

**REVIEW ARTICLE ON OLEA EUROPAEA L. (OLIVE): A
RESERVOIR PLANT WITH PHARMACOLOGICAL ACTIVITY****Chestha Rawat* and Monika Bisht**

School of Pharmaceutical Sciences, Shri Guru Ram Rai University,
Patel nagar, Dehradun (Uttarakhand) India (248001).

Article Received on
05 January 2022,

Revised on 25 January 2022,
Accepted on 15 Feb. 2022

DOI: 10.20959/wjpr20223-23324

Corresponding Author*Chestha Rawat**

School of Pharmaceutical
Sciences, Shri Guru Ram
Rai University, Patel Nagar,
Dehradun (Uttarakhand)
India (248001).

ABSTRACT

Nature has always been a great supporter of human life, even in the worse of situation we are dependent on nature. Apart from scenic beauty, peace, greenery and relaxation of mind, it has also provide us with vast herbal therapy. Various plants, various species each fulfilled with plenty of chemical constituents, every chemical moiety owning the property to heal the suffering, not to mention with the least most side effects, poisonous plants an exception. This review is totally based on the most common plant Olive and its pharmacological activity. Its uses both medicinally as well as commercially is known. Medicinally, it is used as antioxidant, anticarcinogenic, antiinflammatory, antimicrobial, antihypertensive, antidyslipidemic, cardiotonic, laxative,

antiplatelet, antidiabetic, etc where as commercially its used as olive oil, olive stone, olive cake and olive wood. There are three different types of edible olive oil prepared in different ways Kalamata olives (20.3%), green olives (39.7%) and black olives (40%). Awareness about the beneficial uses of olive as a nutritional diet and functional food has given rise to world wide consumption of olive and its products. Different parts of olive tree and their uses had made it a hotspot for researchers, all its seeds, fruits, branches, leaf, roots, stem, none lacks any pharmacological activity.

KEYWORDS: Olive, Antidiabetic, Cardiotonic, Antihypertensive, Antioxidant, Antidyslipidemia, Anticarcinogenic, olive cake.

INTRODUCTION

Basically, found in the tropical and warm temperate regions of the world, a small tree named as olive, Botanically *Olea europaea* L., is the member of oleaceae family. It is famous for its

fruits and in commercial aspect it is the most important in the Mediterranean region, for its oil (Olive oil).^[1] Traditionally, Olive tree denotes the symbol of abundance, glory and peace, in the past even the leaves were used to crown the conqueror of bloody wars. This tree and its vegetative parts like fruits, leaves, stems, roots, seeds all have the history of being a rich source of nutrition, medicines and ceremonial uses.^[2]

Olea europaea L. is the cultivated species that belongs to the oleaceae family, contains 30 genera and 600 species.^[3] Mediterranean Basin produces 98% of the world total (11 million tons approx) olive crop and provides with the economical and dietary benefits to the people.⁴ Even the byproducts of olive, like the olive oil extraction obtained from olive trees are used. The pruning of olive leaves is about ~25 kg per olive tree including 5% of the weight of olive collected at the oilmill.^[5]

Extracts from olive leaf has always proved its way in promoting health and preservation. Ancient Egyptians used to use olive leaf for mummification of Pharaohs. Likewise, it has been used as a folk remedy for malaria and fever.^[6] Though olive and its fruits have nutritional and potential medicinal value, but due to its bitterness it is not palate friendly, so for the reason they are being consumed in two different forms, that is as oil or table olives.^[7] The component responsible for its unacceptable taste is oleuropein.^[8]

Olive is widely distributed in Asia, Africa, Europe, and Oceania which belongs to genus *olea* that approximately contains 30-35 species. *Olea* is the only species of olive that is edible.^[9] Depending on the different place in the world olive is known by different names such as *olivo* (Spain), *elia* (Greece), *jaitun* (India), *olive* (England), *oulivie* (France), and *zaitun* (Pakistan).^[9]

Difference in drupe colour, shape, oil composition, morphology of leaf and phenology helps in its distinction. There are 42 characteristics of leaf, fruit, stone morphology based on their description for their identification.^[10] It's a plant with slow growth and a species with extremely long life expectancy year upto 1000 years. The olive is an advantageous plant which can be consumed whole either as ripe black fruit or as the unripe green fruit.^[10]

Taxonomical classification

Kingdom: Plantae

Division: Angiosperms

Class: Eudicots

Order: Lamiales

Family: Oleaceae

Genus: Olea

Species: europaea^[11]

Chemical constituents

1. Olive fruits

- a. **Flavonols** - Quercetin-3-rutinoside, Luteolin-7-glucoside, Luteolin-5-glucoside, Apigenin-7-glucoside
- b. **Phenolic acids** - Chlorogenic acid, Caffeic acid, p-Hydroxybenzoic acid, Protocatechuic acid, Vanilic acid, Syringic acid, p-Coumaric acid, o-Coumaric acid, Ferulic acid, Sinapic acid, Benzoic acid, Cinnamic acid, Gallic acid
- c. **Phenolic alcohols** - (3,4-Dihydroxyphenyl) ethanol (3,4-DHPEA), (p-Hydroxyphenyl) ethanol (p-HPEA)
- d. **Secoiridoids** - Oleuropein, Demethyloleuropein, Ligstroside, Nuzhenide
- e. **Hydroxycinnamic acid derivatives** – Verbascoside.^[12]

2. Olive leaves

Oleuropein, ligstroside, tyrosol or hydroxytyrosol, out of which major biphenol is oleuropein (ethanolic extract) (20-25% total dry matter).^[13]

3. Olive seeds and olive stone

2- 4 g oil, Lignin (20.63% - 25.11%), hemicellulose (21.45% -27.64%), and cellulose (29.79% - 34.35%).

4. Olive oil

Main phenolic compounds present in olive oil are Hydroxytyrosol and tyrosol, ligustroside and flavonoids.

- 5. Alcohols, ketones, esters, and hydrocarbons are responsible for *O.europaea* aromatic odour.^[14]

Traditional medicinal uses

Olive as a whole is used in traditional system of medicines. Some of its medicinal uses are;

- 1. When taken infusion of olive leaf orally, shows an anti-inflammatory action.

2. Lemon juice along with olive oil is used for treating gallstones.^[15]
3. Oral consumption of extract of olive leaves in hot water is used to induce diuresis and for treating hypertension.
4. External application of fruit for fracture limb, sores, burns, rheumatism and to enhance circulation.^[16]
5. For bronchial asthma hot water extraction of dried leaves is used.
6. To treat Urinary tract infection, diarrhoea and respiratory infection decoction of dried leaves and dried fruits is consumed orally.^[17]
7. Use of olive leaves as mouth cleanser, stomach and intestinal diseases.
8. Daily usage of olive oil is used to prevent hair loss.^[18]
9. Its seed oil is consumed orally as cholagogue, for the removal of gall stones in nephritis which is associated with lead intoxication.
10. Dried leaves decoction oral consumption for diabetes.^[19]
11. Orally when seed oil is consumed it act as a laxative and when applied externally it work as pectoral and an emollient.
12. Its fruit are used for nourishment of skin and as skin cleanser.^[20]
13. Boiled extract is used in inflammation.
14. To cure eye infection ointment prepared from boiled extract is used.^[21]
15. Essential oil, orally taken, to treat liver disease and constipation.
16. Tincture prepared from olive leaves is used for growth of nail and as febrifuge.^[22]

Pharmacological effect

1. Anti-microbial activity

In vitro studies has showed that *Staphylococcus aureus*, *Bacillus subtilis*, *Pseudomonas aeruginosa*, *Escherichia coli*, *Klebsiella pneumoniae*, *B. cereus*, *Candida albicans* and *C. neoformans* Olive leaf extract has shown their antimicrobial activity against all these pathogens.^[23] This means that olive leaves is resistant against insect and microbial attack. OLE has shown strong activity against *E.coli* and *S. aureus* when compared with other pathogenic fungi and bacteria.^[24]

The main component responsible for its antimicrobial activity is oleuropein. In spectrum and potency, hydroxytyrosol showed broader antimicrobial activity than oleuropein, which make it comparable with erythromycin and ampicillin.^[25]

2. Anti-inflammatory activity

Luteolin, apigenin, oleuropein and hydroxytyrosol are the chemical constituents found in olive. All of these has shown anti-inflammatory activity. Like studies have shown that both hydroxytyrosol and oleuropein inhibits leukotriene B₄, which is involved in many inflammatory pathways. Moreover, invitro studies in animal models has proved that luteolin and apigenin possess anti-inflammatory as well as anti-allergic effects.^[26]

It was showed by some researchers that dose dependent effect of OLE on carrageenan induced hind paw edema model in rats is anti-inflammatory and it significantly gave the highest activity. It also possess thromboxane A₂ production inhibition and platelet aggregation properties.^[27]

3. Thyroid activities

Olive leaves aqueous extract when administered for 14 days in rats via feedback mechanism it resulted in reduction of circulating thyroid stimulating hormone levels and increase in T₃ level.^[28]

4. Antioxidant activity

Olive leaves contains high polyphenolic content due to which it plays a major role in trapping the free radicals directly making it the part that have highest oxygen scavenging activity.^[29]

Oleuropein is the potent antioxidant which prevents free radical formation by chelating with metal ions (Fe and Cu) and possess anti-inflammatory property.^[30] Oleuropein content is maximum in leaves.

a) olive oil = 0.05 to 0.12%

b) olive leave = 1 to 14%

Its is already shown using in vitro and ex vivo models in an early investigation that the antioxidant property of olive is higher than that of Vitamin E.^[31]

5. Neuroprotector activity

The concept behind olive and role of its constituents in neuroprotection is not clear. It is suggested that the neuroprotective effect of olive is because of the rich phenolic content present in olive. Many studies were conducted to confirm the mechanism involved.^[32]

One such study showed that A β aggregation was decreased when olive leaf extract rich in oleuropein was given to the rat, it reduces the chances of Alzheimer Disease.^[33]

Oleuropein was also found effective against Parkinson disease. A terpenoid named maslinic acid is present on olive leaf, its effect were studied by different scientists. For neuronal death rats were injected with streptozotocin.^[34] Maslinic acid in dose dependant mode exhibited significant neuroprotective activity.^[35]

6. Cardiovascular disorders

Doxorubicin(DXR) clinically known as adriamycin is an anti-neoplastic drug which is highly effective against many malignant diaseases, but because of its serious cardiotoxic side effects which was leading to congestive heart failure, limited its use.^[36]

An investigation was carried in which the effect of oleuropein on cardiotoxicity was observed. A successful result was obtained according to which oleuropein treated DXR-induced cardiotoxicity by decreasing oxidative stress, by inhibiting lipid peroxidation and by reducing nitric oxide species in cardiomyocytes.^[37]

7. Anti-viral activities

Calcium elenolate a derivative of elenolic acid is the constituent present in olive leaf extract.^[38] It was found active against viruses, which means it work as an antiviral agent. Viruses like rhinovirus, myxoviruses, Herpes simplex type I, Herpes simplex type II, Herpes zoster, Encephalomyocarditis, Polio 1, 2, and 3, two strains of leukemia virus, different strains of influenza and para-influenza viruses were all inhibited by calcium elenolate in vitro.^[39]

Reported mechanism of action includes

- a) Inactivation of virus by preventing its mode of spreading (budding, shedding or assembly at the cell membrane).
- b) By interfering with its amino acid production.
- c) Prevents viral replication by directly penetrating into the infected cell.
- d) Neutralizes the production of reverse transcriptase and protease in case of retroviruses.
- e) By stimulating phagocytosis.^[40]

8. Anti-diabetic effect

Olive is being used as antidiabetic herbal drug. According to the published studies, extract of olive in different dosage were fed to the diabetic rats induced with streptozotocin, 5mg/kg i.p. administration, which provides an in vivo evidence with decreased serum glucose level.^[41]

Other study was carried out using mice diabetic model, OLE dosage 20mg/kg, 40mg/kg or 60mg/kg which showed significant reduction in blood glucose. Such evidences proposed in scientific literature suggested that OLE plays beneficial role in Type 2 Diabetes Mellitus.^[42]

Mechanism included;

- a) By activation of AMPK pathway
- b) Increasing incretin release
- c) By antioxidant activity
- d) Inhibiting glucose transport
- e) Inhibiting digestive enzyme activities
- f) By increasing testosterone levels^[43]

9. Skin protector

For brighten, lighten and spot free skin it must that a diet must be full of nutrition, especially antioxidant. Naturally occurring phenolic compounds are rich source of antioxidant. Olive leaf is one such natural herb enriched with polyphenolic compounds. It contains oleuropein, hydroxytyrosol and squalene (trpane hydrocarbon).^[44] Each constituent plays its role in protection of skin from harmful UV radiation, as reported;

- a) Direct antioxidant action – Oleuropein and Hydroxytyrosol
- b) Filtration of oxygen (at skin level) – Squalene.^[45]

10. Anti-atherosclerotic

Oleuropein and oleacin effect on cardiovascular system was reported. The extract worked up the CVS by dilating coronary blood vessels, also an in vitro evidence showed that it causes platelet activation in non-smoking healthy males.^[46]

Side effect and toxicity

There are many research that prove the pharmacological benefits of olive tree, but limited knowledge about their systemic toxicity. Studies pulled out up to the date present the fact that olive leaf extracts are reliable and does not possess any toxic effect even at high doses.^[47]

When the extract supplemented in male and female rats:

- a) Dose of 2000mg/kg – acute toxicity (for 28 days)
- b) 100,200 and 400 mg/kg – subacute toxicity (for 28 days)

Result = NO TOXICITY^[48]

Some histological changes like fatty cytoplasmic vacuolation, hepatocyte necrosis and slight hemorrhage in both liver and kidneys of rats were reported.^[49]

1mg/ml of extract found no toxic effect in in vitro studies on human cell lines, though regarding the safety of OLE in pregnant and lactating women are not yet conducted.^[50]

CONCLUSION

Olea europaea is growing all over world, its immense uses made it a popular herbal plant that supports multifunctional activities including biological, therapeutic and functional food application. Enrichment with polyphenolic compound that is oleuropein, hydroxytyrosol, verbascoside, tocopherol, etc made it a plant with multiple pharmacological activities. It has been used for its antidiabetic, antimicrobial, antiviral, antiageing, neuroprotective activity and many others. During the processing of olive all the waste produced or olive by-product are still full with high amount of bioactive compounds for nutraceuticals, pharmaceuticals and food industries. OLE extract has shown no sign of toxicity, further studies are required related to the safety profile of olive.

REFERENCES

1. Mohamed A. Diaba, Ghada M. Hadad, Amany K. Ibrahim, Review article on chemical constituents and biological activity of *Olea europaea*, RECORDS OF PHARMACEUTICAL AND BIOMEDICAL SCIENCES, 2020; 4(2): 36-45.
2. Guinda, Á., M. C. Pérez-Camino and A. Lanzón "Supplementation of oils with oleanolic acid from the olive leaf (*Olea europaea*).\" European journal of lipid science and technology, 2004; 106(1): 22- 26.
3. Crisosto, C., L. Ferguson and G. Nanos. Olive (*Olea europaea* L.). Postharvest Biology and Technology of Tropical and Subtropical Fruits, Elsevier, 2011; 63-87e.
4. Procopio, A., S. Alcaro, M. Nardi, M. Oliverio, F. Ortuso, P. Sacchetta, D. Pieragostino and G. Sindona "Synthesis, biological evaluation, and molecular modeling of oleuropein and its semisynthetic derivatives as cyclooxygenase inhibitors.\" Journal of agricultural and food chemistry, 2009; 57(23): 11161-11167.

5. Passi, S., O. De Pità, P. Puddu and G. P. Littarru "Lipophilic antioxidants in human sebum and aging." *Free radical research*, 2002; 36(4): 471-477.
6. Md. Yaseen Khan, Siddharth Panchal, Niraj Vyas, Ameer Butani, Vimal Kumar, *Olea europaea*: A Phyto-Pharmacological Review, *Pharmacognosy Reviews*, 2007; 1: 1.
7. M. Gonzalez et al., Hypoglycemic activity of olive leaf. *Planta Medica*, 1992; 58: 513-515.
8. N. Caturla, J. Perez-Fons, A. Estepa, V. Micol, Differential effects of oleuropein, a biophenol from *Olea europaea*, on anionic and zwitterionic phospholipid model membranes. *Chem Phys Lipids*, 2005; 137: 2-17.
9. S. Z. Hirschman, Inactivation of DNA polymerases of murine leukaemia viruses by calcium elenolate. *Nat New Biol.*, 1972; 238(87): 277-9.
10. F. Visioli et al, The effect of minor constituents of olive oil on cardiovascular disease: new findings. *Nutr Rev*, 1998; 56(5 Pt 1): 142-7.
11. Nilüfer Acar-Tek Duygu Ağagündüz, Olive Leaf (*Olea europaea* L. folium): Potential Effects on Glycemia and Lipidemia, *Ann Nutr Metab*, 2020; 76: 10–15.
12. de Bock M, Thorstensen EB, Derraik JG, Henderson HV, Hofman PL, Cutfield WS. Human absorption and metabolism of oleuropein and hydroxytyrosol ingested as olive (*Olea europaea* L.) leaf extract. *Mol Nutr Food Res*, 2013; 57(11): 2079–85.
13. Peyrol J, Riva C, Amiot MJ. Hydroxytyrosol in the Prevention of the Metabolic Syndrome and Related Disorders. *Nutrients*, 2017; 9(3): 306.
14. Omer SA, Elobeid MA, Elamin MH, Hassan ZK, Virk P, Daghestani MH, et al. Toxicity of olive leaves (*Olea europaea* L.) in Wistar albino rats. *Asian J Anim Vet Adv*, 2012; 7(11): 1175–82.
15. Salah MB, Hafedh A, Manef A. Anti-diabetic activity and oxidative stress improvement of Tunisian Gerboui olive leaves extract on alloxan induced diabetic rats. *J Mater*, 2017; 8: 1359–64.
16. Julia Espeso, Alejandro Isaza, Joo Youl Lee, Pia M. Sørensen, Patricia Jurado, Roberto de Jesús Avena-Bustillos, Mikel Olaizola and Juan Carlos Arboleya, OLIVE LEAF WASTE MANAGEMENT, *Frontiers in Sustainable Food System*, 2021; 5.
17. Athanasiadis, V., Grigorakis, S., Lalas, S., and Makris, D. P. Highly efficient extraction of antioxidant polyphenols from *Olea europaea* leaves using an eco-friendly glycerol/glycine deep eutectic solvent. *Waste Biomass Valori*, 2017; 9: 1985–1992.

18. García Martín, J., Cuevas, M., Feng, C., Álvarez Mateos, P., Torres García, M., and Sánchez, S. Energetic valorisation of olive biomass: olive-tree pruning, olive stones and pomaces. *Processes*, 2020; 8: 511.
19. Bownan, A., and Gansey, G. *The Cambridge Ancient History, The Crisis of Empire*, AD 193-337. (Cambridge, MA: Cambridge University Press), 2009; 12: 717–720.
20. Sato, H., Genet, C., Strehle, A., Thomas, C., Lobstein, A., Wagner, A., et al. Antihyperglycemic activity of a TGR5 agonist isolated from *Olea europaea*. *Biochem. Biophys. Res. Commun.*, 2007; 362: 793–798.
21. Sedef N El and Sibel Karakaya, Olive tree (*Olea europaea*) leaves: potential beneficial effects on human health, *Nutrition Reviews*®, 67(11): 632–638.
22. Komaki E, Yamaguchi S, Maru I, et al. Identification of anti-amylase components from olive leaf extracts. *Food Sci Technol Res*, 2003; 9: 35–39.
23. Dimitrios B. Sources of natural phenolic antioxidants. *Trends Food Sci Technol*, 2006; 17: 505–512.
24. Hollman PC, de Vries JH, van Leeuwen SD, Mengelers MJ, Katan MB. Absorption of dietary quercetin glycosides and quercetin in healthy ileostomy volunteers. *Am J Clin Nutr*, 1995; 62: 1276–1282.
25. Visioli F, Poli A, Galli C. Antioxidant and other biological activities of phenols from olives and olive oil. *Med Res Rev*, 2002; 22: 65–75.
26. Al-Khalil, S., A survey of plants used in Jordanian traditional medicine. *International Journal of Pharmacognosy*, 1995; 33: 317-323.
27. Bellakhdar, J., Claisse, R., Fleurentin, J., Younos, C., Repertory of standard herbal drugs in the Moroccan pharmacopoea. *Journal of Ethnopharmacology*, 1991; 35: 123-143.
28. Bianchi, G., Pozzi, N., Vlahov, G., Pentacyclic triterpene acids in olives. *Phytochemistry*, 1994; 37: 205-207.
29. Boskou, D., Tsimidou, M., Blekas, D., Polar phenolic compounds. In: Boskou, D. (Ed.), *Olive Oil, Chemistry and Technology*. AOCS Press, Champaign, IL, 2006; 73-92.
30. Boudhrioua, N., Bahloul, N., Slimen, I.B., Kechaou, N., Comparison on the total phenol contents and the color of fresh and infrared dried olive leaves. *Industrial Crops and Products*, 2009; 29: 412-419.
31. Maryem Ben Salem, Hanen Affes, Kamilia Ksouda, Zouheir Sahnoun, Khaled Mounir Zeghal and Serria Hammami, PHARMACOLOGICAL ACTIVITIES OF OLEA EUROPAEA LEAVES, *Journal of Food Processing and Preservation*, 2014.

32. WOJCIKOWSKI, K., STEVENSON, L., LEACH, D. et al. Antioxidant capacity of 55 medicinal herbs traditionally used to treat the urinary system: A comparison using a sequential three-solvent extraction process. *J. Altern. Complement. Med*, 2007; 13: 103–109.
33. WATERMAN, E. and LOCKWOOD, B. Active components and clinical applications of olive oil. *Altern. Med. Rev*, 2007; 12: 331–342.
34. VISIOLI, F. et al. Oleuropein protects low density lipoprotein from oxidation. *Life Sci*, 1994; 55: 1965–1971.
35. VISIOLI, F. et al. The effect of minor constituents of olive oil on cardiovascular disease: New findings. *Nutr. Rev*, 1998; 56: 142–147.
36. Rabyah B. Ali¹, Jeffrey Victorino de Jesus, Safya Abdulhamid Ammer, Review Olive Leaves as Anti-diabetic, *Asian Journal of Complementary and Alternative Medicine*, 2019.
37. Cumaoglu A, Rackova L, et al. Effects of olive leaf polyphenols against H₂O₂ toxicity in insulin secreting β -cells. *Acta Biochim Pol*, 2011; 58: 45-50.
38. Al-Azzawie HF, Alhamdani MS. Hypoglycemic and antioxidant effect of oleuropein in alloxandibabetic rabbits. *Life Science*, 2006; 78: 1371–7.
39. Petkov V, Manolov P. Pharmacological analysis 42. of the iridoid oleuropein. *Arzneimittel forschung*, 1972; 22: 1476- 1486.
40. Dzulkifi, A.R., Aishah, A.L., Rahmat, A.M., Isa, A.M., Abas, H.H. and Abu baker, M., Development of a plant information system. Present at the international conference on the use of traditional medicine and other natural products in health-care, Penang, Malaysia, 1993; 8-11.
41. Arshad H Rahmani, Aqel S Albutti, Salah M Aly, Therapeutics role of olive fruits/oil in the prevention of diseases via modulation of anti-oxidant, anti-tumour and genetic activity, *Int J Clin Exp Med*, 2014; 7(4): 799-808.
42. Reynolds T, Dweck AC. Aloe vera leaf gel, A review update. *J Ethnopharmacol*, 1999; 68: 3-37.
43. Owen RW, Mier W, Giacosa A, Hule WE, Spiegelhalder B, Bartsch H. Phenolic compounds and squalene in olive oils: the concentration and antioxidant potential of total phenols, simple phenols, secoroids, lignans and squalene. *Food Chem Toxicol*, 2000; 38: 647-59.
44. Keys A. Mediterranean diet and public health: personal reflections. *Am J Clin Nutr*, 1995; 61: 1321S-1323S.

45. Owen RW, Mier W, Giacosa A, Hule WE, Spiegelhalder B, Bartsch H. Phenolic compounds and squalene in olive oils: the concentration and antioxidant potential of total phenols, simple phenols, secoroids, lignans and squalene. *Food Chem Toxicol*, 2000; 38: 647-59.
46. Rahele Ghanbari, Farooq Anwar, Khalid M. Alkharfy, Anwarul-Hassan Gilani and Nazamid Saari, Valuable Nutrients and Functional Bioactives in Different Parts of Olive (*Olea europaea* L.)—A Review, *Int. J. Mol. Sci*, 2012; 13: 3291-3340.
47. Lavelli, V. Comparison of the antioxidant activities of extra virgin olive oils. *J. Agric. Food Chem*, 2002; 50: 7704–7708.
48. Leenen, R.; Roodenburg, A.J.; Vissers, M.N. Supplementation of plasma with olive oil phenols and extracts: Influence on LDL oxidation. *J. Agric. Food Chem*, 2002; 50: 1290–1297.
49. Carluccio, M.A.; Siculella, L.; Ancora, M.A. Olive oil and red wine antioxidant polyphenols inhibit endothelial activation: Antiatherogenic properties of mediterranean diet phytochemicals. *Arterioscler. Thromb. Vasc. Biol*, 2003; 23: 622–629.
50. Galli, C.; Visioli, F. Antioxidant and other properties of phenolics in olives/olive oil, typical compounds of the mediterranean diet. *Lipids*, 1999; 34: S23–S26.