# Bioactive glass technology in preventive dentistry

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he fundamental principles of preventive dentistry are the preservation of tooth structure and the repair of lesions to minimise the risk of progression of oral disease. The management of the dynamic equilibrium of demineralisation/remineralisation of tooth enamel is key to this process.

Acidic attack on dental tissue is most often caused by bacterial sugar conversion (caries), or by direct dissolution (stomach acid reflux or consumption of acidic foods and drinks). This damage must be mitigated through the protection of the tooth surfaces, neutralisation of oral fluids, and replacement of lost tooth minerals, all tasks where bioactive glass technology can offer tremendous benefits to preventive dentistry. Oral healthcare and dental materials incorporating bioactive glass technology are projected to have a long-term future in oral healthcare.<sup>1</sup>

#### **BIOACTIVE GLASS TECHNOLOGY**

Bioactive glasses are derived from the family of calcium phosphosilicate materials that are degradable in body fluids such as saliva and blood. They act as a vehicle for delivering ions beneficial for both healing and remineralisation. Bioactive glasses were first developed by Hench<sup>2</sup> and his colleagues at the University of Florida in 1969 for the purpose of bone augmentation in orthopaedic surgery. Their use in oral healthcare did not emerge until several decades later with the introduction of Novamin<sup>3</sup>, the active ingredient in Sensodyne Repair and Protect (GlaxoSmithKline Inc., Mississauga ON). This toothpaste additive continues to employ the same glass formulation specifically developed by Hench for bone augmentation.

Whilst reviewing the properties of bioactive glass materials in the early 2000s, Professor Robert Hill, Queen Mary University, London, UK, realised that this original bioactive glass could be optimised specifically for oral healthcare application by:

- 1. Including fluoride<sup>4</sup> within the glass structure to provide controlled, longer-term release.
- 2. Dramatically increasing the phosphate content<sup>5</sup>.
- 3. Reducing the particle size.

Over the subsequent decade, Hill and his coworkers developed fluoro-calcium phosphosilicate<sup>6,7</sup> bioactive glass compositions (fluoride bioactive glass, commonly referred to as FBAG), that are featured in BioMin F toothpaste, and subsequently explored their formulation in dental materials such as remineralising composites, varnishes and cements, innovations that are currently in pre-market development.8

### **TOOTHPASTE APPLICATION**

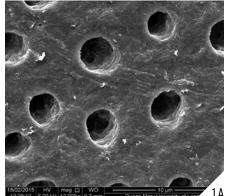
Fluoride bioactive glasses (FBAG) are activated through contact with water. The hydrogen ion of water replaces the calcium ion of the bioactive glass, increasing the pH; this has a neutralising effect on the local microenvironment. It is possible to specifically design the structure of the silicate glass to control the rate of this dissolution. Under acidic conditions, where there is an increased hydrogen ion concentration, this reaction proceeds more rapidly. This dissolution results in the controlled release of calcium, phosphate and fluoride ions which, in turn, combine to form fluorapatite and then crystalise on the tooth surface, particularly at nucleation sites such as those found in the peri-tubular tissues of open dentine tubules.

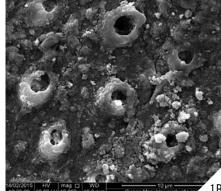
When formulated into toothpastes, FBAG is very effective at occluding these open dentinal tubules, thereby providing increased resistance to subsequent acidic attack. (Fig. 1A,B,C) An adhesive system is included in the toothpaste formulation such that the bioactive glass particles bond to the tooth surface, similarly to glass ionomer cements, allowing the active ingredient to remain in situ for many hours after brushing. The benefits of FBAGs in toothpaste have been validated in recent clinical studies9 and also confirmed by many dental clinicians who have prescribed BioMin F (Fig. 2) to their patients over the past 5 years. In a similar manner, the crystal deposition may also occur on early-stage enamel lesions as reported in scientific studies, 10 and may aid the remineralisation process.

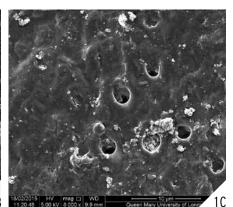
Over the past 60 years, fluoride has become a mainstay of dental public health, incorporated into



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1A. Open Dentine tubules prior to brushing with BioMin. 1B. After brushing with BioMin F. 1C. After acid attack. 2. BioMin F tubes.

toothpastes, varnishes and other treatments, and is added to the community water supply in many regions. The positive effects of fluoride include decreasing caries levels in children, particularly in economically deprived areas.<sup>11</sup> The advent of fluoride treatment has transformed preventive care since its introduction in the late 1940s.

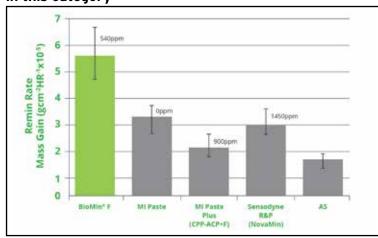
Fluoride is very effective, but its mode of action remains a subject of debate. It is known, however, that one of its main functions is the acceleration of apatite formation, creating the more stable and acidresistant fluorapatite on tooth surfaces. 12 It is also believed that bacteria are less able to adhere to fluorapatite than non-fluoride treated enamel.<sup>13</sup> Unfortunately, concerns have been raised regarding fluoride's potential toxicity;14 however, these allegations are often made without taking into full consideration the relatively low concentrations applied during toothbrush use.

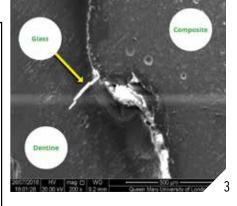
The debate on fluoride concentration continues further. Many clinicians are of the opinion that the more that is applied, the greater the preventive effect. This is not necessarily true (Table 1) as indicated in the 1990s by Ten Cate,14 who stated "For treatments to be effective longer than the brushing and salivary clearance, fluoride needs to be deposited and slowly released." This was further supported by Mohammed et al.15 researching the effect of various fluoride ion concentrations on demineralised enamel by FMAS-NMR<sup>19</sup>. Their results indicated that in (F) solutions at 45 ppm and below fluoride-substituted apatite formation predominates. In (F) solutions above 45 ppm, calcium fluoride forms in increasing proportions. Further increases in fluoride concentration cause no further reduction in demineralization but do increase the proportion of calcium fluoride formation.

There are references in the dental literature claiming calcium fluoride to be a reservoir which can release free fluoride.<sup>16</sup> However, because of the very high insolubility of CaF, in water (0.015 g/L @18°C), which increases only slightly under acidic conditions, this theory seems rather unlikely. Hence, the remineralising effect of fluoride is optimised at relatively low concentration. The specific optimal level is yet to be precisely determined, but Hill has proposed that this value is between 5-45ppm. This concept has been supported by Farooq et al in comparing the remineralisation effects of FBAG and standard 1450ppm sodium fluoride toothpastes. They demonstrated that the FBAG toothpaste provides increased surface micro-hardness, reduced surface roughness, and greater volume gain.17

Sodium fluoride is commonly featured in standard fluoride toothpastes, although other fluorides are also utilised. In all

## Remineralization rates of BioMin vs other products in this category





3. Marginal gap between composite material and dentine filled with Fluorapatite.

Table 1.

cases, the fluoride salts are soluble in water and hence the fluoride concentration diminishes exponentially in the mouth due to salivary dilution after brushing, and even more rapidly if the patient rinses their mouth immediately afterward application.

The half-life of fluoride concentration in the mouth at normal salivary levels is 5 minutes; thus, the length of time that the intraoral fluoride concentration remains in the optimal 5-45ppm range is relatively short. On the other hand, the FBAG toothpaste offers a controlled and sustained fluoride release mode over time. This makes it possible to provide high level protection to dental surfaces using much lower fluoride levels than those found in standard soluble fluoride (sodium monofluorophosphate or amino fluoride) toothpastes. FBAG dentifrices offer a viable alternative for the more efficient delivery of fluoride than what is found in traditional soluble fluoride toothpastes.

#### **RESIN RESTORATIVE MATERIALS**

Many experts agree that the life expectancy of a posterior composite resin restoration is between 5 and 7 years. This relatively limited longevity is largely due to composite shrinkage on setting creating marginal gaps that give rise to leakage and secondary caries. 18 This durability is inadequate as the profession moves away from mercury/silver amalgam which can last significantly longer but has its own drawbacks. By including FBAGs into dental composite formulations, substituting a high percentage of their inert glass content, researchers have shown it is possible to produce a composite resin capable not only of filling a prepared cavity but also of re-mineralizing the dental structures around hard carious lesions that can now be restored after minimally invasive caries removal.<sup>19</sup>

The inclusion of bioactive glass in dental composites is believed to inhibit bacterial decay and promote remineralisation.<sup>20</sup> Kuzic (Oregon State University) states: "The bacteria in the mouth which cause cavities don't seem to like this type of glass and are less likely to colonise on fillings which incorporate it." They are also believed to reduce enzyme-mediated degradation and encourage the remineralisation of demineralised dentin.<sup>21</sup>

The inclusion of FBAGs into dental composite materials creates an apatitelike phase in the marginal gap, (Fig. 3) providing a more effective marginal seal, and preventing acid attack by raising the local pH, thereby reducing the likelihood of composite restorative failure. Bioactive dental composites may improve the longevity and clinical outcomes of composite restorations.<sup>18</sup> There is ongoing research to assess the inclusion of additional ions (such as zinc and magnesium) into the FBAG to further enhance the properties of the restorative material.<sup>22</sup>

There appears to be a bright future for including fluoride bioactive glasses (FBAG) into a wide range of oral healthcare and dental restorative materials. Fluoride bioactive glasses are likely to become mainstream tools for the dental clinician in the development of more preventive approaches to patient care and improved quality of life.

Oral Health welcomes this original article.

## REFERENCES

1. Goldstep F. Proactive intervention dentistry: a model for oral care through life. Compend Contin Educ Dent Suppl 2012;33(6):394-6, 8-402.

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#### REFERENCES

- 2. Hench, L.L. Biomaterials: A forecast for the future. Biomaterials 1998, 19, 1419-1423. Hench L. The story of bioglass®. J Mater Sci Mater Med 2006;17:967-78.
- 3. Greenspan D. NovaMin and tooth sensitivity- an overview. J Clin Dent 2010;21:61-5.
- 4. Brauer D, Karpukhina N, Law R, Hill R. Structure of fluoride-containing bioactive glasses. J Mater Chem 2009;19:5629-36.
- 5. Mneimne M, Hill RG, Bushby AJ, Brauer DS. High phosphate content significantly increases apatite formation of fluoride-containing bioactive glasses. Acta Biomater 2011;7(4):1827-34.
- 6. Brauer DS, Karpukhina N, O'Donnell MD, Law RV, Hill RG. Fluoride-containing bioactive glasses: Effect of glass design and structure on degradation, pH and apatite formation in simulated body fluid. Acta Biomater 2010;6:3275-82.
- 7. Lynch, E.; Brauer, D.S.; Karpukhina, N.; Gillam, D.G.; Hill, R.G.

- Multi-component bioactive glasses of varying fluoride content for treating dentin hypersensitivity. Dent. Mater. 2012, 28, 168-178.
- 8. Tiskaya M, Shahid S, Gillam D, Hill R The use of bioactive glass (BAG) in dental composites: A critical review Dental Materials 2021 v37 is2
- 9. Patel, V.R.; Shettar, L.; Thakur, S.; Gillam, D.; Kamala, D.N. A randomised clinical trial on the efficacy of 5% fluorocalcium phosphosilicatecontaining novel bioactive glass toothpaste. J. Oral Rehabil. 2019, 46, 1121-1126.
- 10. Bakry, A.S.; Abbassy, M.A.; Alharkan, H.F.; Basuhail, S.; Al-Ghamdi, K.; Hill, R. A Novel Fluoride Containing Bioactive Glass Paste is Capable of Re-Mineralizing Early Caries Lesions. Materials 2018, 11, 1636.
- 11. Iheozor-Ejiofor et al; Water fluoridation for the prevention of dental caries Cochrane Database Syst Rev. 2015 Jun; 2015(6): CD010856.
- 12. Aoba T; The effect of fluoride on apatite structure and growth Oral

- Biol Med1997;8(2):136-53.
- 13. Loskil P et al Reduced Adhesion of Oral Bacteria on Hydroxyapatite by Fluoride Treatment Langmuir 2013, 29, 18, 5528-5533.
- 14. Gravitz L The fluoride wars rage on Nature Outlook 21 October 2021.
- 15. Ten Cate J; In vitro Studies on the Effects of Fluoride on De- and Remineralization JDR Feb1990.
- 16. N.R.Mohammed, N.W Kent,
- R.J.M Lynch, N. Karpukhina, R. Hill, P. Anderson. Effects of Fluoride on in vitro Fnamel Demineralization Analyzed by 19F MAS-NMR. Caries Res 2013;47:421-428.
- 17. Rolla G On the role of calcium fluoride in the cariostatic mechanism of fluoride Odontol Scand 1988 Dec:46(6):341-5.
- 18. Faroog et al Enamel Remineralization Competence of a Novel Fluoride-Incorporated Bioactive Glass Toothpaste - A Surface Micro-Hardness, Profilometric, and Micro-Computed Tomographic Analysis. Tomography 2021,7 752-768.

19. Opdam NJ, van de Sande FH,

- Bronkhorst E, Cenci MS, Bottenberg P, Pallesen U, Gaengler P, Lindberg A, Huysmans MC, van Dijken JW. Longevity of posterior composite restorations: a systematic review and meta-analysis J Dent Res. 2014 Oct:93(10):943-9.
- 20. Jefferies S, Fuller A, Boston D. Preliminary evidence that bioactive cements occlude artificial marginal gaps: bioactive cements occlude artificial marginal gaps. J Esthet Restor Dent 2015;27.
- 21. Khvostenko D https://ir.library. oregonstate.edu/concern/graduate\_ thesis\_or\_dissertations/dv13zw59j.
- 22. A Tezvergil-Mutluay, Seseogullari-<u>Dirihan</u> R, <u>V P Feitosa</u> V, <u>G Cama</u> G, DS Brauer D, Sauro S; Effects of Composites Containing Bioactive Glasses on Demineralized Dentin. Dent Res 2017 Aug;96(9):999-1005
- 23. Ali, S.; Farooq, I.; Igbal, K. A review of the effect of various ions on the properties and the clinical a pplications of novel bioactive glasses in medicine and dentistry. Saudi. Dent. J. 2014, 26, 1-5.

There are 2 #13 references which pushed the numbers to 23 (not 22).