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Review Article

APICAL PERIODONTITIS AND EFFECTIVENESS OF ROOT CANAL IRRIGATION. REVIEW**Dr.Karam Salah Eldaour, Dr. Mana Hamad Almunajjim, Dr. Abdullah Bader Alzaid,
Dr. Sami Mohaimeed Alharbi, Dr. Abdulrahman Saleh Aldhuhayan,
Dr.Meshal Hatim alogaly****Article Received:** May 2020**Accepted:** June2020**Published:** July 2020**Abstract:**

In this review we will focus on irrigational solutions and their effectiveness in the elimination of pathogens. We performed a search using electronic databases; MEDLINE, science-direct, and EMBASE, through October, 2019. Search strategies used following MeSH terms in searching: "Apical periodontitis", "root canal irrigation", "root canal". The performance of irrigation in the facility root canal system at the apical root canal stays one of the important restorative needs to deal with AP. Experimental approaches and scientific information revealed that irrigation is one of the most effective way for removal of the root canal germs (planktonic and biofilms). The qualities of irrigation procedure, such as the mixture flow dynamics as well as irrigation substitute, have been lately stood out as the essential elements in successful treatment outcome.

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INTRODUCTION:

Apical periodontitis (AP) is an inflammatory reaction round the origin of tooth to an insult. Generally, AP is triggered by bacteria in root canal area, which happens after irreversible inflammatory reaction of the pulp as well as subsequent pulp death [1]. Medically, AP can emerge in numerous methods, from asymptomatic with simply a periapical radiolucency on an intraoral radiograph to great clinical signs of inflammation such as pain, swelling, soreness, and loss of function. In case of serious pain, the need for management is apparent and also compulsory. Nonetheless, frequently, AP is asymptomatic, and then it can continue to be unnoticed for years unless for some other factor a radiograph is taken of the impacted tooth. However asymptomatic, these lesions (as well as additionally the symptomatic ones) might put stress on the overall health of the influenced people.

The purpose of root canal treatment (RCT) is to prevent or manage AP, that is the sequence of microbial emigration in the root canal system [2]. Chemo-mechanical preparation is considered as one of the most vital procedure of RCT intending to clear and shape the total root canal system, more specifically to obviate of bacteria and pathologic particles from this complex tooth location. A selection of tools as well as methods in mix with decontaminating watering options as well as intracanal medicines have been advised for the chemo-mechanical preparation of contaminated root canals. The cleaning as well as shaping effectiveness of root canal instruments, objective to achieve a well-tapered root canal form, enough for the needed irrigant flow in the whole canal and ideal 3D obturation [2].

Apical periodontitis is an inflammatory disorder of periradicular tissues caused by aetiological agents of endodontic origin. Cleanliness process entails mechanical prep work of root canal, chemical irrigation, as well as if required, intracanal medicines. In this review we will focus on irrigational solutions and their effectiveness in the elimination of pathogens.

METHODOLOGY:

We performed a search using electronic databases; MEDLINE, science-direct, and EMBASE, through October, 2019. Search strategies used following MeSH terms in searching: "Apical periodontitis", "root canal irrigation", "root canal". Then we reviewed the references in our included studies for more relevant articles. Moreover, we restricted our search to only English language studies published with human subjects.

DISCUSSION:**• Apical Symptomatic Diagnoses**

Healthy Apical Structures are not vulnerable to percussion or palpation examining as well as radiographically, the lamina dura bordering the root is undamaged and the gum ligament room is consistent. Just like pulp testing, relative assessments for percussion as well as palpation ought to constantly start with normal teeth as a standard for the individual [3].

Symptomatic AP represents inflammation, generally of the apical periodontium, generating clinical signs entailing an excruciating action to biting and/or percussion or palpation. This may or might not be tagged along by radiographic adjustments (i.e. depending upon the stage of the disease, there might be normal width of the periodontal ligament or there may be a periapical radiolucency) [4]. Severe pain to percussion and/or palpation is extremely characteristic of a deteriorating pulp as well as root canal therapy is needed.

Asymptomatic AP is inflammation and also devastation of the apical periodontium which is of pulpal beginning. It looks like an apical radiolucency as well as does absent professional signs (no pain on percussion or palpation) [5].

Persistent Apical Abscess is an inflammatory reaction to pulpal infectivity and necrosis identified by progressive beginning, little or no discomfort as well as a periodic discharge of pus through a connected sinus tract. Radiographically, there are generally indicators of bony destruction such as a radiolucency. To recognize the source of a straining sinus tract when existing, a guttapercha cone is thoroughly placed with the stoma or opening up until it stops and also a radiograph is utilized [6].

Acute Apical Abscess is an inflammatory reaction to pulpal infection and death identified by rapid onset, spontaneous pain, excessive tenderness of the teeth to stress, pus development and also swelling of connected cells. There may be no radiographic marks of damage and also the individual frequently experiences despair, fever and lymphadenopathy [7]. Condensing Osteitis is a diffuse radiopaque lesion standing for a localized bony response to a low-grade inflammatory stimulus generally seen up the tooth [8].

The two most accepted classification plans are presented in Table 1. These are rather similar, however symptomatic teeth according to the AAE classification might consist of extra cases than teeth with severe AP according to the ICD.

Table 1. Classification of apical periodontitis [9].

AAE	ICD-10
Symptomatic apical periodontitis SAP ¹	K04.4 Acute apical periodontitis of pulpal origin ²
Asymptomatic apical periodontitis AAP	K04.5 Chronic apical periodontitis
Chronic apical abscess	K04.6 Periapical abscess with sinus ³
Acute apical abscess	K04.7 Periapical abscess without sinus
Condensing osteitis ⁴	K04.8 Radicular cyst
Radicular cyst	

¹presents with a broad range of symptoms

²presents with strong pain

³further subdivided in relation to sinus tract location on surfaces

⁴may be seen as a variant of AAP or Chronic apical periodontitis

- **Examination and diagnostic procedures for periapical diseases**

It is vital to collect all of the pertinent info concerning the individual, the oral condition and

also the entailed tooth before making a medical diagnosis. Provisional diagnoses of the status of both the pulp and also periapical tissues must be identified from the patient's description of the symptoms, the history of the problem and also details of any current therapy that has been executed (see Table 2). The findings from the scientific as well as radiographic assessments together with the results of the diagnostic tests are then looked at as well as evaluated to develop a clear-cut diagnosis of the problems of the tissues and also to find out which tooth is involved. It is not always required to do every diagnostic test as well as clinicians should only pick examinations that pertain to the presenting complaint- as an example, a pulp sensibility examination applying warmth is probably not going to help a medical diagnosis unless the individual suffers level of sensitivity to warm [10]. Ideally, there must be at least 2 different indicators and/or symptoms existing to show and verify the illness. If there is any type of question as well as the pain is not serious, then the operator should postpone treatment till the medical diagnosis becomes clear, or otherwise the individual must be referred to a professional for medical diagnosis and therapy.

Table 2. Summary of the examination and diagnostic processes for the assessment of the status of the pulp and periapical tissues [10].

Procedure	Result
1. History and discussion with patient Medical history Dental history Description of presenting complaint Details of any previous treatment of presenting complaint	Provisional diagnosis of presenting condition
2. Clinical Examination Extra-oral signs Intra-oral signs Individual tooth assessment Restoration assessment	Assess possible causative factors Provisional diagnosis of tooth status
3. Clinical Tests Pulp sensibility tests Percussion, mobility, palpation	Provisional diagnosis of the status of the pulp and/or the root canal system Provisional diagnosis of the periapical status
4. Radiographic Examination	Confirm/assess causative factors Provisional diagnosis of periapical status
5. Correlation of the history, clinical, radiographic and test findings	DEFINITIVE DIAGNOSIS - Pulp, root canal and periapical status - Cause(s) of the diseases
6. TREATMENT PLAN Investigation/restoration removal Reassessment of the tooth and its prognosis	Confirm the definitive diagnosis and cause(s) Finalize and continue the treatment plan

- **Root canal instrumentation and bacterial reduction**

Because of its complicated anatomy, with the multiple fins, isthmuses, ramifications as well as additional canals, it is virtually impossible for mechanised root canal instrumentation to form and also cleanse the whole root canal system [11-12]. Intraradicular microbes may be nested in these locations, which are inaccessible to instrumentation [14]. On top of that, this intricate atmosphere stops

irrigates from exerting their complete antimicrobial prospective [13].

Classic Scandinavian researches by Byström and also Sundqvist clearly indicated that mechanical instrumentation has the ability to reduce substantially the variety of microbes in the root canal system [15],[16]. These studies showed that hand equipment using stainless-steel tiers and clean and sterile saline led to a 100- 1000- fold reduction of intraradicular microbes, however it was not possible

to predictably acquire bacteria-free root canals [15]. These outcomes were confirmed by further investigations, which revealed that the sole use of hands-on stainless-steel tools does not make root canals bacteria-free [17],[18]. These examinations well demonstrated the minimal antibacterial impact of mechanical prep work.

The introduction of more versatile alloys like NiTi led to the development of rotating systems. It was presumed that these enhancements would lead to extra effective elimination of microorganisms from infected root canals compared to conventional hand instruments. Lately, several researches have been conducted to contrast the level of intraradicular microbial reduction of hand instruments (stainless-steel or NiTi) compared to rotating NiTi tools with greater tapers than ISO.02 taper. Nearly all of these researches stopped working to show considerable distinctions in between hand and rotary instrumentation [16-18].

In recap, predictable and complete microbial removal does not seem feasible, either with standard hand instrumentation or with newer rotating NiTi-systems [14],[17]. Irrigation of the root canal is vital for efficient elimination of germs.

• Anti-bacterial irrigating solutions

Irrigation fulfils several vital chemical and microbiological features. According to Haapasalo et al irrigation is the only method to get rid of cells residues and germs in planktonic and also biofilm types, from the complex areas of the root canal walls that are not touched by mechanical instrumentation [19]. Taking into consideration with each other these data with the shaping efficiency of various systems/instruments in the today endodontic technique, we can securely approve previous microbiological observations that mechanical preparation cannot accomplish foreseeable disinfection [20]. The elimination of microorganisms after instrumentation of the root canal continues to present as a challenging concern in regards to the commonly acknowledged biological concepts

A couple of crucial elements are chargeable for the performance of a cleansing fluid in the canal system. First, to execute its feature, the irrigation mixture needs to touch with both the cells, which it has to act on, and the microbes to damage. Constant, if not continual, irrigation replacement permits much better outcomes, considering the buffering effect due to the loss of chlorine of the dentin as well as the narrow spaces of the root canal system [21]. An additional vital factor for the performance of irrigate solutions is their time of action. In fact, to attain a full disinfection of the root canal system, one needs to let the NaOCl compound act inside the canal for,

a minimum of, 30 min, because it has been shown that, if used for a much shorter time, the therapy result lowers.

A variable straight affecting the cleaning of the root canal system is the one determined by the mechanical impact of irrigants on canal walls, called shear tension [22]. It is the force applied on the canal surerficies by the circulation of irrigant services.

Table 3. Objectives and requirements of irrigants

<i>The aims of root canal irrigation are:</i>	<i>The following are requirements of a root canal irrigant:</i>
<ul style="list-style-type: none"> • Reduction of intraradicular microorganisms and neutralisation of endotoxins. • Dissolution of vital or necrotic pulp tissue. • Lubrication of canal walls and instruments. • Removal of dentine particles. 	<ul style="list-style-type: none"> • A broad antimicrobial spectrum. • Biocompatibility. • Tissue-dissolution capability

NaOCl

The applience of irrigatory mixtures is a vital aspect of efficient chemomechanical prep work. It enhances microbial removal and facilitates removal of lethal cells as well as dentine chips from the root canal. Irrigants can protect against packaging of the infected hard as well as soft tissue apically in the root canal and also into the periapical location. NaOCl is the extensively used irrigating solution.

NaOCl is best known for its strong anti-bacterial activity; it kills microorganisms very swiftly even at low unities. Waltimo et al. showed that the resistant microbe, *Candida albicans*, was eliminated in vitro in 30 s by either 5% and 0.5% NaOCl, whereas unities 0.05% as well as 0.005% were too weak to kill the yeast also after 24 h of incubation [23]. The great sensitivity of *C. albicans* to the solution of NaOCl was lately additionally verified by Radcliffe et al. [24]. However, Vianna et al. compared these results partially, as 0.5% NaOCl called for 30 minutes to kill *C. albicans*, whereas 5.25% solution eliminated all yeast cells in 15 s [25]. Gomes et al. checked in vitro the result of different concentrations against *E. faecalis* [26]. The microorganism was dead in less than 30 s by the 5.25% remedy, whereas it needed 10 and 30 minutes for total murder of the bacteria by 2.5% and 0.5% mixtures, specifically. The clearly higher resistance to hypochlorite by *E. faecalis* as compared with the yeast *C. albicans* was verified by Radcliffe et al. [24]. Recent laboratory experimentations utilizing 3 Gramnegative anaerobic rods usually separated from main AP, *Porphyromonas gingivalis*, P.

endodontalis, as well as *Prevotella intermedia* demonstrated high vulnerability to NaOCl, and all 3 varieties were killed within 15 s with all unities checked (0.5-- 5%) [25].

Bystrom & Sundqvist revealed that even though 0.5% NaOCl, with or without ethylenediamine-tetra-acetic acid (EDTA), increased the anti-bacterial performance of preparation compared to saline irrigation, all canals could not made germs complimentary also after a number of visits [15],[16].

Chávez de Paz et al. showed that 1% NaOCl harmed the membrane layer stability and got rid of most biofilm cells (24 h old) after 5 min touch with 4 different strains isolated from lingered instances of AP [27]. Nevertheless, fully grown biofilms are extra resistant than their young equivalents. Wang et al. demonstrated that NaOCl was a lot more efficient in killing 1-day-old *Enterococcus faecalis* biofilm than 3- week-old biofilm. This impact was directionally proportional to direct exposure time as well as NaOCl focus [28]. Liu et al. revealed that the resistivity of *E. faecalis* biofilms is affected by the maturity of biofilms and also was additionally dependent on the physical phase of biofilms [29].

Table 4. Summary of NaOCl effectiveness [22-27].

<p>Advantages</p> <ul style="list-style-type: none"> • Its well-established antimicrobial activity. • The most important advantage that makes sodium hypochlorite stands out amongst all irrigants is the tissue solvent capacity. • It acts as a lubricant. • Quick onset.
<p>Disadvantages</p> <ul style="list-style-type: none"> • Toxic • Not substantive • Bad odor and corrosion • Does not dissolve the inorganic part of smear layer

Chlorhexidine (CHX)

CHX is most likely the most commonly utilized biocide in antiseptic products generally. It is able to permeate the cell wall or outer membrane and also assault the microbial cytoplasm or internal membrane layer or the yeast plasma membrane. High concentrations of CHX trigger coagulation of intracellular constituents [33]. CHX gluconate has remained in use for a prolonged period of time in dental care due to its anti-microbial features, its substantivity, and also its reasonably low toxic effect. Regardless of the advantages of CHX, its effectiveness depends on pH and also is considerably lowered in the existence of biological matter [30]. It has a broad anti-microbial feature and also is efficient against both Gram-positive and also Gram-negative microorganisms in addition to

yeasts, whereas mycobacteria and spores of the bacteria are resisting to CHX [31]. CHX is ruled out to be an efficient antiviral agent, and its action is restricted to lipid-enveloped viruses [32].

The retention of CHX to dentin canal wall in an adequate antimicrobial amount was shown to last for at the very least 12 weeks after obturation [34] These results were validated by Souza et al., who showed that 2% CHX in either mixture or gel form could be preserved within root canal dentin for approximately 90 days [35].

A latest systematic evaluation comparing the antimicrobial efficiency of NaOCl and chlorhexidine (CHX) stated variable outcomes [36]. One research study showed NaOCl to be extra efficient than CHX [37]. One more demonstrated contrary findings [38]. Two research studies reported both irrigants are equally reliable, as well as one showed both are ineffective in getting rid of endotoxins [39-41]. The literature is full of comparable research studies with contrasting verdicts.

Table 5. Summary of CHX effectiveness [33-38].

<p>Advantages</p> <ul style="list-style-type: none"> Substantive antimicrobial activity Low key toxicity
<p>Disadvantage</p> <ul style="list-style-type: none"> Lack of tissue dissolving capacity
<p>Mechanism of action</p> <p>CHX is either bacteriostatic or bacteriocidal depending on its concentration. CHX cationic molecules bind to negatively charged bacterial components on cell wall</p>

Chelator Solutions

Although sodium hypochlorite seems the most preferable solitary endodontic irrigant, it cannot dissolve inorganic dentin particles and also thus avoid the development of a smear layer during instrumentation [43]. In addition, calcifications impeding mechanical preparation are frequently run into in the canal system. Demineralizing agents such as ethylenediamine tetraacetic acid (EDTA) and also citric acid have actually therefore been suggested as adjuvants in root canal treatment [43-44]. These are extremely biocompatible as well as are generally made use of in individual treatment products [45]. Although citric acid seems somewhat much more potent at comparable focus than EDTA, either agents reveal high performance in removing the smear layer [46]. Along with their cleaning capacity, chelators may remove biofilms adhering to root canal wall surfaces (Kishor Gulabivala, personal communication). This might discuss why an EDTA irrigant verified to be highly superior to saline in decreasing intracanal microbiota, despite the fact that its disinfectant capability is relatively limited [47-48]. Albeit never displayed in a randomized clinical

trial, an alternating irrigating program of NaOCl and also EDTA may be a lot more effective in reducing microbial burdens in root canal systems than NaOCl alone [16]. Sterilizers like quaternary ammonium compounds (EDTAC) or tetracycline anti-biotics (MTAD) have been included in EDTA and citric acid irrigants, respectively, to raise their antimicrobial capacity [43]. The clinical valuation of this, nevertheless, is doubtful. EDTAC shows similar smear-removing efficiency as EDTA, but it is more caustic [48].

Hydroxyethylidene bisphosphonate (HEBP), additionally named etidronate, is a decalcifying agent that shows only little temporary disturbance with sodium hypochlorite. It has just recently been suggested as a feasible alternative to citric acid or EDTA [49]. HEBP protects against bone traction as well as is utilized systemically in people dealing with osteoporosis or Paget's disease [50]. Nevertheless, whether this representative will improve or abbreviate endodontic cleansing will certainly need to be received future researches.

Hydrogen peroxide

Hydrogen peroxide (H₂O₂) is applied in dental care in various concentrations varying from 1% to 30% [13]. For endodontic therapy, a unity in between 3% and also 5% is chosen. Solutions of H₂O₂ are chemically stable and H₂O₂ is active against bacterias, yeasts, and also viruses due to the production of hydroxy free radicals (\cdot OH) [13]. These free compounds attack some cell structures such as proteins as well as DNA [13]. The anti-microbial performance as well as the tissue dissolving capacity of H₂O₂ are unsatisfactory in comparison with NaOCl. To conclude, there is no clinical proof indicating that H₂O₂ might transcend to various other irrigants [13].

CONCLUSION:

Apical periodontitis (AP) is a common infectious condition around the world as well as is characterized by an inflammatory response and also bone destruction in the periapical tissues triggered by microbial infection in the tooth pulp. It takes place as a series of numerous insults to the teeth pulp, including infection, physical as well as iatrogenic trauma, following endodontic treatment, the damaging impacts of root canal filling components. In response, the host positions an array of defenses, consisting of a number of classes of cells, intercellular messengers, antibodies and effector particles. The microbial aspects as well as host protection drives encounter, clash with, and also damage much of the periapical cells, ending up in a development of different type of AP lesions, which most commonly take the form of responsive granulomas and also cysts, with the concomitant

traction of bone bordering the roots of influenced teeth.

It is shown that mechanical instrumentation is able to minimize substantially the number of germs in the root canal system. However, as a result of complex composition foreseeable as well as complete microbial elimination does not appear to be possible, either with conventional hand instrumentation or with more recent rotating NiTi-systems. Irrigation of the root canal is necessary for efficient removal of pathogens.

The performance of irrigation in the facility root canal system at the apical root canal stays one of the important restorative needs to deal with AP. Experimental approaches and scientific information revealed that irrigation is one of the most effective way for removal of the root canal germs (planktonic and biofilms). The qualities of irrigation procedure, such as the mixture flow dynamics as well as irrigate substitute, have been lately stood out as the essential elements in successful treatment outcome.

REFERENCE:

1. Kakehashi S, Stanley HR, Fitzgerald RJ. The effects of surgical exposures of dental pulps in germ-free and conventional laboratory rats. *Oral Surg Oral Med Oral Pathol* 1965; 20:340–9.
2. Tziafas, D., Alraeesi, D., Al Hormoodi, R., Ataya, M., Fezai, H., & Aga, N. Preparation Prerequisites for Effective Irrigation of Apical Root Canal: A Critical Review. *Journal of Clinical and Experimental Dentistry*, 2017; e1256–e1263. doi:10.4317/jced.54117.
3. Jafarzadeh H, Abbott PV. Review of pulp sensibility tests. Part 1: general information and thermal tests. *Int Endod J* 2010;43:738-62.
4. Jafarzadeh H, Abbott PV. Review of pulp sensibility tests. Part II: electric pulp tests and test cavities. *Int Endod J* 2010;43:945-58.
5. Newton CW, Hoen MM, Goodis HE, Johnson BR, McClanahan SB. Identify and determine the metrics, hierarchy, and predictive value of all the parameters and/or methods used during endodontic diagnosis. *J Endod* 2009;35:1635.
6. Levin LG, Law AS, Holland GR, Abbot PV, Roda RS. Identify and define all diagnostic terms for pulpal health and disease states. *J Endod* 2009;35:1645.
7. Gutmann JL, Baumgartner JC, Gluskin AH, Hartwell GR, Walton RE. Identify and define all diagnostic terms for periapical/periradicular health and disease states. *J Endod* 2009;35:1658.
8. Rosenberg PA, Schindler WG, Krell KV, Hicks ML, Davis SB. Identify the endodontic treatment modalities. *J Endod* 2009;35:1675.

9. Gutmann, J.L. et al. Identify and define all diagnostic terms for periapical/ periradicular health and disease states. *J Endod* 2009;35: 1658–1674.
10. Abbott PV. Classification, diagnosis and clinical manifestations of apical periodontitis. *Endodontic Topics* 2004, 8, 36–54.
11. Hess W. Zur Anatomie der Wurzelkanäle des menschlichen Gebisses mit Berücksichtigung der feinen Verzweigungen am Foramen apikale. *Schweiz Vierteljahresschr Zahnheilk* 1917;1:1-53.
12. Peters OA. Current challenges and concepts in the preparation of root canal systems: a review. *J Endod* 2004;30:559-567.
13. Haapasalo M, Endal U, Zandi H, Coil J. Eradication of endodontic infection by instrumentation and irrigation solutions. *Endodontic Topics* 2005;10:77-102.
14. Trope M, Bergenholtz G. Microbiological basis for endodontic treatment: can a maximal outcome be achieved in one visit? *Endodontic Topics* 2002;1:40-53.
15. Byström A, Sundqvist G. Bacteriologic evaluation of the efficacy of mechanical root instrumentation in endodontic therapy. *Scand J Dent Res* 1981;89:321-328.
16. Byström A, Sundqvist G. The antibacterial action of sodium hypochlorite and EDTA in 60 cases of endodontic therapy. *Int Endod J* 1985;18:35-40.
17. Nair PW, Henry S, Cano V, Vera J. Microbial status of apical root canal system of human mandibular first molars with primary apical periodontitis after “one-visit” endodontic treatment. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod* 2005;99:231-252.
18. Dalton BC, Ørstavik D, Phillips C, Pettiette M, Trope M. Bacterial reduction with nickel-titanium rotary instrumentation. *J Endod* 1998;24:763-767.
19. Haapasalo M, Shen Y, Wang Z, Gao Y. Irrigation in endodontics. *Brit Dent Journal*. 2014;216(6):299–303.
20. Siqueira JF, Lima KC, Magalhaes FA, Lopes HP, de Uzeda M. Mechanical reduction of the bacterial population in the root canal by three instrumentation techniques. *J Endod*. 1999;25:332–5.
21. Arias-Moliz MT, Morago A, Ordinola-Zapata R, Ferrer-Luque CM, Ruiz-Linares M, Baca P, et al. Effects of dentin debris on the antimicrobial properties of sodium hypochlorite and etidronic acid. *J Endod*. 2016;42:771–5.
22. van der Sluis LW. Endodontics in motion: New concepts, materials and techniques 3. The role of irrigants during root canal treatment. *Ned Tijdschr Tandheelkd*. 2015;122:533–8.
23. Waltimo TM, Ørstavik D, Siren EK, Haapasalo MP. In vitro susceptibility of *Candida albicans* to four disinfectants and their combinations. *Int Endod J* 1999; 32: 421–429.
24. Radcliffe CE, Potouridou L, Qureshi R, Hababbeh N, Qualtrough A, Worthington H, Drucker DB. Antimicrobial activity of varying concentrations of sodium hypochlorite on the endodontic microorganisms *Actinomyces israelii*, *A. naeslundii*, *Candida albicans* and *Enterococcus faecalis*. *Int Endod J* 2004: 37: 438–446.
25. Vianna ME, Gomes BP, Berber VB, Zaia AA, Ferraz CC, de Souza-Filho FJ. In vitro evaluation of the antimicrobial activity of chlorhexidine and sodium hypochlorite. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod* 2004; 97: 79–84.
26. Gomes BP, Ferraz CC, Vianna ME, Berber VB, Teixeira FB, Souza-Filho FJ. In vitro antimicrobial activity of several concentrations of sodium hypochlorite and chlorhexidine gluconate in the elimination of *Enterococcus faecalis*. *Int Endod J* 2001: 34: 424–428.
27. Yang Y, Shen Y, Wang ZJ, Huang XY, Maezono H, Ma JZ, et al. Evaluation of the susceptibility of multispecies biofilms in dentinal tubules to disinfecting solutions. *J Endod*. 2016;42(8):1246–50.
28. Liu H, Wei X, Ling J, Wang W, Huang X. Biofilm formation capability of *Enterococcus faecalis* cells in starvation phase and its susceptibility to sodium hypochlorite. *J Endod*. 2010;36(4): 630–5.
29. Du TF, Wang ZJ, Shen Y, Ma JZ, Cao YG, Haapasalo M. Combined antibacterial effect of sodium hypochlorite and root canal sealers against *Enterococcus faecalis* biofilms in dentin canals. *J Endod*. 2015;41(8):1294–8.
30. Russell AD, Day MJ. Antibacterial activity of chlorhexidine. *J Hosp Infect* 1993: 25: 229–238.
31. Shaker LA, Dancer BN, Russell AD, Furr JR. Emergence and development of chlorhexidine resistance during sporulation of *Bacillus subtilis* 168. *FEMS Microbiol Lett* 1988: 51: 73–76.
32. Park JB, Park NH. Effect of chlorhexidine on the in vitro and in vivo herpes simplex virus infection. *Oral Surg* 1989: 67: 149–153.
33. McDonnell G, Russell D. Antiseptics and disinfectants: activity, action, and resistance. *Clin Microbiol Rev* 1999: 12: 147–179.
34. Rosenthal S, Spångberg L, Safavi K. Chlorhexidine substantivity in root canal dentin. *Oral Surgery, Oral Medicine, Oral Pathology, Oral Radiology, and Endodontology*. 2004;98(4):488–92.
35. Souza M, Cecchin D, Farina AP, Leite CE, Cruz FF, Pereira Cda C, et al. Evaluation of chlorhexidine substantivity on human dentin: a

- chemical analysis. *J Endod.* 2012;38(9):1249–52.
36. Goncalves LS, Rodrigues RCV, Andrade CV, Soares RG, Vettore MV. The effect of sodium hypochlorite and chlorhexidine as irrigant solutions for root canal disinfection: a systematic review of clinical trials. *J Endod.* 2016;42(4):527–32.
 37. Vianna ME, Conrads G, Gomes BPFA, Horz HP. Identification and quantification of Archaea involved in primary endodontic infections. *J Clin Microbiol.* 2006;44(4):1274–82.
 38. Ercan ER, Ozekinci T, Atakul F, Gul K. Antibacterial activity of 2% chlorhexidine gluconate and 5.25% sodium hypochlorite in infected root canal: in vivo study. *J Endod.* 2004;30(2):84–7.
 39. Karuvilla JR, Kamath MP. Antimicrobial activity of 2.5% sodium hypochlorite and 0.2% chlorhexidine gluconate separately and combined, as endodontic irrigants. *J Endod.* 1998;24(7):472–6.
 40. Rocas IN, Siqueira JF. Comparison of the in vivo antimicrobial effectiveness of sodium hypochlorite and chlorhexidine used as root canal irrigants: a molecular microbiology study. *J Endod.* 2011;37(2):143–50.
 41. Gomes BPFA, Martinho FC, Vianna ME. Comparison of 2.5% sodium hypochlorite and 2% chlorhexidine gel on oral bacterial lipopolysaccharide reduction from primarily infected root canals. *J Endod.* 2009;35(10):1350–3.
 42. Lester KS, Boyde A. Scanning electron microscopy of instrumented, irrigated and filled root canals. *Br Dent J* 1977;143:359 – 67.
 43. Nygaard Östby B. Chelation in root canal therapy. *Odontol Tidskr* 1957;65:3–11.
 44. Loel DA. Use of acid cleanser in endodontic therapy. *J Am Dent Assoc* 1975;90:148 –51.
 45. Coons D, Dankowski M, Diehl M, et al. Performance in detergents, cleaning agents and personal care products: detergents. In: Falbe J, ed. *Surfactants in consumer products*. Berlin: Springer-Verlag, 1987:197–305.
 46. Zehnder M, Schmidlin P, Sener B, Waltimo T. Chelation in root canal therapy reconsidered. *J Endod* 2005;31:817–20.
 47. Yoshida T, Shibata T, Shinohara T, Gomyo S, Sekine I. Clinical evaluation of the efficacy of EDTA solution as an endodontic irrigant. *J Endod* 1995;21:592–3.
 48. Patterson SS. In vivo and in vitro studies of the effect of the disodium salt of ethylenediamine tetra-acetate on human dentine and its endodontic implications. *Oral Surg Oral Med Oral Pathol* 1963;16:83–103.
 49. Zehnder M, Schicht O, Sener B, Schmidlin P. Reducing surface tension in endodontic chelator solutions has no effect on their ability to remove calcium from instrumented root canals. *J Endod* 2005;31:590 –2.
 50. Russell RG, Rogers MJ. Bisphosphonates: from the laboratory to the clinic and back again. *Bone* 1999;25:97–106.