

REVIEW

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Plant of the Millennium, Caper (*Capparis spinosa* L.), chemical composition and medicinal uses

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Abstract

Background: Caper (*Capparis spinosa* L.) is a common member of the genus *Capparis*, which is a perennial shrub and thorny, and a common aromatic plant in many parts of the world, especially the Mediterranean regions.

Main body: The aim of this mini-review was to outline the most significant health benefits of caper in both traditional and modern pharmaceutical medicine. Scientific databases such as PubMed, Science Direct, Scopus, Research Gate, and Google Scholar with emphasis on Science Direct and Scopus have been used. A review of literature was carried out using the keywords caper, *Capparis spinosa*, health benefits, pharmaceutical benefits, natural products, and caper bush. During the writing of the review, the time period in which the papers were published had not been selected since the focus was on significant researched selected for the areas covered in this mini-review. The main components of its aerial parts are cappariolide A, stachydrin, hypoxanthine, uracil, capparine A, capparine B, flazin, guanosine, 1*H*-indole-3-carboxaldehyde, 4-hydroxy-1*H*-indole-3-carboxaldehyde, kaempferol, thevetiaflavone, tetrahydroquinoline, rutin, kaempferol-3-glucoside, kaempferol-3-rutinoside, kaempferol-3-rhamnorutinoside, isorhamnetin 3-*O*-rutinoside, quercetin 3-*O*-glucoside, ginkgetin, isoginkgetin, sakuranetin and glucocapparin in aerial parts. The main components of root are capparispin, cadabicine 26-*O*-β-D-glucoside, capparispin 26-*O*-β-D-glucoside, and stachydrine, seeds contain glucocapparin. Traditional application of caper is for treatment of headache, fever, convulsions, diabetes, toothache, menstruation, skin disease, kidney disease, liver disease, rheumatism, ulcers, hemorrhoids and sciatica. Fruit and leaves have anti-diabetic effects, fruits have anti-obesity, cholesterol-lowering and anti-hypertensive effects, roots, fruits, stem barks and shoots have antimicrobial effects, leaves, roots and fruits contain anti-inflammatory activity, and aerial parts have antihepatotoxic effects.

Conclusion: On the basis of phytochemical advantages and pharmacological benefits, caper shows its importance as one of the most notable medicinal plant for prevention and treatment of various diseases, however, more researches are need on the usage of caper, especially in modern pharmaceutical science.

Keywords: Caper, Traditional medicine, Pharmaceutical benefits, *Capparis spinosa*, Rutin

Background

Medicinal plants knowledge has its origins in ancient civilizations, and cultures, and its main aim is to return the body to a state of natural balance with its active ingredients (Khoshkharam et al. 2021; Shahrajabian et al. 2021a,b). The main medication functions of medicinal herbs in traditional herbal sciences of countries like Iran, China, India and Japan have been received notable attentions in recent years (Marmitt

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and Shahrajabian 2021; Marmitt et al. 2021; Shahrajabian et al. 2021c,d). Traditional medicine, based largely on herbs, still supports the primary healthcare of more people worldwide than conventional or western medicine (Sun et al., 2021a,b), and they are culturally acceptable and readily available even in modern era (Shahrajabian et al. 2020a,b).

Capparis spinosa L. is a perennial plant typical of the Mediterranean flora and a multipurpose plant used for curing various human ailments (Panico et al. 2005), and its fresh and pickled flower buds were the most studied parts of the plant (Allaith 2016). Although, it is native to the Mediterranean but well grows in Italy, Northern Africa, Greece, Central Asia, Iran, and other parts of the world (Zarei et al. 2021). In traditional medicine and indigenous knowledge, roots are used as diuretic, astringent, and tonic; bark root used as appetizer, astringent, tonic, antidiarrheic and to treat hemorrhoids, and spleen disease; and bark is also used for gout and rheumatism, as expectorant, and for chest diseases. Caper favours a rainy spring and a dry, hot summer with intense sunlight where temperature exceeds 40 °C and average rainfall is 350 mm during spring and winter season (Barbera and Lorenzo 1984). *Capparis spinosa* is called the Plant of the Millennium, and it has highly diverse economic and medicinal value in different system of medicines like in Iranian, Unani, Chinese, Ayurvedic and Greco-Arabi system of medicines (Sher and Alyemeni 2010). The antioxidant activity of various parts of caper was reported from different scholars (Bonita et al. 2002; Yue-lan et al. 2010; Aliyazicioglu et al. 2013; Mazandarani et al. 2014; Kalantari et al. 2018). Caper seeds declared an important source of antioxidant molecules for the food and pharmaceutical industries as they are rich in phenolic compounds and show high antioxidant activity (Tlili et al. 2015). Caper seed is an inexpensive source of omega-6, and its seed oil is regarded as oleic-linoleic oil (Dursun and Dursun 2005; Mahdavi Ara et al. 2014). The consolidated traditional use of its root as remedy against different pains in human is well known since the antiquity (Khatib et al., 2016). *Capparis spinosa* contains bioactive lipids, glucosinolates and flavonoids, and seed oil is rich in unsaturated and rare lipids such as *cis*-vaccenic acid, and the main glucosinolate is glucocapperin, and the main flavonoid is rutin (Sharaf et al. 2000; Argentieri et al. 2012). The most important identified compounds are quercetin, kaempferol and isorhamnetin derivatives in addition to myricetin, eriodictyol, cirsimaritin and galocatechin derivatives (Bakr and El Bishbishy 2016). This article is aimed to have a brief review on pharmaceutical benefits of caper in both traditional

and modern medicinal sciences with considering its chemical constituents.

Main text

Agronomic traits

The *Capparis spinosa* L. is an important medicinal plant in the family Capparaceae, the complete cp genome of *Capparis spinosa* L. has 157,728 bp in length and contains 136 genes, and the plastome contained 80 protein coding genes, 31 tRNA genes and 4 rRNA genes (Alzahrani et al. 2021). This ancient plant is sensitive to salt stress at its primary growth phase, and applying biostimulant may increase plant biomass, proline, soil enzymes activity, increased rutin bioflavonoid of caper, and increased the phenol content of caper (Sadeghi and Taban 2021; Shahrajabian et al. 2021e,f). Caper fruits were characterized to have the exocarp green in all stages of development, and there was a decrease in the protein content with the development of the fruit, while the fruits presented high contents of total phenols, flavonoids and flavanols (Grimaly et al. 2018). The plant is a perennial winter deciduous species that bears founded, fleshy leaves and large white to pinkish flowers, it is a shrub with 30–50 cm height; the caper fruits names as cape berry and it is ellipsoid or ovate shape, and dry heat and intense sunlight make the preferred environment for caper plants. Quaternary ammonium compounds (QACs), as constituents of Capparaceae, play important roles in protecting against abiotic stress which can be found in both roots and leaves (Al-Tamimi et al. 2019). Fertilization induced an increase in productivity and nutrient tissue concentrations of both N and P, and increased K and Ca uptake to maintain the hydric balance, thorny stems and a heavy investment in chemical defenses to prevent grazing, and a deep mycorrhizal root system allow *Capparis* to grown successfully in very infertile soils and to endure environmental stress (Pugnaire and Esteban 1991). Seeds soaked in sulphuric acid and in gibberellins solution, and then dried at room temperature to their original air-dry weight, germinated as well as non-dried seeds (Orphanos 1983). An analysis of caper variability in Tunisia using morphological descriptors revealed intra-specific diversity; two recognized morphotypes were (1) prickly and one that included the inermis and downy caper, and (2) the second one is characterized by creeping shoots and relatively large leaves, low stomatal density, and high numbers of stamens, and the heterogeneous prickly group is characterized by erect shoots and relatively small leaves, high stomatal density, and a small numbers of stamens (Saadaoui et al. 2009). The salient features of the growth mechanism of *Capparis* demonstrate a sophisticated plant response to drought, involving osmotic adjustment, regulated stomatal opening, cell wall properties and

increased root density (Rhizopoulou 1990). Deep roots, with high tip water potentials, may sustain high plant water potentials (Rhizopoulou and Kapolas 2015). Stomata were found on the abaxial surface of sepals, and on both the adaxial and abaxial surfaces of petals, filaments and style were densely packed with small cells, exhibiting an increased density of cell wall material that provides strength, and petals possessed vacuolated parenchyma cells with large intercellular space (Rhizopoulou et al. 2006).

Chemical constituents

It has been reported that *Capparis spinosa* is a rich source, not only of sulphur compounds, but also with phenolic and flavonoid glycosides contributing to its powerful cytotoxic activity (Bakr and El Bishbishy 2016). Berries of *Capparis spinosa* possess carbohydrate content (5%), dietary (3%), protein (2%), and lipids (0.9%), and it contains moderate content of vitamin C (4 mg/100 g fw) (Allaith 2016). Qualitative phytochemical analysis of *Capparis spinosa* (aerial parts) extract, showed, alkaloids, glycosides, carbohydrates, tannins, phenols, and triterpenoids in ethanolic extract, and alkaloids, steroids, carbohydrates, flavonoids, tannins, phenols, and saponins in aqueous extract (Mustafa 2011). Tlili et al. (2011) reported that phytochemicals studies have shown the presence of many beneficial compounds such as spermidine, rutin, quercetin, kaempferol, stigmasterol, campesterol, tocopherols, and carotenoids. Glucocapparin is the main component in extracts of non-fermented berries and is fully degraded upon the fermentation process, and epicatechin amounts are lower and free quercetin is observed after the fermentation (Jimenez-Lopez et al. 2018). Linoleic and oleic acids were established as major fatty acids of seeds oils of caper seed (*Capparis spinosa*) (Ozcan et al. 2012). Elemental analysis values of *Capparis spinosa* L. using EDXRF system are Al ($0.48 \pm 0.05\%$), P ($1.15 \pm 0.01\%$), S ($4.00 \pm 0.06\%$), K ($4.54 \pm 0.03\%$), Ca ($1.18 \pm 0.01\%$), Cl (94.86 ± 25.51 ppm), Ti (55.24 ± 2.30 ppm), Mn (70.04 ± 1.00 ppm), Fe (520.72 ± 4.05 ppm), Ni (24.10 ± 0.05 ppm), Cu (88.27 ± 0.45 ppm), Zn (250.75 ± 0.80 ppm), Br (11.92 ± 0.07 ppm), Rb (79.03 ± 0.19 ppm), Sr (40.20 ± 0.69 ppm), and Pb (5.34 ± 0.13 ppm) (Aliyazicioglu et al. 2013). Constituents of the volatile oils of the leaf, ripe fruit and root of *Capparis spinosa* var. *mucronifolia* growing wild in Isfahan (Iran) were studied by TLC, GC and GC-MS methods, where the leaf oil was composed of isothiocyanates, *n*-alkanes, terpenoids, a phenyl propanoid, an aldehyde and a fatty acid, and the main components of the oil were thymol (26.4%), isopropyl isothiocyanate (11%), 2-hexenal (10.2%), and butyl isothiocyanate (6.3%), and the

volatile oils of the ripe fruit and the root were composed mainly of the methyl, isopropyl and *sec*-butyl isothiocyanates (Afsharypuor et al. 1998). *p*-Methoxy benzoic acid isolated from the methanolic soluble fraction of the aqueous extract of *Capparis spinosa* was found to possess significant antihepatotoxic activity against carbon-tetrachloride and paracetamol induced hepatotoxicity in vivo and thioacetamide and galactosamine induced hepatotoxicity in isolated rat hepatocytes using in vitro technique (Gadgoli and Mishra 1999). The main phytoconstituents of *Capparis spinosa* are, flazin, guanosine, capparine A, capparine B, 1H-indole-3-carboxaldehyde, 4-hydroxy-1H-indole-3-carboxaldehyde, chrysoeriol, apigenin, kaempferol, thevetia-flavone, 5-hydroxymethylfuraldehyde, vanillic acid and cinnamic acid in fruits; cappariside or 4-hydroxy-5-methylfuran-3-carboxylic acid, 5-hydroxymethylfurfural, 5-hydroxymethyl furoic acid, and 2-furoic acid in whole plant; protocatechuic aldehyde, E-butenedioic acid, ethyl 3,4-dihydroxybenzoate, syringic acid, protocatechuic acid, vanillic acid, succinic acid, 4-hydroxybenzoic acid and α -tocopherol in fruits; rutin in aerial parts such as leaves, buds and flowers, and homogalacturonan in fruits (Moufid et al. 2015). Investigation of *Capparis spinosa* of Jordanian origin lead to isolation of two new compounds *g*-sitosterylglucoside-6'-octadecanoate (1) and 3-menthyl-2-butenyl-*g*-glucoside (2) (Khanfar et al. 2003). Three extraction methods, involving maceration, reflux, and ultrasonic extractions were used to evaluate the phytochemical content and antioxidant activity of leaf extracts from *Capparis spinosa* L. collected from five localities in South Tunisia; the maceration extraction method exhibited the highest antioxidant activities using the reducing power assay, the azinobis (3-ethylbenzothiazoline)-6-sulfonic acid (ABTS⁺), and the 2,2-diphenyl-1-picrylhydrazyl (DPPH^o) radical scavenging activity; also, the major identified phenolic acids were quinic acid, gallic acid, and protocatechuic acid, and catechin was the most abundant flavonoid in all detected extracts (Yahia et al. 2020). HPLC analysis showed high levels of rutin and quercetin in leaves of *Capparis*, rutin was $16,939.2 \pm 0.01$ and quercetin was 908.93 ± 0.01 μ g/g fresh weight, and in fruit, 1019.52 ± 0.01 rutin and 97.86 ± 0.01 μ g/g FW quercetin were measured (Mohebbali et al. 2018). Three new alkaloids, capparisine A (1), capparisine B (2), capparisine C (3), and two known alkaloids, 2-(5-hydroxymethyl-2-formylpyrrol-1-yl) propionic acid lactone (4), and N-(3'-maleimidyl)-5-hydroxymethyl-2-pyrrole formaldehyde (5) were isolated from the fruits of *C. spinosa* L. (Yang et al. 2010). In Algeria, the result of chromatographic analysis (CG/MS) leads to the identification of 33 components; palmitic acid (38.19%), nonanal-n (12.61%), cymene-2,5-dimethoxy-para (8.94%), and octacosane

(5.49%) which were the major components of essential oil (Benachour et al. 2020). Aliphatic acids and aldehydes are the most abundant chemical classes in samples from Pantelleria and Salina Islands in Italy, and among sulfur compound, methyl-isothiocyanate is the major one, followed by benzyl-isothiocyanate (Concurso et al. 2016). In one experiment in Turkey, the major volatile compounds found in caper bud oil were benzyl alcohol (20.4%), furfural (7.4%), ethanol methyl pentyl acetal (5.9%), 4-vinyl guaiacol (5.3%), thymol (5.1%), octanoic acid (4.8%), and methyl isothiocyanate (4.5%), while the major volatile compounds found in caper leaves were methyl isothiocyanate (20.0%), thymol (15.5%), 4-vinyl guaiacol (4.3%), hexyl acetate (3.6%), and trans-theaspine (2.6%) (El-Ghorab et al. 2007). The higher phenolic content was determined in the fresh (1843.71 mg/100 g DW), and fermented buds (1198.54–1539.49 mg/100 g DW) rather than the berries (29.72–40.75 mg/100 g DW); quercetin-3-O-rutinoside, kaempferol-3-O-rutinoside, and quercetin-O-gallolyl-O-hexoside were the principal phenolic components in fresh and fermented buds while quercetin-3-O-rutinoside in fresh and fermented berries (Aksay et al. 2021). The occurrence of glucobrassicin, neoglucobrassicin and 4-methoxy-glucobrassicin in roots of *Capparis spinosa* is demonstrated by HPLC and mass spectral methods (Schraudolf 1989). Francesca et al. (2016) reported that high-performance liquid chromatography/electrospray ionization source/mass spectrometry (HPLC–ESI–MS) successfully identified 9 polyphenols, epicatechin was found in untreated fruits, and quercetin in processed caper berries. Three new alkaloids capparispine (1), capparispine 26-O- β -D-glucoside (2), and cadabicine 26-O- β -D-glucoside hydrochloride (3) were isolated from the roots of *C. spinosa* in Xinjiang Uighur Autonomous Region, China (Fu et al. 2008). Quercetin content in different parts of caper are 1.7 mg/g in root, 5 mg/g in stem, 7.92 mg/g in the leaf, 10 mg/g in floral bud, 12.8 mg/g in flower, 9.6 mg/g in fruit and 6.2 mg/g in the seed (Moghaddasian et al. 2012).

Traditional medicinal benefits

Caper is a traditionally used medicinal plant and widely studied for its biological properties. Moroccan sample showed the highest phenolic content across all extraction types followed by Italian and Turkish (Stefanucci et al. 2018). A traditional Persian medicine formulation for diabetes mellitus are *Capparis spinosa*, *Rosa canina*, *Securidaca securigera*, *Silybum marianum*, *Urtica dioica*, *Trigonella foenum-graecum*, and *Vaccinium arctostaphyls* with no notable hepatic, renal and gastrointestinal side effects (Mehrzadi et al. 2021). In Greek popular medicine, a herbal tea made of caper root and young shoots is considered to be beneficial against rheumatism (Mohammad

et al. 2012). Many parts of caper are still being used as drugs and traditional healers in Saudi Arabia describe oral administration of dried fruits of *Capparis spinosa* L. with water to treat hypertension and diabetic complication (Sher and Alyemeni 2010). In Bahrain, fresh caper berries are still eaten by the elderly, whereas in Eastern India, fresh raw caper berries are consumed as an appetizer. Seed clumps of *Capparis spinosa* L. together with shoots, leaves and fruits has been used in eastern part of Central Asia for medicinal purposes (Jiang et al. 2007). A decoction of *Capparis spinosa* L. roots, widely used in traditional folk medicine of southern Italy, and heterocyclic compounds were also recovered from the chloroformic extract of the roots, and it showed an interesting bacteriostatic activity on the growth of *Deinococcus radiophilus* (Boga et al. 2011). The antioxidant, nephroprotective and hepatoprotective effects of methanolic extract of its leaves associated with its phytochemical content, and nine compounds namely rutin, resveratrol, coumarin, epicatechin, luteolin, catechin, kaempferol, vanillic acid and gallic acid are more responsible in support of traditional usage of *Capparis spinosa* to cure kidney and liver diseases (Tlili et al. 2017). In traditional medicinal science, the whole plant was used for rheumatism, roots were used as diuretic, astringent, and tonic, bark root, which has bitter taste, was used as appetizer, astringent, tonic, anti-diarrheic and to treat hemorrhoids and spleen diseases, infusion of stems and root bark were used as anti-diarrheic and febrifuge, fresh fruits were used in sciatica and dropsy, and dried and powdered fruit combined with honey was used in colds, rheumatism, gout, sciatica and backache (Rahnacard and Razavi 2017).

Modern pharmaceutical benefits

Capparis spinosa L. has the potential to down regulate inflammation-involved genes in Alzheimer's disease (AD), due to its high levels of flavonoids and could be beneficial as a dietary complement in AD patients (Mohebbi et al. 2018). *Capparis* species specially *C. spinosa* and *C. decidua* are cultivated commercially and being used in food industries, and *Capparis* species are rich sources of antioxidant and bioactive compounds being responsible for various biological activities (Gull et al. 2015). It has been reported that the methanol extract of *Capparis* (MEC) has antinociceptive effects both at the peripheral and central levels (Arslan et al. 2010). Phytochemical analysis shows that *Capparis spinosa* has high quantities of bioactive constituents, including polyphenolic compounds, which are responsible for its health-promoting effects. The aqueous extract of *Capparis spinosa* L. (20 mg/kg) exhibits a potent lipid lowering activity in both normal and severe hyperglycemic rats after repeated oral administration of its aqueous extract

(Eddouks et al. 2005). The total antioxidant capacity assessed using the molybdate assay ranged between 99.54 ± 0.90 mg AAE (ascorbic acid equivalent)/g in *Capparis spinosa* pollen (Bakour et al. 2020). Synthesis of CuO nanoparticles from *Capparis spinosa* is reported (Samari et al. 2019). The methanolic extract of *C. spinosa* L. demonstrated anti-quorum sensing (QS) and anti-biofilm activity at 0.5–2 mg/mL (Abraham et al. 2011). The antioxidant effect of leaf extract of *Capparis spinosa* is six times greater than fruit, and the use of leaf extract or its active metabolites in the sperm culture medium may be beneficial for maintaining motility, vitality and sperm DNA (Khojasteh Rad et al. 2021). Antioxidant capacities strongly correlated with the total free phenolics, total flavonoids, and total carotenoids (Allaith 2016). The *Capparis spinosa* leaves can be used as a potential, low-cost source of polysaccharide, and the polysaccharides of leaves exhibited significant antioxidant activity; furthermore, a much more antimicrobial activity using the polysaccharide against Gram-negative bacteria (*Escherichia coli*, *Shigella dysenteriae* and *Salmonella typhi*) was found than Gram-positive bacteria (*Bacillus panis* and *Staphylococcus aureus*) (Mazarei et al. 2017). Methyl isothiocyanate was detected as major volatile compounds in caper essential oil, rutin and chlorogenic acid were detected as dominant compounds in caper aqueous infusion which induced to high-inhibitory effect of oil and infusion on HT-29 cell proliferation and NF- κ B activation also caper oil and infusion blocked the cell cycle at G₂/M phase which suggest that both volatile and non-volatile compounds of caper potentially can play an important role in colon cancer prevention (Kulisic-Bilusic et al. 2012). Biodiesel was produced from novel non-edible caper seed oil, and biodiesel showed excellent potential as a sustainable and renewable source of bioenergy (Munir et al. 2021). *Capparis spinosa* reduced brain inflammation, improved LPS-induced cognitive impairment, and by in vivo, *C. spinosa* polarized inflammatory microglial cells towards M2, and by in vitro assay, *C. spinosa* directly shifts lipopolysaccharide (LPS)-induced M1/M2 imbalanced towards M2 cells (Rahimi et al. 2020). The anti-inflammatory, antiarthritic, and anti-thrombotic activities of methanol extract of *Capparis* buds, leaves, stems and fruits has been reported in other studies (Bektas et al. 2012; Ozgun-Acar et al. 2016; Twumasi et al. 2019). Caper extract at 0.1 g kg⁻¹ feed proved to be a potential immunostimulant for fish (Bilen et al. 2016). Eddouks et al. (2004) reported that the oral administration of the aqueous extract of *Capparis spinosa* L. fruit (20 mg/kg) produced a significant decrease on blood glucose levels in streptozotocin (STZ) diabetic rats; the blood glucose levels were nearly normalized 2 weeks after daily repeated oral administration of it, which can

be selected as a potent anti-hyperglycaemic activity in STZ rats without affecting basal plasma insulin concentration. It has been reported that small caper berries had higher, crude protein, phenolic compounds and flavonoids than the big one (El Amri et al. 2019). Biochemical analysis showed nephroprotective and hepatoprotective effects of methanol extracts of *Capparis spinosa* seeds, and histopathological study suggested that the extracts protect against tissues fibrosis (Tir et al. 2019). The regular administration of *Capparis spinosa* leaf or buds normalized all the biochemical parameters and reversed the liver/kidney injury with variable degrees of organ protection, because of antidiabetic and antihyperlipidemic effects of it (Mollica et al. 2017). The putative mechanisms involved in the antihyperglycemic effects of *Capparis spinosa* include reducing carbohydrate absorption from the small intestine, inhibiting gluconeogenesis in the liver, enhancing glucose uptake by tissues, and beta cell protection/regeneration (Vahid et al. 2017). Antioxidant capacities of caper berries strongly correlated with the total free phenolics, total flavonoids, and total carotenoids. *Capparis spinosa* extracts relieved pain related to rheumatoid arthritis and osteoarthritis after single administration (Maresca et al. 2016). *Capparis spinosa* improved the circulating levels of triglyceride and cholesterol, and direct inhibition of gluconeogenesis in liver maybe a probable mechanism of action of this plant, and its administration might be a beneficial therapeutic approach for metabolic syndrome and fatty liver (Jalali et al. 2016). Systematic use of *Capparis spinosa* extract may accelerate bone formation at the expanded suture in rats (Erdogan et al. 2015). Caper fruit extract may be a safe anti-hyperglycemic and anti-hypertriglyceridemic agent for type 2 diabetic patients, with no liver, kidney and other side effects (Fallah Huseini et al. 2013). The ethanolic root bark extract of *C. spinosa* could afford significant dose-dependent protection against CCl₄ induced hepatocellular injury (Aghel et al. 2007). The hydro-ethanolic extract of *Capparis spinosa* (HECS) significantly decreased hepatic phosphoenolpyruvate carboxykinase, increased acetyl CoA carboxylase and non-significantly decreased hepatocyte nuclear factor-4 α (HNF-4 α) as a transactivator of phosphoenolpyruvate carboxykinase (PEPCK) at mRNA expression level in diabetic rats (Assadi et al. 2021). Manikandaselvi et al. (2018) reported the strong antioxidant activity in aqueous extract centered on the outcomes of DPPH radical scavenging (IC-50 1.74 μ g/ml), and inhibition of lipid peroxidation (IC-50 4.01 μ g/ml) assays; and the aqueous extract exhibited anti-inflammatory activity in terms of inhibition of protein denaturation (IC-50 8.01 μ g/ml), inhibition of protease activity (14.24%), and RBC membrane stabilization (IC-50 226.69 μ ml), and also possess antibacterial

activity against *Bacillus cereus*, *Staphylococcus aureus*, *Campylobacter jejuni*, and *Salmonella enteritidis* strains. *Capparis spinosa* fruit extract could be aid prevention of damage to the tissues due to the decreased levels of harmful oxidants in the body in diabetes (Taghavi et al. 2014).

Conclusion

Natural products such as medicinal and aromatic plants have been used as traditional remedies because of their amazing healing characteristics. Caper (*Capparis spinosa* L.) is one of the most important medicinal plant in Iranian, Chinese, Unani, Ayurvedic and Greco-Arabi medicinal systems. The phytochemicals identified in this plant were lipids, flavonoids, alkaloids, saponins, tannins, lignins, glucocapperin, (6S)-hydroxy-3-oxo- α -ionolglucosides and polyphenols. *Capparis spinosa* has anthelmintic, cytotoxic, anti-inflammatory, anti-arthritic, anti-oxidant, anti-microbial, cardiovascular, chondroprotective, anti-diabetic, hypolipidemic, anti-allergic, anti-histaminic, immunomodulatory, anti-carcinogenic and anti-hepatotoxic activities. Caper can be recommended for more clinical experiments to evaluate its clinical efficacy and safety in modern pharmaceutical science by considering its chemical constituents.

Abbreviations

QACs: Quaternary ammonium compounds; ABTS⁺: The azinobis (3-ethylbenzothiazoline)-6-sulfonic acid; DPPH[•]: The 2,2-diphenyl-1-picrylhydrazyle; HPLC: High-performance liquid chromatography; CG/MS: Chromatographic analysis; HPLC-ESI-MS: The high-performance liquid chromatography/electrospray ionization source/mass spectrometry; AD: Alzheimer's disease; MEC: Methanol extract of *Capparis*; AAE: Ascorbic acid equivalent; QS: Anti-quorum; LPS: Lipopolysaccharide; STZ: Streptozotocin; HECS: Streptozotocin (STZ), the hydro-ethanolic extract of *Capparis spinosa*; HNF-4 α : Hepatocyte nuclear factor-4 α ; PEPCK: Transactivator of phosphoenolpyruvate carboxykinase.

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Authors' contribution

M.H.S.: Writing-original draft preparation; W.S.: Writing-original draft preparation; Q.C.: Writing-review and editing. All authors have read and approved to the manuscript.

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Declarations

Ethics approval and consent to participate

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Consent for publication

Not applicable.

Competing interests

The authors declare that they have no potential conflicts of interest.

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