

A Review: Potential Pharmacological Uses of Natural Products from Laminaceae

K. Gayathiri*, M. Sangeetha, V. K. Sharanya, G. Shyam Prakash, R. Vimalavathini, J. Gopi Sudheer Kumar, S. Kavimani

Department of Pharmacology, College of Pharmacy, Mother Theresa Post Graduate and Research Institute of Health Sciences, Indira Nagar, Gorimedu, Puducherry – 605 006, India.

ABSTRACT

Natural products and their derivatives have been used as therapeutic agents from ancient times and in folklore for treatment of many diseases and illness. Many of the natural products from plants have been used as current drug candidates, either in their original or semi synthetic form and secondary metabolites of the plants also be serve as drug precursors, drug prototypes and pharmacological probes. Medicinal plants involve a multifaceted approach in current drug research as combining botanical, phytochemical, biological and molecular techniques. In drug discovery medicinal plants provides continually new and important lead constituents to against various pharmacological targets including cancer, HIV/AIDS, Alzheimer's, Malaria, Pain and Immune alteration and etc. Although, medicinal plants continue provide a new drug leads in drug discovery, and numerous challenges are encountered in procurement and selection of plant materials, screening method and the scale up of active compound, hence this brief review work presents a study of the importance of natural products, especially those derived from higher plants and aims to the highlight the pharmacological significant of Laminaceae family in terms of drug development. Most of the plants in laminaceae family are aromatic plants, they posseses Immunomodulatory activites. Salvia species is one of species in this family, hence aimed to studied the immunomodlatry activity of one of the salvia species.

Keywords: Drug discovery, laiminace, medicinal plants, natural products, pharmacological uses

Received 6 April 2016

Received in revised form 20 April 2016

Accepted 22 April 2016

*Address for correspondence:

K. Gayathiri,

Department of Pharmacology, College of Pharmacy, Mother Theresa Post Graduate and Research Institute of Health Sciences, Indira Nagar, Gorimedu, Puducherry – 605 006, India.

E-mail: gayukrish13@gmail.com

1. INTRODUCTION

Medicinal plants need more attention due to their important role in primary healthcare delivery system for improvement of people's health [1]. For thousands of years, natural products have been the most successful soureces of potential drug leads and played an important role in development of new drug leads against various pharmacological targets including cancer, HIV/AIDS, Alzheimer's, malaria, and pain and the natural sources are using to treat and preventing the various human diseases throughout the world [2]. Natural products are the richest source of biologically active compounds, obtained from plants, animals, minerals etc., about 40% of all medicines were obtained from either natural products or their semi-synthetic derivatives and developed a lead

compound originally obtained from a natural source [3, 4]. Indeed, over the last half century, most clinical applications of secondary metabolites of plants and their derivatives have been applied towards combating several human diseases like cancer and etc.

The importance of natural products is clearly enormous and many higher plants contain novel metabolites with different biological properties, about 25% of the drugs from plants were prescribed worldwide and 121 active compounds being in current use [5]. Natural products research continues to explore a variety of lead structures, which may be used as templates for the development of new drugs by the pharmaceutical industry and will be, important sources of new pharmaceutical

compounds [6]. Several natural product drugs of plant origin have either recently been introduced to the United States market, including *arteether*, *galantamine*, *nitisinone*, and *tiotropium*, or are currently involved in late phase clinical trials.

In the 20th century, revolutionized thinking of pharmaceutical companies and drug development of organic chemistry, in the use of drugs as the receptor theory of drug action, it concluded by the scientists that the some mystical "power of life" are the factors required for the biological activity of the drug rather than individual chemical compounds in natural extracts. This led to the beginning of a totally new era in pharmacology, as pure, isolated chemicals and structural modification to produce potentially more active and safe drug, instead of extracts. Indeed, many bioactive compounds and their chemical structures were elucidated and the economic power of the pharmaceutical companies was increasing [4]. Recently, natural product research has been renewed due to the failure of alternative drug discovery methods to deliver many lead compounds in key therapeutic areas such as immunosuppression, anti-infectives, and metabolic diseases and there has been growing interest in alternative therapies and the therapeutic use of natural products, especially those derived from plants. This interest in drugs of plant origin is due to several reasons, namely, conventional medicine can be inefficient (e.g. side effects and ineffective therapy), abusive and/or incorrect use of synthetic drugs results in side effects and other problems, a large percentage of the world's population does not have access to conventional pharmacological treatment, and folk medicine and ecological awareness suggest that "natural" products are harmless [7].

Nowadays, secondary metabolites of plant such as alkaloids, flavonoids, tannins and other phenolic compounds etc., are the important sources of medicines, especially in developing countries still used plant based traditional medicine for their health care system and there is a tremendous increase in the consumption of herbs as an alternate source of medicine to maintain health and improve the quality of life [8, 9].

In industrialized countries, plant based traditional medicines (Phytotherapeutics are often called as complementary or alternative medicines), and their use has increased steadily over the last 10 years [10]. Based on the analysis, 88 single chemical entities isolated from 72 medicinal plants have been introduced into the modern therapy, many of which have similar therapeutic use and some of these plants derived compounds such as atropine (anticholinergic), codeine (cough suppressant), colchicine (antigout), ephedrine (bronchodilator), morphine (analgesic), pilocarpine (parasympathomimetic), and physostigmine (cholinesterase inhibitor) are still being used widely as single agent or combination formulations in prescription drugs. Most of the natural plant sources are obtained from different families one of this family Lamiaceae is the most important source for lead compound to treat and prevent various diseases [8]. Hence the present work was concentrated on to reveal the pharmacological activities of the Lamiaceae family and their involvement in new drug research in present scenario.

2. Role of Lamiaceae in History of Drug Discovery

The Lamiaceae or Labiatae plant family is one of the largest families among the dicotyledons; it is also termed as mint family, with 236 genera and more than 7,000 species of the order Lamiales. The largest genera of this family are Salvia (900), Scutellaria (360), Stachys (300), Plectranthus (300), Hyptis (280), Teucrium (250), Vitex (250), Thymus (220) and Nepeta (200). [11]. Many plants of this family have been shown to impart an antioxidative effect in foods [12] and rich in terpenoids such as monoterpenes, diterpenes, triterpenes and also sesquiterpenoids, along with flavonoids and other phenolics [13] having a number of biological activities, including antimicrobial, anti-inflammatory, antioxidant, antiviral, cytotoxic, wound healing, neuroprotective, and anticholinesterase [14].

Plants in this family also have an important value in the sociocultural, spiritual and medicinal use in the rural and tribal lives of the developing countries [15]. Many active

essentials oils also obtained from this family and also famous for the presence of diterpenoids among its members. Lamiaceae family species are important for antimicrobial properties which are used in research of antimicrobial activities, for instance, *Salvia argenteneae* L, *Stachysannua* L, *Ballotanigra* L, *Melisa officinalis* L among others [16]. Aromatics plants and species have great importance for food, cosmetics and pharmaceutical industries [17]. Naturally growing Lamiaceae members have been used as tea, spice and for various medicinal purposes such as to treat fever, cough, headaches, wound healing heart diseases and stomach-aches [18]. Among this laminaceae family *Salvia* species are widespread plants in many countries. *Salvia* genera plants commonly as sage. Some of the specials are *S. sclarea*, [19] *S. pratensis* [20], *S. glutinosa*, *S. Officinalis*, *Salvia lavandulaefolia* and *Salvia miltiorrhiza*, these plants are cultivated mostly for its aromatic and widely used as essential oil. A study of the antibacterial activities of the essential oil of *S. officinalis* proved that sage essential oil in higher concentration exhibited a better efficiency than antibiotics [21]. Species belonging to the family Lamiaceae (Labiatae) are important sources of phenolic-type food preservatives and pharmaceuticals. This family also known as mint family due to many of the various herbaceous plants comprising under the genus *Mentha*, a taxon of about 25 species of aromatic, almost exclusively perennial plants in the mint family Lamiaceae. The main chemical constituents of this family are Eugenol, nerol, eugenol methyl ether, caryophyllene, terpinene-4-ol-decylaldehyde, α -selinene, α and α -pinene, camphor and carvacrol, cineole, linalool, omega 3 and 6 fatty acids, dietary fibers, proteins, quercetin because of these constituents they play a vital role in preventing and treating various diseases such as hyperlipidemic, diuretic, anti-inflammatory activity, antimicrobial activity, antiviral, antifungal, immunomodulating activity etc.

3. Role of Laminaceae in Current Drug Research

Developing the therapeutic agents for the future, clearly involve the basic sciences

disciplines that have always been at the core of drug discovery, namely, structural biology to provide information about the target biomacromolecules; chemistry to design and synthesize the drug candidates; and pharmacology to determine the effects of the interaction between drug and target. Taking drug discovery to the next level may require an entirely new approach, but more likely will instead result from the introduction of new disciplines and/or greatly improved technologies into the process seems likely that integrating sophisticated new computational, bioinformatics, pharmacogenomics, engineering, and/or nanotechnology methods into the process will lead to the next stage. Current research in drug discovery from medicinal plants involves a multifaceted approach combining botanical, phytochemical, biological and molecular techniques.

They have been searched for secondary metabolites that have various pharmacological activities such as antioxidant, anticholinesterase, antimicrobial, anti-inflammatory and cytotoxic activity [12]. Many researches were reported various pharmacological activities since 43 year (**Table 1**), reported the list of plants from laminaceae having different pharmacological activities till to date. In the present era increasing of numbers drug-resistant strains were observed due to excess utilization of antibiotic which create an urgent need to develop novel antimicrobial agents [22].

Genus *Salvia* (*Sage*) is one of the most well-known Lamiaceae family, which has wide distribution, with nearly 1000 species in three regions of the world: Central and South America (500 spp.), Central Asia/Mediterranean (250 spp.), and Eastern Asia (90 spp) [23]. *Salvia* (sage) species have been used as folk medicine for cognitive brain function, along with various biological activities, since ancient times. Sage extracts possess antioxidant, estrogenic, and anti-inflammatory properties, anticholinesterase effects and also be used against memory loss. Some researchers have been investigated the anti cholinesterases and anti-alzheimer's effect from *salvia* species and most of the studies

reported the phytoconstituents of salvia species consist with the severap pharmacological uses such as cholinesterase inhibition, antioxidant [24]

and anti-inflammatory, larvicidal activity with respectively monoterpenoids and diterpene (Sclareol), oil and diterpenoids, sesquiterpene such as Germacrene D.

Table 1: List of Laminaceae Plants Involved in Different Potential Pharmacological Uses

S.No.	BIOLOGICAL NAME	COMMON NAME	CHEMICAL CONSTITUENTS	PHARMACOLOGICAL USES	REFERENCES
1.	<i>Ajuga reptans L.</i>	Common bugle	Polyphenols of the Flavonoidic, Polyphenolcarboxylic Acids, Iridoids	Asthma, Gout, Vasoconstrictor, Antioxidant, Antimicrobial, Antiinflammatory and wound healing, Antidiarrhoeaic, Antileucoreic, Hepatoprotecting and Vulnerar.	[25-26]
2.	<i>Nepeta cataria L.</i>	Catnip or Catmint	Caryophyllene, Caryophyllene Oxide, Citronellol, Geraniol, Elemol, Nerol, Nerolidol, Spathulenol, Citronellyl acetate, Nepetalactones, Alkaloids (Actinidine and Iridomyrmecine).	Antispasmodic, Carminative, Stimulant, Colic, Diarrhea, Cough, Asthma, Antibacterial, Antifungal, Analgesic.	[8,27-30]
3.	<i>Thymus vulgaris L.</i>	Thyme	Volatile oil of Thymol, Carvacol	Antiseptic, antimicrobial, anthelmintic, Antibacterial and antifungal, hepatotonic, Appetite stimulant,	[31,32]
4.	<i>Lavandula angustifolia</i>	Lavender English, Ostokhodus	Linalool, Perillyl Alcohol, Linalyl Acetate, Camphor, Limonene, Tannins, Triterpenes, Coumarinscineole and Flavonoids.	Inflammation, Coughing, Digestive Problems and act as a Sedative.	[33,34]
5.	<i>Stachys officinalis</i>	Betony	Glycosides, Betulinic Acid, D-Camphor, Delphinidin, Rutin, Ursolic Acid, Stachydrine, Trigonelline, Betonicine.	Neuro stimulant, Lung, Liver, Gallbladder and Spleen Problems, Digestive system stimulant.	[35]
6.	<i>Coleus aromaticus</i>	Indian borage	Butylaniside, Selenene, Terpinene, Verbenone, Cirsimaritin, Luteolin, Quercetin, Salvigenin, Taxifolin, Crategolic, Euscaphic, Chrysoeriol and Tormentic Acid.	Urolithiasis, Antiepileptic, Antitumor and antimutagenic. Antimicrobial, Asthma, Bronchitis, Urinary Infections.	[36-42]
7.	<i>Ocimum sanctum</i>	Sacred basil, kali tuls, veranda.	Volatile Oil (0.8%), Nerol, Caryophyllene, Terpinene-4-ol-decylaldehyde, A-Selinene, A-Pinene	Smooth muscle relaxants, Cardiac depressants, Antiferlity, Adaptogenic, Immunomodulator y properties	[43,44]
8.	<i>Rosmarinus officinalis</i>	Rosemary	Rosmarinic Acid, Caffeic Acid, Chlorogenic Acid, Carnosolic Acid, Rosmanol, Carnosol, Rosmari-Diphenol, Rosmariquinonem, Ursolic Acid, Glucocolic Acid, Alkaloid Rosmaricine. Oil Contains Esters, Borneol, Cineoles, Chiefly - Pinene and Camphene.	Antispasmodic, antiasthmatic, Pepticulcer, Inflammatory diseases, Hepatotoxicity, Atherosclerosis, Ishaemic heart disease, Anti - cancer, Rheumatoid arthiritis.	[45,46]
9.	<i>Origanum dictamnus</i>	Dittany of crete	Flavonoids, Glycosides, Essential Oil have Carvacrol, A-Terpinene, P -Cymene, Caryophyllene, Borneol, Terpin1-en-4-ol and Carvacrol, Methyl Ether.	Antioxidant, Antibacterial, Antifungal, Diaphoretic, Carminative, Antispasmodic and Analgesic activities.	[16,47-49]
10.	<i>Origanum majorana</i>	Sweet marjoram Knotted marjoram Sampsishia	Polyphenols, Monoterpenes, Anti oxidants, Arbutin, 5,6,3'-Trihydroxy-7,8,4'Trimethoxyflavone, Hesperetin, Hydroquinone, Rosemarinic	Antioxidant, Antifungal, Antimicrobial activities, Antimetastatic and antitumor growth effects on Human Breast Cancer Cells, Anti-proliferative	[50-55]

			Acid	effect.	
11.	<i>Origanum vulgare</i>	Oregano	Sabinene (Monoterpene), (Z)-B Ocine (Monoterpenes), Terpinen-4-ol (Oxygenated Monoterpenes), Spathulenol (Oxygenated Sesquiterpene), Carvacrol and Thymol.	Antioxidant, Antibacterial, Antifungal, Diaphoretic, Carminative, Antispasmodic and Analgesic activities.	[16, 49, 56, 57]
12.	<i>Marrubium vulgare</i>	Horehound or hoarhound	Marrubiin, Marrubenol and Phenylpropanoid Esters, Carvacrol, B-Phellandrene.	Antihypertensive, Antispasmodic, Immunomodulating, Antioedematogenic,	[58-59]
13.	<i>Callicarpa Americana</i>	American Beauty berry	Flavonoids, Essential oils, Humulene epoxide II, A-Humulene.	Antirheumatic, diaphoretic, and febrifuge. Dysentery, dizziness, stomachache, Fever	[60]
14.	<i>Hyssopus officinalis</i>	Hyssop	Quercetin, 7-O-B-Dapiofuranosyl-(1→2)-B-Dxylopyranoside, Quercetin 7-O-B-D-Apiofuranosyl-(1→2)-B-D-Xylopyranoside 30-Ob-D-Glucopyranoside, Apigenin, Luteolin, Diosmin, Phenolic Compounds	Antifungal and insecticidal, Antiviral properties, Myorelaxant, Antiplatelet, α-Glucosidase inhibitory activity.	[61-67]
15.	<i>Agastache Mexicana</i>	Mexican giant hyssop Toronjil morado, toronjil rojo,	Flavonoids -Hesperitin Quercetin, Phenolic Compounds	Anxiogenic or anxiolytic like behaviors, Vasorelaxant, Antioxidant, Antinociceptive and Anti-inflammatory effects, Insomnia and CV disorders	[69-70]
16.	<i>Agastache rugosa</i>	Korean mint, blue liquorice	Methyleugenol, Estragole, Eugenol, Thymol, Pulegone, Limonene, Caryophyllene, Phenylpropanoids.	Nematicidal Activity, Treatment of Anorexia, Vomiting and Antimicrobial, Anti-Fungal and Antiviral Activity.	[71-74]
17.	<i>Minthostachys mollis</i>	Muna	Neomenthol, Menthol, Menthone and Piperitone, Carvacryl Acetate and Pulegone	Insecticide Activity, Antimicrobial Activity, Anthelmintic, Digestive, Aphrodisiac, Anti-Diarrheic, Anti-Rheumatism.	[75-78]
18.	<i>Minthostachys verticillata</i>	Peperina	Pulegone, Menthone, Limonene	Diarrhea and vomiting, Digestive, Sedative, Antispasmodic bronchodilating agent. Insecticidal, fungicidal, and antiparasitic properties, Antimicrobial activity	[79-82]
19.	<i>Melissa officinalis</i>	Melissa, lemon balm, Barangbo Badranj buyeh	Carvacrol, Citral, Sabinene	Blood depurative, Sedative, Diuretic, wound healing, Gout, Dyspepsia, Emmenagogue, Joints pain Diaphoretic Spasm, Ulcer healing properties, Human CNS cholinergic receptor binding, Antiinflammatory, Cytotoxic (MTT), Antiviral, Anti-HSV- 1.	[83-85]
20.	<i>Perilla frutescens</i>	Perilla	Perilloside E 7-(2-O-B-D-Glucuronyl-B-D-Glucuronyloxy)-5,3',4'-Trihydroxyflavone, Shisonin, Scutellarin, Coumaroyl Tartaric Acid, Luteolin, Linolenic Acid, Oleic Acid.	Antioxidant, Analgesic and Abortifacient agent, Asthma, Chronic bronchitis and Vomiting.	[86-90]

21.	<i>Prunella vulgaris</i>	Self heal	Oleanane, Ursane, Lupinane, Ursolic acid, Oleanolic acid, Sterols, Phenyl propanoids, Flavonoids-Luteolin, Homoorientin, Cinaroside, Quercetin Kaempferol, Coumarins, Volatile Oils, Fatty Acids.	Antiinflammation, Breast Cancer, Goiter, Tuberculosis, Immune Modulation, Antioxidative action, Anti-Viral, Anti-bacterial effects.	[91-95]
22.	<i>Plectranthus esculentus</i>	<i>Coleus, Livingstone Potato</i>	Crude Protein, Crude Fibre, Total Ash Content, Carbohydrate Mineral, Sodium, Calcium, Potassium, Iron.	Antibacterial and Antifungal, Treatment of Hypertension, Glaucoma, Asthma, Cancers.	[96,97]
23.	<i>Menthe piperita</i>	Pepper mint, colpermin	Menthol, Menthone, Methofuran, Menthylisovalerate, Limonene, Isopulegone, Cineole, Pinene, Camphene	Anxiety associated with depression, Menstrual cramps, Flatulence, Indigestion, Irritable bowel syndrome Antibacterial, Antifungal and Antiviral.	[98]
24.	<i>Leonotis nepetifolia</i> L.	'Lion ear' Klip dagga.	Leonurine and Stachydrone, Leonuride, Leonurin & Leonuridine, Rutin, Quercetin, Hyperoside, Apigenin, <i>N</i> -Octacosanol, <i>N</i> -Octacosanoic acid, Quercetin, Campesterol- β Sitosterol, Neptaefolin, Neptaefuran, Neptaefuranol, Neptaefolinol, Leonitin, Labdanic, laballenic acid.	Treat Bronchial Asthma, Anti Malarial, Analgesic, Antiinflammatory activity, Antifungal, Antibacterial activity, Antispasmodic activity, Cure of Paralysis, Skin Diseases, Rheumatism, Post natal swelling.	[99-104]
25.	<i>Salvia officinalis</i>	Garden sage	Viridiflorol, Pimaradiene, Salvianolic Acid, Rosmarinic, Bornyl Acetate, Camphene, A and B-Thujone, Linalool, A- and B-Caryophyllene.	Antibacterial, Anti-Inflammatory, Free Radical Scavenging activity, Antitumor Activities, Antioxidant activity	[105-108]
26.	<i>Salvia hispanica</i>	Chia seeds, salba	Omega 3 Fatty Acids, Dietary Fiber, Antioxidants, Amino Acids, Quercetin Kaempferol, Caffeic Acid, Chlorogenic Acids	Antihyperlipidemic, Antitumour, Cardioprotective effects, Anti obesity and anti-diabetic	[109]
27.	<i>Phlomis fruticosa</i>	Jerusalem Sage	Pinene, Iridoids. 1, 8- Cineole and B-Caryophyllene,	Antifungal, Antibacterial activity	[110]
28.	<i>Ocimum basillicum</i>	Sweet basil, sabja, Tukmaria, falloda seeds	Polyphenolic Flavonoids (Orientin and Vicenin), Eugenol, Citronellol Linalool, Limonene, Citral, Terpeneol, B-Carotene, Lutein, Zeaxanthin, Vit A and Vit-K.	Antibacterial, Anti-inflammatory activity	[111-113]

Sclareol also have significant cytostatic and cytotoxic effect. Many studies on the essential oil of *S. fruticosa* reported 1, 8-cineole to be the main component, followed by camphor, α -thujone, β -thujone, (**Figure 1**) and β -caryophyllene. The species is known to contain biologically active sesquiterpenes (**Figure 2**) and diterpenes, besides the high content of oxygenated monoterpenes [114].

Salvinorin - A (Figure 3) is the main active psychotropic molecule in *Salvia divinorum*. Salvinorin A, a neoclerodane diterpene, is the most potent naturally occurring

hallucinogen known and rivals the synthetic hallucinogen lysergic acid diethylamide in potency. Recently, the molecular target of salvinorin - A was identified as the kappa opioid receptor (KOR).

Salvinorin A represents the only known non-nitrogenous KOR selective agonist. Based on the selectivity of salvinorin A for the KOR, this receptor represents a potential molecular target for the development of drugs to treat disorders characterized by alterations in perception, including schizophrenia, Alzheimer's disease and bipolar disorder [115].

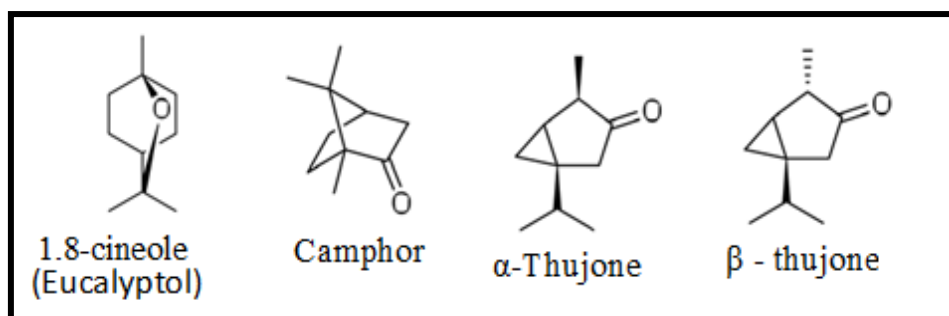


Figure 1: Structures of some monoterpenoids found in *Salvia sepcies*

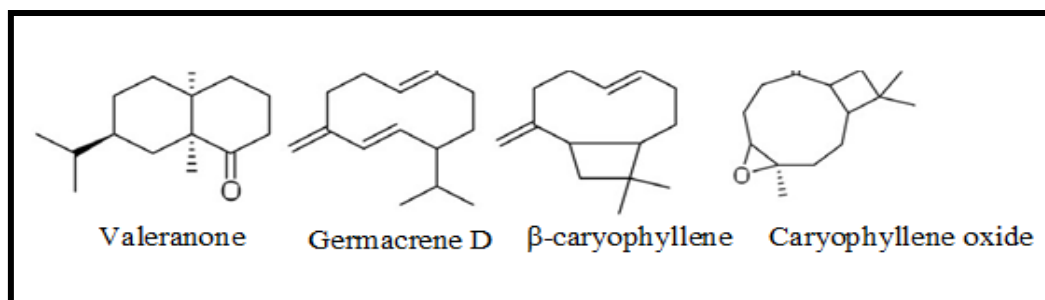


Figure 2: Structures of some sesquiterpenoids found in *Salvia sepcies*

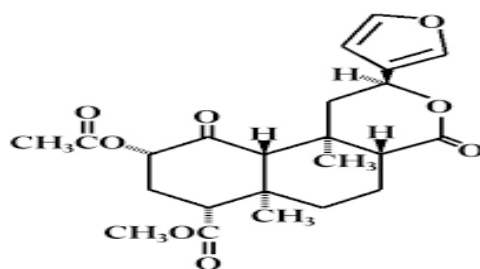


Figure 3: Salvinorin A

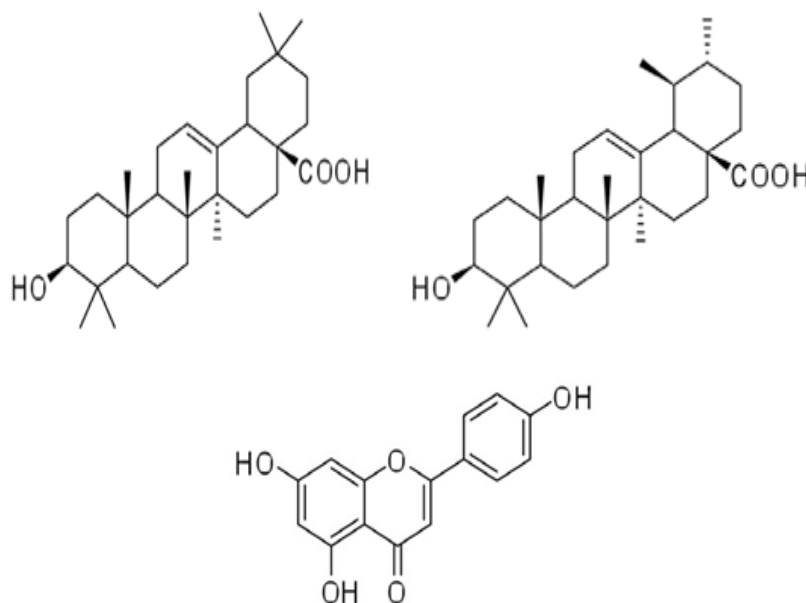
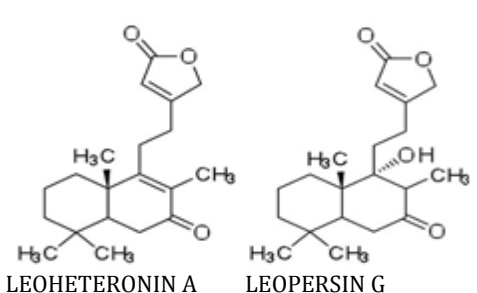
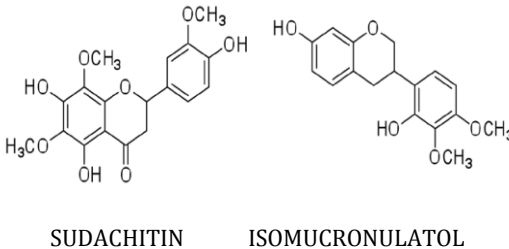
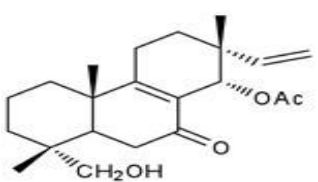
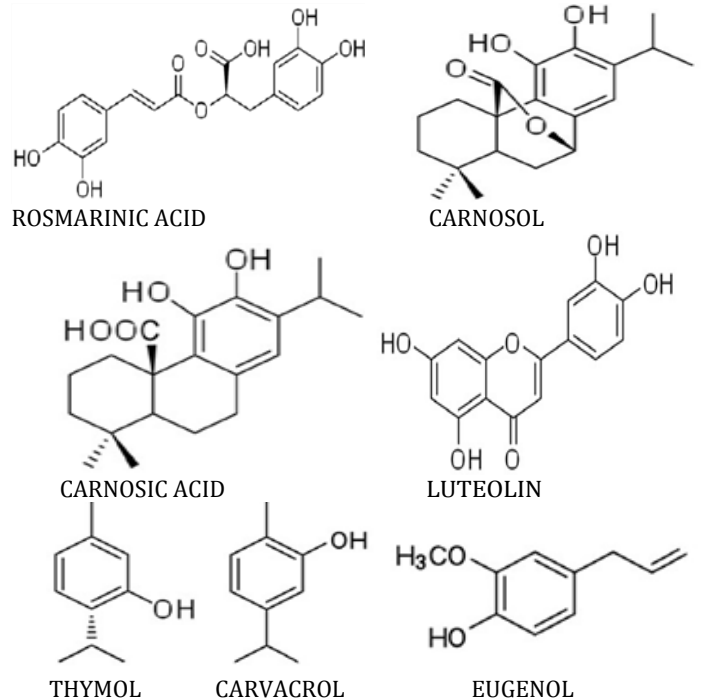


Figure 4: Some of the chemical Constituents and structures of *Rosmarinus officinalis L.*

Table 2: Chemical Constituents and their Pharmacological Activity of Some Laminaceae Plants [118,120]

S. NO.	BIOLOGICAL NAME	CHEMICAL CONSTITUENTS AND STRUCTURE OF THE CONSTITUENTS	PHARMACOLOGICAL ACTIVITY
1.	<i>Leonurus heterophyllus</i> L	 <p>LEOHETERONIN A LEOPERSIN G</p>	AChE enzyme inhibitory activity
2.	<i>Micromeria cilicica</i> (<i>Clinopodium cilicicum</i>)	 <p>SUDACHITIN ISOMUCRONULATOL</p>	BChE enzyme inhibitory activity
3.	<i>Origanum</i> species	 <p>SORGEROLONE</p>	Memory enhancing, Antioxidant, Antibacterial, Anti-inflammatory and antispasmodic effects
4.	<i>Rosmarinus officinalis</i> L.	 <p>ROSMARINIC ACID CARNOSOL</p> <p>CARNOSIC ACID LUTEOLIN</p> <p>THYMOL CARVACROL EUGENOL</p>	<ul style="list-style-type: none"> • Antioxidant and cytotoxic activity, Neuroprotective, antimicrobial, anti-inflammatory activity • Modulate the neuroprotective defense system against cellular stress. • Carnosol have the potential to protect cortical neuronal cells by activation of the keap1/nrf2 pathway. • Monoterpenes, triterpenoids and Apigenin act against alzheimer's disorder by cox-2 inhibition.

One of the most active phytochemical products from the Lamiaceae family plants is rosmarinic acid, particularly rosemary (*Rosmarinus*), sage (*Salvia*), lemon balm (*Melissa officinalis*), catmint (*Nepeta*), and lavender (*Lavandula*), and other phenolics that have strong antioxidant, anti-inflammatory, and neuroprotective effects in the treatment of neurological disorders, particularly AD [116]. As terpenoids, triterpenoids, especially oleanane, ursane, and lupane triterpenoids, Ursolic/oleanolic acid (**Figure 4**) or betulinic acid and their derivatives are important potential agents in the treatment of AD. Diterpenes, especially phenolic ring-containing diterpenes, such as abietanes, were tested for anticholinesterase activity. Ferruginol and taxodione are fairly common abietanes isolated from a number of *Salvia* species that exhibited high BChE inhibition activity and weak-moderate AChE inhibition activity. Several abietane diterpenoids, including Carnosol and Carnosic acid and quinoid abietanes, such as Tanshinones, should also be considered potential anti-Alzheimer agents [117].

Essential oils of some of the salvia (*Satureja*, *Thymus*, *Origanum*, and *Thymbra*) species are known to be rich in the monoterpenes thymol and carvacrol as well as β -pinene. Volatile constituents of the essential oils are likely to readily cross the blood-brain barrier due to their small molecular size and lipophilicity. Their volatile nature may also enable their administration as an inhaled vapor [118]. Therefore, consumption of Lamiaceae plants rich in thymol and carvacrol is useful in the treatment of AD [119].

CONCLUSION

Aromatic plants have possessed several pharmacological activities with presence of several phytoconstituents. However, most of the aromatic plants are not yet exploited for their bioactivities. In reference to the above discussion we focused on investigation of immunomodulatory effect on one of the salvia species with different extraction procedure by employing in-vitro and in-vivo screening methods.

REFERENCES

1. Akerele O. Medicinal plants and primary healthcar, an integrated agenda for action. *Fitoterapia* 1988;59:355-65.
2. Daniel Dias A, Sylvia Urban and Ute Roessner. A Historical Overview of Natural Products in Drug Discovery. *Metabolites* 2012;2:303-36.
3. Jacob EJ. Natural Products-Based Drug Discovery: Some Bottlenecks and Considerations. *Current Science* 2009;96(6):753-54.
4. Mouhssen Lahlou. The Success of Natural Products in Drug Discovery. *Pharmacology & Pharmacy* 2013;4:17-31.
5. Christaki E, Bonos E, Giannenas I et al. Aromatic plants as a source of bioactive compounds. *Agriculture* 2012;2:228-43.
6. Lahlou M. Screening of Natural Products for Drug Discovery. *Expert Opinion on Drug Discovery* 2007;2(5):697-705.
7. Rates SMK. Review: Plants as source of drugs. *Toxicon* 2001;39:603-13.
8. Nostro A, Cannatelli MA, Crisafi G, Alonzo V. The Effect of *Nepeta cataria* extract on adherence and enzyme production of *Staphylococcus aureus*. *International Journal of Antimicrobial Agents* 2001;18:583-85.
9. Edeoga HO, Okwu DE, Mbaebie BO. Phytochemical Constituents of some Nigerian Medicinal plants. *African journal of Biotechnology* 2005;4:685-88.
10. Farnsworth NR, Akerele O, Bingel AS, Soejarto DD, Guo Z. *Bull WHO* 1985;63: 965.
11. Madsen HL, Bertelsen G. Spices as Antioxidants. *Trends in Food Sci. & Tech* 1995;6:271-7.
12. Ulubelen A, Topcu G, and Kolak U. Studies in Natural Product Chemistry, Labiatae Flavonoids and Their Bioactivity (Ed) Attaur-Rahman. Elsevier 2005;30:233-302.
13. Perry E, Howes M-JR. Medicinal plants and dementia therapy: Herbal hopes for brain aging? *CNS. Neurosci. Ther* 2011;17:683-98.
14. Newman DJ, Cragg GM, Snader KM. The influence of natural products upon drug discovery. *Nat Prod Rep* 2000;17:215-34.
15. Venkateshappa SM and Sreenath KP. Potential Medicinal Plants of Lamiaceae. *American International Journal of Research in Formal, Applied & Natural Sciences* 2013 June-August;3(1):82-87.
16. Saraç N, Uğur A, Duru ME, Varol O. Antimicrobial activity, antioxidant activity and chemical composition of *Origanum onites* L. and *Origanum vulgare* L. subsp. *hirtum* (Link) Ietswaart from Mugla (Turkey). *Acta Horti* 2009;826:397-403.

17. Bhattacharjee SK and Jain Shashi. Hand book of medicinal plant. Published by Pointer publisher, Vyas Building SMS. Highway Jaipur 302003 (Raj) India, 2001:220-22.
18. Nurdan Sarac and Aysel Ugur. Antimicrobial activities and usage in folkloric Medicine of some Lamiaceae species growing in Mugla, turkey. *EurAsia J, BioSci* 2007;4:28-37.
19. Lawrence BM. Commercial production of non-citrus essential oils in North America. *Perfumer & Flavorist* 1979;3:21-33.
20. Akbar S, Tariq M, & Nisa M. A study on CNS depressant activity of *Salvia haematodes* Wall. *International Journal of Crude Drug Research*. 1984;22:41-4.
21. Rami K, Li Z. Antimicrobial activity of essential oil of *Salvia officinalis* L. collected in Syria. *Afr J Biotech* 2011;10:8397-402.
22. Kordaly S, Kotan R, Mavi A et al. Determination of the chemical composition and antioxidant activity of the essential oil of *Artemisia dracunculoides* and of the antifungal and antibacterial activities of Turkish *Artemisia absinthium*, *A. dracunculoides*, *Artemisia santonicum* and *Artemisia spicigera* essential oils. *Journal of Agricultural and Food Chemistry* 2005;53:9452-58.
23. Walker JB, Sytsma K, Treutlein J, Wink M. *Salvia* (Lamiaceae) is not monophyletic: implications for the systematics, radiation, and ecological specializations of *Salvia* and tribe *Menthae*. *Am. J. Bot* 2004;91:1115-25.
24. Şenol FS, Orhan IE, Erdem SA, Kartal M, Şener B, Kan Y, et al. Evaluation of cholinesterase inhibitory and antioxidant activities of wild and cultivated samples of Sage (*Salvia frutescens*) by activity guided fractionation. *J. Med. Food* 2011;14:1476-83.
25. Ghita G, Cioanca O, Gille E, Necula R, Zamfirache MM, Stanescu U. Contributions To The Phytochemical Study of Some Samples of *Ajuga reptans* L. and *Ajuga genevensis* L. *Bulletin of The Transilvania University of Brasov Series Vi, Medical Sciences* 2011;4 (53):2.
26. Breschi MC, Martinotti E, Catalano S, Flamini G, Morelli I and Pagni AM. Vasoconstrictor activity of 8-O-acetylharpagide from *Ajuga reptans*. *J. Nat. Prod* 1992;55:1145-8.
27. Baser KHC, Kirimer N, Kurkcuoglu M, and Demirci B. Essential oils of *Nepeta* species growing in Turkey. *Chemistry of Natural Compounds*. 2000;36(4):356-59.
28. Mortuza-Semmani K, Saeedi M. Essential oil composition of *Nepeta cataria* L. and *Nepeta crassifolia* Boiss and Bushe from Iran. *Journal of Essential Oil Bearing Plants*. 2004;7:122-24.
29. Massoco CO, Silva MR, Gorniak SL, Spinosa MS, Bernardi MM. Behavioral effects of acute and long-term administration of catnip (*Nepeta cataria*) in mice. *Veterinary and Human Toxicology* 1995;37:530-33.
30. Aydin S, Beis R, Ozturk Y, Baser KH. Nepetalactone: a new opioid analgesic from *Nepeta caesarea* Boiss. *Journal of Pharmacy and Pharmacology* 1998;50:813-17.
31. Maher Ali Ahmed AlMaqtari, Saeed Mohammed Alghalibi, Ebtessam HasanAlhamzy. Chemical composition and antimicrobial activity of essential oil of *Thymus vulgaris* from Yemen. *Turk J Biochem* 2011;36:342-49.
32. Hompson JJ, Chalcha A, Michet Y, Linhart and Ehlers B. Qualitative and quantitative variation in monoterpene co-occurrence and composition in essential oil of *Thymus vulgaris* chemotypes. *J Chem Ecol* 2003;29(4):859-80.
33. Rychlik M. Quantification of free coumarin and its liberation from glucosylated precursors by stable isotope dilution assays based on liquid chromatography-tandem mass spectrometric detection. *J Agric Food Chem* 2008;2-13, 56(3),7:96-801.
34. Shimizu M, Shogawa H, Matsuzawa T, Yonezawa S, Hayashi T, Arisawa M, et., al. Anti-inflammatory constituents of topically applied crude drugs. IV. Constituents and anti-inflammatory effect of Paraguayan crude drug "alhucema" (*Lavandula latifolia* Vill.). *Chem Pharm Bull (Tokyo)* 1990;38(8):2283-84.
35. Ahmad VU, Arshad S, Bader S, Ahmed A, Iqbal S, and Tareen RB. New phenethyl alcohol glycosides from *Stachys parviflora*. *J Asian Nat Prod Res* 2006;8(1-2):105-11.
36. Warriar PK, Nambiar VP, Ramankutty. *Indian Medicinal Plants*, 1st Ed., Orient Longman Limited, Madras. 1995;4:315-17.
37. Shamachar BR. Use of Pashanabheda *Coleus aromaticus* in ashmari (Urinary calculi). *Natn. Med. J* 1967;9:475-77.
38. Rao BS, Shanbhoge R, Upadhyaya D, Jagetia GC, Adiga SK, Kumar P. Antioxidant, anticlastogenic and radioprotective effect of *Coleus aromaticus* on Chinese hamster fibroblast cells (V79) exposed to gamma radiation. *Mutagenesis* 2006;21:237-42.
39. Perumal G, Subramanyam C, Natrajan D, Srinivasan K, Mohanasundari C, Prabakar K. Antifungal activities of traditional medicinal plant extracts: A preliminary survey. *J Phytolog Res* 2004;17:81-3.
40. Baskar R, Varalaksmi P, Amsaveni. Changes in tissue enzymes produced by *Coleus*

- aromaticus* experimental Urolithiasis. Indian Drugs 1992;29:254-8.
41. Jose MA, Ibrahim, Janardhanan S. Modulatory effect of *Plectranthus amboinicus* Lour on ethylene glycol induced nephrolithiasis in rats. Indian J Pharmacol 2005;37:43-4.
 42. Buznego MT, Perez-Saad H. Antiepileptic effect of *Plectranthus amboinicus* (Lour.). Spreng Rev Neurol 1999;29:229-32.
 43. Annapurani S, Priya R. Antimutagenic, antitumorigenic and antigenotoxic effects of polyphenol extracts of selected medicinal plants. Indian J Nutr Diet 1999;36:431-5.
 44. Ramasubramania Raja R. Medicinally Potential plants of labiatae (Lamiaceae) Family: An Overview. Research journal of medicinal plant 2012;6(3):203-213.
 45. Abe F, Yamauchi T, Nagao T, Kinjo J, Okabe H, Higo H and Akahane H. Ursolic acid as a trypanocidal constituent in rosemary. *Biol. Pharm. Bull* 2002;25:1485-7.
 46. Haloui M, Louedec L, Michel JB and Lyoussi B. Experimental diuretic effects of *Rosmarinus officinalis* and *Centaureum erythraea*. *J. Ethnopharmacol* 2000;71:465-72.
 47. Coelho da Costa A, Cavalcanti dos Santos BE, Santos, FL, Lima EO. Antibacterial activity of the essential oil of *Origanum vulgare* L. (Lamiaceae) against bacterial multiresistant strains isolated from nosocomial patients. *Rev. Bras. Farmacogn* 2009;19:236-41.
 48. Harvala C, Skaltsa H. Contribution a l'etude chimique d'*Origanum dictamnus* L. 1 recommunication. *Plantes Méd. Phytothér.* Tome. 1986;20(4):300-4.
 49. Tommasi L, Negro C, Miceli A, Mazzotta F. Antimicrobial activity of essential oils from aromatic plants grown in the Mediterranean area. *J. Essent. Oil Res* 2009;21:185-89.
 50. Aazza S, Lyoussi B and Miguel MG. Antioxidant activity of some Moroccan hydrosols. *J Med Plants Res* 2011;5:6688-96.
 51. Prakash B, Singh P, Kedia A and Dubey NK. Assessment of some essential oils as food preservatives based on antifungal, antiaflatoxin, antioxidant activities and in vivo efficacy in food system. *Food Res Int* 2012;49:201-8.
 52. Oksal BS, Uysal B, Gencer A and Sozmen F. Antibacterial activity of essential oil obtained from *Origanum majorana* L. By solvent-free microwave extraction. *Planta Med* 2012;78:1267.
 53. Orhan IE, Ozcelik B, Kartal M and Kan Y. Antimicrobial and antiviral effects of essential oils from selected Umbelliferae and Labiatae plants and individual essential oil components. *Turk J Biol* 2012;36:239-46.
 54. Mengulluoglu M and Soyly S. Antibacterial activities of essential oils extracted from medicinal plants against seed-borne bacterial disease agent, *Acidovorax avenae subsp citrulli*. *Res Crop* 2012;13:641-6.
 55. Lamien-Meda A, Lukas B, Schmiderer C, Franz C and Novak J. Validation of a quantitative assay of arbutin using gas chromatography in *Origanum majorana* and *Arctostaphylos uva-ursi* extracts. *Phytochem Anal* 2009;20:416-20.
 56. Souza EL, Stamford TLM, Lima EO, Trajano VN. Effectiveness of *Origanum vulgare* L. essential oil to inhibit the growth of food spoiling yeasts. *Food Control* 2007;18:409-13.
 57. Mastelic J, Jerkovic I, Blazevic I, Poljak-Blazi M, Borovic S, Ivancic-Bace I, Smrecki V, Zarkovic N, Brcic-Kostic K, Vikic-Topic D, Muller N. Comparative study on the antioxidant and biological activities of carvacrol, thymol and eugenol derivatives. *J. Agric. Food Chem* 2008;56:3989-96.
 58. De Souza MM, De Jesus RAP, Cechinel-Filho V and Schlemper V. Analgesic profile of hydroalcoholic extract obtained from *Marrubium vulgare*. *Phytomedicine* 1998;5:103-107.
 59. Meyre-Silva C, Yunes RA, Schlemper V, Campos-Buzzi FV. Cechinel-Filho. Analgesic potential of marrubiin derivatives, a bioactive diterpene present in *Marrubium vulgare* (Lamiaceae). *Farmacogn* 2005;60:321-6.
 60. Mario R. Tellez,* Franck E. Dayan, Kevin K. Schrader, David E. Wedge, and Stephen O. Duke. Composition and Some Biological Activities of the Essential Oil of *Callicarpa americana* (L.) Natural Products Utilization Research Unit, Agricultural Research Service, U.S. Department of Agriculture, Box 8048, Oxford, Mississippi 38677. *J. Agric. Food Chem* 2000;48(7):3008-12.
 61. Wang N, Yang X-W. Two new flavonoid glycosides from the whole herbs of *Hyssopus officinalis*. *J. Asian Natural. Prod. Res.* 2010;12:1044-50.
 62. Varga E, Hajdu Z, Veres K, Mathe I, Nemeth E, Pluhar Z, Bernath J. Investigation of variation of the production of biological and chemical compounds of *Hyssopus officinalis* L. *Acta. Pharm. Hung* 1998a;68:183-88.
 63. Kochan E, Wysokinska H, Chmiel A, Grabias B. Rosmarinic acid and other phenolic acids in hairy roots of *Hyssopus officinalis*. *Biosciences* 1999;54:11-16.
 64. Moro A, Zalacain A, de Hurtado MJ, Carmona M. Effects of agronomic practices on volatile

- composition of *Hyssopus officinalis* L. essential oils. *Molecules* 2011;16:4131-39.
65. Alineza H, Azimi R, Zare M, Ebrahimzadeh MA, Eslami S, Nabavi SF, Nabavi SM. Antioxidant and antihemolytic activities of ethanolic extract of flowers, leaves and stems of *Hyssopus officinalis* L. var. *angustifolius*. *Int. J. Food Prop* 2013;16:1169-78.
 66. Benedec D, Oniga I, Crişan G, Popescu H. Cercetări preliminare asupra unor compuşi polife nolici din *Hyssopus officinalis* L, (Lamiaceae). *Nota 2. Clujul Med* 2003;76:444-49.
 67. Fathiazad F, Hamedeyazdan S. A review on *Hyssopus officinalis* L. Composition and biological activities. *Afr. J. Pharm. Pharmacol.* 2011;14:1959-66.
 68. Molina-Hernández M, Téllez-Alcántara P, Martínez, E. *Agastache mexicana* may produce anxiogenic-like actions in the male rat. *Phytomedicin* 2000;7:199-203.
 69. Estrada-Reyes R, López-Rubalcava C, Ferreyra-Cruz O, DorantesBarrón AM, Heinze G, Moreno Aguilar J, et al. Central nervous system effects and chemical composition of two subspecies of *Agastache mexicana*; an ethnomedicine of Mexico. *J Ethnopharmacol* 2014;153:98-110.
 70. Hernández-Abreu O, Castillo-España P, León-Rivera I, Ibarra-Barajas M, Villalobos-Molina R, González-Christen J, et al. Antihypertensive and vasorelaxant effects of tilianin isolated from *Agastache mexicana* are mediated by NO/cGMP pathway and potassium channel opening. *Biochem Pharmacol* 2009;78:54-61.
 71. Song JH, Kim MJ, Kwon HD, Lee WK, Park IH. Antimicrobial activity and characterization of volatile flavor extracts from *Agastache rugosa*. *J. Food Sci. Nutr.* 1999; 4:97-102.
 72. Chae YA, Hyun-Choong O, Song, JS. Variability of the volatile composition of *Agastache rugosa* in South Korea. *Acta. Hort* 2005;675:59-64.
 73. Shin S, Kang CA. Antifungal activity of the essential oil of *Agastache rugosa* Kuntze and its synergism with ketoconazole. *Lett. Appl. Microbiol.* 2003;36:111-5.
 74. Shin S. Essential oil compounds from *Agastache rugosa* as antifungal agents against *Trichophyton* species. *Arch. Pharm. Res.*, 2004;27:295-99.
 75. Alkire BH, Tucker AO, Maciarelo MJ. Tipo, *Minthostachys mollis* (Lamiaceae): an Ecuadorian mint. *Economic Botany* 1994;48:60-4.
 76. Malagón O, Vila R, Iglesias J, Zaragoza T, Cañigüeral S. Composition of the essential oils of four medicinal plants from Ecuador. *Flavour and Fragrance Journal* 2003;18:527-31.
 77. Hammond GB, Fernández ID, Villegas LF, Viasberg AJ. A survey of traditional medicinal plants from the Callejón de Huaylas, Department of Ancash, Perú. *Journal of Ethnopharmacology* 1998;61:17-30.
 78. Schimidt-Lebuhn AN. Ethnobotany, biochemistry and pharmacology of *Minthostachys* (Lamiaceae). *Journal of Ethnopharmacology* 2008;118:343-53.
 79. De Feo V, Ricciardi A, Biscardi D, Senatore F. Chemical composition and antimicrobial screening of the essential oil of *Minthostachys verticillata* (Griseb.) Epling. Lamiaceae. *J Essential Oil Res* 1998;10:61-5.
 80. Ratera EL, Ratera MO. Plantas que curan. In: EL Ratera, MO Ratera (Eds.), *Plantas de la flora argentina empleadas en medicina popular*. Buenos Aires: Hemisferio Sur 1980;43-85.
 81. Primo V, Rovera M. Determinación de la actividad bacteriana y antiviral del aceite esencial de *Minthostachys verticillata* (Griseb.) Epling. *Rev Argentina Microbiol* 2001; 33:113-7.
 82. Zanón SM, Ceriatti FS. Search for antiviral activity of certain medicina plants from Córdoba, Argentina. *Rev Latino Americana, Microbiol* 1999;41:59-62.
 83. Araujo C, Sousa MJ, Ferreira MF and Leao C. Activity of essential oils from Mediterranean Lamiaceae species against food spoilage yeasts. *J. Food Pro* 2003;66:625-32.
 84. De Sousa AC, Alviano DS, Blank AF, Alves PB, Alviano CS and Gattass CR. *Melissa officinalis* L. essential oil: antitumoral and antioxidant activities. *J. Pharm. Pharmacol* 2004;56:677-81.
 85. Dharmani P, Kuchibhotla VK, Maurya R, Srivastava S, Sharma S and Palit G. Evaluation of anti-ulcerogenic and ulcer-healing properties of *Ocimum sanctum* Linn. *J. Ethnopharmacol* 2004;93:197-206.
 86. Aritomi M, Kumori T, Kawasaki T. Cyanogenic glycosides in leaves of *Perilla frutescens* var. *acuta* *Phytochemistry* 1985;24(10):2438-39.
 87. Ha TJ, Lee JH, Lee MH, Lee BW, Park HSKCH, Fujita T, Funayoshi A, Nakayama MA phenylpropanoid glucoside from *Perilla frutescens*. *Phytochemistry* 1994;37(2):543-46.
 88. Saklani S, Chandra S, Gautam AK. Phytochemical investigation and contribution of *Perilla frutescens* as spices in traditional health care system. *International Journal of Pharmacy Technology* 2011;3(4):3543-54.

89. Takano H, Osakabe N, Sanbongi C, Yanagisawa R, Inoue K, Yasuda A, Natsume M, Baba S, Ichiishi E, Yoshikawa T. Extract of *Perilla frutescens* enriched for rosmarinic acid inhibits seasonal allergic rhinoconjunctivitis in humans. *Exp. Biol. Med.* 2004;229:247-54.
90. Ueda H, Yamazaki C, Yamazaki M. Luteolin as an Antiinflammatory and Anti-allergic Constituent of *Perilla Frutescens*. *Biol. Pharm. Bull* 2002;25(9):1197-1202.
91. Cai ZQ, Ma HY and Zhang M. Content Analysis of Oleanolic Acid and Ursolic Acid in *Prunella vulgaris* Based on Its Provenances and Appearances. *Guangdong Yaoxueyuan Xuebao* 2009;25:256-58.
92. Liu Y, Song SJ and Xu SX. Advances in the Study on the Chemical Constituents and Biological Activities of *Prunella vulgaris* L. *Shenyang Yike Daxue Xuebao* 2003;20:55-9.
93. Sun YM, Xu JZ, Wang ZA and Yu XP. Determination of Ursolic Acid in *Prunella vulgaris* from Different Areas by HPLC. *Shizhen Guoyi Guoyao* 2009;20:1125-26.
94. Wang ZJ, Zhao YY, Chen YY., et al. Triterpenoid Compounds of *Prunella* Genus and Their Features of ¹³C NMR Spectroscopy. *Chinese Materia Medica* 2000;25:583-88;
95. Kojima H and Ogura H. Triterpenoids from *Prunella vulgaris*. *Phytochemistry* 1986;25:729.
96. Kyesmu, PM. and Akueshi, CO. Effects of relative humidity variation on crude protein, crude fibre, fats and ash contents of *Coleus potato (Plectranthus esculentus)*. N.E.Br.) under storage. *Nigerian Journal of Botany* 1989;2:1-7.
97. Richardson P. The chemistry of the Labiatae: An introduction and overview. In: Harley RM and Reynolds T. (Eds.) *Advances in Labiatae Science*, Botanical Garden Kew 1992;291- 97.
98. Baser KHC, Kurkcuoglu M, Tarimcilar G, Kaynak G. Essential oils of *Mentha* Species from Northern Turkey. *J. Essent. Oil Res* 1999;11:579-88.
99. Niteshwar and Kumari H. Ethnomedicinal claims of *Leonotis nepetifolia* (L). R. Br. A Review. *International J. of Res. In Ayurveda and pharmacy* 2012;(3)6:783-85.
100. Veerabadran U, Venkatraman A, Souprayane A, Narayanaswamy M, Perumal DS, Sivalingam S, Devaraj D and Perumal A. Evaluation of antioxidant potential of leaves of *Leonotis nepetifolia* and its inhibitory effects on MCF7 and Hep2 cell lines. *Asian Pacific Journal of Tropical Diseases* 2013;3(2):103-10.
101. Kumar B, Vijayakumar M, Govindarajan R, Pushpangadan P. Ethnopharmacological approaches to wound healing-exploring medicinal plants in India. *J Ethnopharmacol* 2007;114(2):103-13.
102. Nithya V, Brinda P and Anand KV. Wound healing activity of *Leonotis nepetifolia* R. Br., in Wistar Albino rats. *Asian J. of Pharm. Clin. Res* 2011;4(2):23-6.
103. Rastogi RP, Mehrotra BN. Compendium of Indian Medicinal Plants. Central Drug Research Institute, Lucknow and National Institute of Science Communication and Information Resource, New Dehli. 2006;2(411):1970-79.
104. Gunatilaka AAL, Sultanbawa MUS, Balasubramaniam. Chemical investigation of Sri Lankan plants, part 42 a survey of plants of Sri Lanka for alkaloids II. *J NATL Sci councl Sri Lanka*. 1980;8(2):187-207.
105. Veličkovic AS. et al. The possibilities of the application of some species of sage (*Salvia* L.) as auxiliaries in the treatment of some diseases. *Journal of the Serbian Chemical Society*, 2003, 68(1), 435-45.
106. Pierozan MK. et al. Chemical characterization and antimicrobial activity of essential oils of *Salvia* L. species. *Ciencia e Tecnologia de Alimentos* 2009;29(4):764-70.
107. Croteau R, Felton M, Karp F, Kjonas R. Relationship of camphor biosynthesis to leaf development in sage (*Salvia officinalis*) *Plant Physiol* 1981;67:820-4.
108. Khan A, Najeeb-ur- Rahman, Alkharfy K, Gilani A. Antidiarrheal and antispasmodic activities of *Salvia officinalis* are mediated through activation of K⁺ channels. *J Bangladesh Pharmacol Soc* 2011;6:111-6.
109. Avinash Rangaraju, Uppala Mohan Kumar. A Pharmacognostic Study on *Salvia hispanica*. *American journal of pharmacy and health research* 2013;1:9.
110. Sattar AA, Bankova V, Kujumgiev A, Ignatova A, Todorova C, and Popov S. Chemical Composition and Biological Activity of Leaf Exudates from some Lamiaceae Plants. *Bulgarian Academy of Sciences, Bulgaria* 1995;50(1):62-5.
111. Özcan M., Chalchat J.-C. Essential oil composition of *Ocimum basilicum* L. and *Ocimum minimum* L. in Turkey. *Czech J. Food Sci* 2002;20:223-28.
112. Baritoux O, Richard H, Touche J, Derbesy M. Effects of drying and storage of herbs and spices on the essential oil. Part I. Basil, *Ocimum basilicum* L. *Flavour Fragr J* 1992;7:267-71.

113. Lahariya AK, Rao JT. *In vitro* antimicrobial studies of the essential oil of *Cyperus scariosus* and *Ocimum basilicum*. Ind. Drugs 1979;16:150-52.
114. Topçu G, Öztürk M, Kuşman T, Barla Demirköz AA, Kolak U, Ulubelen A. Terpenoids, essential oil composition and fatty acids profile, and biological activities of Anatolian *Salvia fruticosa* Mill. Turk. J. Chem 2013;37:619-32.
115. Douglas J. Sheffler, Bryan L. Roth. Salvinorin A: The 'magic mint' hallucinogen finds a molecular target in the kappa opioid receptor. Trends in pharmacological sciences 2003; 24(3):107-9.
116. Howes MJR, Perry E. The Role of Phytochemicals in the Treatment and Prevention of Dementia. Drug & Aging 2011;28:439-68.
117. Gulacti Topcu, Tuba Kusman "Lamiaceae Family Plants as a Potential Anticholinesterase Source in the Treatment of Alzheimer's Disease" Bezmialem Science 2014;1:1-25.
118. Savelev SU, Okello EJ, Perry EK. Butyryl- and acetylcholinesterase inhibitory activities in essential oils of *Salvia* species and their constituents, Phytother Res 2004;18:315-24.
119. Öztürk M. Anticholinesterase and antioxidant activities of Savoury (*Satureja thymbra* L.) with identified major terpenes of the essential oil, Food Chem 2012;134:48-54.
120. Topcu and Kusman, Lamiaceae Family Plants as a Potential Anticholinesterase Source in the Treatment of Alzheimer's Disease Bezmialem Science 2014;1:1-25.