



REVIEW ARTICLE

Phytochemistry and pharmacological properties of *Ocimum gratissimum* (L.) extracts and essential oil - A critical review

Kaliyaperumal Ashokkumar^{1*}, Arjun Pandian², Muthusamy Murugan¹, M. K. Dhanya¹, and Sampathrajan Vellaikumar³

ABSTRACT

The plant *Ocimum gratissimum* is well-known from the ancient Indian medicine system. *O. gratissimum* has wide variety of therapeutic applications. Folk medicine says that it can help with headaches, fevers, diarrhoea, pneumonia, and other ailments. *O. gratissimum* contains several bioactive constituents widely used as food additives, food colorants, pharmaceuticals, pesticides, and fragrances. This review discusses up to this point data on the phytochemical and pharmacology of *O. gratissimum* extracts and oil from numerous locations worldwide. Pertinent data of *O. gratissimum* was earned from numerous electronic scientific databases, and additional information was obtained from books, thesis and different relevant websites. The yield of the *O. gratissimum* essential oil (OGEO) varied between 0.12% and 1.66%. OGEO was predominantly accumulated phenylpropenes, (55.7%-57.3%) followed by sesquiterpenes (27.5% - 38.1%), and monoterpenes (4.0%-16.1%). Eugenol, germacrene-D, β -ocimene, 1,8-cineole, β -selinene, caryophyllene, γ -murolene, p-cymene, and thymol, are major constituents of OGEO from various origins.

Keywords: *Ocimum gratissimum*; Essential oil; Eugenol; β -ocimene; Phytochemistry; Biological activities

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*Corresponding author email address: biotech.ashok@gmail.com (K. Ashokkumar)

INTRODUCTION

Ocimum gratissimum (L.) is known as Ram tulsi in India. A native of Asia, it is primarily grown in India, Sri Lanka and Nepal (Nadkarni, 1999). This plant's leaves are used to cure anxiety, headaches, and bronchitis (Rabelo et al., 2003; Matasyoh et al., 2007). Traditional medicine has long used *O. gratissimum*. It is also used medicinally, as a condiment, and in cooking. Locally, it is used to treat diarrhoea (Effraim et al., 2003).

Hydrodistillation, steam distillation, microwave, ultrasound-assisted extraction, and supercritical fluid extraction are all methods used to extract essential oils from plants (Azwanida, 2015; Ashokkumar et al. 2020a; Ashokkumar et al., 2020b). Due to the lower cost of the Clevenger apparatus and the use of water as a solvent, the hydrodistillation procedure is the most extensively employed by researchers worldwide (Ashokkumar et al., 2020c). The amount of essential oil produced by *O. gratissimum* ranges from 0.21 to 0.70 percent (Dubey et al., 2000; Matasyoh et al., 2007; Joshi, 2013; Ashokkumar et al., 2020d). Several studies on the OGEO have been carried out all across the world (Matasyoh et al., 2007; Joshi, 2013; Matasyoh et al., 2007; Padalia and Verma, 2011).

The OGEO is rich in phytoneleypropene (eugenol & methyl eugenol), sesquiterpenes (germacrene D & caryophyllene, - muurolene), monoterpenes (-ocimene), and other components. free of charge (Matasyoh et al., 2007; Padalia and Verma, 2011; Joshi, 2013). Many research papers claim that *O. gratissimum* (Offiah and Chikwendu, 1999; Aguiyia et al., 2000; Lahlou et al., 2004; Trevisan et al., 2006; Joshi, 2013; Ajayi et al., 2014; Gontijo et al., 2014; Aderibigbe and Idowu, 2020). This review looks on the phytochemical compositions of *O. gratissimum* and their possible pharmacological activities. However, to better understand the mechanism of

bioactive elements and their intake in animals and humans, which can aid in the prevention of many diseases.

O. GRATISSIMUM ESSENTIAL OIL (OGEO) AND ITS COMPOSITION

The EO yield from *O. gratissimum* varied between 0.12% and 1.66% (Table 1). The profiling of EO of aerial parts of *O. gratissimum* sampled from Western Ghats of southern India predominantly exhibited eugenol (54.42%), germacrene D (15.43%), β -ocimene (12.37%), caryophyllene (4.59%), and γ -muurolene (3.05%), (Ashokkumar et al., 2020d). However, the Brazil grown leaves of OGEO chiefly contained eugenol, 1,8-cineole, and β -selinene. Furthermore, grown aerial parts of OGEO was predominant in p-cymene and thymol (Table 2).

OGEO was predominantly accumulated phenylpropenes, (55.7%- 57.3%) followed by sesquiterpenes (27.5% - 38.1%), monoterpenes (4.0%-16.1%), (Joshi, 2017; Ashokkumar et al., 2020d). The yield of minor constituents of OGEO include β -pinene (0.1%), borneol (0.1%), carvenone (0.1%), α -humulene (0.1%), elemol (1.2%), caryophyllene oxide (0.15%), α -thujene (0.17%), γ -elemene (0.2%), γ -terpinene (0.2%), α -pinene (0.2%), camphor (0.3%), humulene (0.3%), δ -Cadinene (0.4%), sabinene (0.5%), linalool (0.5%), β -bourbonene (0.5%), α -copaene (0.6%), cis-verbenol (0.7%), terpin-4-ol (0.7%), isobornylformate (0.9%), isoledene (1.0%), (Joshi, 2017; Ashokkumar et al., 2020d). Figure 1 depicts the major EO components' molecular structures extracted from *O. gratissimum*.

PHARMACOLOGICAL PROPERTIES OF OGEO AND EXTRACTS

Antioxidant activity

Antioxidants are naturally or artificially derived compounds that could prevent the free radical formation and suppress chronic and degenerative diseases by scavenging free radicals (Haliwell, 2000). Natural antioxidants derived from herbal sources are currently gaining popularity (Velioglu et al., 1998).

Table 1. Yield of EO from *O. gratissimum*

| Technique or method | Oil yield (%) | Authors |
|---------------------|---------------|----------------------------------|
| Hydro-distillation | 1.66 | Melo et al. (2019) |
| Hydro-distillation | 0.60 | Ashokkumar et al. (2020d) |
| Hydro-distillation | 0.65 | Padalia et al. (2014) |
| Hydro-distillation | 1.10 | Verma et al. (2016) |
| Hydro-distillation | 0.12 – 0.78 | Matasyoh et al. (2008) |
| Hydro-distillation | 0.65-0.78 | Kpadonou Kpoviessi et al. (2012) |
| Steam distillation | 1.33 | Ibeh et al. (2017) |

Table 2. Major EO composition of *Ocimum gratissimum* L.

| Origin | Major constituents | Yield (%) | Authors |
|--|--|---|----------------------------------|
| Western Ghats, South India (Kerala) | eugenol germacrene D β-ocimene caryophyllene γ-muurolene | 54.4% 15.4% 12.4% 4.6% 3.1% | Ashokkumar et al. (2020d) |
| Brazil | Eugenol 1,8-cineole β-selinene | 74.3% 15.2% 2.8% | Melo et al. (2019) |
| North India | eugenol germacrene D | 78.0% 4.4% | Padalia et al. (2014) |
| Peninsular India (Karnataka) | eugenol caryophyllene oxide (Z)-β-ocimene | 53.0% 7.2% 3.5% | Verma et al. (2016) |
| Kenya | eugenol methyl eugenol | 68.8 % 13.2% | Matasyoh et al. (2008) |
| Portugal | Thymol p-cymene | 48.1% 12.5% | Martins et al. (1999) |
| Colombia | eugenol 1,8-cineole β-selinene | 43.2% 12.8% 9.0% | Benitez et al. (2009) |
| Benin | p-cymene thymol γ-terpinene α-thujene myrcene | 28.1-53.8% 3.3-29.1% 1.1-10.9% 3.4-10.8% 4.2-8.3% | Kpadonou Kpoviessi et al. (2012) |

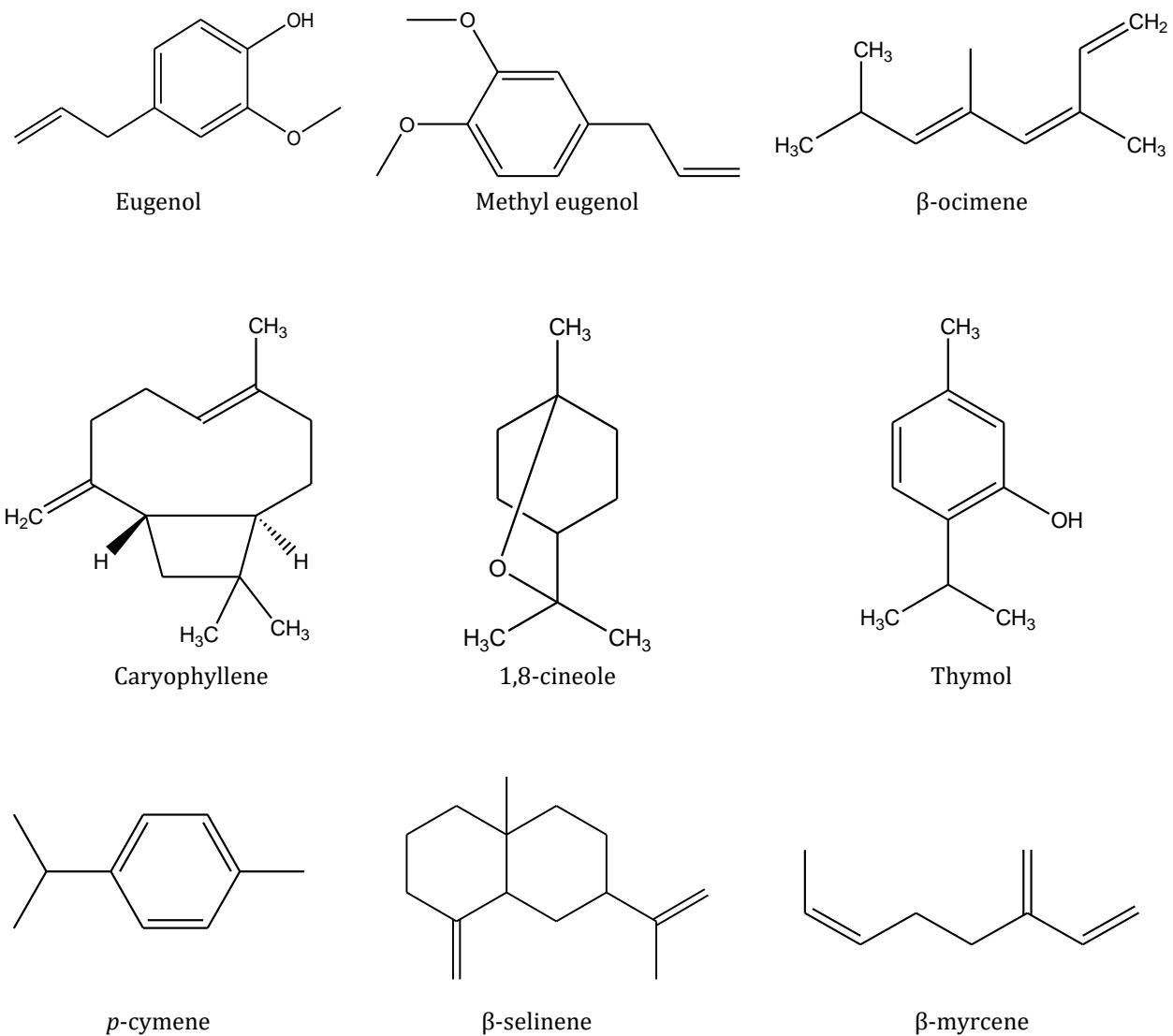


Figure 1. Molecular structures of major OGEO constituents

In another study, DPPH and ABTS models, OGEO demonstrated comparative antioxidant activity with IC₅₀ values of 23.66 and 23.91 g ml⁻¹, respectively. This study also noted that eugenol had marginally lower antioxidant activity than OGEO. In contrast, *O. sanctum* oil had very low antioxidant activity (Joshi et al. 2013)

OGEO has antifungal action against *Candida albicans*. On average, 0.24 mg ml⁻¹ inhibited the most (Kpadonou Kpoviessi et al., 2012). O. ethanolic extract In human dental plaque, gratissimum was antibacterial against *Actinobacillus actinomycetemcomitans*. Compared to 0.2 percent chlorhexidine and dimethyl sulfoxide (DMSO), 0.6 percent extract has potential antibacterial action (Eswar et al., 2016).

Antimicrobial activities

In another investigation, OGEO at 0.24 mg ml⁻¹ and 0.95 mg ml⁻¹ demonstrated considerable

antibacterial action (Kpadonou Kpoviessi et al., 2012). Joshi (2017) tested OGEO and eugenol against 13 bacterial species, including *S. S. S. faecalis*, *M. flavus*, *M. luteus*, *B. subtilis*, *E. O157:H7*, *Proteus mirabilis*. Eugenol had a MIC of $0.33 - 3.33 \text{ mg ml}^{-1}$ and considerable inhibitory action at 1.04 mg ml^{-1} against *S. aureus*. The essential oil had a substantial inhibitory impact at 0.29 mg ml^{-1} against *S. marcescens*.

Insecticidal activity

The insecticidal activity of OGEO was tested against *Sitophilus zeamais* (a major stored pest of maize) by mixtures OGEO (5%) and kaolin (10%). Results remarked that OGEO on the test insects was possessed an 85.7% knockdown effect (Jirovetz et al., 2005; Benelli et al., 2019). Other insecticidal and acaricidal activities were also summarized in Table 3.

Miscellaneous activities

Eugenol from *O. gratissimum* has reported that antiviral activity, which inhibits the HSV-1,2 replication. The bioactive constituent thymol also destructs the virion of HSV-1 (Tshilanda et al., 2020). Other miscellaneous activities were summarized in Table 3.

CONCLUSION

This review discusses the phytochemistry and pharmacology of *O. gratissimum*. Ancient and modern Indians have utilised gratissimum to treat a variety of diseases. *O. gratissimum*, has been used to treat anxiety, headaches, and bronchitis. *O. gratissimum*, say Indian medicinal texts. Historically, gratissimum has been trustworthy, but current pharmacological experiments on *O. gratissimum*.

O. gratissimum, has over 75 secondary metabolites. According to current research, eugenol is the most important bioactive molecule with numerous potential health benefits. Also, *O.* research on gratissimum extracts and OGEO (Table 3). However, *O* research is lacking, gratissimum, and we have suggested some themes for additional research.

First, investigations on metabolite structure in *O.* Phytochemistry reports limit the use of *gratissimum* leaves and aerial portions. Second, little research has been done on preserving OGEO's shelf life.

Future studies need to focus on studying structural characterization of metabolites, shelf-life quality of OGEO, proper experimental setup conduct with negative or positive control and correct MIC values, and finally, clinical investigation implemented with humans is essential.

Table 3. The activities of *O. gratissimum* extracts and essential oil components

| Pharmacological activities | Extract/ essential oil | <i>In vitro</i> / <i>In vivo</i> | Target/ Model | Control(s) | IC 50/ Dosage | Results / Remarks | Reference |
|----------------------------|------------------------|-------------------------------------|------------------------------|--|---|---|----------------------------------|
| Antibacterial activity | Essential oil | <i>In vitro</i> | <i>Staphylococcus aureus</i> | Positive: Doxycycline | MIC : 0.24mg ml ⁻¹ MBC: 0.95mg ml ⁻¹ | Moderate antibacterial activity | Kpadonou Kpoviessi et al. (2012) |
| Antibacterial activity | Essential oil | <i>In vitro</i> | <i>E. coli</i> | Positive: Doxycycline | MIC: 0.48mg ml ⁻¹ | Noteworthy antibacterial activity | Kpadonou Kpoviessi et al. (2012) |
| Antibacterial activity | Essential oil | <i>In vitro</i> | 17 microbial species | Positive: Erythromycin & Amikacin | MIC: 0.29 to 1.51 mg ml ⁻¹ | Best inhibitory effect @ 0.29 mg/ml against <i>S. marcescens</i> | Jhoshi, (2017) |
| Antibacterial activity | Eugenol | <i>In vitro</i> | 17 microbial species | Positive: Erythromycin & Amikacin | MIC: 0.33 to 3.33mg ml ⁻¹ | Best inhibitory effect @ 1.04 mg/ml against <i>S. aureus</i> | Jhoshi, (2017) |
| Antifungal activity | Essential oil | <i>In vitro</i> | <i>Candida albicans</i> | Positive: Nystatin Negative: Solvent | MIC: 0.06 to 0.25 mg ml ⁻¹ MFC: 6.25 to 12.50 mg ml ⁻¹ | Greatest inhibitory effect @ 0.24 mg/ml against <i>Candida albicans</i> | Kpadonou Kpoviessi et al. (2012) |
| Antioxidant activity | Essential oil | <i>In vitro</i> | DPPH | - | IC ₅₀ : 23.66 µg ml ⁻¹ | Significant antioxidant activity observed | Jhoshi, (2013) |
| | | | ABTS | - | IC ₅₀ : 23.91 µg ml ⁻¹ | | |
| Antioxidant activity | Eugenol | <i>In vitro</i> | DPPH | - | IC ₅₀ : 27.16 µg ml ⁻¹ | Eugenol showed lesser antioxidant activity compared to OGE | Jhoshi, (2013) |
| | | | ABTS | - | IC ₅₀ : 32.16 µg ml ⁻¹ | | |

| | | | | | | | |
|----------------------------|------------------------|-----------------|--|--|---|---|------------------------|
| Anthelmintic activity | Acetone extract | <i>In vitro</i> | Adult <i>H. placei</i> nematodes using adult worm motility assay | Negative: Normal Saline | IC ₅₀ : 5-60 mg ml ⁻¹ | Best anthelmintic activity recorded @ 56.04 mg/mL against <i>H. placei</i> | Segun & Sunday (2020) |
| Anti-inflammatory activity | Hydroethanolic extract | <i>In vivo</i> | Rats anesthetized by intraperitoneal injection of 25 mg/kg of thiopental sodium | Positive: - Negative: Distilled water | 100, 200 or 400 mg kg ⁻¹ b.w | The inhibition by the extract was not dose dependent as it was 15.2, 26.7 and 22.4% for 100, 200 and 400 mg/kg respectively | Ajayi, et al. (2014) |
| Anti-inflammatory activity | Aqueous extract | <i>In vivo</i> | Carrageenan-induced paw oedema in rats. | Negative: Distilled water | 100, 200 and 400 mg kg ⁻¹ | Concentration 400 mg kg ⁻¹ substantially increased glutathione level. | Alabi et al. (2019) |
| Antinociceptive activity | Essential oil | <i>In vivo</i> | Swiss albino mice (25-30g) induced pain. Writhing and formalin test | Positive: Indomethacin | 30, 100, 300mg kg ⁻¹ (p.o) | Dose dependent inhibition observed. OGEO possessed antinociceptive properties in the writhing and formalin test | Rabelo et al. (2003) |
| Antinociceptive activity | Aqueous extract | <i>In vivo</i> | Acetic acid-induced nociception in mice. Hot plate test | Negative: Distilled water | 200 and 400mg kg ⁻¹ | Dose dependent Inhibition was observed at TADP concentration of 100–400 mg kg ⁻¹ . | Alabi et al. (2019) |
| Gastroprotective activity | Methanolic extract | <i>In vivo</i> | Stress induced ulcer in rats | - | 200, 400, 800 mg kg ⁻¹ | Decreased ulcer indices in a dose dependent manner | Akah et al. (2007) |
| Antidiabetic activity | Aqueous extract | <i>In vivo</i> | Intraperitoneal administration of (65 mg/kg), Type 1 Diabetes mellitus (DM 1) rats | Negative: Distilled water | Not reported | OG extract showed antidiabetic activity | Okon and Umoren (2017) |

| | | | | | | | |
|-----------------------|---------------|----------------|--|---|--|--|-------------------------|
| Insecticidal activity | Essential oil | <i>In vivo</i> | Cockroach (<i>Nauphoeta cinerea</i>) nymphs (20 days of age) were used | - | LC ₅₀ : 50 to 1000 µg of oil per ml of air. | O GEO has substantial insecticidal properties at lethal concentration (LC ₅₀) of 516 µg ml ⁻¹ | Rodrigues et al. (2020) |
| Acaricidal activity | Essential oil | <i>In vivo</i> | <i>Rhipicephalus microplus</i> (Acari: Ixodidae) | - | LC ₅₀ : 0.84 mg ml ⁻¹ LC ₅₀ : 1.58 mg ml ⁻¹ | Study remarked that seasonal variation in the O GEO and its influences acaricidal activity | Silva Lima (2018) |

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