Review Article



Recent Advancements in Intracanal Medicaments used in Paediatric Dentistry: A Comprehensive Review

Sheenam Ayub, M.D.S., Sonal Gupta, M.D.S., Riya Ojha, M.D.S., Antra Saket, M.D.S.

Department of Pediatric and Preventive Dentistry, KD Dental College and Hospital, Mathura, Uttar Pradesh 281006, India.

Received 23 April 2024 • Revised 19 May 2024 • Accepted 7 June 2024 • Published online 11 December 2024

Abstract:

This review aimed to examine the recent literature concerning conventional intracanal medicaments, and summarize important recent developments. Literature research was performed focusing on articles on safe and effective intracanal medicaments in Google Scholar using the MeSH terms "herbal drugs as intracanal medicaments", "intracanal medicaments for pediatric patients" and "safe intracanal medicaments". A total of 63 studies were included in this review, following inclusion and exclusion criteria. Our literature search revealed that drugs derived from natural sources have become promising alternatives for intracanal medicaments due to their safety profile and potential therapeutic advantages. Furthermore, herbal remedies are frequently well-tolerated by paediatric patients, reducing the likelihood of adverse events. The advantages of herbal drugs in paediatric dentistry extend beyond their antimicrobial efficacy. They offer a holistic approach by harnessing the healing properties of nature, potentially minimizing the need for synthetic chemicals and antibiotics. Recent advancements in intracanal medicaments for paediatric dentistry encompass a spectrum of options, ranging from conventional to herbal-based formulations. While conventional medicaments continue to play a vital role, herbal drugs offer promising alternatives with potential benefits in efficacy, safety, and patient acceptance.

Keywords: herbal drugs as intracanal medicaments, natural products in paediatric dentistry, newer intracanal medicaments

Contact: Dr. Sheenam Ayub, M.D.S.

Department of Pediatric and Preventive Dentistry, Junior Resident, KD Dental College and Hospital, Mathura, Uttar Pradesh 281006, India.

E-mail: dr.sheenamayub@gmail.com

© 2024 JHSMR. Hosted by Prince of Songkla University. All rights reserved.

This is an open access article under the CC BY-NC-ND license

 $\big(http://www.jhsmr.org/index.php/jhsmr/about/editorialPolicies\#openAccessPolicy \big).$

J Health Sci Med Res 2025;43(3):e20241129 doi: 10.31584/jhsmr.20241129 www.jhsmr.org

Introduction

Dental caries is a prevalent concern in the pediatric population, primarily due to dietary habits, inadequate oral hygiene and some other factors such as developmental disorders and drug usage. It begins with the demineralization of the tooth enamel by acids produced by bacteria in dental plaque, which can progress to the point where it affects the dental pulp—the innermost part of the tooth containing nerves, blood vessels, and connective tissue. Depending on the extent of the infection, pulp therapy such as pulpotomy, pulpectomy or root canal therapy (RCT) can be performed to save the tooth, alleviate pain, or treat the infection¹.

The initial foundation for achieving success in endodontic treatments was based on the triad of thorough disinfection, debridement and obturation of the root canal system, with equal importance given to each component. Presently, the success of root canal treatment depends on general principles, irrespective of the specific factors that contribute to its effectiveness.

This encompasses the process of diagnosing and planning treatment, understanding the structure and form of the affected area, adhering to the conventional practices of removing decayed tissue, ensuring thorough disinfection, and filling the root canal, as well as restoring the coronal part of the tooth².

The goal of paediatric endodontics is to maintain the integrity of primary teeth that have suffered pulp tissue damage due to caries, dental trauma, or other causes until they are naturally replaced by permanent teeth. A RCT is necessary when the pulp has become irreversibly infected or necrotic. The procedure is highly dependent on technique, as it can be quite challenging to instrument the complex canals of primary molars that have curved roots and are naturally programmed for resorption. Furthermore, the existence of accessory canals poses challenges in the process of biomechanical preparation,

and achieving complete elimination of necrotic pulp tissue is highly challenging. Due to the challenging nature of achieving thorough mechanical preparation of wide canal walls, it is advised to divide the treatment into two separate appointments, with the application of an antiseptic dressing in-between visits. There is a likelihood of intracanal bacteria multiplying between the appointments when treatment cannot be completed in a single appointment, that is why intracanal medication is recommended for this purpose³.

Walton asserted that intra-canal medicaments have historically been closely associated with endodontics. They are widely recognised as a vital element of the treatment and act as an indispensable factor for the efficacy of RCT. However, the justification for using intra-canal medicaments in modern endodontics is somewhat different. Using an intra-canal medication can be advantageous in minimising the regrowth of bacteria and potentially improving the suppression of bacteria by completely eliminating the bacteria. Interappointment antimicrobial medication functions by impeding the growth of bacteria and eradicating any remaining bacteria, while also reducing the entry of pathogens through a damaged dental restoration⁴. Typically, when treating teeth with a living pulp, intracanal medication is unnecessary. The issue of the function of intracanal medicaments becomes increasingly pertinent and intricate when dealing with cases involving pulp tissue death and inflammation of the tissues surrounding the tooth's root tip.

The main purpose of intra-canal medicaments is to prevent the occurrence of secondary infection and provide a bactericidal effect. It is necessary to attempt to eliminate micro-organisms that survive after obturation; otherwise, they will multiply in the root canals. Chemo-mechanical preparations are ineffective in eliminating the micro-organisms that remain in the dentinal tubules. Improper application of antiseptic intra-canal medicaments

between visits can lead to the proliferation of remaining micro-organisms, resulting in the same number of micro-organisms as in the initial stage of treatment. Thus, it is imperative to use effective intracanal medications between treatments⁵.

Cardinal features of an intracanal medicament⁶

In addition to functioning as a potent disinfectant against germs and fungi, it should also disinfect surfaces. It must not cause irritation to the tissues surrounding the apex of the tooth.

It should maintain stability in the solution. It is expected to exhibit a long-lasting antimicrobial effect.

It must exhibit activity when exposed to blood, serum, and protein derivatives from tissues.

It is desirable for it to exhibit a low surface tension. It should not impede the restoration of periapical tissues.

It should not cause discoloration of the tooth's structure. It should not elicit a cellular immune response.

The objective is to inhibit coronal microleakage and ensure that it does not permeate the temporary restoration.

Rationales of using intra-canal medicaments

Elimination of the pathogens: The primary reason of intra-canal medicament administration is to eliminate the living bacteria and to eradicate all micro-organisms present in the root canal.

To render the contents of the canal inactive: This is achieved by the application of chemicals to preserve or neutralize the tissue or debris that is present in the pulp cavity.

To mitigate post treatment discomfort: Endodontic treatment aims to reduce inflammatory responses, using intracanal medicaments with antibacterial properties to decrease pain and inflammation, though direct application may cause pain in sensory nerves.

Augmenting anaesthesia: Some agents can help reduce pain in inflamed pulp, allowing easy removal between appointments with slight anesthetic issues.

Managing the ongoing periapical infection: Pain and swelling after treatment or draining a canal indicate an active periapical lesion. Intracanal medications are used to restore balance and reduce difficult situations⁷.

Medical and surgical indications of intra-canal medicaments

To desiccate the consistently damp root canal, commonly referred to as weeping canals.

To eradicate any residual micro-organisms in the pulp cavity.

To render the contents of the root canal chemically inactive.

To eliminate tissue debris.

To serve as a protective barrier against leakage from an interappointment dressing in cases with symptoms.

However, concerns over potential drawbacks of intracanal medicaments such as cytotoxicity and antibiotic resistance, have led to the exploration of alternative options. The main objective of this literature review is to search for different treatment outcomes so as to minimize discomfort, especially from the conventional intracanal medicaments, and to explore the plethora of natural products for paediatric patients in dentistry.

Literature research was performed regarding safe and effective intracanal medicaments in Google Scholar and PubMed/MEDLINE articles dated from October 2023 to March 2024 using the MeSH terms "herbal drugs as intracanal medicaments", "intracanal medicaments for pediatric patients" and "safe intracanal medicaments". A total of 63 studies were eventually included in this review following the application of our inclusion and exclusion criteria.

Herbal drugs have been used more or less forever as promising alternatives due to their known high safety profiles and potential therapeutic benefits. Various herbal extracts, including propolis, neem, and aloe vera, have demonstrated antimicrobial properties and the ability to stimulate dentinogenesis and pulp repair. Moreover, herbal medicaments are generally well-tolerated by pediatric patients, reducing the likelihood of adverse reactions.

The advantages of herbal drugs in pediatric dentistry extend beyond their antimicrobial efficacy. They offer a holistic approach by harnessing the healing properties

of nature, minimizing the need for synthetic chemicals and antibiotics. Additionally, herbal medicaments may contribute to patient education and acceptance of natural remedies, aligning with the growing preference for alternative and complementary therapies. Details of various studies regarding the uses of several natural products are enumerated in Table 1.

Table 1 Showing details of natural and newer intracanal medicaments

Herbal drug	Biological source	Application in dentistry	Reference
Curcumin (Turmeric)	Curcuma longa	Intracanal medicament in root canal treatment, periodontal problems, mouthwash, local drug delivery system,	46
		subgingival irrigant, pit and fissure sealant, precancerous	
Propolis	Bee glue	lesion Periodontal disease, mouthwash, irrigant, intracanal	47
	Dee gide	medicament, pulp capping agent, oral ulcers, precancerous	7/
		lesion	
Neem	Azadirachta indica	Toothpaste, mouth wash, intracanal medicament, root canal	48
	7 Zadiraoma maioa	irrigant, periodontal disease, epidermal dysfunction	40
Burdock	Arctium lappa	Intracanal dressing, root canal irrigant, biofilm reduction,	49
	тионат парра	precancerous lesion	.0
Eucalyptus oil	Eucalyptus globulus	Gingivitis, mouthwash, irrigant, intracanal dressing,	50
		toothpaste, lozenges, cold sores, toothpaste, as an	
		anxiolytic, oral candidiasis	
Castor seed oil	Ricinus cvommunis	Storage medium for avulsed tooth, dressing medicament in	51
		pulpectomy & pulpotomy, endodontic sealer	
Guacatonga plant	Casearia cylvestris	Dental plaque treatment, intracanal dressing medicament in	52
		root canal therapy, oral ulcers	
Indian mulberry leaves	Morinda citrifolia	Mouth rinse, intracanal medicament, toothpaste, endodontic	43
		irrigant, disinfecting impressions	
Latex of papaya leaves and	Carica papaya	Gingivitis, toothpaste, mouthwash, intracanal medicament,	53
fruits (Papain)		probiotics, caries removing agent, remineralizing agent	
Tulsi oil	Ocimum sanctum	Obturating material, intracanal medicament, Mouthwash,	54
		toothpaste, gingivitis	
Garlic	Allium sativum	As an irrigant and intracanal medicament, pulpotomy,	55
		mouthwash, periodontal disease	
Lemon solution	Lemon	Intracanal medicament	50
Aloe vera leaf	Aloe vera	Mouthwash, intracanal medicament, root canal irrigant	48
Licorice oil	Glycyrrhiza glabra	Chewing gums, dentifrice, gels, lollipops, intracanal	56
-		medicament, as an irrigant	40
Triphala	Emblica officinalis,	Mouth rinse, intracanal medicament, periodontal problems,	43
(Amalaki, Bibhitaki & Haritaki)		chelating agent, root canal irrigant	
	Terminalia chebula		
Fennel seed	Foeniculum vulgare	Dental plaque and salivary pH reduction, intracanal	57
		medicament, toothpaste, mouthwash	

Commonly used intracanal medicaments in pediatric dentistry

Calcium hydroxide Ca(OH)₂: The introduction of Ca(OH)₂ in dentistry came as, pulp capping material, and it was used in clinical settings for controlling bacteria, decomposing organic residues, managing inflammation, preventing root resorption, restoring tooth structure, and acting as a temporary filling material between appointments. One study reported that the high pH of this substance exhibited antimicrobial properties by deactivating the surface enzymes of microorganisms⁸. Another study reported that *Enterococcus faecalis (E. faecalis)* i.e a gram-positive bacterium which is reported to be a significant pathogen in endodontic infections, was resistant to Ca(OH)₂ and could withstand exposure to Ca(OH)₂ for approximately ten days⁹. Ca(OH)₂ has a minimal impact on facultative anaerobes and Candida species but is effective on obligate anaerobes¹⁰.

Formocresol: Formocresol, a potent antibacterial agent, was initially used by Buckley in 1906. This substance is efficient in combating both aerobic and anaerobic microorganisms within the root canal. It is suitable for emergency endodontic procedures and pain management. However, it exhibits mutagenic and carcinogenic characteristics at elevated levels. In endodontics, a diluted form of Buckley's formocresol formula is used by placing cotton pellets in the tooth's pulp chamber to allow the vapours to spread throughout the entire canal preparation¹¹. The use of this substance as a pulpotomy agent is controversial due to its potential for sensitization, toxicity, mutagenic or carcinogenic effects, and the risk of causing harm to the surrounding oral and para-oral tissues if used carelessly¹².

Glutaraldehyde: The product is offered in a 2% glutaraldehyde concentration. The oil is colourless, has low solubility in water, and shows a slightly acidic reaction. Graven initially suggested using glutaraldehyde as a 2%

concentration intra-canal medicament due to its strong disinfectant and fixative properties¹³. Glutaraldehyde does not trigger an immunologic response, unlike formaldehyde. Glutaraldehyde has bacteriostatic properties, which means it hinders the growth and reproduction of bacteria. The toxicity level is lower than that of formaldehyde. Its high molecular weight prevents it from penetrating periapical tissues, unlike formaldehyde. Direct application of a 2% glutaraldehyde solution to root stumps in a primary tooth is more effective than mixing a drop of the solution with ZOE paste. Its small molecular size prevents it from reaching the apical foramen, causing minimal disturbance to the systemic circulation.

Chlorhexidine gluconate: Chlorhexidine (CHX) gluconate is available in various formulations, such as mouthwash with concentrations of 0.12% and 0.2%, proven to decrease the amount of *Streptococcus mutans* (*S.mutans*) within 24 hours. The product is offered in gel form containing 1% of the active ingredient, as well as in varnishes with concentrations of 1%, 10%, 20%, and 35%. It is minimally toxic and very efficient against *E. faecalis*, a prevalent pathogen in root canal treatments. Two percent CHX gel and a mixture of CHX and Ca(OH)₂ are used in intracanal medicaments. These substances are effective against both aerobic and anaerobic bacteria^{14,15}.

Ledermix paste: Schroeder and Tridon (1960) developed the compound, that is a combination of a glucocorticoid and the antibiotic demeclocycline, a broad-spectrum antibiotic. Triamcinolone acetonide (1%) is the active ingredient present in the Ledermix paste. It is a non-hardening paste material that is soluble in water, utilized as a medication in root canals, or as a pulp capping agent, either applied directly or indirectly. It acts as an anti-inflammatory agent. Ledermix paste releases triamcinolone, a glucocorticoid inside of root canal, and that can reach the bloodstream through diffusion across lateral canals, dentinal tubules and the apical foramen¹⁶. A total of thirty percent

of the release of triamcinolone takes place at the first 24 hours, with the remaining seventy percent released over a 14-week period. It has been also reported that patients treated with Ledermix had superior healing and a more preserved root structure in comparison to a group treated with gutta-percha and sealer¹⁷.

Ozonised oil: Ozone has been increasingly used in dentistry due to its antimicrobial properties and potential benefits in various dental treatments. Ozone can be applied in various forms, like ozonated water, ozonated oil, and ozone gas. It aids in tissue cleaning and encourages the growth of new blood vessels¹⁸. An investigation showed that low concentrations of ozone effectively eradicated Escherichia coli cells, plus the vegetative cells and spores of Bacillus cereus and Bacillus megaterium¹⁹. Siqueira et al. studied the antibacterial effects of ozonized oil and the use of Ca(OH), pastes on different kinds of bacteria commonly found in peri-radicular diseases. Ozone can be utilised as an intracanal medicament by dissolving it in oil for lubrication and disinfection purposes. The success rate for peri-radicular lesions and their antibacterial activity is very high when used as an intracanal medication.

Octenidine: Octenidine (OCT) hydrochloride is an antimicrobial compound classified as a bipyridine. The Sterling-Winthrop Research Institute created it. It acts as an antimicrobial agent in mouthwashes to inhibit the development of dental plaque. It improves the wound healing process. This compound shows potent efficacy against bacteria, fungi, and viruses even at concentrations as low as 0.1% or lower. Topical administration of this substance does not cause any systemic toxicity. Muller et al. conducted a study that determined OCT's antimicrobial effectiveness was not reduced by the presence of organic matter. Makkar et al's research demonstrated that a 0.1% concentration of OCT effectively reduced the colony-forming units of *E. faecalis* following a 3-minute exposure. OCT demonstrated

greater effectiveness than CHX against both planktonic and biofilm microorganisms due to its anti-adhesive characteristic²⁰. OCT hinders bacterial co-aggregation and prevents biofilm formation. OCT lacks tissue solubility. This limits the use of OCT.

Antibiotics: Triple antibiotic paste (TAP) is a combination of three antibiotics. The synergistic action of the three drugs in the paste creates a powerful antimicrobial agent that is effective against microbes. The most widely and effectively used intracanal antibiotic medicament is the combination of Metronidazole, Ciprofloxacin, and Minocycline. Various studies have reported on the efficacy of this combination in eradicating bacteria in the innermost layer of the root canal. Furthermore, it has been found that the administration of this blend of antibiotics on the affected dentinal wall of the root canal reduces the presence of bacteria, with only a few exceptions where they remain undetectable. However, prolonged use of this antibiotic combination in endodontic treatment may cause crown discoloration. This staining property is reported by the use of "minocycline," a tetracycline antibiotic21. TAP is used in baby teeth to disinfect infected root dentin and prevent the growth of resistant microorganisms. The medication penetrates deeply into the dentin, successfully eradicating bacteria. Root canal dressings are effective when used in conjunction with antibiotics. Hence, TAP has broad utility as a medicament within the root canal for pediatric patients. Its intended uses include canals that are consistently damp or secreting fluid, to eliminate tissue debris by breaking down the contents of the root canal, to eliminate excess microorganisms that impede seepage, and to use in cases with many symptoms²².

Nisin: Nisin (bacteriocin) is a naturally occurring anti-microbial peptide derived from the Streptococcus and Lactococcus species, gram-positive bacteria. One of the studies reported about the use of Nisin as an oral

antimicrobial agent, while Howell et al. developed Nisin as an anti-microbial mouth rinse. A study conducted by Tong et al. found that Nisin had the ability to hinder the growth of cariogenic bacteria, such as S. mutans. The biomedical application of this technology has made significant progress, allowing for the prevention of the proliferation of drug-resistant bacterial strains such as methicillinresistant Staphylococcus aureus (S. aureus), Enterococci, Streptococcus pneumoniae, and Clostridium difficile. The antimicrobial activity specifically targets Gram-negative pathogens and a wide range of Gram-positive bacteria, including their spores. It is also effective against drugresistant E. faecalis isolates. The potent and extensive efficacy of this treatment minimises the likelihood of bacterial resistance and reduces cellular toxicity at normal anti-microbial concentrations. Nisin also exhibits antibiofilm properties. The use of Nisin in dentistry has been limited, however, it has been found to be highly effective in eliminating E. faecalis from the root canal system²³.

Anti-microbial photodynamic therapy (aPTD):

A technique known as photosensitization has been used in medicine for a considerable time. It is also called Photoactivated Disinfection (PAD). Due to its antimicrobial efficacy, it is used in endodontic procedures. PAD is composed of abundant amounts of tolonium chloride, also referred to as toluidine blue and a photoactive agent. A red laser at 635 nm is used to stimulate the photosensitizer molecule. Through a small flexible optical fiber, light is delivered to the root canal with a maximum power of 100 mW, while maintaining the safety of adjacent tissue. In addition to disrupting membranes, deactivating enzymes, and damaging genomic and plasmid DNA, the potent oxidizer also causes immediate death in microbial cells. The use of light activated therapy in dentistry could help combat human infections caused by biofilms²⁴.

Endox: Endox endodontic systems sterilize root canals by ejecting high-frequency electrical impulses. In order to achieve sterilization through fulguration, Endox endodontic systems claim to eliminate both pulp and bacteria from the entire root canal. The unit was demonstrated to be incapable of eliminating pulp tissue from root canals without mechanical cleaning, according to a recent study. In spite of the fact that the unit can serve as a substitute for conventional cleaning and shaping methods, the authors do not recommend the use of high-frequency electric pulses exclusively as an endodontic treatment²⁵.

Natural products

Curcumin: Curcumin is a natural compound obtained from the roots of Curcuma longa (Turmeric). It has been recognised for its pleiotropic effects, such as antimicrobial, anti-inflammatory, antioxidant, and anti-cancer. These properties make it relevant in clinical dentistry for the treatment and prevention of various diseases. Concerning its endodontic intracanal medications, in an in vitro study, Curcumin was incorporated into polymeric fibers to evaluate its antimicrobial properties and potential application in root canal disinfection, and the author concluded that photoactivation of curcumin-based medicaments seems to be essential to obtain greater antibiofilm activity²⁶. Recent studies have emphasized curcumin's potential as a viable alternative to TAP for infection control. One study reported that a minimum concentration of 2.5 mg/mL was required to produce a therapeutic response²⁷.

Propolis: Propolis is one of the most superior intracanal medicaments due to its anti-inflammatory and potent antimicrobial properties. Its effectiveness is comparable to NaCl when compared to saline. It can be utilized as a short-term treatment for pulp and periapical pathologies. Propolis contains active components such as flavonoids,

phenolics, and aromatics, contributing to its diverse biological activities like antimicrobial, anti-inflammatory, antioxidant, anaesthetic, and cytotoxic effects. Propolis demonstrates notable antibacterial effectiveness and can serve as an intracanal medicament²⁸.

Azadirachta indica: Commonly referred to as Neem, possesses exceptional antimicrobial properties and the oil of neem seeds has been extensively used in dentistry. The key active chemical constituents responsible for the broad spectrum of antimicrobial effects are azadirachtin, nimbolinin, nimbidin, and nimbin. In dentistry neem leaves and their oil has been evaluated for its antibacterial and antifungal properties and it is quite useful in dentistry for curing gingival problems and maintaining oral health in a natural way. Neem-based pastes and oil have also been studied and thoroughly tested for their intracanal medicament and irrigant purposes removal of oral aerobic and anaerobic pathogens existing in the oral cavity. Nimbidin is the primary extract obtained from the seed kernels of neem. This active ingredient exhibits antibacterial properties. In a recent study, neem oil was reported to have potential antimicrobial effects against S. mutans and E. faecalis, while it was comparably effective against Candida species. It can be used as a substitute for Ca(OH), an intracanal medicament obtained from a chemical source²⁹.

Arctium lappa: It is a plant native to Japan that is found all over the world. It has been extensively utilized as a medicine due to its antibacterial, antioxidant, and antifungal properties. The presence of polyacetylenes in this plant is responsible for their antibacterial activity. *Arctium lappa* has shown significant antimicrobial properties against the various root canal microbes³⁰.

Eucalyptus oil: Eucalyptus oil is an essential oil obtained from the leaves of the Eucalyptus tree. It possesses anti-inflammatory and antimicrobial properties. Its broad antimicrobial spectrum, synergistic effects and

biocompatibility suggest its effectiveness as an intracanal medicament³¹.

Castor oil: Castor oil is obtained from the seeds of *Ricinus communis*. The plant is rich in ricinoleic acid. *Ricinus communis* can be utilised as a root canal irrigant and intracanal medicament due to its composition of alkaloids (oxindole), glycosides, triterpenes, tannin, phenolic compounds, steroids, and flavonoids, which contribute to its antimicrobial properties. One study reported that *Ricinus communis* paste, along with Ca(OH)₂, demonstrated superior antimicrobial efficacy compared to a paste containing Ca(OH)₂ and propylene glycol³². Additionally, another study found that *Ricinus communis* exhibited complete eradication against *Candida albicans* and significant antimicrobial activity against *E. faecalis*³³.

Casearia sylvestris: Casearia sylvestris is a medicinal plant native to tropical America and Brazil that exhibits antiseptic and antimicrobial properties. It can serve as a viable option for short-term intracanal medication in cases of pulp and periapical inflammation. Also known as 'guacatonga,' this plant is recognised for its antimicrobial, diuretic, anti-ulcer, and healing attributes. It includes a substantial quantity of phospholipase A2 inhibitors. Silva et.al. suggest that Casearia sylvestris can be used as a substitute for short-term intracanal medication³⁴.

Morinda citrifolia: It is commonly known as Indian mulberry. It has a large variety of applications due to its analgesic, anti-inflammatory, antioxidant, antibacterial, and antiviral properties. *Morinda citrifolia* shows a significant reduction in the zone produced by the microorganism *E. faecalis*, this property making it suitable for use as an effective intra-canal medicament³⁵.

Papain: Papain is an enzyme and a natural product derived from the latex of papaya leaves and fruits (*Carica papaya*). Papain is a cysteine protease enzyme that exhibits notable antibacterial and anti-inflammatory characteristics

due to its proteolytic properties. Various studies suggest that it can be utilised as an intra-canal medicament. It also possesses a debris-removing effect. Its ability to dissolve tissue makes it useful as a cariostatic agent. Anuj Bhardwaj et al.'s study found that papain treatment was equieffective in comparison with CHX in terms of antimicrobial effectiveness. By virtue of this, it could be a substitute for CHX³⁶.

Tulsi: Tulsi is obtained from the leaves of the plant Tulsi (Ocimum sanctum), a member of the Labiateae family. Tulsi is regarded as sacred plant in subcontinent such as India, and known as sacred basil. It is renowned for its antimicrobial, antifungal, and antiviral characteristics. The essential oil extract of Ocimum sanctum demonstrates strong concentration-dependent antibacterial properties. Using Ocimum sanctum as a medicament inside the tooth canal in paediatric dentistry provides a hopeful natural option. The antimicrobial, anti-inflammatory, and analgesic qualities of this substance make it highly suitable for treating infected root canals in primary teeth. Furthermore, its organic source and limited negative impacts make it well-suited for paediatric patients. Additional research is required to determine the ideal concentration and long-term effectiveness of Tulsi in paediatric endodontic procedures, but it shows promise as a valuable addition to paediatric dental care³⁷.

Garlic: Garlic is obtained from the cloves of *Allium* sativum, and possesses a diverse range of antimicrobial responses, exhibiting both bacteriostatic and bactericidal effects depending on the formulation and concentrations. Its antimicrobial effectiveness stems from its ability to prevent toxin production and the activation of enzymes involved in causing endodontic diseases. It may serve as a substitute intracanal medicament. It has pharmacological properties including antimicrobial, antiplatelet, antithrombotic, and anticancer effects. It has been proven effective against

S. mutans, S. aureus, E. faecalis, and $Escherichia\ coli.$ Research by Eswara K et al. found that garlic had superior antimicrobial effectiveness in comparison to $Ca(OH)_2^{38}$. Another study by Salih et al. found that garlic exhibited greater efficacy against strains of S. aureus in comparison to E. faecalis³⁹.

Lemon solution: Is obtained from the plant *Citrus limon*, from the family Rutaceae. The lemon solution is a natural source of citric acid with a mild level of acidity. One study found that a fresh lemon solution had broad antibacterial properties, even against *E. faecalis*, making it suitable for use as an intracanal medicament⁴⁰.

Aloe vera: Aloe vera is a member of the family Liliaceae. Aloe vera gel is obtained from the mucilaginous tissue found in the center of the Aloe vera leaves. Anthraquinone glycosides found in total leaf extracts of Aloe vera possess potent antibacterial properties. Aloe vera has been studied for its antibacterial effectiveness and examined as a medicament for use inside root canals⁴¹. The leaf extracts contain anthraquinones, which are responsible for their antibacterial properties. The substance also includes various active components such as vitamins, minerals, enzymes, sugars, lignin, saponins, salicylic acids, and amino acids. Alloin and emodin function as analgesics, antibacterials, and antivirals. The effectiveness against Streptococcus pyogenes and E. faecalis is attributed to the presence of anthrax quinine. In 2016, Kurian et al. conducted a study demonstrating that the minimal inhibitory concentration of Aloe vera was more effective than Ca(OH) in eradicating E. faecalis. The antibacterial properties of Aloe vera were found to enhance over time⁴². Aloe vera exhibits wide-ranging antibacterial effects against different oral pathogens when compared to other natural extracts.

Glycyrrhiza glabra: Also known as liquorice, it is recognized for its anti-inflammatory, anticarcinogenic, and antiviral properties. Glycyrrhizin is a chemical constituent,

a triterpenoid compound, that is effective against various strains of *S. mutans*. It exhibits higher bio-compatibility than CHX as an intracanal medicament⁴³.

Triphala: (Haritaki, Bibhitaki, and Amalaki) Triphala, a renowned Ayurvedic blend, has been demonstrated to successfully prevent the formation of biofilm. It is very suitable for use as an intra-canal medicament. It possesses potential antibacterial and anti-inflammatory characteristics. The fruit contains a high concentration of citric acid, which can help remove the smear layer by acting as a chelating agent. A study by Prabhakar et al. found that Triphala displayed antibacterial effectiveness nearly equivalent to Ca(OH)₂ and could serve as a substitute for Ca(OH)₂ in combating *E. faecalis*⁴⁴.

Foeniculum vulgare: This belongs to the Apiaceae family, and its fennel seeds are known for their analgesic, anti-inflammatory, antispasmodic, antioxidant, diuretic, carminative, anti-microbial, and anti-cancer properties. Analysis conducted in a controlled environment indicated its potential application as a medicament within a root canal⁴⁵.

In this study, some of the naturally obtained drugs which have been reported to have utility as intracanal medications and in other endodontic treatments are Curcumin, Propolis, *Aloe Vera*, Neem extracts, and Tulsi oil. Their broad mechanisms and pharmacological actions are summarized in Table 2. Recent advancements in intracanal medicament are adumbrated in Figure 1.

Table 2 Pharmacological properties of natural intracanal medicaments⁵⁸

Pharmacological activity	Biological source	
Analgesics and anti-inflammatory action	Curcuma longa, Arctium lappa, Propolis, Azadirachta indica, Casearia sylvestris, Allium sativum, Citrus limonum, Carica papaya, Morinda citrifolia, and Glycyrrhiza glabra, Triphala,	
Aught underschied auchteur	Ocimum sanctum, Eucalyptus oil	
Anti-microbial action	C. sylvestris, M. citrifolia, Propolis, and Glycyrrhiza glabra, Curcumin, Propolis, Aloe vera,	
Antioxidants	Ocimum sanctum, Eucalyptus oil Curcumin, Grape seed extracts, Quercetin, Morinda citrifolia	
Anti-anxiety	Arctium lappa	
Anti-fungal	Ocimum sanctum, Azadirachta indica	
Anti-viral	Glycyrrhiza glabra, Aloe vera, Ocimum sanctum, Morinda citrifolia	

C. sylvestris=casearia sylvestris

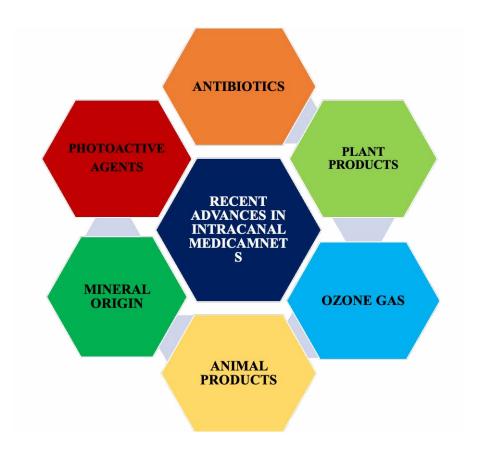


Figure 1 Various sources of intracanal medicaments

Conclusion

Intracanal medicaments play a crucial role in pediatric dentistry by aiding in the disinfection and healing of root canals in primary teeth. Through their antimicrobial properties, they effectively combat bacterial infections, reduce inflammation, and promote the preservation of primary dentition. However, it's essential to recognize the unique considerations of pediatric patients, such as their smaller tooth size and the potential systemic effects of medicaments, necessitating careful selection and administration under professional supervision. Despite the challenges, the judicious use of intracanal medicaments remains a valuable tool in ensuring the long-term oral health and well-being of young dental patients. Continued research and advancements in this area hold promise for further enhancing treatment outcomes and minimizing

potential risks, thereby contributing to the overall success of pediatric dental care. Further research and clinical trials are warranted to elucidate the full therapeutic potential of herbal intracanal medicaments and their integration into pediatric endodontic practice.

Authors' contributions

Authors SA and SG designed and performed the study, RO and AS contributed to proofing and the literature search. All authors also contributed to manuscript writing.

Ethical considerations

Since this was a literature review, no ethical clearance was required. However, we followed all ethical guidelines and practices for the study.

Acknowledgement

We acknowledge the contribution of the supporting staff of the Department of Pediatric & Preventive Dentistry KD Dental College and Hospital, India.

Conflict of interest

There are no conflicts of interest to report.

References

- Shah A, Peacock R, Eliyas S. Pulp therapy and root canal treatment techniques in immature permanent teeth: an update. Br Dent J 2022;232:524–30.
- Byström A, Sundqvist G. Bacteriologic evaluation of the efficacy of mechanical root canal instrumentation in endodontic therapy. Scand J Dent Res 1981;89:321–8.
- Paquette L, Legner M, Fillery ED, Friedman S. Antibacterial efficacy of chlorhexidine gluconate intracanal medication in vivo. J Endod 2007;33:788–95.
- Neelakantan P, Sanjeev K, Subbarao CV. Duration-dependent susceptibility of endodontic pathogens to calcium hydroxide and chlorhexidine gel used as intracanal medicament: an in vitro evaluation. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2007;104:e138-41.
- Almyroudi A, Mackenzie D, McHugh S, Saunders WP.
 The effectiveness of various disinfectants used as endodontic intracanal medications: an in vitro study. J Endod 2002;28:163-7.
- Patro PP. Intracanal medicaments in paediatric endodontics.
 Indian J Forensic Med Toxicol 2020;14:9106–10.
- Walton RE. Intracanal medicaments. Dent Clin North Am 1984:28:783–96.
- Hamidi MR, Mahmoudi E, Moghadamnia AA, Zahedpasha S.
 Effect of calcium hydroxide and chlorhexidine medicaments on the apical seal. Iran Endod J 2012;7:15.
- Kumar H. An in vitro evaluation of the antimicrobial efficacy of Curcuma longa, Tachyspermum ammi, chlorhexidine gluconate, and calcium hydroxide on Enterococcus faecalis. J Conserv Dent 2013;16:144-7.
- Sinha N, Patil S, Dodwad PK, Patil AC, Singh B. Evaluation of antimicrobial efficacy of calcium hydroxide paste, chlorhexidine

- gel, and a combination of both as intracanal medicament: an in vivo comparative study. J Conserv Dent 2013;16:65-70.
- 11. Ellerbruch ES, Murphy RA. Antimicrobial activity of root canal medicament vapors. Journal of Endodontics. 1977;3:189–93.
- Casas MJ, Kenny DJ, Judd PL, Johnston DH. Do we still need formocresol in pediatric dentistry? J Can Dent Assoc 2005;71:749–51.
- Faysal MI, Kany, El Zawahry Mohammed R. Study of the antimicrobial efficiency of some root canal medicaments. Int J Microbiol Res 1987;2:9–17.
- Lee SY, Nam EJ. Clinical efficacy of 1% CHX gluconate gel and 0.12% CHX solution: a randomized controlled trial. Int J Environ Res Public Health 2022;19:9358.
- 15. de Almeida Gomes BP, Vianna ME, Sena NT, Zaia AA, Ferraz CC, de Souza Filho FJ. In vitro evaluation of the antimicrobial activity of calcium hydroxide combined with chlorhexidine gel used as intracanal medicament. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2006;102:544–50.
- Klotz MD, Gerstein H, Bahn AN. Bacteremia after topical use of prednisolone in infected pulps. J Am Dent Assoc 1965;71:871-5.
- Schroeder A. Ledermix 1962--Ledermix today. Evaluation after
 years of experience. Zahnarztl Prax 1975;26:195-6.
- Seidler V, Linetskiy I, Hubálková H, Stanková H, Smucler R, Mazánek J. Ozone and its usage in general medicine and dentistry: a review article. Prague Med Rep 2008;109:5–13.
- 19. Broadwater WT, Hoehn RC, King PH. Sensitivity of three selected bacterial species to ozone. Appl Microbiol 1973;26:391–3.
- Makkar S, Aggarwal A, Pasricha S, Kapur I. Comparative evaluation of octenidine hydrochloride and chlorhexidine as antibacterial root canal irrigant. Indian J Oral Sci 2015;6:10-3.
- Malu K, Khubchandani M. Triple antibiotic paste: a suitable medicament for intracanal disinfection. Cureus 2022;14: e29186.
- Parhizkar A, Nojehdehian H, Asgary S. Triple antibiotic paste: momentous roles and applications in endodontics: a review. Restor Dent Endod 2018;43:1–16.
- Shin JM, Gwak JW, Kamarajan P, Fenno JC, Rickard AH, Kapila YL. Biomedical applications of nisin. J Appl Microbiol 2016;120:1449–65.
- 24. Souza LC, Brito PR, de Oliveira JC, Alves FR, Moreira EJ, Sampaio-Filho HR, et al. Photodynamic therapy with two different photosensitizers as a supplement to instrumentation/irrigation procedures in promoting intracanal reduction of Enterococcus faecalis. J Endod 2010;36:292-6.

- Lendini M, Alemanno E, Migliaretti G, Berutti E. The effect of high frequency electrical pulses on organic tissue in root canals. Int Endod J 2005;38:531–8.
- 26. El-Saadony MT, Yang T, Korma SA, Sitohy M, El-Mageed A, Taia A, et al. Impacts of turmeric and its principal bioactive curcumin on human health: pharmaceutical, medicinal, and food applications: a comprehensive review. Front Nutr 2023;9:1040259.
- 27. Devaraj S, Jagannathan N, Neelakantan P. Antibiofilm efficacy of photoactivated curcumin, triple and double antibiotic paste, 2% chlorhexidine and calcium hydroxide against Enterococcus fecalis in vitro. Sci Rep 2016;6:24797.
- 28. Hossain R, Quispe C, Khan RA, Saikat AS, Ray P, Ongalbek D, et al. Propolis: An update on its chemistry and pharmacological applications. Chin Med 2022;17:1–60.
- Bohora A, Hegde V, Kokate S. Comparison of the antibacterial efficiency of neem leaf extract and 2% sodium hypochlorite against E. faecalis, C. albicans and mixed culture – an in vitro study. Endodontology 2010;22:10–4.
- Wang D, Bădărau AS, Swamy MK, Shaw S, Maggi F, Da Silva LE, et al. Arctium species secondary metabolites chemodiversity and bioactivities. Front Plant Sci 2019;10:834.
- Bachir RG, Benali M. Antibacterial activity of the essential oils from the leaves of Eucalyptus globulus against Escherichia coli and Staphylococcus aureus. Asian Pac J Trop Biomed 2012;2:739–42.
- 32. Garcia LD, De Almeida GL, da CP Pires-de-Souza F, Consani S. Antimicrobial activity of a calcium hydroxide and Ricinus communis oil paste against microorganisms commonly found in endodontic infections. Rev Odonto Ciência 2009;24:406-9.
- 33. Valera MC, Maekawa LE, Oliveira LD, Jorge AO, Shygei É, Carvalho CA. In vitro antimicrobial activity of auxiliary chemical substances and natural extracts on Candida albicans and Enterococcus faecalis in root canals. J Appl Oral Sci 2013;21:118–23.
- 34. Silva FB, Almeida JM, Sousa SM. Natural medicaments in endodontics: a comparative study of the anti-inflammatory action. Braz Oral Res 2004;18:174-9.
- 35. Nirmala S, Surender LR, Reddy N, Reddy SD, Chukka RR, Kumar N. Antimicrobial efficacy of morinda citrifolia, Nisin, and 2% chlorhexidine against enterococcus faecalis: an in-vitro

- study. Cureus 2022;14:e23206.
- 36. Bhardwaj A, Ballal S, Velmurugan N. Comparative evaluation of the antimicrobial activity of natural extracts of Morinda citrifolia, papain and aloe vera (all in gel formulation), 2% chlorhexidine gel and calcium hydroxide, against Enterococcus faecalis: an in vitro study. J Conserv Dent Endod 2012;15:293–7.
- 37. Ahirwar P, Shashikiran ND, Sundarraj RK, Singhla S, Thakur RA, Maran S. A clinical trial comparing antimicrobial efficacy of essential oil of Ocimum sanctum with triple antibiotic paste as an intracanal medicament in primary molars. J Indian Soc Pedod Prev Dent 2018;36:191–7.
- 38. Eswar K, Venkateshbabu N, Rajeswari K, Kandaswamy D. Dentinal tubule disinfection with 2% chlorhexidine, garlic extract, and calcium hydroxide against Enterococcus faecalis by using real-time polymerase chain reaction: In vitro study. J Conserv Dent Endod 2013;16:194–8.
- Salih JM, Monawer AT, Abdullkahar IM. Antibacterial activity of garlic against multi-drug resistant Staphylococcus aureus and Enterococcus faecalis in Duhok city. J Univ Duhok 2016;19:114–22.
- 40. Varshini R, Subha A, Prabhakar V, Mathini P, Narayanan S, Minu K. Antimicrobial efficacy of Aloe vera, lemon, Ricinus communis, and calcium hydroxide as intracanal medicament against Enterococcus faecalis: a confocal microscopic study. J Pharm Bioallied Sci 2019;11(Suppl 2):S256.
- Samarh SN, Khalaf NA, Hajhamad MM. Evidence based medical use of aloe vera extracts: short review of literature. Int J Res Med Sci 2017;5:4198–4202.
- 42. Kurian B, Swapna DV, Nadig RR, Ranjini MA, Rashmi K, Bolar SR. Efficacy of calcium hydroxide, mushroom, and Aloe vera as an intracanal medicament against Enterococcus faecalis: An in-vitro study. Endodontology 2016;28:137-42.
- 43. Wahab S, Annadurai S, Abullais SS, Das G, Ahmad W, Ahmad MF, et al. Glycyrrhiza glabra (Licorice): a comprehensive review on its phytochemistry, biological activities, clinical evidence and toxicology. Plants 2021;10:2751.
- 44. Prabhakar J, Senthilkumar M, Priya MS, Mahalakshmi K, Sehgal PK, Sukumaran VG. Evaluation of antimicrobial efficacy of herbal alternatives (Triphala and green tea polyphenols), MTAD, and 5% sodium hypochlorite against Enterococcus faecalis biofilm formed on tooth substrate: an in vitro study. J Endod 2010;36:83-6.
- 45. Hemmanur S, Nasim F. Evaluation of antimicrobial efficacy of

- foeniculum vulgare essential oil: an in vitro study. Plant Cell Biotechnol Mol Biol 2020;21:83-9.
- 46. Nagpal M, Sood S. Role of curcumin in systemic and oral health: an overview. J Nat Sci Biol Med 2013;4:3.
- 47. Jain S, Rai R, Sharma V, Batra M. Propolis in oral health: a natural remedy. World J Pharm Sci 2014;2:90-4.
- 48. Salman BN, Vahabi S, Rad MM. Use of herbs and medicinal plants in dentistry: a review. J Dent Sch 2017;35:133-49.
- 49. Gentil M, Pereira JV, Sousa YT, Pietro R, Neto MD, Vansan LP, et al. In vitro evaluation of the antibacterial activity of Arctium lappa as a phytotherapeutic agent used in intracanal dressings. Phytother Res 2006;20:184–6.
- Dagli N, Dagli R, Mahmoud RS, Baroudi K. Essential oils, their therapeutic properties, and implication in dentistry: a review.
 J Int Soc 2015;5:335-40.
- Nabavizadeh M, Abbaszadegan A, Khodabakhsi A, Ahzan S, Mehrabani D. Efficiency of castor oil as a storage medium for avulsed teeth in maintaining the viability of periodontal ligament cells. J Dent 2018;19:28.
- 52. Ribeiro SM, Fratucelli ÉD, Bueno PC, de Castro MK, Francisco AA, Cavalheiro AJ, et al. Antimicrobial and antibiofilm activities of Casearia sylvestris extracts from distinct Brazilian biomes

- against Streptococcus mutans and Candida albicans. BMC Complement Altern Med 2019;19:1-6.
- Bussadori SK, Castro LC, Galvão AC. Papain gel: a new chemo-mechanical caries removal agent. J Clin Pediatr Dent 2005;30:115.
- 54. Agarwal S, Gupta S, Tandon S, Mathur R, Rai TS, Kumar M, et al. Comparative evaluation of Ocimum sanctum and calcium hydroxide mix as an obturating material in deciduous molars: an in vivo study. Int J Clin Pediatr Dent 2020;13:617.
- 55. Shooriabi M. Effects of Allium sativum (Garlic) and Its derivatives on oral diseases: a narrative review. J Res Dent Maxillofac Sci 2021;6:36–44.
- Tharakan AP, Pawar M, Kale S. Effectiveness of licorice in preventing dental caries in children: a systematic review.
 J Indian Soc Pedod Prev Dent 2020;38:325–31.
- 57. Shirahatti RV, Ankola AV, Nagesh L. Effect of fennel seeds on dental plaque and salivary pH-a clinical study. J Oral Health Community Dent 2010;4:38-41.
- 58. Kishan KV, Shah NC, Das DT, Parikh M. Herbal medicaments in endodontics-current guidelines for in vivo studies in India. Restor Dent Endod 2019;22:411-4.