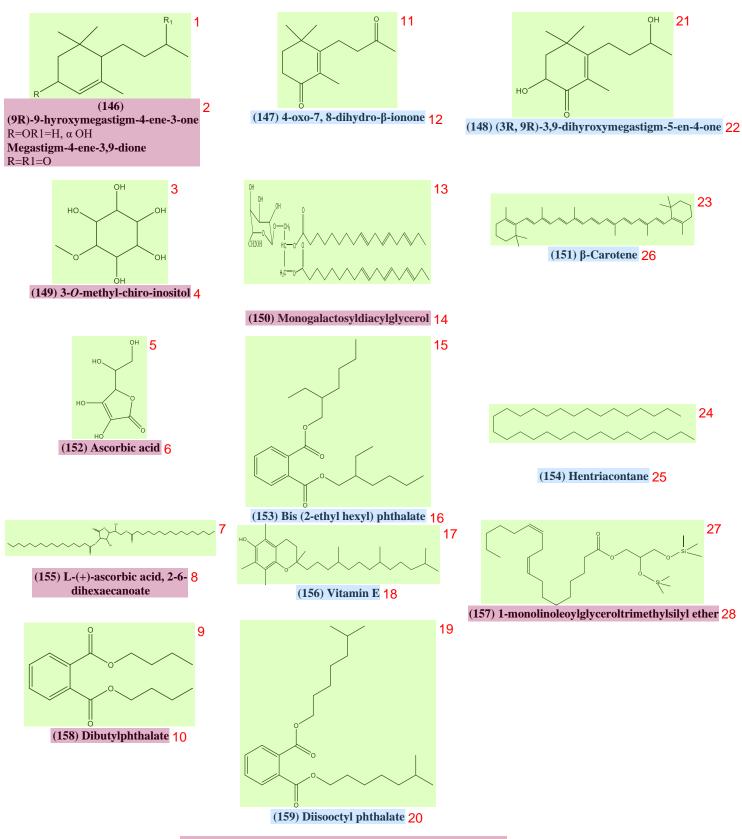
(125) 7-Hydroxy-1-(4-hydroxy-3methoxyphenyl)-N2, N3-bis(4hydroxyphenethyl)-6-methoxy-1,2dihydronaphthalene-2,3dicarboxamide

(126) Trans-4-hydroxy 22 prolinebtaine

(127) Pyrrolo[1,2-A] Pyrazin-1,4-dione, 24 hexahydro-3-(phenylmethyl)

(143) a- Humulene





Structures of isolated metabolites from family Aizozceae 29

*T. decandra* and *Sceletium spp.* was evaluated against 1 formaldehyde-induced arthritis in rats, and a significant inhibition of chemically-induced arthritis, indicates anti-inflammatory potential<sup>17-22-24-51</sup>.

Anti-inflammatory activity was evaluated in 2 and chronic models. Significant antiinflammatory activity was observed for chloroform extract of T. decandra in both carrageenan, dextran, and mediators induced edema models. The chloroform extract showed maximum inhibition of 58.36% at the dose of 200 mg/kg after 3 hrs of drug treatment in carrageenan-induced paw edema. The chloroform extract of T. decandra also exhibited significant antiinflammatory properties in dextran-induced paw edema model. Dextran-induced paw edema is known to be mediated both by histamine and serotonin. Dextran induces fluid accumulation, which contains little protein and few neutrophils, whereas carrageenan induces protein rich exudation containing large number of neutrophils. The extract effectively suppressed the inflammation produced by both carrageenan and dextran<sup>22</sup>.

# Antimicrobial activity 3

It has been reported that the methanolic extract 4 of M. nodiflorum, M. crystallinum, M. forsskaolii, and canariense showed a broad-spectrum antibacterial and antifungal activity against: Bacillus subtilis, K. pneumonia, S. aureus, S. pyogenes, E. coli, A. fumigatus, A. nigar, C. albicans, and Mucor spp<sup>48</sup>. For C. edulis five bioactive flavonoid compounds, rutin, neohesperidin, hyperoside, cactichin, and ferulic acid were isolated from the ethyl acetate fraction and individually or collectively were responsible for the antibacterial against 11 known human pathogenic bacteria, five Gram-positive: Staphylococcus epidermidis, Staphylococcus aureus, Bacillus subtilis, Streptococcus pneumoniae, Streptococcus pyogenes, and six Gram-negative: Pseudomonas aeruginosa, Haemophilus influenzae, Eschericha coli, Klebsiella pneumoniae, Acinetobacter baumanii, Moraxella catarrhalis <sup>34</sup>. Moreover, the isolated compounds from the ethanolic extract of Galenia Africana showed a remarkable antimycobacterial activity against M. smegmatis and M. tuberculosis<sup>55</sup>.

The extracts and the essential oil from the 5 fresh leaves of *S. portulacastrum* showed antibacterial, antifungal. The ethanolic extract showed potential antibacterial activity against the causative agents and pathogens related to various gastrointestinal disorders leading to indigestion, dysentery, and diarrhea<sup>15</sup>. Moreover, the ethanolic extract of the *S. portulacastrum* showed potential against the causative agents of nosocomial infections, *Staphylococcus aureus* and *E. coli*, while the essential oil exhibited notable antibacterial activity against both Gram-positive and

Gram-negative bacteria as well as significant 6 antifungal<sup>15</sup>. In addition, *S. portulacastrum*, showed positive activity against human immunodeficiency viruses<sup>15</sup>. Finally, it was reported that the new isolated flavonoid 2 - (3', 4' dihydroxy-phenyl) - 3, 5, 7-trihydroxy-chromen-4-one isolated from *T. decandra* showed antibacterial activity against *Pseudomonas aeruginosa* and by molecular docking it was found that FAS II β-hydroxyacyl-ACP (*FabZ*) of *P. aeruginosa* is a potential target of the isolated compound<sup>40</sup>. There are many other reported biological activities of members of family Aizoaceae as shown in **Table 11.** 

#### **CONCLUSIONS 7**

From this review, it can be deduced that the 8 major compounds of family Aizoaceae are alkaloids, triterpenes, flavonoids, sterols, lignans, fatty acids, phenolic acids, and essential oils. The review also showed that different extracts of aerial parts of plants of family Aizoaceae posse diverse biological activities as anti-acetylcholinesterase, butyrylcholinesterase, anti-infective, antihyperlipidemic, antipyretic, antifertility, diuretic, nephroprotective and others. It would, therefore, be important extensively investigate to their phytochemicals and pharmacologically determine their activities for future drug discovery and development.

#### Conflict of interest 9

The authors declare that there is no conflict of 10 interest regarding the publication of this paper.

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