BSP Implementation of Prevention and Treatment of Peri-implant Diseases – The EFP S3 Level Clinical Practice Guideline

Nicola West, Iain Chapple, Shauna Culshaw, Nikos Donos, Ian Needleman, Jeanie Suvan, Luigi Nibali, Amit Patel, Philip M Preshaw, Moritz Kebschull, On behalf of the EFP workshop participants and methodological consultant

 PII:
 S0300-5712(24)00150-7

 DOI:
 https://doi.org/10.1016/j.jdent.2024.104980

 Reference:
 JJOD 104980



To appear in: Journal of Dentistry

Received date:11 March 2024Accepted date:28 March 2024

Please cite this article as: Nicola West, Iain Chapple, Shauna Culshaw, Nikos Donos, Ian Needleman, Jeanie Suvan, Luigi Nibali, Amit Patel, Philip M Preshaw, Moritz Kebschull, On behalf of the EFP workshop participants and methodological consultant, BSP Implementation of Prevention and Treatment of Peri-implant Diseases – The EFP S3 Level Clinical Practice Guideline, *Journal of Dentistry* (2024), doi: https://doi.org/10.1016/j.jdent.2024.104980

This is a PDF file of an article that has undergone enhancements after acceptance, such as the addition of a cover page and metadata, and formatting for readability, but it is not yet the definitive version of record. This version will undergo additional copyediting, typesetting and review before it is published in its final form, but we are providing this version to give early visibility of the article. Please note that, during the production process, errors may be discovered which could affect the content, and all legal disclaimers that apply to the journal pertain.

© 2024 Elsevier Ltd. All rights reserved.

i. TITLE

BSP Implementation of Prevention and Treatment of Peri-implant Diseases – The EFP S3 Level Clinical Practice Guideline

ii. SHORT RUNNING TITLE

Prevention and Treatment of Peri-implant Diseases

iii. Full names of the authors

Nicola West^{1,2}, Iain Chapple^{3,4}, Shauna Culshaw⁵, Nikos Donos⁶, Ian Needleman⁷, Jeanie Suvan⁷, Luigi Nibali⁸, Amit Patel⁹, Philip M Preshaw^{10, 11}, Moritz Kebschull^{3,4,12,13}

* On behalf of the EFP workshop participants and methodological consultant (listed below).

iv. Author's institutional affiliations

1. Restorative Dentistry, Bristol Dental School, University of Bristol, Bristol BS1 2LY, UK.

2. Restorative Dentistry, Bristol Dental Hospital, University Hospitals Bristol and

Weston NHS Foundation Trust, Marlborough Street, Bristol, BS1 3NU.

Secretary General of the European Federation of Periodontology

3. Birmingham NIHR Biomedical Research Centre in Inflammation. The University of Birmingham, Birmingham, United Kingdom.

4. Birmingham Community Healthcare NHS Foundation Trust, Birmingham, United Kingdom.
5. University of Glasgow Dental School, College of Medical, Veterinary and Life Sciences,
University of Glasgow, Glasgow, UK.

6. Centre for Oral Clinical Research, Institute of Dentistry, Faculty of Medicine and Dentistry, Queen Mary University of London (AMUL), Turner Street, London, E1 2AD.

7. Unit of Periodontology, UCL Eastman Dental Institute, Bloomsbury Campus, Rockefeller Building, 21 University Street, London, WC1E 6DE.

8. Periodontology Unit, Centre for Host-Microbiome Interactions, Faculty of Dentistry, Oral & Craniofacial Sciences, King's College London, Guys Hospital, Great Maze Pond, London, SE1 9RT

9. Birmingham Dental Specialists, Associate Professor, University of Birmingham,

Birmingham, UK. President of the Association of Dental Implantology

10. School of Dentistry, University of Dundee, Dundee UK

11. School of Dental Sciences, University of Newcastle, Newcastle upon Tyne, UK

12. Division of Periodontics, Section of Oral, Diagnostic and Rehabilitation Sciences, College

of Dental Medicine, Columbia University, New York, NY, USA.

13. School of Dentistry, University of Birmingham, Birmingham B5 7EG, UK. President-Elect of

the European Federation of Periodontology

Corresponding Authors

Professor Nicola West, BDS PhD FDS RCS (Eng) FDS (Rest Dent) FDS RCS Ed FHEA

Professor of Restorative Dentistry (Periodontology), Honorary Consultant in Restorative Dentistry, Director of Dental Clinical Trials Unit, Bristol Dental School, University of Bristol, Lower Maudlin Street, Bristol BS1 2LY, UK Mobile: +44 (0)78 258 313 52 <u>N.X.West@bristol.ac.uk</u>

Professor Moritz Kebschull, Dr. med. dent., Dr. med. dent. habil., MBA

Chair of Restorative Dentistry, School of Dentistry, University of Birmingham, 5 Mill Pool Way, Birmingham B5 7EG, UK. Honorary Consultant in Periodontics, Birmingham Community Healthcare NHS Trust/Birmingham Dental Hospital, 5 Mill Pool Way, Birmingham B5 7EG, UK. <u>moritz@kebschull.me</u>

Abstract

Objectives: to adapt the supranational European Federation of Periodontology (EFP) Prevention and Treatment of Peri-implant Diseases – The EFP S3 Level Clinical Practice Guideline for UK healthcare environment, taking into account a broad range of views from stakeholders and patients.

Sources: This UK version, based on the supranational EFP guideline [1] published in the *Journal of Clinical Periodontology*, was developed using S3-level methodology, combining assessment of formal evidence from 13 systematic reviews with a moderated consensus process of a representative group of stakeholders, and accounts for health equality, environmental factors and clinical effectiveness. It encompasses 55 clinical recommendations for the Prevention and Treatment of Peri-implant Diseases, based on the classification for periodontal and peri-implant diseases and conditions [2].

Methodology: The UK version was developed from the source guideline using a formal process called the GRADE ADOLOPMENT framework. This framework allows for adoption (unmodified acceptance), adaptation (acceptance with modifications) and the *de novo* development of clinical recommendations. Using this framework, following the S3-process, the underlying evidence was updated and a representative guideline group of 111 delegates from 26 stakeholder organisations was assembled into four working groups. Following the formal S3-process, all clinical recommendations were formally assessed for their applicability to the UK and adoloped accordingly.

Results and Conclusion: Using the ADOLOPMENT protocol, a UK version of the EFP S3-level clinical practice guideline for the Prevention and Treatment of Peri-implant Diseases was developed. This guideline delivers evidence- and consensus-based clinical recommendations of direct relevance to the UK healthcare community including the public.

Clinical Significance

The S3-level-guidelines combine evaluation of formal evidence, grading of recommendations and synthesis with clinical expertise of a broad range of stakeholders. The international S3level-guideline was implemented for direct clinical applicability in the UK healthcare system, facilitating a consistent, interdisciplinary, evidence-based approach with public involvement for the prevention and treatment of peri-implant diseases.

Key words: peri-implant diseases, peri-implant mucositis, peri-implantitis, prevention, treatment, clinical guideline, dental implant, patient participation.

Commentary – UK Implementation

This guideline is the UK implementation of the S3-level guideline "Prevention and Treatment of peri-implant diseases" originally developed by the European Federation of Periodontology (EFP). The implementation process in the UK followed the GRADE ADOLOPMENT framework [3].

The EFP guideline was developed by an international working group of periodontists and expert stakeholders. The guideline document was finalised and formally voted upon in a structured consensus conference format during the XVIII European Workshop in Periodontology in La Granja de San Ildefonso, Segovia, Spain, on November 6th- 9th, 2022. The guideline text and the underlying systematic reviews were published in open access format in a special issue of the *Journal of Clinical Periodontology* [1]

Information about the authors of the EFP guideline, their institutions, their declared interests, the workshop participants, the involved stakeholder societies and organisations, the abstract and the description of the clinical relevance can be found in section 9 of this guideline document.

The authors and workshop participants of the British ADOLOPMENT project are listed below.

* Workshop Participants, in alphabetic order

Tameam Alaubidie, Koula Asimakopoulou, Paul Baker, Avijit Banerjee, Daniel Benson, Rickesh Bhopal, Igor Blum, Deborah Bomfim, Nuno Borges, Colin Burns, Elena Calciolari, Iain Chapple, Marilou Ciantar, Rachael Clampin, Peter Clarke, Nick Claydon, Rebecca Cooper (observer), Shauna Culshaw, Andrew Cundy, Francesco D'Aiuto, Radhika Desai, Thomas Dietrich, Nikos Donos, Henry Duncan, Ian Dunn, Kenneth Eaton, Julian Ekiert, Harriet Elsworthy, Gamze Eroglu, Sally Ferrier, Roger Fisher, Claire Forbes-Haley, Ana Beatriz Gamboa, Pynadath George, Mandeep Ghuman, Nikolaos Gkranias, Victoria Griffiths, Alina Grossman (Observer), Kasia Gurzawska-Comis, Stephen Hancocks, Martina Hayes, Debbie Hemington, Mark Ide, Rhiannon Jones, Elaine Judd, Roshni Karia, Moritz Kebschull, Annika Kroeger, Gerry Linden, Emily Lu, Isobel Madden, Kathryn Mayo, Claire McCarthy, Gerry McKenna, William McLaughlin, Payvand Menhadji, Imogen Midwood, Mike Milward, Aidan Moran, Margaret Morgan, Madeleine Murray, Rajan Nansi, Ian Needleman, Luigi Nibali, Eimear O'Connell, Sunkanmi Oladeji Olaore, Amit Patel, Divyash Patel, Rupali Patel, Vipul Patel, Viraj Patel, Michael Paterson, Jeniffer Perussolo, Elliot Pound, Philip Preshaw, Devan Raindi, Raj Rattan, Anthony Roberts, Simone Ruzario, Shazad Saleem, Ross Scales (observer), Sasha Scambler, Joon Seong, Mitul Shah, Shakeel Shahdad, Praveen Sharma, Paul Shenfine, Brinder Singh Shergill, Rajiv Sheth, Andrew Smith, Sami Stagnell, Charlotte Stilwell, Claire Storey, Vanessa Sousa, Jeanie Suvan, Manoj Tank, Joel Thomas, Aradhna Tugnait, Wendy Turner, Jaimini Vadgama, Jiten Vaghela, Bobby Varghese, Viren Vithlani, Jenny Walker, Natasha West, Nicola West, Paul Weston, Matthew Wright, Roger Yates, Albert Yeung, Zehra Yonel.

Methodological Consultant

Professor Ina Kopp

Workshop Organization

British Society of Periodontology and Implant Dentistry

Professor Nicola West & Professor Moritz Kebschull

Scientific societies involved in the guideline development process

Association of Clinical Oral Microbiologists

Association of Dental Implantology

British Association of Dental Therapists

British Association for the Study of Community Dentistry (Public health)

British Dental Association

British Endodontic Society

British Society of Dental Hygiene and Therapy

British Society of Periodontology and Implant Dentistry

British Society of Prosthodontics

British Society of Restorative Dentistry

European College of Gerodontology

European Organisation for Caries Research

European Society of Endodontology

International Team for Implantology

Osteology Foundation

Restorative Dentistry UK

Scottish Antimicrobial Prescribing Group

Other organisations involved in the guideline development process

British Dental Journal

British Society of Periodontology and Implant Dentistry Patient Forum

College of General Dentistry

Council of European Chief Dental Officers

Dental Protection

General Dental Council (Observer)

NHS England and NHS Improvement

NHS Education for Scotland

Platform for Better Oral Health in Europe

Acknowledgements

We would like to thank Nicola Hellin and Natasha West for their assistance in preparation of the manuscript and Roshni Karia for proof reading the manuscript.

We would like to thank all patient representatives and members of the British Society of Periodontology and Implant Dentistry Patient Forum for their participation in the process.

The authors express their gratitude to all reviewers involved in the preparation of the systematic reviews. In addition, we wish to thank the organisations which agreed to participate in the guideline ADOLOPMENT process: Association of Clinical Oral Microbiologists, Association of Dental Implantology, British Association of Dental Therapists, British Association for the Study of Community Dentistry (Public health), British Dental Association, British Dental Journal, British Endodontic Society, British Society of Dental Hygiene and Therapy, British Society of Periodontology and Implant Dentistry, British Society of Periodontology and Implant Dentistry, College of General Dentistry, Council of European Chief Dental Officers, Dental Protection, European College of Gerodontology, European Organisation for Caries Research, European

Society of Endodontology, General Dental Council (Observer), International Team for Implantology, NHS England and NHS Improvement, NHS Education for Scotland, Osteology Foundation, Platform for Better Oral Health in Europe, Restorative Dentistry UK, Scottish Antimicrobial Prescribing Group. We are extremely grateful that no organisations we approached declined to take part in this process.

Journal Presson

1 | INTRODUCTION

1.1 | The Health Problem

1.1.1 | Definition

Peri-implant diseases are inflammatory conditions that affect the peri-implant tissues and are induced by peri-implant biofilms. There are two distinct conditions: peri-implant mucositis and peri-implantitis.

Peri-implant mucositis is "an inflammatory lesion of the peri-implant mucosa, in the absence of continuing marginal bone loss" [4]. It is characterised clinically by bleeding on gentle probing. Other clinical signs of inflammation may be present, such as erythema, swelling and/or suppuration, and an increase in probing depth is frequently observed in the presence of peri-implant mucositis due to oedema or a decrease in probing resistance [5]. Peri-implant mucositis is primarily caused by a disruption of host-microbial homeostasis at the implant-mucosa interface and is a reversible condition when assessed indirectly at the host biomarker level [4]. Additional factors associated with the onset and progression of peri-implant mucositis include biofilm accumulation, smoking, and radiation therapy [5].

Peri-implantitis has been defined as a "peri-implant biofilm-associated pathological condition, occurring in tissues around dental implants, and characterized by inflammation in the periimplant mucosa and subsequent progressive loss of supporting bone" [5]. Clinically, periimplantitis sites exhibit inflammation, bleeding on probing and/or suppuration, increased probing depths, and/or recession of the mucosal margin, in addition to radiographic bone loss compared to previous examinations [5]. The primary etiological factor for peri-implantitis onset and progression is the accumulation of a peri-implant plaque biofilm. Important risk factors/indicators have been identified, including a history of severe periodontitis, poor plaque control, and no regular supportive peri-implant care (SPIC) following implant therapy. Less conclusive evidence was found for smoking and diabetes, or local factors such as the presence of submucosal cement following prosthetic restoration of the implant, or positioning of implants limiting access to oral hygiene and maintenance. Other factors, such as the absence of peri-implant keratinized mucosa, occlusal overload, presence of titanium particles within

peri-implant tissues, bone compression necrosis, overheating, micromotion, or biocorrosion have been proposed as risk factors for peri-implant diseases onset and/or progression, but further research is required to clarify their true roles [6].

Peri-implant diseases, especially peri-implantitis, represent a growing public health problem due to their high prevalence and the associated consequences (implant and implant-supported prosthesis loss), including dental care costs, which are substantial.

1.1.2 | Pathophysiology

To better understand the pathophysiology of peri-implant diseases, knowledge of the pathophysiology of periodontal diseases has been extensively used, and findings on peri-implant mucositis have been likened to those of biofilm-induced gingivitis. The same applies to peri-implantitis and periodontitis. However, when compared to periodontal tissues, peri-implant tissues lack cementum and periodontal ligament; thus, there are only two peri-implant tissue layers, alveolar bone and peri-implant mucosa. Additional differences are found in the peri-implant mucosa: the peri-implant epithelial attachment is usually longer; the connective tissue exhibits no fibres inserting into the supra-crestal area; and vascularization is lower.

Peri-implant biofilms are considered to be the primary etiological factor for peri-implant mucositis, based on strong evidence derived from animal and human studies [5]. Such biofilms form on the hard, non-shedding surfaces of the implant and implant-supported restorations, similar to the formation of dental plaque biofilms on teeth [7, 8]. Histologically, peri-implant mucositis is similar to gingivitis: a well-defined inflammatory lesion, adjacent to the junctional/pocket epithelium, richly infiltrated by vascular structures, plasma cells and lymphocytes, but not extending apically to the junctional/pocket epithelium, or into the supracrestal area [4, 5].

Evidence exists to support the contention that peri-implant mucositis is treatable, and can be successfully managed by careful control of the peri-implant biofilm. However, if allowed to persist, peri-implantitis develops, as it is believed that peri-implant mucositis always precedes peri-implantitis [4, 5].

9

The primary etiological agent for peri-implantitis is also the accumulation of the peri-implant biofilm, with human observational studies demonstrating a higher risk of incident peri-implantitis in patients with poor biofilm control and/or non-adherence to maintenance care, and based on intervention studies using anti-infective approaches [5].

Peri-implantitis lesions are larger than those associated with peri-implant mucositis or with periodontitis and are characterised by greater number of neutrophils and larger proportions of B cells when compared with peri-implant mucositis. Consistent with periodontitis lesions, plasma cells and lymphocytes predominate within the immune-inflammatory infiltrate [6]. However, these characteristic histological features have not been associated with specific bacteria [9] or proinflammatory cytokine profiles [5].

1.1.3 | Prevalence

During the XI European Workshop in Periodontology (2014), entitled "Effective Prevention of Periodontal and Peri-implant Diseases", a systematic review was specifically commissioned to address the prevalence of peri-implant diseases. Eleven studies were selected, and the meta-analyses demonstrated a patient-level prevalence estimate of 43% (95% confidence interval – CI [32; 54]) for peri-implant mucositis and 22% (95% CI [14; 30]) for peri-implantitis [10]. Another systematic review comprising 47 studies, reported a prevalence of 46.83% (95% CI [38.30; 55.36]) for peri-implant mucositis and of 19.83% (95% CI [15.38; 24.27]) for peri-implantitis [11].

1.1.4 | Consequences of failure to treat peri-implant diseases

As previously explained, peri-implant mucositis can be treated and resolved, but if left untreated, can progress to peri-implantitis; peri-implant mucositis is widely believed to precede peri-implantitis. Peri-implantitis can be initiated rapidly following prosthetic restoration and loading of the fixture during function, and if no treatment is provided, it is likely to progress in a non-linear accelerating pattern [5] and at a faster rate than is typically seen in periodontitis lesions [6].

Progression of peri-implantitis will most likely lead to the loss of the affected implant and the implant-supported prosthesis.

Limited information is available on the morbidity associated with peri-implant diseases or their impact on quality of life. One study concluded that neither peri-implantitis nor surgical treatment of the same had any impact on Oral Health Related Quality of Life (OHRQoL) [12], whilst another study assessing morbidity after non-surgical and surgical treatment of peri-implantitis, concluded that pain levels were low to moderate and most pronounced in the first two days [13].

1.1.5 | Financial aspects

According to a market analysis report [14], the global market size of dental implants is estimated at \$4.6 billion USD in 2022 and is expected to grow at an annual rate of around 10%, up to 2030. The increase is based upon the demand for treatment with dental implants by the population and on the widening range of clinicians providing implant therapy. It is also associated with the growing need for longer-term supportive care to avoid/control biological and mechanical complications, including managing complications with implant-supported restorations and maintaining peri-implant tissue health [15]. There is increasing awareness of the need to plan long-term supportive care programs during the treatment planning phase, and of the financial, biological and legal consequences of not doing so. For example, patients may be able to cover the initial cost of dental implants and their associated restorations at the time of implant placement, when they are employed and earning a living, but the long-term cost of supportive care may not be explained clearly to patients and may impact when they are no longer economically active [15]. A Swedish study of 514 subjects recently calculated such costs [16], including the costs of preventive measures and of procedures to treat implant complications, over a period of 8.2 years. The mean cost ranged from €878 Euro's (single-tooth restoration) to €1,210 (full-arch restoration), the larger proportion of the cost being for prevention (\in 741), whilst implant loss was the most expensive complication (\in 1,508), followed by peri-implantitis (€1,244).

A cost-effectiveness analysis was undertaken to assess preventive, non-surgical and surgical interventions ([17], with the model assuming that each implant was followed for 20 years. The annual provision of supportive peri-implant care (SPIC) was dichotomized, and the risk profile of patients was also considered, with implant loss and cost as primary outcomes. For management of peri-implantitis, 11 approaches (non-surgical and surgical instrumentation alone or with adjuncts) were compared. The authors concluded that, within the limitations of

11

their study methodology, not providing annual SPIC increased the risk of peri-implant diseases. Conversely, providing SPIC could prevent or delay the onset of disease and was cost-effective, especially in high-risk groups.

Cost-effectiveness has also been evaluated for non-surgical treatment approaches of periimplantitis [18]. Change in probing depth (PD) was the primary outcome when comparing eight interventions. Instrumentation alone, use of an air polishing device, or combining instrumentation with local antiseptics/antibiotics provided better value for money than Er:YAG laser, a specific ultrasonic device (Vector[®]), photodynamic therapy or instrumentation combined with chlorhexidine.

Of relevance is the cost comparison of SPIC with that of the supportive care of teeth. This was assessed in a private practice in Norway [19] in 43 patients with 847 teeth and 119 implants. The mean number of "disease-free years" was 8.66 for implants, 9.08 for neighbouring teeth, and 9.93 for teeth on the contra-lateral side of the mouth, with no statistically significant differences. However, due to the high prevalence of peri-implantitis, the extra cost of maintaining implants was five times higher than for teeth.

Finally, financial considerations should include the economic impact of edentulism. Whilst not yet clearly established, at least two factors may support its importance: firstly, the need for rehabilitation and the associated costs; secondly, and in case of lack of rehabilitation, the negative consequences for quality of life, nutrition, systemic health and wellbeing. In addition, it is also widely contended that individual- and community-level social inequalities strongly impact on levels of edentulism [20].

2 | AIM OF THE GUIDELINE

This guideline aims to identify best-practice interventions for preserving the health of periimplant tissues and, thereby, extending the longevity of complication-free survival of dental implants when used to replace missing teeth. The main objective, therefore, is to summarize the evidence-based recommendations for individual interventions used in the management

(both prevention and treatment) of peri-implant diseases, based on the best available evidence and/or expert consensus. In so doing, this guideline aims to: (i) inform sound preventive/therapeutic approaches to the management of peri-implant diseases, and thereby improve the overall quality of peri-implant interventions undertaken in Europe and worldwide; (ii) reduce dental implant loss arising due to peri-implantitis, and (iii) ultimately reduce medical and dental costs and improve the quality of life of patients.

2.1 | Target users of the guideline

Oral health professionals, together with stakeholders related to oral health care. In addition, this clinical practice guideline (CPG) aims to inform medical professions, health systems, policymakers, patients and the public.

2.2 | Targeted environments

Academic/hospital environments, community-based dental clinics, and practices.

2.3 | Targeted patient population

People awaiting dental implant rehabilitation.

People receiving dental implant rehabilitation.

People with dental implants and, therefore, at risk of developing peri-implant diseases.

People with peri-implant mucositis.

People with peri-implantitis.

People with peri-implant mucositis, following successful peri-implant treatment.

People with peri-implantitis, following successful peri-implant treatment.

2.4 | Exceptions from the guideline

This guideline does not consider in detail the health/economic cost-benefit ratio of the proposed therapies, since (i) the target users and patient populations include people in different countries with diverse, not readily comparable health care systems, and (ii) there is a paucity of sound scientific data available addressing this issue.

This guideline does not consider the management of other peri-implant tissue conditions, such as hard- and soft-tissue deficiencies around dental implants [21], unusual peri-implant

problems (such as peri-implant peripheral giant-cell granuloma, pyogenic granuloma, squamous cell carcinoma, metastatic carcinomas, malignant melanoma) or implant fractures, that may mimic or share certain clinical features with biofilm-associated peri-implant conditions [22].

3 | METHODOLOGY

3.1 | General framework

This guideline was developed following methodological guidance published by the Standing Guideline Commission of the Association of Scientific Medical Societies in Germany (AWMF) (https://www.awmf.org/ leitlinien/awmf-regelwerk/awmf-guidance.html) and the Grading of Recommendations Assessment, Development and Evaluation (GRADE) Working Group (https://www.gradeworkinggroup.org/).

The guideline was developed under the auspices of the European Federation of Periodontology (EFP) and overseen by the EFP Workshop Committee. This guideline development process was steered by an Organizing Committee and a methodology consultant designated by the EFP. All members of the Organizing Committee participated in the EFP Workshop Committee.

To ensure adequate stakeholder involvement, the EFP established a guideline panel involving dental professionals representing national periodontal societies within the EFP, together with experts in Prosthodontics, Implant Dentistry and Oral Surgery (Table 1). These delegates were nominated and selected by the Organizing Committee and participated in the guideline development process with voting rights in the consensus conference. For the guideline development process, delegates were assigned to four Working Groups that were chaired by selected members of the Organizing Committee and guided by the methodology consultant. This panel was supported by key stakeholders from European scientific societies with a strong professional interest in periodontal care and from European organizations representing key groups within the dental profession (Table 2), and key experts from non-EFP member regions, such as North America and Australia.

Commentary – UK Implementation

The UK implementation followed the GRADE ADOLOPMENT framework [3].

Table 1. Guideline panel.

Scientific society/organisation	Delegate(s)
European Federation of Periodontology	Organising Committee, Working Group
(EFP)	Chairs (in alphabetic order):
	Tord Berglundh, Iain Chapple, David
	Herrera, Søren Jepsen, Moritz
	Kebschull, Panos Papapanou, Mariano
	Sanz, Frank Schwarz, Anton Sculean,
	Maurizio Tonetti
	Methodologist:
	Ina Kopp
	Clinical Experts (in alphabetic order):
0	Mario Aimetti
	Juan Blanco
	Nagihan Bostanci
	Philippe Bouchard
	Nurcan Buduneli
	Elena Calciolari
	María Clotilde Carra
	Raluca Cosgarea
	Jan Cosyn
	Bettina Dannewitz
	Beatriz de Tapia
	Yvonne de Waal
	Jan Derks
	Henrik Dommisch
	Nikos Donos
	Peter Eickholz
	Bahar Eren Kuru
	Elena Figuero
	Moshe Goldstein
	Filippo Graziani
	Jasmin Grischke
	Fernando Guerra

Г	·
	Lisa Heitz-Mayfield
	Karin Jepsen
	Odd Carsten Koldsland
	France Lambert
	Antonio Liñares
	Bruno Loos
	Phoebus Madianos
	Paula Matesanz
	Ana Molina
	Virginie Monnet Corti
	Eduardo Montero
	Frauke Müller
	Luigi Nibali 👞
	Andrés Pascual
	Ioannis Polyzois
	Marc Quirynen
	Ausra Ramanauskaite
	Stefan Renvert
	Mario Roccuzzo
.0	Philipp Sahrmann
	Giovanni Salvi
	Nerea Sánchez
	Ignacio Sanz
	Lior Shapira
	Andreas Stavropoulos
	Meike Stiesch
	Wim Teughels
	Cristiano Tomasi
	Leonardo Trombelli
	Anders Verket
	Asaf Wilensky
Scientific Societies	
European Dental Hygienists Federation	Gitana Rederiene
EFP – Executive Committee	Darko Božić
EFP – Executive Committee	Monique Danser
EFP – Executive Committee	Spyros Vassilopoulos
EFP – Executive Committee	Nicola West
European Society of Endodontology	Lise-Lotte Kirkevang
Other organisations	-
Council of European Dentists	Paulo Melo
European Dental Students' Association	Ieva Tamošiūnaitė

Platform for Better Oral Health in Europe Kenneth Eaton

Table 2. Key stakeholders contacted and participants.

Institution / Society	Acronym	Answer*	Representative
Association for Dental Education in Europe	ADEE	no proposal	none
Continental European Division of IADR	CED-IADR	no proposal	none
Council of European Chief Dental Officers	CECDO	no answer	none
Council of European Dentists	CED	participant	Paulo Melo
European Association for Osseointegration	EAO	participant	cancelled
European Association of Dental Public Health	EADPH	no answer	none
European Dental Hygienists Federation	EDHF	participant	Gitana Rederiene
European Dental Students' Association	EDSA	participant	leva Tamošiūnaitė
European Federation of Conservative Dentistry	EFCD	no answer	none
European Orthodontic Society	EOS	no answer	none
European Prosthodontic Association	EPA	no answer	none
European Society of Endodontology	ESE	participant	Lise Lotte Kirkevang
Platform for Better Oral Health in Europe	РВОНЕ	participant	Kenneth Eaton

*Messages sent on April 4th, 2022

Contributors to the UK Implementation	
	-
Scientific Societies/Organisations	Clinical expert/ Representative (in
$\mathbb{C}^{\mathbb{C}}$	alphabetical order)
Association of Clinical Oral Microbiologists	Professor Andrew Smith
Association of Dental Implantology	Daniel Benson, Rickesh Bhopal, Colin
	Burns, Eimear O'Connell, Nikos Donos,
	Kasia Gurzwaska-Comis, Claire McCarthy,
	Payvand Menhadji, Eimear O'Connell,
	Sunkanmi Oladeji Olaore, Amit Patel, Viraj
	Patel, George Pynadath, Eddie Scher, Mitul
	Shah, Paul Shenfine, Rajiv Sheth, Sami

	Stagnell, Joel Thomas, Jiten Vaghela, Jaimini
	Vadgama, Nicola West
British Association of Dental Therapists	Debbie Hemmington, Claire McCarthy,
	Jeanie Suvan
British Association for the Study of	Albert Yeung
Community Dentistry (Public Health)	
British Dental Association	Avijit Banerjee, Nicola West
British Endodontic Society	Henry Duncan
British Society of Dental Hygiene and	Simone Ruzario, Claire McCarthy, Rhiannon
Therapy	Jones, Victoria Griffiths, Harriet Elsworthy
British Society of Periodontology and Implant	Tameam Alaubidie, Paul Baker, Deborah
Dentistry	Bomfim, Nuno Borges, Iain Chapple, Marilou
	Ciantar, Peter Clarke, Nick Claydon,
0	Shauna Culshaw, Andrew
	Cundy, Francesco D'Aiuto, Radhika Desai,
	Thomas Dietrich, Nikos Donos, Ian Dunn,
	Ken Eaton, Gamze Eroglu, Harriet
	Elsworthy, Claire Forbes-Haley, Ana Beatriz
	Gamboa, Mandeep Ghuman, Nikolaos
	Gkranias, Debbie Hemington, Mark Ide,
	Roshni Karia, Moritz Kebschull, Annika
	Kroeger, Gerry Linden, Emily Lu, Isobel
	Madden, Kathryn Mayo, Claire
	McCarthy, William McLaughlin, Payvand
	Menhadji, Imogen Midwood, Mike
	Milward, Madeleine Murray, Rajan Nansi, Ian
	Needleman, Luigi Nibali, Rupali Patel, Vipul
	Patel, Michael Paterson, Philip M
	Preshaw , Devan Raindi, Raj Rattan, Anthony
	Roberts, Mitul Shah, Shazad Saleem,

	Praveen Sharma, Shakeel Shahdad, Sami
	Stagnell, Claire Storey, Joon Seong, Manoj
	Tank, Joel Thomas, Aradhna Tugnait, Wendy
	Turner, Bobby Varghese, Viren
	Vithlani, Jenny Walker, Natasha West,
	Nicola West, Paul Weston, Matthew
	Wright, Roger Yates, Zehra Yonel.
British Society of Prosthodontics	Claire Forbes-Haley, Gerry McKenna
British Society of Restorative Dentistry	Mark Ide, Claire Storey
European College of Gerodontology	Martina Hayes, Devan Raindi, Mitul Shah
European Organisation for Caries Research	Avijit Banerjee
European Society of Endodontology	Henry Duncan
International Team for Implantology	Colin Burns, Victoria Griffiths, Charlotte
	Stilwell
Osteology Foundation	Elena Calcioari
Restorative Dentistry UK	Claire Forbes-Haley, Mark Ide, Claire Storey
Scottish Antimicrobial Prescribing Group	Andrew Smith
Other organisations involved in the guideline	
development process	
British Dental Journal	Stephen Hancocks
British Society of Periodontology and Implant	Rachael Clampin, Andrew Cundy, Roger
Dentistry Patient Forum	Fisher, Julian Ekiert
<u> </u>	
(Dental Patient Representatives)	(Sally Ferrier, Elaine Judd, Margaret Morgan)
College of General Dentistry	Claire Forbes-Haley, Roshni Karia
Council of European Chief Dental Officers	Kenneth Eaton
Dental Protection	Raj Rattan
General Dental Council (Observers)	Rebecca Cooper, Alina Grossman, Ross
	Scales
NHS England and NHS Improvement	Claire Forbes-Haley, Divyash Patel
NHS Education for Scotland	Isobel Madden

Platform for Better Oral Health in Europe

In addition, the EFP engaged an independent guideline methodologist to advise the panel and facilitate the consensus process (Prof. Dr. med. Ina Kopp [I.K.]). The guideline methodologist had no voting rights.

The EFP and the guideline panel attempted to involve patient forums/organizations but were unable to identify any groups focused on periodontal diseases at a pan-European level. In future updates, efforts will be undertaken to include the perspectives of citizens/patients [23]. National societies will be encouraged to involve patient groups within individual countries as key stakeholders for the *Adaptation, Adoption, De Novo Development* – "ADOLOPMENT" of this CPG [3].

Commentary – UK Implementation

For the development of the UK version of the guideline, a broad range of potential addressees of the guideline from 26 organisations was asked to participate by the BSP.

In contrast to the EFP S3-level guideline, the British version benefitted strongly from the input and advice of the BSP Patient Forum delegates. Delegates were supported throughout the process with 1:1 meetings as appropriate to explain process.

3.2 | Evidence Synthesis

3.2.1 | Systematic search and critical appraisal of guidelines

To assess and utilize existing guidelines during the development of the present guideline, we performed electronic searches in a range of well-established guideline registers and the websites of large periodontal societies:

- Guideline International Network (GIN)
- Guidelinecentral.com
- The National Institute for Health and Clinical Excellence (NICE)
- Canadian Health Technology Assessment (CADTH)
- European Federation for Periodontology (EFP)

- American Academy of Periodontology (AAP)
- American Dental Association (ADA)
- BIGG International database of GRADE guidelines
- ECRI Guidelines Trust
- DynaMed database
- US Preventive Services Task Force
- Scottish Intercollegiate Guidelines Network, Healthcare Improvement Scotland

(SIGN-HIS)

The last search was performed on 13th January 2023. Search terms used were: "implant", "dental implant", "peri-implant*", "guidelines, "clinical practice guidelines". In addition, content was screened by hand searches, see Table 3.

Table 3. Results of the guideline search.

Database	Identified, potentially relevant guidelines	Critical appraisal
Guideline International Network (GIN) International Guidelines Library (#1)	No thematically relevant hits	Not applicable
	Insertion of customised exposed titanium implants,	Focus on oro-facial implants, therefore potentially relevant
The National Institute for Health and Clinical Excellence (NICE) (#2)	without soft tissue cover, for complex orofacial reconstruction	But: Data more than a decade old, does not directly address biological complications
	(July 2013)	Not applicable
	Insertion of customised titanium implants, with soft tissue cover, for orofacial	Focus on oro-facial implants, therefore potentially relevant But: Data more than a decade
	reconstruction (July 2013)	old, does not directly address biological complications Not applicable
	Soft-palate implants for simple snoring	Focus on oral implants, therefore potentially relevant

	(November 2007)	But: Data more than 15 years old, focus on palatal implants, does not directly address biological complications
		Not applicable
		Focus on oral implants, therefore potentially relevant
	Soft-palate implants for obstructive sleep apnoea (November 2007)	But: Data more than 15 years old, focus on palatal implants, does not directly address biological complications
		Not applicable
Guidelinecentral.com "Dentistry" category	Antibiotic Prophylaxis for Prevention of Prosthetic Joint Infection	Does not readily address per- implant diseases
	(January 2015)	Not applicable
	Prevention of Orthopaed c Implant Infection in Patients Undergoing Dental Procedures	Does not readily address per- implant diseases
	(December 2012)	Not applicable
Agency for Healthcare Research and Quality (#3)	No thematically relevant hits	Not applicable
Canadian Health Technology Assessment (CADTH) (#4)	Biological Mesh: A Review of Clinical Effectiveness, Cost- Effectiveness and Guidelines – An Update	Focus on implants in other areas, no direct relation to oral diseases
	(August 2015)	Not applicable
JIC	Osseointegrated Prosthetic Implants for Lower Limb Amputation: A Review of Clinical Effectiveness, Cost- Effectiveness and Guidelines	Focus on implants in other areas, no direct relation to oral diseases
	(February 2017)	Not applicable
	Immediate Osseointegrated Implants for Cancer Patients: A Review of Clinical and Cost- Effectiveness	Focus on dental implants in very specific, selected patient group, peri-implantitis not directly addressed, 7 years old data
	(January 2015)	Not applicable
European Federation of Periodontology (EFP) (#5)	EFP S3-Level Clinical Practice Guideline for Stage I-III Periodontitis	Indirectly applicable, high quality
	EFP S3-Level Clinical Practice Guideline for Stage IV Periodontitis	Indirectly applicable, high quality
American Academy of Periodontology (AAP) (#6)	AAP Best Evidence Consensus: Biologics in Clinical Practice (Oct 2022)	Focus on periodontal defects only – peri-implantitis not addressed

		Not applicable
	AAP Best Evidence Consensus: Periodontal Phenotype (January 2020)	Focus on tissues around teeth, rather than dental implants
		Not applicable
	AAP Best Evidence Consensus: Laser Therapy	Potentially relevant: Two SRs address adjunctive laser use and photo-dynamic therapy, respectively, for peri- implant mucositis and peri- implantitis
	(April 2018)	But: More than four years old, superseded by new SRs in current guideline
	AAP Best Evidence Consensus:	Not directly applicable Does not readily address per-
	Cone-Beam Computed Tomography	implant diseases
	(October 2017)	Not applicable
American Dental Association (ADA) (#7)	No thematically relevant hits	Not applicable
BIGG International database of GRADE guidelines (#8)	Antibiotic prophylaxis is not indicated prior to dental procedures for prevention of periprosthetic joint infections	Does not readily address per- implant diseases
	(2017)	Not applicable
ECRI Guidelines Trust (#9)	No thematically relevant hits	Not applicable
DynaMed (#10)	Anaerobic Bacterial Infections	Does not readily address per- implant diseases
		Not applicable
	Gingivitis and Periodontitis in Adults	Does not readily address per- implant diseases
		Not applicable
3		Potentially applicable, as it addresses an important risk factor
	Oral Healthcare in Persons With Diabetes	But: No specific recommendations, no standardised methodology, no guideline Not applicable
US Preventive Services Task	Dental and Periodontal Disease: Counselling	More than two decades old, does not readily address peri- implant conditions
Force (#11)	(1996)	Not applicable

Scottish Intercollegiate Guidelines Network, Healthcare Improvement Scotland (SIGN- HIS) (#12)	No thematically relevant hits	Not applicable
---	-------------------------------	----------------

#1. https://guidelines.ebmportal.com/

#2. <u>https://www.nice.org.uk/guidance/published?type=csg,cg,mpg,ph,sg,sc</u>

#3. https://www.ahrq.gov/gam/index.html

#4. https://www.cadth.ca/

#5. http://www.efp.org/publications/index.html

#6. https://www.perio.org/research-science/best-evidence-consensus-bec/

- #7. https://ebd.ada.org/en/evidence/guidelines
- #8. https://sites.bvsalud.org/bigg/en/biblio/
- #9. https://www.ecri.org/solutions/ecri-guidelines-trust

#10. https://www.dynamed.com/

#11. https://www.uspreventiveservicestaskforce.org/uspstf/topic_search_results

#12. https://www.sign.ac.uk/our-guidelines/

Only guidelines published in English and with full texts available were included. The methodological quality of these guideline texts was critically appraised using the AGREE II framework (<u>https://www.agreetrust.org/agree-ii/</u>).

We did not identify guidelines/documents directly relevant to the current guideline development process due to: (i) their publication time, (ii) their methodological approach, or (iii) their stated inclusion criteria. We have referenced the EFP S3-level Clinical Practice Guidelines [24, 25] where applicable.

Commentary – UK Implementation

The guideline search was not re-done due to the workshops taking place 7 months after the publication of the EFP paper.

3.2.2 | Systematic search and critical appraisal of the literature

For this guideline, a total of 13 systematic reviews (SRs) were conducted to support the guideline development process [26-38]. The corresponding manuscripts are published within this special issue of the Journal of Clinical Periodontology.

All SRs were conducted following the "Preferred Reporting Items for Systematic Reviews and Meta-Analyses" (PRISMA) framework [39] and were prospectively registered in PROSPERO.

Commentary – UK Implementation

For all systematic reviews, the same search (using the same databases) and screening processes was repeated. If relevant papers were identified, they were retrieved, to confirm inclusion. If that was the cases, risk of bias was assessed and a critical evaluation of the possible influence of the new evidence in the already reported results was made.

3.2.3 | Focused questions

In all 13 systematic reviews, focused questions in PICOS format [40, 41] were proposed by the authors in February-March 2022 to a panel comprising the working group chairs and the methodological consultant in order to review and approve them (Table 4a-d). The panel took great care to avoid overlaps between the SRs or significant thematic omissions in order to ensure that they encompass the main interventions currently undertaken in the management of peri-implant diseases.

Table 4. PICOS questions addressed by each Systematic Review, listed according to working group: (a) Peri-implant health & Prevention; (b) Management of Peri-implant mucositis; (c) Management of Peri-implantitis - non-surgical; (d) Management of Peri-implantitis - surgical.

(a)

leference	Systematic Review title	PICOS question (as written in the original article)
[42]	Primary prevention of peri- implant diseases: A systematic review and meta-analysis	What is the efficacy of preventive interventions, involving risk factor control, in patients i) awaiting dental implant rehabilitation (primordial prevention), or ii) already having dental implant(s) with healthy peri-implant tissues (primar prevention)?
Supportive care for the prevention of disease recurrence/progression following peri-implantitis	 #1. In patients treated for peri-implantitis (P) what is the efficacy of supportive care (I) in comparison with no supportive care (C), in terms of peri-implant tissue stability (O), as reported in prospective and retrospective studies o at least 3 years duration (S). #2. In patients treated for peri-implantitis (P) what is the efficacy of supportive care with adjunctive local antiseptic agents (I) in comparison with supportive care without local antiseptic agents (C), in terms of peri-implant tissue stability (O). 	
	treatment: A systematic review.	 (O), as reported in prospective and retrospective studies of at least 3 years duration (S)? #3. In patients treated for peri-implantitis (P) what is the efficacy of supportive care with a frequency of more than once a year (I) in comparison with supportive care with a frequency of once a year or less (C) in terms of peri-implant tissue stability (O), as reported in prospective and retrospective studies of at least 3 years duration (S)?

26

Reference	Systematic Review title	PICOS question (as written in the original article)
	Non-surgical therapy of peri- implant mucositis –	 #1. In human subjects suffering peri-implant mucositis (P), has professionally administered non-surgical mechanical/physical therapy (I) any effect over no treatment (C), in terms of clinical/radiographic parameters and invasiveness (O), as shown in clinical randomized controlled trials (RCTs) (S)? #2. In human subjects suffering peri-implant mucositis (P), is any single mode of professionally administered non-surgical mechanical/physical therapy (I) superior to other single modes of professionally administered non-surgical mechanical/physical therapy (C), in terms of clinical/radiographic parameters and invasiveness (O), as shown in (RCTs) (S)
[44]	mechanical/physical approaches: a systematic review.	#3. In human subjects suffering peri-implant mucositis (P), are combinations of treatment modes of professionally administered non-surgical mechanical/physical therapy (I) superior to single modes of professionally administered non- surgical mechanical/physical therapy (C), in terms of clinical/radiographic parameters and invasiveness (O), as shown in (RCTs) (S)?
		#4. In human subjects suffering peri-implant mucositis (P), does repetition of professionally administered non-surgical mechanical/physical therapy (I) provide added benefits over single administration (C), in terms of clinical/radiographic parameters and invasiveness (O), as shown in (RCTs) (S)?
[45]	Efficacy of chemical approaches during non-surgical submarginal instrumentation in the management of peri-implant mucositis: a systematic review.	In patients with peri-implant mucositis (P), what is the efficacy of (I) professionally administered topical antibiotics (with unsustained drug release), topical antispetics (hydrogen peroxide, chlorhexidine, delmopinol hydrochloride, sodium hypochlorite, chitosan, acids) or photodynamic therapy during non-surgical submarginal peri-implant instrumentation compared to (C) non-surgical submarginal peri-implant instrumentation with or without additional control/placebo treatment in terms of (O) reduction of bleeding on probing (BOP) in (S) randomized controlled clinical trials (RCTs) controlled clinical trials, prospective and retrospective case-control-studies, and case series with a follow-up of ≥ 3 month
[46]	Efficacy of adjunctive measures in peri-implant mucositis. A systematic review and meta- analysis.	In systemically healthy humans with PiM, what is the efficacy of patient-performed or administered (by prescription) measures used adjunctively to submarginal instrumentation, as compared to submarginal instrumentation alone or combined with a negative control, in terms of reducing bleeding on probing (BOP), in randomized controlled clinical trial (RCTs) with at least 3-month follow up?
	301	

Reference	Systematic Review title	PICOS question (as written in the original article)	
[47]	Efficacy of mechanical/physical approaches for implant surface decontamination in nonsurgical submarginal instrumentation of peri-implantitis. A systematic review.	 #1. In patients with peri-implantitis, what is the efficacy of non-surgical submarginal peri-implant instrumentation with mechanical/physical decontamination methods (e.g. air- polishing, sonic/ultrasonic devices, lasers) alone or combinations thereof, compared to non-surgical submarginal instrumentation with placebo decontamination (non-aiming at mechanical/physical decontamination, e.g., scalers to emove hard deposits with adjunctive saline irrigation) in terms of change in peri-implant PD and/or change in BOP, in parallel-arm and split-mouth RCTs with ≥ 10 recruited/randomized subjects per treatment arm, in controlled clinical trials and prospective cohort-studies with ≥ 30 recruited subjects with ≥ 6 months duration? #2. In patients with peri-implantitis, what is the efficacy of non-surgical submarginal peri-implant instrumentation with mechanical/physical decontamination methods (e.g. air- polishing, sonic/ultrasonic devices, lasers) alone or combinations thereof and additional measures/interventions (e.g. irrigation with antiseptics), compared to non-surgical submarginal instrumentation with placebo decontamination (non-aiming at mechanical/physical decontamination decontami	
[48]	Efficacy of chemical approaches for implant surface decontamination in conjunction with sub-marginal instrumentation, in the non- surgical treatment of peri- implantitis. A systematic review and meta-analysis.	In adult patients with peri-implantitis (P), what is the efficacy of sub-marginal instrumentation combined with chemica surface decontamination (I) in comparison with sub- marginal instrumentation with or without placebo (C), in terms of changes in probing depths (PD) and/or bleeding on probing (BOP) (O), as reported in randomized clinical trials (RCT), nonrandomized controlled clinical trials (CCT) or prospective cohort studies, with a minimum of 6-month "follow-up" (S)?	
[49]	Efficacy of adjunctive measures in the non-surgical treatment of peri-implantitis. A systematic review.		

(d)

cacy of access flap and pocket imination procedures in the nagement of peri-implantitis – systematic review and meta- analysis	#1. In patients requiring treatment of peri-implantitis (P), what is the effect of surgical therapy including access flap or pocket elimination procedures (I), when compared to non-surgical therapy (C), in terms of reduction of probing depth (PD) and/or of bleeding on probing (BOP) (O), as observed in randomized controlled trials with a follow-up of ≥6 months and a sample size of ≥10 patients per arm (S)? #2. In patients requiring treatment of peri-implantitis, what are the long term outcomes of surgical access flap or
•	#2. In patients requiring treatment of peri-implantitis, what are the long-term outcomes of surgical access flap or
a systematic review and meta- analysis	pocket elimination procedures based on prospective studies (interventional or observational) with a sample of \geq 20 patients and a follow-up of \geq 12 months?
The efficacy of bone constructive therapies in the nagement of peri-implantitis.	#1. In patients with peri-implantitis, what is the efficacy of different bone reconstructive therapies compared to acces flap surgery in terms of pocket reduction and change in bleeding/suppuration on probing, at a minimum of 12-month of follow-up?
•	#2. In patients with peri-implantitis, what is the long-term (≥12 months) performance of reconstructive therapies in terms of pocket reduction, change in bleeding on probing/suppuration?
Mechanical and physical implant surface decontamination approaches in conjunction with surgical peri-implantitis treatment: A systematic review	#1. In patients with peri-implantitis (population), what is the efficacy of adjunctive or alternative mechanical/physical measures for implant surface decontamination in conjunction with surgical peri-implantitis treatment (intervention) compared with standard surface instrumentation (comparison) in changing signs of inflammation (outcomes), as reported in RCTs and CCTs with a follow-up period of at least 6 months (study design)?
	#2. In patients with peri-implantitis (population), what is the efficacy of adjunctive or alternative mechanical/physical measures for implant surface decontamination in conjunction with surgical peri-implantitis treatment (intervention) compared with standard surface instrumentation including additional measures performed for both test and control groups (e.g., local application of antimicrobials and/or additional mechanical/physical measures) (comparison) in changing signs of inflammation (outcomes), as reported in RCTs and CCTs with a follow-up period of at least 6 months (study design)?
e Efficacy of Implant Surface Decontamination Using Chemicals during Surgical eatment of Peri-implantitis: A stematic Review and Meta- Analysis.	In adult patients with peri-implantitis, what is the efficacy of surgical therapy with adjunctive chemical surface decontamination of implant surfaces in comparison with surgical therapy alone or with placebo, in terms of probing depth (PD) reduction and bleeding on probing (BoP)/suppuration on probing (SoP) as reported in randomized (RCTs) and non-randomized controlled clinical trials (non-RCTs) with a follow-up of at least 6 months?
Adjunctive locally and systemically delivered ntimicrobials during surgical eatment of peri-implantitis.	In patients with peri-implantitis, what is the efficacy of surgical therapy combined with systemic or local antimicrobial in comparison with surgical therapy alone, in terms of pocket probing depth reduction, as assessed in randomized controlled trials (RCTs) with at least 6 months of follow-up?
	nagement of peri-implantitis. ystematic review and meta- analysis chanical and physical implant surface decontamination proaches in conjunction with surgical peri-implantitis atment: A systematic review e Efficacy of Implant Surface Decontamination Using Chemicals during Surgical atment of Peri-implantitis: A stematic Review and Meta- Analysis. Adjunctive locally and systemically delivered timicrobials during surgical

3.2.4 | Relevance of outcomes

For the present guideline, the recommendations of the "Implant Dentistry Core Outcome Set and Measurements" (ID-COSM) initiative were followed [55-58], specifically the conclusions of the systematic review dealing with the outcome measures used in clinical studies [55]. As expected, and since the report of the strongest outcome (dental implant/implant-supported prosthesis survival) was not frequently found, surrogate parameters were selected, in parallel with the previous EFP guidelines on the treatment of periodontitis [24, 25].

The primary outcomes selected were parameters capturing the inflammatory component of the peri-implant tissues: probing depths (PD) and bleeding on probing (BOP)/suppuration on probing (SOP), since they were the most consistently reported outcomes.

The selected secondary outcomes were radiographic marginal bone loss (MBL), composite outcomes including the primary outcomes and MBL, dental implant/implant-supported prosthesis survival/loss, and patient-reported outcome measures (PROMs).

3.2.5 | Search strategy

All SRs utilized a comprehensive search strategy of at least two different databases, supplemented by a hand search of periodontology-focused journals and the reference lists of included studies. In all SRs, the electronic and manual search, as well as the data extraction, was undertaken in parallel by two or more investigators.

3.2.6 | Quality assessment of included studies

In all SRs, the risk of bias of controlled clinical trials was assessed using the Cochrane Risk-of bias tool (https://methods.cochrane.org/bias/resources/rob-2-revised-cochrane-risk-bias-tool-randomized-trials). For observational studies, the Newcastle-Ottawa-Scale was used (http://www.ohri.ca/programs/clinical_epidemiology/ oxford.asp).

3.2.7 | Data synthesis

Where applicable, the available evidence was summarized by means of a meta-analysis.

3.3 | From Evidence to Recommendation: Structured Consensus Process

The structured consensus development conference was held during the XVIII European Workshop in Periodontology in La Granja de San Ildefonso Segovia, Spain, on November 6th-9th, 2022. Using the 13 SRs as background information, evidence-based recommendations were formally debated by the guideline panel using the format of a structured consensus development conference. This consisted of small group discussions and open plenary discussions, where the proposed recommendations were presented, voted upon and adopted by consensus [59]. Delegates with conflicts of interest abstained from voting and abstentions were recorded. Prior to the in-person meeting, three online meetings were organized (one at the plenary level, and two at the working group level) in September and October 2022, to advance the process of guideline development to a mature stage prior to the face-to-face consensus meeting.

In the small group phase, delegates convened in four working groups (WGs) directed by twothree chairpersons belonging to the EFP Workshop Committee, addressing the following subtopics:

- WG #1. Peri-implant health & Prevention (chairs lain Chapple and Søren Jepsen).
- WG #2. Management of Peri-implant mucositis (chairs Mariano Sanz and Anton Sculean).
- WG #3. Management of Peri-implantitis non-surgical (chairs David Herrera, Moritz Kebschull and Maurizio Tonetti).
- WG #4. Management of Peri-implantitis surgical (chairs Tord Berglundh, Panos Papapanou and Frank Schwarz).

With the support of the methodology expert, recommendations and draft background texts were generated and subsequently presented, debated, and subjected to a vote in the plenary sessions with all delegates present. During these plenary sessions, the guideline development process and discussions and votes were overseen and facilitated by the independent guideline methodologist (I.K.). The plenary votes were recorded using an electronic voting system, checked for accuracy, and then introduced into the guideline text.

The consensus process was conducted as follows:

3.3.1 | Plenary session 1 (online session, 1st September 2023)

Introduction to guideline methodology (presentation, discussion) by the independent guideline methodologist (I.K.) and the chair of the workshop (D.H.).

3.3.2 | Working groups 1-4 (online sessions, 18th October 2023 – 20th November 2023)

- Initial evaluation of declarations of interest and management of conflicts of interest.
- Presentation of the evidence (SR results) by group chairs and reviewers.
- Invitation of all members of the working group to reflect critically on the quality of available evidence by group chairs, considering the GRADE criteria.
- Structured group discussions:
 - initial discussions for the development of draft recommendations and their grading, considering the GRADE-criteria.
 - initial discussions for the development of draft background texts, considering the GRADE-criteria.
 - invitation to comment on draft recommendations and background text to suggest reasonable amendments by group chairs.
 - o collection and merging of amendments by group chairs.

3.3.3 | Plenary session 2 (online session 21st November 2023)

- Presentation of working group results (draft recommendations and background text) by Working Group chairs.
- Invitation to formulate questions, statements, and reasonable amendments of the plenum by the independent guideline methodologist /facilitator.
- Answering questions by working group chairs.
- Collection and merging of amendments by an independent moderator.
- Preliminary vote on all suggestions provided by the working groups and all reasonable amendments.
- Assessment of the strength of consensus.
- Recording of abstentions made due to potential conflicts of interest.
- Opening debate, where no consensus was reached or reasonable need for discussion was identified.

- Formulation of tasks to be solved within the working groups.
- Discussion of tasks and potential amendments raised by the plenum.
- Formulation of reasonable and justifiable amendments, considering the GRADE framework.
- Initial voting within the working group on recommendations and guideline text in preparation for the plenary session.

3.3.4 | Plenary session 2 (online session, 23rd November 2023)

- Presentation of working group results by working group chairpersons.
- Invitation to formulate questions, statements, and reasonable amendments of the plenary by the independent moderator.
- Collection and merging of amendments by an independent moderator.
- Preliminary vote.
- Assessment of the strength of consensus.
- Opening debate, where no consensus was reached or reasonable need for discussion was identified.
- Formulation of reasonable alternatives.
- Final vote of each recommendation, recording the consensus and abstentions due to potential conflicts of interest.

Commentary – UK Implementation

The UK implementation process followed the process described above.

The entire implementation process was performed using a video conference system.

In brief, the following steps were performed:

- 1. Assess necessity of updating searches for potential new guidelines and evidence
- 2. Critical assessment of all clinical recommendations following the GRADE ADOLOPMENT process
- 3. Draft proposals for the implementation of each clinical recommendation were introduced at working group level
- 4. Working group meeting with external, independent moderator and formal consensus process
- 5. Plenary sessions with formal consensus process and final voting

3.4 | Definitions: Rating the Quality of Evidence, Grading the Strength of Recommendations and Determining the Strength of Consensus

For all recommendations and statements, this guideline makes transparent:

- the underlying quality of evidence, reflecting the degree of certainty / uncertainty of the evidence and robustness of study results.
- the grade of the recommendation, reflecting the criteria considered to make the judgement; the strength of consensus, indicating the degree of agreement within the guideline panel; the number of abstentions due to potential conflicts of interest.

3.4.1 | Quality of Evidence

The quality of evidence was assessed using a recommended rating scheme [60, 61].

3.4.2 | Strength of Recommendations

The grading of the recommendations used the grading scheme (Table 5) by the German Association of the Scientific Medical Societies (AWMF) & Standing Guidelines Commission, [62], taking into account not only the quality of evidence, but also considering a judgement guided by the following criteria:

- relevance of outcomes and quality of evidence for each relevant outcome
- consistency of study results
- direct applicability of the evidence to the target population/PICOS specifics
- precision of effect estimates using confidence intervals
- magnitude of the effects
- balance of benefit and harm
- ethical, legal, economic considerations
- patient preferences

Table 5. Strength of Recommendations: Grading Scheme [63].

Grade of recommendation grade*	Description	Syntax
А	Strong recommendation	We recommend (↑↑) /
		We recommend not to ($ ightarrow ightarrow$)
В	Recommendation	We suggest to (↑) /
		We suggest not to ($ ightarrow$)
0	Open recommendation	May be considered (↔)

*If the group felt that evidence was not clear enough to support a recommendation, Statements were formulated, including the need (or not) for additional research.

Journal Pression

The grading of the quality of evidence and the strength of a recommendation may therefore differ, but where they do, the justification and context is clearly documented in the background narrative that follows each recommendation table.

Commentary – UK Implementation The aforementioned procedures and criteria have also been used for the UK implementation.

3.4.3 | Strength of Consensus

The consensus determination process followed the recommendations by the (German Association of the Scientific Medical Societies (AWMF) & Standing Guidelines Commission, 2012) [62]. Where consensus could not be reached, different points of view were documented in the guideline text (see Table 6).

Table 6. Strength of Consensus: Determination Scheme [63].

Unanimous consensus	Agreement of 100 % of participants
Strong consensus	Agreement of > 95 % of participants
Consensus	Agreement of 75 – 95 % of participants
Simple majority	Agreement of 50 – 74 % of participants
No consensus	Agreement of <50 % of participants

Con	Commentary – UK implementation										
The	aforementioned	procedures	and	criteria	have	also	been	used	for	the	UK
imp	lementation.										

3.5 | Editorial Independence

3.5.1 | Funding of the guideline

The development of this guideline and its subsequent publication was financed entirely by internal funds of the European Federation of Periodontology (EFP), without any support from industry or other organisations.

Commentary – UK Implementation

The UK implementation has been funded entirely by funds from the BSP.

3.5.2 | Declaration of Interests and Management of Conflicts

All members of the guideline panel declared secondary interests using the standardized form provided by the International Committee of Medical Journal Editors (ICMJE) (International Committee of Medical Editors).

Management of conflicts of interests (Cols) was discussed in the working groups and the plenary sessions, following the principles provided by the Guidelines International Network [64]. According to these principles, panel members with relevant, Cols abstained from voting on guideline statements and recommendations within the consensus process. Those abstentions were recorded in each recommendation table.

Commentary – UK Implementation

All participants of the implementation process were asked to declare their interests using the ICMJE form, as outlined above.

Summarised accounts of the interests can be found in section 9.

3.6 | Peer review

All 13 systematic reviews (SR) underwent a multi-step peer review process. First, the draft documents were evaluated by members of the EFP Workshop Committee and the methodological consultants using a custom-made appraisal tool to assess: (i) the methodological quality of the SRs using the AMSTAR 2 checklist [65] and (ii) whether all PICOS questions were addressed as planned. Detailed feedback was then provided for the SR authors. Subsequently, all 13 systematic reviews underwent the regular editorial peer review process defined by the *Journal of Clinical Periodontology*.

The guideline text was drafted by the chairs of the working groups, in close cooperation with the methodological consultant, and circulated among the members of the guideline group prior to the Workshop. The methodological quality was formally assessed by an external consultant

using the AGREE framework. The guideline was subsequently peer-reviewed for its publication in the *Journal of Clinical Periodontology* following the standard evaluation process of the journal.

Commentary – UK Implementation

The UK implementation has been peer-reviewed by an external, independent dental scientist with experience in S3-level guideline development.

3.7 | Implementation and dissemination plan

For this guideline, a multi-stage dissemination and implementation strategy will be established and implemented by the EFP, supported by a communication campaign.

This will include:

- Publication of the guideline and the underlying systematic reviews as an Open Access special issue of the *Journal of Clinical Periodontology*.
- Commentary, Adoption, or Adaptation [3] by national societies.
- Generation of educational material for dental professionals and patients, and dissemination via the EFP member societies.
- Dissemination via educational programs at dental conferences.
- Dissemination via the EFP through European stakeholders via National Society members of the EFP.
- Long-term evaluation of the successful implementation of the guideline by a survey of EFP members.

The timeline of the guideline development process is detailed in Table 7.

Time point	Action
April 2018	Decision by European Federation of Periodontology (EFP)
	General Assembly to develop comprehensive treatment
	guidelines for periodontitis and peri-implant diseases

May - September 2018	EFP Workshop Organizing Committee (WOC) assesses merits		
	and disadvantages of various established methodologies and		
	their applicability to the field		
November 2021	EFP WOC decides on (i) topics covered by proposed guideline,		
	(ii) working groups and chairs, (iii) systematic reviewers, and (iv)		
	outcome measures		
February 2022	EFP WOC decides invited systematic reviewers		
March 2022	Decision on consensus group, invitations sent to participants		
	invitations sent to stakeholders		
March 2022	Submission of PICO(S) questions by systematic reviewers to		
	group chairs for internal alignment		
March 30 th , 2022	Online meeting with consultant, WOC and reviewers, to better		
	define PICOS.		
	Final decision by WOC on PICOS		
April 2022	Decision on PICO(S) and information sent to reviewers		
June - August 2022	Submission of Systematic reviews to WOC by the reviewers,		
	initial quality assessment		
August - September 2022	Submission to Journal of Clinical Periodontology, peer review		
	and revision process		
September – December	Peer review and revision process in Journal of Clinical		
2022	Periodontology		
September 26 th , 2022	Online plenary meeting		
September 28 th , 2022	Online working group meetings		
September – October 2022	Submission of declarations of interest by all delegates		
October 19 th , 2022	Online working group meetings		
October 2022	Electronic circulation of reviews		
November 6 th -9 th , 2022	Workshop in La Granja with moderated formalized consensus		
	process		
November 2022 – January	Formal stakeholder consultation, finalise guideline method,		
2023	report and background text		
January 18 th , 2023	Online Plenary meeting		
February 2023	Submission of guideline document to the Journal of Clinical		
	Periodontology		
April 2023	Publication of guideline and underlying Systematic Reviews in		
	the Journal of Clinical Periodontology		
April-September 2023	Processes of adaptation/adoption by National Societies		

Commentary – UK Implementation				
Time point	Action			
June 2023	Decision by the BSP executive board to implement the EFP S3 guideline in the United Kingdom using the GRADE ADOLOPMENT framework. Invitation of project leads and chairpersons, methodologists and independent moderators (Prof. Ina Kopp), and stakeholder organisations. Assessment of potential Cols for all guideline group members.			
July 2023	Update of the 13 systematic review searches assessed. This was not needed regarding timelines			
September 2023?	Assessment of the updated evidence base and the clinical recommendations of the original guideline document by the group chairs. Draft of possible adaptations. Distribution of the draft clinical recommendations within the working groups.			
October 2023 - November 2023	Working group phase – Three working groups prepared adoloped clinical recommendations with independent moderation and formal CoI management. Subsequently, all recommendations from all working groups were circulated to all guideline group members for critical assessment.			
21 st & 23 rd November 2023	Plenary session with independent moderation and formalised consensus process (virtual)			
February 2024	Final editing			
February 2024	Formal approval by all stakeholders and involved organisations			
February 2024	Submission to Journal of Dentistry for peer review			
March 2024	Publication and start of the dissemination phase (articles, webinars, interactive materials			

3.8 | Validity and update process

The guideline is valid until 2028. However, the EFP, represented by the members of the Organizing Committee, will continuously assess current developments in the field. Where there

are major changes of circumstances, e.g., new relevant evidence, this will trigger an update of the guideline to potentially amend the recommendations. It is planned to update the current guideline regularly on demand and consistent with the format of a living guideline.

Commentary – UK Implementation

The UK implementation will be updated at regular intervals, following the updates of the underlying supra-national EFP S3-guideline. The authors of the implementation are in close contact with the EFP Workshop Committee to facilitate this update process.

4 | MANAGEMENT OF PERI-IMPLANT DISEASES – PREVENTION, DIAGNOSIS AND TREATMENT SEQUENCE

4.1 |Specific approaches in the management of peri-implant diseases

Dental implants and dental implant abutments are class lib medical devices [66], according to the 1993 Medical Device Directive (MDD, 93/42/EEC), which is maintained in the 2017 Medical Device Regulation (MDR, Council Regulation 2017/745) [67]. This class of medical devices considers "implantable devices and long-term (> 30 days) surgically invasive devices", and applies to most implants used in the orthopaedic, dental, ophthalmic, and cardiovascular fields. Implantable devices are "partially introduced into the human body through surgical intervention and intended to remain in place after the procedure for at least 30 days" [66]. They can be further classified according to their expected "duration", either as short-term (normally intended for continuous use for not more than 30 days), or long-term (normally intended for continuous use for more than 30 days). In the current MDR regulation from the European Parliament and the Council of the European Union, published in 2017 [67], and enforced in May 2022, dental implants and dental implant abutments are considered within the category MDN 1103 (non-active dental implants and dental materials) as "non-active implants and long term surgically invasive devices" [67]. Other non-active implants are classified in different categories as "non-active cardiovascular, vascular and neurovascular implants" (MDN 1101), "non-active osteo- and orthopaedic implants" (MDN 1102) and "non-active soft tissue and other implants" (MDN 1104).

When developing a clinical practice guideline (CPG) related to dental implants (in the present case, on the management of peri-implant conditions), the CPG structure could be based on

similar guidelines on other "long-term surgically invasive devices"; however, the clinical use of dental implants has a fundamental difference, since these medical devices are partially inserted in the jaws. Since the oral cavity is one of the most diverse and microbially abundant niches in the human body [68], the intraoral part of the implant will always be exposed to this contaminated environment. Therefore, dental implants have been specifically designed to withstand biofilm formation on the non-shedding transmucosal abutment surface, which will be covered by the appropriate prosthetic devices to serve as tooth replacements, then subject to the same measures of infection prevention control as natural teeth (oral hygiene practices). Another strategy that could have been followed in the development of this guideline was to implement a parallel process to that undertaken for the treatment of periodontal diseases [24, 25]. However, the major anatomical and histological differences between periodontal and peri-implant tissues (reported in Section 1.1.2) and the histopathological dissimilarities between periodontitis and peri-implantitis lesions [5, 6, 9] necessitated a different approach.

The structure of the present guideline, therefore, must recognize the specific features of the "implantable medical devices" and the biological distinctions between the peri-implant and periodontal diseases. Specifically, interventions for the prevention and treatment of peri-implant diseases may be implemented prior to inserting the medical device (dental implant), at the time of placement and restoration (implant/prosthesis placement), as well as post-rehabilitation, in recognition of the high incidence of peri-implant diseases.

Consequently, interventions were first organized according to the stage of implant therapy, applicable to:

- patients awaiting dental implant rehabilitation
- patients receiving dental implant rehabilitation
- patients already rehabilitated using dental implant(s)

Subsequently, interventions were organized according to the clinical status of the peri-implant tissues:

- before dental implant placement
- healthy peri-implant tissues

- peri-implant mucositis
- peri-implantitis
- following treatment of peri-implant mucositis
- following treatment of peri-implantitis

This guideline has been organised into interventions following these different stages of periimplant tissue management:

- Risk factor control before implant placement
- Risk factor control during implant/prosthesis placement
- Maintenance of peri-implant tissue health
- Treatment of peri-implant mucositis
- Treatment of peri-implantitis (non-surgical)
- Treatment of peri-implantitis (surgical)
- Secondary prevention of peri-implant mucositis
- Secondary prevention of peri-implantitis

4.2 | Management according to the stage of implant therapy

Three different clinical scenarios exist (Figure 1):

- patients awaiting dental implant rehabilitation (pre-operative)
- patients receiving dental implant rehabilitation (peri-operative)
- patients already having dental implant/s (post-operative)

Figure 1. Management of peri-implant diseases, according to the stage of implant therapy.

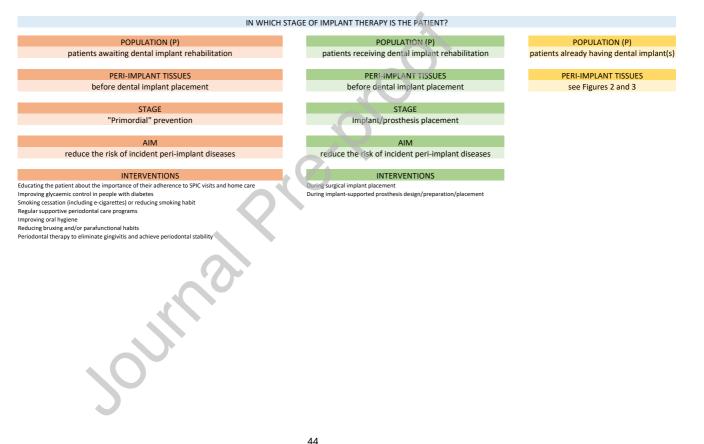


Figure 2. Management of peri-implant diseases, according to the diagnosis of the peri-implant condition: *healthy peri-implant tissues* and *peri-implant mucositis*.

PERI-IMPL	ANT CONDITION DIAGNOSIS - healthy peri-implant tissues and peri-implant m	ucositis				
POPULATION (P)	POPULATION (P)	POPULATION (P)				
patients already having dental implant(s)	patients already having dental implant(s)	patients already having dental implant(s)				
PERI-IMPLANT TISSUES	PERI-IMPLANT TISSUES	PERI-IMPLANT TISSUES				
healthy peri-implant tissues	patients with peri-implant mucositis	patients treated for peri-implant mucositis				
STAGE	STAGE	STAGE				
Primary prevention	Peri-implant mucositis treatment	Secondary prevention of peri-implant mucositis				
AIM	AIM	AIM				
reduce the risk of incident peri-implant diseases	bleeding on probing reduction	not directly assessed				
INTERVENTIONS	INTERVENTIONS	INTERVENTIONS				
vision of regular supportive care sation of agents moking gmentation of peri-implant soft tissues aroved oral hygiene bucing bruxing/parafunctional habits	Professionally administered adjunctive topical antibiotics with un-sustained release Professionally administered adjunctive topical antiseptics Photodynamic therapy used adjunctively. Professionally administered (prome used) probiotics Self-administered (prome used) probiotics Self-administered (or alom tratication by prescription) systemic antibiotics Self-administered (prome used) probiotics Self-administered (prome used) probiotics Self-admini					
	45					

Figure 3. Management of peri-implant diseases, according to the diagnosis of the peri-implant condition: peri-implantitis.

PERI-IMPLANT CONDITION DIAGNOSIS - peri-implantitis

patients already having dental implant(s) PERI-IMPLANT TISSUES patients with peri-implantitis

POPULATION (P)

STAGE Peri-implantitis treatment (non-surgical step of peri-implantitis therapy)

> AIM inflammation reduction (PD, BoP, SoP)

INTERVENTIONS Non-surgical submarginal instrumentation - mechanical/physical cleaning/decontamination Submarginal instrumentation with lasers Submarginal instrumentation with lasers Submarginal instrumentation with air-polishing Non-surgical submarginal instrumentation - chemical approaches for cleaning/decontamination Submarginal instrumentation with antimicrobial photodynamic therapy

Submarginal instrumentation with animulation approximate to Submarginal instrumentation with antiseptic desicant solution Non-surgical submarginal instrumentation – adjunctive therapies Adjunctive systemically administered antimicrobials Adjunctive systemically administered antibiotics Adjunctive probiotics

- ournal

POPULATION (P) patients already having dental implant(s) PERI-IMPLANT TISSUES patients with peri-implantitis STAGE Peri-implantitis treatment (surgical step of peri-implantitis therapy) AIM inflammation reduction (PD, BoP, SoP) INTERVENTIONS

INTERVENTIONS Access flap or resective procedures Reconstructive approaches Additional methods for implant surface decontamination Photo-metanical and physical implant surface decontamina-Chemical implant surface decontamination procedures Adjunctive used flocil/systemic antimicrobials Adjunctive systemically administered antibiotics Adjunctive locally administered antibiotics ation procedures



Regular supportive peri-implant care (SPIC) Professional mechanical plaque removal (PMPR) Specific oral hygiene instructions (OHI) Adjunctive local antiseptic agents in SPIC

46

4.2.1 | Pre-operative interventions

Due to the high prevalence of peri-implant diseases (described in Section 1), any patient receiving dental implants should be considered at risk of developing some form of peri-implant disease. Once the dental implant/abutment complex is exposed to the oral environment, and once the dental implant has been prosthetically loaded and is in function, biofilms can accumulate on their surface, and the ensuing inflammatory process can lead to the onset of peri-implant diseases. Therefore, interventions to prevent peri-implant diseases should commence during the treatment plan stage and continue during implant placement and prosthetic rehabilitation. These pre-operative interventions should focus on controlling the known risk factors associated with the development of peri-implant diseases, such as smoking, diabetes, uncontrolled or untreated periodontitis, and inadequate oral hygiene practices. These interventions are described in Section 5, and the term "primordial" prevention of peri-implant diseases refers to those interventions that can be implemented at the treatment plan stage and target the above risk factors. The concept of "primordial" prevention was first introduced by Strasser in 1978 [69], as prevention attained through a self-directed lifestyle that precludes the development of risk factors in a population. More recently, the American Heart Association [70], has defined the term on a population-wide basis, where primordial prevention is conceived as a strategy to prevent whole societies from experiencing epidemics, while the corresponding strategy on the individual level is to prevent the development of risk factors, consistent with the use of the term in the present guideline, as described in Section 5.

4.2.2 | Peri-operative interventions

There is evidence in the scientific literature that "dental implants placed under less than ideal circumstances" are often encountered in day-to-day practice [6], which may result in an increased prevalence of peri-implantitis [5]. There is also evidence that prosthetic factors may also increase the risk of onset/progression of peri-implant diseases [6]. In fact, the consensus report from the 2017 Workshop on the Classification of Periodontal and Peri-implant diseases stated that "there is some limited evidence linking peri-implantitis to factors such as the post-restorative presence of submucosal cement and the positioning of implants in a manner that does not facilitate oral hygiene and maintenance" [5].

Based on these facts, prevention of peri-implant diseases must also be a focus when:

- placing the dental implant, i.e., aiming at optimal implant positioning and considering local factors preventing an ideal placement.
- designing and installing the prosthetic reconstruction, i.e.., considering local risk factors that may prevent access for oral hygiene, or if possible, electing screw-retained restorations.

4.2.3 | Post-operative interventions

Once the implants have been exposed to the oral environment, and the prosthetic reconstruction has been installed and is in function, the clinical condition of the peri-implant tissues should guide its management. Given the reported high incidence/prevalence of peri-implant diseases (described in Section 1), patients should be immediately enrolled into a supportive peri-implant care (SPIC) program. SPIC programs should include interventions for primary prevention of peri-implant diseases, such as professional supra- and sub-marginal plaque biofilm removal and oral hygiene motivation and coaching, as well as early detection of pathological conditions.

4.3 | Diagnosis of peri-implant conditions

Successful implant-supported rehabilitation requires enrolment in a SPIC, where patients are routinely assessed to facilitate early diagnosis of peri-implant diseases.

The 2018 World Workshop on the Classification of Periodontal and Peri-Implant Diseases and Conditions [5, 22] has established clear case definitions for peri-implant health [71], periimplant mucositis [4] and peri-implantitis [6].

4.3.1 | Diagnosis of healthy peri-implant tissues

According to this 2018 classification [5, 71], a diagnosis of peri-implant health requires:

- Absence of clinical signs of inflammation.
- Absence of bleeding or suppuration on gentle probing.
- No increase in probing depth compared to previous examinations.

• Absence of bone loss beyond crestal bone level changes resulting from initial bone remodelling.

The present guideline has also adopted the recent ID-COSM initiative consensus [58] and the slightly modified definition of peri-implant health, that allows for the presence of a single bleeding spot around the implant.

4.3.2 | Diagnosis of peri-implant mucositis

For a diagnosis of peri-implant mucositis, the 2018 classification requires [4, 5]:

Presence of bleeding and/or suppuration on gentle probing with or without increased probing depth compared to previous examinations.

• Absence of bone loss beyond crestal bone level changes resulting from initial bone remodelling.

Following the modification of the ID-COSM initiative consensus [58], this definition has been updated as follows: presence of bleeding (more than one spot at a location around the implant or presence of a line of bleeding or profuse bleeding at any location) and/or suppuration on gentle probing, in the absence of bone loss beyond crestal bone level changes resulting from initial bone remodelling.

4.3.3 | Diagnosis of peri-implantitis

A diagnosis of peri-implantitis requires [5, 6]:

Presence of bleeding and/or suppuration on gentle probing.

- Increased probing depth compared to previous examinations.
- Presence of bone loss beyond crestal bone level changes resulting from initial bone remodelling.

However, in the absence of previous examination data, the diagnosis of peri-implantitis can be based on the combination of:

- Presence of bleeding and/or suppuration on gentle probing.
- Probing depths of ≥ 6 mm.

 Bone levels ≥3 mm apical of the most coronal portion of the intraosseous part of the implant.

4.4 | Specific care pathways according to diagnosis of the peri-implant condition

Almost 25 years ago, at the Food and Drug Administration (FDA)/AAP consensus conference in 1997, Lang and co-workers [72-74] proposed the Cumulative Interceptive Supportive Therapy (CIST) concept for the management of peri-implant diseases. This protocol was based on a combination of early detection, and implementation of preventive and therapeutic interventions, aimed first to prevent the onset, and then to treat peri-implantitis as early as possible to arrest its progression and thus prevent loss of the implant. Whist the interventions recommended in the current guideline are different, the overall strategy and philosophy are similar.

Depending on the clinical diagnosis, distinct care pathways can be followed (Figures 2 and 3). However, the important overarching principle portends that peri-implant mucositis is treatable and leads to the restoration of peri-implant tissue health. Therefore, primary prevention of periimplant diseases and secondary prevention of peri-implant mucositis (after peri-implant mucositis treatment) share identical interventions. Moreover, since the treatment of periimplant mucositis is the primary intervention in the prevention of peri-implantitis, this treatment should also be considered a preventive strategy. The maintenance of health and function of dental implants and the associated implant-supported prostheses through prevention and treatment of peri-implantitis is, therefore, the primary aim of this guideline. However, once periimplant tissue support, even if the inflammation is successfully controlled. Therefore, specific clinical definitions following the treatment of peri-implantitis need to be established.

4.4.1 | Specific care pathways in healthy peri-implant tissues

In cases of peri-implant tissue health, interventions for primary prevention should be implemented as part of a SPIC program, including periodical professional supra-and sub-marginal plaque biofilm removal.

4.4.2 | Specific care pathways in peri-implant mucositis

Interventions for the management of peri-implant mucositis are detailed in Section 6 and focus on biofilm control, either self-administered or professionally delivered. Treatment outcomes should be evaluated after 2-3 months, and if relevant endpoints have not been achieved, retreatment is recommended. These endpoints reflect the re-establishment of peri-implant health; if peri-implant health is re-established, then the primary prevention of peri-implant diseases and the secondary prevention of peri-implant mucositis are essentially identical. Furthermore, since the treatment of peri-implant mucositis is central to the prevention of the onset of peri-implantitis [75], this treatment is in fact the most important preventive intervention for peri-implantitis and, as such, represents the main component of professional interventions during SPIC.

4.4.3 | Specific care pathways in peri-implantitis

Once a diagnosis of peri-implantitis has been established, two points must be recognized:

- Peri-implantitis is an irreversible condition; therefore, even after successful periimplantitis therapy, a diagnosis of "stable" peri-implantitis is assigned at the particular implant.
- Peri-implantitis treatment outcomes depend upon a multitude of factors (implant and prosthetic characteristics, patient factors, local factors, disease severity, bone defect configuration). Consequently, customized interventions specifically targeting one or several of the above factors are used in its management (as reported in the systematic reviews). The treatment outcomes of these interventions are variable.

Based on these care pathways, the management of peri-implantitis should encompass the following steps:

- Upon diagnosis, a decision must be made whether the affected implant is treatable.
- If so, an initial *non-surgical therapy step*, that includes sub-marginal instrumentation, is performed.
- Following the non-surgical step, re-evaluation of clinical outcomes, based on a set of pre-established criteria for success, will guide the decision whether to enrol the patient

in a secondary prevention SPIC program, or to proceed with the *surgical step*, provided the affected implant continues to be deemed treatable.

- The *surgical step* of peri-implantitis treatment must always include sub-marginal instrumentation after elevating a surgical flap.
- Following evaluation of clinical outcomes after the surgical step, and provided that a set
 of pre-established criteria for success are met, the patient is enrolled into a secondary
 prevention SPIC program. If these criteria are not fulfilled, and the affected implant is
 still deemed to be maintainable, the implant should be re-treated.
- SPIC programs for secondary prevention following peri-implantitis treatment may be different from programs designed for primary prevention.

4.5 | Key aspects in the management of peri-implant diseases

In addition to the chronological flow of interventions (see Figure 4) and the different steps of therapy depending on the specific peri-implant condition diagnosed, we highlight the following **key messages**:

- Appropriate interventions for the preservation and/or restoration of peri-implant tissue health should be considered before, in conjunction with, and after the placement of dental implants.
- Risk factor assessment and control, and diagnosis and monitoring of the health/disease status of the peri-implant tissues, are critical in selecting the appropriate care pathway for the individual patient.
- Successful, long-term maintenance of peri-implant tissue health encompasses behavioural modification, health monitoring, appropriate preventive interventions and, when necessary, careful treatment planning and execution.
- Peri-implant tissue health, peri-implant mucositis, and peri-implantitis represent a continuum. Changes are driven by inflammatory changes subsequent to microbial biofilm accumulation. Controlling inflammation through removal of the plaque biofilm is key to both preserving health and preventing and treating peri-implant diseases.
- Preventive and treatment interventions are organized into specific needs-based care pathways.

- Prevention aims to attain and preserve peri-implant tissues that are free of clinical inflammation. This is achieved by enabling adequate self-performed and professionally delivered oral hygiene measures that need to be customized according to the design of implant-supported restorations.
- Supportive peri-implant care is an essential component of implant dentistry; it is critical for
 preserving peri-implant tissue health / preventing disease onset and must be offered to
 every patient who receives dental implants.
- The aim of treatment is to arrest the inflammatory processes within the peri-implant tissues and to control local and systemic risk factors that may sustain it. Disruption of the locally accumulating microbial biofilms is a key target.
- Treatment of peri-implant mucositis is considered a key strategy in the prevention of the onset of peri-implantitis.
- Treatment of peri-implantitis is performed sequentially, and encompasses an initial nonsurgical step, followed by a surgical step, depending on the outcomes of the initial treatment. SPIC should always be instituted, particularly upon completion of periimplantitis treatment.

The first part of this CPG document (sections 1-4) was prepared by the steering group with the help of the methodology consultant. Section 4, forming the basis for the specific recommendations, was subsequently evaluated by the experts participating in the consensus workshop and voted in a plenary session.

Strength of consensus Unanimous consensus (0% of the group abstained due to potential Col)

5 | RECOMMENDATIONS FOR THE PREVENTION OF PERI-IMPLANT DISEASES

Risk assessment and risk factor control are necessary to prevent the development of periimplant diseases in patients who are candidates for dental implant(s), and in patients who have received dental implant/s and currently have healthy peri-implant tissues.

The purpose of "primordial" prevention (see Section 4.2.1) in the context of the current workshop is to prevent risk factor development prior to dental implant placement. The goal is to attain and maintain optimal oral health to prevent the development of peri-implant diseases over time. There is no current definition of what the optimal oral and general health status of a patient should be prior to dental implant placement, or of which metrics should be included in such a definition. Therefore, no study directly addressing primordial prevention of peri-implant diseases was found, and any recommendations regarding primordial prevention are based upon indirect evidence and expert-based consensus.

The purpose of primary prevention is to prevent disease onset following dental implant placement and loading. The goal is to achieve an optimal oral condition and to maintain dental implant health over time by controlling risk factors for the disease.

The relationship between primordial, primary, secondary, and tertiary prevention is represented in Figure 5, which documents the approach taken by the workshop to interpret the different forms of prevention in the context of peri-implant diseases.

No studies were identified that provided direct evidence for primary prevention. The recommendations are therefore inferred from observational and interventional studies with various working hypotheses that were not originally developed to test the efficacy of a preventative measure on the occurrence of peri-implant diseases. Therefore, the recommendations regarding primary prevention are both evidence-based and expert-based.

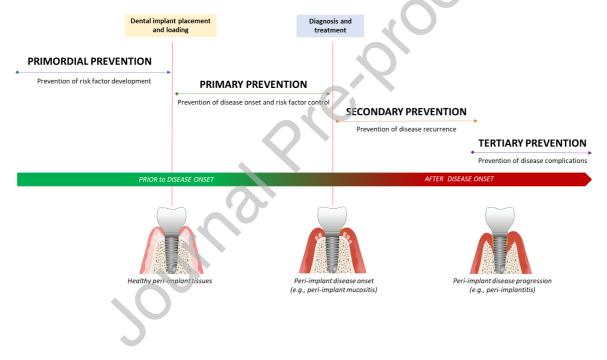
In the present guideline, the term supportive peri-implant care (SPIC) is used to comprise an individually tailored follow-up program which has been described in the available studies with the terms: (1) supportive care; (2) supportive peri-implant care; (3) supportive peri-implant

54

therapy; (4) supportive periodontal therapy; (5) supportive periodontal and peri-implant therapy; (6) supportive therapy.

Junalprendicio

Figure 5. Levels of prevention for peri-implant diseases. The present guideline deals with primordial, primary and secondary prevention. Primordial prevention involves preventing the development of risk factors for peri-implant diseases, including those introduced at the time of implant placement, e.g. position of the implant and cleansability of the prosthesis. Thus, primordial prevention also applies to patients with implants who have healthy peri-implant tissues and no risk factors. However, for the purpose of this guideline, the presence of an implant was regarded as a risk factor for peri-implant diseases (e.g. due to plaque accumulation on a non-shedding surface). Therefore, in the above scheme, primordial prevention does not apply once a dental implant has been placed.



56

5.1 | Recommendations for primordial prevention of peri-implant diseases

The overall objective of this section is to answer the question: in patients awaiting implant placement, does primordial prevention involving the control of lifestyle and behavioural risk factors prevent the development of peri-implant mucositis and peri-implantitis?

R5.1 | In patients awaiting implant placement, do the following behaviours or interventions, prior to implant placement reduce the incidence of peri-implant mucositis and peri-implantitis?

- Educating the patient about the importance of their adherence to SPIC visits and home care
- Improving glycaemic control in people with diabetes
- Smoking cessation (including e-cigarettes) or reducing smoking habit
- Participation in regular supportive periodontal care programs
- Improving oral hygiene
- Reducing bruxing and/or parafunctional habits
- Periodontal therapy to eliminate gingival inflammation and achieve periodontal stability

PICOS question addressed by a systematic review

R5.1 – Expert consensus-based recommendation

In patients awaiting implant placement, **we recommend**:

- 1. thorough assessment of the patient's risk profile to identify and manage modifiable risk factors/indicators for peri-implant diseases.
- 2. Guideline conformed treatment of gingivitis and periodontitis to a stable endpoint and adherence to a supportive care program prior to implant placement.

Supporting literature [26]

Quality of evidence Very low

Grade of recommendation Grade A – $\uparrow\uparrow$

Strength of consensus Unanimous consensus (0% of the group abstained due to

potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

In patients awaiting implant placement, we recommend:

- 1. thorough assessment of the patient's risk profile to identify and manage modifiable risk factors/indicators for peri-implant diseases
- 2. Guideline conformed treatment of gingivitis and periodontitis to a stable endpoint and adherence to a supportive care program prior to implant placement.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

This question was an additional question that was not specifically addressed by the systematic review and therefore relies upon indirect evidence from studies included the review and on expert opinion.

R5.2 | Prior to and during implant placement, what are the considerations related to implant positioning to reduce the risk of incident peri-implant diseases?

Additional question addressed by the WG

R5.2 – Expert consensus-based recommendation

We recommend that treatment planning for 3-dimensional implant positioning should meet the following conditions:

- adequate buccal/lingual bone thickness to allow the implant to be placed in a prosthetically guided position with good primary stability and surrounded circumferentially by bone.
- adequate mesio-distal distance between an implant and adjacent tooth/implant to allow adequate space for prosthetic components and access for oral hygiene aids.

 appropriate apical-coronal position of the implant platform (shoulder) to allow adequate space for prosthetic components and to avoid an excessively deep mucosal sulcus ("tunnel").

Supporting literature [5, 6, 75-80]

Quality of evidence Low

Grade of recommendation Grade $A - \uparrow \uparrow$

Strength of consensus Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adapted.

We recommend that treatment planning for 3-dimensional implant positioning should meet the following conditions:

- adequate buccal/lingual bone thickness to allow the implant to be placed in a prosthetically guided position with good primary stability and surrounded circumferentially by bone.
- adequate mesio-distal distance between an implant and adjacent tooth/implant to allow adequate space for prosthetic components and access for oral hygiene aids, and adequate space for PMPR.
- appropriate apical-coronal position of the implant platform (shoulder) to allow adequate space for prosthetic components and to avoid an excessively deep mucosal sulcus ("tunnel").

Updated Evidence: No new applicable evidence was identified Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

This question was an additional question that was not specifically addressed by the systematic review and therefore relies upon indirect evidence and on expert opinion.

R5.3 | During implant-supported prosthesis design and placement, are there specific considerations to reduce the risk of incident peri-implant diseases?

Additional question addressed by the WG

R5.3 – Expert consensus-based recommendation

In order to facilitate optimal plaque control around implants and to prevent incident periimplant diseases, **we recommend** prosthetic treatment planning should provide for:

- good access for oral hygiene aids used by the patient to remove plaque
- good access for professional monitoring (peri-implant probing) and professional mechanical plaque removal
- a prosthesis contour with a favourable emergence angle and profile to facilitate optimal plaque control

Supporting literature [75, 81-91]

Quality of evidence Moderate

Grade of recommendation Grade A – $\uparrow\uparrow$

Strength of consensus Unanimous consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

In order to facilitate optimal plaque control around implants and to prevent incident peri-

implant diseases, **we recommend** prosthetic treatment planning should provide for:

- good access for oral hygiene aids used by the patient to remove plaque
- good access for professional monitoring (peri-implant probing) and professional mechanical plaque removal
- a prosthesis contour with a favourable emergence angle and profile to facilitate optimal plaque control

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

This question was not addressed by the systematic review and therefore represents an expert consensus-based recommendation, derived from indirect evidence using the cited supporting literature, which may change in the future as new evidence emerges. Expert opinion based on experience is that implant-supported fixed prostheses should have smooth, polished, convex intaglio surfaces, avoid "ridge lap" designs, and, in general, avoid an over-contoured prosthesis, thus facilitating optimal plaque biofilm removal.

5.2 | Recommendations for primary prevention of peri-implant diseases

The overall objective of this section is to answer the question: in patients with dental implants and peri-implant tissue health, does primary prevention involving control of lifestyle and behavioural risk factors prevent the development of peri-implant mucositis and periimplantitis?

R5.4| How should the peri-implant health status be assessed at each clinical examination?

Additional question addressed by the WG

R5.4 - Expert consensus-based recommendations

We recommend peri-implant probing to assess the presence of bleeding on probing, and to monitor changes in probing depth, and changes in the mucosal margin level. The following are advised:

- 1. baseline probing within 3-months of prosthesis delivery
- 2. re-probe at every clinical examination
- 3. use a probe with a 0.5 mm diameter tip and a light probing force (0.2 N)
- 4. record peri-implant probing depths circumferentially (ideally at 6 sites) and bleeding on probing/suppuration
- 5. assess and record the width of keratinised/attached peri-implant mucosa

In addition, **we recommend** a baseline intraoral radiograph be obtained at the completion of physiological remodelling to document marginal bone levels (MBL). At subsequent visits,

if there is an increase in PD in conjunction with BOP/suppuration, we recommend an intraoral radiograph to evaluate the MBL.

Supporting literature

[5, 22, 75, 92, 93]

Grade of recommendation Grade A – $\uparrow\uparrow$

Strength of consensus Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

We recommend peri-implant probing to assess the presence of bleeding on probing, and to monitor changes in probing depth, and changes in the mucosal margin level. The following are advised:

- 1. baseline probing within 3-months of prosthesis delivery
- 2. re-probe at every clinical examination
- 3. use a probe with a 0.5 mm diameter tip and a light probing force (0.2 N)
- 4. record peri-implant probing depths circumferentially (ideally at 6 sites) and bleeding on probing/suppuration
- 5. assess and record the width of keratinised/attached peri-implant mucosa

In addition, **we recommend** a baseline intraoral radiograph be obtained at the completion of physiological remodelling to document marginal bone levels (MBL). At subsequent visits, if there is an increase in PD in conjunction with BOP/suppuration, we recommend an intraoral radiograph to evaluate the MBL.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

This question was not addressed by the systematic review and therefore represents an expert consensus-based recommendation, derived from indirect evidence using the cited supporting literature.

R5.5 | In patients with diabetes and healthy peri-implant tissues, does glycaemic control reduce the risk of incident peri-implant diseases?

PICOS question addressed by a systematic review

R5.5 – Evidence-based recommendation

In patients with diabetes who have healthy peri-implant tissues, **we recommend** glycaemic control to maintain peri-implant health.

Supporting literature [26]

Quality of evidence Low

Grade of recommendation Grade A – $\uparrow\uparrow$

Strength of consensus Consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

We recommend glycaemic control to maintain peri-implant health in patients with diabetes who have healthy peri-implant tissues.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

<u>Intervention</u>

The exposure/risk factor of interest for peri-implantitis is diabetes, and the preventative intervention is glycaemic control (as measured by % of HbA1c).

Available evidence

Number and design of included studies

Eleven observational studies including 6 case-control studies and 5 cohort studies [26].

Risk of bias

According to the Newcastle Ottawa Scale (NOS), 8 studies were at low risk of bias and 3 studies were at high risk of bias.

Effect sizes and their clinical relevance

Pooled data analyses revealed a significantly lower rate of peri-implantitis (OR=0.16; 95% CI [0.03; 0.96]; p=0.004; I2: 0%; analysis based on two studies including 385 implants), and significantly lower marginal bone level (MBL) changes over time (-0.36 mm; 95% CI [-0.65; -0.07; p<0.0001; I2: 95%; analysis based on six studies including 591 implants) in patients with good glycaemic control compared with poor glycaemic control. The mean difference in PD and BOP was not significantly different between the groups. With respect to dental implant survival, diabetes patients with poor glycaemic control were found to have a 7.59 increased risk of dental implant failure compared to patients with good glycaemic control (OR=7.59; 95% CI [1.63; 35.3]; p=0.01; I2: 0%; based on two studies including 524 implants). The estimated mean implant survival was 99% (95% CI [97.8%; 100%]; based on five studies including 253 dental implants) in patients with good glycaemic control and 95.6% (95% CI [91.4%; 99.8%]; based on five studies including 271 dental implants) in patients with poor glycaemic control.

The effect size of these findings is considered clinically relevant, but it must be highlighted that the results are based on a limited number of studies with small sample sizes, that the analyses were performed at the implant level only, and that the definition of good and poor glycaemic control was not consistent among the studies (i.e., good glycaemic control was defined as HbA1c between 6.1% and 8% in five studies, <7% in one study, and <6% in another study; poor glycaemic control was defined as HbA1c level ranging between 8.1% and 10% in 5 studies, as HbA1c >8% in one study, and as HbA1c ranging between 7% and 9% in another study; three studies also included a group of very poorly controlled type-2 diabetes patients, as HbA1c >9 or >10%).

Consistency

Consistency was found in the overall results, favouring good glycaemic control over poor glycaemic control. However, the definition of good and poor glycaemic control was not consistent among the available studies.

64

Balance of benefit and harm

Not assessed. However, glycaemic control in patients with diabetes is advised independently of implant therapy.

Overall certainty of the evidence

No study provided direct evidence. The results are inferred from studies with various working hypotheses that were not originally developed to test the effectiveness of a preventative measure on the occurrence of peri-implant diseases. Further research is needed to provide confidence in the estimated effect of glycaemic control on the risk of peri-implant diseases.

From evidence to recommendation- additional considerations Not applicable.

R5.6| In patients with healthy peri-implant tissues, does provision of regular supportive peri-implant care (SPIC) reduce the risk of incident peri-implant diseases?

PICOS question addressed by a systematic review

R5.6 – Evidence-based recommendation

We recommend regular supportive peri-implant care in patients who have healthy periimplant tissues, to reduce the risk of incident peri-implant diseases, emphasising to the patient the importance of their adherence to SPIC visits and home care.

Supporting literature [26]

Quality of evidence Moderate

Grade of recommendation Grade A – ↑↑

Strength of consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

We recommend regular supportive peri-implant care in patients who have healthy periimplant tissues, to reduce the risk of incident peri-implant diseases, emphasising to the patient the importance of their adherence to SPIC visits and home care. Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

The risk factor/exposure is a lack of appropriate patient follow-up, including periodontal and peri-implant care, and the preventative intervention is promoting and attaining adequate/regular patient adherence to the supportive periodontal/peri-implant care (SPC/SPIC). Various interventions were employed (Table 3 in [26]). The term "supportive peri-implant care" (SPIC) covers the following terms used by the authors of individual studies: (1) supportive care (1 study); (2) supportive peri-implant care (2 studies); (3) supportive peri-implant therapy (4 studies); (4) supportive periodontal therapy (2 studies); (5) supportive periodontal and peri-implant therapy (3 studies); (6) supportive therapy (2 studies). For regular supportive care, the interval between the intervention sessions was: (1) tailored (3 studies); (2) 3 months (1 study); (3) 4 months (1 study); (4) 3 to 6 months (1 study); (5) \leq 6 months (1 study); (6) \leq 12 months (3 studies); (7) unknown (4 studies).

Available evidence

Number and design of included studies

Fourteen studies, 13 observational studies, and 1 RCT [26].

Risk of bias

According to NOS: 7 studies were at low risk of bias and 6 studies were at high risk of bias. According to RoB-II-RCT: 1 study was at some concern.

Effect sizes and their clinical relevance

Twelve studies compared patients regularly attending the recommended SPIC program (adherent) versus non-attending patients or those attending SPIC visits irregularly. Pooled data analyses revealed that patients attending SPIC regularly were at significantly lower risk of presenting with peri-implant diseases (including both peri-implant mucositis and peri-implantitis) (OR=0.42; 95% CI [0.24; 0.75]; p=0.003; I2: 57%; analysis based on six studies including 736 patients) during the study follow-up period (ranging from 1 to 20 years). This was

also observed for the specific diagnosis of peri-implantitis, both at the patient (OR=0.45; 95% CI [0.30; 0.68]; p=0.0002; I2: 51%; analysis based on 6 studies including 736 patients) and implant level (OR=0.26; 95% CI [0.15; 0.46]; p<0.0001; I2: 21%; analysis based on 6 studies including 1337 implants). No significant differences were observed between regular and irregular adherence to SPIC for the diagnosis of peri-implant mucositis.

In a sensitivity analysis excluding those studies that involved patients with a history of periodontitis, dental implants undergoing regular SPIC showed an OR=0.23 (95% CI [0.08; 0.64]; p=0.005; I2: 0%) of developing peri-implantitis compared to dental implants with no SPIC (based on two studies).

When dental implants were used as the statistical unit of analysis, those subjected to regular SPIC demonstrated a lower PD (mean difference: -0.48 mm; 95% CI [-0.67; -0.29]; p<0.0001; I2: 32%; analysis based on five studies including 867 implants) and a reduced risk of exhibiting a MBL > 2 mm (OR: 0.4; 95% CI [0.25; 0.66]; p=0.0003; I2: 73%; analysis based on three studies including 689 implants). Irregular SPIC was associated with a 3.76 increased risk of implant failure (95% CI [1.50; 9.45]; p=0.005; I2: 0%) compared to regular SPIC.

All studies reporting dental implant survival evaluated study samples that included a proportion of patients with a history of periodontitis. Overall, the estimated mean implant survival was 99.3% (95% CI [98.6%; 100%]) in the regular SPIC group (based on 564 implants), and 97.8% (95% CI [95.6%; 99.9%]) in the irregular SPIC group (based on 454 implants) (follow-up ranging from 4.5 to 20 years after implant loading).

The RCT that was evaluated compared four different SPIC protocols (including a 3-monthly SPIC with curettes, with sonic scalers or air polishing, and with or without chlorhexidine varnish application) and found no significant differences between the groups in terms of PD, BOP, and survival at 1 year [94].

When comparing patients with a history of generalized moderate-to-severe periodontitis presenting with deep residual pockets (>6 mm) during SPC, with patients who had a history of generalized moderate-to-severe periodontitis but without residual deep pockets, a significantly

67

higher occurrence of peri-implantitis (3.5% vs. 15.2%, implant level analysis) was observed when deep residual pockets were present [95].

Consistency

All selected studies were overall consistent, favouring regular SPIC over irregular SPIC.

Balance of benefit and harm

Not assessed. However, the importance and clinical relevance of SPIC should be reinforced, given that regular SPIC carries little risk compared to the benefits it brings.

Overall certainty of the evidence

Moderate. Results are inferred from studies with various working hypotheses that were not originally developed to test the effectiveness of a preventative measure on the occurrence of peri-implant diseases. Further research, including clinical trials with strict inclusion criteria, may have an impact on confidence in the estimated effect of regular versus irregular SPIC on the risk of peri-implant diseases.

From evidence to recommendation- additional considerations

Not applicable.

R5.7 In patients who smoke and have healthy peri-implant tissues, does the cessation of cigarette smoking reduce the risk of incident peri-implant diseases?

PICOS question addressed by a SR

R5.7 – Expert consensus-based recommendation

In patients with healthy peri-implant tissues, **we recommend** validated smoking cessation

interventions (by conformance with guidelines) to reduce the risk of peri-implant diseases.

Supporting literature [26]

Quality of evidence Very low

Grade of recommendation Grade A – $\uparrow\uparrow$

Strength of consensus Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

We recommend validated smoking cessation interventions (by conformance with guidelines) In patients with healthy peri-implant tissues to reduce the risk of peri-implant diseases.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

The risk factor is smoking, and the preventative intervention is promotion of smoking cessation advice/strategies.

Available evidence

Number and design of included studies

Four studies, including three case-control and one cohort study [26]. Clear similarities between the three case-control studies conducted by the same research team were noted.

Risk of bias

According to NOS the three case-control studies were at high risk of bias, and the cohort study was at low risk of bias.

Effect sizes and their clinical relevance

Only one study described the occurrence of peri-implant diseases as a clinical diagnosis, reporting a lower rate of peri-implant mucositis (43.9% vs. 48.6%) and peri-implantitis (19.7% vs. 30.5%) in former smokers compared to current smokers [96]. The authors observed a direct association between cumulative smoking exposure and the risk for peri-implantitis, as well as with the time span since smoking cessation.

All studies reported significant clinical differences between former smokers, e-cigarette users, waterpipe smokers, and current smokers. The former smoker category exhibited less periimplant mucosal inflammation, PD and MBL compared to the other categories.

Consistency

There is insufficient evidence to determine whether cigarette smoking cessation decreases the risk for peri-implant diseases. There is little evidence to support the contention that using e-cigarettes, or the habit of water pipe smoking is associated with a decreased risk for peri-implant diseases compared to cigarette smoking.

Balance of benefit and harm

Not assessed. However, because of the several harmful consequences of smoking, smoking cessation should be advised and promoted for every patient irrespective of implant therapy.

Overall certainty of the evidence

Low. No interventional studies were found to provide direct evidence. The results are inferred from studies with various working hypotheses that were not originally developed to test the effectiveness of smoking cessation on the occurrence of peri-implant diseases. Further research is very likely to have an impact on confidence in the estimate of the effects of cigarette cessation on the reduction of the risk of incident peri-implant diseases. Regarding the use of non-cigarette smoking, any estimate of effect is very uncertain.

From evidence to recommendation- additional considerations Not applicable.

R5.8 | In patients with healthy peri-implant tissues, does augmentation of peri-implant soft tissues lower the likelihood of incident peri-implant diseases?

PICOS question addressed by a SR

R5.8 – Evidence-based recommendation

In patients who have dental implants with an absence or deficiency (narrow width) of keratinized/attached mucosa, and where the patient experiences discomfort on brushing,

increasing peri-implant keratinised/attached mucosal width to maintain peri-implant health

may be considered.

Supporting literature [26]

Quality of evidence Low

Grade of recommendation Grade O - \leftrightarrow

Strength of consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

In patients who have dental implants with an absence or deficiency (narrow width) of keratinized/attached mucosa, and where the patient experiences discomfort on brushing, increasing peri-implant keratinised/attached mucosal width to maintain peri-implant health **may be considered**.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (6.8% abstentions due to potential Col)

Background

Intervention

The risk factor is the deficiency of peri-implant keratinized mucosa (PIKM) (PIKM < 1, 2 or 3 mm according to the studies), and the preventative intervention is the augmentation of PIKM by a free gingival graft (FGG).

Available evidence

Number and design of included studies

Six of the studies included in the systematic review and meta-analysis (3 RCTs, 1 NRCT, 1 case-control, and 1 cohort study) were considered. They compared peri-implant tissue health parameters between sites with a deficiency in PIKM and receiving a FGG to increase PIKM width versus no intervention. No study was specifically designed to assess the impact of FGG on the prevention of peri-implant diseases.

Risk of bias

According to RoB-II-RCT: the three RCTs presented some concerns. According to RoBins-NRTC: the selected study was at moderate risk of bias. According to NOS: the two studies were at low risk of bias.

Effect sizes and their clinical relevance

Indirect evidence based on the evaluation of peri-implant health parameters in the short term showed a non-significantly different PPD between the PIKM-augmented and non-augmented sites but a significantly lower clinical soft tissue inflammation index (BOP/GI) (standardized mean difference - SMD=-1.18; 95% CI [-1.85; -0.51]; p=0.0006; I²: 69%) around the dental implants receiving FGG to augment PIKM. Concerning the mean MBL, based on data from four studies, a significant difference in favour of PIKM-augmented sites (SMD: -0.25; 95% CI [-0.45; -0.05]; p=0.01; I²: 62%) was also noted. When excluding from pooled data analysis cohort and case-control studies, the results were consistent with no statistical heterogeneity. No difference in PPD (SMD: -0.25; 95% CI [-0.63; -0.13]; p=0.20; I²: 0%; based on 107 implants), whereas a significant difference in BOP (SMD: -1.5; 95% CI [-1.93; -1.06]; p<0.0001; I²: 0%; based on 107 implants) and MBL changes (SMD: -0.33; 95% CI [-0.55, -0.11; p=0.003; I²: 0%; based on two studies, 66 implants) were noted between PIKM-augmented sites vs. non-augmented sites.

Only two studies reported the occurrence of PIDs [97, 98]. The first study defined periimplantitis as the presence of BOP, PPD \geq 5 mm, and a radiographic bone loss \geq 3.5 mm [97]. During a mean follow-up of 12 years, 3 groups receiving FGG, CTG, or no intervention were compared. No statistical differences were found between groups. The second study, a 10-year prospective cohort, observed a significantly higher rate of PIDs for dental implants with PIKM deficiency compared to implants surrounded by PIKM (51.4% vs. 12.7%; p<0.0001) [98]. The authors also reported a significantly lower soreness for implants surrounded by PIKM or placed in the alveolar mucosa receiving FGG compared to implants surrounded by alveolar mucosa and not receiving FGG [98].

72

Consistency

Results are based on heterogeneous studies with, most of the time, small sample sizes and short follow-ups. Consistency is low.

Balance of benefit and harm

Not assessed. However, the decision-making process concerning surgical procedures to augment PIKM should consider the general risks associated with periodontal and implant surgery.

Overall certainty of the evidence

Low. No study design provided direct evidence. Results are inferred from studies with various working hypotheses that were not originally developed to test the effectiveness of peri-implant soft tissue augmentation procedures on the prevention of peri-implant diseases over time.

Care must be taken regarding the interpretation of the study results due to the high clinical heterogeneity of the included studies. Most of the studies described clinical peri-implant outcomes in the short-term (6-12 months follow-up), whereas only two observational studies reported the occurrence of peri-implant diseases over a 10-year (low risk of bias) and 12-year (high risk of bias) follow-up.

However, a reduced width of keratinized tissue is associated with an increased prevalence of peri-implantitis, plaque accumulation, soft-tissue inflammation, mucosal recession, marginal bone loss, and greater patient discomfort [99]. The effectiveness of increasing PIKM as a preventative measure for peri-implant diseases requires longitudinal studies designed with a long-term follow-up, to evaluate the outcome of interest (i.e., peri-implant diseases).

From evidence to recommendation- additional considerations Not applicable. R5.9| In patients with healthy and thin peri-implant tissues (< 2 mm in thickness), does soft tissue augmentation lower the likelihood of incident peri-implant diseases?

PICOS question addressed by a SR

R5.9 – Evidence-based statement

We **do not know** if undertaking procedures to augment soft tissue thickness prevents the development of peri-implant diseases, since there is lack of evidence to support an association between increasing soft tissue thickness and peri-implant tissue health.

Supporting literature

[26, 100, 101]

Quality of evidence - Low

Strength of consensus Consensus (7.8% of the group abstained due to potential Col)

BSP Implementation

This evidence-based statement is adapted.

In patients who have existing dental implants with healthy, but thin (<2mm thickness) peri-implant tissues, **we do not know** whether increasing soft tissue thickness at this stage lowers the future likelihood of peri-implantitis.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (2.5% abstentions due to potential Col)

Background

Intervention

Peri-implant soft tissue augmentation to increase PIKM thickness, including the following surgical procedures: (1) connective tissue graft (CTF), (2) free gingival graft (FGG), (3) the use of xenogenic collagen matrix (XCM), (4) or acellular dermal matrix allograft.

Available evidence

Number and design of included studies Eight studies, including one NRCT and six RCTs [26]

Risk of bias

According to NOS: 1 study is at high risk of bias According to RoB-II-RCT: 2 studies were at low risk of bias, and 4 studies presented some concern. According to RoBins-NRTC: 1 study is at moderate risk of bias

Effect sizes and their clinical relevance

Pooled data analyses were based on 4 studies, including 179 implants, and found no difference between CTG/FGG vs. XCM for mean PPD, MBL, and BOP. One controlled clinical trial with a small sample size (19 patients) observed a 4.3% rate of peri-implantitis in the control group compared to 0% in the test group receiving CTG (partial split-mouth design) [102]. Metaanalysis was performed pooling together two studies comparing CTG vs. no intervention [97, 102], and including 37 implants in CTG-augmented sites vs. 69 implants in non-augmented sites. It showed no significant difference between the two groups for the rate of incident periimplantitis (OR=1.97; 95% CI [0.20; 19.72]; p=0.56; I²: 0%).

Consistency

Data are consistent, although based on a very limited number of studies.

Balance of benefit and harm

Not assessed. However, the decision-making process should balance the risks associated with the different surgical procedures aimed at increasing PIKM thickness against the risks of surgery and the additional related costs, in people with peri-implant mucosal health.

Overall certainty of the evidence

No study design provided direct evidence. Care must be taken regarding the interpretation of the results, due to the high clinical heterogeneity of the included studies, in particular the high variability of the timeline at which the augmentation procedure was performed (before or after dental implant placement, after dental implant loading, simultaneously to the dental implant placement, at the stage 2 surgery, etc.). Most of the studies described clinical peri-implant outcomes in the short-term (6-12 months follow-up).

From evidence to recommendation- additional considerations

Not applicable.

R5.10 | In patients with healthy peri-implant tissues, does improved oral hygiene prevent incident peri-implant diseases?

PICOS question addressed by a SR

R5.10 – Expert consensus-based recommendation

In patients who have dental implants **we recommend** specific, individually-tailored oral

hygiene instructions to reduce the risk of incident peri-implant diseases

Supporting literature [26]

Grade of recommendation Grade A – $\uparrow\uparrow$

Strength of consensus Unanimous consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

In patients who have dental implants **we recommend** specific, individually-tailored oral hygiene instructions to reduce the risk of incident peri-implant diseases

Updated Evidence: No new applicable evidence was identified.

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

The risk factor is inadequate oral hygiene (OH), and the preventative intervention, improving OH behaviours. The following toothbrushes were evaluated: (1) counter-rotational powered toothbrush, (2) sonic toothbrush, and (3) manual toothbrush. The following frequencies were evaluated: brushing at least twice/day or brushing at most once/day.

<u>Available evidence</u>

Number and design of included studies

Three studies were selected: two RCTs, and one case-control study [26].

Risk of bias

According to NOS: one study was at low risk of bias. According to RoB-II-RCT: two studies were at some concern.

Effect sizes and their clinical relevance

Due to the heterogeneity in reporting outcome measures, no analysis of pooled data was possible. One study found a significant difference in favour of a counter-rotational powered toothbrush in terms of peri-implant mucosal inflammation and implant survival compared to manual toothbrushing [10]). One case-control study indicated that the frequency of tooth brushing (at least twice a day vs. at most once a day) had no impact on peri-implant PD, MBL, and BOP [104].

Consistency

The three studies included were inconclusive regarding the type of toothbrush to use (e.g. powered or manual toothbrush), or the frequency of toothbrushing that was most effective in maintaining peri-implant health.

Balance of benefit and harm

Not assessed in the studies considered. However, advising patients about OH and promoting OH behaviour improvements (in terms of techniques and frequency) carry little risk compared to the benefit it brings.

Overall certainty of the evidence Low.

From evidence to recommendation- additional considerations Not applicable. R5.11 | In patients with healthy peri-implant tissues, does reducing bruxing/ parafunctional habits reduce the risk of incident peri-implant diseases?

PICOS question addressed by a SR

R5.11 – Expert consensus-based statement

We do not know whether in patients with healthy peri-implant tissues, controlling bruxing/parafunctional habits reduces the risk of incident peri-implant diseases.

Supporting literature [26]

Quality of evidence no studies met the inclusion criteria in the review

Grade of recommendation Grade O – ↔ Statement, additional research needed

Strength of consensus Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based statement is adopted.

We do not know whether in patients with healthy peri-implant tissues, controlling bruxing/parafunctional habits reduces the risk of incident peri-implant diseases.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

There were no studies that investigated the control of bruxing/parafunctional habits in patients with healthy peri-implant tissues in preventing the risk of peri-implant diseases.

5.3 | Secondary and tertiary prevention: recommendations for supportive peri-implant care

This section aims to answer the following questions: in patients treated for peri-implantitis, what is the efficacy of: 1) supportive care, 2) supportive peri-implant care with adjunctive local antiseptic agents, and 3) of supportive peri-implant care with a frequency of more than once a year in achieving peri-implant tissue stability.

A systematic review [35] was designed to evaluate the efficacy of providing supportive periimplant care (SPIC), as well as specific SPIC protocols and frequency upon peri-implant stability after a minimum recall period of 3-years. Fifteen studies were identified that met the inclusion criteria, which included a minimum of 20 volunteers. No studies were specifically designed to evaluate SPIC provision, protocol or frequency, and all studies were surgical intervention trials that included SPIC as part of their design. Therefore, there were no studies that compared specific SPIC protocols or frequency of provision, or the use of adjunctive therapies versus none, or studies that compared the provision of SPIC versus no SPIC.

There were 10 prospective and 5 retrospective studies, 14 of which provided SPIC using various techniques for professional mechanical plaque removal (PMPR) in combination with (n=10) or without (n=4) oral hygiene instruction. Disease recurrence/progression outcomes were defined by the authors of the respective studies (n=13), or were based upon progressive deterioration in BOP, PD or marginal bone level (n=2). Stability outcomes and disease recurrence were reported at both the implant and the patient level.

The three PICOS questions documented below could not be answered by the systematic review, and a meta-analysis was inappropriate due to the high heterogeneity of the data. However, risk of bias was deemed low in 87 % of the studies. The working group participants felt there was sufficient data to address the overarching question of whether regular provision of SPIC improved peri-implant tissue stability following surgical treatment of peri-implantitis, in an evidence based manner, however most recommendations are based upon expert consensus. There were additional questions deemed to be of importance to clinical practice that were not directly informed by the systematic review, but for which the workshop formulated recommendations based on the literature base.

Given the paucity of available studies (n=15), the background study characteristics provided following the recommendation tables are deemed applicable to all recommendations.

79

R5.12 | In patients treated for peri-implantitis, does supportive peri-implant care (SPIC) prevent recurrence of peri-implantitis in the medium to long-term (≥ 3 years)?

PICOS question addressed by a SR

R5.12 – Evidence-based recommendation

We recommend the provision of SPIC to reduce the risk of recurrence of peri-implantitis

and consequent implant loss, emphasising to the patient the importance of their adherence

to SPIC visits and home care.

Supporting literature [35, 105]

Quality of evidence Low (indirect evidence)

Grade of recommendation Grade A – ↑↑

Strength of consensus Unanimous consensus (0% of the group abstained due to potential

Col)

BSP Implementation

This evidence-based recommendation is adopted.

We recommend the provision of SPIC to reduce the risk of recurrence of peri-implantitis and consequent implant loss, emphasising to the patient the importance of their adherence

to SPIC visits and home care.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

See background text in R5.16, common for recommendations R5.11-16.

R5.13 | In patients treated for peri-implantitis, what is the recommended frequency of supportive peri-implant care (SPIC)?

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

See background text in R5.16, common for recommendations R5.11-16.

R5.14 | In patients treated for peri-implantitis, what is the appropriate protocol for supportive peri-implant care provision (SPIC)?

PICOS question addressed by a SR

R5.14 – Expert consensus-based recommendation

We recommend that the implementation of a patient-centred SPIC protocol should include the following components:

- Interview (medical, social & oral history update, risk assessment, patient feedback)
- Assessment of: oral, including peri-implant tissue health, prosthetic components, patient competence to undertake oral hygiene
- Reinforce risk factor control (e.g. smoking, oral dryness, glycaemic control)
- Professional intervention: individualised oral healthcare plan, including oral hygiene coaching, PMPR to include entire dentition/implants)
- determination of next recall interval tailored according to patient-, implant- and restoration-based risk factors.*

Supporting literature [26, 35]

Quality of evidence Very low (indirect evidence for some components)

Grade of recommendation Grade $A - \uparrow \uparrow$

Strength of consensus Unanimous consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

We recommend that the implementation of a patient-centred SPIC protocol should include

the following components:

Interview (medical, social & oral history update, risk assessment, patient feedback)

• Assessment of: oral, including peri-implant tissue health, prosthetic components, patient competence to undertake oral hygiene

• Reinforce risk factor control (e.g. smoking, oral dryness, glycaemic control)

• Professional intervention: individualised oral healthcare plan, including oral hygiene coaching, PMPR to include entire dentition/implants)

• determination of next recall interval tailored according to patient-, implant- and restoration-based risk factors.*

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

*The protocol applies to any patient with dental implants.

Background

See background text in R5.16, common for recommendations R5.11-16.

R5.15 | In patients treated for peri-implantitis is there a specific regime for professional mechanical plaque removal (PMPR) that reduces risk of disease recurrence?

PICOS question addressed by a SR

R5.15 – Expert consensus-based statement

We do not know which specific PMPR regime is most effective in reducing the risk of recurrent peri-implantitis. However, based upon the periodontal literature and indirect evidence, the following approaches for dental implant biofilm removal can be used alone or in combination:

- Titanium or stainless-steel area-specific curettes
- Ultrasonic/sonic instruments
- Rubber cup or brushes
- Air-polishing devices with glycine powder or erythritol alone or in combination.

Supporting literature [35]

Quality of evidence No studies were identified to compare different PMPR regimes

Grade of recommendation Grade $O - \leftrightarrow$ Statement, additional research needed

Strength of consensus Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based statement is adopted.

We do not know which specific PMPR regime is most effective in reducing the risk of recurrent peri-implantitis. However, based upon the periodontal literature and indirect evidence, the following approaches for dental implant biofilm removal can be used alone or in combination:

- Titanium or stainless-steel area-specific curettes
- Ultrasonic/sonic instruments
- Rubber cup or brushes
- Air-polishing devices with glycine powder or erythritol alone or in combination.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (10% abstentions due to potential Col)

Background

See background text in R5.16, common for recommendations R5.11-16.

R5.16 | In patients treated for peri-implantitis is there a specific oral hygiene method that reduces risk of disease recurrence?

PICOS question addressed by a SR

R5.16 – Expert-based consensus statement (1), Evidence-based recommendation (2) **1. We do not know** which specific oral hygiene method is most effective in reducing the risk of recurrent peri-implantitis. However, based upon the periodontal literature, indirect evidence and expert opinion, we recommend care individually tailored for the patient including at least:

- twice daily brushing of dental implants and teeth using either manual or rechargeable power brushes;
- once daily use of interproximal brushes of an appropriate size;

2. We recommend oral hygiene methods be demonstrated by the patient to the oral

healthcare professional and periodically reinforced.

Supporting literature [35]

Quality of evidence No studies were identified to compare different oral hygiene methods

Grade of recommendation Grade O – \leftrightarrow Statement, additional research needed (1); Grade A – $\uparrow\uparrow$ (2)

Strength of consensus Unanimous consensus (10.9% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based statement (1), evidence-based recommendation

(2) is adopted.

1. **We do not know** which specific oral hygiene method is most effective in reducing the risk of recurrent peri-implantitis. However, based upon the periodontal literature, indirect evidence and expert opinion, we recommend care individually tailored for the patient including at least:

- twice daily brushing of dental implants and teeth using either manual or rechargeable power brushes;
- once daily use of interproximal brushes of an appropriate size;

2. **We recommend** oral hygiene methods be demonstrated by the patient to the oral healthcare professional and periodically reinforced.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (25% abstentions due to potential Col)

Background

See background text in R5.16, common for recommendations R5.11-16.

R5.17 | In patients treated for peri-implantitis does the professional administration* of adjunctive local antimicrobial agents as part of a supportive peri-implant care (SPIC) program reduce the risk of disease recurrence?

Question addressed by a SR

R5.17 – Expert-based consensus recommendation

We suggest not to use professional application* of adjunctive local antimicrobial agents in SPIC to reduce the risk of recurrent peri-implantitis.

Supporting literature [35]

Quality of evidence No studies were identified to specifically evaluate local antimicrobial agent use in secondary prevention of peri-implantitis

Grade of recommendation Grade $B - \psi$

Strength of consensus Strong consensus (3.8% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

We recommend not to use professional application* of adjunctive local antimicrobial

agents in SPIC to reduce the risk of recurrent peri-implantitis.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (6.3% abstentions due to potential

Col)

*Professional administration is by the oral healthcare professional within the dental office.

Background

Intervention

Supportive peri-implant care (SPIC) provided after completion of active peri-implantitis therapy

(i.e. any intervention during a supportive care visit). These interventions include:

• reinforcement of systemic risk factor control (e.g. metabolic, inflammatory, and hormonal diseases, medications, tobacco use, stress)

- management of remaining local risk factors (site-related factors, e.g. keratinized tissue width), implant- and prosthesis-related factors)
- reinforcement of self-performed mechanical plaque control regimes (with or without antiseptic agents)
- professional mechanical plaque removal (PMPR):
 - removal of supra- and sub-mucosal biofilm by hand or mechanical instruments
 - removal of supra- and sub-mucosal hard deposits (calculus) by hand or mechanical instruments

Available evidence

Number and design of included studies

A total of 15 studies were included in this systematic review [35]. The studies included were of prospective (n=10) and retrospective (n=5) design reporting on a single treatment group (n=9) or multiple treatment groups (n=6), conducted in a university (n=12) or private practice (n=3). All studies that fulfilled the inclusion criteria regarding patient number (\geq 20 patients) and follow-up time (\geq 3 years), were focused on the medium to long term outcomes of peri-implantitis treatment. None of the studies meeting the inclusion criteria were specifically designed to evaluate or compare different supportive peri-implant care protocols or SPIC frequencies and only one study was designed to evaluate the effect of SPIC on the secondary prevention of peri-implantitis.

Risk of bias

Most studies (87%) were assessed as having a low risk of bias, two studies (13%) showed some concerns, mainly regarding the inclusion of participants (lack of randomization information), treatment standardization, or definition of treatment success and disease recurrence. There was considerable heterogeneity between studies with respect to study design including: peri-implantitis case definitions, outcomes reported, outcome definitions for success and disease recurrence, peri-implantitis treatment methods, and supportive care protocols.

Effect sizes and their clinical relevance

Definitions for peri-implantitis, treatment success, and recurrence of disease varied considerably across the fifteen studies, contributing significantly to the heterogeneity of the data. While all definitions of peri-implantitis included clinical parameters such as bleeding on probing, probing depth, and radiographic bone loss, the defined thresholds for bone loss and probing depth were heterogeneous.

Definitions for success were reported by 13 of the studies but also varied between studies. Therefore, a quantitative assessment of implant- and patient-level success was not possible. In 9 studies, success was defined as PD < 5 mm with no bleeding on probing or suppuration and no further bone loss. In one study, success was defined as PD < 4 mm with no bleeding on probing or suppuration and no mobility. One study defined success as PD reduction, favourable soft tissue parameters and BOP decrease. Another study defined success as no further bone loss of > 1.0 mm and no implant removal, and a further study defined success as radiographic evidence of > 25% bone fill.

The definition of disease "recurrence" also varied significantly between the studies. In 8 studies, "further bone loss" was defined as one important criterion for recurrence, together with implant loss (two studies). In 4 studies, BOP was a criterion for recurrence and in one study disease recurrence included clinical outcomes not meeting the success criteria.

Consistency

The review found that peri-implant tissue stability reported at the patient-level and at the implant-level varied widely and that recurrence of peri-implantitis was reported in up to 65.2 % of treated implants receiving supportive peri-implant care in studies with a follow-up of 3 years or more. While the systematic review [35] aimed to identify the most effective supportive care protocol in maintaining peri-implant tissue stability after peri-implantitis treatment, no comparison of protocols could be made. Furthermore, as the studies were not specifically designed to evaluate supportive care protocols, detailed information regarding supportive care was lacking. Therefore, it was not possible to make any conclusion regarding the most effective supportive care protocols included similar preventative and therapeutic principles of supportive periodontal care as described in the EFP S3-level treatment guideline

for stages I-III periodontitis [25]. Regular removal of plaque from the treated implant was common to all protocols described. Several studies also specified the provision of full-mouth professional plaque removal and the reinforcement of oral hygiene instructions.

Balance of benefit and harm

The results of this review confirm that SPIC may result in peri-implant tissue stability after periimplantitis treatment. However, disease recurrence may occur, requiring additional treatment or, in some cases, implant removal. The undesirable effects of SPIC have not been described in the included studies.

Overall certainty of the evidence

Currently, there is no high-quality evidence available to answer the PICOS of the systematic review. Based on the available literature a meta-analysis was not possible. The overall evidence on the effect of SPIC on the secondary prevention of peri-implantitis is based on one RCT, seven prospective and five retrospective clinical trials. Provision of SPIC following peri-implantitis therapy may prevent disease recurrence or progression. Insufficient evidence is available to identify (i) a specific supportive care protocol for secondary prevention of peri-implantitis (ii) the effect of adjunctive local antiseptic agents in the secondary prevention of peri-implantitis and (iii) the impact of frequency of supportive care provision. Future prospective randomized controlled studies designed to evaluate supportive care protocols are needed.

From evidence to recommendation- additional considerations

Acceptability

In most of the identified studies, the number of drop-outs were few and the study participants seemed to be compliant. Based on the findings of the systematic review [35] it may be assumed that the provision of SPIC with a frequency between 3 to 6 monthly over a time span of three years is acceptable for patients following peri-implantitis treatment.

Feasibility

There were no perceived barriers.

Ethical Considerations

As an example, in Germany, neither implant therapy nor SPIC is part of the statutory health insurance. Patients only receive access to SPIC through private health insurance or self-payment.

Economic Considerations

As SPIC may prevent peri-implantitis recurrence, it is an important tool to support overall oral health and well-being of patients with implants. The loss of an implant may be associated with bone loss, psychological distress, pain, and costly and time-demanding retreatments which may require specialist management.

Legal Considerations

There were no legal constraints.

6 | RECOMMENDATIONS FOR THE MANAGEMENT OF PERI-IMPLANT MUCOSITIS

6.1 | Introduction - general recommendations in the management of peri-implant mucositis

R6.1 | In patients with peri-implant mucositis, which are the goals/endpoints of treatment?

Additional question addressed by the WG

R6.1 – Expert consensus-based recommendations

We **recommend** that clinicians use as the endpoint of peri-implant mucositis treatment at implant level: \leq 1 point of BOP* and absence of suppuration

We recommend that clinicians evaluate these endpoints 2-3 months after the intervention,

and in presence of \geq 2 BOP sites, or \geq 1 sites with profuse BOP, or presence of suppuration, re-treatment should be rendered.

Supporting literature

[76, 106-109]

Grade of recommendation Grade A – $\uparrow\uparrow$

Strength of consensus Unanimous consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

We recommend that clinicians use as the endpoint of peri-implant mucositis treatment at implant level: \leq 1 point of BOP* and absence of suppuration.

We recommend that clinicians evaluate these endpoints 2-3 months after the intervention, and in presence of \geq 2 BOP sites, or \geq 1 sites with profuse BOP, or presence of suppuration, re-treatment should be rendered.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

*: BOP should be a spot rather than profuse bleeding using gentle forces (0.2 N) with a manual periodontal probe (0.5 mm tip), provided the contours of the restoration allow for adequate probing. In sites where probing is not feasible, peri-implant mucosal inflammation should be assessed through the modified sulcus bleeding index (Mombelli, van Oosten, Schurch, & Land, 1987).

Background

This recommendation is an expert-based recommendation supported by experimental studies [107], experimental peri-implant mucositis studies [76, 108, 109] and studies evaluating the probe penetration and bleeding on probing in healthy periodontal versus peri-implant tissues [106]. All these studies have assessed the similarities and differences between peri-implant and periodontal tissues, how peri-implant tissues respond to biofilm accumulation and, which is the degree of reversibility when the biofilm is eliminated (experimental peri-implant mucositis model).

R6.2 | In patients with peri-implant mucositis, what is the effect of oral hygiene as an adjunct to professional mechanical plaque removal (PMPR)?

PICOS question addressed by a SR

R6.2 – Expert consensus-based recommendation

In patients with peri-implant mucositis, we recommend self-performed effective oral

hygiene along with PMPR

Supporting literature [37]

Quality of evidence No clinical studies were identified.

Grade of recommendation Grade A – ↑↑

Strength of consensus Unanimous consensus (0% of the group abstained due to

potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

In patients with peri-implant mucositis, **we recommend** self-performed effective oral hygiene along with PMPR.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

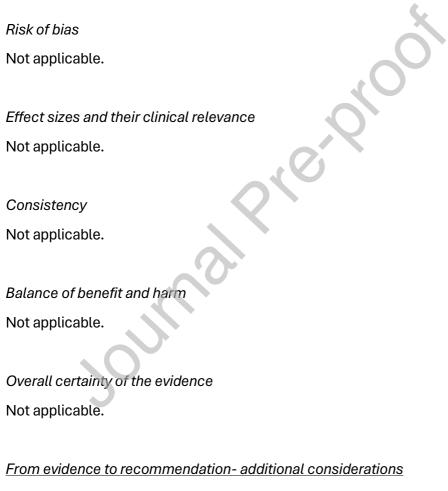
Background

Intervention

There are no available clinical studies with an arm with oral hygiene alone without PMPR. Furthermore, for obvious ethical reasons there are no studies without implementing oral hygiene measures. However, there is indirect evidence from experimental mucositis studies demonstrating that oral hygiene can revert the inflammatory signs in the peri-implant mucosa. This evidence has concluded that experimental peri-implant mucositis is caused by biofilm accumulation and that it may be reversible by means of oral hygiene reinforcement alone [76, 108, 109]. Due to this microbial aetiology, there is a clear rationale to combine professionally administered non-surgical mechanical/physical therapy with patient-performed oral hygiene reinforcement in the treatment of peri-implant mucositis. This combination results in biofilm disruption and leads to improved clinical outcomes.

Available evidence

There are no RCTs, nor observational studies (with n=30 patients or more), or single arms from RCTs (with n=10 patients or more) evaluating the efficacy of oral hygiene reinforcement alone as treatment for peri-implant mucositis. Similarly, there are no RCTs where professionally administered nonsurgical mechanical/physical instrumentation was implemented without oral hygiene reinforcement.



Acceptability

Self-performed oral hygiene measures are generally well-accepted by individuals.

Feasibility

Non-surgical mechanical/physical treatment of peri-implant mucositis can be performed by dental hygienists, general dentists as well as specialist.

Ethical Considerations

Not applicable.

Economic Considerations Not applicable.

Legal Considerations Not applicable.

R6.3 | In patients with peri-implant mucositis, what is the efficacy of oral irrigators adjunctively used to PMPR?

PICOS question addressed by a SR

R6.3 – Evidence-based recommendation

In patients with peri-implant mucositis the self-use of oral irrigation devices with water may

be considered as an adjunct to PMPR.

Supporting literature [31]

Quality of evidence Low (two RCTs, one with low and the other with moderate risk of bias)

Grade of recommendation Grade $O - \leftrightarrow$ need for further research

Strength of consensus Consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

In patients with peri-implant mucositis the self-use of oral irrigation devices with water may

be considered as an adjunct to PMPR.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (21.4 % abstentions due to potential Col)

Background

Intervention

Oral irrigators can be used regularly as adjuvants to PMPR in addition to regular oral hygiene practices.

Available evidence

Number and design of included studies

The systematic review [31] included two RCTs evaluating the effect of oral irrigators used by the patient adjunctively to PMPR compared to PMPR, demonstrating significant BOP reduction at 3 months in patients with peri-implant mucositis.

Risk of bias

The overall risk of bias of the included studies was judged as 'moderate' (Rob2 tool), with one study with a low risk of bias and one with a moderate risk of bias.

Effect sizes and their clinical relevance

These two RCTs show imprecision in the effect estimates, the results are not consistent, and publication bias could not be assessed.

Consistency

The reported results are not consistent.

Balance of benefit and harm

It could not be assessed.

Overall certainty of the evidence

Two RCTs have evaluated the adjunctive self-use of oral irrigators, one using 0.06% CHX as the irrigating fluid and the other water, one study was at low and the other at moderate risk of bias. Furthermore, the imprecision of the effect estimates, the lack of consistency of the results and the potential risk of publication advises downgrading the quality of the evidence.

Acceptability

Oral irrigators are usually well accepted by patients.

Feasibility

There are no perceived barriers.

Ethical Considerations

There are no perceived ethical considerations.

Economic Considerations

There is an additional cost on buying the irrigator.

Legal Considerations

There are no perceived legal considerations.

R6.4 | In patients with peri-implant mucositis, what is the effect of any single mode of PMPR, compared to other single modes of PMPR?

PICOS question addressed by a SR

R6.4 – Evidence-based recommendation

In patients with peri-implant mucositis, ultrasonics with plastic coated tips or airpolishing devices with glycine powder or titanium curettes or chitosan brushes **may**

be considered as a single mode of PMPR.

Supporting literature [37]

Quality of evidence Low (two RCTs demonstrating positive effects within the single

mode of PMPR, but without differences among them)

Grade of recommendation: Grade $O - \leftrightarrow$ (need for further research)

Strength of consensus (5.3% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adapted.

In patients with peri-implant mucositis, ultrasonics with plastic coated or carbon fibre tips or air-polishing devices with glycine powder or titanium curettes or chitosan brushes **may be considered** as a single mode of PMPR.

Updated Evidence: No new applicable evidence was identified Strength of Consensus: Unanimous consensus (9.4% abstentions due to potential Col)

Background

Intervention

PMPR aims at reducing soft tissue inflammation by removing hard and soft deposits from the surface of the dental implant and/or its supra-structure without scratching the surface of the smooth transmucosal element (implant collar, abutment). Several modalities including ultrasonics with carbon fibre or plastic tip, air-polishing, curettes of plastic, carbon or titanium or rotating/oscillating brushes and lasers have been used within PMPR. The endpoint of treatment is to eliminate inflammation, evaluated by bleeding on probing and suppuration.

Available evidence

Two RCTs comparing two single modes of mechanical therapies were identified [37]. One is a 12-month parallel group RCT (n=37 patients) comparing glycine powder air polishing and ultrasonic with plastic coated tips. The mean BOP reductions were 31.8% and 35.1%, respectively at 12 months, without statistically significant differences between both modes of therapy. The other is a 6-month split-mouth RCT (n=11 patients) comparing titanium curettes and chitosan brushes after a period of oral hygiene. The mean reduction in BOP severity (modified sulcus bleeding index), was 0.84 and 0.61, respectively. The mean disease resolution at implant level (up to one spot BOP) was 50% and 35% at 6 months.

Risk of bias

Study quality assessment identified some concerns of risk of bias in one study and high risk of bias the other.

Effect sizes and their clinical relevance

One study reported disease resolution/treatment success in 8.3-16.7 % at 6 months, and BOP severity of 0.70-0.74. In this study, oral hygiene instruction was performed before the baseline examination. Another study reported BOP extent at 12.1-18.6 % at 12 months.

Consistency

Evidence was consistent in the two studies with limited reduction in BOP. The only patientreported outcome showed no difference in pain during treatment when titanium curettes were compared to chitosan brush.

Balance of benefit and harm

An overall consideration of the benefit versus harm of professionally administered non-surgical mechanical/physical therapy supports the recommendation.

Overall certainty of the evidence Low.

From evidence to recommendation- additional considerations

Acceptability

Patients usually accept and understand the need for treatment.

Feasibility

Non-surgical mechanical/physical treatment of peri-implant mucositis can be performed by dental hygienists, general dentists as well as specialist.

Ethical Considerations Not applicable.

Economic Considerations Not applicable.

Legal Considerations

Not applicable.

R6.5 | In patients with peri-implant mucositis, what is the effect of combinations of PMPR

procedures, compared to single modes?

PICOS question addressed by a SR

R6.5 – Evidence-based recommendations

In patients with peri-implant mucositis, we suggest not to add air-polishing devices

to conventional PMPR (curettes, ultrasonics, or both), even though these devices have shown efficacy when used as a single mode of treatment.

In patients with peri-implant mucositis, **we suggest not to** add diode lasers with conventional PMPR (curettes, ultrasonics, or both).

Supporting literature [37]

Quality of evidence Moderate (three RCTs, n=313 patients)

Grade of recommendation Grade $B - \downarrow$

Strength of consensus Consensus (15.4% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

We do not recommend the addition of air-polishing devices to conventional PMPR (curettes, ultrasonics, or both), in patients with peri-implant mucositis, even though these devices have shown efficacy when used as a single mode of treatment.

In patients with peri-implant mucositis, we suggest not to add diode lasers with conventional PMPR (curettes, ultrasonics, or both).

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (15.2% abstentions due to potential Col)

Background

Intervention

Professionally administered PMPR therapy aims at reducing soft tissue inflammation by removing hard and soft deposits from the surface of dental implants and/or its supra-structure. Combinations of PMPR therapy have been used and include laser adjunctive to ultrasonics and curettes, and air-polishing adjunctive to ultrasonics. The endpoint of treatment is absence of inflammation, i.e., BOP and suppuration.

Available evidence

Number and design of included studies

Three RCTs addressed the PICOS question (n=313 patients). Two RCTs analysed the effect of laser therapy adjunctive to ultrasonics and curettes (n=289), and one RCT analysed the effect of air-polishing adjunctive to ultrasonics (n=24), all with a 3-month follow-up. One study compared ultrasonics with carbon fibre tip plus glycine powder air polishing versus ultrasonics alone (n=24). The results on mean BOP severity were 1.1 and 1.0, respectively. The second study (n=220) compared ultrasonics with carbon fibre tip and titanium coated curettes with and without diode laser (980 nm) application. Results were 34.5% and 30.9% disease resolution, respectively. BOP extent at 3-months were 23.2% and 26.8%, respectively. The third study (n=69) compared ultrasonic with plastic tips and plastic curettes with and without diode laser (810 nm). The reported BOP extent was 0.26 and 0.57 respectively at 3 months, being this difference statistically significant.

Risk of bias

Study quality assessment identified some concerns of risk of bias in two studies, and a third had a high risk of bias.

Effect sizes and their clinical relevance

One RCT reported disease resolution/treatment success in 30.9-34.6 % and 23.2-26.8 % bleeding on probing extent at 3 months. Another RCT reported BOP extent of 0.26 and 0.57 in favour of adjunctive laser at 3 months, which was statistically significant. The third RCT reported BOP severity of 1.0 and 1.1 at 3 months.

Consistency

Evidence was consistent in the studies with a reduction in BOP, but statistically significant only in one of the RCTs with laser therapy adjunctive to ultrasonics and curettes. No patientreported outcomes were reported.

Balance of benefit and harm

An overall consideration of the benefit versus harm of professionally administered non-surgical mechanical/physical therapy supports the recommendation.

Overall certainty of the evidence Moderate.

From evidence to recommendation- additional considerations

Acceptability

Patients usually accept and understand the need for treatment.

Feasibility

Non-surgical mechanical/physical treatment of peri-implant mucositis can be performed by dental hygienists, general dentists as well as specialists.

Ethical Considerations

Not applicable.

Economic Considerations

Additional costs associated with adjunctive laser therapy may not be justified.

Legal Considerations Not applicable. R6.6 | In patients with peri-implant mucositis, what is the effect of repeating PMPR procedures, compared to a single administration of PMPR?

PICOS question addressed by a SR

R6.6 – Expert consensus-based recommendation

In patients with peri-implant mucositis, **we recommend** repeating PMPR if the endpoints of therapy have not been achieved within 3 months after the administration of PMPR. These endpoints and the evaluation times should be modified according to the patient's oral hygiene, risk factor profile and the cleansability of the prosthesis.

Supporting literature No studies evaluating the impact of repeated PMPR on periimplant mucositis outcomes were identified [37]

Quality of evidence No evidence from clinical studies identified.

Grade of recommendation Grade $A - \uparrow \uparrow$

Strength of consensus Unanimous consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

In patients with peri-implant mucositis, **we recommend** repeating PMPR if the endpoints of therapy have not been achieved within 3 months after the administration of PMPR. These endpoints and the evaluation times should be modified according to the patient's oral hygiene, risk factor profile and the cleansability of the prosthesis.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

If the endpoint of professionally administered non-surgical mechanical/physical therapy is not met following an intervention, it may be advisable to repeat the treatment.

Available evidence

There are no available RCTs or any observational study (with n=30 patients or more), or single arms from RCTs (with n=10 patients or more) evaluating the effect of repeated PMPR in the treatment of peri-implant mucositis.

Risk of bias

Not applicable.

Effect sizes and their clinical relevance

No RCTs were available, but in one of the included trials [110], results were reported at multiple time-points after providing repeated mechanical instrumentation. After an initial reduction of 20.9-28.6% in BOP extent, the effect of further repetitions was limited (1.9-6.3%, and 0.0-11.3%, respectively).

Consistency

Not applicable.

Balance of benefit and harm Not applicable.

Overall certainty of the evidence Not applicable.

From evidence to recommendation- additional considerations Acceptability Not applicable.

Feasibility

Non-surgical mechanical/physical treatment of peri-implant mucositis can be performed by dental hygienists, general dentists as well as specialist.

Ethical Considerations

Not applicable.

Economic Considerations Not applicable.

Legal Considerations

Not applicable.

R6.7| In patients with peri-implant mucositis, what is the effect of modifying the implant-

supported prosthesis to enable oral hygiene access?

Question not addressed by the SR

R6.7 – Evidence -based recommendation

In patients with peri-implant mucositis where the implant-supported prosthesis does not allow for proper self-performed and/or professional cleansability, **we recommend** cleaning/removal/modification of the prosthesis.

Supporting literature [111, 112]

Quality of evidence High (one RCT with low risk of bias, n=45)

Grade of recommendation Grade A – $\uparrow\uparrow$

Strength of consensus Unanimous consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

In patients with peri-implant mucositis where the implant-supported prosthesis does not allow for proper self-performed and/or professional cleansability, **we recommend** cleaning and/or removal/modification of the prosthesis as necessary.

BSP implementation additional background text:

Feasibility

Prosthesis modification may be implemented by general dentists as well as specialists.
Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

Modification of the implant-supported prosthesis to improve accessibility for oral hygiene and biofilm removal in surfaces of dental implants and restorative components.

Available evidence

There is one RCT (n=45) [112] evaluating the adjunctive effect of modifying the prosthesis to enable adequate oral hygiene. An additional publication reports on the 30-month follow up of the same study [111].

Risk of bias

Low risk of bias.

Effect sizes and their clinical relevance

Results at 6 months demonstrated reductions in the modified bleeding index of 1.14 and 0.50 for test and control groups, respectively, being these differences statistically significant; and, at 6 months, disease resolution was 66.6% and 9.6%, respectively.

Consistency

Not applicable.

Balance of benefit and harm

There is a clear benefit and minimal harm in the prosthesis modification to improve access for biofilm control.

Overall certainty of the evidence

Limited due to the scarcity of the available evidence.

From evidence to recommendation- additional considerations

Acceptability

Well accepted intervention, although patients may complain for a short time of food entrapment.

Feasibility

Prosthesis modification should be implemented by general dentists as well as specialist.

Ethical Considerations Not applicable.

Economic Considerations Not applicable.

Legal Considerations

Not applicable.

R6.8 | In patients with peri-implant mucositis, what is the efficacy of locally administered antibiotics adjunctive to PMPR?

PICOS ques	tion ac	ldressed by a	SR						
R6.8 – Evidence-based recommendation									
In patients administere			mucositis,	we	recommend	not	to	use	locally
Supporting	literat	ure [29, 113]							
Quality of evidence No direct evidence available									
Grade of re	comm	endation Gra	de A – ↓↓						

Strength of consensus Unanimous consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

We **recommend not to** use locally administered antibiotics in patients with periimplant mucositis.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

Professional administration of topical antibiotics, with sustained drug release, following nonsurgical mechanical/physical therapy in patients with peri-implant mucositis.

<u>Available evidence</u>

Number and design of included studies

No study could be identified when considering the inclusion criteria outlined in the systematic review [29]. However, one RCT (n=32) evaluated the adjunctive effect of minocycline microspheres in the treatment of peri-implant mucositis/incipient peri-implantitis (bone loss less or equal to three threads). Results showed a significant added effect in reducing BOP and PD at 6 months. However, BOP relapsed after 9 months (Renvert et al., 2006).

Risk of bias Not applicable.

Effect sizes and their clinical relevance Not applicable. Consistency

Not applicable.

Balance of benefit and harm

Harm versus benefit considerations on the use of antibiotics need to be undertaken. The use of antibiotics should always meet the antibiotic stewardship guideline.

Overall certainty of the evidence Not applicable.

From evidence to recommendation- additional considerations Acceptability Not applicable.

Feasibility

Not applicable.

Ethical Considerations

The use of antibiotics should always meet the antibiotic stewardship guideline.

Economic Considerations

High economic costs and limited availability of products in European countries need to be considered.

Legal Considerations Not applicable.

R6.9 | In patients with peri-implant mucositis, what is the efficacy of other locally administered agents adjunctive to PMPR?

PICOS question addressed by a SR

R6.9 – Evidence-based recommendation

In patients with peri-implant mucositis, we suggest not to use locally administered

agents (antiseptics, "postbiotics", desiccant gel) as adjuncts to PMPR

Supporting literature [29, 114, 115] and expert opinion

Quality of evidence Low

Grade of recommendation Grade B – \downarrow

Strength of consensus Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

In patients with peri-implant mucositis, **we suggest not to** use professionally applied locally administered agents (antiseptics, "postbiotics", desiccant gel) as adjuncts to PMPR

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (7.7% abstentions due to potential Col)

Background

Intervention

Professional administration of topical antiseptics (hydrogen peroxide, chlorhexidine, delmopinol hydrochloride, sodium hypochlorite, chitosan, acids, "postbiotics") following nonsurgical mechanical/physical therapy in patients with peri-implant mucositis. "Postbiotics" are products of the metabolic activity of the micro-organism, which, by exerting an antioxidant action, lead to a positive effect on the host [11]; in contrast with probiotics, they do not include alive microorganisms.

Available evidence

Number and design of included studies

Two RCTs were selected by the systematic review [29]. One of them (n=37 patients) assessed the professional administration of 0.12% CHX in 119 implants over the time periods of 1, 3, and

6 months in non-smokers. In the control group, CHX was not professionally applied. Outcome measures compared PD, BOP, and visible plaque index (PII). Disease resolution, SOP, and PROMs were not reported. In both the control and test group, significant reductions in PD, BOP, and visible PII were observed when comparing values at baseline with values at the 3- and 6-month follow-up. The inter-group comparison revealed no differences when comparing test and control groups [117]. The second one (n=46 patients) tested the professional administration of 0.95% NaOCl in 68 implants over time-period of 1, 3 and 6 months. In the control group, NaOCl was not applied. Outcome measures compared reduction in BOP, PD, and a modified PII. In addition, disease resolution was evaluated. Significant reductions in BOP, PD, and the modified PII for oral implants in both the test and control group at the 6-month follow-up. The inter-group comparison did not show differences among groups regarding BOP, disease resolution, PD, or the modified PII [118]. Changes in SOP as well as PROMs were not reported. Thus, the main finding of the RCTs identified in the systematic review were that 0.12% CHX or 0.95% NaOCl did not additionally improve clinical outcomes.

Apart from the evidence included in the systematic review, three additional RCTs were considered, evaluating the adjunctive effect of an antiseptic (CHX chip), a "postbiotic" (*"Lactobacillus Ferment"*), and a desiccant gel/liquid (concentrated aqueous mixture of hydroxybenzenesulphonic and hydroxymethoxybenzene acids, together with sulphuric acid), as adjunctives to mechanical therapy, compared with mechanical therapy plus application of 1% CHX gel [114, 115, 119]. Their quality of evidence was considered as low. The results reported for the "postbiotic", a significant reduction of PD and BOP and gingival bleeding index scores at 6 months, with no significant differences between groups [114]. For the desiccant gel/liquid, a significant reduction BOP and modified bleeding index was reported [119]. For the CHX-chip, BOP and PD were significantly reduced in CHX-chip group at 6 months [115]. The effect of the 1% CHX was heterogeneous, being beneficial in two studies [114, 119] but not in the third [115].

Risk of bias

For CHX and NaOCl, study quality assessment using the RoB 2 tool identified a low risk of bias for both studies included [117, 118]. For the other three RCTs, risk of bias was not evaluated, since they were not included in the systematic review.

Effect sizes and their clinical relevance

For CHX, based on one RCT (n=37 patients), no additional effect of 0.12% CHX was demonstrated regarding reductions in BOP, PD, and PlI. For NaOCl, based on one RCT (n=46 patients), no additional effect of 0.95% NaOCl was identified regarding reductions in BOP, PD, and PlI. For the "postbiotic" gel, based on one RCT (n=20 patients), no additional effect was demonstrated [114]. For the desiccant solution, based on one RCT (n=23 patients), significant differences between groups were only observed for plaque indices [119]. In this study, no additional effect of the desiccant was shown when compared to control CHX gel application [119]. The effect of the desiccant on Pl was unclear [119]. For the CHX chip, based on one RCT (n=32 patients), significant additional benefits in BOP were observed in the test group, but the statistically significant differences observed at baseline precluded a strong conclusion on the adjunctive effect [115].

Consistency

Not applicable.

Balance of benefit and harm

In both identified studies, the adjunctive professional administration of 0.12% CHX or 0.95% NaOCl did not cause unintentional side effects that suggest harm to the patient [117, 118]. Thus, formulations of both CHX and NaOCl may be considered as a professional treatment adjunctive to non surgical mechanical/physical therapy in the treatment of peri-implant mucositis. Future studies are needed to further investigate the efficacy of the given and other concentrations of CHX and NaOCl. For CHX, several adverse effects, such as taste alteration, mouth numbness, xerostomia, and tooth discoloration have been reported [120]. For NaOCl, the occurrence of potential adverse effects is uncertain for various concentrations. Potential adverse side effects must be considered to balance benefits and harms. For the "postbiotic gel", potential side effects, such as allergic reactions, cannot be excluded. For the CHX chip, numerous unintentional side effects are listed in the product information, but they are reported to be not frequent and usually mild. For the desiccant, no unintentional side effect was reported [119], however, potential side effects of sulfuric acid are listed by the company,

and thus, the application is not recommended in patients if allergic to sulphur in any form and in the case of pre-existing skin disorders.

Overall certainty of the evidence

The certainty is weak, and the quality of evidence is graded as very low based on the lack of studies.

From evidence to recommendation- additional considerations

Acceptability

In general, the application of antiseptics is well accepted by patients when understanding the pathogenesis of peri-implant mucositis.

Feasibility

CHX gels, CHX chips, desiccant materials, "postbiotics", and NaOCl formulations can be professionally applied by the general dentist or specialist. Their adjunctive use is not clinically demanding or time-consuming. For the NaOCl formulation (PeriSolv®, RLS Global AB, Mölndal, Sweden); CHX chip (PerioChip, Karr Dental, Wollerau, Switzerland), the "postbiotic" (Biorepair Parodontgel Intensive, Coswell SPA, Funo di Argelato, BO, Italy), and the desiccant liquid (HybenX® Oral Tissue decontaminant™, EPIEN Medical Inc., Saint Paul, MN, USA), specific brands were tested and the information provided may only be valid for those products, that may not be available in all markets.

Ethical Considerations

Based on the available evidence, no evaluation of ethical aspects could be performed.

Economic Considerations

CHX gels, CHX chips, desiccant materials, "postbiotics", and NaOCl formulations are associated with additional costs to the patient as well as to the dental professional team. The application of any antiseptic treatment adjunctive to non-surgical mechanical/physical therapy may lead to additional costs for the patients depending on individual health insurance plans in the individual countries. As examples, the additional costs associated with the use of the desiccant material, in Germany, are approximately $100 \in$ for two syringes of 1 mL each, and for

the use of CHX chips is approximately 300 € for 20 applications. No information on costeffectiveness could be retrieved from the RCTs [114, 115, 117-119].

Legal Considerations

The NaOCl formulation (PeriSolv[®]) is approved as Class I medical device in the European Union, and the desiccant material (HybenX[®]) has also been approved as Class I medical device in the European Union and Canada. The implications of the use in other geographical locations or the use for indications besides the ones approved are unclear.

R6.10 | In patients with peri-implant mucositis, what is the efficacy of locally administered

photodynamic therapy adjunctive to PMPR?

PICOS question addressed by a SR

R6.10 – Evidence-based recommendation

In patients with peri-implant mucositis, we suggest not to use photodynamic therapy

adjunctively to PMPR.

Supporting literature [29]

Quality of evidence Low (5 RCTs)

Grade of recommendation Grade B – \downarrow

Strength of consensus Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

In patients with peri-implant mucositis, **we suggest not to** use photodynamic therapy adjunctively to PMPR.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (1.8% abstentions due to potential Col)

Background

Intervention

Application of antimicrobial photodynamic therapy (aPDT) adjunctive to non-surgical mechanical/physical therapy in patients with peri-implant mucositis.

Available evidence

Number and design of included studies

For the application of aPDT adjunctive to submarginal instrumentation, 5 RCTs (in total, n=204 patients) analysed an estimated number of 231 implants over a time-period of 3 months [29]. Of these five studies on adjunctive application of aPDT four included patients with habitual tobacco intake (cigarette smokers, smoke-less tobacco chewers and vaping individuals). In the control groups, aPDT was not applied. In the test groups, the intervention varied in terms of a range in the applied wavelength between 660 nm and 670 nm, power density between 100 milliwatts (mW) and 150 mW. One study did not report on treatment modalities. Outcome measures compared BOP, PD, and Pll. Disease resolution and PROMs were not reported. In the synthesis of data, three studies were evaluated for changes in BOP and PD and four studies for changes in Pll, comparing test and control groups. For BOP and PD, no difference was identified between test and control groups, whereas for PI, a significant difference was shown in favour of aPDT adjunctive to submarginal instrumentation [29]. High heterogeneity as well as a high level of asymmetry were evident [29]. Two RCTs were excluded due to the lack of reporting mean and standard deviation or assessing a modified bleeding index instead of BOP. Changes in SOP as well as PROMs were not reported. The main findings were that aPDT did not additionally improve clinical outcomes for changes in BOP, PD, or Pll.

Risk of bias

For aPDT, study quality assessment using the RoB 2 tool identified a low risk of bias for one study, whereas some concerns indicated a risk of bias in four studies on aPDT.

Effect sizes and their clinical relevance

For aPDT, based on three RCTs (204 patients) included in the meta-analysis, no additional effect of the adjunct application of aPDT was demonstrated regarding reduction in BOP and PD [29]. A significant reduction of PII was identified in the meta-analysis, however, clinically, this

reduction was not related to the reduction of surrogate parameters for disease resolution (reduction or absence of BOP, reduction in PD) [29].

Consistency

For aPDT, the identified RCTs included male patients only, and from these 5 RCTs 4 focused on patients with habitual tobacco intake (cigarette smokers, smoke-less tobacco chewers and vaping individuals). The analysis of data revealed high heterogeneity among the studies [2]). This inconsistency among the studies may be explained by the heterogeneity of reported outcome parameters as well as regarding the variation of tobacco intake habits, even though only male patients were evaluated. In addition, the intervention varied in terms of a range in the applied wavelength between 660 nm and 670 nm, power density between 100 milliwatts (mW) and 150 mW, and choice of photosensitizer (phenothiazine chloride, methylene blue) in the respective test groups.

Balance of benefit and harm

For the additional application of aPDT adjunctive to submarginal instrumentation, no benefit was identified in the meta-analysis [29]. Potential harm of aPDT adjunctive to submarginal instrumentation has not been studied to date. However, potential adverse effects cannot be entirely ruled out due to various wavelength, power density, and photosensitizer available on the market.

Overall certainty of the evidence

The overall certainty regarding the additional effect of aPDT is weak. The quality of evidence is low.

From evidence to recommendation- additional considerations

Acceptability

The adjunct application of aPDT is accepted by patients when understanding the pathogenesis of peri-implant mucositis.

Feasibility

The application of aPDT can only be performed by a trained operator and appropriate eye protection must be used by the dental professional team and the patient.

Ethical Considerations

Not applicable.

Economic Considerations

The application of aPDT causes comparatively high costs for the dental team with regard to the acquisition and maintenance of the corresponding equipment. For the patient, aPDT adjunctive to submarginal debridement may lead to additional costs depending on individual health insurance plans in the individual countries. No information on cost-effectiveness could be retrieved from the five selected RCTs. Additional costs associated with adjunctive laser therapy may not be justified.

Legal Considerations Not applicable.

R6.11 | In patients with peri-implant mucositis, what is the efficacy of patient selfadministered antiseptics adjunctive to PMPR?

PICOS question addressed by a SR

R6.11 – Evidence-based recommendation

In patients with peri-implant mucositis the time limited self-administration of oral rinse

antiseptics (chlorhexidine and herbal-based) adjunctive to PMPR may be considered.

Supporting literature [31]

Quality of evidence Moderate (6 RCTs, using different antiseptic agents, CHX and herbalbased).

Grade of recommendation Grade O – \leftrightarrow

Strength of consensus Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

In patients with peri-implant mucositis the time limited self-administration of oral rinse antiseptics (chlorhexidine and herbal-based) adjunctive to PMPR **may be considered**.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (26.2% abstentions due to potential Col)

Background

Intervention

Application of antiseptics adjunctive to PMPR in patients with peri-implant mucositis.

Available evidence

Number and design of included studies

The systematic review [31] included five RCTs evaluating the effect of antiseptics used by the patient adjunctively to submarginal instrumentation compared to submarginal instrumentation alone or combined with a negative control or a placebo in terms on BOP reduction at 3 months in patients with peri-implant mucositis.

In these five RCTs, self-administered antiseptics as adjuvant to PMPR were used in the format of gels (0.5% CHX) or mouth rinses. In this latter delivery format (mouth rinses), the following active agents have been tested: CHX at different concentrations (0.03%, 0.12% or 0.2%) alone or combined with CPC (0.05%); herbal-based mouth rinses; delmopinol (0.2%).

Risk of bias

The overall risk of bias of the included studies was judged as 'low' (Rob2 tool), with all five studies with a low risk of bias.

Effect sizes and their clinical relevance

For CHX gel: 29 received PMPR at the implant sites were instructed to brush around the implant twice daily using a chlorhexidine gel (0.5%) (n=15) or a placebo gel (n=14) for a period of 4 weeks, and there were significant reductions in the mean number of sites with BOP from baseline to 1 month for both test and control groups (P <0.05), with little apparent change

between 1 and 3 months (p>0.1); there was no statistically significant difference in the changes in BOP between the test and control groups at 1 month or at 3 months (p>0.1).

For CHX mouth rinses, four RCTs with 166 patients compared the efficacy of self-administered CHX mouth rinses versus distilled water/saline or placebo, for 2 weeks, 1 month or 1 year, and the results showed significant reductions over time of BOP, with conflicting results in terms of superiority versus control. Statistically significant differences in BOP or in modified gingival index (MGI) were noted after 3 months, while no statistically significant differences in terms of BOP were reported at 1 month or with the usage of 0.05% CHX plus 0.05% cetylpyridinium chloride (CPC) at 1 year [121].

For herbal mouth rinses, two RCTs with 62 patients were managed with self-administered herbal-based mouth rinses for 2 weeks or NaCl/distilled water. At 3 months, statistically significant differences in BOP and in MGI, between test and control groups were reported, with better performance in the herbal mouth rinse groups.

For delmopinol, one RCT analysed the efficacy of 1-month self-performed delmopinol mouth rinse versus placebo, with 59 patients. Both treatments showed reduction on BOP with no differences among test and control groups.

Consistency

Conflicting results were reported when using CHX.

Balance of benefit and harm

In the included studies, some antiseptics have been associated with undesirable side effects, such as transient anaesthetic sensation in the oral mucosa (delmopinol) or higher levels of staining on the teeth or tongue (CHX). Moreover, other rarer side effects cannot be excluded.

Overall certainty of the evidence Low.

From evidence to recommendation- additional considerations

Acceptability

Antiseptics are widely accepted by the population.

Feasibility

There are no perceived barriers.

Ethical Considerations

The issue has not been addressed. There are no perceived ethical considerations.

Economic Considerations

For dentifrices, it may not be relevant since it is always combined with mechanical tooth brushing. For mouth rinses use, the extra cost should be taken into consideration.

Legal Considerations

It should also be noted that the evidence base contains studies using products that may no longer be available.

R6.12 | In patients with peri-implant mucositis, what is the efficacy of patient selfadministered probiotics adjunctive to PMPR?

PICOS question addressed by a SR

R6.12 – Evidence-based recommendation

In patients with peri-implant mucositis the professionally guided self-administration of

probiotics may be considered as adjunctive to PMPR.

Supporting literature [31]

Quality of evidence Moderate (6 RCTs)

Grade of recommendation Grade O – \leftrightarrow

Strength of consensus Consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

In patients with peri-implant mucositis, professionally guided self-administration of probiotics in patients with peri-implant mucositis, **may be considered** as adjunctive to *PMPR*.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (2.2% abstentions due to potential Col)

Background

Intervention

Adjunctive probiotic tablets containing *Lactobacillus reuteri*. In two trials, the adjunctive measurement was combined with a 0.12% CHX mouth rinse, 15 days before starting probiotics intake. The most frequent posology was one tablet per day for one month. In contrast, the shortest posology was two tables per day for three weeks and the longest, twice per day for three months.

Available evidence

Number and design of included studies

The systematic review [31] included six RCTs evaluating the effect of systemic probiotic used by the patient, adjunctively to submarginal instrumentation, compared to submarginal instrumentation alone or combined with a negative control or a placebo, in terms of BOP reduction at 3 months in patients with peri-implant mucositis.

Risk of bias

The overall risk of bias of the included studies was judged as 'low' (Rob2 tool), with three studies with a low risk of bias and three with a moderate risk of bias.

Effect sizes and their clinical relevance At 3 months, results revealed:

- Statistically significant greater reduction in BOP (%) for probiotics (L. reuteri) than controls (n_{studies}=6; n_{patients}=260; WMD=12.11%; 95% CI [3.20; 21.03]; p=0.008; l²=93.3%).
- Statistically significant greater reduction in plaque (%) for probiotics (L. reuteri) than controls (n_{studies}=6; n_{patients}=260; WMD=14.20%; 95% CI [3.46; 29.94]; p=0.01; I2=92.4%).
- No statistically significant differences in PD reductions.
- Complete disease resolution was only reported in one study (32% after 135 days, without differences between test and control groups).

At 6 months, no statistically significant differences were found when comparing probiotics versus control groups for any study outcome. No adverse events were reported due to the adjunctive use of L. reuteri tablets.

Consistency

All studies reported the same tendency.

Balance of benefit and harm

No adverse events have been reported. Clear benefits observed at 3 months, although they were not sustained at 6 months.

Overall certainty of the evidence

Moderate.

From evidence to recommendation- additional considerations

Acceptability

Systemic probiotics are still not widely accepted by the population.

Feasibility

There are no perceived barriers.

Ethical Considerations

There are no perceived ethical considerations.

Economic Considerations

There are no perceived economic considerations, although an extra economic cost is derived from the prescription of the probiotics.

Legal Considerations

There are no perceived legal considerations.

R6.13 | In patients with peri-implant mucositis, what is the efficacy of the oral administration of systemic antibiotics when used adjunctively to PMPR?

PICOS question addressed by a SR

R6.13 – Evidence-based recommendation

Due to concerns about patients' health and the impact of systemic antibiotic use to

public health, in patients with peri-implant mucositis we recommend not to use

Supporting literature [25, 31] and antibiotic stewardship.

Quality of evidence Low (3 RCTs)

Grade of recommendation Grade A – $\downarrow \downarrow$

Strength of consensus Unanimous consensus (0% of the group abstained due to potential Col)

potential COI)

BSP Implementation

This evidence-based recommendation is adopted.

Due to concerns about patients' health and the impact of systemic antibiotic use to

public health, in patients with peri-implant mucositis **we recommend not to** use

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

The following systemic antibiotics (prescribed as oral administration) as adjuvants to submarginal instrumentation have been tested in the treatment of peri-implant mucositis:

- Azithromycin (500 mg the first day and 250 mg, from the 2nd to 4th day).
- Amoxicillin (500 mg, thrice daily for one week).

Available evidence

Number and design of included studies

The systematic review [31] included three RCTs evaluating the effect of systemic antibiotics prescribed as oral administration adjunctively to submarginal instrumentation. In one study, amoxicillin was compared to submarginal instrumentation combined with probiotics. In another study, the adjunctive administration of azithromycin was compared with instrumentation alone. In the third study, the adjunctive use of azithromycin plus a 0.12% CHX mouth rinse was compared with instrumentation plus a 0.12% CHX mouth rinse. Outcomes evaluated in these three studies were the percentage of BOP, plaque index, and PD.

Risk of bias

The overall risk of bias in the included studies was judged as 'moderate' (Rob2 tool), with all the three studies with a moderate risk of bias.

Effect sizes and their clinical relevance

At 3 months, results revealed:

- Statistically significant greater reduction in BOP (%) for antibiotics than controls (n_{studies}=3; n_{patients}=101; WMD=5.97%; 95% CI [1.34; 10.59]; p=0.012; I2=58.1%).
- Statistically significant greater reduction in plaque (%) for antiseptics than controls (n_{studies}=3; n_{patients}=101; WMD=14.74%; 95% CI [3.83; 25.65]; p=0.008; I2=83.2%).
- Statistically significant differences in the reduction in PD (mm) for the use of systemic antibiotics than controls only for one study (n_{studies}=1; n_{patients}=28; mean difference [MD]=1.8 mm; 95% CI [1.37; 2.23]; p<0.001).

- Complete disease resolution was rarely reported. One study reported at three months an OR of 4.5 (95% CI [1.2; 17.0]; p<0.05) of favourable treatment in favour of systemic azithromycin in comparison with the control group.

At six months, the results were the following:

- Statistically significant greater reduction in BOP (%) for antibiotics than controls (n_{studies}=2; n_{patients}=71; WMD=20.79%; 95% CI [15.24; 26.34]; p<0.001; I2=30.60%).
- Statistically significant greater reduction in plaque (%) for antiseptics than controls (n_{studies}=2; n_{patients}=7; WMD=13.97%; 95% CI [4.10; 23.84]; p=0.006; I2=30.6%).
- Only one study using amoxicillin reported statistically significant differences with control group (n_{studies}=1; n_{patients}=28; MD=2.60 mm; 95% CI [2.20; 3.00]; p<0.001).

No studies reported a longer follow-up than six months.

Consistency

All studies reported the same tendency.

Balance of benefit and harm

In one study that collected side effects, no adverse events were observed after antibiotic intake. No specific concerns can be raised for antibiotics as adjunctive use for treating periimplant mucositis.

Overall certainty of the evidence Moderate.

From evidence to recommendation- additional considerations

Acceptability

The population widely accepts antibiotics. Nevertheless, there is an issue related to the need of diminishing the usage of antibiotics due to the potential risks associated with antibiotic resistance.

Feasibility

There are no perceived barriers.

Ethical Considerations

The issue has not been addressed. There are no perceived major ethical considerations. Yet it must be reiterated the need of containing prescription of antibiotics for the population at large.

Economic Considerations

The specific economic considerations can be stated.

Legal Considerations

No specific legal consideration can be stated.

7 | RECOMMENDATIONS FOR NON-SURGICAL MANAGEMENT OF PERI-IMPLANTITIS

7.1 | Introduction - general recommendations in the *non-surgical step* of peri-implantitis treatment

The management of peri-implantitis is a relatively new area of research and clinical practice. Although key differences impacting care between peri-implantitis and periodontitis have been identified, the theoretical foundation of peri-implantitis treatment is based on the successful approaches developed for the treatment of periodontitis. Therefore, a step-by-step approach may be appropriate, as it has been suggested for the treatment of periodontitis [25], and described in Section 4 of the present CPG. Thus, the interventions included in the systematic reviews of Working Group #3 [27, 28, 33] are part of the *non-surgical step* of peri-implantitis treatment.

This stepwise approach mirrors the one used in periodontal therapy [25], and the included interventions are also similar to those proposed for periodontitis. The main objective of the *non-surgical step* of peri-implantitis treatment is to control peri-implant biofilms and inflammation, and therefore the central intervention would be submarginal instrumentation. In addition, interventions focusing on supramarginal biofilm control or on risk factor control, are also part of the *non-surgical step* of peri-implantitis treatment.

After delivery of treatment, progress in controlling inflammation and suppuration should be monitored, and the outcomes should be re-assessed. While in periodontitis treatment, endpoints of therapy have been well established, and success of steps 1 and 2 of treatment is a reasonable expectation [122], comparable evidence for the treatment of peri-implantitis is still scarce. The rationale for using a stepwise approach and for a non-surgical phase of peri-implantitis treatment, therefore, comes from i) attempting biofilm and inflammation control with relatively simple approaches before escalating treatment complexity and invasiveness; ii) the fact that subjects with peri-implantitis frequently present with poorly controlled periodontitis that requires a concomitant stepwise treatment approach; and iii) the ability to deliver any surgical treatment at a later step and in a subject with better biofilm and risk factor control.

R7.1 | Is peri-implantitis treatable?

Additional question addressed by the WG

R7.1 – Expert consensus-based recommendation

In patients with peri-implantitis, **we recommend** therapy to retain an individually acceptable implant/prosthesis as the first line of treatment.

We recommend that peri-implantitis therapy starts with a non-surgical step, followed by re-evaluation and, depending on the outcomes, progress to the surgical step or to SPIC.

Supporting literature [27, 28, 33] and Expert opinion

Quality of evidence Moderate – indirect evidence derived from 15 RCTs, with at least

6-month follow up (10 with low, 3 with some concerns and 2 with high risk of bias)

Grade of recommendation Grade $A - \uparrow \uparrow$

Strength of consensus: Strong consensus (0% of the group abstained due to

potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

In patients with peri-implantitis, **we recommend** therapy to retain an individually acceptable implant/prosthesis as the first line of treatment.

We recommend that peri-implantitis therapy starts with a non-surgical step, followed by re-evaluation and, depending on the outcomes, progress to the surgical step or to SPIC.

Additional BSP Comment

Certain aspects of the treatment of PI may require specific training.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

The interventions for treating peri-implantitis differ among studies, but they most commonly include submarginal instrumentation and peri-implant biofilm control [27, 28, 33] in both test and control groups.

Available evidence

Number and design of included studies

In the SRs prepared for the present project [27, 28, 33], 15 RCTs with at least 6-month follow up, were considered as valid for developing recommendations. For the present recommendation, outcomes from both test and control groups are considered.

Risk of bias

Ten presented with low risk of bias, three with some concerns and two with high risk.

Effect sizes and their clinical relevance

The observed improvements after treatment are significant in magnitude and consistent across the considered RCTs. Taken together, the evidence is unlikely to arise from the placebo or the

Hawthorne effect. Still, it is not possible to assess the relative contribution of the different components that have been tested.

Consistency

Not applicable.

Balance of benefit and harm

Benefits were observed in both the test and control groups. Out of 17 test groups, statistically significant benefits were observed in 11 for PD reduction and 9 for BOP. Out of 17 control groups, statistically significant benefits were observed in 11 for PD reduction and 7 for BOP. The percentage of disease resolution was provided by seven test groups (ranging 0%-65%) and seven control groups (ranging 14%-55%). Limited evidence of harm was presented.

Overall certainty of the evidence Moderate.

From evidence to recommendation-additional considerations

Acceptability

The interventions for treating peri-implantitis seem to be acceptable for patients, health providers, and health authorities, although no direct evidence is available.

Feasibility

The interventions for treating peri-implantitis are feasible, although some of them may need specific training.

Ethical Considerations

The interventions for treating peri-implantitis may negatively impact equity, if public services are not covering the cost, and those will need to be directly covered by patients.

Economic Considerations

Limited evidence is available, see Section 1.

Legal Considerations

Not applicable.

R7.2 | Which interventions should be provided as part of the non-surgical step of periimplantitis treatment?

Additional question addressed by the WG

R7.2 – Expert consensus-based recommendation

We recommend that the following interventions should be provided as part of the nonsurgical step of peri-implantitis:

- Oral hygiene instructions and motivation.
- Risk factor control.
- Prosthesis cleaning/removal/modification including controlling biofilm retentive factors and evaluation of the components of the prosthesis, whenever needed and feasible.
- Supramarginal and submarginal instrumentation.
- Concomitant periodontal therapy as needed.

Supporting literature [27, 28, 33] and Expert opinion

Quality of evidence Low – indirect evidence derived from 15 RCTs, with at least 6-month follow up (10 with low, 3 with some concerns and 2 with high risk of bias)

Grade of recommendation Grade A – ↑↑

Strength of consensus: Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

We recommend that the following interventions should be provided as part of the nonsurgical step of peri-implantitis:

- Oral hygiene instructions and motivation.
- Risk factor control.
- Prosthesis cleaning/removal/modification including controlling biofilm retentive factors and evaluation of the components of the prosthesis, whenever needed and feasible.

- Supramarginal and submarginal instrumentation.
- Concomitant periodontal therapy as needed.

Clarification for BSP implementation:

Prosthesis cleaning/removal/modification including controlling biofilm retentive factors and evaluation of the components of the prosthesis, whenever needed and feasible (e.g., if the prosthesis does not allow proper biofilm removal and if the treating dentist has the necessary tools and expertise to remove/modify it).

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

The group identified interventions within those detailed and performed in test and control groups of the 15 RCTs included in the three SRs [27, 28, 33]. Among them, the most relevant were selected, and placed in chronological sequence:

- Oral hygiene instructions and motivation, see Section 5.
- Risk factor control, see Section 6.
- Prosthesis cleaning/removal/modification, including controlling biofilm retentive factors and evaluation of the components of the prosthesis, whenever needed and feasible. If renewal is necessary, additional evaluation of the overall treatment planning should be made, considering the added costs and the cost-effectiveness ratio [16]
- Supramarginal and submarginal instrumentation. For the latter, for the present work, instrumentation performed with curettes and/or sonic/ultrasonic devices was considered as the basic/control intervention. Additional or alternative methods to clean/decontaminate the implant surface are discussed in the following recommendations.
- Concomitant periodontal therapy as needed. If periodontal diseases are detected, they should be properly managed, in particular periodontitis, which is a recognized risk factor

for peri-implantitis [5, 6]. Concomitant treatment of periodontitis should follow available guidelines [25].

Available evidence

Number and design of included studies

In the SRs prepared for the present project [27, 28, 33], 15 RCTs with at least 6 months of follow-up, were considered as valid for developing recommendations. For the present recommendation, both test and control groups are considered.

Risk of bias

Ten presented with a low risk of bias, three with some concerns, and two with high risk.

Effect sizes and their clinical relevance Not applicable.

Consistency

Not applicable.

Balance of benefit and harm

Benefits were observed in both test and control groups (see Background text of previous recommendation). Limited evidence of harm was presented.

Overall certainty of the evidence Low.

From evidence to recommendation- additional considerations

Acceptability

The interventions for treating peri-implantitis seem to be acceptable for patients, health providers and health authorities, although no direct evidence is available.

Feasibility

The interventions for treating peri-implantitis are feasible, although some of them may need specific training.

Ethical Considerations

The interventions for treating peri-implantitis may negatively impact equity if public services are not covering the cost, as in these situations they will need to be directly covered by patients.

Economic Considerations

Limited evidence is available, see Section 1.

Legal Considerations Not applicable.

R7.3 | Which are the endpoints of the non-surgical step of peri-implantitis treatment, and when and how should they be evaluated?

Additional question addressed by the WG

R7.3 – Expert consensus-based recommendation

1. To assess the outcome of the non-surgical step of peri-implantitis treatment, **we recommend** monitoring residual inflammation/suppuration and probing depths. Patient satisfaction, good oral hygiene, and prosthesis cleansability should also be considered.

2. We recommend using, at implant level, residual probing depths ≤ 5 mm with no BOP at more than one point* and no suppuration, as therapy endpoints.

3. If they are not achieved, we recommend considering additional treatment.

4. **We recommend** evaluating the outcome (re-evaluation) of the non-surgical step of therapy after 6-12 weeks; it may be prudent to monitor cases frequently during healing.

Supporting literature [27, 28, 33] and Expert opinion

Quality of evidence Low – indirect evidence derived from 15 RCTs, with at least 6 months of follow up (10 with low, 3 with some concerns and 2 with high risk of bias)

Grade of recommendation Grade A – $\uparrow\uparrow$

Strength of consensus: Unanimous consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

1. To assess the outcome of the non-surgical step of peri-implantitis treatment, **we recommend** monitoring residual inflammation/suppuration and probing depths. Patient satisfaction, good oral hygiene, and prosthesis cleansability should also be considered.

2. We recommend using, at implant level, residual probing depths ≤ 5 mm with no BOP at more than one point* and no suppuration, as therapy endpoints.

3. If they are not achieved, we recommend considering additional treatment.

4. **We recommend** evaluating the outcome (re-evaluation) of the non-surgical step of therapy after 6-12 weeks; it may be prudent to monitor cases frequently during healing.

Clarification for BPS implementation:

standard manual periodontal probe (0.5 mm tip).

Regarding point 2: threshold of ≤5 mm may not always be achievable dependent on baseline probing depth following implant placement)

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

* BOP present at a single spot, not line or profuse bleeding, is compatible with control of inflammation. It should be assessed at six sites per implant using gentle forces (0.2 N) with a

Background

Intervention

The group identified follow up intervals and outcomes among those described in test and control groups of the 15 RCTs included in the three SRs [27, 28, 33]. In addition, the findings of the ID-COSM project (see Section 2) were also considered [5]).

Available evidence

Number and design of included studies

In the SRs prepared for the present project [27, 28, 33], 15 RCTs with at least 6 months of follow-up, were considered as valid for developing recommendations.

Risk of bias

Ten presented with low risk of bias, three with some concerns and two with high risk.

Effect sizes and their clinical relevance Not applicable.

Consistency

Not applicable.

Balance of benefit and harm

Not applicable.

Overall certainty of the evidence Low.

From evidence to recommendation- additional considerations

Acceptability

The evaluation of the outcomes after the non-surgical step of peri-implantitis treatment seems to be acceptable for patients, health providers and health authorities, although no direct evidence is available.

Feasibility

The evaluation of the outcomes after the non-surgical step of peri-implantitis treatment seems to be feasible.

Ethical Considerations Not applicable.

Economic Considerations Not applicable.

Legal Considerations Not applicable.

7.2 | Non-surgical submarginal instrumentation - mechanical/physical cleaning/decontamination

The systematic review by Cosgarea and co-workers [27] focused on mechanical/physical approaches for implant surface cleaning/decontamination. Three PICOS questions were formulated, one to understand the efficacy of submarginal instrumentation versus no treatment or supramarginal instrumentation (PICOS #3) and two PICOS questions aimed to evaluate different mechanical/physical decontamination methods (e.g. air- polishing, sonic/ultrasonic devices, lasers), alone or in combination, compared to non-surgical submarginal instrumentation with/without placebo decontamination (non-aiming at mechanical/physical decontamination, e.g., scalers to remove hard deposits with adjunctive saline irrigation) with (PICOS #2) or without (PICOS #1) other concomitant interventions.

The review initially identified nine RCTs, but for the consensus report seven RCTs were finally considered, five [123-127] assessing various types of laser therapy (i.e. Nd:YAG, diode laser, Er,Cr:YSGG, Er:YAG), and two [128, 129] assessing an air-abrasive decontamination system. Two presented a high risk of bias, and the other five a low risk of bias.

R7.4 | What is the efficacy of submarginal instrumentation in the non-surgical step of periimplantitis treatment?

PICOS question addressed by a SR

R7.4 – Evidence -based recommendation

In patients with peri-implantitis, we recommend performing non-surgical supra- and sub-

marginal instrumentation with curettes and/or sonic/ultrasonic devices.

Supporting literature [27, 28, 33] and Expert opinion

Quality of evidence Moderate - indirect evidence derived from 15 RCTs, with at least 6

months of follow up (10 with low, 3 with some concerns and 2 with high risk of bias)

Grade of recommendation Grade A – $\uparrow\uparrow$

Strength of consensus: Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence -based recommendation is adopted.

In patients with peri-implantitis, **we recommend** performing non-surgical supra- and submarginal instrumentation with curettes and/or sonic/ultrasonic devices.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (16.2% abstentions due to potential Col)

Background

Intervention

For the present CPG development process, the control intervention to evaluate non-surgical submarginal instrumentation approaches was defined as those approaches not aiming at mechanical/physical decontamination, which includes scalers or sonic/ultrasonic devices to remove hard deposits with/without adjunctive irrigation with an inactive solution (i.e. saline). For answering the proposed question, studies comparing control decontamination with no treatment or supragingival instrumentation were searched for. Since no direct evidence was found, indirect evidence derived from the control groups of the selected studies was used: in some control groups, in addition to submarginal instrumentation, additional interventions were included (that were also part of the treatment protocol in the test group), such as adjunctive decontamination with chlorhexidine digluconate as subgingival irrigation (0.1-0.2%), as

subgingival application (1% chlorhexidine digluconate gel) or as mouth rinsing (two weeks with 0.1-0.2% chlorhexidine digluconate) [126-128].

<u>Available evidence</u> Number and design of included studies No study was found answering this question.

Risk of bias Not applicable

Effect sizes and their clinical relevance

Due to the lack of studies, indirect evidence was used, analysing the clinical impact in control groups in the 15 RCTs identified in the three systematic reviews [27, 28, 33]. Out of 17 control groups, statistically significant benefits were observed in 11 for PD reduction and in seven for BOP. The percentage of disease resolution was provided for seven control groups, and it ranged 14%-55%. Limited evidence of harm was presented.

Consistency

Most control groups found a statistically significant impact of the treatment, this was similar to that reported in test groups.

Balance of benefit and harm

No proper evaluation of patient-reported outcome measures (PROMs) was carried out.

Overall certainty of the evidence Moderate.

From evidence to recommendation- additional considerations

Acceptability

There is no evidence so far for clinicians' or patients' acceptability.

Feasibility

Implementation of therapy may be negatively influenced by the lack of retrievability and/or shape of the prosthetic suprastructure.

Ethical Considerations

No data are available to address ethical considerations.

Economic Considerations

Cost-effectiveness has not been evaluated in these studies.

Legal Considerations

So far, if the manufacturer`s indications are respected, there are no legal considerations.

R7.5 | What is the efficacy of lasers in the submarginal instrumentation of the non-surgical step of peri-implantitis treatment?

PICOS question addressed by a SR

R7.5 – Evidence-based recommendation

We suggest not to use lasers, either adjunctively or as monotherapy, for non-surgical submarginal peri-implant instrumentation.

Supporting literature [27, 28, 33]

Quality of evidence Low – 5 RCTs (n=178 patients, n=225 implants) with a minimum

follow-up of 6 months (2 studies at high risk and 3 studies with low risk of bias)

Grade of recommendation Grade $B - \psi$

Strength of consensus: Unanimous consensus (0% of the group abstained due to potential Col)

001)

BSP Implementation

This evidence-based recommendation is adopted.

We suggest not to use lasers, either adjunctively or as monotherapy, for non-surgical

submarginal peri-implant instrumentation.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (2.3% abstentions due to potential Col)

Background

Intervention

Lasers have received significant attention as a method for submarginal instrumentation as they may enhance biofilm removal and/or surface decontamination. Lasers are a wide class of biomedical instruments, each one of them working based on specific principles. In the selected studies, different lasers have been tested, either alone as monotherapy (three studies) or as an adjunct to conventional submarginal instrumentation (two studies).

Available evidence

Five RCTs (n=178 patients, n=225 implants) with a minimum follow-up of 6 months, with various types of laser (Nd:YAG, diode laser, Er,Cr:YSGG, Er:YAG) assessed the submarginal peri-implant instrumentation with lasers alone or in combination with additional chlorhexidine irrigation [(123-127]. Two of them used Er:YAG laser as monotherapy, one study used Nd:YAG laser as monotherapy, two studies used diode laser adjunctive to mechanical decontamination with curettes, of which one study also had a group using Er,Cr:YSGG laser as an adjunctive treatment.

Risk of bias

Two studies were at high risk, and three studies at low risk of bias.

Effect sizes and their clinical relevance

Due to heterogeneity in the treatment protocol, no meta-analysis was carried out. All studies showed improvements in both test and control groups in PD and BOP, at 3 and/or 6 months compared to baseline. In general, studies showed no additional benefit from the application of lasers at 6 months, in terms of either PD or BOP reductions. Only in one study did the adjunctive application of a Er,Cr:YSGG laser show statistically significantly larger PD reductions at 6 months, compared to submarginal instrumentation alone [123-127]. An Er:YAG laser as

monotherapy [123-127] led to statistically significant differences in BOP. Their magnitude, however, was small.

Consistency

Positive results for the primary outcomes were observed in all five RCTs, for both control and test groups.

Balance of benefit and harm

No proper evaluation of patient-reported outcome measures (PROMs) was carried out in the studies.

Overall certainty of the evidence Low.

From evidence to recommendation- additional considerations

Acceptability

None of the included studies provides evidence of superior patients` acceptance of laser application as compared to mechanical instrumentation with curettes. There is no evidence so far for clinicians` acceptability.

Feasibility

Implementation of therapy may be negatively influenced by the lack of retrievability and/or shape of the prosthetic suprastructure.

Ethical Considerations

No data are available to address ethical considerations.

Economic Considerations

Cost-effectiveness has not been evaluated in these studies.

Legal Considerations

So far, the manufacturer`s indications are respected, there are no legal considerations.

R7.6 | What is the efficacy of submarginal instrumentation with air-polishing in the nonsurgical step of peri-implantitis treatment?

PICOS question addressed by a SR

R7.6 – Evidence-based recommendation

We suggest not to use air polishing for non-surgical submarginal peri-implant

instrumentation.

Supporting literature [27, 130]

Quality of evidence Very low - 2 RCTs (n=64 patients, n=75 implants) with a minimum

follow-up of 6 months, with low risk of bias

Grade of recommendation Grade $B - \downarrow$

Strength of consensus: Consensus (13.7% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

We suggest not to use air polishing for non-surgical submarginal peri-implant instrumentation.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (13.3% abstentions due to potential Col)

Background

Intervention

To overcome challenges with conventional submarginal instrumentation alternative approaches have been assessed. Among them, air-polishing systems have been tested both as monotherapy and as adjuncts to conventional submarginal instrumentation.

Available evidence

Number and design of included studies

Two RCTs (n_{patients}=64, n_{implants}=75) assessed the submarginal peri-implant instrumentation with air-polishing (128, 129). One used air-polishing as monotherapy (128), while the other combined ultrasonics and air-polishing (129).

Risk of bias

Both studies had low risk of bias.

Effect sizes and their clinical relevance

Due to the heterogeneity of the treatment protocols, no meta-analysis was carried out. Both studies on air-abrasive decontamination showed PD and BOP reductions but no statistically significant differences. Inter-group differences for BOP were observed with air-polishing as monotherapy [128].

Consistency

Not feasible to be assessed.

Balance of benefit and harm

One study reported higher levels of pain values during treatment and after one week for the glycine powder group as compared to mechanical instrumentation with ultrasonics [129]. Cases of subcutaneous emphysema have been reported after the use of air-polishing devices [131-133]. Among members of the expert panel, three groups had experienced such adverse events.

Overall certainty of the evidence Very low.

From evidence to recommendation- additional considerations

Acceptability

Patient perception and acceptance were assessed in one study, showing no statistically significant differences [129].

Feasibility

Implementation of therapy may be negatively influenced by the lack of retrievability and/or shape of the prosthetic suprastructure. Sometimes sub-marginal delivery may not be possible due to the size of the nozzle.

Ethical Considerations

Consider that the additional clinical benefit, if present, is small; that there is a potential risk of harm (subcutaneous emphysema); and that no clear benefit in terms of patient acceptability has been demonstrated.

Economic Considerations

Cost-effectiveness has not been evaluated in these studies.

Legal Considerations

So far, the manufacturer 's indications are respected, there are no legal considerations.

7.3 | Non-surgical submarginal instrumentation - chemical approaches for cleaning/decontamination

The systematic review by de Waal and co-workers evaluated chemical approaches for implant cleaning/decontamination, aiming to answer the following PICOS question: "in adult patients with peri-implantitis (P), what is the efficacy of sub-marginal instrumentation combined with chemical surface decontamination (I) in comparison with sub-marginal instrumentation with or without placebo (C), in terms of changes in probing depths (PD) and/or bleeding on probing (BOP) (O), as reported in randomized clinical trials (RCT), nonrandomized controlled clinical trials (CCT) or prospective cohort studies, with a minimum of 6-month follow-up (S)?".

Three RCTs were identified: two with low risk of bias and one with some concerns. Two RCTs assessed the benefits of antimicrobial photodynamic therapy (aPDT) as an adjunct to submarginal instrumentation, using either toluidine blue [134] or methylene blue [135] as photosensitizers. One RCT assessed the efficacy of a desiccant material consisting of a gel of concentrated aqueous mixture of hydroxybenzenesulphonic and hydroxymethoxybenzene acids and sulphuric acid [129].

R7.7 | What is the efficacy of adjunctive antimicrobial photodynamic therapy in the nonsurgical step of peri-implantitis treatment?

PICOS question addressed by a SR

R7.7 – Evidence-based recommendation

We suggest not to use antimicrobial photodynamic therapy, adjunctively to submarginal instrumentation or as monotherapy, in non-surgical peri-implantitis therapy.

Supporting literature [28]

Quality of evidence Very low – for adjunctive use, two 6-months RCTs, one with some

concerns and one with low risk of bias; as monotherapy, no studies were considered.

Grade of recommendation Grade $B - \psi$

Strength of consensus: Unanimous consensus (1.9% of the group abstained due to

potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

We suggest not to use antimicrobial photodynamic therapy, adjunctively to submarginal instrumentation or as monotherapy, in non-surgical peri-implantitis therapy.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (2.5% abstentions due to potential Col)

Background

Intervention

Antimicrobial photodynamic therapy (aPDT) involves the local application of light and a photosensitizing compound. Photosensitizers are generally applied sub-marginally (in the periimplant pocket). Photons with specific energy (wavelength) interact with the specific photosensitizer and release electrons that catalyse an oxidative reaction which has an antibacterial effect. The rationale for application of this method in the control of peri-implantitis is based on its potential antibacterial effect on the microbial biofilm associated with the implant [136].

Available evidence

Number and design of included studies

Two RCTs assessing antimicrobial photodynamic therapy as adjunct to sub-marginal instrumentation, using either toluidine blue (66/66 patients) [134 or methylene blue (25/26 patients and 30/33 implants) [135], with appropriate wavelengths for the photosensitizers (635 nm for toluidine blue, 670 nm for methylene blue). As expected, no studies were found assessing aPDT as monotherapy, since aPDT can not remove biofilm.

Risk of bias

One study was considered at low risk of bias, the other had some concerns in terms of bias.

Effect sizes and their clinical relevance

Although both studies reported some favourable results in terms of PD reduction for aPDT as adjunct to submarginal instrumentation, over submarginal instrumentation alone, results were inconsistent and/or showed no differences for other outcome variables (BOP, MBL and/or CAL). No meta-analysis could be performed due to the limited number of studies identified and their heterogeneity.

Consistency

Substantial heterogeneity was observed in study design, interventions (laser type, photosensitizer, pre-treatment), populations studied, and reported results of the studies.

Balance of benefit and harm

No adverse effects were reported.

Overall certainty of the evidence

Due to the heterogeneity in study design, interventions, populations studied and reported outcomes, the certainty of evidence is very low.

From evidence to recommendation- additional considerations

Acceptability

There is insufficient data to support or refute the use of aPDT as adjunct to submarginal instrumentation in the non-surgical treatment of peri-implantitis.

Feasibility

The adjunctive use of aPDT following submarginal instrumentation is not clinically demanding or time consuming but requires the availability of a laser.

Ethical Considerations

There is no evidence for ethical considerations. The studied photosensitizers are generally considered as safe.

Economic Considerations

The additional cost associated with aPTD may not be justified.

Legal Considerations

There are no obvious legal considerations.

R7.8 | What is the efficacy of an adjunctive antiseptic desiccant solution in the nonsurgical step of peri-implantitis treatment?

PICOS question addressed by a SR

R7.8 – Evidence-based recommendation

We suggest not to use a desiccant antiseptic gel, adjunctively to submarginal instrumentation or as monotherapy, in non-surgical peri-implantitis therapy.

Supporting literature [28]

Quality of evidence Very low – one RCT with 6 months follow-up, with low risk of bias, on

adjunctive use. No studies as monotherapy were considered.

Grade of recommendation Grade B – \downarrow

Strength of consensus: Unanimous consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

We suggest not to use a desiccant antiseptic gel, adjunctively to submarginal instrumentation or as monotherapy, in non-surgical peri-implantitis therapy.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (4.7% abstentions due to potential Col)

Background

Intervention

In some studies, patients diagnosed with chronic periodontitis were treated with a desiccant material, consisting of a gel or liquid of concentrated aqueous mixture of hydroxybenzenesulphonic and hydroxymethoxybenzene acids, together with sulphuric acid. Results were promising regarding improvements in clinical parameters, microbiological variables, and inflammatory mediators when compared to subgingival instrumentation alone [137, 138]. The same principles were used for its application as an adjunct to submarginal instrumentation in the treatment of peri-implantitis.

Available evidence

Number and design of included studies

One factorial design RCT with two control and two test groups (16/16 patients and 16/16 implants) assessed the adjunctive desiccant antiseptic gel and the method of sub-marginal instrumentation [129]. No studies were found testing efficacy as monotherapy.

Risk of bias

The study was considered at low risk of bias.

Effect sizes and their clinical relevance

PD and CAL reduction were greater in patients treated with the desiccant material, regardless of the submarginal instrumentation method (ultrasonic scaler alone or combined with glycine powder air-polishing). The magnitude of the additional improvements in PD was 0.5 mm. There were no significant differences for any of the other outcomes reported.

Consistency

Not applicable.

Balance of benefit and harm

No adverse effects were reported. However, since the product is an acid, a negative impact on the surrounding tissues may happen (caustic effect on the soft tissues).

Overall certainty of the evidence

Due to the limited number of studies, the certainty of the evidence is very low.

From evidence to recommendation-additional considerations

Acceptability

There is insufficient data to support the use of desiccant material as an adjunct to submarginal instrumentation in the non-surgical treatment of peri-implantitis.

Feasibility

The adjunctive use of desiccant material following submarginal instrumentation is not clinically demanding or time-consuming. Currently, there is only one brand name/manufacturer for this material (HybenX[®], EPIEN Medical Inc., Saint Paul, MN, USA).

Ethical Considerations

There is no evidence for ethical considerations.

Economic Considerations

There are additional costs associated with the use of the desiccant material (e.g. in Germany the cost are approximately 100 euro for two syringes of 1 mL each).

Legal Considerations

The product has been approved as Class I medical device in the European Union and Canada. The implications of the use in other geographical locations or the use for indications besides the ones approved are unclear.

7.4 | Non-surgical submarginal instrumentation – adjunctive therapies

The systematic review by Liñares and co-workers [33] explored the added value of adjunctive therapies by answering the following PICOS question: "in patients diagnosed with periimplantitis (population), which is the efficacy of patient-performed or administered adjunctive measures to non-surgical therapy (intervention) as compared to no adjunct (comparison), in terms of probing depth and/or bleeding on probing reductions (primary outcomes), reported in RCTs or CCTs with at least 6 months of follow-up (study design)?".

Initially, eight studies were identified, but for the consensus development, five RCTs were finally considered: two on local antimicrobials, two on systemic antimicrobials and one on probiotics. Two studies presented some concerns and three studies a low risk of bias. The other studies were excluded due to different reasons: non-sustained release for local antimicrobials; inadequate control group (treated with aPDT) and inclusion criteria (abscess) for systemic antimicrobials; antimicrobials; and antibiotic intake in test and control groups, when assessing probiotics.

R7.9 | Do adjunctive locally administered antimicrobials improve the clinical outcome of subgingival instrumentation?

R7.9 – Evidence-based recommendation

We suggest not to use locally administered antimicrobials, adjunctively to submarginal instrumentation or as monotherapy, in non-surgical peri-implantitis therapy.

Supporting literature [33, 113, 139)]

Quality of evidence Low – 2 RCTs for chlorhexidine "chips" with low risk of bias and 2 RCTs for minocycline microspheres

Grade of recommendation Grade B – \downarrow

Strength of consensus: Consensus (1.9% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

We suggest not to use locally administered antimicrobials, adjunctively to submarginal instrumentation or as monotherapy, in non-surgical peri-implantitis therapy.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (4.7% abstentions due to potential Col)

Background

Intervention

Locally delivered antimicrobials may be used as an adjunct to subgingival instrumentation in patients with periodontitis, particularly in non-responding and recurrent sites [140]. The same principle may apply for non-surgical therapy of peri-implantitis.

Available evidence

Number and design of included studies

Two placebo-controlled RCTs with 6-month follow-up assessed the adjunctive effect of locally applied chlorhexidine "chips" to the non-surgical submarginal instrumentation [141, 142]. These studies used an intense regime with multiple, repeated applications during the observation period. In addition, although they were not included in the systematic review, two RCTs evaluating locally applied minocycline microspheres were considered in the discussions [33, 113, 139].

Risk of bias

Two RCTs with low risk of bias.

Effect sizes and their clinical relevance

Results of two studies evaluating multiple applications of a biodegradable matrix containing chlorhexidine were pooled for meta-analyses, showing a statistically significant improved PD

reduction (WMD=0.2 mm; 95% CI [0.0; 0.5]; p=0.031; I²=0.0%; p=0.570). No or very limited information was available for BOP or disease resolution.

Consistency

Not feasible due to the limited information available.

Balance of benefit and harm

No increase in adverse effects were observed. PROMs were not reported. Harm versus benefit considerations on the use of locally delivered antibiotics need to be considered.

Overall certainty of the evidence .

Low.

From evidence to recommendation- additional considerations

Acceptability

No specific information is available; however, local antimicrobials are normally easy to use by the practitioners. Conversely, some patients/clinicians may not be willing to use antimicrobial products.

Feasibility

Some of the evaluated products may not be commercially available in some countries. For chlorhexidine "chips", only one brand/manufacturer is available (PerioChip®, Dexcel Pharma, Or Akiva, Israel). For minocycline microspheres, the brand tested in the considered studies was Arestin® (OraPharma, Bridgewater, NJ, USA).

Ethical Considerations No applicable.

Economic Considerations

Economic costs and cost-effectiveness should be considered before their use. Economic cost may be relatively high (for chlorhexidine "chips", one chip may cost around 30 euro, while for

minocycline microspheres, one cartridge costs around 100 \$, especially if multiple applications are needed). Some additional information is presented in Section 1.

Legal Considerations

Some of the evaluated products have not been registered for use in some countries, and/or may not have been approved for this specific indication.

R7.10 | Do adjunctive systemically administered antibiotics improve the clinical outcomes of non-surgical treatment?

PICOS question addressed by a SR

R7.10 – Expert consensus-based recommendation

Due to concerns about patients' health and the impact of systemic antibiotic use on public health, its routine use as an adjunct to non-surgical treatment in patients with peri-

implantitis is **not recommended**.

Supporting literature [33]

Quality of evidence Low – two RCTs, one with some concerns, and another with low risk of bias.

Grade of recommendation Grade A – $\psi\psi$

Strength of consensus - Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adapted.

Due to concerns about patients' health and the impact of systemic antibiotic use on public health, **we recommend not to use** systemically administered antibiotics during non-surgical treatment of peri-implantitis in a non-specialist setting.

Their routine use as an adjunct to non-surgical treatment in patients with peri-implantitis in specialist setting is **not recommended**.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

The adjunctive use of systemic antimicrobials has been extensively evaluated in the treatment of periodontitis [143]. The same principles may apply for its adjunctive use in the *non-surgical step* of the treatment of peri-implantitis.

The expert group evaluated, first, the adjunctive benefit of systemic antibiotics to submarginal instrumentation alone. The effect was both statistically significant and clinically relevant. In the included studies [144, 145] the effect tended to be more pronounced at cases with initially deeper lesions and to improve over time up to one year. At least in one study [144], the benefit included improvements in marginal bone levels. The size of the benefit may allow achievement of the stipulated treatment endpoints in a significant number of cases and hence avoid surgical intervention. The clinical recommendation that antibiotics cannot be recommended as a routine is, therefore, based on the general principles of antibiotic stewardship and the public health objective of limiting unnecessary use of antibiotics in dentistry. Rationale for limitation is twofold: the public health considerations related to spread of antibiotic resistance and the potential individual harms related to dysbiosis of the individual patient microbiome. The panel felt that clinicians should avoid use of systemic antibiotics for the management of periimplantitis and limit it to cases at the end of the severity spectrum (e.g. deep pockets \geq 7 mm, extensive suppuration) and/or with multiple and/or strategically affected implants that could respond well and be retained over time (the suggested protocol in these cases would be metronidazole 500 mg/8 hours/7 days). However, the use of systemic antimicrobials should be avoided in palliative care of lost implants.

Available evidence

Two studies were included in the systematic review [33], both showing statistically significant benefits in PD reduction at 6 months and up to 12 months after the prescription of systemic antimicrobials. These results were more pronounced when the deepest site of each implant was considered for the analysis. A significant effect for the use of systemic antimicrobials in radiographic bone gain (≈1.2 mm) was observed on rough-surface implants [144]. However, no changes in marginal bone levels were reported on machined implants [144, 145].

153

In both studies, PD reductions improved from 3 to 12 months suggesting that, if at the reevaluation (6-12 weeks) the recommended endpoints are not achieved at implant level (i.e. residual PD ≤5 mm with no BOP at more than one site point and no suppuration), but a clear improvement in PD reduction is detected, it may be adequate to wait longer before a decision to perform additional treatment is made.

Number and design of included studies

RCTs (n=2) with a double-blind, placebo controlled, parallel design with follow-up up to 12 months [144, 145]. One evaluated amoxicillin plus metronizadole (n=40 patients/40 implants) [144, 145] and the other, metronizadole alone (n=32 patients/62 implants) [144, 145].

Risk of bias

Risk of bias was low for one study, while the other study presented some concerns.

Effect sizes and their clinical relevance

Systemic antimicrobials showed a greater PD reduction when compared to mechanical debridement alone at 6 months and up to the 12 months follow-up (≈1.5 mm). These results were more pronounced when the deepest site of each implant was considered for the analysis.

Consistency

Not applicable.

Balance of benefit and harm

One study assessed the potential side effects of systemic antibiotics, with 6 subjects (38%) in the test group (systemic metronidazole) and 5 (31%) in the control group (placebo) reporting either gastrointestinal disorders, headaches, or metallic taste, without significant differences among groups. Global concerns regarding the overuse of antibiotics and the development of antibiotic resistance must be considered. Benefit versus harm analysis includes considerations on the overall use of antibiotics for the individual patient and public health. Systemic antibiotic regimens have shown long-lasting impacts on the faecal microbiome, including an increase in genes associated with antimicrobial resistance.

Overall certainty of the evidence

Limited evidence is available.

From evidence to recommendation- additional considerations

Acceptability

Due to concerns for patient's health and the impact of systemic antibiotic use on public health, its routine use as an adjunct to submarginal peri-implant instrumentation in patients with peri-implantitis is not recommended.

Feasibility

Adjunct systemic antimicrobials to non-surgical peri-implant therapy are a feasible procedure since these antimicrobials may be prescribed in most countries. Moreover, the procedure does not demand high clinical skills.

Ethical Considerations

Important concerns are related to patient's health and the impact of systemic antibiotic use to public health.

Economic Considerations

Although economic considerations have not been analysed in the included studies, some indications can be given. The cost of systemic antimicrobials is low, particularly in comparison to other potential adjuncts (e.g. local antimicrobials or probiotics). Although there is not enough evidence to provide any strong recommendation, the prescription of systemic antimicrobials in specific cases may reduce the need for additional treatment, including surgical procedures, reducing added costs and morbidity.

Legal Considerations

There are no specific legal considerations.

R7.11 | What is the efficacy of adjunctive probiotics in the non-surgical step of periimplantitis treatment?

PICOS question addressed by a SR R7.11 – Evidence-based recommendation We suggest not to use probiotics as an adjunct to submarginal instrumentation, in non-surgical peri-implantitis therapy. Supporting literature [33] Quality of evidence Very low - one RCT with some concerns in risk of bias Grade of recommendation Grade B - 4 Strength of consensus: Strong consensus (0% of the group abstained due to potential Col) **BSP** Implementation This evidence-based recommendation is adopted. We suggest not to use probiotics as an adjunct to submarginal instrumentation, in non-surgical peri-implantitis therapy. Updated Evidence: No new applicable evidence was identified Strength of Consensus: Unanimous consensus (2.2% abstentions due to potential Col)

Background

Intervention

Probiotics are defined as "live microorganisms which, when administered in adequate amounts, confer a health benefit on the host" [146]. Probiotics have been proposed to modulate oral microbiota and host immune response [147, 148]. While it has been suggested that probiotics may not be used as an adjunct to subgingival instrumentation in the treatment of stages I-III periodontitis [2]), regarding peri-implantitis, available studies reveal contradictory results.

Available evidence

Number and design of included studies

One placebo-controlled RCT assessed the adjunctive effect of probiotics to non-surgical submarginal instrumentation [149], with a preparation containing *Lactobacillus reuteri*, to be applied both locally and systemically.

Risk of bias Some concerns.

Effect sizes and their clinical relevance No adjunctive effect of the use of probiotics was observed on PD or BOP.

Consistency

Not applicable.

Balance of benefit and harm

No proper evaluation of patient-reported outcome measures (PROMs) was carried out, although the extrapolation from the periodontal field suggests that this formulation is safe, and patients do not frequently report adverse effects.

Overall certainty of the evidence

Very low.

From evidence to recommendation- additional considerations

Acceptability

No specific information is available. However, probiotics are normally easy to use by the practitioners. Conversely, some patients/clinicians may not be willing to use these products.

Feasibility

Adjunctive probiotics to non-surgical peri-implant therapy are a feasible approach since these products can be prescribed in many countries. Moreover, the procedure does not demand high clinical skills.

Ethical Considerations

Not applicable.

Economic Considerations

There is an additional cost associated with the use of probiotics that is borne by the patient.

Legal Considerations

There are no specific legal considerations.

8 | RECOMMENDATIONS FOR THE SURGICAL MANAGEMENT OF PERI-IMPLANTITIS

8.1 | Introduction - general recommendations in the surgical *step* of peri-implantitis treatment

The purpose of a surgical approach in the management of peri-implantitis is to provide access to the implant to facilitate surface decontamination. The goal is to achieve the resolution of the inflammatory lesion. Target sites for surgical treatment are those presenting with persisting signs of pathology after non-surgical therapy, i.e., deep pockets together with BOP/SOP.

A *standard surgical procedure* includes, in addition to flap elevation and removal of inflamed tissue, cleaning/ decontamination of the implant surface using e.g., small pieces of gauze soaked in saline and removal of mineralized deposits with curettes.

Additional procedures in the surgical treatment of peri-implantitis may include: (i) the management of peri-implant osseous defects using reconstructive approaches, (ii) additional methods for implant surface decontamination and (iii) the adjunctive use of local/systemic antibiotics.

R8.1 | What is the importance of adequate self-performed oral hygiene in the context of surgical treatment of peri-implantitis?

Additional question addressed by the WG

R8.1 – Expert consensus-based recommendation

We recommend not to perform surgical treatment of peri-implantitis in patients not

achieving and maintaining adequate levels of self-performed oral hygiene.

Supporting literature Expert opinion

Quality of evidence Not applicable

Grade of recommendation Grade $A - \downarrow \downarrow$

Strength of consensus Strong consensus (0% of the group abstained due to potential

Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

We recommend not to perform surgical treatment of peri-implantitis in patients not achieving and maintaining adequate levels of self-performed oral hygiene.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Studies have shown the detrimental effects of surgical treatment of periodontitis in patients with insufficient levels of self-performed oral hygiene [25]. Since bacterial biofilms are considered the primary etiological factor for both periodontitis and peri-implantitis, the importance of adequate self-performed levels of oral hygiene needs to be emphasized also in the context of surgical treatment of peri-implantitis. Similar to the periodontal scenario, studies on surgical treatment of peri-implantitis have also indicated unfavourable outcomes in patients not achieving and maintaining adequate levels of self-performed oral hygiene [150, 151].

R8.2 | What is the level of professional expertise required for surgical treatment of periimplantitis?

Additional question addressed by the WG

R8.2 – Expert consensus-based recommendation

We recommend that dental teams offering implant therapy also possess the professional expertise to manage peri-implantitis. Since surgical treatment of peri-implantitis is complex, we **recommend** that it is provided by dentists with specific training or by specialists.

Supporting literature Expert opinion

Quality of evidence Not applicable

Grade of recommendation Grade A – ↑↑

Strength of consensus Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

We recommend that dental teams offering implant therapy also possess the professional expertise to manage peri-implantitis. Since surgical treatment of peri-implantitis is complex, **we recommend** that it is provided by dentists with specific training or by specialists.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Recognition of peri-implantitis as a disease entity is relatively recent and the armamentarium of surgical approaches is constantly evolving. The dental team must be continuously updated on the most effective treatment modalities. Treatment of peri-implantitis lies within the scope of the speciality of periodontology.

R8.3 | What are the end points of successful surgical therapy of peri-implantitis?

Additional question addressed by the WG

R8.3 – Expert consensus-based recommendations

- 1. We recommend that, at implant level, clinicians use \leq 1 point of BOP, absence of SOP, PD \leq 5 mm and absence of progressive bone loss compared to pre-treatment bone levels to verify disease resolution.
- 2. **We recommend** that clinical parameters be recorded 6 months post-treatment and that radiographs be obtained at 12 months.
- 3. **We suggest** that complication-free survival of the implant and implant-supported prosthesis and patient satisfaction (e.g. aesthetic appreciation) be included in the long-term evaluation of treatment outcomes.

Supporting literature Expert consensus

Quality of evidence Not applicable

Grade of recommendation Grade A – $\uparrow\uparrow$ (1,2); Grade B – \uparrow (3)

Strength of consensus

(1) Strong consensus (0% of the group abstained due to potential Col)

(2) Strong consensus (0% of the group abstained due to potential Col)

(3) Strong consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

- We recommend that, at implant level, clinicians use ≤ 1 point of BOP, absence of SOP, PD ≤5 mm and absence of progressive bone loss compared to pre-treatment bone levels to verify disease resolution.
- 2. **We recommend** that clinical parameters be recorded 6 months post-treatment and that radiographs be obtained at 12 months.
- 3. **We suggest** that complication-free survival of the implant and implant-supported prosthesis and patient satisfaction (e.g. aesthetic appreciation) be included in the long-term evaluation of treatment outcomes.

BSP implementation additional text:

Regarding the thresholds defined above, the BSP implementation refers to literature in section 4.3.3 defining cases of peri-implant disease [5, 6].

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Studies [152-154] demonstrate that progression of peri-implantitis occurs in the presence of clinical signs of inflammation, and is manifested through reduction of peri-implant bone levels. In contrast, shallow peri-implant probing depths and absence of BOP/SOP have been associated with stable peri-implant support in longitudinal studies.

R8.4 | What considerations should be made about the implant-supported prosthesis when performing surgical treatment of peri-implantitis?

Additional question addressed by the WG

R8.4 – Expert consensus-based recommendations

- 1. **We recommend** that implant-supported prostheses that do not allow access for self-performed oral hygiene be adjusted prior to surgical therapy of peri-implantitis.
- 2. **We suggest** that implant-supported prostheses be removed, if feasible, in conjunction with surgical treatment of peri-implantitis to facilitate access and peri-implant tissue healing.

Supporting literature: Not applicable

Quality of evidence Not applicable

Grade of recommendation Grade A – $\uparrow \uparrow$ (1); Grade B – \uparrow (2)

Strength of consensus

(1) Unanimous consensus (0% of the group abstained due to potential Col)

(2) Consensus (0% of the group abstained due to potential Col)

BSP Implementation

This expert consensus-based recommendation is adopted.

1. **We recommend** that implant-supported prostheses that do not allow access for self-performed oral hygiene be adjusted prior to surgical therapy of peri-implantitis.

2. **We suggest** that implant-supported prostheses be removed, if feasible, in conjunction with surgical treatment of peri-implantitis to facilitate access and peri-implant tissue healing.

BSP implementation additional literature citation

Supporting literature refer to R6.7 [111, 112]

BSP implementation additional text:

Consent should be taken from patient for the potential risk of damage to prosthesis during modification and or removal (refer to R6.7).

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Adequate levels of self-performed oral hygiene are a prerequisite for successful outcomes of surgical treatment for peri-implantitis. Studies have shown that inadequate access for oral hygiene around implants is associated with higher risk for peri-implantitis [87, 155] therefore, adjustment of the implant-supported prosthesis with the aim to facilitate access for oral hygiene is an important measure prior to surgical treatment of peri-implantitis.

8.2 | Indications of the surgical treatment of peri-implantitis and efficacy of access/resective approaches

R8.5 | When is surgical treatment of peri-implantitis indicated?

PICOS question addressed by a SR

R8.5 – Evidence-based recommendation

In peri-implantitis patients in whom endpoints of non-surgical therapy (PD \leq 5 mm & \leq 1

point of BOP) have not been achieved, we recommend performing surgical therapy.

Supporting literature [30, 32]

Quality of evidence Moderate

Grade of recommendation Grade A – ↑↑

Strength of consensus Consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adapted

In peri-implantitis patients in whom endpoints of non-surgical therapy (PD \leq 5 mm & \leq 1

point of BOP) have not been achieved, we suggest performing surgical therapy

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Consensus (0% abstentions due to potential Col)

Background

Intervention

Surgical therapy of peri-implantitis may consist of different approaches, including simple access flap, pocket elimination or reconstructive procedures. All modalities incorporate flap elevation, removal of inflamed tissues and implant surface debridement/decontamination.

Available evidence

Number and design of included studies

Data from 13 prospectively collected studies (649 patients) with a follow-up ranging from 1 to 5 years, addressed access flap and resective surgery. Seven RCTs assessed the efficacy of reconstructive surgery (194 patients) compared to access flap surgery. The respective datasets were evaluated in two systematic reviews [30, 32]. All studies reported on reduction of PD and BOP. Clinically relevant end points (e.g., PD <6 mm), PROMs, health-economic parameters and adverse events were not consistently reported.

Risk of bias

The 13 studies on access flap and resective surgery were generally found to be at low RoB, while multiple studies evaluating reconstructive measures were judged to show high RoB.

Effect sizes and their clinical relevance

For access flap and resective surgery, the estimated reduction of PD was 2.2 mm (95% CI [1.8; 2.7]). Reconstructive surgery resulted in similar PD reduction (additional effect relative to access flap alone: -0.39 (95% CI [-1.16; 0.24]) at 12 months. For access flap and resective surgery, reduction of standardized mean %BOP was estimated at 27.0 (95% CI [19.8; 34.2]) and an overall bone gain of 0.2 mm (95% CI [0.0; 0.5]) was noted. Reconstructive surgery resulted in an additional bone gain of 0.75 mm (95% CI [-1.39; -0.11]) over access flap alone at 12 months (confidence interval is presented with negative values, since in the original analyses positive values indicated more gain for access flap and negative for reconstructive procedures). Over 5-year observation periods, disease recurrence/progression was observed at 32% to 44% of treated implants. Corresponding implant loss was low in the short term but after 5 years ranged from 14% to 21%.

Consistency

Results were consistent across studies for changes of PD and MBL. Reduction of BOP was heterogenous across studies. Data were generated in various clinical settings, including university centres and private clinics.

165

Balance of benefit and harm

In general, considerable improvements in clinical and radiographic parameters were noted. However, disease recurrence and implant loss were not uncommon events after 5 years. Data on PROMs and adverse events were rarely reported.

Overall certainty of the evidence

The certainty of evidence is graded as moderate based on the lack of direct comparisons between surgical and non-surgical therapy of peri-implantitis.

From evidence to recommendation- additional considerations

Acceptability

PROMs were rarely reported. Limited data suggest a high degree of patient satisfaction at 1 year following surgical therapy. Adverse events reported were mostly related to the use of systemic antibiotics.

Feasibility

Related procedures are clinically demanding.

Ethical Considerations

Some decontamination procedures and grafting materials evaluated in the studies included have not been tested for safety.

Economic Considerations

Health-economic parameters were not evaluated in the identified studies. In general, surgical therapy of peri-implantitis is a costly procedure. Some decontamination procedures and grafting materials may generate additional costs in the absence of documented benefit.

Legal Considerations

Some decontamination procedures and grafting materials evaluated in the studies included have not been tested for safety and are considered off-label.

R8.6 | What is the efficacy of surgical treatment of peri-implantitis using access flap or resective procedures (resection of hard / soft peri-implant tissues aiming at reducing or eliminating pockets)?

PICOS question addressed by a SR

R8.6 – Evidence-based recommendation

In peri-implantitis patients in whom endpoints of non-surgical therapy (PD \leq 5 mm & \leq 1 point of BOP) have not been achieved, **we recommend** performing access flap or resective surgery as both modalities are effective.

Supporting literature [30, 32]

Quality of evidence Moderate

Grade of recommendation Grade A – $\uparrow\uparrow$

Strength of consensus Consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

In peri-implantitis patients in whom endpoints of non-surgical therapy (PD \leq 5 mm & \leq 1 point of BOP) have not been achieved, **we recommend** performing access flap or resective surgery as both modalities are effective.

BSP implementation additional background text:

Where a decision of surgery has been made

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

Surgical therapy of peri-implantitis includes flap elevation, removal of inflamed tissues and implant surface debridement/decontamination. In access flap procedures, soft tissue flaps are

simply repositioned, while resective approaches aim at apically displacing flaps through soft tissue and/or hard tissue recontouring.

Available evidence

Number and design of included studies

Thirteen studies (n=649 patients), with a follow up range from 1 to 5 years (only two studies with a 5-year follow-up), were included [30, 32]. One study was an RCT comparing surgical therapy to non-surgical intervention. All datasets were prospective and a total of ten originated from control arms within RCTs, while the remaining two were case series. All studies reported on reduction of PD and BOP. Clinically relevant end points (e.g. PD <6 mm), PROMs, health-economic parameters and adverse events were not consistently reported.

Risk of bias

The 13 studies were generally found to be at low RoB. In the two evaluations covering longer follow-ups [\geq 5 years; [154, 156], loss to follow-up exceeded 20% and the overall rating was downgraded to "fair".

Effect sizes and their clinical relevance

Based on 18 studies (n=661 implants), the estimated reduction of PD was 2.2 mm (95% CI [1.8; 2.7]). Based on 8 studies (n=477), reduction of standardized mean BoP% was estimated at 27.0 (95% CI [19.8; 34.2]). Based on 12 studies (n=637), a standardized mean bone gain of 0.2 mm (95% CI [0.0; 0.5]) was estimated. Over 5-year observation periods, disease recurrence/progression was observed at 32% to 44% of treated implants. Corresponding implant loss was low in the short term but after 5 years ranged from 14% to 21%.

Consistency

Results were consistent across studies in regard to changes of PD and MBL. Reduction of BOP was heterogenous across studies. Data were generated in various clinical settings, including university centres and private clinics.

168

Balance of benefit and harm

In general, considerable improvements in clinical and radiographic parameters were noted. However, disease recurrence and implant loss were not uncommon events after 5 years. Data on PROMs (two studies) and adverse events (three studies) were rarely reported.

Overall certainty of the evidence

The certainty of evidence is graded as moderate based on the lack of direct comparisons between surgical and non-surgical therapy of peri-implantitis.

From evidence to recommendation- additional considerations

Acceptability

PROMs were reported in two studies, only. Limited data suggest a high degree of patient satisfaction at 1 year following surgical therapy. Adverse events reported in three studies were mostly related to the use of systemic antibiotics.

Feasibility

Related procedures are clinically demanding.

Ethical Considerations

Some decontamination procedures evaluated in the studies included have not been tested for safety.

Economic Considerations

Health-economic parameters were not evaluated in the identified studies. In general, surgical therapy of peri-implantitis is a costly procedure. Some decontamination procedures may generate additional costs in the absence of documented benefit.

Legal Considerations

Some decontamination procedures evaluated in the studies included have not been tested for safety and are considered off-label.

8.3 | Management of peri-implant osseous defects using reconstructive approaches

R8.7 | Do reconstructive procedures used in the management of osseous defects (e.g., bone substitute materials) as part of surgical treatment of peri-implantitis result in superior outcomes when compared to access flap alone?

PICOS question addressed by a SR

R8.7 – Evidence-based recommendation

In the surgical management of osseous defects in peri-implantitis patients, access flap with or without reconstructive procedures **may be considered**; no evidence demonstrating

superiority of any specific surgical technique was identified.

Supporting literature [30]

Quality of evidence Low

Grade of recommendation Grade $O \leftrightarrow$ (need for further research)

Strength of consensus Consensus (19.0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

In the surgical management of osseous defects in peri-implantitis patients, access flap with or without reconstructive procedures **may be considered**; no evidence demonstrating superiority of any specific surgical technique was identified.

BSP implementation of additional new evidence :

Since the publication of the treatment guideline, one new RCT was published [157], which confirmed similar clinical improvements in 40 peri-implantitis patients undergoing either reconstructive surgery (DBBM and collagen membrane) or access flap surgery (mean PPD reduction in the deepest site 3.7±1.9 mm vs. 4.2±1.8 mm).

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (5.4% abstentions due to potential Col)

Background

Intervention

Reconstructive procedures aim to regenerate the bony defect, achieve re-osseointegration, and limit peri-implant soft-tissue recession [158]. Reconstructive therapy of peri-implant bone defects includes the use of bone grafts, bone replacement grafts, barrier membranes, bioactive agents (growth factors, autologous platelet concentrates and amelogenin), or combinations thereof.

Available evidence

Number and design of included studies

Seven RCTs assessed the efficacy of reconstructive surgery (total of 200 implants in 194 patients) compared to access flap surgery (total of 188 implants in 184 patients) [30]. Different types of reconstructive surgeries were documented, including the use of titanium granules, amelogenin, deproteinized bovine bone mineral (DBBM or DBBM graft with 10% collagen) alone or combined with a native bilayer collagen membrane, or a beta-tricalcium phosphate graft formulated with prolonged release of local doxycycline.

Risk of bias

Based on RoB2, there was concern for four studies in one domain (predominantly due to bias in measurement of the outcome), while three studies were considered at high risk of bias, mainly due to the combination of missing outcomes and bias in selection of the reported results.

Effect sizes and their clinical relevance

Meta-analysis (4 studies; 262 patients and 272 implants) showed an estimated mean difference in PD changes between access flap surgery and reconstructive surgery of -0.39 (95% CI [-1.16; 0.24]; p=0.325, l²=66.4%) at 12 months. No evidence of small-study effects was detected. Amongst the five studies that reported on BOP changes at 12 months, one study showed a statistically significant improvement for reconstructive therapy as compared with access flap surgery. No differences were indicated in relation to the change in SOP. At 12 months, implant survival was similar between the two treatment procedures, ranging from 85.7% to 100% for access flap and from 95% to 100% for reconstructive therapy. Meta-

analysis for changes in radiographic mean bone levels (4 studies; 262 patients and 272 implants) showed a statistically significant benefit of reconstructive compared to access flap surgery of -0.75 mm (95% CI [-1.39; -0.11]; p=0.022; l²=83.4%). The confidence interval is presented with negative values, since in the original analyses positive values indicated more gain for access flap and negative for reconstructive procedures. Irrespective of the surgical approach and biomaterial employed, resolution of peri-implantitis is unpredictable and a significant difference between the two treatment approaches was not consistently shown.

Consistency

Overall, inconsistency in the direction of effect was noticed for the included studies, as only one showed a significant improvement in PD change and one in BOP change, when reconstructive procedures were employed.

Balance of benefit and harm

A similar number of adverse events and complications was associated with reconstructive and access flap surgeries. In the long-term a number of implants is expected to develop disease recurrence, which may require additional surgical procedures or could lead to implant loss.

Overall certainty of the evidence

The certainty of evidence is low based on the quality of the studies (RoB) and inconsistency of outcomes.

From evidence to recommendation- additional considerations

Acceptability

Only two studies considered PROMs, with no significant differences in terms of pain scores, number of tablets taken and satisfaction.

Feasibility

Related procedures are clinically demanding.

Ethical Considerations

Some decontamination procedures applied in the studies have not been tested for safety.

Economic Considerations

No study addressed health economic outcomes on this topic (Donos et al., 2023). Reconstructive surgery represents an additional financial burden for the patient, which should be discussed with the patient.

Legal Considerations

Not applicable.

R8.8 | What are the specific prerequisites (e.g. dimensions of intra-bony defects) for a reconstructive approach?

PICOS question addressed by a SR

R8.8 – Evidence-based recommendations

We suggest that reconstructive procedures preferably be applied at intra-osseous

defects with a depth of \geq 3 mm.

Supporting literature [30]

Quality of evidence Low

Grade of recommendation Grade B – \uparrow

Strength of consensus Consensus (13.3% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

We suggest that reconstructive procedures preferably be applied at intra-osseous

defects with a depth of ≥ 3 mm.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (4.8% abstentions due to potential Col)

Background

Intervention

See previous section.

Available evidence

Number and design of included studies

None of the identified studies in the systematic review was designed to investigate the site prerequisites for a reconstructive [30]. Nevertheless, the 4 RCTs of the network meta-analysis included \geq 3 mm, angular peri-implant bone defects, which showed significant improvements in clinical and radiographic parameters from baseline to 12 months post-reconstructive therapy. Deeper defects are more likely to result in radiographic defect fill and 3- and 4-wall defects result in higher reduction in PD and BOP.

Risk of bias

Based on RoB2, the risk of bias varied from low to high in the relevant studies.

Effect sizes and their clinical relevance

Not applicable.

Consistency

Despite the three identified studies showed consistency on the impact of defect morphology on the treatment outcome, none of these studies was designed to answer this question.

Balance of benefit and harm Not applicable.

Overall certainty of the evidence Low.

From evidence to recommendation- additional considerations Acceptability Not applicable.

Feasibility

Not applicable.

Ethical Considerations

Not applicable.

Economic Considerations

Not applicable.

Legal Considerations

Not applicable.

R8.9 | What are the preferred materials to be used in reconstructive procedures?

PICOS question addressed by a SR

R8.9 – Evidence-based recommendation

Bone grafts with or without barrier membranes **may be considered** in reconstructive procedures.

Supporting literature [30]

Quality of evidence Low

Grade of recommendation Grade O \leftrightarrow

Strength of consensus Consensus (19.0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

Bone grafts with or without barrier membranes **may be considered** in reconstructive procedures.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (11.4% abstentions due to potential Col)

Background

Intervention

A variety of bone substitutes, barriers and bioactive agents have been proposed for reconstructive procedures.

Available evidence

Number and design of included studies

Five RCTs and six prospective case series assessed the efficacy of reconstructive periimplantitis therapy [30].

Risk of bias

Based on RoB2, two out of the five included RCTs were at high risk of bias, some concerns were raised for two studies and one was at low risk of bias. Based on ROBINSI, one CCT was at serious risk of bias, three prospective cohort studies were considered at serious risk of bias and two prospective cohort studies were at critical risk of bias [30].

Effect sizes and their clinical relevance

Regardless of the biomaterials applied, reconstructive therapy led to a mean PD reduction ranging from 2.0 to 4.5 mm and to a mean reduction in BOP ranging from 44.8% to 86% at 12 months post therapy. Studies reporting on SOP, showed a significant reduction at 12 months) and 5 years post-surgery. Based on one study (45 patients and 75 implants), included in the network meta-analysis (4 studies; 160 patients and 190 implants) [30], an improved PD reduction was shown when a xenogenic rather than an autologous graft was applied in combination with a collagen membrane. Implant survival at 12 months ranged from 92% to 100%, but when considering composite outcomes for peri-implantitis resolution the range reported by the included studies was considerably wider (0% to 91% at 12 months).

Consistency

All reconstructive procedures improved clinical and radiographic outcomes as compared to baseline regardless of the biomaterials employed.

Balance of benefit and harm

None of the different reconstructive approaches was associated with early side effects or adverse events beyond what would be expected for this type of surgical procedure. Notably, the use of a combination of membrane and bone graft was associated with an increased risk for flap dehiscence in two studies.

Overall certainty of the evidence Low.

From evidence to recommendation- additional considerations

Acceptability

Based on one study, the use of a graft alone was associated with significantly less pain at 2 weeks as compared to the combined use of a graft and collagen membrane.

Feasibility

Not applicable.

Ethical Considerations Not applicable.

Economic Considerations

No study addressed health economic outcomes on this topic. However, it should be noted that reconstructive surgery represents an additional financial burden for the patient.

Legal Considerations Not applicable.

R8.10 | What is the preferable mode of healing (submerged versus transmucosal) to be used in reconstructive procedures?

Additional question addressed by the WG

R8.10 – Expert statement

We do not know whether a submerged or transmucosal healing protocol would influence

the outcomes of reconstructive procedures.

Supporting literature [30]

Quality of evidence: Quality of evidence not applicable; no studies identified

Grade of recommendation Grade O \leftrightarrow

Strength of consensus Strong consensus (1.9% of the group abstained due to potential

Col)

BSP Implementation

This expert statement is adopted.

We do not know whether a submerged or transmucosal healing protocol would influence

the outcomes of reconstructive procedures.

BSP implementation additional background text:

During healing a high level of infection control should be maintained.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

In reconstructive procedures, submerged and transmucosal healing have been documented.

Available evidence

Number and design of included studies

No focused question in the current systematic review [30] was formulated to address this topic. Nevertheless, none of the included studies compared submerged to unsubmerged healing protocol.

Risk of bias

Not applicable.

Effect sizes and their clinical relevance Not applicable.

Consistency

Not applicable.

Balance of benefit and harm

The main advantage of submerged healing would be to achieve primary wound closure and to promote an aseptic healing environment, which are crucial factors for stabilizing the blood clot, improving graft stability, and maximizing the regenerative potential of the intrabony compartment. On the other hand, unsubmerged healing eliminates the need for prosthesis removal, reduces treatment time, costs and possibly the overall complexity of treatment.

Overall certainty of the evidence

Very low.

From evidence to recommendation- additional considerations

Acceptability

It should be noted that a submerged healing protocol may result in the need of temporary tooth replacement.

Feasibility Not applicable.

Ethical Considerations Not applicable.

Economic Considerations

No study addressed health economic outcomes on this topic. It should be noted that unsubmerged healing eliminates the need of prosthesis removal, thus reducing treatment time and possibly costs.

Legal Considerations Not applicable.

8.4 | Additional methods for implant surface decontamination

R8.11 | Do photo-/mechanical and physical implant surface decontamination procedures improve outcomes of surgical treatment?

PICOS question addressed by a SR

R8.11 – Evidence-based recommendations (1, 2) and statement (3)

1. **We suggest not** to use air-polishing or Er:YAG laser for implant surface decontamination during surgical treatment of peri-implantitis.

2. Titanium brushes **may be considered** as an alternative/adjunct to standard decontamination.

3. There is insufficient evidence to make any recommendation regarding the use of implantoplasty.

Supporting literature [34]

Quality of evidence Low

Grade of recommendation Grade $B - \psi$ (1); Grade $O - \leftrightarrow$ (2); Statement (3)

Strength of consensus

- (1) Consensus (7.8% of the group abstained due to potential Col)
- (2) Consensus (0% of the group abstained due to potential Col)

(3) Consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendations (1&2) and statement (3) is adopted.

1. **We suggest not to** use air-polishing or Er:YAG laser for implant surface decontamination during surgical treatment of peri-implantitis.

2. Titanium brushes **may be considered** as an alternative/adjunct to standard decontamination.

3. There is insufficient evidence to make any recommendation regarding the use of implantoplasty.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (8.3% abstentions due to potential Col)

Background

Intervention

As substantial evidence supports the bacterial aetiology of peri-implantitis, removal of the biofilm from contaminated implant surfaces is a crucial treatment step in obtaining disease resolution [5, 6, 159].

Available evidence

Number and design of included studies

In total, five RCTs (4 two-armed and 1 three-armed; 183 patients/242 implants) with follow-up periods ranging from 6 to 24 months were included [34]. One RCT investigated the alternative use of air polishing with erythritol powder in conjunction with non-reconstructive surgical periimplantitis therapy compared to standard instrumentation [160]; another RCT, with 3-arms, assessed the efficacy of titanium brushes (test 1) and air polishing with glycine powder (test 2) as alternative decontamination measures for implant surface decontamination compared to standard instrumentation [161]; two RCTs investigated the efficacy of Er:YAG laser compared to either standard instrumentation [162] or debridement with piezoelectric scaler and stainless-steal

scaler [163] during reconstructive therapy and as an adjunct to implantoplasty; and one RCT evaluated the added value of a titanium brush, on top of ultrasonic decontamination and hydrogen peroxide in regenerative surgery [164].

Risk of bias

Based on ROB 2, two RCTs were judged to have an overall low risk of bias, two RCTs had an overall high risk of bias, and one RCT had an unclear risk of bias.

Effect sizes and their clinical relevance

Based on two RCTs with 6- to 12-month follow-ups, the adjunctive/alternative use of an air abrasive device with glycine or erythritol powders did not result in improved BOP reductions compared to the control during surgical therapy of peri-implantitis [160, 161]. One RCT indicated a significantly higher PD reduction following the alternative use of air polishing with glycine powder and titanium brushes compared to the standard decontamination [161]. Based on one RCT, after 6 months, alternative use of titanium brush resulted in significantly higher BOP reduction compared to either air polishing or the standard instrumentation (i.e., curettes to remove hard deposits plus gauze soaked in saline/saline irrigation) [161].

During reconstructive therapy, a titanium brush resulted in significantly greater reduction of the deepest PD values compared to the control group (i.e., mechanical and chemical implant surface decontamination) [164]. An Er:YAG laser resulted in significantly higher PD reductions after 6-months in one RCT, but was not associated with improved BOP reductions over respective control measures (i.e., implantoplasty and standard instrumentation or debridement with piezoelectric scaler and stainless-steal scaler) as shown in two RCTs [162, 163].

Consistency

Two RCTs reported on no benefit of air polishing either with erythritol or glycine powder on the reduction of BOP values [160, 161]. A beneficial effect of the use of a titanium brush was reported in two RCTs in terms of BOP [161] and PD reductions [164]. Two RCTs consistently reported on no benefits of Er:YAG laser on changing BOP values after 6- and 12-months in conjunction with reconstructive therapy and as an adjunct to implantoplasty [162, 163].

Inconsistencies were found between the studies with respect to the PD changes following Er:YAG laser application. In fact, significantly higher PD reduction following the use of Er:YAG laser was reported after 6-months in one RCT [163], whereas after 24-months another RCT indicated no benefits of ER:YAG laser in reducing PD values [162].

Balance of benefit and harm

Harms have not been explicitly reported and evaluated in two RCTs. A slight pigmentation of peri-implant soft-tissues was observed in one out of 30 patients treated with implantoplasty. One RCT reported on adverse events observed in one out of 16 patients associated with persistence of suppuration and swelling following air polishing. Another RCT reported on membrane exposure during the healing, following reconstructive therapy of peri-implantitis, however, without providing the number of implants/patients experiencing this complication.

Overall certainty of the evidence

The evidence was graded as low due to a low number of studies with a considerable heterogeneity.

From evidence to recommendation- additional considerations

Acceptability

None of the studies investigated PROMs.

Feasibility

Certain decontamination protocols may be considered as technically demanding.

Ethical Considerations

Certain decontamination protocols have not been tested for safety.

Economic Considerations

Economic aspects could not be assessed due to the lack of reporting.

Legal Considerations Not applicable.

R8.12 | Do chemical implant surface decontamination procedures improve outcomes of surgical treatment?

PICOS question addressed by a SR

R8.12 – Evidence-based recommendation

We suggest not to use chlorhexidine or photodynamic therapy for implant surface

decontamination during surgical therapy of peri-implantitis.

Supporting literature [38]

Quality of evidence Very low

Grade of recommendation Grade B – \downarrow

Strength of consensus Consensus (1.7% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

We suggest not to use chlorhexidine or photodynamic therapy for implant surface decontamination during surgical therapy of peri-implantitis.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (11.4% abstentions due to potential Col)

Background

Intervention

Adjunctive antimicrobial chemical therapy is an approach used to improve the standard implant surface decontamination methods. Chlorhexidine (CHX) has antiseptic properties that kills bacteria. Photodynamic therapy (PDT) functions by light activation of a photosensitizing dye to generate reactive oxygen species that destroy those bacteria.

Available evidence

Number and design of included studies

Evidence was available for PDT from 2 RCTs (n=43) and for CHX from 2 RCTS (n=130) (Wilensky et al., 2023). Both with a follow-up of ≥ 6 and up to 12 months. Only RCTs reporting mean PD changes and BOP changes were included.

Risk of bias

For PDT the risk of bias was low to unclear, and for CHX it was unclear to high risk of bias.

Effect sizes and their clinical relevance

No benefits were observed with the adjunctive application of CHX; no improvement was observed for PDT in terms of PD reduction, and only minor reductions in BOP (mean difference - MD=7.4).

Consistency

For PDT heterogeneity was low, and for CHX it was medium to high.

Balance of benefit and harm

One study did not report on adverse events, while three studies reported no to minor adverse effects. One study reported gastrointestinal problems in five patients that were taking systemic antibiotics. One study reported no adverse effects, and another study reported two patients with one complication.

Overall certainty of the evidence

The GRADE analysis showed a very low certainty of evidence for both adjunctive treatments in all the tested parameters.

From evidence to recommendation- additional considerations

Acceptability

None of the studies reported patient-reported outcomes and there is no evidence supporting one approach over the other, including the standard therapy.

Feasibility

While CHX solution is affordable and easily available, PDT results in additional costs without any documented clinical benefit.

Ethical Considerations

The lack of efficacy together with possible side effects, such as hypersensitivity, suggest that these treatments are not justified.

Economic considerations

The additional costs associated with adjunctive PDT therapy are not justified.

Legal Considerations

PDT is an off-label use during surgery, with no clear benefits.

8.4 | Adjunctive use of local/systemic antimicrobials

R8.13 | Do adjunctive systemically administered antibiotics improve clinical outcomes of surgical treatment of peri-implantitis?

PICOS question addressed by a SR

R8.13 – Evidence-based recommendation

Due to concerns about patient health and the impact of systemic antibiotic use on public

health and inconsistent evidence, its use as adjunct to surgical therapy of peri-implantitis is

not recommended.

Supporting literature [36]

Quality of evidence Low

Grade of recommendation Grade A – $\downarrow \downarrow$

Strength of consensus (0% of the group abstained due to potential Col)

BSP Implementation

This evidence-based recommendation is adopted.

Due to concerns about patient health and the impact of systemic antibiotic use on public health and inconsistent evidence, its use as adjunct to surgical therapy of peri-implantitis is **not recommended**.

BSP implementation additional text:

This recommendation does not necessarily apply to complex procedures, for example significant augmentations.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Strong consensus (0% abstentions due to potential Col)

Background

Intervention

Tissue destruction at peri-implantitis sites is more pronounced than periodontitis around teeth due to anatomical differences, larger size of the inflammatory lesion, and extent of the lesion to the bone crest. Therefore, clinicians are tempted to use systemic antibiotics in addition to the surgical treatment of peri-implantitis.

Available evidence

Number and design of included studies

Two RCTs including 49 patients (25 test, 24 control) and 39 patients (20 test, 19 control) and followed for one year showed inconsistent results in terms of PD, BOP, bone level changes: one assessed the systemic application of amoxicillin, 750 mg, twice per day for 10 days, and starting 3 days prior to surgery [165]; the other evaluated the systemic application of azithromycin, 500 mg at the day of surgery, and 250 mg, once per day, during 4 additional days [166].

Risk of bias

Some concerns [165] and high risk [166], as evaluated with RoB 2.

Effect sizes and their clinical relevance

Disease resolution (based on <5 mm PDs, no BOP and no additional bone loss >5 mm) was consistent between studies and favoured systemic antibiotics: 56% test vs. 29.2% control [165]; 46.7% test vs. 25% control group [166]. Two implant losses occurred in the control group of the first study [165].

Consistency See previous section.

Balance of benefit and harm

The potential benefit of the use of systemic antibiotics needs to be balanced with the overall risks, which include adverse events (e.g. allergic reactions) and antibiotic resistance.

Overall certainty of the evidence Low.

From evidence to recommendation-additional considerations

Acceptability

Due to concerns about patients' health and the impact of systemic antibiotic use on public health and inconsistent evidence, its use as adjunct to surgical therapy of peri-implantitis is not recommended.

Feasibility

Not applicable.

Ethical Considerations

Harms related to the intake of systemic antibiotics must be balanced with potential benefits.

Economic Considerations Not applicable. Legal Considerations

Not applicable.

R8.14 | Do adjunctive locally administered antibiotics improve clinical outcomes of surgical treatment of peri-implantitis?

PICOS question addressed by a SR

R8.14 – Evidence-based statement

There is insufficient evidence to make any recommendation on the use of local antibiotics

as adjuncts in the surgical treatment of peri-implantitis.

Supporting literature [36, 38]

Quality of evidence Very low

Grade of recommendation Statement

Strength of consensus Unanimous consensus (2.1% of the group abstained due to potential Col)

BSP Implementation

This evidence-based statement is adopted.

There is insufficient evidence to make any recommendation on the use of local antibiotics as adjuncts in the surgical treatment of peri-implantitis.

Updated Evidence: No new applicable evidence was identified

Strength of Consensus: Unanimous consensus (0% abstentions due to potential Col)

Background

Intervention

Tissue destruction at peri-implantitis sites is more pronounced than periodontitis around teeth due to anatomical differences, larger size of the inflammatory lesion, and extent of the lesion to the bone crest. Therefore, clinicians are tempted to use antibiotics in addition to the surgical treatment of peri-implantitis.

Available evidence

Number and design of included studies

Two RCTs were identified: one assessing local minocycline application at the time of surgery in 50 patients (25 test, 25 control), and repeated at 1, 3 and 6 months, with all patients also receiving systemic amoxicillin thrice per day, 500 mg, for 3 days [167] and another evaluating local doxycycline application in 27 patients (14 test, 13 control), formulated in a bone graft, at the time of surgery [168].

Risk of bias

High risk of bias for both RCTs.

Effect sizes and their clinical relevance Not applicable.

Consistency

Not applicable.

Balance of benefit and harm

The potential benefit of the use of local antibiotics needs to be balanced with the overall risks, which include adverse events (e.g. allergic reactions) and antibiotic resistance.

Overall certainty of the evidence Very low.

From evidence to recommendation- additional considerations

Acceptability

Not applicable.

Feasibility

Related products may not be available in all European countries.

Ethical Considerations

Harms related to the intake of local antibiotics must be balanced with potential benefits.

Economic Considerations

Additional costs related to the medical product must be considered.

Legal Considerations Not applicable.

Journal Prevention

9. Information about the source document, The EFP S3 Level Clinical Practice Guideline

Authors of the Prevention and Treatment of Peri-implant Diseases – The EFP S3 Level Clinical Practice Guideline

David Herrera¹, Tord Berglundh², Frank Schwarz³, Iain Chapple^{4,5}, Søren Jepsen⁶, Anton Sculean⁷, Moritz Kebschull^{4,5,8}, Panos N. Papapanou⁸, Maurizio S. Tonetti⁹, Mariano Sanz¹, * On behalf of the EFP workshop participants and methodological consultant (listed below).

iv. Author's institutional affiliations

1. ETEP (Etiology and Therapy of Periodontal and Peri-implant Diseases) Research Group, University Complutense of Madrid, Spain.

2. Department of Periodontology, Institute of Odontology, The Sahlgrenska Academy, University of Gothenburg, Gothenburg, Sweden.

3. Department of Oral Surgery and Implantology, Johann Wolfgang Goethe-University, Carolinum, Frankfurt, Germany.

4. Periodontal Research Group, Institute of Clinical Sciences, College of Medical & Dental Sciences, The University of Birmingham, Birmingham, United Kingdom.

5. Birmingham Community Healthcare NHS Foundation Trust, Birmingham, United Kingdom.

6. Department of Periodontology, Operative and Preventive Dentistry. University Hospital Bonn, Bonn, Germany.

7. Department of Periodontology, School of Dental Medicine, University of Bern, Bern, Switzerland.

8. Division of Periodontics, Section of Oral, Diagnostic and Rehabilitation Sciences, College of Dental Medicine, Columbia University, New York, NY, USA.

8. Shanghai PerioImplant Innovation Center, Department of Oral and Maxillo-facial Implantology, Shanghai Key Laboratory of Stomatology, National Clinical Research Centre for Oral Diseases, Shanghai Ninth People Hospital, Shanghai Jiao Tong University School of Medicine, Shanghai, China.

* Workshop Participants, in alphabetic order

Mario Aimetti, Juan Blanco, Nagihan Bostanci, Philippe Bouchard, Darko Božić, Nurcan Buduneli, Elena Calciolari, María Clotilde Carra, Raluca Cosgarea, Jan Cosyn, Bettina Dannewitz, Monique Danser, Beatriz de Tapia, Yvonne de Waal, Jan Derks, Henrik Dommisch, Nikos Donos, Kenneth Eaton, Peter Eickholz, Bahar Eren Kuru, Elena Figuero, Moshe Goldstein, Filippo Graziani, Jasmin Grischke, Fernando Guerra, Lisa Heitz-Mayfield, Karin Jepsen, Lise-Lotte Kirkevang, Odd Carsten Koldsland, France Lambert, Antonio Liñares, Bruno Loos, Phoebus Madianos, Paula Matesanz, Paulo Melo, Ana Molina, Virginie Monnet Corti, Eduardo Monter, Frauke Müller, Luigi Nibali, Andrés Pascual, Ioannis Polyzois, Marc Quirynen, Ausra Ramanauskaite , Gitana Rederiene, Stefan Renvert, Mario Roccuzzo, Philipp Sahrmann, Giovanni Salvi, Nerea Sánchez, Ignacio Sanz-Sánchez, Lior Shapira, Andreas Stavropoulos, Meike Stiesch, Ieva Tamošiūnaitė, Wim Teughels, Cristiano Tomasi, Leonardo Trombelli, Spyros Vassilopoulos, Anders Verket, Nicola West, Asaf Wilensky.

Methodological Consultant

Workshop Organization European Federation of Periodontology

Scientific societies involved in the guideline development process European Dental Hygienists Federation European Society for Endodontology

Other organisations involved in the guideline development process Council of European Dentists European Dental Students' Association Platform for Better Oral Health in Europe

v. Acknowledgments

The authors express their gratitude to all reviewers involved in the preparation of the systematic reviews and sincerely thank those organizations that participated in the guideline development

process: Council of European Dentists, European Dental Hygienists Federation, European Dental Students' Association, European Society for Endodontology, Platform for Better Oral Health in Europe.

Conflict of Interest Statement

Individual potential conflict of interest forms were completed by all participants and are available on file at the European Federation of Periodontology and extracted in the Supporting Information, available online [CPGperiimplant-Supporting Information_Potential conflict of interest (20230114)]. Potential conflicts of interest, in the previous 36 months, reported by the chairs of the workshop (in alphabetic order) are listed here:

Tord Berglundh (Chair) reports - Grants or contracts from any entity: Dentsply Implants IH, Osteology Foundation (University institution grants). Consulting fees: Dentsply Implants IH (Personal). Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Dentsply Implants IH, Spanish Society of Periodontology (SEPA), French Society of Periodontology (Lecture, personal). Support for attending meetings and/or travel: Dentsply Implants IH, Spanish Society of Periodontology (SEPA), French Society of Periodontology, European Federation of Periodontology (EFP) (Support for travel when participating as an invited speaker). Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: National Board of Health and Welfare, Sweden (National guidelines in Dentistry) (University institution grant); Swedish Quality Registry for Caries and Periodontal Disease (Personal fee).

Iain Chapple (Chair) reports - Grants or contracts from any entity: Grant from GSK (PhD studentship & other grant); Two grants from DEBRA (Epidermolysis Bullosa Charity), one grant from Unilever (Research Grants); NIHR Biomedical Research Centre Grant (Research infrastructure grant). Royalties or licenses: Quintessence (Book Royalties). Consulting fees: J&J, GSK, Unilever, Philips (Consultancy Fees). Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: J&J, GSK, Unilever, Philips (Sponsored Lecture Fees); Lectures to Swiss, German, Italian, Portuguese, Turkish, Polish, Danish, Austrian, British societies, IAP Colombia (Expenses paid for meetings). Support for attending meetings and/or travel: P&G, J&J (Support to attend international conference).

Patents planned, issued or pending: 65 patents (Patents on saliva diagnostics). Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: European Workshop Committee of the European Federation of Periodontology – co-chair; Invited member of EFP Executive Board (Unpaid); Chair of FDI working group on Oral & Systemic Health (Unpaid); Scientific Advisor to the British Society of Periodontology Executive Board (Unpaid). Other financial or non-financial interests: non-financial (my wife runs Oral Health Innovations that has the license for PreViser and DEPPA risk assessment software in the UK).

David Herrera (Chair) reports - Grants or contracts from any entity: Dentaid, Kulzer, Lacer, ZIZ Dental, Isdin, Affinity (Research contract via university); University of Bristol, University of Pisa (Research contract via university). Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Oral-B, Straumann, Klockner, Dexcel, Dentaid, Colgate, Lacer Spanish Society of Periodontology (SEPA), European Federation of Periodontology (EFP), Saudi Society of Periodontology (SSP), Ukrainian Association of Periodontology (UAPerio), Iberopanamerican Federation of Periodontology (FIPP), Dansk Parodontologisk Selskab (DPS) (Personal honoraria for lectures); Dentaid (Manuscript writing). Support for attending meetings and/or travel: Oral-B, Straumann, Klockner, Dexcel, Dentaid, Colgate (Support for attending meetings and/or travel when participating as invited speaker). Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: Patron of the Fundación SEPA de Periodoncia e Implantes Dentales (Unpaid); Chair of the European Workshop Committee of the European Federation of Periodontology (Unpaid). Receipt of equipment, materials, drugs, medical writing, gifts or other services: Dentaid, Lacer, ZIZ Dental, Isdin, Affinity (Receipt of equipment, materials, drugs, in association with research contracts).

Sören Jepsen (Chair) reports - Grants or contracts from any entity: Osteology Foundation (Research Contract with University); Innovationsfond, GBA, German Ministry of Health (Research Contract with University); GlaxoSmithKline Consumer Health (Research Contact with University). Consulting fees: KZV Nordrhein (Consultancy Fees). Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Geistlich Pharma (Personal honoraria for lectures). Leadership or fiduciary role in other board, society,

committee or advocacy group, paid or unpaid: German Society of Dentistry and Oral Medicine (DGZMK), Advisory Council on continuing dental education.

Moritz Kebschull (Chair) reports - Grants or contracts from any entity: National Institutes for Health Research, Genolytic, Bredent, Unilever (To University of Birmingham). Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Camlog, Curaden, Dexcel, Geistlich, Hu-Friedy, NSK, Procter&Gamble, Resolve Biosciences, Unilever (For lecturing, to me); Several universities and public bodies, learned societies (For lecturing, to me). Patents planned, issued or pending: With commercial partner Genolytic (filed) (Perio diagnostics). Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: EFP, BSP, DG PARO (Unpaid). Receipt of equipment, materials, drugs, medical writing, gifts or other services: Geistlich, Hu-Friedy, NSK, Straumann (Support for courses / material and instruments/devices).

Panos Papapanou (Chair) reports - Grants or contracts from any entity: National Institutes of Health/ National Institute of Aging (NIH/NIA R1 AG7615; 9/3/221-8/31/226) (A Longitudinal Study of Periodontal Infections and Alzheimer's Disease: The WHICAP Ancillary Study of Oral Health); Colgate-Palmolive, Inc., Piscataway, NJ, USA (Cerebrospinal fluid microbiome, oral health and cognitive aging). Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: New York Society of Periodontics (Lecture honorarium); Straumann (Lecture honorarium). Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: Editor in Chief, Journal of Clinical Periodontology (EiC honorarium from Wiley and EFP).

Mariano Sanz (Chair) reports - Grants or contracts from any entity: Straumann, Nobelbiocare, Sweden & Martina, Dentsply Implants, TiCare Implants, Klockner Implants, DENTAID, Sunstar, Geistlich Pharma (Research contract via university); Osteology Foundation, Oral Reconstruction Foundation, ITI Foundation (Member). Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Straumann, Nobelbiocare, Sweden & Martina, Dentsply Implants, TiCare Implants, Klockner Implants, Dentaid, Sunstar, Geistlich Pharma, Osteology Foundation, ITI, ORF (Personal honoraria for lectures); Dentaid (Personal honoraria for manuscript writing). Support for attending

meetings and/or travel: Straumann, Nobelbiocare, Sweden & Martina, Dentsply Implants, TiCare Implants, Klockner Implants, Dentaid, Sunstar, Geistlich Pharma, Osteology Foundation, ITI, ORF (Support for attending meetings and/or travel when participating as invited speaker). Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: President ORF, Fellow ITI, Honorary Member OF (Unpaid).

Frank Schwarz (Chair) reports - Grants or contracts from any entity: Osteology Foundation, ITI, Oral Reconstruction Foundation (Contract via University); Camlog, Straumann, Geistlich, Dentsply (Contract via University); BMBF and Dentsply (Contract via University -Grant to investigate surgical peri-implantitis therapy by combination of a plasma and a water jet). Consulting fees: Henry Schein (Personal Honoraria). Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Osteology Foundation, ITI, Oral Reconstruction Foundation (Personal lecture fees); Camlog, Straumann, Geistlich (Personal lecture fees). Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: Osteology Foundation (Annual compensation fee); EAO (unpaid). Receipt of equipment, materials, drugs, medical writing, gifts or other services: Straumann, Camlog, Geistlich, Dentsply (Receipt of equipment, materials in association with research contracts via university).

Anton Sculean (Chair) reports - Grants or contracts from any entity: ITI, Straumann, Geistlich, GC, Regedent AG, Research Grant by the University of Bristol and GlaxoSmithKline, Oral Reconstruction Foundation, Association for Dental Infection Control (Research contract via university. Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Osteology Foundation, Geistlich, Straumann, Regedent AG, Camlog, Oral Reconstruction Foundation, EMS, German Society of Periodontology (DGParo), Danish Dental Association (Honoraria for lectures). Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: Osteology Foundation (Honorarium for Board Meeting); Swiss Dental Society (SSO) (Bernese Section) (Honorarium for Board Meeting).

Maurizio Tonetti (Chair) reports - Grants or contracts from any entity: Straumann AG, Geistlich AG, Sunstar SA (Grant). Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Straumann AG (Sunstar SA); Nobel Biocare

(Hu-friedy); Geistlich AG. Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: European Research Group on Periodontology, Executive Director (Unpaid). Receipt of equipment, materials, drugs, medical writing, gifts or other services: Straumann AG; Geistlich AG.

Corresponding author

David Herrera

ETEP (Etiology and Therapy of Periodontal and Peri-implant Diseases) Research Group Faculty of Odontology, University Complutense of Madrid Plaza Ramón y Cajal s/n (Ciudad Universitaria) 28040 Madrid, Spain davidher@ucm.es +34 91 3941907

ORCID

Mariano Sanz https://orcid.org/0000-0002-6293-5755 David Herrera https://orcid.org/0000-0002-5554-2777 Moritz Kebschull https://orcid.org/0000-0003-1863-0679 Iain Chapple https://orcid.org/0000-0003-2697-7082 Søren Jepsen https://orcid.org/0000-0002-4160-5837 Tord Beglundh https://orcid.org/0000-0001-5864-6398 Anton Sculean https://orcid.org/0000-0003-2836-5477 Maurizio S. Tonetti https://orcid.org/0000-0002-2743-0137

Abstract

Background: The recently published Clinical Practice Guidelines (CPGs) for the treatment of periodontitis in stages I-IV provided evidence-based recommendations for the treatment of periodontitis patients, defined according to the 2018 classification. Peri-implant diseases were also re-defined in the 2018 classification, and it is well-establish that both peri-implant

mucositis and peri-implantitis are highly prevalent and, in addition, peri-implantitis is a challenging to manage condition, with important consequences in terms of morbidity.

Aim: To develop an S3 Level CPG for the management of peri-implant diseases, focusing on the implementation of interdisciplinary prevention and treatment approaches required to prevent peri-implant disease development or recurrence and to treat/rehabilitate patients with dental implants following peri-implant disease development.

Material and Methods: This S3 Level CPG was developed by the European Federation of Periodontology (EFP), following methodological guidance from the Association of Scientific Medical Societies in Germany and the Grading of Recommendations Assessment, Development and Evaluation (GRADE) process. A rigorous and transparent process included synthesis of relevant research in 13 specifically commissioned systematic reviews, evaluation of the quality and strength of evidence, the formulation of specific recommendations and a structured consensus process with leading experts and a broad base of stakeholders.

Results: The S3 Level CPG for the management of peri-implant diseases culminated in recommendations for different interventions, to be implemented before, during and after implant placement/loading. Prevention of peri-implant diseases should start when dental implants are planned, surgically placed and prosthetically loaded. Once the implants are loaded and in function, a supportive peri-implant care program should be organised, including periodical assessment of peri-implant tissue health. If peri-implant mucositis or peri-implantitis are detected, recommendations for their management are provided.

Conclusion: The present S3 Level CPG informs clinical practice, health systems, policymakers and, indirectly, the public on the available and most effective modalities maintain healthy periimplant tissues, and to manage peri-implant diseases, according to the available evidence at the time of publication.

Key words: peri-implant diseases, peri-implant mucositis, peri-implantitis, clinical guideline, dental implant

Conflicts of Interest Declarations

Supplied as a separate document for the purposes of blind review see Supplementary material.

CRediT authorship contribution statement

All authors contributed to development of the guideline. All authors contributed to the development of the manuscript, approved the final version for submission and agreed to be accountable for the work.

Declaration of Competing Interest and Funding for the BSP adolopment of the S3 guideline

Individual potential conflict of interest forms were completed by all participants and are available on file at the British Society of Periodontology and Implant Dentistry. In addition, potential conflict of interest information of the Chairs of the workshop is listed here (in alphabetical order):

Professor lain Chapple (Chair) reports - Grants or contracts from any entity:

Grant from GSK - PhD studentship & other grant, Grants from DEBRA (Epidermolysis Bullosa Charity) - Two Research Grants, NIHR Biomedical Research Centre Grant- Research infrastructure grant, Grant from Unilever - Research Grant. Royalties or licenses: Quintessence - Book Royalties. Consulting fees: J&J, GSK, Unilever & Philips. Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: P&G, J&J, Unilever & Philips - Sponsored Lecture Fees. Lectures to Swiss, German, Italian, Portuguese, Turkish, Polish, Danish, Austrian, Australian/New Zealand, Egyptian and British societies, IAP Colombia - Expenses paid for meetings. Support for attending meetings and/or travel: P&G and J&J -Support to attend international conferences. Patents planned, issued or pending: 65 patents - Patents on saliva diagnostics. Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: European Workshop Committee of the European Federation of Periodontology – co-chair (Unpaid). Chair of FDI working group on Oral & Systemic Health (Unpaid), Invited member of EFP Executive Board (Unpaid), Scientific Advisor to the British Society of Periodontology Executive Board (Unpaid) Secretary to EFP Workshop Committee (Unpaid).

Professor Shauna Culshaw (Chair) reports - Grants or contracts from any entity: Wellcome Trust - Research grant, Haleon - Research grant, BBSRC/Haleon - Research grant, Dentsply Sirona - Research grant, NHS Research Scotland - Research grant, Public Health Scotland -Research grant, NIHR - Research grant. Consulting fees: Haleon - Advisory Board. Support for

attending meetings and/or travel: Haleon, Dentsply Sirona. Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: SDCEP Revision of Guidelines for Periodontal Treatment in Primary Care, British Society of Periodontology Patient Forum. Receipt of equipment, materials, drugs, medical writing, gifts or other services: Dentsply Sirona - Materials to support research projects.

Professor Nikos Donos (Chair) reports – Grants or contracts from any entity: Research grants from: Geistlich, Straumann, GSK, ITI, Osteology Foundation, NIH, MRC, Barts Charity & St Barts Medical trust. Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Lecture fees from – Geistlich, ITI foundation &

Osteology Foundation . Participation on a Data Safety Monitoring Board or Advisory Board: SEED - Previously advisory board. Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: Osteology Foundation board - Board member Lead Science committee, ADI - Academic Rep. Stock or stock options: SEED - Stock options. Receipt of equipment, materials, drugs, medical writing, gifts or other services: BTI – equipment, EMS – equipment, Geistlich – biomaterials, Straumann - Biomaterials, Implants, Hu-friedy - Surgical instrument. Other financial or non-financial interests: Private practice limited to periodontics and implant dentistry.

Professor Moritz Kebsch ull (Guidelines Lead) reports – Grants or contracts from any entity: National Institute for Health and Care Research, UK; Unilever, UK; Genolytic, Germany; Bredent, Germany, EMS, Switzerland. Payment or honoraria for lectures, presentations, speakers' bureaus, manuscript writing or educational events: Camlog, Curaden, CP GABA, Dexcel, EMS, Geistlich, Hu-Friedy, NSK, Procter&Gamble. Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid – President-Elect, European Federation of Periodontology. Receipt of equipment, materials, drugs, medical writing, gifts or other services: EMS, Geistlich, Hu-Friedy, NSK, Straumann (all equipment for teaching and research). Patents planned, issued or pending: One patent family with Genolytic on microbial diagnosis. Other financial or non-financial interests: Unpaid director in my wife's dental company, KVW Periodontal Consulting Ltd.

Professor Ian Needleman (Chair) reports - Grants or contracts from any entity: GSK/Haleon Oral healthcare - Research funding to institute. Consulting fees: GSK/Haleon - Consultancy related to prevention of periodontal diseases. Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: European Federation of Periodontology - Registration, travel and accommodation for EuroPerio 10, Copenhagen 2022. European Research Group on Periodontology (ERGOPerio) -Accommodation for Implant Dentistry Core Outcome Set consensus meeting Copenhagen 2022. Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid - British Society of Periodontology Patient Forum – Lead. Stock or stock options: Vedanta Biosciences Potential institutional payments related to microbial therapies for periodontitis. Other financial or non-financial interests: Cochrane Oral Health – Editor, Delivering Better Oral Health Guidance: Public Health England - Development role related to periodontal and peri-implant health.

Professor Luigi Nibali (Chair) reports – Grants or contracts from any entity: Research grants from Unilever. Royalties or licenses: Royalties from books published by Wiley Blackwell. Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: British Society of Periodontology, Spanish Society of Periodontology, DG Paro, Finnish Dental Society, Association Dentaire Francaise, Planden, Autran Dental Academy. Support for attending meetings and/or travel: PER IADR, PanEndo, EFP. Receipt of equipment, materials, drugs, medical writing, gifts or other services: Loan units from EMS and Mectron, laboratory kits in kind from Biotechne. Other financial or non-financial interests: Director of limited company providing periodontal clinical care (private practice).

Dr. Amit Patel (Chair) reports - Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Oral B lectures. Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: President of the Association of Dental Implantology. Other financial or non-financial interests: My own company – Amit Patel Dentistry Ltd

Professor Philip Preshaw (Chair) reports - Grants or contracts from any entity: FDI World Development Fund – Research grant - Implementing a post-pandemic Health Coaching training

programme for empowering dental professionals in disease prevention Co-Investigator. (Unpaid - grant awarded to institution). NIHR HTA programme – Research Grant - The ENHANCE-D Study – ENHANCing smoking cessation interventions in Dentistry Co-Investigator. (Unpaid - grant awarded to institution). Royalties or licenses: Wiley - Book royalties (paid). Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Springer Nature - Journal editorial role (paid). Patents planned, issued or pending: Patents filed with Philips Oral Healthcare (unpaid). Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: British Society of Periodontology – President (2024) and Council member (unpaid), Journal of Clinical Periodontology - Editorial Board member (unpaid), European Federation of Periodontology Executive Committee member (unpaid). Other financial or non-financial interests: Salvivo Ltd - Director (unpaid).

Dr Jeanie Suvan (Chair) reports - Grants or contracts from any entity: Association for Dental Infection Control Research grant to institution, Engineering and Physical Science Research Council - Research grant to institution. Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events: Hu-Friedy Group - Personal lecture honorarium, Perio Campus Foundation Italy - Personal lecture honorarium, Health Education England - Personal lecture honorarium, Osteology - Personal lecture honorarium. Support for attending meetings and/or travel: Support for travel when lecturing at above. Participation on a Data Safety Monitoring Board or Advisory Board - P&G Global Dental Schools Advisory Board - Personal honorarium. Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: UCL High Risk Research Ethics Committee- No financial payment.

Professor Nicola West (Guidelines Lead) reports – Grants or contracts from any entity: Research grants from Unilever, Haleon, 3 Shape, Align. Consulting fees: Haleon, Unilever, P&G, J&J. Payment or honoraria for lectures, presentations, speakers bureaus, manuscript writing or educational events : British Society of Periodontology, Haleon, P&G. Leadership or fiduciary role in other board, society, committee or advocacy group, paid or unpaid: Secretary General of the European Federation of Periodontology, President elect of the British

Society of Periodontology and Implant Dentistry. Other financial or non-financial interests: Director of limited company - The Bristol Perio Clinic Ltd.

ORCID

 Iain Chapple https://orcid.org/0000-0002-9653-5629

 Nikos Donos https://orcid.org/0000-0002-4117-9073

 Moritz Kebschull https://orcid.org/0000-0003-1863-0679

 Ian Needleman https://orcid.org/0000-0003-4696-1651

 Luigi Nibali https://orcid.org/0000-0003-1712-5662

 Philip M Preshaw https://orcid.org/0000-0002-7153-4865

 Jeanie Suvan https://orcid.org/0000-0002-6256-6235

 Nicola West https://orcid.org/0000-0002-9127-5530

Funding of the guideline

The development of this guideline and its subsequent publication was financed entirely by internal funds of the British Society of Periodontology and Implant Dentistry.

Appendix A. Supplementary data

Supplementary material related to this article can be found, in the online version.

References

- [1] Herrera D, Berglundh T, Schwarz F, Chapple I, Jepsen S, Sculean A, et al. Prevention and treatment of peri-implant diseases—The EFP S3 level clinical practice guideline. Journal of Clinical Periodontology. 2023;50(S26):4-76.
- [2] Caton JG, Armitage G, Berglundh T, Chapple ILC, Jepsen S, Kornman KS, et al. A new classification scheme for periodontal and peri-implant diseases and conditions -Introduction and key changes from the 1999 classification. J Clin Periodontol. 2018;45 Suppl 20:S1-s8.
- [3] Schünemann HJ, Wiercioch W, Brozek J, Etxeandia-Ikobaltzeta I, Mustafa RA, Manja V, et al. GRADE Evidence to Decision (EtD) frameworks for adoption, adaptation, and de novo development of trustworthy recommendations: GRADE-ADOLOPMENT. J Clin Epidemiol. 2017;81:101-10.
- [4] Heitz-Mayfield LJA, Salvi GE. Peri-implant mucositis. J Periodontol. 2018;89 Suppl 1:S257-S66.
- [5] Berglundh T, Armitage G, Araujo MG, Avila-Ortiz G, Blanco J, Camargo PM, et al. Periimplant diseases and conditions: Consensus report of workgroup 4 of the 2017 World Workshop on the Classification of Periodontal and Peri-Implant Diseases and Conditions. J Periodontol. 2018;89 Suppl 1:S313-S8.
- [6] Schwarz F, Derks J, Monje A, Wang HL. Peri-implantitis. J Periodontol. 2018;89 Suppl 1:S267-S90.
- [7] Bermejo P, Sánchez MC, Llama-Palacios A, Figuero E, Herrera D, Sanz M. Topographic characterization of multispecies biofilms growing on dental implant surfaces: An in vitro model. Clin Oral Implants Res. 2019;30(3):229-41.
- [8] Sánchez MC, Llama-Palacios A, Fernández E, Figuero E, Marín MJ, León R, et al. An in vitro biofilm model associated to dental implants: structural and quantitative analysis of in vitro biofilm formation on different dental implant surfaces. Dent Mater. 2014;30(10):1161-71.
- [9] Sahrmann P, Gilli F, Wiedemeier DB, Attin T, Schmidlin PR, Karygianni L. The Microbiome of Peri-Implantitis: A Systematic Review and Meta-Analysis. Microorganisms. 2020;8(5).
- [10] Derks J, Tomasi C. Peri-implant health and disease. A systematic review of current epidemiology. J Clin Periodontol. 2015;42 Suppl 16:S158-71.
- [11] Lee CT, Huang YW, Zhu L, Weltman R. Prevalences of peri-implantitis and peri-implant mucositis: systematic review and meta-analysis. J Dent. 2017;62:1-12.
- [12] Rustand K, Aass AM, Sen A, Koldsland OC. Oral health-related quality of life following peri-implantitis surgery: A prospective study. Clin Oral Implants Res. 2022;33(8):858-67.
- [13] Norum T, Aass AM, Koldsland OC. Pain and morbidity after non-surgical and surgical treatment of peri-implantitis. Acta Odontol Scand. 2019;77(8):624-9.
- [14] Research GV. Dental Implants Market Size, Share & Trends Analysis Report By Implant Type (Zirconium, Titanium), By Region (North America, Europe, Asia Pacific, Latin America, MEA), And Segment Forecasts, 2024 - 2030. 2022. p. 125.
- [15] Alani A, Kelleher M, Bishop K. Peri-implantitis. Part 1: Scope of the problem. Br Dent J. 2014;217(6):281-7.
- [16] Karlsson K, Derks J, Wennström JL, Petzold M, Berglundh T. Health economic aspects of implant-supported restorative therapy. Clin Oral Implants Res. 2022;33(2):221-30.

- [17] Schwendicke F, Tu YK, Stolpe M. Preventing and Treating Peri-Implantitis: A Cost-Effectiveness Analysis. J Periodontol. 2015;86(9):1020-9.
- [18] Listl S, Frühauf N, Dannewitz B, Weis C, Tu YK, Chang HJ, et al. Cost-effectiveness of non-surgical peri-implantitis treatments. J Clin Periodontol. 2015;42(5):470-7.
- [19] Fardal Ø, Grytten J. A comparison of teeth and implants during maintenance therapy in terms of the number of disease-free years and costs -- an in vivo internal control study. J Clin Periodontol. 2013;40(6):645-51.
- [20] Ito K, Aida J, Yamamoto T, Ohtsuka R, Nakade M, Suzuki K, et al. Individual and community-level social gradients of edentulousness. BMC Oral Health. 2015;15:34.
- [21] Hämmerle CHF, Tarnow D. The etiology of hard- and soft-tissue deficiencies at dental implants: A narrative review. J Clin Periodontol. 2018;45 Suppl 20:S267-S77.
- [22] Renvert S, Persson GR, Pirih FQ, Camargo PM. Peri-implant health, peri-implant mucositis, and peri-implantitis: Case definitions and diagnostic considerations. J Clin Periodontol. 2018;45 Suppl 20:S278-S85.
- [23] Brocklehurst PR, McKenna G, Schimmel M, Kossioni A, Jerković-Ćosić K, Hayes M, et al. How do we incorporate patient views into the design of healthcare services for older people: a discussion paper. BMC Oral Health. 2018;18(1):61.
- [24] Herrera D, Sanz M, Kebschull M, Jepsen S, Sculean A, Berglundh T, et al. Treatment of stage IV periodontitis: The EFP S3 level clinical practice guideline. J Clin Periodontol. 2022;49 Suppl 24:4-71.
- [25] Sanz M, Herrera D, Kebschull M, Chapple I, Jepsen S, Beglundh T, et al. Treatment of stage I-III periodontitis-The EFP S3 level clinical practice guideline. J Clin Periodontol. 2020;47 Suppl 22(Suppl 22):4-60.
- [26] Carra MC, Blanc-Sylvestre N, Courtet A, Bouchard P. Primordial and primary prevention of peri-implant diseases: A systematic review and meta-analysis. J Clin Periodontol. 2023;50 Suppl 26:77-112.
- [27] Cosgarea R, Roccuzzo A, Jepsen K, Sculean A, Jepsen S, Salvi GE. Efficacy of mechanical/physical approaches for implant surface decontamination in non-surgical submarginal instrumentation of peri-implantitis. A systematic review. J Clin Periodontol. 2023;50 Suppl 26:188-211.
- [28] de Waal YCM, Winning L, Stavropoulos A, Polyzois I. Efficacy of chemical approaches for implant surface decontamination in conjunction with sub-marginal instrumentation, in the non-surgical treatment of peri-implantitis: A systematic review. J Clin Periodontol. 2023;50 Suppl 26:212-23.
- [29] Dommisch H, Hoedke D, Valles C, Vilarrasa J, Jepsen S, Pascual La Rocca A. Efficacy of professionally administered chemical agents as an adjunctive treatment to submarginal instrumentation during the therapy of peri-implant mucositis. J Clin Periodontol. 2023;50 Suppl 26:146-60.
- [30] Donos N, Calciolari E, Ghuman M, Baccini M, Sousa V, Nibali L. The efficacy of bone reconstructive therapies in the management of peri-implantitis. A systematic review and meta-analysis. J Clin Periodontol. 2023;50 Suppl 26:285-316.
- [31] Gennai S, Bollain J, Ambrosio N, Marruganti C, Graziani F, Figuero E. Efficacy of adjunctive measures in peri-implant mucositis. A systematic review and metaanalysis. J Clin Periodontol. 2023;50 Suppl 26:161-87.
- [32] Karlsson K, Trullenque-Eriksson A, Tomasi C, Derks J. Efficacy of access flap and pocket elimination procedures in the management of peri-implantitis: A systematic review and meta-analysis. J Clin Periodontol. 2023;50 Suppl 26:244-84.

- [33] Liñares A, Sanz-Sánchez I, Dopico J, Molina A, Blanco J, Montero E. Efficacy of adjunctive measures in the non-surgical treatment of peri-implantitis: A systematic review. J Clin Periodontol. 2023;50 Suppl 26:224-43.
- [34] Ramanauskaite A, Schwarz F, Cafferata EA, Sahrmann P. Photo/mechanical and physical implant surface decontamination approaches in conjunction with surgical peri-implantitis treatment: A systematic review. J Clin Periodontol. 2023;50 Suppl 26:317-35.
- [35] Stiesch M, Grischke J, Schaefer P, Heitz-Mayfield LJA. Supportive care for the prevention of disease recurrence/progression following peri-implantitis treatment: A systematic review. J Clin Periodontol. 2023;50 Suppl 26:113-34.
- [36] Teughels W, Seyssens L, Christiaens V, Temmerman A, Castro AB, Cosyn J. Adjunctive locally and systemically delivered antimicrobials during surgical treatment of periimplantitis: A systematic review. J Clin Periodontol. 2023;50 Suppl 26:359-72.
- [37] Verket A, Koldsland OC, Bunaes D, Lie SA, Romandini M. Non-surgical therapy of periimplant mucositis-Mechanical/physical approaches: A systematic review. J Clin Periodontol. 2023;50 Suppl 26:135-45.
- [38] Wilensky A, Shapira L, Limones A, Martin C. The efficacy of implant surface decontamination using chemicals during surgical treatment of peri-implantitis: A systematic review and meta-analysis. J Clin Periodontol. 2023;50 Suppl 26:336-58.
- [39] Moher D, Liberati A, Tetzlaff J, Altman DG. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. J Clin Epidemiol. 2009;62(10):1006-12.
- [40] dissemination Cfra. Systematic Reviews: CRD's Guidance forUndertaking Reviews in Health Care: Centre for Reviews and Dissemination, University of York, 2008 Published by CRD, University of York January 2009 2009 [This publication presents independent guidance produced by the Centre for Reviews and

Dissemination (CRD). The views expressed in this publication are those of CRD and not

necessarily those of the NHS, the NIHR or the Department of Health.

All rights reserved. Reproduction of this book by photocopying or electronic means for

non-commercial purposes is permitted. Otherwise, no part of this book may be reproduced,

adapted, stored in a retrieval system or transmitted by any means, electronic, mechanical,

- photocopying, or otherwise without the prior written permission of CRD.]. Available from: https://www.york.ac.uk/media/crd/Systematic_Reviews.pdf.
- [41] Guyatt GH, Oxman AD, Kunz R, Atkins D, Brozek J, Vist G, et al. GRADE guidelines: 2.
 Framing the question and deciding on important outcomes. J Clin Epidemiol. 2011;64(4):395-400.
- [42] Carra MC, Blanc-Sylvestre N, Courtet A, Bouchard P. Primary prevention of periimplant diseases: A systematic review and meta-analysis. J Clin Periodontol. 2023;(submitted).
- [43] Stiesch M, Grischke J, Schaefer P, Heitz-Mayfield LJA. Supportive care for the prevention of disease recurrence/progression following peri-implantitis treatment: A systematic review. J Clin Periodontol. 2023;(submitted).

- [44] Verket A, Koldsland OC, Bunæs D, Lie SA, Romandini Mx. Non-surgical therapy of periimplant mucositis – mechanical/physical approaches: a systematic review. J Clin Periodontol. 2023;(submitted).
- [45] Dommisch H, Hoedke D, Vallès Vegas C, Vilarrasa J, Jepsen S, Pascual La Rocca Ax. Efficacy of chemical approaches during non-surgical submarginal instrumentation in the management of peri-implant mucositis: a systematic review. . J Clin Periodontol. 2023;(submitted).
- [46] Gennai S, Bollain J, Ambrosio N, Marruganti C, Graziani F, Figuero E. Efficacy of adjunctive measures in peri-implant mucositis. A systematic review and metaanalysis. J Clin Periodontol. 2023;(submitted).
- [47] Cosgarea R, Roccuzzo A, Jepsen K, Sculean A, Jepsen S, Salvi GE. Efficacy of mechanical/physical approaches for implant surface decontamination in nonsurgical submarginal instrumentation of peri- implantitis. A systematic review. J Clin Periodontol. 2023;(submitted).
- [48] de Waal YCM, Winning L, Stavropoulos A, Polyzois I. Efficacy of chemical approaches for implant surface decontamination in conjunction with sub-marginal instrumentation, in the non-surgical treatment of peri-implantitis. A systematic review and meta-analysis. J Clin Periodontol. 2023;(submitted).
- [49] Liñares A, Sanz-Sánchez I, Dopico J, Molina A, Blanco J, Montero Ex. Efficacy of adjunctive measures in the non-surgical treatment of peri-implantitis. A systematic review. J Clin Periodontol. 2023;(submitted).
- [50] Karlsson K, Trullenque-Eriksson A, Tomasi C, Derks J. Efficacy of access flap and pocket elimination procedures in the management of peri-implantitis a systematic review and meta-analysis. J Clin Periodontol. 2023;(submitted).
- [51] Donos N, Calciolari E, Ghuman M, Baccini M, Sousa V, Nibali Lx. The efficacy of bone reconstructive therapies in the management of peri-implantitis. A systematic review and meta-analysis. J Clin Periodontol. 2023;(submitted).
- [52] Ramanauskaite A, Schwarz F, Cafferata Chea EA, Sahrmann P. Mechanical and physical implant surface decontamination approaches in conjunction with surgical peri-implantitis treatment: A systematic review. J Clin Periodontol. 2023;(submitted).
- [53] Wilensky A, Shapira L, Limones A, Martin C. The Efficacy of Implant Surface Decontamination Using Chemicals during Surgical Treatment of Peri-implantitis: A Systematic Review and Meta-Analysis. J Clin Periodontol. 2023;(submitted).
- [54] Teughels W, Seyssens L, Christiaens V, Temmerman A, Castro A, Cosyn J. Adjunctive locally and systemically delivered antimicrobials during surgical treatment of periimplantitis. J Clin Periodontol. 2023;(submitted).
- [55] Derks J, Ichioka Y, Dionigi C, Trullenque-Eriksson A, Berglundh J, Tomasi C, et al. Prevention and management of peri-implant mucositis and peri-implantitis: A systematic review of outcome measures used in clinical studies in the last 10 years. J Clin Periodontol. 2023;50 Suppl 25:55-66.
- [56] Needleman I, Sanz M, Carrillo de Albornoz A, Safii S, Hassan NHM, Qian S, et al. Involving people with lived experience in developing a core outcome set for implant dentistry research. The Implant Dentistry-Core Outcomes Sets and Measures (ID-COSM) project. J Clin Periodontol. 2023;50 Suppl 25:96-106.
- [57] Sanz M, Carrillo de Albornoz A, Martín C, Needleman I, Tonetti MS. Multi-stakeholder contribution to the identification of a core outcome set and measurements in implant

dentistry (ID-COSM initiative) using the Delphi methodology. J Clin Periodontol. 2023;50 Suppl 25:107-21.

- [58] Tonetti MS, Sanz M, Avila-Ortiz G, Berglundh T, Cairo F, Derks J, et al. Relevant domains, core outcome sets and measurements for implant dentistry clinical trials: The Implant Dentistry Core Outcome Set and Measurement (ID-COSM) international consensus report. J Clin Periodontol. 2023;50 Suppl 25:5-21.
- [59] Murphy MK, Black NA, Lamping DL, McKee CM, Sanderson CF, Askham J, et al. Consensus development methods, and their use in clinical guideline development. Health Technol Assess. 1998;2(3):i-iv, 1-88.
- [60] Balshem H, Helfand M, Schünemann HJ, Oxman AD, Kunz R, Brozek J, et al. GRADE guidelines: 3. Rating the quality of evidence. J Clin Epidemiol. 2011;64(4):401-6.
- [61] Schünemann HJ, Zhang Y, Oxman AD, Group EEiG. Distinguishing opinion from evidence in guidelines. BMJ. 2019;366:14606.
- [62] German Association of the Scientific Medical Societies (AWMF) SGC. AWMF Guidance Manual and Rules for Guideline Development. 2012 [Available from: http://www.awmf.org/leitlinien/awmf-regelwerk.html.
- [63] German Association of the Scientific Medical Societies (AVMF), Standing Guidelines Commission. AWMF Guidance Manual and Rules for Guideline Development.2012. Available from: http://www.awmf.org/leitlinien/awmf-regelwerk.html.
- [64] Schünemann HJ, Al-Ansary LA, Forland F, Kersten S, Komulainen J, Kopp IB, et al. Guidelines International Network: Principles for Disclosure of Interests and Management of Conflicts in Guidelines. Ann Intern Med. 2015;163(7):548-53.
- [65] Shea BJ, Reeves BC, Wells G, Thuku M, Hamel C, Moran J, et al. AMSTAR 2: a critical appraisal tool for systematic reviews that include randomised or non-randomised studies of healthcare interventions, or both. BMJ. 2017;358:j4008.
- [66] Commission TE. *MEDICAL DEVICES: Guidance document Classification of medical devices*. 2010.
- [67] Union. TEPatCotE. REGULATION (EU) 2017/745 OF THE EUROPEAN PARLIAMENT AND OF THE COUNCIL of 5 April 2017 on medical devices, amending Directive 2001/83/EC, Regulation (EC) No 178/2002 and Regulation (EC) No 1223/2009 and repealing Council Directives 90/385/EEC and 93/42/EEC.: Official Journal of the European Union, 117/1 2017.
- [68] Gupta VK, Paul S, Dutta C. Geography, Ethnicity or Subsistence-Specific Variations in Human Microbiome Composition and Diversity. Front Microbiol. 2017;8:1162.
- [69] Strasser T. Reflections on Cardiovascular Diseases. Interdisciplinary Science Reviews. 1978;3(3):225-30.
- [70] Lloyd-Jones DM, Hong Y, Labarthe D, Mozaffarian D, Appel LJ, Van Horn L, et al. Defining and setting national goals for cardiovascular health promotion and disease reduction: the American Heart Association's strategic Impact Goal through 2020 and beyond. Circulation. 2010;121(4):586-613.
- [71] Araujo MG, Lindhe J. Peri-implant health. J Clin Periodontol. 2018;45 Suppl 20:S230-S6.
- [72] Lang NP, Mombelli A, Tonetti MS, Brägger U, Hämmerle CH. Clinical trials on therapies for peri-implant infections. Ann Periodontol. 1997;2(1):343-56.
- [73] Lang NP, Wilson TG, Corbet EF. Biological complications with dental implants: their prevention, diagnosis and treatment. Clin Oral Implants Res. 2000;11 Suppl 1:146-55.

- [74] Mombelli A, Lang NP. The diagnosis and treatment of peri-implantitis. Periodontol 2000. 1998;17:63-76.
- [75] Jepsen S, Berglundh T, Genco R, Aass AM, Demirel K, Derks J, et al. Primary prevention of peri-implantitis: managing peri-implant mucositis. J Clin Periodontol. 2015;42 Suppl 16:S152-7.
- [76] Chan D, Pelekos G, Ho D, Cortellini P, Tonetti MS. The depth of the implant mucosal tunnel modifies the development and resolution of experimental peri-implant mucositis: A case-control study. J Clin Periodontol. 2019;46(2):248-55.
- [77] Valles C, Rodríguez-Ciurana X, Clementini M, Baglivo M, Paniagua B, Nart J. Influence of subcrestal implant placement compared with equicrestal position on the periimplant hard and soft tissues around platform-switched implants: a systematic review and meta-analysis. Clin Oral Investig. 2018;22(2):555-70.
- [78] Schuldt Filho G, Dalago HR, Oliveira de Souza JG, Stanley K, Jovanovic S, Bianchini MA. Prevalence of peri-implantitis in patients with implant-supported fixed prostheses. Quintessence Int. 2014;45(10):861-8.
- [79] Kumar PS, Dabdoub SM, Hegde R, Ranganathan N, Mariotti A. Site-level risk predictors of peri-implantitis: A retrospective analysis. J Clin Periodontol. 2018;45(5):597-604.
- [80] Farina R, Filippi M, Brazzioli J, Tomasi C, Trombelli L. Bleeding on probing around dental implants: a retrospective study of associated factors. J Clin Periodontol. 2017;44(1):115-22.
- [81] De Ry SP, Roccuzzo A, Lang NP, Heitz-Mayfield LJ, Ramseier CA, Sculean A, et al. Evaluation of the implant disease risk assessment (IDRA) tool: A retrospective study in patients with treated periodontitis and implant-supported fixed dental prostheses (FDPs). Clin Oral Implants Res. 2021;32(11):1299-307.
- [82] Grischke J, Szafrański SP, Muthukumarasamy U, Haeussler S, Stiesch M. Removable denture is a risk indicator for peri-implantitis and facilitates expansion of specific periodontopathogens: a cross-sectional study. BMC Oral Health. 2021;21(1):173.
- [83] Heitz-Mayfield LJA, Heitz F, Lang NP. Implant Disease Risk Assessment IDRA-a tool for preventing peri-implant disease. Clin Oral Implants Res. 2020;31(4):397-403.
- [84] Katafuchi M, Weinstein BF, Leroux BG, Chen YW, Daubert DM. Restoration contour is a risk indicator for peri-implantitis: A cross-sectional radiographic analysis. J Clin Periodontol. 2018;45(2):225-32.
- [85] Mattheos N, Vergoullis I, Janda M, Miseli A. The Implant Supracrestal Complex and Its Significance for Long-Term Successful Clinical Outcomes. Int J Prosthodont. 2021;34(1):88-100.
- [86] Mattheos N, Janda M, Acharya A, Pekarski S, Larsson C. Impact of design elements of the implant supracrestal complex (ISC) on the risk of peri-implant mucositis and peri-implantitis: A critical review. Clin Oral Implants Res. 2021;32 Suppl 21:181-202.
- [87] Serino G, Ström C. Peri-implantitis in partially edentulous patients: association with inadequate plaque control. Clinical Oral Implants Research. 2009;20(2):169-74.
- [88] Soulami S, Slot DE, van der Weijden F. Implant-abutment emergence angle and profile in relation to peri-implantitis: A systematic review. Clin Exp Dent Res. 2022;8(4):795-806.
- [89] Staubli N, Walter C, Schmidt JC, Weiger R, Zitzmann NU. Excess cement and the risk of peri-implant disease – a systematic review. Clinical Oral Implants Research. 2017;28(10):1278-90.

- [90] Strauss FJ, Siegenthaler M, Hämmerle CHF, Sailer I, Jung RE, Thoma DS. Restorative angle of zirconia restorations cemented on non-original titanium bases influences the initial marginal bone loss: 5-year results of a prospective cohort study. Clin Oral Implants Res. 2022;33(7):745-56.
- [91] Yi Y, Koo KT, Schwarz F, Ben Amara H, Heo SJ. Association of prosthetic features and peri-implantitis: A cross-sectional study. J Clin Periodontol. 2020;47(3):392-403.
- [92] Lindhe J, Meyle J, Periodontology GDoEWo. Peri-implant diseases: Consensus Report of the Sixth European Workshop on Periodontology. J Clin Periodontol. 2008;35(8 Suppl):282-5.
- [93] Sanz M, Schwarz F, Herrera D, McClain P, Figuero E, Molina A, et al. Importance of keratinized mucosa around dental implants: Consensus report of group 1 of the DGI/SEPA/Osteology Workshop. Clin Oral Implants Res. 2022;33 Suppl 23:47-55.
- [94] Ziebolz D, Klipp S, Schmalz G, Schmickler J, Rinke S, Kottmann T, et al. Comparison of different maintenance strategies within supportive implant therapy for prevention of peri-implant inflammation during the first year after implant restoration. A randomized, dental hygiene practice-based multicenter study. Am J Dent. 2017;30(4):190-6.
- [95] Cho-Yan Lee J, Mattheos N, Nixon KC, Ivanovski S. Residual periodontal pockets are a risk indicator for peri-implantitis in patients treated for periodontitis. Clin Oral Implants Res. 2012;23(3):325-33.
- [96] Costa FO, Lages EJP, Cortelli SC, Cortelli JR, Mattos-Pereira GH, Lima RPE, et al. Association between cumulative smoking exposure, span since smoking cessation, and peri-implantitis: a cross-sectional study. Clin Oral Investig. 2022;26(7):4835-46.
- [97] Frisch E, Ziebolz D, Vach K, Ratka-Krüger P. The effect of keratinized mucosa width on peri-implant outcome under supportive postimplant therapy. Clin Implant Dent Relat Res. 2015;17 Suppl 1:e236-44.
- [98] Roccuzzo M, Grasso G, Dalmasso P. Keratinized mucosa around implants in partially edentulous posterior mandible: 10-year results of a prospective comparative study. Clin Oral Implants Res. 2016;27(4):491-6.
- [99] Ramanauskaite A, Schwarz F, Sader R. Influence of width of keratinized tissue on the prevalence of peri-implant diseases: A systematic review and meta-analysis. Clin Oral Implants Res. 2022;33 Suppl 23:8-31.
- [100] Tavelli L, Barootchi S, Avila-Ortiz G, Urban IA, Giannobile WV, Wang HL. Peri-implant soft tissue phenotype modification and its impact on peri-implant health: A systematic review and network meta-analysis. J Periodontol. 2021;92(1):21-44.
- [101] Valles C, Vilarrasa J, Barallat L, Pascual A, Nart J. Efficacy of soft tissue augmentation procedures on tissue thickening around dental implants: A systematic review and meta-analysis. Clin Oral Implants Res. 2022;33 Suppl 23:72-99.
- [102] Hosseini M, Worsaae N, Gotfredsen K. Tissue changes at implant sites in the anterior maxilla with and without connective tissue grafting: A five-year prospective study. Clin Oral Implants Res. 2020;31(1):18-28.
- [103] Truhlar RS, Morris HF, Ochi S. The efficacy of a counter-rotational powered toothbrush in the maintenance of endosseous dental implants. J Am Dent Assoc. 2000;131(1):101-7.
- [104] Alhakeem M, Kanounisabet N, Nowzari H, Aslroosta H, Moslemi N. Risk indicators of long-term outcome of implant therapy in patients with a history of severe periodontitis or no history of periodontitis: A retrospective cohort study. Int J Dent Hyg. 2023;21(1):227-37.

- [105] Roccuzzo A, Imber JC, Marruganti C, Salvi GE, Ramieri G, Roccuzzo M. Clinical outcomes of dental implants in patients with and without history of periodontitis: A 20-year prospective study. J Clin Periodontol. 2022;49(12):1346-56.
- [106] Monje A, Amerio E, Farina R, Nart J, Ramanauskaite A, Renvert S, et al. Significance of probing for monitoring peri-implant diseases. Int J Oral Implantol (Berl). 2021;14(4):385-99.
- [107] Reinedahl D, Chrcanovic B, Albrektsson T, Tengvall P, Wennerberg A. Ligature-Induced Experimental Peri-Implantitis-A Systematic Review. J Clin Med. 2018;7(12).
- [108] Zitzmann NU, Berglundh T, Marinello CP, Lindhe J. Experimental peri-implant mucositis in man. J Clin Periodontol. 2001;28(6):517-23.
- [109] Salvi GE, Aglietta M, Eick S, Sculean A, Lang NP, Ramseier CA. Reversibility of experimental peri-implant mucositis compared with experimental gingivitis in humans. Clin Oral Implants Res. 2012;23(2):182-90.
- [110] Riben-Grundstrom C, Norderyd O, André U, Renvert S. Treatment of peri-implant mucositis using a glycine powder air-polishing or ultrasonic device: a randomized clinical trial. J Clin Periodontol. 2015;42(5):462-9.
- [111] de Tapia B, Bonnin M, Valles C, Mozas C, Herrera D, Sanz M, et al. Clinical outcomes and associated factors in the treatment of peri-implant mucositis, combining mechanical debridement and prosthesis modification: A 30-month follow-up prospective case series. J Clin Periodontol. 2022;49(12):1357-65.
- [112] de Tapia B, Mozas C, Valles C, Nart J, Sanz M, Herrera D. Adjunctive effect of modifying the implant-supported prosthesis in the treatment of peri-implant mucositis. J Clin Periodontol. 2019;46(10):1050-60.
- [113] Renvert S, Lessem J, Dahlén G, Lindahl C, Svensson M. Topical minocycline microspheres versus topical chlorhexidine gel as an adjunct to mechanical debridement of incipient peri-implant infections: a randomized clinical trial. J Clin Periodontol. 2006;33(5):362-9.
- [114] Butera A, Pascadopoli M, Pellegrini M, Gallo S, Zampetti P, Cuggia G, et al. Domiciliary Use of Chlorhexidine vs. Postbiotic Gels in Patients with Peri-Implant Mucositis: A Split-Mouth Randomized Clinical Trial. Applied Sciences [Internet]. 2022; 12(6).
- [115] Sahrmann P, Bettschart C, Wiedemeier DB, Al-Majid A, Attin T, Schmidlin PR. Treatment of Peri-Implant Mucositis with Repeated Application of Chlorhexidine Chips or Gel during Supportive Therapy - A Randomized Clinical Trial. Dent J (Basel). 2019;7(4).
- [116] Żółkiewicz J, Marzec A, Ruszczyński M, Feleszko W. Postbiotics-A Step Beyond Preand Probiotics. Nutrients. 2020;12(8).
- [117] Menezes KM, Fernandes-Costa AN, Silva-Neto RD, Calderon PS, Gurgel BC. Efficacy of 0.12% Chlorhexidine Gluconate for Non-Surgical Treatment of Peri-Implant Mucositis. J Periodontol. 2016;87(11):1305-13.
- [118] Iorio-Siciliano V, Blasi A, Stratul SI, Ramaglia L, Sculean A, Salvi GE, et al. Antiinfective therapy of peri-implant mucositis with adjunctive delivery of a sodium hypochlorite gel: a 6-month randomized triple-blind controlled clinical trial. Clin Oral Investig. 2020;24(6):1971-9.
- [119] Lombardo G, Signoriello A, Corrocher G, Signoretto C, Burlacchini G, Pardo A, et al. A Topical Desiccant Agent in Association with Manual Debridement in the Initial Treatment of Peri-Implant Mucositis: A Clinical and Microbiological Pilot Study. Antibiotics (Basel). 2019;8(2).

- [120] Poppolo Deus F, Ouanounou A. Chlorhexidine in Dentistry: Pharmacology, Uses, and Adverse Effects. Int Dent J. 2022;72(3):269-77.
- [121] Pulcini A, Bollaín J, Sanz-Sánchez I, Figuero E, Alonso B, Sanz M, et al. Clinical effects of the adjunctive use of a 0.03% chlorhexidine and 0.05% cetylpyridinium chloride mouth rinse in the management of peri-implant diseases: A randomized clinical trial. J Clin Periodontol. 2019;46(3):342-53.
- [122] Suvan J, Leira Y, Moreno Sancho FM, Graziani F, Derks J, Tomasi C. Subgingival instrumentation for treatment of periodontitis. A systematic review. J Clin Periodontol. 2020;47 Suppl 22:155-75.
- [123] Abduljabbar T, Javed F, Kellesarian SV, Vohra F, Romanos GE. Effect of Nd:YAG laserassisted non-surgical mechanical debridement on clinical and radiographic periimplant inflammatory parameters in patients with peri-implant disease. J Photochem Photobiol B. 2017;168:16-9.
- [124] Alpaslan Yayli NZ, Talmac AC, Keskin Tunc S, Akbal D, Altindal D, Ertugrul AS. Erbium, chromium-doped: yttrium, scandium, gallium, garnet and diode lasers in the treatment of peri-implantitis: clinical and biochemical outcomes in a randomized-controlled clinical trial. Lasers Med Sci. 2022;37(1):665-74.
- [125] Roccuzzo A, Klossner S, Stähli A, Imber JC, Eick S, Sculean A, et al. Non-surgical mechanical therapy of peri-implantitis with or without repeated adjunctive diode laser application. A 6-month double-blinded randomized clinical trial. Clin Oral Implants Res. 2022;33(9):900-12.
- [126] Schwarz F, Bieling K, Bonsmann M, Latz T, Becker J. Nonsurgical treatment of moderate and advanced periimplantitis lesions: a controlled clinical study. Clinical Oral Investigations. 2006;10(4):279-88.
- [127] Schwarz F, Sculean A, Rothamel D, Schwenzer K, Georg T, Becker J. Clinical evaluation of an Er:YAG laser for nonsurgical treatment of peri-implantitis: a pilot study. Clin Oral Implants Res. 2005;16(1):44-52.
- [128] Sahm N, Becker J, Santel T, Schwarz F. Non-surgical treatment of peri-implantitis using an air-abrasive device or mechanical debridement and local application of chlorhexidine: a prospective, randomized, controlled clinical study. J Clin Periodontol. 2011;38(9):872-8.
- [129] Merli M, Bernardelli F, Giulianelli E, Carinci F, Mariotti G, Pini-Prato G, et al. Short-term comparison of two non-surgical treatment modalities of peri-implantitis: Clinical and microbiological outcomes in a two-factorial randomized controlled trial. J Clin Periodontol. 2020;47(10):1268-80.
- [130] Renvert S, Lindahl C, Roos Jansåker AM, Persson GR. Treatment of peri-implantitis using an Er:YAG laser or an air-abrasive device: a randomized clinical trial. J Clin Periodontol. 2011;38(1):65-73.
- [131] Alonso V, García-Caballero L, Couto I, Diniz M, Diz P, Limeres J. Subcutaneous emphysema related to air-powder tooth polishing: a report of three cases. Aust Dent J. 2017;62(4):510-5.
- [132] Bassetti M, Bassetti R, Sculean A, Salvi GE. [Subcutaneous emphysema following non-surgical peri-implantitis therapy using an air abrasive device: a case report]. Swiss Dent J. 2014;124(7-8):807-17.
- [133] Bruckmann C, Bruckmann L, Gahleitner A, Stavropoulos A, Bertl K. Pneumocephalus as result of nonsurgical peri-implantitis treatment with an air-polishing device for submucosal debridement-A case report. Clin Exp Dent Res. 2022;8(3):632-9.

- [134] Wang H, Li W, Zhang D, Wang Z. Adjunctive photodynamic therapy improves the outcomes of peri-implantitis: a randomized controlled trial. Aust Dent J. 2019;64(3):256-62.
- [135] Alasqah MN. Influence of adjunctive non-surgical peri-implant therapy on clinical and salivary cytokine profile in obese patients. Photodiagnosis Photodyn Ther. 2022;37:102721.
- [136] Vohra F, Al-Rifaiy MQ, Lillywhite G, Abu Hassan MI, Javed F. Efficacy of mechanical debridement with adjunct antimicrobial photodynamic therapy for the management of peri-implant diseases: a systematic review. Photochem Photobiol Sci. 2014;13(8):1160-8.
- [137] Isola G, Matarese G, Williams RC, Siciliano VI, Alibrandi A, Cordasco G, et al. The effects of a desiccant agent in the treatment of chronic periodontitis: a randomized, controlled clinical trial. Clin Oral Investig. 2018;22(2):791-800.
- [138] Lombardo G, Signoretto C, Corrocher G, Pardo A, Pighi J, Rovera A, et al. A topical desiccant agent in association with ultrasonic debridement in the initial treatment of chronic periodontitis: a clinical and microbiological study. New Microbiol. 2015;38(3):393-407.
- [139] Renvert S, Lessem J, Dahlén G, Renvert H, Lindahl C. Mechanical and repeated antimicrobial therapy using a local drug delivery system in the treatment of periimplantitis: a randomized clinical trial. J Periodontol. 2008;79(5):836-44.
- [140] Herrera D, Matesanz P, Martín C, Oud V, Feres M, Teughels W. Adjunctive effect of locally delivered antimicrobials in periodontitis therapy: A systematic review and meta-analysis. J Clin Periodontol. 2020;47 Suppl 22:239-56.
- [141] Machtei EE, Frankenthal S, Levi G, Elimelech R, Shoshani E, Rosenfeld O, et al. Treatment of peri-implantitis using multiple applications of chlorhexidine chips: a double-blind, randomized multi-centre clinical trial. J Clin Periodontol. 2012;39(12):1198-205.
- [142] Machtei EE, Romanos G, Kang P, Travan S, Schmidt S, Papathanasiou E, et al. Repeated delivery of chlornexidine chips for the treatment of peri-implantitis: A multicenter, randomized, comparative clinical trial. J Periodontol. 2021;92(1):11-20.
- [143] Teughels W, Feres M, Oud V, Martín C, Matesanz P, Herrera D. Adjunctive effect of systemic antimicrobials in periodontitis therapy: A systematic review and metaanalysis. J Clin Periodontol. 2020;47 Suppl 22:257-81.
- [144] Blanco C, Pico A, Dopico J, Gándara P, Blanco J, Liñares A. Adjunctive benefits of systemic metronidazole on non-surgical treatment of peri-implantitis. A randomized placebo-controlled clinical trial. J Clin Periodontol. 2022;49(1):15-27.
- [145] Shibli JA, Ferrari DS, Siroma RS, Figueiredo LC, Faveri M, Feres M. Microbiological and clinical effects of adjunctive systemic metronidazole and amoxicillin in the nonsurgical treatment of peri-implantitis: 1 year follow-up. Braz Oral Res. 2019;33(suppl 1):e080.
- [146] Hill C, Guarner F, Reid G, Gibson GR, Merenstein DJ, Pot B, et al. Expert consensus document. The International Scientific Association for Probiotics and Prebiotics consensus statement on the scope and appropriate use of the term probiotic. Nat Rev Gastroenterol Hepatol. 2014;11(8):506-14.
- [147] Gatej SM, Marino V, Bright R, Fitzsimmons TR, Gully N, Zilm P, et al. Probiotic Lactobacillus rhamnosus GG prevents alveolar bone loss in a mouse model of experimental periodontitis. J Clin Periodontol. 2018;45(2):204-12.

- [148] Invernici MM, Furlaneto FAC, Salvador SL, Ouwehand AC, Salminen S, Mantziari A, et al. Bifidobacterium animalis subsp lactis HN019 presents antimicrobial potential against periodontopathogens and modulates the immunological response of oral mucosa in periodontitis patients. PLoS One. 2020;15(9):e0238425.
- [149] Laleman I, Pauwels M, Quirynen M, Teughels W. The usage of a lactobacilli probiotic in the non-surgical therapy of peri-implantitis: A randomized pilot study. Clin Oral Implants Res. 2020;31(1):84-92.
- [150] de Waal YC, Raghoebar GM, Meijer HJ, Winkel EG, van Winkelhoff AJ. Implant decontamination with 2% chlorhexidine during surgical peri-implantitis treatment: a randomized, double-blind, controlled trial. Clin Oral Implants Res. 2015;26(9):1015-23.
- [151] Koldsland OC, Wohlfahrt JC, Aass AM. Surgical treatment of peri-implantitis: Prognostic indicators of short-term results. J Clin Periodontol. 2018;45(1):100-13.
- [152] Karlsson K, Derks J, Håkansson J, Wennström JL, Petzold M, Berglundh T. Interventions for peri-implantitis and their effects on further bone loss: A retrospective analysis of a registry-based cohort. J Clin Periodontol. 2019;46(8):872-9.
- [153] Carcuac O, Derks J, Abrahamsson I, Wennström JL, Petzold M, Berglundh T. Surgical treatment of peri-implantitis: 3-year results from a randomized controlled clinical trial. J Clin Periodontol. 2017;44(12):1294-303.
- [154] Carcuac O, Derks J, Abrahamsson I, Wennström JL, Berglundh T. Risk for recurrence of disease following surgical therapy of peri-implantitis-A prospective longitudinal study. Clin Oral Implants Res. 2020;31(11):1072-7.
- [155] Tormena M, Matarazzo F, de Oliveira BM, Pereira O, Previdelli I, Araújo MG. The effect of inadequate access to peri-implant hygiene on marginal bone level. A 4-year cohort prospective study. Clin Oral Implants Res. 2020;31(9):836-45.
- [156] Heitz-Mayfield LJA, Salvi GE, Mombelli A, Loup PJ, Heitz F, Kruger E, et al. Supportive peri-implant therapy following anti-infective surgical peri-implantitis treatment: 5-year survival and success. Clin Oral Implants Res. 2018;29(1):1-6.
- [157] Heitz-Mayfield LJA, Heitz F, Koong B, Huang T, Chivers P. Surgical peri-implantitis treatment with and without guided bone regeneration. A randomized controlled trial. Clinical Oral Implants Research. 2023;34(9):892-910.
- [158] Jepsen S, Schwarz F, Cordaro L, Derks J, Hämmerle CHF, Heitz-Mayfield LJ, et al. Regeneration of alveolar ridge defects. Consensus report of group 4 of the 15th European Workshop on Periodontology on Bone Regeneration. J Clin Periodontol. 2019;46 Suppl 21:277-86.
- [159] Lang NP, Berglundh T, Periodontology WGoSEWo. Periimplant diseases: where are we now?--Consensus of the Seventh European Workshop on Periodontology. J Clin Periodontol. 2011;38 Suppl 11:178-81.
- [160] Hentenaar DFM, De Waal YCM, Stewart RE, Van Winkelhoff AJ, Meijer HJA, Raghoebar GM. Erythritol air polishing in the surgical treatment of peri-implantitis: A randomized controlled trial. Clin Oral Implants Res. 2022;33(2):184-96.
- [161] Toma S, Brecx MC, Lasserre JF. Clinical Evaluation of Three Surgical Modalities in the Treatment of Peri-Implantitis: A Randomized Controlled Clinical Trial. J Clin Med. 2019;8(7).
- [162] Schwarz F, John G, Mainusch S, Sahm N, Becker J. Combined surgical therapy of periimplantitis evaluating two methods of surface debridement and decontamination. A two-year clinical follow up report. J Clin Periodontol. 2012;39(8):789-97.

- [163] Wang CW, Ashnagar S, Gianfilippo RD, Arnett M, Kinney J, Wang HL. Laser-assisted regenerative surgical therapy for peri-implantitis: A randomized controlled clinical trial. J Periodontol. 2021;92(3):378-88.
- [164] de Tapia B, Valles C, Ribeiro-Amaral T, Mor C, Herrera D, Sanz M, et al. The adjunctive effect of a titanium brush in implant surface decontamination at peri-implantitis surgical regenerative interventions: A randomized controlled clinical trial. J Clin Periodontol. 2019;46(5):586-96.
- [165] Carcuac O, Derks J, Charalampakis G, Abrahamsson I, Wennström J, Berglundh T. Adjunctive Systemic and Local Antimicrobial Therapy in the Surgical Treatment of Periimplantitis: A Randomized Controlled Clinical Trial. J Dent Res. 2016;95(1):50-7.
- [166] Hallström H, Persson GR, Lindgren S, Renvert S. Open flap debridement of periimplantitis with or without adjunctive systemic antibiotics: A randomized clinical trial. J Clin Periodontol. 2017;44(12):1285-93.
- [167] Cha JK, Lee JS, Kim CS. Surgical Therapy of Peri-Implantitis with Local Minocycline: A 6-Month Randomized Controlled Clinical Trial. J Dent Res. 2019;98(3):288-95.
- [168] Emanuel N, Machtei EE, Reichart M, Shapira L. D-PLEX500: a local biodegradable prolonged release doxycycline-formulated bone graft for the treatment for periimplantitis. A randomized controlled clinical study. Quintessence Int. 2020;51(7):546-53.

Figure 4. Chronological flow of interventions, according to implant therapy stage and to the diagnosis of the peri-implant condition.

STAGE OF IMPLANT THERAPY	patients awaiting dental implant rehabilitation	patients receiving dental implant rehabilitation	patients already having dental implant(s)				
PERI-IMPLANT CONDITION DIAGNOSIS	before dental implant placement		healthy peri-implant tissues	peri-implant mucositis	treated peri-implant mucositis	peri-implantitis	treated peri-implantitis
Preventive interventions	"Primordial" prevention						
	Interventions during implant/prosthesis placement						
			Primary prevention of peri-implant mucositis	Secondary prevention of peri-implant mucositis / Primary prevention of peri-implantitis			Secondary prevention of peri-implantitis
Therapeutic interventions	\mathbf{O}			peri-implant mucositis therapy		non-surgical step of peri- implantitis therapy	
	3			re-evaluation		re-evaluation	
						surgical step of peri- implantitis therapy	

* "Primordial prevention" involves preventing the development of risk factors for peri-implant diseases. Therefore, primordial prevention also applies to patients with implants who have healthy peri-implant tissues and no risk factors. However, for the purpose of this guideline, the presence of an implant was regarded as a risk factor for peri-implant diseases (e.g. due to plaque accumulation on a non-shedding surface). Therefore, in the above scheme, primordial prevention does not apply once a dental implant has been placed.