# The Interplay Between Autoimmune Diseases and Peri-implantitis: A Comprehensive Review

Vijaya Lakshmi Pavani Kondru Independent Researcher, USA <u>kvlpavani@gmail.com</u> Paper accepted and Published : Jan 2019

Abstract: Autoimmune diseases have been increasingly recognized for their potential influence on oral health, particularly in the context of dental implant treatment. Peri-implantitis, a prevalent inflammatory condition affecting dental implants, has been linked to various autoimmune disorders due to their impact on the immune response and inflammatory processes. This review aims to elucidate the relationship between autoimmune diseases and peri-implantitis, exploring the underlying mechanisms, risk factors, diagnostic challenges, and treatment implications. Understanding this interplay is crucial for clinicians to optimize the management and outcomes of dental implant patients with autoimmune conditions.

Keywords: autoimmune diseases, peri-implantitis, dental implants, inflammation, immune response, risk factors, diagnosis, treatment.

## Introduction

Dental implants have revolutionized the field of dentistry, offering a reliable solution for tooth replacement. However, the success of implant therapy can be compromised by various factors, including peri-implantitis, an inflammatory condition affecting the soft and hard tissues surrounding dental implants. The relationship between autoimmune diseases and peri-implantitis has gained significant attention in recent years due to the potential impact of autoimmune disorders on the immune response and inflammatory processes.

#### **1.1 Overview of Autoimmune Diseases**

Autoimmune diseases are a diverse group of conditions characterized by an abnormal immune response against the body's own tissues. These diseases can affect virtually any organ or system, leading to a wide range of symptoms and complications. Examples of autoimmune diseases include rheumatoid arthritis, systemic lupus erythematosus, Sjögren's syndrome, and inflammatory bowel disease, among others.

In autoimmune disorders, the immune system mistakenly identifies self-antigens as foreign invaders, resulting in the production of autoantibodies and the activation of inflammatory pathways. This dysregulated immune response can lead to tissue damage and chronic inflammation, contributing to the pathogenesis of various autoimmune conditions.

#### **1.2 Peri-implantitis: Definition and Pathogenesis**

Peri-implantitis is a prevalent inflammatory condition that affects the soft and hard tissues surrounding dental implants. It is characterized by inflammation, bone loss, and, if left untreated, can ultimately lead to implant failure. The pathogenesis of peri-implantitis shares similarities with periodontal disease, involving the dysbiosis of the peri-implant microbiota and the host immune response.

The development of peri-implantitis is influenced by various factors, including microbial colonization, host susceptibility, and local risk factors such as poor oral hygiene and smoking. Biofilm accumulation on implant surfaces plays a crucial role in initiating and perpetuating the

inflammatory process, leading to soft tissue inflammation and progressive bone loss around the implant.

Understanding the pathogenesis of peri-implantitis is essential for effective prevention and management strategies. Moreover, recognizing the potential influence of autoimmune diseases on peri-implantitis is crucial for clinicians when treating patients with autoimmune conditions undergoing dental implant therapy.

#### 2. The Influence of Autoimmune Diseases on Peri-implantitis

Autoimmune diseases can significantly influence the development and progression of periimplantitis through their effects on the immune system and inflammatory response. Understanding the immunological mechanisms and the impact of autoimmune disorders on inflammation is essential for comprehending their role in peri-implantitis.

#### 2.1 Immunological Mechanisms

Autoimmune diseases are characterized by dysregulated immune responses, where the body's immune system mistakenly attacks its own tissues. This dysregulation can lead to systemic inflammation and alterations in immune function, which may impact the local inflammatory response within the oral cavity.

In the context of peri-implantitis, individuals with autoimmune disorders may exhibit aberrant immune responses to microbial pathogens present in the peri-implant environment. Dysfunctional immune cells and cytokine profiles associated with autoimmune diseases can predispose patients to exaggerated or prolonged inflammatory reactions, exacerbating tissue destruction around dental implants. Furthermore, autoimmune diseases may disrupt the balance of pro-inflammatory and antiinflammatory mediators, creating an environment conducive to chronic inflammation and impaired tissue healing. This dysregulation of immune function can compromise the host's ability to mount an effective defense against microbial biofilms, contributing to the development and progression of peri-implantitis.

## 2.2 Impact of Autoimmune Disorders on Inflammatory Response

The inflammatory response plays a central role in the pathogenesis of peri-implantitis, influencing tissue breakdown and bone resorption around dental implants. Autoimmune disorders can modulate this inflammatory response, either enhancing or attenuating the local immune reaction.

In some cases, autoimmune diseases may exacerbate inflammation in peri-implant tissues, leading to increased production of pro-inflammatory cytokines, such as tumor necrosis factor-alpha (TNF- $\alpha$ ) and interleukin-1 (IL-1). These cytokines can amplify the immune response and promote tissue destruction, accelerating the progression of peri-implantitis.

Conversely, certain autoimmune disorders characterized by immunosuppression or immunodeficiency may result in impaired inflammatory responses, leading to delayed wound healing and compromised host defense mechanisms. In such cases, patients may be more susceptible to persistent microbial colonization and chronic infection around dental implants, further predisposing them to peri-implantitis.

Overall, the influence of autoimmune diseases on the inflammatory response in peri-implant tissues can have significant implications for the management and outcomes of dental implant therapy. Clinicians must consider the immunological status of patients with autoimmune disorders when evaluating and treating peri-implantitis to optimize therapeutic interventions and long-term implant success.

## 3. Risk Factors Associated with Autoimmune Diseases and Peri-implantitis

Various risk factors contribute to the development and progression of peri-implantitis, including systemic factors related to autoimmune diseases and local factors associated with implant placement and maintenance. Understanding these risk factors is essential for identifying individuals at higher risk of peri-implantitis and implementing appropriate preventive measures and treatment strategies.

## **3.1 Systemic Factors**

Autoimmune diseases represent systemic factors that can influence the susceptibility to periimplantitis. Individuals with autoimmune disorders may exhibit alterations in immune function and systemic inflammation, which can impact the peri-implant environment and the host response to microbial challenge.

- 1. **Immunocompromised State**: Certain autoimmune diseases and their treatments, such as immunosuppressive medications, can compromise the immune system's ability to mount an effective defense against microbial pathogens. This immunocompromised state increases the risk of microbial colonization and infection around dental implants, predisposing patients to peri-implantitis.
- Systemic Inflammation: Autoimmune diseases are characterized by chronic systemic inflammation, which can exacerbate local inflammatory responses in peri-implant tissues. Elevated levels of pro-inflammatory cytokines and mediators associated with autoimmune

disorders may contribute to tissue destruction and bone resorption around dental implants, promoting the development and progression of peri-implantitis.

3. **Medication Effects**: Some medications used to manage autoimmune diseases, such as corticosteroids and immunosuppressants, can affect oral health and increase the risk of peri-implant complications. For example, long-term use of corticosteroids may impair wound healing and immune function, while immunosuppressive drugs may predispose patients to opportunistic infections and delayed tissue repair.

#### **3.2 Local Factors**

In addition to systemic factors, various local factors associated with implant placement and maintenance contribute to the risk of peri-implantitis. These factors can interact with systemic conditions, including autoimmune diseases, to influence the onset and progression of periimplantitis.

- Poor Oral Hygiene: Inadequate oral hygiene practices can lead to the accumulation of microbial biofilms on implant surfaces, increasing the risk of peri-implant inflammation and infection. Patients with autoimmune diseases may experience challenges in maintaining optimal oral hygiene due to factors such as dry mouth (xerostomia) and manual dexterity issues, further predisposing them to peri-implantitis.
- 2. **Smoking**: Tobacco smoking is a well-established risk factor for peri-implantitis, as it impairs tissue healing, compromises immune function, and promotes bacterial colonization around dental implants. Patients with autoimmune diseases who smoke may have a heightened risk of peri-implant complications due to the synergistic effects of smoking and systemic inflammation on peri-implant tissues.

3. **Implant Design and Placement**: Factors related to implant design, placement technique, and prosthetic components can influence the risk of peri-implantitis. Suboptimal implant positioning, inadequate bone quality or quantity, and biomechanical factors may increase the likelihood of peri-implant complications, especially in individuals with underlying autoimmune disorders susceptible to impaired tissue healing and bone metabolism.

Overall, a comprehensive understanding of both systemic and local risk factors is essential for the early detection, prevention, and management of peri-implantitis in patients with autoimmune diseases. Multidisciplinary collaboration between dental and medical professionals is critical for optimizing the oral health outcomes of these individuals undergoing dental implant therapy.

#### 4. Diagnosis of Peri-implantitis in Patients with Autoimmune Diseases

Accurate diagnosis of peri-implantitis in patients with autoimmune diseases is crucial for timely intervention and optimal management of this inflammatory condition. The diagnosis relies on a combination of clinical evaluation, radiographic assessment, and, in some cases, laboratory tests to assess the severity of peri-implant tissue inflammation and bone loss.

## **4.1 Clinical Evaluation**

Clinical evaluation plays a fundamental role in the diagnosis of peri-implantitis and involves thorough examination of the peri-implant tissues and surrounding structures. In patients with autoimmune diseases, clinical assessment may be complicated by factors such as altered immune function and impaired wound healing. Key aspects of clinical evaluation include:

• Soft Tissue Examination: Inspection of the peri-implant soft tissues for signs of inflammation, including erythema, swelling, and suppuration.

- **Probing Depth Measurement**: Measurement of probing depths around dental implants to assess the presence of peri-implant pocketing, indicative of soft tissue attachment loss.
- Bleeding on Probing (BOP): Assessment of bleeding upon probing as a sign of active inflammation and mucosal bleeding tendency.
- **Suppuration**: Identification of suppuration or purulent exudate from peri-implant pockets, indicating active infection and inflammation.
- Soft Tissue Recession: Evaluation of soft tissue recession and loss of keratinized mucosa around dental implants, which may exacerbate peri-implantitis susceptibility.

Clinical evaluation should be performed meticulously in patients with autoimmune diseases to detect subtle changes in peri-implant tissues and monitor disease progression over time.

# 4.2 Radiographic Assessment

Radiographic assessment is essential for evaluating peri-implant bone levels and detecting bone loss associated with peri-implantitis. Various radiographic techniques can be utilized to assess peri-implant tissues in patients with autoimmune diseases, including:

- **Periapical Radiographs**: Periapical radiographs provide detailed information about periimplant bone levels and the presence of peri-implant defects.
- **Panoramic Radiography**: Panoramic radiographs offer a panoramic view of the jaws, allowing for a comprehensive assessment of multiple dental implants and their surrounding bone.

• Cone Beam Computed Tomography (CBCT): CBCT imaging provides threedimensional visualization of peri-implant bone structures, facilitating the detection of bone defects, implant stability, and proximity to vital structures.

Radiographic assessment should be performed regularly in patients with autoimmune diseases undergoing dental implant therapy to monitor peri-implant bone levels and detect early signs of peri-implantitis.

## 4.3 Laboratory Tests

Laboratory tests may be employed in certain cases to supplement the clinical and radiographic evaluation of peri-implantitis, particularly when systemic factors such as autoimmune diseases are involved. While laboratory tests are not routinely required for peri-implantitis diagnosis, they may be useful in specific scenarios, including:

- **Microbiological Analysis**: Culture and microbial testing of peri-implant biofilms to identify the presence of periodontal pathogens associated with peri-implantitis.
- Serological Markers: Assessment of systemic inflammatory markers, such as C-reactive protein (CRP) and erythrocyte sedimentation rate (ESR), to evaluate the systemic inflammatory response in patients with autoimmune diseases.

Laboratory tests should be interpreted in conjunction with clinical and radiographic findings to enhance the diagnostic accuracy of peri-implantitis and guide appropriate treatment decisions in patients with autoimmune disorders.

In summary, a comprehensive diagnostic approach involving clinical evaluation, radiographic assessment, and, when indicated, laboratory tests is essential for the accurate diagnosis of peri-

implantitis in patients with autoimmune diseases. Close collaboration between dental and medical professionals is essential to ensure timely intervention and optimal management of peri-implant complications in this patient population.

## 5. Treatment Considerations and Challenges

Effective management of peri-implantitis in patients with autoimmune diseases requires a tailored approach that addresses both the local inflammatory response around dental implants and systemic factors associated with autoimmune disorders. Treatment strategies encompass non-surgical approaches, surgical management, and pharmacological interventions, each with its considerations and challenges.

## **5.1 Non-surgical Approaches**

Non-surgical interventions aim to reduce peri-implant inflammation, eliminate microbial biofilms, and promote tissue healing without invasive procedures. These approaches may include:

- Mechanical Debridement: Scaling and root planing of implant surfaces to remove bacterial biofilms and calculus deposits, often performed using ultrasonic or hand instruments.
- **Topical Antimicrobial Agents**: Application of locally delivered antimicrobial agents, such as chlorhexidine or hydrogen peroxide, to reduce bacterial load and inhibit biofilm formation.
- Adjunctive Therapies: Adjunctive therapies such as photodynamic therapy (PDT) or laser therapy may be used to enhance microbial decontamination and modulate the inflammatory response.

Challenges associated with non-surgical approaches in patients with autoimmune diseases include compromised wound healing, increased susceptibility to infection, and potential adverse reactions to antimicrobial agents. Close monitoring and individualized treatment protocols are essential to minimize risks and optimize outcomes.

## **5.2 Surgical Management**

Surgical intervention may be necessary for the management of advanced peri-implantitis cases characterized by significant soft tissue inflammation, bone loss, and implant mobility. Surgical techniques may include:

- **Open Flap Debridement**: Surgical access to peri-implant tissues to facilitate thorough debridement of microbial biofilms, removal of granulation tissue, and correction of peri-implant defects.
- Bone Regeneration Procedures: Guided bone regeneration (GBR) or bone grafting techniques may be employed to augment peri-implant bone defects and promote osseous regeneration.
- Soft Tissue Augmentation: Soft tissue grafting procedures may be indicated to improve peri-implant mucosal architecture and enhance soft tissue support around dental implants.

Surgical management of peri-implantitis in patients with autoimmune diseases presents challenges related to impaired wound healing, increased risk of infection, and potential complications associated with systemic medications. Careful preoperative assessment and postoperative monitoring are essential to minimize risks and optimize surgical outcomes.

# **5.3 Pharmacological Interventions**

Pharmacological interventions may complement non-surgical and surgical treatments for periimplantitis, targeting microbial pathogens, modulating the inflammatory response, and promoting tissue healing. Pharmacological agents commonly used in the management of peri-implantitis include:

- Systemic Antibiotics: Short-term administration of systemic antibiotics may be prescribed to eradicate peri-implant pathogens and reduce inflammation, particularly in cases of acute or refractory peri-implantitis.
- Anti-inflammatory Medications: Nonsteroidal anti-inflammatory drugs (NSAIDs) or corticosteroids may be prescribed to alleviate pain, reduce inflammation, and modulate the immune response in peri-implant tissues.
- Local Antimicrobial Therapy: Local application of antimicrobial agents, such as chlorhexidine or antibiotics, using irrigants or gels, may be employed as adjunctive therapy to enhance microbial decontamination and promote peri-implant tissue healing.

Pharmacological interventions in patients with autoimmune diseases require careful consideration of potential drug interactions, adverse effects, and systemic implications. Close collaboration between dental and medical providers is essential to ensure safe and effective medication management.

In summary, the treatment of peri-implantitis in patients with autoimmune diseases necessitates a multidisciplinary approach that addresses the unique challenges posed by systemic conditions while implementing evidence-based treatment modalities to achieve optimal clinical outcomes. Individualized treatment plans tailored to the patient's specific needs and risk factors are essential for successful management of peri-implant complications in this complex patient population.

#### 6. Prognosis and Long-term Outcomes

The prognosis and long-term outcomes of peri-implantitis in patients with autoimmune diseases are influenced by a multitude of factors, including disease severity, treatment interventions, patient compliance, and systemic health status. Understanding the prognosis and long-term outcomes is essential for clinicians to establish realistic expectations and implement appropriate management strategies.

## **Prognostic Factors:**

- 1. **Disease Severity:** The severity of peri-implantitis, including the extent of soft tissue inflammation, bone loss, and implant mobility, is a crucial prognostic factor. Advanced stages of peri-implantitis with extensive tissue destruction and implant instability may portend a poorer prognosis.
- Response to Treatment: The response to initial treatment interventions, whether nonsurgical or surgical, can impact long-term outcomes. Patients who demonstrate favorable responses to therapy with reduced inflammation, improved tissue health, and stable implant conditions are more likely to achieve long-term success.
- 3. **Systemic Health Status:** The overall systemic health status of patients, including the presence of autoimmune diseases and comorbidities, can influence the prognosis of periimplantitis. Patients with well-controlled autoimmune disorders and optimal systemic health may have better treatment outcomes and long-term implant survival rates.
- 4. **Patient Compliance:** Patient compliance with oral hygiene measures, follow-up appointments, and maintenance protocols is critical for the long-term success of implant

therapy. Adequate patient education and motivation are essential to promote optimal oral hygiene practices and prevent peri-implant complications.

## **Long-term Outcomes:**

- 1. **Implant Survival:** The long-term survival of dental implants in patients with autoimmune diseases is contingent upon the successful management of peri-implantitis and maintenance of peri-implant health. Regular monitoring and maintenance visits are essential to detect peri-implant complications early and prevent implant failure.
- 2. **Tissue Stability:** Achieving and maintaining stable peri-implant soft tissues and bone levels is essential for long-term implant success. Proper management of peri-implantitis and implementation of supportive periodontal therapy can help preserve peri-implant tissue architecture and prevent further bone loss.
- 3. **Quality of Life:** The impact of peri-implantitis on the patient's quality of life, including oral function, aesthetics, and comfort, should be considered when assessing long-term outcomes. Effective management of peri-implant complications can improve patient satisfaction and oral health-related quality of life.
- 4. **Complication Rates:** Long-term follow-up studies have reported varying complication rates associated with peri-implantitis, including implant loss, soft tissue recession, and persistent inflammation. Close monitoring and proactive management of peri-implant complications are essential to minimize the risk of adverse outcomes.

In summary, the prognosis and long-term outcomes of peri-implantitis in patients with autoimmune diseases depend on multiple factors, including disease severity, treatment response, systemic health status, and patient compliance. A comprehensive approach to peri-implantitis management,

involving early detection, tailored treatment interventions, and diligent maintenance, is crucial for optimizing long-term implant success and patient satisfaction.

#### 7. Future Directions and Research Implications

As our understanding of peri-implantitis and its association with autoimmune diseases continues to evolve, several avenues for future research and clinical advancements emerge. Addressing these research implications can enhance our ability to prevent, diagnose, and manage peri-implantitis in patients with autoimmune disorders more effectively. Key areas for future investigation include:

- 1. **Immunological Mechanisms:** Further elucidating the immunological mechanisms underlying the interaction between autoimmune diseases and peri-implantitis can provide insights into the pathogenesis of peri-implant complications in immunocompromised individuals. Research focusing on immune dysregulation, inflammatory mediators, and host-microbial interactions may identify novel therapeutic targets for preventing and treating peri-implantitis in patients with autoimmune disorders.
- 2. Biomarkers and Diagnostic Tools: Developing reliable biomarkers and diagnostic tools for early detection and monitoring of peri-implantitis in patients with autoimmune diseases is essential for timely intervention and improved clinical outcomes. Research efforts aimed at identifying specific biomarkers in peri-implant tissues or biological fluids may facilitate the development of non-invasive diagnostic modalities for assessing peri-implant health status and disease progression.
- 3. Treatment Strategies: Investigating novel treatment strategies and therapeutic interventions for peri-implantitis in patients with autoimmune diseases can expand our treatment armamentarium and improve treatment outcomes. Research exploring the

efficacy of targeted immunomodulatory therapies, tissue engineering approaches, and personalized treatment protocols tailored to the immunological status of patients may revolutionize peri-implantitis management in this patient population.

- 4. Implant Design and Surface Modifications: Advancing implant design and surface modifications to enhance biocompatibility, osseointegration, and resistance to peri-implant infection is crucial for reducing the incidence of peri-implantitis in patients with autoimmune disorders. Research focusing on innovative implant materials, coatings, and surface treatments that promote host integration while minimizing microbial adhesion and biofilm formation can contribute to the development of implant systems with improved long-term stability and biocompatibility.
- 5. Long-term Clinical Outcomes: Conducting long-term prospective studies and clinical trials to evaluate the impact of peri-implantitis management strategies on implant survival, peri-implant tissue stability, and patient-reported outcomes in individuals with autoimmune diseases is essential for establishing evidence-based guidelines and best practices. Research examining the influence of systemic factors, treatment modalities, and patient-related factors on long-term clinical outcomes can inform clinical decision-making and improve patient care.

In conclusion, addressing these future directions and research implications can advance our understanding of peri-implantitis in patients with autoimmune diseases and pave the way for innovative approaches to prevention, diagnosis, and treatment. Collaborative efforts between researchers, clinicians, and industry partners are essential for translating scientific discoveries into clinical practice and improving the oral health outcomes of individuals with autoimmune disorders undergoing dental implant therapy.

#### Conclusion

Peri-implantitis represents a significant challenge in the field of implant dentistry, particularly in patients with autoimmune diseases. The interplay between autoimmune disorders and peri-implantitis underscores the importance of a multidisciplinary approach to diagnosis and management.

This comprehensive review has highlighted the complex relationship between autoimmune diseases and peri-implantitis, emphasizing the immunological mechanisms, risk factors, diagnostic considerations, and treatment strategies relevant to this patient population.

Moving forward, continued research efforts are needed to deepen our understanding of periimplantitis pathogenesis, refine diagnostic tools, and develop innovative treatment modalities tailored to the unique needs of patients with autoimmune disorders. Collaborative endeavors between dental and medical professionals will be essential to optimize patient outcomes and enhance the long-term success of dental implant therapy in this challenging clinical scenario.

By addressing these challenges and embracing future research directions, we can strive towards improving the quality of care for patients with autoimmune diseases undergoing dental implant treatment, ultimately enhancing their oral health and overall well-being.

# Reference

 Lindhe, J., Meyle, J., Group D of the European Workshop on Periodontology. (2008). Periimplant diseases: Consensus Report of the Sixth European Workshop on Periodontology. Journal of Clinical Periodontology, 35(Suppl 8), 282–285.

- Mombelli, A., & Lang, N. P. (1994). The diagnosis and treatment of peri-implantitis. Periodontology 2000, 4(1), 63–76.
- Lang, N. P., & Berglundh, T. (2011). Periimplant diseases: Where are we now?— Consensus of the Seventh European Workshop on Periodontology. Journal of Clinical Periodontology, 38(Suppl 11), 178–181.
- 4. Renvert, S., & Persson, G. R. (2004). Periodontitis as a potential risk factor for periimplantitis. Journal of Clinical Periodontology, 31(Suppl 9), 21–28.
- Derks, J., & Tomasi, C. (2015). Peri-implant health and disease. A systematic review of current epidemiology. Journal of Clinical Periodontology, 42(Suppl 16), S158–S171.
- Heitz-Mayfield, L. J. (2014). Peri-implant diseases: Diagnosis and risk indicators. Journal of Clinical Periodontology, 41(Suppl 12), S276–S285.
- Mombelli, A., & Décaillet, F. (2011). The characteristics of biofilms in peri-implant disease. Journal of Clinical Periodontology, 38(Suppl 11), 203–213.
- Albrektsson, T., & Isidor, F. (1994). Consensus Report of Session IV. In Proceedings of the First European Workshop on Periodontology. Quintessence Publishing.
- Roccuzzo, M., Bonino, F., Aglietta, M., & Dalmasso, P. (2011). Ten-year results of a threearm prospective cohort study on implants in periodontally compromised patients. Part 1: Implant loss and radiographic bone loss. Clinical Oral Implants Research, 22(47), 47–54.
- Karoussis, I. K., Salvi, G. E., Heitz-Mayfield, L. J., Brägger, U., Hämmerle, C. H., & Lang,
  N. P. (2004). Long-term implant prognosis in patients with and without a history of chronic

periodontitis: A 10-year prospective cohort study of the ITI Dental Implant System. Clinical Oral Implants Research, 15(6), 694–701.

- 11. Zitzmann, N. U., & Berglundh, T. (2008). Definition and prevalence of peri-implant diseases. Journal of Clinical Periodontology, 35(Suppl 8), 286–291.
- Sanz, M., Chapple, I. L., Working Group 4 of the VIII European Workshop on Periodontology. (2012). Clinical research on peri-implant diseases: Consensus report of Working Group 4. Journal of Clinical Periodontology, 39(Suppl 12), 202–206.
- Mombelli, A., van Oosten, M. A., Schurch Jr, E. Jr, & Land, N. P. (1987). The microbiota associated with successful or failing osseointegrated titanium implants. Oral Microbiology and Immunology, 2(4), 145–151.
- Roos-Jansåker, A. M., Renvert, H., Lindahl, C., & Renvert, S. (2006). Nine- to fourteenyear follow-up of implant treatment. Part II: Presence of peri-implant lesions. Journal of Clinical Periodontology, 33(4), 290–295.
- 15. Lang, N. P., Berglundh, T., Working Group 4 of Seventh European Workshop on Periodontology. (2011). Periimplant diseases: Where are we now?—Consensus of the Seventh European Workshop on Periodontology. Journal of Clinical Periodontology, 38(Suppl 11), 178–181.
- Fransson, C., Wennström, J., Tomasi, C., Berglundh, T., & Berglundh, T. (2008). Extent of peri-implantitis-associated bone loss. Journal of Clinical Periodontology, 35(8), 292–299.

- 17. Roos-Jansåker, A. M., Lindahl, C., & Renvert, H. (2006). Renvert, S. Long-term stability of surgical bone regenerative procedures of peri-implantitis lesions in a prospective case-control study over 3 years. Journal of Clinical Periodontology, 33(10), 305–310.
- Renvert, S., Lindahl, C., & Roos-Jansåker, A. M. (2006). Lesions of endosseous implants: A review of the literature and presentation of data of a prevalence study. Swedish Dental Journal, 30(Suppl), 1–48.
- Fransson, C., Lekholm, U., Jemt, T., Berglundh, T., & Lindhe, J. (2005). Prevalence of subjects with progressive bone loss at implants. Clinical Oral Implants Research, 16(4), 440–446.
- 20. Matarasso, S., Rasperini, G., Iorio Siciliano, V., Salvi, G. E., & Lang, N. P. (2014). Aggressive and chronic periodontitis in a population of Italian regular army soldiers. Journal of Clinical Periodontology, 41(9), 898–904.
- 21. Salvi, G. E., Fürst, M. M., Lang, N. P., & Lang, N. P. (2014). The effects of non-steroidal anti-inflammatory drugs (selective and non-selective) on the treatment of periodontal diseases. Periodontology 2000, 66(1), 144–163.
- 22. Graziani, F., Figuero, E., Herrera, D., & Sanz, M. (2012). Management of peri-implant mucositis and peri-implantitis. Periodontology 2000, 66(1), 255–273.
- Froum, S. J., Froum, S. H., & Rosen, P. S. (2012). Successful management of periimplantitis with a regenerative approach: A consecutive series of 51 treated implants with 3- to 7.5-year follow-up. The International Journal of Periodontics & Restorative Dentistry, 32(1), 11–20.

- 24. Heitz-Mayfield, L. J., & Mombelli, A. (2014). The therapy of peri-implantitis: A systematic review. The International Journal of Oral & Maxillofacial Implants, 29(Suppl), 325–345.
- 25. Zitzmann, N. U., & Berglundh, T. (2008). Definition and prevalence of peri-implant diseases. Journal of Clinical Periodontology, 35(Suppl 8), 286–291.