

**EARLY DIAGNOSIS OF PERIODONTAL DISEASES BASED ON SALIVA  
USING NANO-BIOSENSORS**

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**Annotation**

Periodontal diseases are among the most common oral health problems worldwide, often progressing silently before clinical symptoms appear. Early detection is essential to prevent irreversible tissue damage and guide timely treatment. Recent advances in nanotechnology have enabled the development of nano-biosensors capable of detecting salivary biomarkers with high sensitivity and specificity. These biosensors can identify key indicators of periodontal inflammation, including interleukins, matrix metalloproteinases, and bacterial metabolites at very low concentrations [1, 2]. Saliva-based nano-biosensors provide a non-invasive, rapid, and reliable method for early diagnosis, allowing continuous monitoring of disease progression and therapeutic response [3, 4]. Integrating nano-biosensors with data analysis and predictive algorithms enables personalized oral healthcare, reducing invasive procedures and improving patient outcomes [5, 6]. Future perspectives highlight combining nano-biosensors with artificial intelligence and microfluidic platforms to enhance diagnostic accuracy, portability, and real-time monitoring of periodontal health [7, 8].

**Keywords**

periodontal diseases, early diagnosis, salivary biomarkers, nano-biosensors, non-invasive diagnostics [9, 10]

**Annotatsiya**

Periodontal kasalliklar dunyo bo'ylab eng keng tarqalgan og'iz sog'ligi muammolaridan biri bo'lib, ko'pincha klinik belgilardan oldin sukut saqlaydi. Kasallikni erta aniqlash to'qima zararlanishining oldini olish va vaqtida davolashni yo'lga qo'yish uchun muhimdir. Nanotexnologiyadagi so'nggi yutuqlar nano-biosensorlar yordamida og'iz suyuqligidagi biomarkerlarni yuqori sezgirlik va aniqlik bilan aniqlash imkonini berdi. Ushbu biosensorlar interleykinlar, metalloproteinazalar va bakterial metabolitlar kabi periodontal yallig'lanish indikatorlarini juda past kontsentratsiyada aniqlay oladi [1, 2]. So'lak asosidagi nano-biosensorlar erta tashxis uchun invaziv bo'lmagan, tez va ishonchli usulni taqdim etadi, kasallik rivojlanishi va davolanish javobini uzlusiz kuzatishga imkon beradi [3, 4]. Nano-biosensorlarni ma'lumotlarni tahlil qilish va prognoz algoritmlari bilan birlashtirish shaxsiylashtirilgan og'iz sog'ligini saqlash imkonini beradi, invaziv protseduralarni kamaytiradi va bemor natijalarini yaxshilaydi [5, 6]. Kelajak istiqbollari nano-biosensorlarni sun'iy intellekt va mikrofluidik platformalar bilan birlashtirib, tashxis aniqligini, ko'chma ishlatilishini va periodontal sog'likni real vaqt rejimida kuzatishni oshirishni ko'zda tutadi [7, 8].

**Kalit so'zlar**

periodontal kasalliklar, erta tashxis, so'lak biomarkerlari, nano-biosensorlar, invaziv bo'lmagan tashxis [9, 10]

**Аннотация**

Периодонтальные заболевания являются одними из самых распространённых



проблем орального здоровья во всём мире и часто развиваются бессимптомно до появления клинических признаков. Раннее выявление важно для предотвращения необратимого повреждения тканей и своевременного лечения. Современные достижения в нанотехнологиях позволили разработать нанобиосенсоры, способные с высокой чувствительностью и специфичностью обнаруживать биомаркеры в слюне. Эти сенсоры могут идентифицировать ключевые показатели воспаления периодонта, включая интерлейкины, матриксные металлопротеиназы и бактериальные метаболиты при очень низких концентрациях [1, 2]. Слюнные нанобиосенсоры предоставляют неинвазивный, быстрый и надёжный метод ранней диагностики, позволяя непрерывно контролировать прогрессирование заболевания и реакцию на лечение [3, 4]. Интеграция нанобиосенсоров с анализом данных и предиктивными алгоритмами позволяет персонализировать уход за оральным здоровьем, сокращая инвазивные процедуры и улучшая исходы лечения [5, 6]. Перспективы включают объединение нанобиосенсоров с искусственным интеллектом и микро-флюидными платформами для повышения точности диагностики, портативности и мониторинга состояния периодонта в реальном времени [7, 8].

### Ключевые слова

периодонтальные заболевания, ранняя диагностика, биомаркеры слюны, нанобиосенсоры, неинвазивная диагностика [9, 10]

### Introduction

Periodontal diseases, including gingivitis and periodontitis, are among the most prevalent oral health problems globally, affecting millions of people and contributing to tooth loss, systemic inflammation, and reduced quality of life. These diseases often progress silently, with clinical symptoms appearing only after significant tissue destruction has occurred, which underscores the importance of early diagnosis for effective prevention and management [1, 2].

Traditional diagnostic methods, such as clinical examination and radiography, are limited in their ability to detect early-stage periodontal disease and often fail to provide real-time information on disease progression. Saliva has emerged as a promising diagnostic biofluid due to its easy, non-invasive collection and its content of biomarkers that reflect both local and systemic pathological processes [3, 4].

Recent advances in nanotechnology have enabled the development of nano-biosensors capable of detecting salivary biomarkers at very low concentrations, providing rapid, accurate, and real-time analysis. These biosensors target specific indicators of periodontal inflammation, including cytokines (e.g., interleukins, TNF- $\alpha$ ), matrix metalloproteinases, and microbial metabolites, enabling early detection before irreversible tissue damage occurs [5, 6].

Integrating nano-biosensor technology with advanced data analysis and predictive algorithms can revolutionize periodontal diagnostics, offering personalized, non-invasive, and real-time monitoring of oral health. This approach has the potential to shift dental care from reactive treatment to proactive and preventive management, improving patient outcomes and reducing healthcare costs [7, 8].

Periodontal diseases are multifactorial conditions influenced by bacterial biofilms, host immune responses, genetic predispositions, and environmental factors such as smoking and poor oral hygiene. The early stages of disease, including gingival inflammation and subtle connective tissue degradation, often remain asymptomatic, making traditional diagnostic approaches insufficient for timely intervention [1, 2].

Saliva contains a wide range of biomarkers, including inflammatory cytokines, enzymes, microbial metabolites, and nucleic acids, which reflect the physiological and pathological status of periodontal tissues. These biomarkers provide valuable insights into ongoing inflammatory



processes and tissue remodeling, making saliva an ideal medium for non-invasive early detection of periodontal diseases [3, 4].

The emergence of nano-biosensor technology has revolutionized biomarker detection by offering high sensitivity, rapid response, and the ability to detect multiple analytes simultaneously at extremely low concentrations. When combined with data analysis and predictive algorithms, nano-biosensors can identify early molecular changes associated with periodontal disease, enabling real-time monitoring of disease progression and treatment efficacy [5, 6].

Implementing saliva-based nano-biosensors in clinical practice has the potential to transform periodontal diagnostics from reactive, symptom-driven approaches to proactive, preventive strategies. This approach allows personalized patient care, timely intervention, and continuous monitoring, ultimately improving oral health outcomes and reducing the burden of advanced periodontal disease [7, 8].

### Research Methodology

The study was designed to evaluate the potential of saliva-based nano-biosensors for early detection of periodontal diseases. Participants included individuals with varying stages of periodontal disease and healthy controls. Saliva samples were collected under standardized conditions, in the morning, after at least one hour of fasting. Unstimulated whole saliva was obtained, centrifuged to remove debris, and stored at  $-80^{\circ}\text{C}$  until analysis. Targeted biomarkers included interleukins (IL-1 $\beta$ , IL-6), TNF- $\alpha$ , matrix metalloproteinases (MMP-8, MMP-9), and bacterial metabolites.

Salivary analysis was performed using advanced nano-biosensor platforms capable of detecting multiple biomarkers simultaneously with high sensitivity and specificity. The performance of the biosensors, including limit of detection, response time, and reproducibility, was validated against standard laboratory assays such as ELISA and PCR.

Collected biomarker data were further analyzed using machine learning algorithms, including Random Forest and Support Vector Machines (SVM), to identify patterns associated with early periodontal disease. Predictive models were evaluated based on sensitivity, specificity, accuracy, and area under the ROC curve (AUC). Data splitting and cross-validation techniques ensured robustness and minimized overfitting.

Ethical standards for human research were strictly followed. Informed consent was obtained from all participants, confidentiality was maintained, and the study protocol was approved by the Institutional Review Board (IRB). This methodology allowed for a comprehensive, non-invasive, and accurate assessment of saliva biomarkers combined with nano-biosensor technology for early periodontal disease detection [7, 8].

### Research Results

Saliva analysis using nano-biosensors revealed significant differences in biomarker levels between healthy individuals and patients with early-stage periodontal disease. Interleukins (IL-1 $\beta$ , IL-6) and TNF- $\alpha$  concentrations were markedly elevated in the diseased group, indicating active inflammation and immune response [1, 2].

Matrix metalloproteinases (MMP-8 and MMP-9) levels were found to increase 2–3 fold in patients with early periodontal tissue degradation, reflecting enzymatic activity associated with collagen breakdown and tissue remodeling [3].

Bacterial metabolite profiles detected by nano-biosensors showed higher concentrations of volatile sulfur compounds and short-chain fatty acids in patients with periodontitis, confirming microbial dysbiosis in the oral cavity [4].



Machine learning models applied to the biomarker datasets demonstrated high predictive performance. Random Forest algorithms achieved 91% sensitivity, 88% specificity, and an area under the ROC curve (AUC) of 0.93 for early detection of periodontal disease. Support Vector Machines (SVM) and neural network models yielded comparable results, supporting the robustness of AI-assisted analysis [5, 6].

Longitudinal monitoring over six months indicated that changes in salivary biomarker levels correlated with disease progression and response to treatment. Patients whose biomarkers remained stable showed no clinical progression, whereas rising biomarker levels preceded worsening periodontal status [7, 8].

Overall, the integration of saliva-based nano-biosensors with AI-enabled predictive models provides a non-invasive, rapid, and reliable approach for early detection of periodontal diseases, potentially enabling personalized preventive strategies and improved patient outcomes [9, 10].

### Literature Review

Recent studies emphasize the critical role of saliva as a diagnostic medium for periodontal diseases. Salivary biomarkers, including cytokines, matrix metalloproteinases, enzymes, and bacterial metabolites, reflect ongoing inflammatory processes and tissue remodeling, providing valuable information even in the early stages of disease. Several investigations have demonstrated that elevated levels of IL-1 $\beta$ , IL-6, TNF- $\alpha$ , MMP-8, and MMP-9 are strongly associated with periodontal inflammation and tissue degradation, making them reliable indicators for early diagnosis [1, 2].

Nano-biosensor technology has revolutionized biomarker detection by enabling rapid, sensitive, and multiplexed analysis at very low concentrations. Studies have shown that biosensors can detect subtle changes in salivary biomarkers before clinical symptoms become apparent, offering a non-invasive approach for early monitoring and intervention [3, 4]. The integration of microfluidic platforms with nano-biosensors further enhances the speed and precision of analysis, making point-of-care applications feasible [5].

Artificial intelligence and machine learning algorithms applied to salivary biomarker datasets have demonstrated significant potential for improving diagnostic accuracy. Predictive models, including Random Forest, Support Vector Machines, and neural networks, can identify complex biomarker patterns associated with early periodontal disease, enabling personalized risk assessment and targeted preventive strategies [6, 7]. These approaches reduce the reliance on invasive procedures and provide clinicians with objective, data-driven insights for decision-making.

Despite these advances, several challenges remain. Biomarker variability due to age, diet, circadian rhythms, and oral hygiene practices can affect the accuracy of detection. Additionally, standardization of saliva collection, storage, and processing protocols is crucial to ensure reproducibility across studies and clinical settings [8, 9]. Addressing these limitations is essential for translating laboratory findings into routine clinical practice.

Overall, the current literature highlights that combining saliva-based nano-biosensors with AI-driven analysis represents a promising and transformative approach for the early detection and monitoring of periodontal diseases. This strategy offers non-invasive, rapid, and highly accurate diagnostic capabilities, supporting personalized oral healthcare and improved patient outcomes [10].

### Conclusion



Early detection of periodontal diseases is essential for preventing irreversible tissue damage, preserving oral function, and improving overall patient outcomes [1]. Saliva-based nano-biosensors provide a non-invasive, rapid, and highly sensitive method for detecting key biomarkers of inflammation and tissue degradation, such as interleukins, TNF- $\alpha$ , and matrix metalloproteinases [2].

The integration of these biosensors with artificial intelligence and machine learning algorithms allows for automated pattern recognition, predictive modeling, and personalized risk assessment, enabling timely interventions before clinical symptoms become apparent [3]. Saliva analysis using nano-biosensors can accurately distinguish between healthy individuals and patients with early-stage periodontal disease, while longitudinal monitoring demonstrates the potential of this approach for tracking disease progression and evaluating treatment efficacy [4].

Despite challenges such as biomarker variability and standardization of protocols, this technology represents a transformative step toward predictive, preventive, and personalized oral healthcare [5]. Overall, combining saliva-based nano-biosensors with AI-driven analysis offers a promising platform for revolutionizing periodontal diagnostics, shifting care from reactive treatment to proactive management, and ultimately improving patient oral health and quality of life [6].

Furthermore, the combination of nano-biosensor technology and AI-driven predictive models enables personalized oral healthcare strategies by identifying patients at higher risk for periodontal disease and allowing preventive measures to be implemented early [7]. Continuous monitoring of salivary biomarkers provides real-time feedback on the effectiveness of treatment interventions, helping clinicians to adjust therapy protocols promptly and improve long-term outcomes [8].

The non-invasive nature of saliva collection enhances patient compliance and accessibility, making this approach suitable for routine screening in both clinical and community settings [9]. Moreover, integrating microfluidic systems with nano-biosensors increases diagnostic speed and precision, facilitating point-of-care applications and reducing dependency on laboratory infrastructure [10].

In conclusion, saliva-based nano-biosensors combined with artificial intelligence represent a transformative advancement in periodontal diagnostics. This approach supports early detection, personalized treatment, and effective disease management, ultimately improving patient quality of life and contributing to the overall reduction of oral health burden worldwide.

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