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THE SCIENCE BEHIND BIOACTIVE DENTAL ADHESIVE SYSTEMS: BENEFITS AND APPLICATIONS

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ABSTRACT

The integration of bioactive dental adhesive systems represents a significant advancement in restorative dentistry, offering enhanced therapeutic benefits beyond traditional adhesive materials. This review explores the underlying science, benefits, and clinical applications of bioactive dental adhesives. Bioactive adhesives are designed to interact favorably with biological tissues, promoting remineralization, reducing bacterial colonization, and improving the longevity of dental restorations. These adhesives incorporate various bioactive compounds, such as calcium phosphate, fluoride, and antimicrobial agents, which contribute to the regeneration of dentin and the prevention of secondary caries. The application of bioactive dental adhesives spans a wide range of restorative procedures, including direct and indirect restorations, sealants, and cavity liners. Clinical studies have demonstrated improved outcomes in terms of bond strength, durability, and patient satisfaction. By leveraging the bioactive properties of these advanced materials, dental practitioners can provide more effective and sustainable treatments, ultimately leading to better oral health and patient care. This review underscores the potential of bioactive dental adhesive systems to revolutionize dental practice and highlights the need for further research to optimize their formulations and applications.

KEYWORDS: Bioactive dental adhesives, Restorative dentistry, Remineralization, Dental materials, Dental restorations.

INTRODUCTION

Modern restorative dentistry prioritizes the thorough removal of carious tissue while maintaining healthy tooth structure to improve mechanical retention ^[1]. Dental adhesives have been a cornerstone of restorative dentistry for decades, facilitating the bonding of restorative materials to tooth structures. Dental bonding systems act as an intermediary substance that adheres restorative materials to hard dental tissue, enhancing retention, marginal sealing, and resistance at the tooth-restoration interface ^[2]. However, polymerization stress is the most significant obstacle for composite materials and a major factor in the clinical failure of current adhesion methods ^[3]. During the curing process of composite resins, polymerization shrinkage occurs as monomers combine to form polymers, leading to a reduced

overall volume. This shrinkage can create internal contraction tensions and stresses at the edges of the restoration ^[4]. The success of restorative procedures hinges on the bonding effectiveness of adhesive materials. As a result, dental adhesives have become crucial in restorative dentistry and are among the most important biomaterials in Health Sciences ^[3].

Efforts to streamline complex multistep dental adhesives and enhance their user-friendliness have resulted in the creation of "universal" adhesives. These adhesives have gained popularity in dentistry due to their reduced toxicity, versatility in both self-etching and etch-and-rinse procedures, flexibility, and simplified application steps ^[5]. In 2012, the inaugural universal adhesive entered the market in Japan under the name Scotchbond Universal Adhesive, produced by 3M Oral Care in St. Paul, Minnesota. Subsequently, universal adhesives were introduced with capabilities to be utilized alongside resin luting cements ^[6], various substrates without surface treatment ^[7], reduced treatment durations ^[8], or differing levels of surface moisture on enamel and dentin surfaces ^[9]. Despite their remarkable versatility, some reports indicate that several universal adhesives exhibit lower bond strength compared to self-etching two-step adhesives. This discrepancy is especially noticeable in dentin when used in self-etch mode ^[10] and in enamel with etch-and-rinse and self-etch modes ^[11]. Nevertheless, the clinical importance of universal adhesives has grown ^[12], underscoring the necessity for further research to enhance their application.

Bioactive dental adhesives represent a significant innovation in the field of dentistry. Unlike traditional Bioactive dental adhesives, bioactive adhesives are designed to interact positively with biological tissues, offering therapeutic benefits that extend beyond mere adhesion. These advanced materials are formulated with bioactive compounds, such as calcium phosphate, fluoride, and antimicrobial agents, which contribute to the regeneration of dentin, promote remineralization, and help prevent secondary caries ^[13]. Numerous investigations have explored the impact of incorporating nanoparticles like silver, calcium phosphate, zinc oxide, and titanium dioxide into different restorative materials, including glass ionomer cements, resin-based materials, and dental adhesives ^[14-16]. The addition of antibacterial components has been shown to have positive effects on interfacial bonding ^[17], reduce bacterial viability ^[18], inhibit matrix metalloproteinase activity, and importantly, mitigate the impact of factors that initiate chemical degradation of dental adhesives ^[19, 20]. The bioactive properties of these adhesives support the natural healing processes of the tooth, thereby enhancing the overall health and longevity of dental restorations.

The purpose of this review is to provide a comprehensive overview of bioactive dental adhesive systems, highlighting their underlying science, benefits, and clinical applications. By examining the composition and mechanisms of these adhesives, this review aims to elucidate how they contribute to improved dental health and restoration outcomes. Additionally, the review will explore the various clinical applications of bioactive dental adhesives, from direct restorations to sealants and cavity liners, and discuss the evidence from clinical studies that supports their efficacy. Finally, this review will address the potential impact of bioactive dental adhesives on dental practice and oral health, and outline the future directions for research and development in this promising area.

Underlying Science of Bioactive Dental Adhesive Systems

Dentin demineralization typically occurs due to organic acids from oral bacteria or acid etching during adhesive procedures, leading to the deterioration of the dentin matrix ^[21]. Exposed demineralized collagen fibers can be degraded by endogenous matrix metalloproteinases (MMPs) activated by etching

^[22]. While it's ideal to shield the exposed collagen network with dentin adhesive to prevent collagen degradation post-bonding, complete encapsulation of all demineralized collagen fibers in the hybrid layer is impractical ^[23]. To remineralize dentin and prevent collagen degradation, various agents such as fluoride, sodium trimetaphosphate, zinc oxide, titanium nanoparticles, chlorhexidine, and benzalkonium chloride have been integrated into dentin adhesives ^[24-27]. Additionally, biomaterials like tricalcium phosphate and tricalcium silicates have been proposed as additives to restorative materials to hinder the demineralization cascade and subsequent collagen degradation ^[28]. Dentin permeability, as noted by Pashley et al. ^[29], influences the adhesive layer and may diminish bond strength. In the totaletch approach, phosphoric acid etchant removes hydroxyapatite crystals from dentin, heightening the permeability of demineralized dentin. Consequently, total-etch dentin adhesives may exhibit higher permeability than self-etch dentin adhesives ^[30].

A robust connection between the adhesive and dentin greatly prolongs the restoration's lifespan, and its efficacy is intricately linked to the varied characteristics of dentin adhesive. Bioactive dental adhesives are composed of several key components that together provide the necessary mechanical, chemical, and biological properties required for effective dental restorations. The primary constituents include a polymer matrix, bioactive compounds, fillers, and additional modifiers ^[31]. The polymer matrix forms the bulk of the adhesive and is responsible for its mechanical properties and bonding capability. Common polymers used include Bisphenol A-Glycidyl Methacrylate (Bis-GMA), Urethane Dimethacrylate (UDMA), and Triethylene Glycol Dimethacrylate (TEGDMA) ^[32]. Bioactive Compounds impart biological activity to the adhesive, promoting remineralization and providing antibacterial effects. Key bioactive compounds include ^[31, 33-37]: Calcium Phosphate (such as Hydroxyapatite, Tricalcium Phosphate (TCP), Amorphous Calcium Phosphate), Fluoride (Such as Sodium Fluoride (NaF2), Fluoride-Releasing Glass Fillers), and Antimicrobial Agents (Such as as Chlorhexidine, Quaternary Ammonium Compounds (e.g., QAS), Silver Nanoparticles). Additional modifiers are incorporated to enhance the properties and performance of the adhesive ^[31]. Bioactive dental adhesives are complex formulations designed to provide not only strong and durable bonds to dental tissues but also additional therapeutic benefits. The combination of a polymer matrix, bioactive compounds, fillers, photoinitiators, and other modifiers allows these adhesives to enhance the restoration's longevity, promote remineralization, and offer antibacterial protection, ultimately contributing to better oral health outcomes.

Benefits of Bioactive Dental Adhesive Systems

Bioactive dental adhesive systems are a significant advancement in dental materials technology, providing numerous benefits over traditional adhesives. They enhance bonding and longevity of restorations while promoting remineralization of tooth structures. These adhesives possess antibacterial properties, reducing the risk of secondary caries. Their biocompatibility and ability to stimulate tissue healing make them ideal for various dental applications.

Promotion of remineralization

Bioactive dental adhesive systems play a crucial role in promoting the remineralization of tooth structures, offering numerous benefits that extend the lifespan and health of dental restorations. These adhesives are designed to release beneficial ions, such as calcium and phosphate, which integrate into the tooth structure ^[38]. This process not only rebuilds enamel but also strengthens it, enhancing the tooth's natural defenses against decay ^[39]. By actively contributing to the remineralization process,

bioactive adhesives provide a protective barrier that traditional adhesives lack. One of the primary benefits of promoting remineralization is the strengthening of tooth enamel. Enamel is the outermost layer of the tooth and is crucial for protecting against cavities and other forms of decay ^[40]. Bioactive adhesives release ions that help rebuild and fortify this essential layer, making teeth more resistant to acid attacks from bacteria and food ^[41, 42]. This fortification is particularly important in areas that have been demineralized due to poor oral hygiene or acidic diets, helping to restore the tooth's natural hardness and durability ^[43]. Another significant advantage is the reduction of tooth sensitivity. Sensitivity often occurs when enamel is worn away, exposing the underlying dentin and its microscopic tubules that lead to the tooth's nerve ^[44]. Bioactive adhesives promote the remineralization of these areas, effectively sealing the dentinal tubules and reducing sensitivity ^[45]. This not only improves patient comfort but also enhances the overall success of dental restorations by ensuring that they are both functional and comfortable.

Promoting remineralization also helps prevent secondary caries, which are cavities that form around the margins of dental restorations. Secondary caries is a common cause of restoration failure and often require additional dental work to address ^[46]. By continuously releasing remineralizing ions, bioactive adhesives help repair early demineralized areas and prevent the progression of decay. This proactive approach to oral health can significantly extend the life of dental restorations and reduce the need for future dental interventions. Lastly, the long-term durability and overall oral health benefits of bioactive adhesives cannot be overstated. By continuously supporting the remineralization of tooth structures, these adhesives help maintain the integrity of both the tooth and the restoration over time. Patients benefit from fewer dental visits and a lower risk of complications, while dentists can achieve more reliable and long-lasting results. In essence, bioactive dental adhesive systems represent a significant advancement in dental care, offering a comprehensive solution that supports both the immediate and long-term health of teeth.

Reduction of bacterial colonization and prevention of secondary caries

Bioactive dental adhesive systems offer substantial benefits in reducing bacterial colonization and preventing secondary caries, making them a valuable advancement in dental technology. Traditional adhesives often lack the ability to combat bacterial growth, which can lead to the development of secondary caries around dental restorations ^[47]. Bioactive adhesives, however, are designed to release antibacterial agents that inhibit bacterial activity and growth ^[48]. This property significantly reduces the risk of new cavities forming around existing restorations, ensuring longer-lasting dental work and better oral health.

One of the key advantages of bioactive adhesives is their ability to create an unfavorable environment for bacteria. These adhesives can release ions, such as fluoride, calcium, and phosphate, which are detrimental to bacterial survival ^[49]. Fluoride, for example, is well-known for its antibacterial properties and its ability to enhance remineralization. By incorporating fluoride-releasing bioactive adhesives, dentists can offer additional protection against bacteria, thereby reducing the likelihood of secondary caries and maintaining the integrity of the restoration ^[50].

Another important benefit is the improved sealing capability of bioactive adhesives. Effective marginal sealing is crucial for preventing bacterial infiltration and microleakage, which are common causes of secondary caries. Bioactive adhesives form a stronger and more durable bond with the tooth structure, minimizing gaps where bacteria can enter ^[51]. This enhanced sealing ability not only protects against

decay but also ensures that the restoration remains intact and functional over a longer period, reducing the need for replacement or repair ^[52]. In summary, the integration of bioactive dental adhesive systems into clinical practice represents a forward-thinking approach that prioritizes both prevention and treatment, ultimately fostering better long-term oral health.

Improved bond strength and durability of dental restorations

Bioactive dental adhesive systems represent a major advancement in dental technology, particularly in enhancing the bond strength and durability of dental restorations. Traditional adhesives often suffer from weaknesses over time due to degradation and wear, leading to restoration failures ^[53]. Bioactive adhesives, however, are formulated to create stronger and more resilient bonds with both the tooth structure and restorative materials ^[54]. This increased bond strength ensures that restorations remain securely in place, reducing the risk of dislodgement and the need for frequent repairs.

One of the primary benefits of improved bond strength in bioactive adhesives is their ability to withstand the mechanical stresses encountered in the oral environment ^[55]. Chewing, grinding, and other daily activities exert significant pressure on dental restorations. Bioactive adhesives enhance the mechanical properties of the bond, making it more resistant to these forces ^[56]. This resilience is crucial for maintaining the integrity of restorations such as fillings, crowns, and bridges, ensuring they remain functional and effective for longer periods [57]. The durability of bioactive dental adhesives is also significantly enhanced by their ability to release beneficial ions, such as calcium and phosphate, which support the ongoing remineralization of the tooth structure. This continuous release helps to repair and reinforce the tooth around the restoration, creating a more stable and durable interface ^[58]. By promoting a healthier and more robust tooth-restoration interface, bioactive adhesives help to prevent the marginal breakdown that can lead to restoration failure and secondary caries ^[59]. Another important aspect of improved bond strength and durability is the reduction of microleakage. Microleakage occurs when there are gaps or weak spots in the adhesive bond, allowing bacteria and oral fluids to penetrate and cause decay. Bioactive adhesives create a more complete and secure seal around restorations, effectively eliminating these vulnerabilities ^[60, 61]. This superior sealing capability not only enhances the longevity of the restoration but also contributes to overall oral health by preventing the onset of secondary caries. In summary, the improved bond strength and durability offered by bioactive dental adhesive systems represent a substantial benefit, ensuring that dental restorations are both long-lasting and effective, thereby promoting better oral health outcomes.

Enhanced patient outcomes and satisfaction

One of the key factors contributing to enhanced patient outcomes is the reduction in post-operative sensitivity. Bioactive adhesives effectively seal dentinal tubules, which helps prevent the discomfort often associated with dental restorations ^[62]. By promoting remineralization and maintaining a healthy tooth structure, these adhesives minimize the chances of sensitivity, ensuring a more comfortable recovery period for patients. This comfort level is crucial for patient satisfaction, as it reduces anxiety and improves the overall perception of dental procedures ^[63]. Enhanced aesthetic results also play a vital role in patient satisfaction ^[64]. Bioactive dental adhesives help maintain the natural appearance of restorations by resisting discoloration and ensuring a seamless integration with the surrounding tooth structure ^[65]. Patients appreciate the improved aesthetics, which can boost their self-esteem and confidence in their smile ^[66]. The long-lasting nature of these adhesives means that patients can enjoy these aesthetic benefits for an extended period, further enhancing their satisfaction with the treatment.

Lastly, the overall convenience and cost-effectiveness of bioactive dental adhesive systems contribute to improved patient outcomes. By reducing the need for frequent repairs and replacements, these adhesives save patients time and money ^[67]. Patients can rely on the durability and effectiveness of their restorations, knowing that they are investing in a long-term solution for their dental health. This reliability fosters a sense of trust and satisfaction with the dental care provider, as patients feel they are receiving high-quality, value-driven treatment. As a result, bioactive adhesives represent a valuable advancement in dental technology, offering both immediate and long-term benefits for patients and dental practitioners alike.

In summary, bioactive dental adhesive systems offer substantial benefits, including enhanced bonding, remineralization, antibacterial properties, and improved biocompatibility. These advantages make them an excellent choice for both patients and dental practitioners aiming for long-lasting and health-promoting dental restorations.

Clinical Applications of Bioactive Dental Adhesives

Bioactive dental adhesives have a wide range of clinical applications, transforming how dental restorations are performed and maintained. Their unique properties not only enhance the bond strength and durability of restorations but also promote oral health through remineralization and antibacterial action [68]. This versatility makes them an invaluable tool in various dental procedures, from routine fillings to complex restorations, ensuring better patient outcomes and longer-lasting results (Table 1).

Clinical	Description	Benefits	Examples	References
Application				
Caries Prevention	Preventing and treating	Reduces caries	Adhesives	[69, 70]
and Treatment	dental caries by halting	incidence, enhances	containing	
	demineralization and	remineralization of	fluoride, calcium	
	enhancing	enamel and dentin.	phosphate.	
	remineralization.			
Dentin	Reducing sensitivity in	Provides immediate and	d Adhesives with	[71, 72]
Hypersensitivity	dentin by occluding	long-lasting relief from	bioactive glass,	
	dentinal tubules and	sensitivity.	strontium.	
	forming a protective layer.			
Pulp Capping	Protecting the dental pulp	Promotes healing and	Calcium	[73, 74]
	after exposure or near-	formation of reparative	hydroxide-based	
	exposure during	dentin.	adhesives.	
	restorative procedures.			

Table 1 (Clinical applications of bioactive dental adhesives in modern dentistry.)

Orthodontic	Enhancing bonding	Improves bracket	Fluoride-releasing	[75, 76]
Applications	strength of brackets and	adhesion, minimizes	adhesives.	
	reducing white spot	enamel decalcification.		
	lesions around brackets.			
		Adding amorphous	Orthodontic	[77]
	It has desirable properties	tricalcium phosphate	composite	
	that are not easily	nanoparticles to an	containing silver	
	separated during the	orthodontic composite	nanoparticles	
	treatment but are easily	does not significantly	/orthodontic	
	removed after the	affect shear bond	composite	
	treatment.	strength, whereas silver	containing	
		nanoparticles reduce the	amorphous	
		shear bond strength of	tricalcium	
		the orthodontic	phosphate	
		composite.	nanoparticles.	
Periodontal	Enhancing periodontal	Supports regeneration of	Adhesives with	[78, 79]
Applications	tissue regeneration and	periodontal tissues,	growth factors,	
	promoting healing after	reduces inflammation.	antimicrobial	
	periodontal surgery.		agents.	
Crown and Bridge	Improving the bonding of	Enhances retention and	Resin-modified	[80, 81]
Cementation	crowns and bridges to	stability, reduces	glass ionomer	
	natural tooth structure.	microleakage.	cements.	
Restorative	Bonding restorative	Provides strong, durable	Universal	[82, 83]
Dentistry	materials to tooth	bond, reduces secondary	adhesives, self-	
	structure effectively and	caries.	etch adhesives.	
	durably.			
Endodontic	Sealing root canals and	Ensures a tight seal,	Bioactive root	[84, 85]
Applications	promoting periapical	reduces post-treatment	canal sealers.	
	healing after root canal	complications.		
	treatment.			

One primary application of bioactive dental adhesives is in restorative dentistry, particularly for direct restorations like composite fillings ^[86]. Traditional composite fillings can sometimes fail due to inadequate bonding and secondary caries ^[87]. Bioactive adhesives address these issues by forming a strong bond with both the tooth structure and the composite material, while releasing ions that promote remineralization and inhibit bacterial growth. This dual action ensures the restoration remains intact and functional for a longer period, reducing the need for frequent replacements ^[88, 89]. Indirect restorations, such as crowns, bridges, and veneers, also benefit significantly from bioactive dental adhesives ^[90]. These adhesives provide superior adhesion to both the tooth structure and the restoration greaterial, ensuring that restorations remain securely in place under the mechanical stresses of chewing and biting. The enhanced sealing properties of bioactive adhesives prevent microleakage, a common issue with indirect restorations that can lead to decay and restoration failure ^[32].

Bioactive dental adhesives are also highly beneficial in pediatric dentistry. Children are particularly susceptible to dental caries, and the use of bioactive adhesives in restorative treatments can help prevent the progression of decay ^[91]. The remineralization properties of these adhesives strengthen the remaining tooth structure and protect against future caries. Additionally, the antibacterial action of bioactive adhesives reduces the risk of infection, promoting better oral health for young patients and reducing the need for more invasive treatments later ^[92, 93]. In endodontic procedures, bioactive dental adhesives play a crucial role in ensuring the long-term success of root canal treatments. They can be used to seal the root canal space, preventing bacterial re-entry and promoting the healing of periapical tissues ^[95, 95]. The ability of bioactive adhesives to release calcium ions can stimulate the formation of a hard tissue barrier, enhancing the sealing of the root canal and improving the overall prognosis of endodontic treatments. This application helps in achieving better long-term outcomes and reducing the risk of treatment failure.

Periodontal therapy also benefits from the use of bioactive dental adhesives. During procedures such as root planing and scaling, bioactive adhesives can be used to seal exposed root surfaces and prevent bacterial colonization ^[96]. The antibacterial properties of these adhesives help to reduce the microbial load in periodontal pockets, promoting healing and reducing the risk of recurrence ^[97]. By supporting the regeneration of periodontal tissues, bioactive adhesives contribute to the overall success of periodontal treatments and improve patient outcomes.

Bioactive dental adhesives are particularly valuable in prosthodontics, where they can be used to improve the retention and stability of dentures and other prosthetic devices ^[98]. These adhesives form a strong bond with the prosthetic material and the underlying tissue, ensuring a secure fit. The release of remineralizing ions can help to maintain the health of the supporting tissues, reducing the risk of complications such as denture stomatitis ^[99]. This application enhances the comfort and functionality of prosthetic devices, improving the quality of life for patients. In cosmetic dentistry, bioactive dental adhesives are used to enhance the aesthetic outcomes of treatments such as bonding and veneers ^[100]. These adhesives provide a strong bond and a seamless interface between the tooth and the restoration, ensuring that the final result looks natural and is resistant to staining and discoloration ^[101, 102]. The ability of bioactive adhesives to promote remineralization and protect against decay ensures that cosmetic restorations remain beautiful and functional for an extended period.

It is important to use orthodontic composites with favorable properties, which are easily removed after the end of the treatment but not easily debonded during treatment. Nanoparticles have drawn attention for their antibacterial properties when added to composite resins. results of a study indicated that adding amorphous tricalcium phosphate nanoparticles to an orthodontic composite does not significantly decrease the shear bond strength. In contrast, silver nanoparticles reduce the shear bond strength of the orthodontic composite ^[77]. During orthodontic treatment, a reduction in shear bond strength can cause brackets to detach from the tooth surface, complicating the procedure ^[103]. Additionally, the inclusion of silver nanoparticles in the composite can lead to discoloration, which is aesthetically unpleasing ^[104]. In contrast, composites containing amorphous tricalcium phosphate nanoparticles maintain a shear bond strength similar to that of composites without nanoparticles. They can release phosphorus and calcium ions during treatment, aiding in the remineralization of decayed lesions ^[77]. Furthermore, these nanoparticles can raise the pH of the environment from 4 to 6.5, helping to neutralize bacterial acids and reduce decay ^[105].

Finally, bioactive dental adhesives are instrumental in preventive dentistry. They can be used in sealant applications to protect the occlusal surfaces of teeth from decay ^[106]. This preventive application is especially important for children and adolescents, helping to reduce the incidence of dental caries and promoting overall oral health from an early age ^[107]. In summary, bioactive dental adhesives have diverse and significant clinical applications across various branches of dentistry. Their ability to provide strong, durable bonds, promote remineralization, and prevent bacterial colonization makes them an essential tool for enhancing the success and longevity of dental treatments. From restorative and pediatric dentistry to endodontics, periodontics, prosthodontics, cosmetic, and preventive dentistry, bioactive adhesives offer numerous benefits that improve patient outcomes and satisfaction.

Challenges and Considerations

Bioactive dental adhesive systems leverage advanced materials science to provide significant benefits over traditional adhesives. They release ions such as calcium, phosphate, and fluoride, which promote remineralization and inhibit bacterial growth. However, the integration of these bioactive components poses several challenges. Ensuring a controlled and sustained release of these ions while maintaining the adhesive's mechanical properties is complex ^[108]. Researchers must balance the bioactivity with the need for a strong, durable bond, which requires sophisticated formulations and manufacturing processes.

One of the primary challenges is achieving a consistent and prolonged ion release without compromising the adhesive's structural integrity ^[109]. Bioactive adhesives must release ions at a rate that is effective for remineralization and antibacterial action, but this release should not weaken the adhesive over time ^[31, 110]. Variability in the oral environment, such as pH fluctuations and exposure to different food and drink substances, can affect the ion release rates ^[111]. Manufacturers need to develop formulations that are robust enough to perform consistently under these varying conditions.

Another significant consideration is the compatibility of bioactive adhesives with different restorative materials and tooth structures. The adhesive must form a strong bond with both the tooth and the restorative material, which can vary widely in composition and properties ^[112]. This requires extensive testing and optimization to ensure that the adhesive works effectively across a range of clinical scenarios. Additionally, the adhesive must be easy to handle and apply in a clinical setting, requiring formulations that balance bioactivity with practical usability for dental professionals.

Long-term clinical performance is also a crucial factor ^[113]. While bioactive dental adhesives have shown promising results in laboratory and short-term clinical studies, their long-term durability and effectiveness need continuous evaluation. Dentists and patients need to be confident that these

adhesives will maintain their beneficial properties over many years. This necessitates ongoing research and long-term clinical trials to monitor the performance of bioactive adhesives in real-world conditions. Understanding potential long-term issues, such as the stability of bioactive components and their interaction with oral tissues, is essential for ensuring the reliable performance of these advanced materials. In conclusion, while bioactive dental adhesive systems offer significant advantages in promoting oral health and enhancing the durability of dental restorations, they also present several challenges and considerations. Achieving a balance between bioactivity and mechanical strength, ensuring consistent ion release, compatibility with various materials, and long-term performance are key factors that need to be addressed. Ongoing research and development are critical to overcoming these challenges and fully realizing the potential of bioactive dental adhesives in clinical practice.

Future Directions and Research Needs

The evolution of bioactive dental adhesive systems is poised to reshape the landscape of restorative dentistry, offering promising avenues for future research and development. One crucial area of exploration lies in optimizing the bioactivity and longevity of these adhesives through advanced materials science and engineering ^[114]. Researchers aim to fine-tune the composition and structure of bioactive adhesives to achieve controlled and sustained release of therapeutic ions, such as calcium, phosphate, and fluoride. By harnessing nanotechnology and biomimetic principles, scientists seek to create bioactive adhesives that mimic the natural processes of enamel remineralization and antibacterial defense, resulting in more resilient and biocompatible restorations ^[115, 116].

Advancements in understanding the complex interactions between bioactive adhesives and the oral environment are essential for improving their clinical efficacy. Future research endeavors should focus on elucidating the effects of salivary constituents, pH fluctuations, and microbial biofilms on the performance of bioactive adhesives ^[117, 118]. By gaining insights into these dynamic interactions, researchers can develop strategies to enhance the stability and therapeutic efficacy of bioactive adhesives in diverse clinical scenarios, ensuring consistent and long-lasting benefits for patients.

In addition to enhancing bioactivity, future research should address the practical challenges associated with the clinical application of bioactive dental adhesive systems. Developing user-friendly formulations that are easy to handle and apply in a dental setting is crucial for widespread adoption. Furthermore, researchers should explore novel delivery mechanisms and application techniques to optimize the effectiveness of bioactive adhesives while minimizing procedural complexities and patient discomfort.

Longitudinal clinical studies with extended follow-up periods are paramount for assessing the longterm performance and durability of bioactive dental adhesive systems. By monitoring patients over several years, researchers can evaluate the stability of adhesive bonds, the incidence of secondary caries, and the overall success rates of restorations ^[119]. These studies will provide valuable insights into the real-world performance of bioactive adhesives and identify areas for further refinement and improvement.

Collaboration between multidisciplinary teams of researchers, clinicians, and industry partners is essential for advancing the science behind bioactive dental adhesive systems. By fostering partnerships and knowledge exchange, researchers can accelerate innovation and translate scientific discoveries into clinically impactful solutions. Moreover, initiatives aimed at standardizing testing protocols and establishing evidence-based guidelines will facilitate the evaluation and adoption of bioactive adhesives in routine clinical practice.

In summary, the future of bioactive dental adhesive systems holds immense promise, driven by ongoing research efforts to optimize their bioactivity, understand their interactions with the oral environment, address practical challenges, and evaluate their long-term clinical performance. Through interdisciplinary collaboration and a commitment to translational research, bioactive adhesives have the potential to revolutionize restorative dentistry, offering patients durable, biocompatible, and preventive solutions for maintaining optimal oral health.

CONCLUSION

Bioactive dental adhesive systems represent a groundbreaking advancement in restorative dentistry, offering multiple benefits surpassing traditional adhesive materials. These innovative adhesives provide robust bonding and durability actively promote remineralization and reduce bacterial colonization, thereby enhancing the longevity and effectiveness of dental restorations. By incorporating bioactive compounds such as calcium phosphate, fluoride, and antimicrobial agents, these adhesives support dentin regeneration and prevent secondary caries, leading to improved patient outcomes and satisfaction. The clinical applications of bioactive dental adhesives are extensive, encompassing direct and indirect restorations, sealants, and cavity liners. This versatility makes them a valuable tool in a wide range of dental procedures, contributing to more effective and sustainable oral health treatments. The integration of bioactive properties into dental adhesives signifies a shift towards materials that not only restore but also actively maintain and improve oral health.

The potential impact of bioactive dental adhesive systems on dental practice and oral health is profound. By providing materials that interact beneficially with biological tissues, dental practitioners can enhance the quality and longevity of restorations, ultimately leading to better patient care. The adoption of these advanced materials can lead to a reduction in recurrent dental issues, less invasive treatments, and improved overall oral health outcomes. The future of bioactive dental adhesives in restorative dentistry is promising. Continued research and development are essential to optimize their formulations and expand their applications. Innovations in bioactive materials and technologies will likely lead to even more effective and accessible solutions for dental care. As these systems become more widely adopted, they have the potential to significantly elevate the standards of dental practice and patient care, heralding a new era in restorative dentistry.

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