

A Review on Triple Antibiotic Paste as an antimicrobial agent in Endodontics

Dr. Harleen Aujla

Submitted: 25-01-2022	Revised: 01-02-2022	Accepted: 04-02-2022

ABSTRACT: The favorable outcome of an endodontic treatment depends on the elimination of the microbial colonies present in the root canal and periapical region. Also, if any necrotic tissue is left remaining in the root canals, the efficacy of the root canal medicaments decreases. Triple antibiotic paste (TAP) containing metronidazole, ciprofloxacin and minocycline has been proposed as a suitable root canal medicament due to its antimicrobial effects. This review article discusses the properties of TAP and its efficiency on the root canal disinfection, both in primary and permanent dentition, along with its relation in regenerative endodontic procedures. This paper also reviews the present disadvantages of the Triple antibiotic paste. Endodontics Keywords: treatment, Triple antibiotic paste, root canal medicaments

I. INTRODUCTION:

Micro-organisms are a predominant factor in Endodontics infections. Several Microbial colonies have been identified in root canal infections which include both aerobic and anaerobic organisms. Keeping in mind the complexity of the root canal systems, a combination of antibiotics is required along with mechanical instrumentation to target the diverse flora and create a bacteria-free environment. For this purpose, the antibiotic combination that appears most effective is that of metronidazole, ciprofloxacin and minocycline, commonly known as the Triple Antibiotic Paste.

Dating back to 1951, Grossman was one of the first to use an antibiotic paste in Endodontics. He formulated a poly-antibiotic paste known as PSBC, which was a combination of Penicillin, bacitracin, streptomycin and caprylate sodium in a silicone vehicle. Although the PSBC showed significant effects, it was not quite effective against the anaerobic microbes. As a result the use of PSBC was very limited.

Presently, antibiotic preparations containing corticosteroids as anti-inflammatory agents are also being used commercially, which include Ledermix (Lederle Pharmaceuticals, Wolfratshausen, Germany) and Septomixine Forte (Septodont, Saint- Maur, France). But because of their narrow spectrum of activity none are rendered suitable for use in Endodontics.

Hoshino and colleagues initially developed the Triple Antibiotic Paste (TAP) and investigated the efficacy of the paste in eliminating microorganisms from the root canal system. TAP is a combination of Metronidazole, ciprofloxacin and minocycline with 1:1:1 ratio. The outcome showed excellent results in the elimination of the bacteria from the radicular system.

Metronidazole is a broad spectrum nitroimidazole. It is known for its antibacterial activity against anaerobic cocci and gram-negative as well as gram-positive bacilli.

Tetracyclines such as minocycline and doxycycline are broad spectrum antibiotics with bacteriostatic activity against gram-positive and gram-negative organisms. They are also considered to be effective against anaerobic and facultative bacteria.

Ciprofloxacin is a Fluoroquinolone with a bactericidal mode of action. It is highly effective against gram-negative microorganisms. Due to the tendency of resistance development against ciprofloxacin, it is usually combined with metronidazole in the treatment of poly microbial infections.

A study by Sato et al showed that TAP was very effective in killing bacteria in the deep layers of radical dentin. This antibiotic combination was also used by Banchs and Trope in a recent clinical report, as well as by Iwaya et al in a case report demonstrating revascularization of an immature tooth with Apical periodontitis.

II. APPLICATIONS OF TAP:

Root canal disinfection



Root canal infections consist of both aerobic and anaerobic bacteria. Keeping this in mind, the most effective and practical combination of antibacterial is TAP. As previously discussed, an in situ study by Sato et al evaluated the potency of TAP and as per the results, no micro organisms were found in the infected radicular dentin within 24 hrs of TAP application. Thus putting forth the opinion to treat periodical lesions first with the Triple antibiotic paste, followed by other endodontic procedures. Özan and Er found the combination of drugs in TAP are also helpful in treating large cyst-like lesions and periodical lesions.

<u>Regeneration and revascularization</u>

Tap is effective in eliminating microbes from the infected pulps, thus creating a favourable environment for new vital tissue to regenerate. Endodontics Regenerative procedures (ERPs) are considered the latest advancement in Endodontics. TAP has shown to present with thickening of radicular dentin walls, apical foramen closures and continued root development. Studies have also shown that the percentage of increased dentin thickness was higher with TAP as compared to Calcium Hydroxide or formocresol.

<u>As an Intracanal Medicament</u>

During the root canal treatment, TAP acts as an antiseptic agent for the root canal systems. In a study, Murvindran and James exhibited the potential of TAP to remove the microbial colonies and create a suitable microbe free environment. At the same time, Kim and Kim investigated that the triple antibiotic paste also showed characteristics of inhibiting E. Faecalis. Additionally, various clinical studies and case reports of horizontally fractured teeth, where TAP was used as an intracranial medicament and MTA as the coronal obturator showed repair of the fracture on radiographs with disappearance of symptoms within 12 months from initiating the treatment.

• <u>Treatment of Primary teeth</u>

The lesion sterilization and tissue repair (LSTR) therapy, employs the TAP and has shown to be successful in eliminating pathogens from the root canals effectively. But the complete eradication of infection from the infected primary teeth is still a challenge, because of the complexity of the root canals leading to difficulty in mechanical debridement of the canals. Nakornchai et al demonstrated that both Vitapex and TAP were effective in root canal treatment of infected primary dentition. The study showed a success rate of 100% and 96% respectively for TAP and Vitapex.

III. DRAWBACKS:

Post treatment with the triple antibiotic paste, studies were conducted and tooth discolouration emerged to be a major concern. And Minocycline was considered the leading cause of discolouration. Reynolds et al before placement of the TAP used a dentin bonding agent to prevent the discolouration, but the discolouration was only reduced and not completely eliminated.

Also, numerous studies have shown that the concentration of TAP used in regeneration and revascularization procedures can cause notable loss of radicular dentin and a considerable increase in roughness of dentin.

To further add to the list, removal of the Triple antibiotic paste from the radicular canals is a tedious process. This is primarily because TAP penetrates and binds with the dentinal structures, making it difficult to be removed. Chlorhexidine is the least effective of all and ultrasonic activation of 5.25 % Sodium Hypochlorite is considered the most effective. Recently, an investigation by Turkaydin et al showed the use of XP-Endo Finisher, which removed a larger amount of TAP from the root canal space than syringe or ultrasonic irrigation techniques.

IV. CONCLUSION :

A root canal treatment is considered successful when the microbial colonies are either reduced or eliminated completely from the root canal system. Although a crucial part of endodontic treatment, Biomechanical instrumentation alone cannot provide a sterile environment in the canal system. To aid in the eradication of the microbiota, TAP can effectively be used for sterilization of the root canals. The triple antibiotic paste has shown to be a good medicament in revascularization and regeneration. While we continue to research and find the ideal antimicrobial material, TAP has truly shown to be one of the top contenders in Endodontics.

REFERENCES

[1]. Triple antibiotic paste in root canal therapy Rangasamy Vijayaraghavan, Veerabathran Mahesh Mathian, Alagappan Meenakshi Sundaram, Ramachandran Karunakaran, and Selvaraj Vinodh



- [2]. A Review on Triple Antibiotic Paste as a Suitable Material Used in Regenerative Endodontics Zahed Mohammadi, Hamid Jafarzadeh, Sousan Shalavi, Shapour Yaripour, Farid Sharifi, and Jun-Ichiro Kinoshita
- [3]. Sundqvist G. Ecology of the root canal flora. J Endod. 1992;18:427–30. [PubMed] [Google Scholar]
- [4]. Sato I, Ando-Kurihara N, Kota K, Iwaku M, Hoshino E. Sterilization of infected rootcanal dentine by topical application of a mixture of ciprofloxacin, metronidazole and minocycline in situ. Int Endod J. 1996;29:118–24. [PubMed] [Google Scholar]
- [5]. 5.Trope M. Treatment of immature teeth with non-vital pulps and apical periodontitis. Endodod Topics. 2006;14:51–9. [Google Scholar]
- [6]. Reynolds K, Johnson JD, Cohenca N. Pulp revascularization of necrotic bilateral bicuspids using a modified novel technique to eliminate potential coronal discolouration: A case report. Int Endod J. 2009;42:84–92. [PubMed] [Google Scholar]
- [7]. Hoshino E, Takushige T. LSTR 3Mix-MP method-better and efficient clinical procedures of lesion sterilization and tissue repair (LSTR) therapy. Dent Rev. 1998;666:57–106. [Google Scholar]
- [8]. Kim J, Kim Y, Shin S, Park J, Jung I. Tooth discoloration of immature permanent incisor associated with triple antibiotic therapy: A case report. J Endod. 2010;36:1086–91. [PubMed] [Google Scholar]
- [9]. Thibodeau B, Trope M. Pulp revascularization of a necrotic infected immature permanent tooth: Case report and review of the literature. Pediatr Dent. 2007;29:47–50. [PubMed] [Google Scholar]
- [10]. Banchs F, Trope M. Revascularization of immature permanent teeth with apical periodontitis: New treatment protocol. J Endod. 2004;30:196–200. [PubMed] [Google Scholar]
- [11]. Bansal R, Bansal R. Regenerative endodontics: A state of art. Indian J Dent Res. 2011;22:122–31. [PubMed] [Google Scholar]
- [12]. Mohammadi Z, Jafarzadeh H, Shalavi S, Palazzi F. Recent Advances in Root Canal Disinfection: A Review. Iran Endod J. 2017;12(4):402–6. [PubMed] [Google Scholar]

- [13]. Disinfection of Immature Teeth with a Triple Antibiotic Paste
- [14]. William Windley, III, DDS, MS, Fabricio Teixeira, DDS, MSc, PhD, Linda Levin, DDS, PhD, Asgeir Sigurdsson, DDS, MS, Martin Trope, DMD
- [15]. Infection Control in Regenerative Endodontic Procedures By Su-Min Lee, D.D.S., M.S., DScD
- [16]. Comparative Evaluation of Antimicrobial effects of Triple Antibiotic Paste and Amox and its derivatives against E. faecalis: An in vitro study
- [17]. Manjeet Kaur, Shrikant Kendre, Parmod Gupta, Navneet Singh, Harsimran Sethi, Neha Gupta, Rushil Acharya
- [18]. Microbiological Aspects of Root Canal Infections and Disinfection Strategies: An Update Review on the Current Knowledge and Challenges
- [19]. Jasmine Wong, Daniel Manoil, Peggy Näsman, Georgios N. Belibasakis and Prasanna Neelakantan
- [20]. Endodontic treatment of chronically infected primary teeth using triple antibiotic paste: An in vivo study - G. Anuradha Reddy,E. Sridevi, A. J. Sai Sankar, K. Pranitha, M. J. S Pratap Gowd, and C. Vinay
- [21]. Bystrom A, Sundqvist G. Bacteriologic evaluation of the efficacy of mechanical root canal instrumentation in Endodontics therapy. Scand j dent res. 1981;89(4):321-8 [Pubmed]] [Google Scholar]
- [22]. Windley W, 3rd , Teixeira F, Levin L, Sigurdsson A, Trope M. Disinfection of immature teeth with a triple antibiotic paste. J Endod. 2005;31(6):439–43. [PubMed] [Google Scholar]
- [23]. Kim ST, Abbott PV, McGinley P. The effects of Ledermix paste on discolouration of mature teeth. Int Endod J. 2000;33(3):227–32.
- [24]. Yilmaz S, Dumani A, Yoldas O. The effect of antibiotic pastes on microhardness of dentin. Dent Traumatol. 2016;32(1):27–31.[PubMed] [Google Scholar]
- [25]. Yassen GH, Eckert GJ, Platt JA. Effect of intracanal medicaments used in endodontic regeneration procedures on microhardness and chemical structure of dentin. Restor Dent Endod. 2015;40(2):104–12. [[PubMed]]
- [26]. Cvek M, Nord CE, Hollender L. Antimicrobial effect of root canal debridement in teeth with immature roots. A clinical and microbiologic study. Odontol Revy 1976; 27:1–10.



- [27]. Fouad AF. Microbial Factors and Antimicrobial Strategies in Dental Pulp Regeneration. J Endod. 2017;43(9S):S46-S50
- [28]. Abbott PV. Selective and intelligent use of antibiotics in Endodontics. Aust Endo J 2000;26:30-9
- [29]. Gilad JZ, Teles R, Goodson M, White RR, Stashenko P. Development of a Clindamycin impregnated fiber as an intracanal medication in endodontic therapy. J Endod 1999;25:722-7.
- [30]. ADA Council on Scientific Affairs. Antibiotics used in dentistry. J Am Dent Assoc 1997;128:648
- [31]. da Silva LA, Nelson-Filho P, da Silva RA, Flores DS, Heilborn C, Johnson JD, et al. Revascularization and periapical repair after endodontic treatment using apical negative pressure irrigation versus conventional irrigation plus triantibiotic intracanal dressing in dogs' teeth with apical periodontitis. Oral Surg Oral Med Oral Pathol Oral Radiol Endod 2010;109:779-87
- [32]. Mouton Y. The antibiotics. Role of antibiotics in the prevention of infectious diseases. Lille Med 1975;20:914-915. [[PubMed]
- [33]. Wynn RL, Bergman SA, Meiller TF, Crossley HL. Antibiotics in treating oralfacial infections of odontogenic origin: an update. Gen Dent 2001;49:238-244. [PubMed]
- [34]. Özan U, Er K. Endodontic treatment of a large cyst-like periradicular lesion using a combination of antibiotic drugs: a case report. J Endod 2005;31:898-900. [PubMed]
- [35]. Akman M, Akbulut MB, Aydinbelge HA, Belli S. Comparison of different irrigation activation regimens and conventional irrigation techniques for the removal of modified triple antibiotic paste from root canals. J Endod 2015;41:720-724. [PubMed]
- [36]. Berkhoff JA, Chen PB, Teixeira FB, Diogenes A. Evaluation of triple antibiotic paste [PubMed]
- [37]. Kim B, Song MJ, Shin SJ, Park JW. Prevention of tooth discoloration associated with triple antibiotics. Restor Dent Endod 2012;37:119-122.
- [38]. Kirchhoff AL, Raldi DP, Salles AC, Cunha RS, Mello I. Tooth discoloration and internal bleaching after the use of triple antibiotic paste. Int Endod J 2015;48:1181-1187. [PubMed]

- [39]. Pai S, Vivekananda Pai AR, Thomas MS, Bhat V. Effect of calcium hydroxide and triple antibiotic paste as an intracanal medicaments on the incidence of interappointment flare-up in diabetic patients: an in vivo study. J Conserv Dent 2014;17:208-211. [PubMed]
- [40]. Trope M. Endodontic considerations in dental trauma. In: Ingle JI, Bakland LK, Baumgartner JC, editors. Ingle's endodontics. Hamilton (ON): BC Decker; 2008. p1330-1357.
- [41]. Prather BT, Ehrlich Y, Spolnik K, Platt JA, Yassen GH. Effects of two combinations of triple antibiotic paste used in endodontic regeneration on root microhardness and chemical structure of radicular dentine. J Oral Sci 2014;56:245-251. [PubMed]
- [42]. Dhillon JS, Amita, Saini SK, Bedi HS, Ratol SS, Gill B. Healing of a large periapical lesion using triple antibiotic paste and intracanal aspiration in nonsurgical endodontic retreatment. Indian J Dent 2014;5:161-165. [PubMed]
- [43]. Sabrah AH, Yassen GH, Spolnik KJ, Hara AT, Platt JA, Gregory RL. Evaluation of residual antibacterial effect of human radicular dentin treated with triple and double antibiotic pastes. J Endod 2015;41:1081-1084.
- [44]. Trairatvorakul C, Detsomboonrat P. Success rates of a mixture of ciprofloxacin, metronidazole, and minocycline antibiotics used in the non-instrumentation endodontic treatment of mandibular primary molars with carious pulpal involvement. Int J Paediatr Dent 2012;22:217-227.
- [45]. Manuel ST, Parolia A, Kundabala M, Vikram M. Non-surgical endodontic therapy using triple-antibiotic paste. Kerala Dental Journal. 2010;33:88–90. [Google Scholar]
- [46]. Gomes-Filho JE, Duarte PCT, de Oliveira CB, Watanabe S, Lodi CS, Cintra LTA, et al. Tissue reaction to a triantibiotic paste used for endodontic tissue self-regeneration of nonvital immature permanent teeth. J Endod. 2012;38:91–4. [PubMed]
- [47]. Lin LM, Skribner JE, Gaengler P. Factors associated with endodontic treatment failures. J Endod. (1992) 18:625–7.[Pubmed] [Google Scholar]
- [48]. Pinheiro ET, Gomes BP, Ferraz CC, Sousa EL, Teixeira FB, Souza-Filho FJ. Microorganisms from canals of root-filled teeth with periapical lesions. Int Endod J. (2003) 36:1–11. [PubMed] [Google Scholar]



- [49]. Hoshino E, Kurihara-Ando N, Sato I, Uematsu H, Sato M, Kota K, Iwaku M. Invitro antibacterial susceptibility of bacteria taken from infected root dentine to a mixture of ciprofloxacin, metronidazole and minocycline. Int Endod J. 1996;29(2):125– 30. [PubMed] [Google Scholar]
- [50]. Albuquerque MT, Ryan SJ, Munchow EA, Kamocka MM, Gregory RL, Valera MC, Bottino MC. Antimicrobial Effects of Novel Triple Antibiotic Paste-Mimic Scaffolds on Actinomyces naeslundii Biofilm. J Endod. 2015;41(8):1337–43. [PubMed]