Bioactive glasses as group of biomaterials used in dentistry

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Abstract: Bioactive glasses are able to bond to both soft and hard tissue and promote the bone growth. The bioactivity behavior of these glasses is related to the formation of a biologically active hydroxyapatite layer on the surface of the glasses. The mechanism of bonding of bioactive glasses to tissues includes a series of surface reactions that occur when the glass is exposed to an aqueous environment. Bioactive glasses have a wide range of applications, such as bone grafts, scaffolds, coating materials, and are used for hypersensitivity treatment. One of the most important properties of bioactive glasses is their ability to exhibit antibacterial activity, which creates a bacteria-free environment while healing and regenerating the defect area. These potentials of bioactive glass make it a unique material to be widely used in dentistry. Such materials can stimulate bioactive behavior around the fixed restorations margins and provide a bioactive surface. Therefore, they can develop periodontal tissue attachment and create complete sealing of the marginal gap. This sealing can prevent the failure of fixed ceramic restorations by eliminating the secondary caries, micropenetration of oral bacteria and their adhesion on the cement surface. [Meshari H. Alanzi, Atif M. Almadani, Alyan Alazemi. Bioactive glasses as group of biomaterials used in dentistry. Life Sci J 2017;14(4):111-116]. ISSN: 1097-8135 (Print) / ISSN: 2372-613X (Online).

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1. Introduction

BAGs were introduced in dentistry: As substitutes for reconstruction of voids and defects of facial bones (Schepers, et al., 1991; Schepers, et al., 1993; Suominen and Kinnunen 1996), in rehabilitation of the dentoalveolar complex, including BAG implants (Wilson, et al., 1994) and regeneration of periodontal bone support (Larmas, et al., 1995; Shapoff, et al., 1997; Lovelace, et al., 1998). BAGs may have therapeutic value as mineralizing agents in caries prophylactics, and also as a desensitizing agent in clinical situations where opened dentinal tubules lead to hypersensitive teeth (Orchardson, et al., 1994).

Furthermore, in implantology, a coating of technically adequate BAG on the fixture surface may serve as a means to attach mucosal or dermal soft tissues to the osseointegrated construction by an HCA bridge (Linde and Berglundh 1998). In addition, BAGs may also have an application in root canal therapy providing a biological seal in the form of mineral deposition inducing materials in the root canal and at the apex (Sarin and Rekhi 2016). The base components are usually SiO2, Na2O, CaO, and P2O5 and given below are percentages in weight of the most common BAGs.

Bioglass composition in wt %:

SiO2-45 wt% Na2O - 24.5 wt% CaO - 24.5 wt% P2O5-6 wt%.

The most studied is the Bioglass **(B** 45S5. The abbreviation indicates that it contains 45% in weight of SiO2 (oxide creator) and the molar rate between Ca/P is of 5:1. Glasses with significantly lower molar rate (in the form of CaO and P2O5) do not generate connections with the bone (Sarin and Rekhi 2016).

The ability of BAG to support osteogenesis:

When the glass composition exceeds 52% by weight of SiO2, the glass will bond to the bone but not to soft tissues. This finding provided the basis for clinical use of Bioglass for implants to maintain the alveolar ridge of edentulous patients. BAG is used extensively in dentistry in the treatment of bone defects, ridge preservation, and periodontal bone defects (Subbaiah and Thomas 2011).

Mengel et al., (2006) conducted a clinical and a radiological study to compare the long-term effectiveness of a bioabsorbable membrane and a BAG in the treatment of intrabony defects in patients with generalized aggressive periodontitis, results showing significant improvement in probing depth and clinical attachment loss. Radio graphically, the defects were found to be filled significantly more in BAG group (Subbaiah and Thomas 2011). The material has also been used for repair of alveolar bone defects in humans (Mengel, et al., 2006) and recently it has been used for sinus floor augmentation in humans, showing bone regenerative activity (Furusawa and Mizunuma 1997).

BAGs of the SiO2-Na2O-CaO-P2O5 type have recently been suggested as topical root canal disinfectants (Zehnder, et al., 2004). The ability of BAG to support osteogenesis is well known but recent work has also shown its proangiogenic potential, which should provide benefits for the application of BAG to soft tissue repair which are being seen in recent works regarding the tissue-engineered regeneration of structures such as the synovial joint condyle, bone tendon complex, bone ligament junction, and the periodontium.

The potential of these materials for remineralization of both enamel and dentin has been studied in vitro and in situ and holds promise (Mukai and ten Cate 2002; Madan, et al., 2011). In addition, the unique ionic reactions and potential antimicrobial and anti-inflammatory properties might prove useful in treating gingivitis.

2. Clinical Applications

Bioactive-glass (B-G) is a material known for its favorable biological response when in contact with surrounding fibro-osseous tissues, due not only to an osteoconductive property, but also to an osteostimulatory capacity, and superior biocompatibility for use in human body, the possibility of employing these biomaterials to fill and repair dentoalveolar defects in a rapid and controllable way has been thoroughly investigated by oral and maxillofacial surgeons as well as dentists (Gosain 2004). Since its introduction, the original B-G has been released as PerioGlas® (now sold by NovaBone Products LLC, Alachua, FL, USA) for periodontal regeneration, and NovaBon (NovaBone Products LLC) or Biogran® (BIOMET 3i, Palm Beach Gardens, FL, USA) used in oral and maxillofacial surgery. Other commercial products based on meltderived calcium sodium phosphosilicates, BonAlive® (BonAlive Biomaterials, Turku, Finland) and StronBoneTM (RepRegen Ltd, UK), are also available for bone reconstructive surgery (Gosain 2004).

One of the first commercial applications of B-Gs in dentistry was to prevent the resorption of alveolar bone after tooth removal and to maintain or enhance bony ridge form for subsequent restorative treatments with implant supported prostheses (Margonar, et al., 2012). Root cones of B-G, placed in fresh residual extraction cavities as well as into artificial sockets produced by bone splitting of previous extraction sites, were able to recreate original alveolar ridge dimensions prior to dental implant surgery (Margonar, et al., 2012). In the re-entry procedure 12 months after insertion, bone formation was clearly visible providing evidence of the superiority of this material compared to other bone graft substitutes (Profeta and Prucher 2015).

There has been a number of clinical studies that have demonstrated consistent results in a variety of alternative treatments including the surgical modification/reduction (elevation) of the maxillary sinus (Stavropoulos, et al., 2012), the regeneration of inter proximal bone defects in periodontal therapies (Lovelace, et al., 1998), periapical application during endodontic microsurgery (Pantchev, et al., 2009), management of cystic defects (El-Ghannam, et al., 2004), as well as reconstructive procedures for treating peri-implantitis (Talreja, et al., 2013). All these clinical applications have one thing in common -the proven efficacy and effectiveness to bond with hard tissue and enhance its growth due to the osteoconductive and osteostimulatory properties of the glass. Osteoconduction refers to the ability to support the migration of bone starting from the walls of the defect toward the central portion of the graft (Schepers and Duchevne 1997).

New bone formation has been demonstrated histologically in human oral bone defects treated with B-G (Margonar, et al., 2012). There is also evidence that the replacement and infiltration of osseous tissue start at 4 months, and all B-G particles completely disappear at 16 months following the grafting procedure (Tadjoedin, et al., 2000). Consequently bone formation occurred in multiple growth sites, rapidly filling the bone defect. This new bone had the histologic and biomechanical properties of the surrounding bone as soon as 6–7 months after grafting (Merkx, et al., 2003). Furthermore, histology revealed rare inflammatory cells and absence of giant cells even around the remaining particles, which confirmed the biocompatibility of B-G (Margonar, et al., 2012). Antimicrobial activity of B-G (Peltola, et al., 2008). might contribute to the resolution of inflammatory responses and provide extraordinarily favorable conditions for an uneventful healing process (Tadjoedin, et al., 2000; Stavropoulos, et al., 2012). Wilson et al., (1981) was the first to document the safety of use of B-G and long-term studies confirmed that it is well tolerated in children (Lindfors 2009) and adults (Lindfors, et al., 2010).

NovaMin®

NovaMin® is technically described as an inorganic amorphous calcium sodium phosphosilicate (CSPS) material that was designed based on a class of materials known as bioactive glasses. It comprises

45% SiO2, 24.5% Na2O, 24.5% CaO and 6% P2O5 (Kobayashi, et al., 2010). The particular composition of NovaMin® is identical to that of the best known bioactive glass material, Bioglass®, and contains only calcium, sodium, phosphate and silica, all as an amorphous matrix. Its chemical formula is CaNaO6PSi. NovaMin® delivers silica and ionic calcium, phosphorus and sodium, which are necessary for bone and tooth mineralization. It was developed and patented by NovaMin Technology, Inc., (Gjorgievska and Nicholson 2011) III. Uses Of Novamin At present, bioactive glasses have a wide range of clinical applications in both medicine and dentistry (Melek, et al., 2013). It has been proposed as the material of choice for bone regeneration (Debnath, et al., 2014), hypersensitivity (Burwell, et al., 2010), antigingivitis and antiplaque effect (Tai, et al., 2006). It is also used for implant coating, as air-abrasive particles to remove carious enamel and dentine (Farooq, et al., 2012), as a dental material to improve the bonding of the restorative material to dentin (Goudouri 2011), stain removing agent (Kakodkar, et al., 2013) and in endodontic treatments (Brannstrom 1986) IV. Bone And Tissue Regeneration NovaMin® has the property to promote osteogenesis by allowing rapid formation of bone. It also may act as a barrier retarding epithelial down growth and demonstrates antimicrobial property in vivo. Due to the merits of bioglass, it is one of the preferred alloplast (Ong, et al., 1998; Gerhardt and Boccaccini 2010). Mechanism of Action In bone regeneration, bioglass particles not osteoconductivity, only show but also an osteostimulatory effect. In the treatment for dentinal hypersensitivity the physical occlusion of Novamin® particles begins when the material is subjected to an aqueous environment (Kumar, et al., 2015).

Recent studies have also demonstrated a potential for Novamin® to prevent demineralization and/or aid in remineralization of white-spot lesions. NUPRO® NUSolution TM with NovaMin® is the newest Dentsply Professional prophy paste. It is currently the only product powered by NovaMin, delivering the triple benefit of tooth desensitization, tubule occlusion and stain removal (Milleman, et al., 2012). The potential of these materials for remineralization of both enamel and dentin has been studied in vitro and in situ and holds promise. In addition, the unique ionic reactions and potential antimicrobial and anti-inflammatory properties might prove useful in treating gingivitis (Kumar, et al., 2015).

Bioactive glasses/Bioglass scaffolds

Bioactive glasses/Bioglass are very attractive materials for producing scaffolds devoted to bone regeneration due to their versatile properties, which can be properly designed depending on their composition. An important feature of bioactive glasses, which enables them to work for applications in bone tissue engineering, is their ability to enhance revascularization, osteoblast adhesion, enzyme activity and differentiation of mesenchymal stem cells as well as osteoprogenitor cells.

Many trace elements have also been incorporated in the glass network to obtain the desired properties, which have beneficial effects on bone remodeling and/or associated angiogenesis (Kaur, et al., 2014).

Europium-Containing Mesoporous Bioactive Glass Scaffolds for Stimulating in Vitro and in Vivo Osteogenesis

Europium, as an important rare earth element, has been used as a solid-state lighting material. The prepared Eu-MBG scaffolds have highly interconnective large pores, a high specific surface area, and well-ordered mesopores, as well as uniformly distributed Eu. The incorporation of 2-5 mol % Eu into MBG scaffolds gives them a luminescent property. The in vitro degradation of Eu-MBG scaffolds has a functional effect on the change of the luminescence intensity. In addition, Eu-MBG can be used for labeling bone marrow stromal cells (BMSCs) in vitro and still presents a distinct luminescence signal in deep bone tissues in vivo to label new bone tissue via release of Eu ions. Wu et al., (2016) study for the first time reports that the incorporation of the rare earth element Eu into bioscaffolds has the ability to accelerate bone regeneration in vivo, and thus, the prepared Eu-MBG scaffolds possess bifunctional properties with biolabeling and bone regeneration (Wu, et al., 2016).

S53P4 Bioactive Glass in Bone Healing and Osteomyelitic Treatment

Treatment of osteomyelitis with S53P4 bioactive glass is safe and effective even in one-stage treatment options, without the addition of local antibiotics. Adequate debridement, proper defect filling, and adequate containment of the bioactive glass granules are essential (Wu, et al., 2016).

Bioactive glass fillers reduce bacterial penetration into marginal gaps for composite restorations:

Bioactive glass (BAG) has been shown to have both an antimicrobial effect on oral bacteria and the ability to remineralize adjacent mineralized tissues (Zehnder, et al., 2004; Vollenweider, et al., 2007; Waltimo, et al., 2007; Brown, et al., 2011). This suggests BAG containing composites may have the potential to slow the development and propagation of secondary tooth decay at restoration margins. It has been shown that composites containing up to 15% by weight non-silanated BAG filler can have mechanical properties comparable to, or superior to, commercial composites (Khvostenko, et al., 2013). Tauböck et al., (2014) showed that BAG particles embedded in a resin matrix can still induce bioactivity and increase the pH of a buffered saline solution. BAG containing resin dental composites reduce biofilm penetration into marginal gaps of simulated tooth restorations (Khvostenko, et al., 2016).

References

- 1. Brannstrom M (1986). "The cause of postrestorative sensitivity and its prevention." J Endod 12(10): 475-81.
- 2. Brown ML, Davis HB, Tufekci E, Crowe JJ, Covell DA and Mitchell JC (2011). "Ion release from a novel orthodontic resin bonding agent for the reduction and/or prevention of white spot lesions. An in vitro study." Angle Orthod 81(6): 1014-20.
- 3. Burwell A, Jennings D, Muscle D and Greenspan DC (2010). "NovaMin and dentin hypersensitivity--in vitro evidence of efficacy." J Clin Dent 21(3): 66-71.
- Debnath T, Chakraborty A and Pal TK (2014). "A clinical study on the efficacy of hydroxyapatite - Bioactive glass composite granules in the management of periodontal bony defects." J Indian Soc Periodontol 18(5): 593-600.
- El-Ghannam A, Amin H, Nasr T and Shama A (2004). "Enhancement of bone regeneration and graft material resorption using surface-modified bioactive glass in cortical and human maxillary cystic bone defects." Int J Oral Maxillofac Implants 19(2): 184-91.
- Farooq I, Imran Z, Farooq U, Leghari A and Ali H (2012). "Bioactive glass: a material for future." 3(2): 199-201.
- 7. Furusawa T and Mizunuma K (1997). "Osteoconductive properties and efficacy of resorbable bioactive glass as a bone-grafting material." Implant Dent 6(2): 93-101.
- 8. Gerhardt L and Boccaccini A (2010). "Bioactive glass and glass ceramic scaffolds for bone tissue engineering." Materials 3: 3867-910.
- 9. Gjorgievska E and Nicholson JW (2011). "Prevention of enamel demineralization after tooth bleaching by bioactive glass incorporated into toothpaste." Aust Dent J 56(2): 193-200.
- Gosain AK (2004). "Bioactive glass for bone replacement in craniomaxillofacial reconstruction." Plast Reconstr Surg 114(2): 590-3.
- 11. Goudouri O (2011). "Dental ceramics/bioactive glass composites: characterization and

mechanical properties investigation." Bioceramics Develop Appl 1: 1-4.

- Kakodkar G, Lavania A and Ataide Ide N (2013). "An In vitro SEM Study on the Effect of Bleaching Gel Enriched with NovaMin on Whitening of Teeth and Dentinal Tubule Occlusion." J Clin Diagn Res 7(12): 3032-5.
- 13. Kaur G, Pandey OP, Singh K, Homa D, Scott B and Pickrell G (2014). "A review of bioactive glasses: Their structure, properties, fabrication and apatite formation." J Biomed Mater Res A 102(1): 254-74.
- 14. Khvostenko D, Hilton TJ, Ferracane JL, Mitchell JC and Kruzic JJ (2016). "Bioactive glass fillers reduce bacterial penetration into marginal gaps for composite restorations." Dent Mater 32(1): 73-81.
- 15. Khvostenko D, Mitchell JC, Hilton TJ, Ferracane JL and Kruzic JJ (2013). "Mechanical performance of novel bioactive glass containing dental restorative composites." Dent Mater 29(11): 1139-48.
- Kobayashi M, Saito H, Mase T, Sasaki T, Wang W, Tanaka Y, Nakamura M, Nagai A and Yamashita K (2010). "Polarization of hybridized calcium phosphoaluminosilicates with 45S5-type bioglasses." Biomed Mater 5(2): 25001.
- Kumar A, Singh S, Thumar G and Mengji A (2015). "Bioactive Glass Nanoparticles (NovaMin®) for Applications in Dentistry." IOSR-JDMS 14(8): 30-35.
- Larmas E, Sewon L, Luostarinen T, Kangasniemi I and Yli-Urpo A (1995). Bioactive glass in periodontal defects. Initial clinical findings of soft tissue and osseous repair. Biometrics. J. Wilson, L. Hench and D. Greenspan. Oxford, Elsevier Science. 8: 279-284.
- Linde J and Berglundh (1998). The peri- implant mucosa. Clinical Periodontology and Implant Dentistry. J. Linde and N. Lang. Munksgaard, Copenhagen: 862-872.
- 20. Lindfors NC (2009). "Treatment of a recurrent aneurysmal bone cyst with bioactive glass in a child allows for good bone remodelling and growth." Bone 45(2): 398-400.
- 21. Lindfors NC, Koski I, Heikkila JT, Mattila K and Aho AJ (2010). "A prospective randomized 14-year follow-up study of bioactive glass and autogenous bone as bone graft substitutes in benign bone tumors." J Biomed Mater Res B Appl Biomater 94(1): 157-64.
- 22. Lovelace TB, Mellonig JT, Meffert RM, Jones AA, Nummikoski PV and Cochran DL (1998). "Clinical evaluation of bioactive glass in the

treatment of periodontal osseous defects in humans." J Periodontol 69(9): 1027-35.

- Madan N, Sharma V, Pardal D and Madan N (2011). "Tooth remineralization using bio-active glass - A novel approach." J Acad Adv Dent Res 2: 45-9.
- 24. Margonar R, Queiroz T, Luvizuto E, Marcantonio E, Lia R, Holzhausen M and Marcantonio-Júnior É (2012). "Bioactive glass for alveolar ridge augmentation." J Craniofac Surg 23: e220-222.
- 25. Melek ET, Zheng K and Boccaccini A, Int. J. Appl. Glass Sci. 2013: 1–13. (2013). "Novel bioactive glasses in medical applications." 1-13.
- 26. Mengel R, Schreiber D and Flores-de-Jacoby L (2006). "Bioabsorbable membrane and bioactive glass in the treatment of intrabony defects in patients with generalized aggressive periodontitis: results of a 5-year clinical and radiological study." J Periodontol 77(10): 1781-7.
- Merkx MA, Maltha JC and Stoelinga PJ (2003). "Assessment of the value of anorganic bone additives in sinus floor augmentation: a review of clinical reports." Int J Oral Maxillofac Surg 32(1): 1-6.
- Milleman JL, Milleman KR, Clark CE, Mongiello KA, Simonton TC and Proskin HM (2012). "NUPRO sensodyne prophylaxis paste with NovaMin for the treatment of dentin hypersensitivity: a 4-week clinical study." Am J Dent 25(5): 262-8.
- 29. Mukai Y and ten Cate JM (2002). "Remineralization of advanced root dentin lesions in vitro." Caries Res 36(4): 275-80.
- Ong MM, Eber RM, Korsnes MI, MacNeil RL, Glickman GN, Shyr Y and Wang HL (1998). "Evaluation of a bioactive glass alloplast in treating periodontal intrabony defects." J Periodontol 69(12): 1346-54.
- Orchardson R, Gangarosa LP, Sr., Holland GR, Pashley DH, Trowbridge HO, Ashley FP, Kleinberg I and Zappa U (1994). "Dentine hypersensitivity-into the 21st century." Arch Oral Biol 39 Suppl: 113S-119S.
- Pantchev A, Nohlert E and Tegelberg A (2009). "Endodontic surgery with and without inserts of bioactive glass PerioGlas--a clinical and radiographic follow-up." Oral Maxillofac Surg 13(1): 21-6.
- Peltola MJ, Aitasalo KM, Aho AJ, Tirri T and Suonpaa JT (2008). "Long-term microscopic and tissue analytical findings for 2 frontal sinus obliteration materials." J Oral Maxillofac Surg 66(8): 1699-707.

- 34. Profeta A and Prucher G (2015). "Bioactiveglass in periodontal surgery and implant dentistry." Dent Mater J 34(5): 559-571.
- Sarin S and Rekhi A, (2016). "Bioactive glass: A potential next generation biomaterial." SRM J Res Dent Sci 7: 27-32.
- 36. Schepers E, de Clercq M, Ducheyne P and Kempeneers R (1991). "Bioactive glass particulate material as a filler for bone lesions." J Oral Rehabil 18(5): 439-52.
- 37. Schepers EJ and Ducheyne P (1997). "Bioactive glass particles of narrow size range for the treatment of oral bone defects: a 1-24 month experiment with several materials and particle sizes and size ranges." J Oral Rehabil 24(3): 171-81.
- 38. Schepers EJ, Ducheyne P, Barbier L and Schepers S (1993). "Bioactive glass particles of narrow size range: a new material for the repair of bone defects." Implant Dent 2(3): 151-6.
- 39. Shapoff CA, Alexander DC and Clark AE (1997). "Clinical use of a bioactive glass particulate in the treatment of human osseous defects." Compend Contin Educ Dent 18(4): 352-4, 356, 358 passim.
- 40. Stavropoulos A, Sima C, Sima A, Nyengaard J, Karring T and Sculean A (2012). "Histological evaluation of healing after transalveolar maxillary sinus augmentation with bioglass and autogenous bone." Clin Oral Implants Res 23(1): 125-31.
- Subbaiah R and Thomas B (2011). "Efficacy of a bioactive alloplast, in the treatment of human periodontal osseous defects-a clinical study." Med Oral Patol Oral Cir Bucal 16(2): e239-44.
- 42. Suominen E and Kinnunen J (1996). "Bioactive glass granules and plates in the reconstruction of defects of the facial bones." Scand J Plast Reconstr Surg Hand Surg 30(4): 281-9.
- 43. Tadjoedin ES, de Lange GL, Holzmann PJ, Kulper L and Burger EH (2000). "Histological observations on biopsies harvested following sinus floor elevation using a bioactive glass material of narrow size range." Clin Oral Implants Res 11(4): 334-44.
- 44. Tai BJ, Bian Z, Jiang H, Greenspan DC, Zhong J, Clark AE and Du MQ (2006). "Anti-gingivitis effect of a dentifrice containing bioactive glass (NovaMin) particulate." J Clin Periodontol 33(2): 86-91.
- 45. Talreja PS, Gayathri GV and Mehta DS (2013). "Treatment of an early failing implant by guided bone regeneration using resorbable collagen membrane and bioactive glass." J Indian Soc Periodontol 17(1): 131-6.

- 46. Taubock TT, Zehnder M, Schweizer T, Stark WJ, Attin T and Mohn D (2014).
 "Functionalizing a dentin bonding resin to become bioactive." Dent Mater 30(8): 868-75.
- 47. Vollenweider M, Brunner TJ, Knecht S, Grass RN, Zehnder M, Imfeld T and Stark WJ (2007). "Remineralization of human dentin using ultrafine bioactive glass particles." Acta Biomater 3(6): 936-43.
- 48. Waltimo T, Brunner TJ, Vollenweider M, Stark WJ and Zehnder M (2007). "Antimicrobial effect of nanometric bioactive glass 45S5." J Dent Res 86(8): 754-7.
- Wilson J, Clark A, Dou E, Crier J, Smith W and Summit J (1994). Clinical applications of bioglass implants. Bioceramics. O. Andersson

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and R. Happonen. Cambridge, Butterworth-Heinemann. 7: 415-422.

- Wilson J, Pigott GH, Schoen FJ and Hench LL (1981). "Toxicology and biocompatibility of bioglasses." J Biomed Mater Res 15(6): 805-17.
- 51. Wu C, Xia L, Han P, Mao L, Wang J, Zhai D, Fang B, Chang J and Xiao Y (2016). "Europium-Containing Mesoporous Bioactive Glass Scaffolds for Stimulating in Vitro and in Vivo Osteogenesis." ACS Appl Mater Interfaces 8(18): 11342-54.
- 52. Zehnder M, Soderling E, Salonen J and Waltimo T (2004). "Preliminary evaluation of bioactive glass S53P4 as an endodontic medication in vitro." J Endod 30(4): 220-4.