

A Review on Endodontic Irrigants: Herbal versus Conventional

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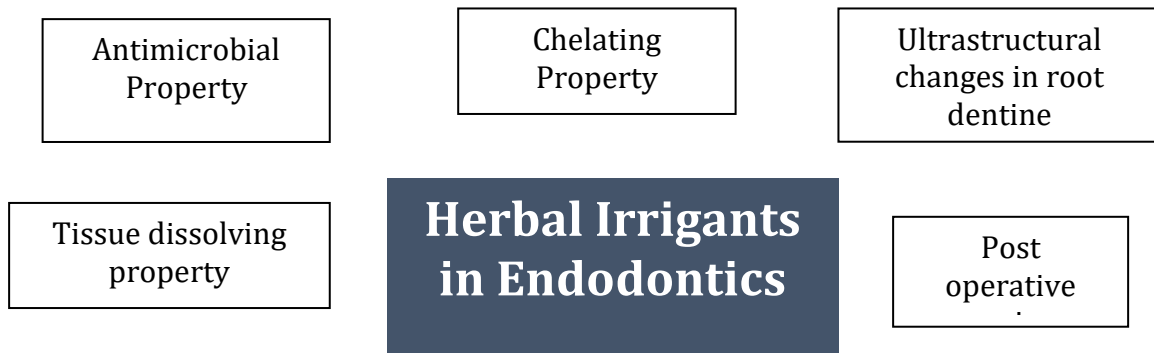
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ABSTRACT

The intricate structure of the root canal system and the resistance mechanisms of multi-species microbial biofilms are the main challenges of root canal therapy, which provide substantial obstacles for irrigants. Herbal irrigants, derived from natural sources, have gained attention for their potential antimicrobial properties and reduced side effects compared to conventional irrigants. This article presents a comprehensive and comparative analysis of herbal irrigants over conventional irrigants in root canal therapy.

Keywords: Root canal, irrigation, herbal, Sodium hypochlorite, Ethylenediamine tetraacetic acid, M. citrifolia juice, Triphala, Green Tea, Proanthocyanidins, Chlorhexidine, Azadirachta Indica, Arnica montana, anti-oxidant, oregano extract, cyclooxygenase

Graphical Abstract:



INTRODUCTION

Endodontics is a specialty that primarily focuses on the study of structure and function of the tooth, the pulp and

periradicular tissues, and its necessary treatment. Mechanical shaping of the root canal system is required for effective placement of intracanal medicaments and

subsequent obturation. However, irrigation performed before, during and post instrumentation, is crucial for thorough disinfection and elimination of contaminated necrotic tissue.

Over the past two decades, irrigation has become increasingly vital for successful root canal therapy. Challenges for effective root canal irrigation are multifaceted.

Aside from the flushing action, biologically effective irrigation solutions are necessary to thoroughly clean the main canal and its intricacies. These areas can harbor biofilm and dentin debris, protecting underlying biofilm from instruments and irrigants. The presence of multi-species microbial biofilms with the ability to adhere to dentin surfaces poses a significant challenge to conventional irrigation strategies.

Therefore, an ideal irrigant should possess antimicrobial properties, an ability to dissolve the remnant necrotic pulp tissue, and should pose minimal irritation to the periapical tissue.¹

Currently used Irrigants:

Tissue dissolving agent

- Sodium Hypochlorite

Sodium hypochlorite (NaOCl) is primarily used as a root canal irrigant owing to tissue dissolving properties. It has an antibacterial effect, particularly against bacteria in biofilms. The principal ingredients of a concentrated hypochlorite solution are hypochlorite and sodium hydroxide which exhibits actions based on the pH and the concentration of available chlorine. Hypochlorous acid (HOCl) is a weak acid and dissociates to the hypochlorite ion (-OCl) and proton (H⁺) depending on the solution pH. Chlorine (Cl) atoms in HOCl and OCl⁻ act on various biological molecules. It is generally believed that HOCl is the active species in the germicidal action, whereas the concentration of -OCl is a key factor determining the cleaning efficiency.²

Regarding concentration of the solution there is no consensus on the ideal concentration, with recommendations

varying from 0.5 to 8.25%³. While higher concentrations may be beneficial, evidence supporting this is limited. It is well known that NaOCl is caustic, and its accidental extrusion into periapical tissues should be avoided. There is a lack of clinical evidence linking NaOCl concentration to the severity of such accidents.

Pre-heating to 50–60°C prior to irrigation has shown to enhance the effectiveness of low-concentration NaOCl solutions⁴. However, rapid cooling occurs after application in vivo, limiting any potential short-term benefits.⁵

Chelating Agents

- Ethylenediamine tetraacetic acid (EDTA)

NaOCl has limited action on the inorganic smear layer components and hard tissue debris thus necessitating the additional action of a demineralizing agent. Studies have shown that NaOCl acts on the organic content of the smear layer facilitating its removal. Most widely used chelating agent is ethylenediamine tetraacetic acid (EDTA^{6,7}. After instrumentation, a 15–17% solution of its disodium salt, with a pH of 7-8, is a strong chelator that dissolves the inorganic content of smear layer and hard tissue fragments^{8,9}

EDTA also appears to disturb the biofilm matrix, which encourages its separation¹⁰. This suggests that EDTA may also function in tandem with NaOCl's anti-biofilm activity & smear layer removal. Other chelators that have been used instead of EDTA are Maleic acid, citric acid and HEDP (1-hydroxy ethylidene 1,1-diphosphonate) also known as etidronic acid.

Antimicrobial agents

- Chlorhexidine (CHX)

Earlier studies have shown that CHX is as effective as or even more successful than NaOCl against bacteria¹¹. These results were based on the overuse of *Enterococcus faecalis* as a test species in most of the studies. (Swimberghe et al., 2019)¹².

However current literature has shown that endodontic microbiology in root canal is polymicrobial and multispecies biofilm plays a major role. Therefore, it is concluded that CHX is not as effective as NaOCl as an antibacterial agent and has limited action against the EPS matrix^{10,13}. The possibility of CHX binding to dentine and providing a long-lasting antibacterial action (substantivity) is one of the primary arguments in favor of the drug, since it may help to prevent bacterial recolonization following root canal therapy¹⁴

Combination Irrigants

No one irrigant exists that has optimal tissue dissolving, demineralizing and anti-bacterial action. In order to improve the characteristics, combination irrigants are available in the market. To improve wettability and decrease the surface tension, surfactants are occasionally added. Other formulations have combined irrigant with antibiotic, chelating agent, and one or more surfactants (Figure 1). One example is Bio Pure MTAD (Dentsply Sirona, Charlotte, NC, USA), QMix, Tetra clean.

Dioguardi, et al.: Methods for endodontic irrigation

Table 2: Advantages/disadvantages of irrigants mainly used in the clinical practice							
Irrigants	Parameters						
	Efficacy on organic residues	Efficacy on inorganic residues	Antibacterial activity	Damage to the surrounding periodontal tissues	Manageability (bad smell, clothes staining and others)	Cost	Enlargement of dental tubules
NaOCl	Yes	No	Yes	Yes	Bad	Low	High
EDTA	No	Yes	No	Very low	Good	Low	High
Citric acid	Yes	Yes	Yes	Yes	N/A	Low	Low
Digluconate chlorhexidine	No	No	Yes	No	Good	Low	Not influenced
Tetraclean	Yes	Yes	Yes	No	Good	High	High
BioPure MTDA	Yes	Yes	Yes	No	Good	High	High
QMix	Yes	Yes	Yes	No	Good	High	High

EDTA: Ethylenediaminetetraacetic acid, MTDA: Mixture of Tetracycline Detergent Acid, N/A: Not reported

Figure 1. Methods for endodontic irrigation¹⁵

Disadvantages of Conventional Irrigants:

- Owing to the caustic nature of NaOCl, its inadvertent extrusion of even 1% concentration into periapical tissue can have adverse reactions^{16,17}
- EDTA has a weak antimicrobial effect. When used in combination with NaOCl, there is a loss of free available chlorine¹⁸, reducing the efficacy of NaOCl. Furthermore, this combination adversely brings about erosion and decalcification of root dentin which has an overall negative effect on dentin flexural strength.
- Maleic acid and Citric acid^{19, 20}. in combination with NaOCl have shown to reduce the available chlorine [18]. The antimicrobial activity of citric acid is very limited²¹.
- CHX when used in similar concentration as NaOCl, is equally or more cytotoxic²². Additionally, it reacts with

residual NaOCl in the root canal and forms para-chloroaniline which is a potentially toxic, carcinogenic orange-brown precipitate²³ that may also cause discoloration²⁴.

Herbal Endodontic Irrigants

Herbal and natural products have been used in dentistry and medicine for thousands of years. Renewed awareness and popularity in recent years may be owing to the advantages - affordability, ease of use, longer shelf life, biocompatibility, antioxidant and anti-inflammatory properties. There are a variety of herbal products available which have the potential to match the effect or to be used as adjuncts to conventional irrigants.

Kale and Raut have proposed classification based on the properties of the currently available herbal irrigants (Figure 2)

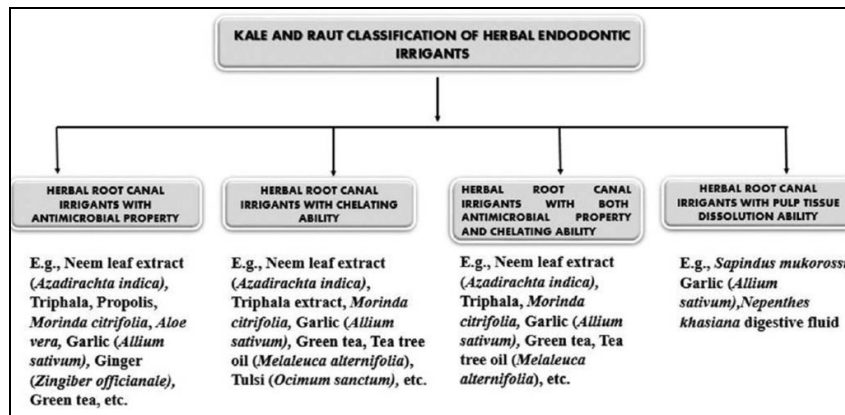


Figure 2. Herbal endodontic Irrigant classification²⁵

Properties of Herbal Irrigants in Endodontics

1. Smear layer removal
2. Antimicrobial efficacy
3. Post operative pain
4. Ultrastructural changes in root dentine
5. Tissue dissolving property

1. Smear layer removal:

Elimination of the smear layer allows for deeper penetration of irrigants into the root canal intricacies and facilitates the proper contact of intracanal medication.²⁶

Regarding smear layer removal efficiency, Murray et al., 2008 demonstrated that 6% *M. citrifolia* juice (MCJ) was equally effective as 6% NaOCl when 17% liquid EDTA was used as final flush.²⁷ Another study by Ok et al., 2015 oregano extract 1 or 2% had a similar smear layer eradication efficacy to 5.25% NaOCl followed by 17% EDTA.

In a recent systematic review²⁸ authors stated that additional factors also play a role such as volume of the irrigant used and the contact time. The smear layer removal efficiency of oregano extract was found to be similar to that of the conventional agent (NaOCl, EDTA). However, it was concluded that herbal agents show inferior smear layer removal when compared to EDTA.

2. Antimicrobial efficacy:

- *M. citrifolia* juice (MCJ) –
M. citrifolia juice (MCJ) is a biocompatible antioxidant and contains antibacterial

compounds such as L-asperuloside and alizari. It has shown potential to be used as an endodontic irrigant. Unlike NaOCl, accidental extrusion does not bring about any adverse effects. It has been shown to be more effective than 2% CHX against *E. faecalis*. However, MCJ and CHX are not effective when used in combination because they neutralize each other's properties.

An in-vitro study on extracted teeth evaluated for the antimicrobial activity of 2% CHX gel, propolis, MCJ and Ca (OH)₂ in infected root canal dentin. It was concluded that propolis and *M. citrifolia* were more effective against *E. faecalis*.

- Triphala

Triphala is a popular ayurvedic herbal formulation consisting of dried and powdered fruits of three medicinal plants namely *Terminalia bellerica*, *Terminalia chebula*, and *Embilica officinalis*.²⁹ Triphala constitutes a number of compounds such as Tannin, Quinones, flavins, flavonoids, flavanols, gallic acid and Vitamin C. Tannins have the ability to inactivate microbial adhesins, enzymes, and cell envelope transport proteins whereas quinones target the microbial cell by acting on surface-exposed adhesins, cell wall polypeptides and membrane-bound enzymes.

A systematic review compared the antimicrobial action of herbal irrigants and NaOCl. Amongst the herbal agents used were triphala, green tea polyphenol, and *Morinda citrifolia*. Triphala was found to be

the most effective antimicrobial agent, followed by green tea polyphenol, and *Morinda citrifolia*.²⁷

- Green Tea

Green tea polyphenols (GTPs) are prepared from the young shoots of the tea plant *Camellia sinensis*. The antimicrobial activity is due to inhibition of bacterial enzyme gyrase by binding to ATP B subunit.

Green tea has been shown to exhibit antibacterial activity on *E. faecalis*³⁰ but has shown reduced antibacterial effect in comparison to NaOCl.

The antibacterial properties have been associated with the polyphenol catechin fraction of green tea leaves which constitute up to 30%. There are four main types of catechins: Epigallocatechin-3-gallate (EGCG), epigallocatechin, epicatechin-3-gallate and epicatechin. EGCG has been shown to cause irreversible membrane disruption in both Gram-positive and Gram-negative bacteria and also to inhibit bacterial DNA gyrase, leading to bacterial cell death.³¹

- *Azadirachta Indica*

It is known as the Indian Neem or Margosa tree. It has antibacterial properties against *E. faecalis* and *Candida albicans*. It has strong antioxidant and antimicrobial properties, making it a potential agent for root canal irrigation.³²

The antimicrobial efficacy of 2.5% sodium hypochlorite and 0.2% chlorhexidine gluconate were compared with an experimental irrigant formulated from *A. indica* and found that neem irrigant has potent antimicrobial efficacy and can be considered for endodontic use.³³

3. Post operative pain

Postoperative discomfort immediately following the root canal therapy is an important concern for both patients and professionals (Kherlakian et al., 2016 and Farzaneh et al., 2018). 1.7 to 70% of patients experienced postoperative

discomfort. Furthermore, within the first 24 hours, that percentage went up to 80%³⁴

According to Hosny et al, an experimental study where Neem versus 2.5% NaOCl as irrigants were used to record the intensity of postoperative pain and number of endotoxins following root canal treatment of mandibular molars.

It was seen that the intensity of the pain gradually decreased over time, with no discernible difference between them at any of the follow-up times except for the first 24 hours where neem group recorded less pain than the control group. Also, Neem showed a positive correlation between high antibacterial effect with reduced endotoxin levels.

It is proposed that the potent antibacterial, anti-inflammatory, antipyretic, analgesic, and biocompatible properties of Neem play a vital role in reducing pain. Additionally, Neem inhibits bacterial adhesion and their capacity to colonize (Hannah et al. 2013)³⁵ According to Khetarpal³⁶, the active components of neem such as nimbidin and nimbolide, can trigger bacterial cell wall lysis when present. By controlling pro-inflammatory enzymes including cyclooxygenase (COX) and lipoxygenase (LOX), neem also functions as an anti-inflammatory agent.

4. Ultrastructural changes in root dentine

Endodontic treatment aims to disinfect the root canal system and achieve 3D hermetic seal to prevent the entry of microorganisms³⁷

Preservation of collagen present in root dentin has shown to have beneficial bonding at core material, sealer and root dentin. However, commonly used root canal irrigants such as NaOCl & EDTA have shown deleterious effect on the root dentin by causing degradation of collagen.³⁸

According to Kumar PS et al, chloramines and protein-derived by-products dissociate when NaOCl reacts with organic portion of the root dentin. These by-products have adverse effects on the pyridinoline cross-links present within the type 1 collagen

leading to lower bond strength. Therefore, irrigation with NaOCl leads to structurally compromised collagen in the root dentin.³⁹ Currently the focus is on stabilization of dentin collagen and neutralization of bacterial collagenase activity which would result in improved mechanical integrity of root filled teeth.

Research into increasing collagen cross-linking has been applied to improve the mechanical and biological stability of dentin collagen. Synthetic collagen cross-linking agents are available which can induce exogenous cross-links but have disadvantages such as high cytotoxicity and incompatible mechanical properties.⁴⁰

Thus, an endodontic disinfectant that kills *E. faecalis* whilst not compromising the biomechanical properties of dentine would be beneficial for endodontic irrigation regimes. Herbal irrigants used during instrumentation have antioxidant properties in addition to being an anti-bacterial & anti-inflammatory agent. The antioxidant property can lead to scavenging of the oxidative by-products thereby increasing the adhesion of sealer to the root dentin (Houde V et al., 2006)⁴¹. Previous studies have shown that they may cause inhibition of collagen breakdown along with stiffening and cross-linking of the collagen.

A study was done to assess the effect of the pushout bond strength (PBS) of epoxy resin-based root canal sealer to root dentin following the use of two novel herbal irrigants, 10% proanthocyanidins (PAs) and 2% *Azadirachta indica* (neem extract) with 3% sodium hypochlorite. It was concluded that the PBS of 10% Proanthocyanidins was the highest, followed by 3% NaOCl and *Azadirachta indica*. Polyphenolic structures

of proanthocyanidins that are capable of forming stable hydrogen bond structures and produce non-biodegradable collagen matrices.⁴²

Another study conducted by Khanvilkar et al found that the binding between AH Plus and dentin was greatly increased by irrigation with *Arnica montana*. At a dosage of 5 mg/mL, *Arnica montana* demonstrates 63.68% total antioxidant activity and 71.52% DPPH scavenging capability using the phosphomolybdate method. It demonstrated the maximum binding capacity to resin sealer due to its anti-inflammatory and antioxidant properties, which mitigate the deleterious effects of free radicals.⁴³

5. Tissue dissolving property

Sodium hypochlorite is a potent tissue solvent and antibacterial agent, however this chemical solution is harmful to periapical tissues only if it is extruded from the apical foramen⁴⁴ and its toxicity increases with the concentration. Numerous studies have shown clinically that sodium hypochlorite damages the periodontal and periapical tissues.

A study conducted by. Guçluer et al⁴⁵ showed extract solutions prepared from different solvents of *S. mukorossi* displayed a statistically significant effect as a tissue solvent. According to the study, the effective cause of the solvent ability might be linked to the structure of saponin in *S. mukorossi*, which reduces surface tension, thus increasing membrane permeability. Many studies revealed that the surfactant addition to NaOCl performed a synergistic action and affected the tissue dissolving specification positively.⁴⁶

Properties	Herbal Irrigants
Herbal root canal irrigants with antimicrobial property – Extracts have demonstrated antimicrobial property against common endodontic pathogens (Kale and Raut et al., 2021) ²⁵	<ul style="list-style-type: none"> • Neem leaf extract (<i>Azadirachta indica</i>), • Triphala, Propolis, • <i>Morinda citrifolia</i>, • Aloe vera, • Garlic (<i>Allium sativum</i>), • Ginger (<i>Zingiber officinal</i>),

	<ul style="list-style-type: none"> • Green tea, Tea tree oil (Melaleuca alternifolia), • Carvacrol • Turmeric (Curcuma longa), • Arctium lappa, • Babool (Acacia nilotica), • Tulsi (Ocimum sanctum), • Cinnamon (Cinnamomum zeylanicum), • Miswak (Salvadora persica), • Passion fruit juice, Jeeryin, • Clove (Syzygium aromaticum), • Hybanthus enneaspermus • Guava (Psidium guajava), • Terminalia chebula seed extract, • Mangifera indica L. kernel (Mango kernel), • Zataria multiflora Essential oil, • Andrographis ukorossi, • Thymus vulgaris, • Calendula arvensis, • Mimusops elengi, • Syzygium cumini
<p>Herbal root canal irrigants with chelating ability – These extracts possess the ability to remove the smear layer formed on the dentin surface after instrumentation (Kale and Raut, 2021) ²⁵</p>	<ul style="list-style-type: none"> • Neem leaf extract (A. indica) • Triphala extract • M. citrifolia, garlic (A. sativum) • green tea, tea tree oil (M. alternifolia) • Tulsi leaf extract (O. sanctum) • Miswak (S. persica) • Grapefruit (Citrus paradisi) • Passion fruit juice, orange oil • German chamomile (Marticaria recutita) • T. chebula seed extract • Amla (Emblica officinalis) • Pomegranate peel (Punica granatum), • Citrus aurantifolia-Sapindus ukorossi extracts • Moringa oleifera • Lemon grass • Turmeric (C. longa)
<p>Herbal root canal irrigants with both antimicrobial property and chelating ability – These extracts have demonstrated antimicrobial property as well</p>	<ul style="list-style-type: none"> • Neem leaf extract (A. indica)

as the ability to remove the smear layer ²⁵	<ul style="list-style-type: none"> • Triphala • M. citrifolia • garlic (A. sativum) • green tea, tea tree oil (M. alternifolia) • Tulsi leaf extract (O. sanctum) • Miswak (S. persica) • passion fruit juice, • T. chebula seed extract • turmeric (C. longa)
Herbal root canal irrigants with pulp tissue dissolution ability – These extracts possess the ability to dissolve the pulp tissue ²⁵	<ul style="list-style-type: none"> • Sapindus mukorossi extract, • Garlic (A. sativum), • Nepenthes khasiana digestive fluid

Table 1. Classification of herbal irrigants based on their property

CONCLUSION

Herbal irrigants have shown potential to be used either as alternatives or as conjunctive agents in root canal disinfection. The advantages coupled with the lack of adverse reaction outweigh the disadvantages.

Herbal medicines performed less effectively in terms of the antibacterial property & smear layer removal than various sodium hypochlorite & EDTA concentrations except few which showed similar efficacy. Continued research into possible combinations for effective disinfection is the need of the hour.

Declaration by Author

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