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Title picture: From the case report of Karin Groß and Stefan Wolfart, here Figure 1: Fixation of the lateral incisors – dimensioned and shaped according to the wax-up – for attaining a harmonious dental arch in the congenital absence of 12 and 22, p. 160–167; (Fig. 1: K. Groß)

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Are mini-implants an alternative to standard-diameter implants?

What must I be aware of when using mini-implants?
Dental implants are an indispensable part of the dental treatment spectrum and represent an integral component of contemporary prosthodontic treatment concepts. The use of implants for both removable and fixed prosthodontics offers clinically relevant advantages, which have been recorded for patients in the form of an improvement in the oral health-related quality of life and masticatory function, as well as, better long-term restorative treatment prognosis [9].

Implant-prosthetic concepts
The prognosis for fixed restorations on implants is very good (5-year prognosis 96.4 %, 10-year prognosis 93.9 %) and corresponds to the prognosis for short-span restorations on teeth [29]. Thus, through the use of implants, the grinding of the natural teeth and long-span bridges can be avoided, while free-end edentulous spaces can also be treated. Moreover, it is possible to restore the edentulous jaw with a complete denture using 4–8 implants [25, 26].

In the case of removable dentures, the use of implants in the edentulous jaw (1–6 implants in the mandible and 4–8 implants in the maxilla) can significantly improve the masticatory function and satisfaction with dental restorations. Patients prefer good fixation and low rotation of dentures [9]: special consideration is needed to provide an utmost quadrangular support, while the axes of rotation and selection of the attachment system concept must be thoroughly thought out. In order to achieve the best possible retention and support for a denture in a partially edentulous dentition, so-called strategic implants, which are placed in strategically important positions, can be used. In cases of complete edentulism, one or two implants may be used to reduce the denture’s rotation. A rigidly supported, removable denture, or more specifically, a denture that wobbles less, not only leads to better patient satisfaction, but also to less wear and tear of the attachment systems, and thus to a better long-term prognosis.

Narrow-Diameter-Implants (NDIs)
The described implant-prosthetic concepts for standard diameter implants (> 3.5 mm) have been investigated and are evidence-based. The results, however, cannot be generalized 1:1 with regard to reduced diameter implants.

Reduced diameter implants are also called Narrow-Diameter-Implants (NDIs). They have diameters that range between 1.8–3.5 mm and can be divided into 3 categories based on their diameter [20]:

Category 1 is comprised of mini-implants (MDI, 1.8–2.5 mm) which are basically one-piece. Category 2 (diameter 2.5–3.25 mm) and category 3 (diameter 3.3–3.5 mm) consist of two-piece implants.

Category 3 NDIs that are made of pure titanium (titanium grade IV) have only one fifth of the mechanical load capacity (200 N vs. 1000 N) in comparison to standard-diameter implants (4.1 mm) [5]. This reduction of implant diameter consequently leads to an increased risk of fracture, at least in theory. Therefore, the susceptibility to fracture of the two-piece category 2 and 3 implants can be reduced by making modifications to the implant-abutment connection, which allows for thicker implant wall thickness, or by employing titanium-zirconium alloys with a higher fracture resistance. Category 1 implants cannot be made of pure titanium due to the high fracture potential; instead, they are produced from a titanium alloy (Titanium Grade V, Ti6Al-4V ELI) and are a single piece, as a two-piece design would reduce the wall thickness.
It is also known from materials science and finite element studies that a change in implant geometry leads to a changed force distribution into the peri-implant cortical bone [19]. In this regard, a change in implant diameter has a greater effect than a change in implant length [8]: an increase in diameter from 2.5 mm to 3.3 mm reduces the stress on the cortical bone by 30.7 %, whereas an increase in implant length from 8.5 mm to 15 mm reduces it by only 1.7 %.

This raises the question of the survival prognosis of reduced diameter implants, especially in cases of compromised bone supply. It should be noted that in spite of the fact that very good implant survival and success rates have been documented for standard-diameter implants [24], these were not achieved in cases of poor bone quality and quantity [7, 15].

The scientific data published to date on NDIs shows very good results with regard to implant survival and success which is comparable to that of standard implants (90–100 %) [6, 35, 36]. Nonetheless, a recent review indicates that there are significant differences between the 3 categories of NDIs in terms of failure rates: category 2 and 3 show very good prognoses, which are comparable to standard-diameter implants. In contrast, category 1 implants which are mini-implants with a diameter of 2.5 mm or less, show a significantly higher risk of implant loss with an odds ratio of 4.54 (CI: 1.51–13.65) [33]. Therefore, if possible, a category 2 and 3 NDI or a category 1 implant which is thicker should preferably be chosen.

Modern category 3 NDIs show very good results and, with regard to the prosthetic concepts described above, can probably be employed similarly to standard implants in many cases. The use of category 2 and 3 NDIs is unproblematic for the indication of lower incisor and upper lateral incisor single-tooth implants, but their use in the molar region, where high masticatory loads exist, is not recommended [19, 33]. If these principles are followed, a very good implant survival rate (90–100 %) has been documented for fixed restorations on reduced diameter implants [36].

**Mini-implants**

The extremely resorbed alveolar bone can be so thin, however, that only category 1 implants are possible without bone augmentation. There are thus clinically relevant indications for this implant group: in a recent review of category 1 implants, very good survival rates (98 %) and success rates (93 %) were reported for the indication stabilization of a complete mandibular denture [27]. For the same indication, in a recently published prospective 5-year study performed at the University of Bern, a survival and success rate of 100 % for immediately loaded 1.8 mm diameter implants [12, 13] was documented. However, the use of category 1 NDIs for the indication of fixed restorations does not seem advisable. Although mini-implants can function well as described above, they should not be taken lightly. The success of therapy with mini-implants depends on patient selection and the experience of the dental practitioner [31]. In a study conducted in 5 centers and on 1029 examined implants, it was shown that the average implant survival rate was 91 %. When the data was assessed for each particular center, however, the situation was as follows: Four centers achieved success rates of over 90 % while the fifth achieved a success rate of only 69 %, as 13 from 42 implants were lost [10]. An intensive analysis of the surgical procedure therefore seems advisable: the surgical procedure is in principle very straightforward, but the drilling...
The use of mini-implants has been shown to considerably improve the masticatory function and biting power in both older and younger patients [14], although elderly patients needed more time to attain the improvements in masticatory function [12]. In spite of these benefits, the retention of the implant overdentures using the O-rings of the MDI matrices is less rigid than the anchorage provided by a milled parallel-walled bar. This represents a certain loss of comfort which must be taken into account during individual therapy planning. Nevertheless, in a prospective 5-year study, it was shown that the oral health-related quality of life improved significantly through the use of 4 interforaminal implants [30]. The mobility can also bring about a clinically relevant benefit: the spherical ball-shaped patrices of the mini-implants display hardly any tartar build-up and the peri-implant mucosa is usually healthy, which can be attributed to the self-cleaning effect of the rubber O-rings [37]. It should be emphasized that the absence of peri-implant keratinized mucosa did not lead to increased bone resorption rates [13]. For prophylactic reasons, however, a peri-implant keratinized mucosa is still recommended for easier implant hygiene.

In the edentulous mandible, mini-implants have been proven to be a safe therapy option for better retention of removable dentures. [10, 13, 35]. In addition to their indication in edentulous patients, partially edentulous patients can also benefit from the use of mini-implants.

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Clinical tips for mini-implant use

The simple surgical and prosthetic procedure, which requires minimal material and time, coupled with less follow-up care, is highly valued by dental practitioners [34]. For prosthetic planning, it is important to note that mini-implants should be well distributed and respect a minimum distance of 4 mm from each other. If the distance is less than 4 mm, difficulties arise in positioning the matrices side by side due to the matrix housing size. In patients with strong masticatory forces and thin mandibular dentures, where the plastic coating of the matrix is less than 2 mm thick, there is an increased denture fracture risk in the area of the matrix housing [11]. In a study by Mundt et al., a fracture rate of 20 % was determined for MDI-implant overdentures which had no model cast reinforcement [28]. The incorporation of a lingual 1 mm thick reinforcement band made of cast alloy is recommended in such cases [13]. In a recent prospective study with 5-year results, a 35 % fracture rate was determined for mandibular dentures without model cast reinforcement. After the model cast reinforcement was adopted, no further fractures occurred [13]. A lingual reinforcement, or more specifically, a lingual thickening of a mandibular prosthesis by about 2-4 mm is well tolerated by many patients and is not perceived as disturbing [4].

The recommendation of a model cast prosthesis after a fracture has oc-
curred can lead to disagreements with the patient: therefore, it is advisable to offer a model cast reinforcement as a matter of principle, which can even be integrated at a later point in mandibular dentures. In the maxilla, a model cast reinforcement should be planned from the outset, as the thickness of the denture is consistently reduced.

According to the clinical protocol for mini-implant systems, 4 implants in the mandible and 6 implants in the maxilla are needed for implant overdenture retention. If a minimum insertion torque of 35 Ncm is achieved, mandibular MDIs may be loaded immediately. In the case of lower primary stability values, immediate loading in the mandible is not advisable and a soft relining is recommended instead; just after a 3-month healing period, the matrix housings can be polymerized and the delayed loading of the implants performed. In the maxilla, a 6-month healing period is generally recommended before loading. However, data from the University of Montreal in Canada show that in the mandible, a torque of 15 Ncm would be sufficient for immediate loading [21].

Despite the good implant survival outcomes in the mandible, the same results should not readily be assumed for the maxilla. As the rates of implant loss are higher in the edentulous maxilla, Shatkin et al. report a survival rate of 95.1 % for mandibular and 83.2 % for maxillary overdentures [35]. The working group for implantology and biomaterials research at the University of Bonn reported similar results with respect to the differences between the upper and lower jaws, although these are momentarily still being scientifically evaluated. With regard to the increased rates of implant loss in the maxilla, there appears to be a cluster effect, meaning that more implants are lost in single patients. Increased caution is therefore required in the edentulous maxilla. This assessment is consistent with the recommendations of the International Team for Implantology (ITI) and its SAC system: the edentulous mandible is classified surgically as being straightforward and the edentulous maxilla as being complex.

**Conclusion**

The simplicity of treatment and low material costs result in the fact that social indications can also be treated with mini-implants. Using MDIs, it is possible to come close to the goal, as formulated in various scientific statements, of restoring the edentulous mandible in patients using implant-supported overdentures [16]. Treatment with 4–6 mini-implants also opens up new possibilities for a minimally invasive approach. Future scientific studies are necessary in order to explore the limits for mini-implant indication. Mini-implants already represent an established and well documented treatment option for treating the edentulous mandible. On the other hand, they are not suitable for the indication of fixed restorations in masticatory load bearing areas. Their mechanical inferiority compared to standard-diameter implants has been proven and it must be taken into account when determining their indication [5, 17, 19].

**Conflicts of interest**

In the past, Prof. Dr. Norbert Enkling has given paid lectures at scientific conferences and lectures with workshops for implant companies such as Nobel Biocare, SIC Invent, Dentalium Implants, 3M Espe and Condent.

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Karin Groß, Stefan Wolfart

Interfaces between orthodontics and prosthodontics: interdisciplinary possibilities and “innovations”

Introduction: This article presents possibilities regarding how orthodontists and prosthodontists can cooperate together to create conditions for an esthetically pleasing and functionally oriented treatment result. When planning the prosthetic treatment of anterior gaps, the gaps can be adjusted in advance by means of orthodontic treatment so that they can subsequently be treated in the best possible esthetic and functional manner.

Material and Methods: Adhesive bridges represent a suitable treatment option in adolescents for congenitally missing lateral incisors or after anterior tooth loss due to trauma. When designed with one wing, adhesive bridges can be used before the completion of jaw bone development and, if necessary, they can be orthodontically positioned together with the abutment tooth. Prior to complex prosthetic rehabilitation in the esthetic area, malpositioned abutment teeth can be orthodontically positioned or aligned to generate space for an ideal restoration design. Moreover, orthodontic extrusion provides the possibility of achieving sufficient abutment heights in the case of deeply destroyed or fractured teeth, thus permitting the adequate treatment of these critical teeth according to prosthetic guidelines. Oftentimes, in the case of tilted, protruded or retruded teeth, an orthodontic intervention previous to tooth preparation is useful for tooth substance conservation and maintaining the vitality of abutment teeth.

Conclusion: It can be concluded that a specific combination of orthodontic and prosthetic treatment measures can achieve a better restorative treatment result in terms of function, esthetics and long-term stability. Often, this is accompanied by a reduction in treatment duration and costs.

Keywords: interdisciplinary treatments; gap management; extrusion; esthetics; adhesive bridges
Introduction

In order to achieve esthetically pleasing and functionally optimized treatment results in complex treatment cases, close cooperation between the orthodontist and prosthodontist is often advisable. In ideal circumstances, this teamwork extends from treatment planning to joint treatment implementation and is rewarded by satisfied patients and high-quality treatment results. The overlap, or interface, between the two disciplines is multifaceted. This article discusses 4 of these interfaces and presents clinical examples to illustrate them:

1. If individual teeth are missing, the question of which specialty domain should have pre-eminence, and accordingly the responsibility, often arises: Should an existing dental gap be orthodontically closed, or be restored by prosthetic means, such as a bridge, adhesive bridge or implant? Each of these approaches has advantages and disadvantages; they can best be discussed in a joint consultation between the dentist, orthodontist and patient.

2. In the case that a dental gap is to be treated by prosthetic means, it is necessary to first check the gap size and decide whether or not it is acceptable, as corrective orthodontic treatment may be required beforehand for an esthetically pleasing or functionally reasonable solution. The same applies for the restoration of multiple teeth in the esthetic zone. Often, this is only useful if the abutments are spaced at a distance that allows esthetically pleasing length-width ratios to be anticipated for each individual restoration and between adjacent anterior teeth.

3. For the crowning of teeth, it is necessary to have adequate abutment height and sufficient healthy tooth structure after tooth preparation. If this is not the case due to biological width considerations and a high degree of destruction, an orthodontic extrusion can be considered.

4. Following a crowning procedure, approximately 15 % of teeth show a negative response to sensitivity testing after 10 years [12]. A remaining dentin thickness of 0.5 mm should be aimed for in order to reduce the risk of pulp damage [15]. Therefore, it is important to remove as little healthy tooth structure as possible during preparation. Currently, preparation designs which preserve tooth substance have gained in importance. In this respect, the position and axial inclination of teeth play a decisive role. In clinical situations where the prospective abutment teeth require a common insertion path, but differ greatly in their axial inclinations, a conservative tooth preparation design is not attainable. In these cases, in order to avoid exceedingly invasive tooth preparations, orthodontically active components can be used in combination with interim dentures. By combining the temporary prosthetic phase with

Figure 1 Fixation of the lateral incisors – dimensioned and shaped according to the wax-up – for attaining a harmonious dental arch in the congenital absence of 12 and 22

Figure 2 Situation after removal of the brackets; width of the gaps optimally shaped

Figure 3 Placement of adhesive bridges to replace teeth 12 and 22

Figure 4 Two mesially interconnected adhesive bridges simultaneously adopt the function of a retainer.
orthodontic dental alignment, a lower amount of tooth substance removal can be facilitated.

First interface: missing teeth
Anterior region gaps are most frequently caused by the congenital absence of teeth, especially maxillary lateral incisors, or traumatic loss. Around one quarter of young people experience front tooth trauma by the age of 14 and about 7% of these front teeth are lost [23]. Circa 1.5–1.8% of people in Germany are affected by upper lateral incisor agenesis [17]. Hence, it can be assumed that approximately 3–3.5% of young people aged 14 years old are missing anterior teeth [7]. A number of treatment options can then be chosen. If the lateral incisors are congenitally missing, patients, or rather their parents, are often presented with the options of either an orthodontic gap closure or an implant-supported restoration. Orthodontists often favor gap closure by means of canine mesialization. Gap closure not only has functional implications, but also esthetic ones. This is because canines differ noticeably from lateral incisors, in both shape and color, as canines usually appear darker than incisors. Consequently, in the majority of cases, the treatment is not fully complete after orthodontic gap closure due to esthetic reasons; for a natural appearance, canines then require corrective treatment to imitate the appearance of lateral incisors. This implicates adapting the tooth's shape and color by means of applying a veneer, preparation of the enamel or whiteness of the enamel by bleaching [18, 20, 26]. Particularly in the case of unilateral agenesis, asymmetrical tooth positions may lead to esthetic impairments. For this reason, the symmetrical alignment of the front teeth should always be an important objective.

Implant-supported restorations are contraindicated in adolescence. Nevertheless, even after transversal bone growth and canine eruption have been completed, vertical bone growth may continue into adulthood.
Implants which are placed too early are often located in an infra position after a few years, which not only results in step formation at the incisal edges, but also in poor red-white esthetics in the area of the gingival line. Especially in the case of a high smile line, the timing for implant placement should be carefully considered. Adhesive bridges are often overlooked, or are solely considered a more or less temporary solution, until the age is reached when implants can be inserted. Lately, however, very good long-term results have been reported when using adhesive bridges. One-wing adhesive bridges have proven to be more reliable than two-wing bridges [2, 4, 10, 19, 24]. The clinical survival rate of one-wing adhesive bridges made of aluminum or zirconium oxide ceramics is between 95.4 and 98 % after 10 years [6, 9]. Thus, the survival rates of implant-supported crowns and one-wing adhesive bridges can therefore be regarded as being equivalent [11].

With reference to legally consistent patient education, the law governing the improvement of patients’ rights, anchored in the Federal Law Gazette 2013, specifies that the patient should be completely informed regarding all possible treatment options [1]. Accordingly, adhesive bridges must definitely be discussed as a treatment option. This necessity is also underlined by the fact that adhesive bridges are listed as a standard care procedure in the service catalogue of statutory health insurance companies since 2016. The key advantages of one-wing adhesive bridges are the minimally invasive preparation design which is only confined to the enamel and the fact that the restoration can be applied regardless of the stage of jaw bone development. The requirement is the presence of a completely erupted, caries-free neighboring tooth which borders the gap, having an adhesive surface in the enamel of at least 30 mm², as well as, sufficient space relative to the antagonist jaw for the purpose of being able to design a connector with a height of at least 3 mm [8]. Additionally, sufficient space should be generated for wing insertion in the case of maxillary adhesive bridges. For this purpose, the corresponding abutment tooth is protruded by approximately 0.7 mm using an aligner or multiband therapy. This creates sufficient space to attach the wing of the bridge without the need to perform any occlusal adjustments by grinding of antagonist teeth in the antagonist jaw. These necessary space conditions should be taken into consideration early on when planning an orthodontic treatment.

If only one incisor is missing, the contralateral tooth in the same jaw determines the size and shape of the intended restoration. A timely consultation with the prosthodontist is helpful here. Thus, a wax-up/set-up can be prepared at an early stage in order to model a suitable tooth. A try-in of the tooth wax-up can be made, which is thereafter converted into composite. The model tooth is fixed in the dental arch in order to give the orthodontist the chance to adjust the dental gap exactly to the desired size and position for the subsequent esthetic restoration. A similar procedure is useful if both lateral incisors are missing (Fig. 1). In this case, too, an ideal situation can be created through precise prosthetic planning.
before, or at the latest, during orthodontic treatment (Fig. 2). Above all, the ideal width of the lateral incisors must be carefully considered. When the width of the lateral incisors is between 50 and 74 % of that of the central incisors, they are perceived as being esthetic [25] (Fig. 3). However, if the gaps are set-up as being too small or too large, the dental restoration can either
• then be too narrowly or too widely designed,
• arranged in an overlapping position,
• be fitted in, but with a remaining gap
• or orthodontic measures must be carried out once again after the prosthetic treatment.

Directly after the removal of the multiband appliance, a retention splint must be worn; it is essential to ensure that the patient wears it continuously until the adhesive bridge is introduced. Otherwise, there is a risk that the teeth could move in the short time it takes for the adhesive bridge to be finished. Directly after the adhesive bridge is cemented, a retainer must be applied, as is common for all orthodontic treatments. The splitting together of the two adhesive wings can take over this function (Fig. 4).

Second interface: setting ideal tooth spaces before beginning with the restoration
Provided that several teeth in the esthetic zone must be restored with crowns, or veneers, it is always useful to prepare a wax-up in advance to check if the ideal length-width ratios and proportions of the teeth with respect to each other can be achieved. In the abrasive dentition shown here, conspicuous diastemas existed between all anterior teeth. Given the tooth positions, it was not possible to close the spaces and make sufficient vertical bite correction, and yet, still be able to create esthetically pleasing anterior esthetics (Fig. 5–7). For this purpose, anterior tooth lengths averaging 11.7 mm for the central incisors, 9.5 mm for the lateral incisors and 10.8 mm for the canines are desirable [13]. Additionally, a harmonious anterior overjet, a length-to-width ratio of the central incisors corresponding to 75–85 %, and a central to lateral incisor width ratio between 50 and 74 % should be present [25]. In order to achieve this, a gap closure by means of mesialization and a gap expansion in the premolar region took place. Based on this orthodontic pre-treatment, which lasted about 1 year, an esthetically pleasing and functionally appropriate prosthetic restoration could be created (Fig. 8–11).

Third interface: orthodontic extrusion for tooth preservation
The extrusion of roots often makes it possible to preserve and restore fractured or deeply destroyed teeth [3, 5]: The background behind this has to do with the planned/existing dental post in non-vital teeth having to be surrounded by at least 2 mm of healthy hard substance (adequate ferrule design). For vital abutment teeth, however, a remaining abutment height

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**Figure 18** Extrusion of approximately 4 mm after 4 weeks

**Figure 19** Initial situation, distally inclined canines

**Figures 20 and 21** Interim denture with attached elastic bands at the beginning of treatment; frontal and occlusal views.
of at least 3 mm is required. Furthermore, an adequate biological width must be maintained around the abutment teeth. In the existing case, the ceramic crown on tooth 21 was para gingivally fractured. The remaining abutment did not fulfill the conditions just described for a new crown which would have a favorable long-term prognosis. The neighboring teeth 11 and 22 displayed adequate restorations. Surgical crown lengthening was not an option due to esthetic reasons, as the previous gingival profile showed a harmonious appearance. In order to preserve the tooth, the root was extruded with the help of magnets. For this purpose, a magnet with flowable composite was attached to the vital abutment tooth. The magnet had to be reduced in size beforehand so that it matched the remaining abutment tooth (Fig. 12). Using a 1 mm thick placeholder, a second magnet was positioned on top of the first magnet and fixed to a previously fabricated Miniplast splint (Fig. 13). The supracrestal periodontal fibers were additionally cut with a scalpel under local anesthesia. In one to two weeks, the root was orthodontically extruded to the desired millimeter using magnetism. By moving the magnet twice inside the splint, with a renewed adjustment of 1 mm space between the magnets, the root could be extruded approximately 3 mm over a period of 6 weeks. The costs of this preliminary measure are not included in the service catalogue of the statutory health insurance companies, and must therefore, like any other orthodontic treatments in adulthood, be covered by the patients themselves. Subsequent to this procedure, a core build-up with composite, preparation of the abutment and fabrication of a temporary crown ensued (Fig. 14). In order to prevent the tooth from migrating again, the temporary crown was fixed to the adjacent teeth with retainer wire for 4 months. After the retention period, the definitive zirconia crown could be fabricated with vestibular veneering and adhesively bonded to the abutment. Figure 15 shows the restoration half a year after placement. The tooth continues to react positively to the cold sensitivity test.

Another method to mobilize teeth is through forced extrusion using rubber bands. For this purpose, if required, bars are fixed occlusally to the adjacent teeth [14, 16]. The abutment itself is fitted with a dumbbell-shaped device in the oro-vestibular direction. Elastic bands can then be hooked in with varying amounts of force (Fig. 16–18), so that the tooth root usually extrudes within a few weeks. The supracrestal separation of the gingival fibers prevents the fixed gingiva from moving with the root. When a coronal displacement of the soft tissue is also desired, the supracrestal fibers are not cut [16]. This applies to all extrusions, regardless of the procedure.

Fourth interface: conservative preparation by uprighting teeth before crown preparation

A well-known procedure and typical example includes the classical
uptightening of a molar that is inclined towards a mesial gap. We would like to present the case of an innovative, simple, and patient-oriented approach for aligning two canines. A 67-year-old patient with a remaining dentition consisting of 33, 43, and 44 in the lower jaw required a new prosthetic restoration. Owing to the inclination of the canines, the patient could hardly insert and remove the existing claspgretainable removable partial denture (Fig. 19). Besides this, the patient no longer wished for the new restorative treatment to have visible clasps. After a detailed consultation, he chose a telescopic denture. In order to have achieved a common insertion path during the preparation of the 3 abutment teeth, the canines should have been devitalized. A treatment alternative was represented by the uprighting of the two teeth in a mesial direction. The orthodontic appliance could be inte-grated into the already existing interim denture by applying orthodontic elements into its structure. Also, in this case, the orthodontic treatment had to be privately covered by the patient and an additional private agreement for the treatment and cost plan was created, which contained just the orthodontic measures. In order to use the time to test the new vertical bite position by means of the interim denture, a guide groove for the elastic band was prepared in cooperation with the department of orthodontics on the canines in the distal, lingual and vestibular areas. In addition, the denture teeth 31 and 41 each received a knob vestibularly to avoid protrusion or reinsertion. In this way, the patient was able to attach elastics on his own, which he changed daily (Fig. 20 and 21). After 3 months, the two canines were upright enough so that their preparation could be successfully accomplished, while maintaining a common insertion path and the vitality of the pulp (Figs. 22 and 23). The retention of the two teeth was ensured by the telescopic restoration. Figure 24 shows the patient after 3 years. The two canines still respond positively to sensitivity testing.

**Conclusion**

In summary, a purposeful combination of prosthetic and orthodontic treatment measures can achieve better esthetics and function in many cases. This is often accompanied by a reduction in treatment duration and costs. For optimal treatment outcomes, coordinated treatment planning between the orthodontist and prosthodontist is recommended before beginning treatment.

**Conflicts of Interest**
The authors declare that there is no conflict of interest as defined by the guidelines of the International Committee of Medical Journal Editors.

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Estimating the masking effect of post-orthodontic white spot lesions before resin infiltration

Richard J. Wierichs*, Celine Kobbe*, Hendrik Meyer-Lückel

**Introduction:** Orthodontic treatment with fixed elements increases the risk of developing white spot lesions due to additional retention opportunities for biofilm. One approach for the esthetic treatment of these lesions is caries infiltration. In order to estimate the final masking effect of the resin infiltration, re-wetting with water or ethanol is often performed just after the etching procedure.

**Patient cases:** Four patients who were diagnosed with white spot lesions in the esthetically visible area during treatment with fixed orthodontic appliances were informed about the possibility of optically masking these lesions using resin infiltration. The infiltration (Icon, DMG) was performed according to the manufacturer’s instructions, but the etching procedure was performed up to three times. The decision regarding whether to repeat the etching procedure was made subjectively by the practitioner during the re-wetting test with ethanol. The masking effect generated by ethanol and resin infiltration was analyzed using digital photographs before treatment, before and during the first 9 sec of re-wetting as well as one week after treatment.

**Result:** In all 4 patient cases, the observed minimum color difference during re-wetting appears to be a good indicator for assessing whether a satisfactory masking effect can be achieved. Furthermore, in the total of 55 treated white spot lesions, lesions with higher color value differences were etched more frequently than lesions with less pronounced color value differences. Overall, the masking effect one week after resin infiltration was satisfactory in all cases.

**Discussion:** During the re-wetting test, the minimum color difference between the initial lesion and the intact enamel which surrounds the lesion seems to be a good predictor of the final outcome of resin infiltration of post-orthodontic white spot lesions. Furthermore, white spot lesions with a higher initial color difference appear to require multiple etching procedures.

**Conclusion:** Resin infiltration is a useful microinvasive method for masking white spot lesions which develop during orthodontic treatment with fixed appliances. Moreover, the temporary masking effect achieved by the re-wetting test with ethanol can be used as an indicator for estimating the number of etching procedures required.

**Keywords:** caries infiltration; post-orthodontic treatment; enamel lesion; white spot lesion; esthetics
Introduction
Orthodontic treatment with fixed appliances increases the risk of developing initial non-cavitated carious lesions – also called white spot lesions [3]. The more difficult oral hygiene around the brackets represents an additional retention opportunity for biofilm and therefore increases the caries risk [2]. A rapid and progressive development of white spot lesions could then potentially become anesthetic concern for patients [15]. The characteristic white, opaque appearance of the lesions can be physically explained by the stronger scattering of light within the lesion’s body as a result of air and saliva inclusions in comparison to the surrounding healthy enamel [13]. In literature, the prevalence of white spot lesions after treatment with fixed elements varies between 23 %, 50 % and even 97 % [10].

White spot lesions remineralize once the brackets, which represent a retention site for plaque, have been removed. Although fluoride-containing agents can be used to enhance remineralization, the esthetic appearance is usually not sufficiently improved [19]. For this reason, different prevention strategies have been employed during treatment with fixed elements, e.g. fluoride-releasing sealants or bonding materials as well as the daily use of fluoride-containing or chlorhexidine-containing mouthwashes. However, these strategies cannot prevent the development of white spot lesions [21]. After removal of the fixed elements, microabrasion represents another treatment option which is most suitable for very superficial lesions because concave tooth surfaces can develop in the case of deeper lesions [22]. Direct and indirect restorations also lead to satisfactory and predictable results, but they should only be used in cavitated lesions due to their invasive nature [18].

Another therapeutic approach for the treatment of initial, non-cavitated carious lesions is caries infiltration. This involves the obturation of the microporous enamel areas of a caries by means of flowable resins (“infiltrants”) so as to arrest further caries progression [17]. Furthermore, the optical appearance is positively changed by caries infiltration [11]. Yet, it is difficult to predict the masking effect during the clinical application of the procedure. The varying masking results can be explained in relation to the variable thickness of the surface layer [8, 14]. In inactive lesions, the layer is usually thicker and less permeable than in newly formed active ones [1]. In contrast, the pores of active carious lesions appear larger and the surface layer less mineralized, thus allowing the resin to penetrate deeper into the lesion’s body [13]. However, to date, there is no way of estimating the ability of the infiltrant to diffuse into the remaining surface layer after etching. This would be desirable because it is difficult to remove the infiltrant after its application if the final result is considered not satisfactory. At the same time, a high degree of demineralization of the carious lesion is also unfavorable for attaining the most complete masking result possible.

The refractive indices of liquids vary closer to the refractive index of air. Therefore, the application of a liquid after the etching process should permit a (temporary) assessment of whether the surface layer has been sufficiently removed in order to allow a sufficiently deep diffusion [6]. Consequently, a liquid could theoretically be used to assess the masking effect before the infiltrant is applied. In this context, based on the four cases presented in this study, it was investigated if the final infiltration result is predictable using the re-wetting test.

Case studies
Four patients who complained about “white spots” on their teeth in the esthetically visible area after the removal of the fixed appliance were treated in a clinical study [10]. These areas were diagnosed as post-orthodontic white spot lesions. After the patient’s consent, the infiltration procedure was performed according to the following treatment scheme (Fig. 1):

- a. Cleaning the teeth with fluoride-free polishing paste (Cleanic; Kerr, Bioggio, Schweiz)
Estimating the masking effect during post-orthodontic white spot lesions before resin infiltration

WIERICH ET AL.: Estimating the masking effect of post-orthodontic white spot lesions before resin infiltration

Results

After removal of the brackets, 55 white spot lesions (ICDAS code 2, upper jaw: n = 29; lower jaw: n = 26) were diagnosed on the anterior teeth and first premolars in the 4 patients (14–16 years). The initial situation and the final treatment result are shown in Figure 2 and Figure 5. Satisfactory masking effects were achieved in all cases. This was independent of the number of etching procedures. The evaluation of the digital images after treatment showed that the number of etching procedures correlated with the initial color differences. The stronger the (remaining) color difference was estimated during the treatment, the more etching procedures were performed. Furthermore, it was shown that the subjectively perceived minimum color difference during the re-wetting test is well suited for estimating the color difference one week after infiltration. Teeth which did not show a satisfactory masking result during the first 3–5 sec of the re-wetting test were etched again (Fig. 4). Furthermore, the evaluation of the digital images confirmed that a difference in the practitioner’s color perception existed; it was significantly lower one week after infiltration as compared to during re-wetting (Fig. 5).

Discussion

Based on the presented cases, the color change during the re-wetting tests, their influence on the number of etching procedures and their correlation with the masking results one week after infiltration were described. It was observed that the minimum color difference during re-wetting seems to be suitable for estimating the color difference one week after infiltration. Especially during the first

Figure 2A-D After orthodontic treatment with fixed elements, multiple white spot lesions dominated the appearance of the upper and lower jaw (A). Appearance after the etching procedure (B). Estimation of the masking effect during the re-wetting test (C). A satisfactory final result was obtained one week after infiltration (D). The color difference one week after infiltration at the cervical margins of teeth 11 and 21 (D) is even lower than the minimum color difference during re-wetting (C). It should be noted that the partial image (C) only shows the re-wetting test on teeth 11 and 21. Before recording the final result (D), the cavitated lesion on tooth 23 was treated with a restoration.

- b. Application of a resin-based, light-curable barrier (OpalDam; Ultradent, South Jordan, USA) to keep the working area dry and protect the gingiva.
- c. Etching the teeth with 15% HCl gel – 2 min per lesion (ICON etch; DMG, Hamburg, Deutschland)
- d. Removing the HCl gel and spraying with water – 30 sec per tooth
- e. Air drying the teeth with compressed air – 30 sec per tooth
- f. Re-wetting the teeth with ethanol – 30 sec per tooth (ICON dry; DMG, Hamburg, Deutschland)
- g. Air drying the teeth with compressed air – 10 sec per tooth

During the re-wetting process, the practitioner (C.K.) decided if the masking result was satisfactory. If the subjective assessment was not satisfactory, steps c–g were repeated. The etching process was performed a maximum of three times. Afterwards, the treatment scheme was continued:
- h. Application of the infiltrant – 3 min exposure time (ICON Infiltrant; DMG, Hamburg, Deutschland)
- i. Removing the excess with a foam pellet
- j. Light-curing – 40 sec per tooth
- k. Application of the infiltrant – 1 min exposure time
- l. Removing excess with a foam pellet
- m. Light-curing – 40 sec per tooth
- n. A final polishing of the infiltration area was performed (SofLex; 3 M, Saint Paul, USA and Occlubrush; Kerr, Orange, USA).

In order to evaluate the masking effect, standardized digital photographs (shutter speed: 1/250, aperture: F29, ISO sensitivity: 100, white balance: 6250 K, tooth/lens distance: 20 cm) were taken at the following times: before treatment, 1 sec before ethanol application (treatment step f), in the following 9 sec (1 image per second) and one week after treatment (Fig. 2–4). Care was taken to avoid a color change due to the dehydration of the teeth.

Figure 3A-B Cervical white spot lesion on tooth 33. The gray chart on the images was used to standardize colorimetric evaluation. A: initial situation; B: situation one week after resin infiltration.
3 sec of re-wetting, there is a strong reduction in the color difference. If no considerable reduction in the color difference is subjectively detected, etching should be repeated. In any case, after resin infiltration the color difference is (even) further reduced when compared to the color difference during re-wetting. This might be explained by the higher refractive index of the infiltrant (RI = 1.51), which is closer to the refractive index of healthy enamel (RI = 1.63) than to the refractive index of ethanol (RI = 1.36). Furthermore, it could be observed that the number of etching procedures correlated with the initial color difference. The data showed a positive albeit weak correlation between the number of etching procedures and the subjective evaluation of the severity of the white spot lesion [10].

In order to achieve an esthetically satisfactory masking result for white spot lesions, it is necessary to infiltrate the body of the lesion as completely as possible [13]. For this purpose, the surface layer of the lesion has to be modified during the etching procedure so that the resin or ethanol can diffuse into the lesion. However, the thickness of this surface layer varies greatly [14]. This may help explain why the number of etching procedures, and thereby the length of the etching procedure varied, not only in these patient cases, but also in other studies [7, 8]. Depending on the aesthetic appearance, the number of etching procedures and the duration of etching varied both in the previous studies and in the present patient cases. However, the maximum number of etching procedures was limited to three in order to not remove too much enamel. This was also done if the esthetic result of the re-wetting test was not satisfactory after the third etching procedure. Since each etching procedure removes between 35 μm and 45 μm of the enamel surface [14], 3 etching procedures lead to a maximum removal of 105–135 μm. This probably results in the complete removal of the surface layer in most active and also inactive lesions. A further etching procedure would therefore remove an unnecessary amount of enamel because, after the removal of the surface layer, the success of masking is likely to depend on the depth, or degree of mineralization of the lesion, instead of the remaining less porous surface layer.

The cases presented here were treated within the framework of a study which had been approved by the Ethics Committee of the RWTH Aachen University (EK 110/13) and which had already been published [10]. In this study, a total of 221 lesions in 29 patients were infiltrated and resin infiltration noticeably reduced the color difference of white spot lesions after treatment both subjectively and objectively. The final color differences were only slightly above the perceivable thresh-
These color differences are not perceived by the human eye from a normal social distance. Furthermore, a subgroup analysis showed that the color difference was below this threshold for 73% of lesions that were etched just once, whereas lesions that were etched two or three times were below the threshold in only 62% and 32% of the cases, respectively. The final result, for lesions etched two or three times, was therefore less satisfactory than for lesions etched once. Furthermore, a strong correlation between the minimum color difference during re-wetting and the final color value one week after infiltration could be observed. It was also demonstrated that the minimum color difference during re-wetting is also well suited to assess if another etching step is necessary. The results are also consistent with previous studies [7, 8], which also showed a good masking effect using this treatment procedure for post-orthodontic white spot lesions. Further clinical studies which investigated the masking effect after resin infiltration and their results are also shown in Table 1.

Interestingly, the studies also suggest that the time interval between bracket removal and resin infiltration appears to play an important role for the successful masking of white spot lesions [8]. The shorter this time interval, the more successful the masking effect seems to be. Consequently, patients who had undergone orthodontic treatment more than 12 months ago were excluded in the present study. Furthermore, the question as to how the masking effect could be further optimized by infiltrating the white spot lesions during orthodontic treatment was raised. Thus far, however, this question has only been investigated in a non-controlled study [16] and a case report [12]. In both publications white spot lesions were infiltrated immediately after detection without interrupting orthodontic treatment. Although the results in both publications were promising, this approach has not yet been widely explored.

**Table 1** Clinical studies investigating the masking success of white spot lesions, mostly after orthodontic treatment.

<table>
<thead>
<tr>
<th>Authors</th>
<th>Year of publication</th>
<th>Time interval until follow-up examination</th>
<th>Number of patients treated/number of infiltrated lesions</th>
<th>Treated teeth</th>
<th>Results</th>
</tr>
</thead>
<tbody>
<tr>
<td>Kim et al. [7]</td>
<td>2011</td>
<td>1 week</td>
<td>9 patients, 18 teeth</td>
<td>Anterior teeth</td>
<td>61% complete, 33% partial, 6% no masking effect</td>
</tr>
<tr>
<td>Hammad et al. [4]</td>
<td>2012</td>
<td>Immediately</td>
<td>18 patients, 108 teeth</td>
<td>Maxillary anterior teeth (ICDAS 2+3)</td>
<td>Significant increase of gray scale value (= masking)</td>
</tr>
<tr>
<td>Knosel et al. [8]</td>
<td>2013</td>
<td>6 months</td>
<td>20 patients, 111 infiltrated and 108 control teeth</td>
<td>Anterior teeth and canines</td>
<td>ΔΔE: Infiltration -2,3 Controls +0,2 Significant improvement</td>
</tr>
<tr>
<td>Knösel et al.⁶ [9]</td>
<td>2019</td>
<td>2 years</td>
<td>8 patients, 45 teeth</td>
<td>Anterior teeth and canines</td>
<td>ΔΔE: Infiltration -3,2</td>
</tr>
<tr>
<td>Senestraro et al. [20]</td>
<td>2013</td>
<td>8 weeks</td>
<td>20 patients, 46 infiltrated and 20 control teeth</td>
<td>Maxillary anterior teeth</td>
<td>Reduction of lesion’s surface 61% Infiltration 1% Controls</td>
</tr>
</tbody>
</table>

⁶Post evaluation of Knösel et al. 2013

**Conclusion**

The patient cases illustrate that the temporary masking effect during the re-wetting test with ethanol, before subsequent resin infiltration, seems to be a good predictor of the final masking result and can be used to determine the necessary number of etching procedures. Furthermore, white spot lesions with a higher initial color difference seem to require more etching procedures. Overall, resin infiltration has been shown to be a useful method for masking white spot lesions occurring during orthodontic treatment.

**Note from the authors**

The cases presented in this article have been treated in a clinical study published in the Journal of Dentistry titled „Evaluation of the value of re-wetting prior to resin infiltration of post-orthodontic caries lesions“ [10].

**Conflicts of interest**

R. Wierichs and C. Kobbe declare that there is no conflict of interest as defined by the guidelines of the International Committee of Medical Journal Editors.
Estimating the masking effect of post-orthodontic white spot lesions before resin infiltration

The Charité – Universitätsmedizin Berlin (Germany) holds US and European patent for an infiltration technique for dental caries lesion in which one of the authors (Hendrik Meyer-Lückel, HML) is appointed as inventor. HML receives royalties as well as research funding from DMG (Hamburg, Germany), the manufacturer of Icon.

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Orthodontics for adults – current innovations and interdisciplinary possibilities

Introduction: At this point in time, treatment of adult patients is part of the everyday lives of orthodontists. There is no difference when determining the indication for therapy in children and adolescents: the goal is the prevention and treatment of misalignments of jaws and teeth, as well as maintaining a healthy masticatory system.

Discussion: Nevertheless, there are some characteristics that need to be taken into consideration: The biomechanics should be adjusted in accordance to the progressing loss of attachment. Simultaneously, there is a great desire for preferably subtle treatment options. Furthermore, interdisciplinary treatment concepts play a more important role in adults compared to children and adolescents.

Results: The digitalization in orthodontics resulted in new and innovative therapy options for all these requirements. Generated data records using intraoral scans offer various options in diagnosis as well as planning in order to improve treatments and patient comfort. Special orthodontic software solutions were necessary to establish individualized therapy concepts. Treatment appliances with different systems can be printed using the export of STL data and made available for orthodontists. Further advances are expected in the next few years, which makes digital orthodontics an exciting work field.

Keywords: digital orthodontics; orthodontics for adults; 3D-print; CAD/CAM restorations; interdisciplinary orthodontics
Introduction
Orthodontics is no longer limited to the treatment of children and adolescents. At this point, the treatment of adult patients is part of the clinical everyday life of this field of dentistry. In the years 2012 to 2014, every fourth patient undergoing orthodontic treatment in the US and Canada was over 18 years old [1]; the tendency is increasing. There are a number of reasons that explain this tendency:

An increased awareness for oral health exists in today’s society and tooth position is also especially important for adult patients. It is socially established that straight and white teeth are considered healthy and attractive, whereas dysgnathia can lead to negative reactions in the social environments [18].

The optimized dental supply situation in the last several decades and the demographic change leads to the inevitable increase in average age in the orthodontic practice: an aging patient collective has a higher demand to optimize the position of their own natural teeth. However, adults do not just turn to orthodontists for aesthetic reasons: From a prophylactic periodontal point of view, it is advisable for older patients to correct misaligned teeth in order to prevent changes in bone structure caused by unequal load distribution.

Even with interdisciplinary questions such as pre-prosthetic distribution or pre-surgical formation of dental arches, the dental disciplines should work hand in hand in order to achieve the best possible results for the patients.

Orthodontic treatment can be subtly designed by using established treatment methods such as lingual therapy or aligner therapy, which is desired by patients that reject a conventional therapy due to the temporary unaesthetic impairment. With the advances in digital orthodontics, new exciting approaches for individualized treatment planning and execution are available [6, 10, 11, 23].

Special requirements in orthodontics for adults
The orthodontic treatment of adult patients has underlying special features. Generally, as with any patient, a periodontal screening should be done before orthodontic treatment and the patient should be referred to a dentist if there is any demand for treatment. This is indispensable, because otherwise the orthodontic therapy contributes to a progression of periodontal destruction on inflamed periodontium [20].

Usually, there is a thicker compact bone, increased attachment loss and less vascularisation of the bone with increasing age, which can to some extent be accompanied with restriction of tooth movement. Additionally, adults more often take medications that can affect the bone metabolism and therefore also impact the orthodontic therapy (Tab. 1).

Following the attachment loss, the center of resistance of the teeth is displaced more apically, and the orthodontic forces have to be adjusted. Further anchoring measures should be taken into consideration, because periodontally weakened teeth cannot compensate for the opposing forces in orthodontic therapy anymore. Mini implants can provide remedy, and will be discussed further down in this article.

The adult patients’ desire
Adult patients seek out orthodontists especially with the desire to carry out a subtle treatment in a foreseeable time period. The lingual multi bracket therapy offers patients a very popular alternative to vestibular braces. Tooth movements can be carried out efficiently in all 3 spatial planes using a completely individualized apparatus (Fig. 1–3).

By establishing aligner therapy, orthodontists were given another tool to reach the desired treatment result step-by-step in a gentle manner. Nevertheless, it shall be noted that a selection of suitable patients as well as the dentists’ experience with the system are imperative for successful treatment using aligners.

Current innovations in orthodontics
In order to meet the desire of adult patients, digital technologies are useful to carry out accurate and individualized planning. Digital orthodontics consists of 3 components:
1. Scan system (intraoral scanner or model scanner)
2. CAD software
3. 3D-print

Scan systems
The basis for a digital workflow is always a three-dimensional data set, usually generated by an intraoral scan. If no intraoral scanner is available, a model scanner can always be used as well. Different scan systems have been established for a few years in orthodontics and contribute to drive out the conventional impressions further and further. The precision of powder-free intraoral scans complies with that of the classic impression [13], however, it offers significantly greater patient comfort.

CAD software
This generated data set can then be imported into a suitable CAD-soft-
ware that takes care of further processing (Fig. 4). The all-digital procedure of the diagnosis, the planning and the subsequent implementation of the therapy is sketched out below and the advantages of digitalization are shown.

In our clinic we use the software OnyxCeph™ (image instruments). The workflow in digital orthodontic diagnosis includes digital photography, digital x-rays and the digital impressions mentioned above.

A full evaluation of patient documents can be done based on the generated data and then has to be supplemented with a clinical examination. The intraoral and extraoral images as well as the cephalometric x-rays can be measured and analyzed on a computer. Also the three-dimensional model can be evaluated via software (Fig. 5). Studies show that the accuracy of digital models is equal to the conventional evaluation of plaster models and sooner or later will depict the new gold standard in orthodontic diagnosis [17].

When diagnosis is completed and a type of therapy has been chosen, it can be planned and designed digitally. Lingual therapy and aligner therapy belong to the subtle treatment methods that are preferred by adult patients as described above. These usually assume a set-up which simulates the desired final position of the teeth. In the past, the well known and complex manual steps using a saw-cut model was necessary.

This is now digitally possible in a substantially precise and effective approach (Fig. 6). Teeth can be individually segmented and subsequently moved freely in the spatial planes according to the desired target position using the appropriate software.

In lingual therapy, individualized arches can be calculated using the set-up produced specifically for the patient. These can be manually bent using a printed drill template. Alternatively, the arch geometry in the CSV format can be transmitted to a bending robot, which leads to a reduction in time and increase in precision. The therapeutic transversal expansion of dental arches depicts a risk in stabilization of orthodontic

Figure 4 Segmented digital model after intraoral scan (3Shape TRIOS 3)

Figure 5 Orthodontic analysis of a digital model using the orthodontic software OnyxCeph™

Figure 6 Creation of a digital set-up using the orthodontic software OnyxCeph™
treatments [22]. First initial data show that a more stabilized tooth position can be achieved following individualized therapy [Wolf et al., submitted].

Clinical expertise is indispensable for digital planning of aligner therapy in order to assess limits of movements and plan helpful tools such as attachments correctly. Furthermore, in many cases an additional anchoring is required. Should this anchoring demand be maximal, a skeletal anchoring in the form of mini implants (TAD; temporary anchorage device) can be chosen. Because of this, tooth movements can be executed that usually can only be carried out with severe side effects. Examples of this are mesial and distal movements of molars, intrusions of individual elongated teeth, mass retractions and distractions of individually impacted teeth [3].

Ideally, the insertion of such a TAD is planned digitally. Depending on the intended application, the mini implant must be inserted in another region. Anatomic structures in the immediate topographic surroundings such as roots of teeth, inferior alveolar nerve and palatal artery have to be spared.

In order to guarantee this, a drilling template can be used. These can be manufactured conventionally in the lab or using CAD/CAM procedure. For this, an x-ray is superimposition of intraoral scans of the upper jaw with a cephalometric x-ray

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<table>
<thead>
<tr>
<th>Substance</th>
<th>Appearance/Application</th>
<th>Effects</th>
<th>Impact on bone and tooth movement</th>
</tr>
</thead>
<tbody>
<tr>
<td>NSAID (Aspirin, Ibuprofen)</td>
<td>Analgesic, anti-inflammatory</td>
<td>Bone resorption</td>
<td>Tooth movement</td>
</tr>
<tr>
<td>Paracetamol</td>
<td>analgesic</td>
<td>No effects</td>
<td>No impact</td>
</tr>
<tr>
<td>Bisphosphonates</td>
<td>Osteoporosis, cancer treatment</td>
<td>Bone resorption</td>
<td>Tooth movement</td>
</tr>
<tr>
<td>Insulin, Metformin</td>
<td>Diabetes mellitus</td>
<td>Bone stabilization</td>
<td>Tooth movement</td>
</tr>
<tr>
<td>Vitamin D</td>
<td>Pregnancy, breastfeeding period, depressions</td>
<td>Bone resorption</td>
<td>Tooth movement</td>
</tr>
<tr>
<td>Eicosanoids (prostaglandins, leukotrienes etc.)</td>
<td>mediators</td>
<td>Bone resorption</td>
<td>Tooth movement</td>
</tr>
<tr>
<td>Fluoride</td>
<td>Osteoporosis, caries prophylaxis</td>
<td>Bone resorption, bone mass, bone density</td>
<td>Tooth movement</td>
</tr>
</tbody>
</table>

Table 1 Medications and their effect on the bone and tooth movement
(Fig. 1-16, Tab. 1: L. Brämswig)
imposed with the intraoral scan (Fig. 7). For palatal mini implants a cephalometric x-ray is usually used. If a TAD is inserted interradicular, a CBCT (cone beam CT) is required for the superimposition process. With this, an accurate positioning of the mini implants can be planned digitally under consideration of the immediate anatomic structures. As soon as the desired position and angulation of the mini implant is determined, a separate model is generated that defines the direction of insertion. It can then be used to design a drilling template in the desired extent (Fig. 8–10).

Using a CAD/CAM-drilling template guarantees better control during insertion and reduces the risk of deviation of the planned path of insertion, especially with inexperienced practitioners [2].

Difficult anatomic structures are another indication for a CBCT while planning mini implants. In our own investigations we were able to show significant superimposition of mucosa of the anterior palate in patients with unilateral cleft lip and palate. The best region of insertion on the affected half of the jaw is the transition of premaxilla and maxilla [14] (Fig. 11 and 12).

A combination of mini pins and aligners has been proven to be clinically practical and therefore broadens the indication spectrum of aligner treatments in selected cases [21].

Digital tools are available for the traditional vestibular bracket treatment. The correct positioning of the brackets at the beginning of the treatment have a significant influence on the orthodontic treatment success. The conventional tooth banding of the patient is usually done adhesively on the patient directly, where every bracket is placed on each tooth individually, positioned, and then cured. Alternatively, transfer trays can be manufactured to transfer into the patient’s mouth. This requires a previous laboratory process, however, it offers advantages for patient and practitioner by reducing chair time. Studies could also show that indirect adhesion guarantees a more precise bracket positioning in the vertical plane [12].

Meanwhile, a transfer tray such as this can also be planned and designed digitally. The basis here is also a segmented model via software that simulates the roots of teeth. Based on the axis of the tooth, brackets can be positioned correctly virtually. On top of the correct position of the bracket, a positioning template can be designed, saved as an STL data set and submitted to the printer (Fig. 13). The brackets can be placed in the CAD-CAM drilling template and transferred indirectly into the patient’s mouth. Initial investigations from our clinic show that a good transfer of the digitally planned bracket position can be achieved.

The planning and design of individualized metallic devices is another option to design treatment devices using a suitable software offer. A patient-specific adjustment of treatment devices was already possible before, however, the usage of pre-fabricated goods in anatomic conditions that vary widely from the norm is always associated with restrictions.

Digital methods also show efficient solution approaches, so that, ideally, delicate devices can be ad-
justed individually and subsequently manufactured (Fig. 14 and 15). Besides the advantage of the highly precise fit, it can also lead to shortened appointments because the separation for conventional bands is not necessary in digitally produced devices.

**3D-print**

All previously depicted procedures of therapy planning and design of treatment devices via a CAD software lead to a 3D print. Sprockets for aligners, drill templates for mini implants or bonding trays for bracket transfers can be saved as STL data sets and submitted to a printer. Depending on the indication there are various procedures available, and in dentistry especially the SLA (stereolithography), DLP (Digital Light Processing) and the FDM print (Fused Deposition Modeling) have become established. Resin-printers can usually impress with shorter printing times, whereas the post processing in form of alcoholisation and post-curing of filament printers is omitted. Orthodontic questions can be achieved with any of the systems [5, 9], even though further advances can be expected in the next few years. Another form of the 3D print is the SLM Printing (selective laser melting), which can be used to manufacture metallic devices. The acquisition of a printer like that is currently extremely cost-intensive for the single practice, but an external printer on larger labs is already provided. In an additive production process the material to process in powder form is melted layer by layer. A cobalt chrome alloy is usually used in orthodontic devices.

**Interdisciplinary orthodontics**

Besides digital innovations in orthodontics mentioned above, there are also benefits offered in the interdisciplinary exchange. Dental colleagues can simultaneously consult on corresponding questions in their workspace using a digital model without having to refer to the plaster model.

Thus, operation splints were designed and subsequently printed using a 3D-data set in a CAD/CAM procedure. Studies could show that an equal or higher precision in production was achieved in less time compared to conventional splints [7, 8, 24, 25].

Furthermore, a VTO (visual treatment objective) can be created in patients treated by orthognathic surgery in order to set up a pre-therapeutic prognosis for the soft tissue profile after corrective osteotomy. This tool can be used for demonstration and patient communication, however, it should be implemented carefully to not incite unrealistic expectations [15]. Such a VTO can be generated either using two-dimensional or three-dimensional data sets, where the latter produces more precise prognoses [19].

In addition, in another working field between surgical dentists and orthodontists, there are new and exciting approaches to improve care of patients. In patients with agenesis or premature tooth loss, orthodontic space closure is not always the first choice. Prosthetic restorations, which definitely yield great results, are often needed to completely close such spaces. In selected cases, the autogenous transplantation of a tooth into the aplastic region offers a great predictable therapy option [16]. If a three-dimensional x-ray is present, that tooth that is to be transplanted can be deducted and printed pre-operatively using a DICOM data set of the CBCT. The generated template is then tried on in the region that needs to be treated during the surgery. This has the advantage, that the bone can be prepared optimally, which is accompanied with little bone loss. Also, the extra-alveolar time of the tooth as well as the damage of PDL cells is minimized during the operation, which leads to an improved prognosis of the treatment [4].

There are also new paths of treatment seen in other dental treatments. If a dentist wishes to treat
subgingival defects of a tooth by extruding that tooth, a strong anchoring should be chosen in patients that are not banded in order to avoid unwanted side effects on neighboring teeth. Through the SLM printing, the rigid and also delicate apparatus as described above can be designed with the desired tooth movements without loss of anchoring (Fig. 16). After successful extrusion the dentist can proceed with the treatment of the tooth.

Conclusions
Using digital treatment methods, various new approaches are given in orthodontics that facilitate and improve therapy. Moreover, new ranges of treatment appeared that could be used in interdisciplinary questions and therapy planning and therefore contribute to treatment success. Thanks to the advances in 3D-printing technology it will lead to exciting innovations in the future.

Conflicts of interest
The authors declares that there is no conflict of interest within the meaning of the guidelines of the International Committee of Medical Journal Editors.

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Philipp Winterhalder, Nassim Ayoub, Stephan Möhlhenrich, Ali Modabber, Frank Hölzle

Advances and limits in planning and implementing orthognathic surgery

Introduction: Orthognathic surgery has steadily developed since its establishment in 1849, and is characterized by consistent collaboration with orthodontics.

Materials and Methods: Computer-based operation planning, that also takes resulting changes in soft tissue into account, can now be carried out through three-dimensional X-ray imaging, face scans and powerful data processing. This procedure can also be used to produce individualized osteosynthesis material and osteotomy templates, which in combination even enable orthognathic surgery without the use of an occlusal wafer. Piezo surgery represents a minimally invasive alternative to conventional methods for the osteotomy. As an alternative to titanium, resorbable polymers can also be used as osteosynthesis material.

Conclusion: Due to modern orthodontics and computer-based operation planning, the surgery-first concept enables orthognathic surgery without prior orthodontic treatment for eligible cases. The effects of orthognathic surgery on the upper airway must be considered during planning. In addition to treating dysgnathia, orthognathic surgery expands the upper airway and has become established as a treatment option for obstructive sleep apnea.

Keywords: computer-based surgery planning; prediction of soft tissue changes; individualized osteosynthesis material; resorbable osteosynthesis material; surgery first; piezosurgical osteotomies; effects on the upper airways
Introduction
Orthognathic surgery has been characterized by continuous advancement since its origin. In order to classify the current advancements and limitations, it is worth taking a brief look at the history of orthognathic surgery [38]. The first orthognathic surgery was performed by the American surgeon Hullihen in 1849. He normalized the mandibular length of a female patient with burn-related deformities with anterior segmental osteotomy [13]. Among others, two principle pioneers of early orthognathic surgery are considered to be Angle and Blair, who together described standardized mandibular osteotomies in 1907 as a treatment for mandibular prognathism [5, 38]. Blair also began classifying types of dysgnathia and realized the importance of orthodontics [38]. Many other surgeons described various different procedures for mandible correction, however, due to high rate of recurrence and significant risks they were unable to implement these at the time [38].

This was followed by the development of osteotomies that are used today (Fig. 1): Wassmund displaced the maxilla with an anterior segmental osteotomy in 1935 to treat maxillary protrusion [43]. In 1955, Trauner performed the reversed L-shaped osteotomy of the mandibular ramus to treat mandibular prognathism with shortened mandibular ramus [41]. Schuchardt described the posterior maxillary osteotomy in the same year, as well as the diagonal, sagittal osteotomy of the mandibular ramus to increase bone apposition [33]. Köle performed osteotomies of the alveolar process in both jaws in 1959, as well as genioplasty in 1968 [20, 21]. In 1955, Obwegeser developed the revolutionary sagittal split osteotomy of the mandibular ramus [30], which was modified by Dal Pont in 1958 [9] and von Hunsuck in 1968 [14]. In 1969, Obwegeser performed the protrusion of the maxilla with a Le Fort-I osteotomy and combined this with a mandible correction in 1970, making this the first corrective osteotomy of both jaws [28, 29]. In addition to the development of the effective osteotomies, the postoperative stability of the jaws is mainly due to the advancement of osteosynthesis procedures. A working party for osteosynthesis founded in 1958 in Switzerland is still at the forefront of this research today.

In addition to surgical aspects, orthodontic treatment and interdisciplinary treatment planning is absolutely crucial for a successful therapy of dysgnathia. Major progress in orthodontics was described in 1984, specifically the possibility of using mini screws as means of skeletal anchoring [8]. When compared to its beginnings, modern day orthodontics is more gentle, faster and can even supersede lengthy preoperative tooth movement in certain cases.

The basis of interdisciplinary treatment planning is still individualized cephalometry, which was introduced in 1974 by Hasund and further developed with Segner [10, 35]. Additionally, detailed model analyses and clinical investigations were also carried out. Treatment plans tend to mostly structure in preoperative measures, for example orthodontic pretreatment, followed by orthognathic surgery to displace one or both jaws, and orthodontic follow-up treatment. Close collaboration between orthodontics and maxillary surgery is necessary during all treatment stages. After completing pre-treatment, a model operation is performed to plan the orthognathic surgery, where the planned displacement can be implemented from previous analysis. Besides the reliable model operation using plaster models in articulators, digital operation planning is gaining more importance. Current progress in orthognathic surgery is characterized by digitalization, new materials, minimal invasiveness and an individually optimized treatment result. In the following, computer-based operation planning, prediction of soft tissue changes, individualized and absorbable osteosynthesis material, the surgery-first concept, piezosurgical orthognathic surgery and the effects on the upper airways are explained.

Computer-based operation planning
The analog operation planning using plaster models, face bow, articulators and wax bites has proven itself reliable over decades. Computer-based operation planning has become established as an alternative in research and patient care due to the increasing availability and advancement of digital technologies.

Apart from the analog and digital approach, the exact recording of dental, skeletal, functional and aesthetic difficulties is an absolute necessity for
successful planning. In addition to orthodontic and maxillofacial expertise, individualized cephalometry, model analysis and the clinical investigation are the basis of the treatment planning.

The operation planning should not be viewed as an isolated part, but rather as part of the treatment plan. Similar to analog operation planning, the digital method leads to success when the practitioner has the ability to gather and understand the underlying individual dental, skeletal, aesthetic and functional parameters and their deviation. Recent findings of cephalometric fundamental principles, such as the harmony box, enable the optimal use of benefits of 3-dimensional, computer-based dysgnathia-planning.

The individualized cephalometry sets the basis for a successful treatment, despite new analysis and planning methods. The holistic diagnosis to differentiate between dental and skeletal causes is made possible with individualized cephalometry. Furthermore, this enables the analysis of facial proportions as well as the assessment of mutual dependencies in sagittal and vertical direction. The individualized cephalometry helps to adjust the Curve of Spee and Wilson, correctly plan the axial position of the front teeth in jaw rotations and estimate the autorotatory effects when changing the vertical relation. However, software programs for digital operation planning offer reduced determination of cephalometric parameters, or parameters that are not comparable with classic parameters. Program designers should offer a complete cephalometric analysis. After completing comprehensive diagnostics, it is not only possible to create operation splints, but also to create analyses and predictions for soft tissue. Especially in the case of asymmetries digital operation planning is helpful because mirror images can be made for optimal adjustment. Movements of the skeletal bases can only be assessed roughly, which is why cephalometric analysis of required movements is helpful at the beginning of treatment planning. The patient should be informed about measurement and prediction errors of the used equipment and programs during the analysis and prediction of soft tissue changes, in order to avoid unrealizable, specific expectations and even legal concerns. The computer-based operation planning combined 3-dimensional x-ray data from a DVT or CT with a digital impression of the dental arches. Additionally, a face scan was done to analyze the soft tissue. These data sets are used to create a complete model which simulates the operation using a planning software.

After successful virtual displacement of the jaws the operation splints are created digitally and 3D-printed and after the osteotomy, the new jaw positions can be defined using teeth impressions (Fig. 2, 3 and 4). During planning, the jaw movements can be positioned in the 1/100 mm-range, which can lead to successful treatment of complex cases, such as vertical height correction [15]. The benefits of computer-based operation planning regarding the quality of results include the consideration of the individual temporomandibular joint axis and the expected change of the soft tissue. The planning time period can be shortened in computer-based operation planning compared to analog planning. For the planning of a sole mandibular displacement, the planning time period of 195 minutes can be reduced by 41 % and when displacing both jaws, the planning time period of 385 minutes can be reduced by 62 % [31].

Shortening of the planning period described in literature compared to analog operation planning is questionable when considering the additional survey of fundamental dysgnathia parameters and creating splints in 3D print. In the alternative definition of the necessary movements and submission of implementation to a service provider or software program, the planning know-
how and understanding of specific surgical circumstances would be lost. A shortened planning period would therefore reduce the practitioners influence and produce additional costs. Other disadvantages of computer-based operation planning are high investment costs and the necessary staff training in the new workflow.

**Prediction of soft tissue changes**

Displacement of the jaws also causes displacement of the soft tissue. While the displacement of the jaws can be planned and implemented precisely, the resulting displacement of the soft tissue is more difficult to predict. The reason for this is that the soft tissues do not follow the displacement in a 1:1 ratio but rather in a different ratio depending on their anatomic region. Even with the knowledge of these specific displacement factors, the individual patients’ reaction complicates a prediction and does not allow a precise statement concerning displacement of soft tissues. Because soft tissues play an important role in the aesthetic and functional result, it is rewarding to consider the most optimal estimated displacement during planning and evaluating. The mean values of displacement factors were determined for anatomic regions in many studies. Lateral x-rays and photographs were used as underlying two-dimensional data, whereas new studies are mostly based on three-dimensional data from CT, DVT and face scans. Displacement factors were described as follows, whereas a displacement factor smaller than 1 means a smaller displacement than the according bone reference point: 0.78 in the upper lip region, 0.77 in the lower lip region, 0.74 on the tip of the nose 0.70 at the base of the nose and 0.73 on the cheeks [23] as well as 0.94 on the chin [6].

Due to the postoperative swelling, an assessment of the actual soft tissue displacement is advisable after a few weeks. An investigation of prediction accuracy of 3 planning programs showed a mean error of 1.8 mm +/- 0.8 mm for 3D (Dolphin Imaging & Management Solutions, Chatsworth, CA, USA), 1.2 mm +/- 0.4 mm for ProPlan CMF (Dentsply-Sirona, York, PA, USA) and 1.3 mm +/- 0.4 mm for the prediction based on a Finite-Elemente-Simulation [18]. However, the measurement of the soft tissues already has a rather high error, which is why a mean error of up +/- 2 mm is needed to collect 90 % of the measurement points in a face scan [17].

**Individualized osteosynthesis material**

New manufacturing technologies have made it possible to create patient-specific osteosynthesis material and osteotomy templates. The basis for this is computer-based operation planning. Osteosynthesis material can therefore be directly created based on the virtual shaping via laser sintering. A less complex alternative is to print synthetic 3D models of the bone using the virtual situation and to individualize ready-made osteosynthesis materials by bending it into the correct position. With computer-based operation planning, osteotomy templates from synthetic materials can be 3D-printed and used to implement osteotomies. These osteotomy templates can additionally help keep the safety distance to the dental roots or nerves. Using patient-specific osteosynthesis materials and osteotomy templates is supposed to make exact implementation of planned jaw displacement possible. This is an advantage in the vertical dimension, because the implementation of pure splint-based methods is surgically demanding. In combination with fitting laser sintering osteosynthesis materials, the screw holes of the osteotomy templates can be used after the osteotomy for osteosynthesis material. Thus, the position is transferred accurately and an operation splint is not absolutely essential [39]. However, templates and individually manufactured osteosynthesis materials, especially when sintered when splinted, are inapplicable with minimally-invasive surgical access because of the larger measurements. Another possibility for surgery without splints is using intraoperative navigation [4]. There are optical and electromagnetic systems available that are not widely applied due to high equipment costs despite good results. The disadvantage of an operation without splints is that the occlusion is unstable right after the displacement, because the orthodontic follow-up treatment still has to take place. Therefore, a splint is useful in many cases to ensure postoperative results during the healing phase, even when it is not essential for intraoperative positioning.

A significant disadvantage of patient-specific osteosynthesis material and osteotomy templates is the increased planning effort. Specifically, the laser sintering of patient-specific osteosynthesis material for orthognathic surgery may be the equivalent of the entire case compensation and has not yet established itself as a standard process. With further developments of manufacturing technologies and more frequent implementation, cost reduction and a wider application of these innovative processes is anticipated in the future.

**Resorbable osteosynthesis material**

Back in 1932 an absorbable osteosynthesis material with pure magnesium was available, which did not stand the test of time due to its biochemical instability. Absorbable osteosynthesis material took a back seat, because the introduction of stainless steel in the 1940s and titanium in the 1950s has been successful until today [34]. However, persisting osteosynthesis material can lead to foreign body sensation, bacterial colonization and growth limitation. Titanium causes artefacts in x-ray or MRI imaging, which complicates diagnosis. Even with new accidents or other necessary surgeries, persisting osteosynthesis materials can be a source of complications. The osteosynthesis material is often removed after bone healing is completed, however, this second procedure also comes with all the risks of surgery and is more complicated due to the existing scars. The biochemical industry has developed absorbable polymers since the 1960s and absorbable osteosynthesis material since the 1990s, among other things based on lactic acid, glycolic acid and polydioxanone. Depending on the composition, the degradation period takes weeks to years, whereas the
degradation is mostly based on hydrolysis of ester bonds [34]. In order to partially compensate reduced stability when compared to titanium, absorbable osteosynthesis material is designed larger and requires more invasive surgical access. In clinical studies, the osteosynthesis material made out of resorbable polymer in orthognathic surgeries in the maxilla or mandible is sufficiently stable and causes similar complication rates like osteosynthesis material made out of titanium [1, 16, 32].

Resorbable osteosynthesis made out of polymers has the additional special feature that it is not radiopaque. This offers the benefit that no artefacts are created in imaging, however, it also prevents position and integrity control. Another option are new magnesium alloys with zirconium and strontium, which are significantly more stable compared to polymers, but need to be improved concerning biocompatibility [34].

Surgery first

When developing dysgnathia, the position of the teeth usually changes to compensate the underlying skeletal cause. For example, in mandibular retrognathia the incisors are often severely protruded. In such a compensated situation a normalization of the skeletal base by jaw displacement would cause a malocclusion. Therefore, in the conventional treatment concept the tooth positions are normalized with regard to the alveolar ridge and the dental arch. This causes the dysgnathia-based malocclusion to become fully visible. The dental arches that now match each other are optimally aligned when planning the surgery and relocated accordingly during the operation.

In contrast to this, a treatment concept called surgery first has been used since 2009, where no or minimal orthodontic treatment occurs before the orthognathic surgery [7]. The best and definite occlusion is aspired after surgery, with orthodontic treatment. The procedure was made possible with advances in orthodontic treatment, often using skeletal anchoring elements such as mini screws. Benefits of the surgery-first concept include shortening the treatment period by several months as well as reduced strain on the patient by skipping the pretreatment phase [36]. The surgery-first concept is used in class-II and class-III dysgnathia patients for displacement of one or both jaws. The treatment planning is demanding and often involves computer-supported operation planning. There is no consensus of indications of the surgery-first concept. Patients with mandibular prognathism, less teeth crowding and little dental compensation are particularly suitable [7, 22]. Patients with a mandibular retrognathia, deep bite, narrow palate or pronounced axis deviation of the front teeth seem unsuitable for the surgery-first concept [7, 22]. Treatment using the surgery-first concept should be critically questioned with increasing asymmetry of the patient case [11], even though successful treatments of pronounced asymmetry have also been described [42]. The treatment procedures of the surgery-first concept are continually being developed, which can lead to extended indications. For example, the treatment of protrusion of lower incisors as a surgery-first concept was described as a piezosurgical subapical osteotomy [12]. In order to facilitate the extensive, postoperative orthodontic treatment in a surgery-first concept, intraoperative skeletal anchoring elements such as mini-plates or mini-screws were inserted if needed. Postoperative, combined with increased bone remodeling [36], tooth movements can occur within a few months. Orthodontists and surgeons must individually discuss every patient to decide if the surgery-first concept depicts a promising therapy option. Even though all dislocations in digital operation planning are done very precisely, the spatial positioning of the segments provides a challenge for the user. Besides surgical aspects, orthodontic expertise is indispensable to determine positions of the jaws and to assess the postoperative orthodontic movements. Based on the absent preoperative decompensation the postoperative occlusion is only minimally supported in the surgery-first concept. Unfavorable occlusion contacts can even lead to postoperative mandibular malpositions [7]. Postoperative stabilizing with an operation splint is therefore recommended especially during multiple segmentations of the maxilla. The splint can be strengthened additionally for intraoperative application.
with an incorporated transpalatal bow.

Piezosurgical osteotomies

The basis of piezography is the observation by the brothers Pierre and Jacques Curie in 1880, that a crystal converts electrical energy into micromovements, which is considered the piezoelectric effect (Greek, piezein, ‘to push’). The clinical use has developed since the 1950s from isolated applications in dentoalveolar surgery to the modern piezography gadgets used today, where the first one in Germany was approved in 2002 [40].

In piezography, micro movements of the piezo crystal are transferred to the tissue with different serrated or grained top pieces. The vibrations amount to 60–210 µm with a frequency of 25–29 kHz [2]. Hereby, only hard tissue is removed and soft tissue such as nerves are spared. Piezography can be used to displace the maxilla in the Le Fort-I level as well as the displacement of the mandible in sagittal segmental osteotomy. A significant reduction of operation time is possible by piezosurgical osteotomies [19], while in addition the saw cuts can be designed more delicately. A review with 799 patients showed no prolonged duration of the piezosurgical segmental osteotomy of the mandible [37]. The frequency of nerve impairment at least 6 months after the surgery with 4.7 % of the patients was significantly less compared to the 61.6 % when a saw was used. In another study, the impact of piezography on the fracture pattern of sagittal segmental osteotomy was examined, whereby no difference to conventional osteotomy was found in comparison [27]. The selection of different osteotomy top pieces makes piezography versatile, and a curved top piece was used to weaken the base of the lower jaw. However, similarly to conventional weakening, this showed a poorer fracture pattern.

Effects on upper airways

Repositioning of the jaws with the adjacent soft tissues inevitably affects the upper respiratory tract. A measurement of the upper airways and their transformations is possible using three-dimensional imaging with DVT or CT and has presented superior to measurements based on lateral radiographs [25]. In case of impaired nasal breathing the airway resistance in this region can be determined with rhinomanometry. A palate expansion causes an expansion of the main nasal cavity and a reduced airway resistance during nasal breathing [3].

A mandibular retraction reduces the airways and bears the risk of respiratory problems in patients with pronounced class-III malocclusions. This is particularly true, when the tongue is large compared to the jaw. As an alternative to the sole mandibular retraction in patients with class-III malocclusion, an additional maxillary protrusion should be considered. Even this approach causes changes in the airways, which was described in a study with 22 patients [24]. The volume of the nasopharynx changed from 5.4 cm³ preoperative to 5.2 cm³ postoperative. The volume of the oropharynx reduced from 17.8 cm³ to 11.9 cm³, and the hypopharynx from 7.2 cm³ to 4.6 cm³. Despite the significant changes no patient presented respiratory problems 6 months postoperative.

The effects of jaw corrections on the airways are not only of importance in orthognathic patients. In obstructive sleep apnea syndrome reduced muscle tone causes blocked upper airways due to the weight of the soft tissues. The patient wakes up repeatedly because of respiratory distress, often without even realizing. Although there are usually only a few symptoms besides fatigue, it can lead to serious health issues such as arterial hypertension, heart disease or diabetes mellitus. In therapy-resistant obstructive sleep apnea the protrusion of both jaws is an important treatment option [26]. In order to displace the mandibular soft tissues even more anteriorly, an additional genioplasty is often performed.

Conflicts of interest

The authors declares that there is no conflict of interest within the meaning of the guidelines of the International Committee of Medical Journal Editors.

Literatur

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**Overview:** The update of the guideline on “diagnostics and management of precursor lesions of oral squamous cell carcinoma in dental and oral medicine” began in 2017 and was finalized in April 2020 after a total of 3 formal consensus processes. It was coordinated by the German Society of Dental and Oral Medicine (DGZMK) and the German Society for Oral and Maxillofacial Surgery. Specifically, the guideline updates the knowledge and recommendations, particularly the following aspects:

- the classification of potentially malignant oral lesions considering the updated WHO classification of 2017
- special status of proliferative verrucous leukoplakia
- definition of “suspicious” lesions under observation of clinical evidence of a malignant transformation
- specific designation of examinations, whose significance is not supported with reliable study data
- topical corticoid therapy of lichen, especially intralesional therapies

Furthermore, the existing recommendations were updated and complemented by statements and new recommendations.

**Keywords:** precursor lesion; dental and oral medicine; malignant transformation; early detection; screening; oral mucosa; WHO classification
Introduction

It’s the guidelines’ aim to record the current state of knowledge for a relevant problem in health care and if possible, draft key statements in the form of clear recommendations for action. For this, regular updates are necessary in order to keep up with the development of scientific knowledge. Having said this, the update of the guideline “diagnostics and management of precursor lesions of oral squamous cell carcinoma in dental and oral medicine” was started in 2017 and finalized in April 2020. It was carried out by the German Society for Dental and Oral Medicine (DGZMK) and the German Society for Oral and Maxillofacial Surgery. The update of the guideline was added to the list of prioritized topics of the task force of DGZMK, BZÄK and KZBV, which consisted of representatives of DGZMK, KZBV and BZÄK.

The authors of the guideline conducted current literature research (Medline) until 2018 to draft the recommendations and background text and included the relevant literature in the guidelines. Based on the existing guidelines, the coordinators revised the document in the first step and added current literature. Simultaneously, certain wordings were clarified that had led to misunderstandings in the past, and a formal division between recommendations and statements was included. The referencing of other guidelines was also updated.

This draft was the basis of a formal Delphi method with two Delphi rounds and a conclusive consensus conference with the participation of elected representatives of professional societies on 23.01.2019 in Cologne (see Tab. 1) under methodical moderation of AWMF. Within this consensus conference, the key statements and additions in the context of literature were discussed and a formal and structured consent added to the methodology of a nominal group process.

Because this is an S2k guideline, a more detailed evaluation of studies in the sense of an evidence grading or even weighting and synthesis of study results did not take place.

Updated precisely, the guideline specifies the current knowledge and recommendations, particularly the following aspects:

- classifications of potentially malignant oral lesions regarding the updates WHO classifications of 2017
- special status of proliferative verrucous leukoplakia
- definition of “suspicious” lesions under observation of clinical evidence of a malignant transformation
- specific designation of examinations, whose significance is not supported with reliable study data
- topical corticoid therapy of lichen ruber mucosa, especially intraleisional therapy
- recommendations

The state of research and the decision criteria of recommendations were renewed in form of background texts as in the previous version, which were included in the extended version of the guideline. Since these texts create references to relevant literature, they are displayed here for information.

Classifications of potentially malignant oral lesions in consideration of the upgraded WHO classification of 2017

In the current WHO classification of head and neck tumors 2017 [7], mostly classification of dysplasia degrees are used. However, the term squamous intraepithelial neoplasia (abbreviated: SIN) will continue to be used as a synonym of potentially malignant lesions (previously: oral precursor lesions) of squamous cell carcinoma. The potentially neoplastic character of the lesion is depicted in

<table>
<thead>
<tr>
<th>WHO 2017: Dysplasia</th>
<th>WHO 2005: Dysplasia</th>
<th>Ljubljana classification squamous intraepithelial lesions (SIL)</th>
<th>Squamous intraepithelial neoplasia (SIN)</th>
<th>Reduced squamous intraepithelial neoplasia (SIN)</th>
</tr>
</thead>
<tbody>
<tr>
<td>low grade dysplasia</td>
<td>low grade dysplasia</td>
<td>basal and parabasal hyperplasia</td>
<td>SIN I</td>
<td>SIN: low risk</td>
</tr>
<tr>
<td>moderate dysplasia</td>
<td>moderate dysplasia</td>
<td>atypical hyperplasia (risk epithelium)</td>
<td>SIN II</td>
<td>SIN: high risk</td>
</tr>
<tr>
<td>high grade dysplasia</td>
<td>high grade dysplasia</td>
<td>carcinoma in situ</td>
<td>SIN III</td>
<td>SIN: high risk</td>
</tr>
<tr>
<td>carcinoma in situ</td>
<td>carcinoma in situ</td>
<td>carcinoma in situ</td>
<td></td>
<td></td>
</tr>
<tr>
<td>invasive carcinoma</td>
<td>invasive carcinoma</td>
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</tr>
</tbody>
</table>

*Since both moderate dysplasia and high grade dysplasia are considered “high risk” lesions, the graduation can be modified to a binary structure “low grade” and “high grade” (with “high grade” including moderate and high grade dysplasia)
the nomenclature. In the following, the term of potentially malignant oral lesions classification (2017) instead of precursor lesions and other terms is used according to WHO (precancerous condition, precancerous lesion, potentially malignant lesion, precursor lesion etc.).

Compared to earlier versions of the WHO classification, a reduction of degrees of dysplasia is introduced in the form of a binary classification. Ultimately, the three traditional degrees of dysplasia are reduced to a “low grade” group and a “high grade” group, which essentially corresponds to the clinical risk assessment and signals a distribution into “low risk” SIN and “high risk” SIN. In earlier nomenclature, high grade intraepithelial neoplasia (SIN 3) included the carcinoma in situ of earlier classifications (carcinoma risk of 90 %) [3, 5, 15, 27], (see overview 1).

Special status of proliferative verrucous leukoplakia
The proliferative verrucous leukoplakia (PVL) has a special status, because the morphological degree of dysplasia does not correlate with the clinical risk potential. Even though low degrees of dysplasia are typically found in PVL or can be missing completely, a highly malignant transformation rate (about 70 %) and consequently high tumor-related mortality is expected with this entity.

Definition of “suspicious” lesions while describing clinical features of a malignant transformation
The recommendations for action are based on the fact that oral lesions that are evaluated as “suspicious” for malignant transformation. Apart from the chronological development (persistence of a lesion for more than 2 weeks) the following clinical criteria should be regarded as indicators suggestive of a malignant transformation, specifically:
- newly occurred and of unknown duration
- thick hyperkeratosis
- inhomogeneity
- erosion
- bleeding on contact or light mechanical stress
- missing cause
- pathological vascular dilation/vessels

Examinations, whose significance is not supported with reliable study data:
The description of research methodology was upgraded and newly structured. Specifically, a group of examinations is designated whose significance is not supported with reliable study data.
Necessary examinations for therapy decisions:
• inspection: using a systematic examination procedure it is ensured that all relevant regions of the oral mucosa can be investigated and critically assessed
• palpation

Further investigations:
• review of causes of mechanical irritation
• sensitivity test of neighboring teeth to record odontogenic inflammatory causes
• determining periodontal parameters to record periodontal causes

Helpful examinations in justified individual cases:
• swab for microbiological diagnostics
• virological diagnostics
• reviewing a reaction to dental materials

Examinations, whose significance is not supported with reliable study data:
• x-ray examination to record dental and osseous inflammatory causes
• examination of lymph node status to evaluate accompanying inflammatory reactions or to recognize the spread of the tumor
• sensitivity test (lingual nerve and mental nerve) to evaluate sensation of pain or to recognize the spread of the tumor

Table 1 Listing of the professional societies involved/organizations and the elected representatives

<table>
<thead>
<tr>
<th>Professional societies involved/ Organizations</th>
<th>Abbreviation</th>
<th>Elected representatives</th>
</tr>
</thead>
<tbody>
<tr>
<td>Working Group for Oral and Maxillofacial Surgery</td>
<td>AGOKI</td>
<td>Prof. Dr. A. M. Schmidt-Westhausen</td>
</tr>
<tr>
<td>Working Group Oral Pathology and Oral Medicine</td>
<td>AKOPOM</td>
<td>Prof. Dr. Dr. U. Müller-Richter</td>
</tr>
<tr>
<td>Federal Association of German Oral Surgeons</td>
<td>BDO</td>
<td>Prof. Dr. J. Jackowski Prof. Dr. T.M. Remmerbach</td>
</tr>
<tr>
<td>Federal Association of Dentists</td>
<td>BZAK</td>
<td>Dr. J. Beck</td>
</tr>
<tr>
<td>Professional Association of German Pathologists</td>
<td>BDP</td>
<td>§</td>
</tr>
<tr>
<td>German Dermatological Society</td>
<td>DDG</td>
<td>Prof. Dr. F. Kiesewetter</td>
</tr>
<tr>
<td>German Society for Otorhinolaryngology, Head and Neck Surgery</td>
<td>DGHNO-KHC</td>
<td>Prof. Dr. J.P. Klußmann Prof. Dr. C. Wittekindt (deputy)</td>
</tr>
<tr>
<td>German Society of Implantology</td>
<td>DGI</td>
<td>Prof. Dr. F. Schwarz</td>
</tr>
<tr>
<td>German Society for oral and maxillofacial surgery</td>
<td>DGMKG</td>
<td>Prof. Dr. K. Hertrampf (coordination) Prof. Dr. Dr. M. Kunkel (mandate)</td>
</tr>
<tr>
<td>German Society of Peridontology</td>
<td>DGPARO</td>
<td>PD Dr. C. Graetz</td>
</tr>
<tr>
<td>German Society of Pathology</td>
<td>DGP</td>
<td>§</td>
</tr>
<tr>
<td>German Society for Dental Prosthetics and Materials Science</td>
<td>DGPro</td>
<td>Prof. Dr. H.-J. Wenz, MME</td>
</tr>
<tr>
<td>German Cancer Society (working group ENT and OMF-surgery in oncology)</td>
<td>DKG</td>
<td>Prof. Dr. J.P. Klußmann</td>
</tr>
<tr>
<td>National Association of Statutory Health Insurance Dentists</td>
<td>KZBV</td>
<td>Dr. J. Beck</td>
</tr>
</tbody>
</table>

§) The elected representative of the German Society of Pathology (DGP) and the Professional Association of German Pathologists (BDP) left the guideline group during the creation process. Both specialist groups were given the opportunity to comment on the finalized draft guideline and both approved.

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were able to show a high sensitivity of oral mucosa, due to the loss of fluorescence. In autofluorescence diagnostics were examined the application of toluidine blue in order to detect oral mucosa lesions that have not already been previously recognized clinically. In addition, more authors are critical of the low accuracy for potentially malignant oral lesions with a sensitivity of under 50 % [9, 16, 18].

Overall, the big effort involved with chairside application has prevented the clinical implementation of staining and the expansion of the methods and usage in dental practices. In literature, there is no data on the application in primary care. After more than five decades, a relevant development and expansion of the method is not to be expected. b) Chemiluminescence and autofluorescence diagnostics

In the last few years, the procedures for chemiluminescence and autofluorescence diagnostics were examined as supportive visual tools in detecting potentially malignant oral lesions and oral carcinomas in different studies. In the process of the chemiluminescence source of light, the oral mucosa is pretreated with 1 % acetic acid. Possible modifications in the keratinization are supposed to show up in white and then stand out after subsequent treatment with toluidine blue. In autofluorescence diagnostics, dysplastic or neoplastic lesions are supposed to show up darker compared to healthy (green) oral mucosa, due to the loss of fluorescence. Koch et al. (2010) were able to show a high sensitivity of 93 % in a patient collective (N = 78) with conspicuous clinical mucosa lesions in the examination using autofluorescence diagnosis, however, the specificity was only at 13–17 % [11]. In the study of Mehrota et al. (2010) both visual methods came into effect and showed significantly worse results [17]. The procedure with autofluorescence showed a sensitivity of 50 % and a specificity of 38.9 % in 156 examined lesions. When applying chemiluminescence in 102 examined lesions, the sensitivity was at 0 and the specificity at 75.5 %. Further studies that investigated these procedures showed similar critical, unsatisfactory results [1, 2, 8, 21, 22]. The inhomogeneous and inadequate data shows no scientific basis for the application of either visual procedure in early detection of potential malignant oral lesions and oral carcinomas.

c) Narrow band imaging

Another visual method that was evaluated in studies for early detection of oral carcinomas and potentially malignant oral lesions in the last few years is the narrow band imaging from other areas of surface diagnostics in the oral cavity in studies. The method used two narrow-banded frequency domains (400–430 nm and 525–555 nm) in order to depict differences in vessel plexus instead of continuous frequency spectrum of white light. Yang et al. (2012 and 2013) showed a sensitivity of 96.3 % for the narrow band imaging using a patient collective of n = 317 and a specificity of 60.1 % compared to white light with 87 % sensitivity and 93.5 % specificity. However, the transferability of the results and a potential recommendation is only possible to a limited extent, because in studies from Asian countries, many lesions are buccal lesions, caused by the enjoyment of betel products and therefore many localisations for lesions were underrepresented [28, 29]. A systematic review from 2014 [26] came to the conclusion that this method has diagnostic potential, however, a statement for recommendation in the field of early detection is not possible due to the insufficient and inadequate data.

Topical corticoid therapy of mucosal lichen ruber, particularly intralesional therapy

Also the background text on measures of demarcation of inflammatory/irritating phenomena was extended and specifically, the intralesional therapy with corticoid was included.

Local corticoid therapy

For a symptomatic lichen ruber the local treatment using steroids is the therapy of choice [13]. There is not enough evidence for a recommendation of a specific steroid therapy regarding the outcome of “pain reduction” [24]. This was also confirmed by the study of Liu et al. (2013) [14]. They could, however, determine a positive therapy effect of an intraleisional therapy with betamethasone compared to therapy with triamcinolone in their randomized, controlled study regarding the outcome “recurrence of a lesion within three months”.

Unresponsiveness to steroid application confirms the indication of a biopsy [23].

Recommendations

Since the key statements of the guideline are formulated in the recommendations, all recommendations of the guideline are written out in the following. An explanation of modifications was included when significant changes have been made to the previous version.

**Recommendation 1:**

Within the recommended systematic examination of the oral cavity every 6 months it should be ensured that all regions of the oral mucosa, the lips and the neighboring tissues are critically investigated. When changes are observed, further diagnostics should take place.

Expert consensus
Recently incorporated recommendation

**Recommendation 2:** When the cause of mucosal changes is assumed, e.g. a mechanical irritation of inflammatory systemic disease, the cause should be eliminated first, and if necessary this includes treating the systemic disease.  
*Expert consensus*

**Recommendation 6:** If a biopsy might not be representative of the whole lesion, a complete diagnostic excision should take place.  
*Strong consensus*

**Recommendation 3:** In case of suspecting a manifested carcinoma, the patient should be referred immediately to introduce further diagnostics and therapy.  
*Expert consensus*

**Recommendation 7:** If a mucosal lesion is considered non-malignant and there is no need for an biopsy, nonetheless monitoring is intended since some uncertainty remains regarding the dignity of oral mucosa lesions, a brush cytology should be used.  
*Consensus*

**Recommendation 4:** When suspecting a malignant transformation of the mucous membrane, a histological clarification should take place.  
*Expert consensus*

**Recommendation 8:** In extended oral mucosa lesions, where a diagnostic excision would lead to a highly perioperative morbidity, an extensive brush biopsy is an alternative to multiple simultaneous biopsies.  
*Strong consensus*

**Recommendation 5:** For a cytological diagnosis, the harvesting procedure should be done using brushes, because they can collect superficial as well as deeper cell layers.  
*Expert consensus*

**Recommendation 9:** For extensive oral mucosal lesions with chronic progression (for example with a proliferative verrucous leukoplakia) there is the problem that on one hand, representative localisations (e.g. most advanced in tumor progression) can sometimes not be defined, and on the other hand, a complete diagnostic excision especially in cross-regional lesions can imply the perioperative morbidity of a tumor resection or is not technically feasible in multifocal lesion. In these rare cases, the large surface coverage of the brush biopsy (reduction of “sampling errors”) must be weighted against the greater diagnostic accuracy of the excision biopsy (limited to the excised tissue).  
*Strong consensus*

**Recommendation 10:** A histological clarification (biopsy) should take place**, if there is a beginning regression in the first two weeks, but not a complete healing after two more weeks.  
*Expert consensus*

**Recommendation 11:** According to prevailing opinion, lesions that are clinically homogeneous, and evaluated as histologically “low grade” (previously SIN I or low dysplastic) can be primarily monitored.

Lesions that are histologically classified as “high grade” (previously SIN II or III, or moderately or highly dysplastic) should be excised completely.  
*Consensus*

*/*The timeframe of recommendations 9 and 10 apply to patients where normal wound healing can be expected.

**Statement 1:** There is not enough evidence for a recommendation regarding application of further technology in cytology (immunohistology, DNA cytometry etc.) in early recognition of oral precursor lesions.  
*Expert consensus*
S2k guideline “Diagnostics and management of precursor lesions of oral squamous cell carcinoma in dental and oral medicine”

Recommendation 12:
When there is a discrepancy between the clinical appearance and the histological evaluation (for example, inhomogeneous leukoplakia without histological dysplasia), another histological review or a transfer for a second opinion/introduction of further diagnostics and therapy should follow.

Expert consensus

Recommendation 13:
After removal or monitoring of low grade dysplastic lesions an inspection interval of 6 months should be followed. In all other manifestations of dysplastic lesions, a check-up interval of 3 months should be followed.

Specific recommendations exist for mucosal lichen ruber for the necessity of constant monitoring. The check-up interval should not exceed four months.

Expert consensus

Recommendation 14:
A check-up should always be recommended to the patient, independent from the type of therapy.

Expert consensus

Recommendation 15:
In general, an outpatient treatment under local anesthesia is sufficient. A treatment in general anesthesia/sedation can be indicated depending on localisation or due to expected problems in cooperation of the patient (e.g. gag reflex), in patients with large overall extent of mucocutineal measures, in manifested local risk factors or after consideration of these criteria based on the preference of the patient.

In-patient treatment can be indicated in severe systemic diseases or particular surgery developments.

Expert consensus

There is additional information available in the form of a more detailed guideline report. The documents can be downloaded from the websites of the AWMF, BZÄK and the DGZMK. The next update is planned for 2025.

Conflict of interest
The authors state that there is no conflict of interest within the guidelines of the International Committee of Medical Journal Editors.

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Dental implants are an integral part of modern dental, oral and maxillofacial medicine. Apart from rehabilitating the physiological masticatory function and increasing the quality of life, dental implants have a positive influence on the general health condition [31]. For successful treatment, a key prerequisite is the osseous healing of the inserted implants (osseointegration). This strongly depends on an adequately functioning immune system, as it is involved in regulating postoperative wound healing and it has an important modulating effect on the tendency of subsequent peri-implantitis development. In this respect, immunocompromised patients exhibit a particularly higher risk profile.

Given the current demographic development of higher life expectancy and continuously improving health care, it has become apparent that the number of patients displaying immunodeficiency, who wish to be treated with implant-supported restorations, is increasing. Furthermore, due to the ongoing improvement in long-term immunosuppressive therapies, a significant increase in this category of patients can be anticipated in the future [1, 13, 17].

Moreover, in deciding when dental implants are indicated in immunocompromised patients, it must be considered that the patient population is very heterogeneous with varying degrees of immune system impairment. For the dental practice, this means a high degree of uncertainty with respect to the indication for treatment, as well as the associated treatment sequence, which includes preparation, conducting the operative intervention and postoperative aftercare.

For a better overview, the classification of immunodeficiency can be grouped as follows:

1. Primary, congenital immunodeficiency*
2. Secondary, acquired immunodeficiency, e.g. HIV infection**
3. Secondary, drug-induced immunosuppression, e.g. steroids (cortisone)

**Autoimmune diseases**

Studies performed in the last 10 years corroborate that there is a steady increase in the frequency of autoimmune diseases [17]; at present, their prevalence in Europe and North America is presumed to be as high as 12.5 % [13, 17]. Accordingly, there is reason to suspect that the number of patients with autoimmune diseases, who wish for dental implants, will grow. The mechanisms which underlie the reduction of the body’s immunological tolerance to one’s own body molecules, and subsequently, to a lowered immune response and to different forms of autoimmune diseases, is not yet fully understood [28]. Socioeconomic, genetic and environmental factors as well as certain types of infections are discussed as being triggering factors for autoimmune reactions.

Autoimmune diseases occur more frequently in women. They represent the predominant group among patients suffering from autoimmune diseases (75 %) [13]. In a systematic review, the influence of autoimmune diseases and their therapy on the survival rates of dental implants was investigated. The outcome revealed a clear trend towards female patients who accounted for 98 % of the patient population. The frequent occurrence of different coexisting autoimmune diseases was also conspicuous; examples include the combination of rheumatoid arthritis and another autoimmune disease.
Sjögren’s syndrome or dermatomyositis as well as oral lichen planus and Sjögren’s syndrome.

With regard to the modalities for treating autoimmune diseases such as rheumatoid arthritis, polymyalgia rheumatica, pemphigus vulgaris, scleroderma, Sjögren’s syndrome and systemic lupus erythematosus, mainly steroid medication such as prednisone or other glucocorticoid derivatives are used for therapy. Neither the effect of the drug nor the underlying autoimmune disease was found to affect implant survival [8].

In principle, due to the possible risk of malignant transformation of the oral manifestation of the underlying diseases, stringent implant aftercare should be respected. There is evidence to suggest that the very rare peri-implant carcinomas occur with particular frequency in patients with oral lichen planus [22].

**Special role of Crohn’s disease**

The chronic inflammatory bowel disease, Crohn’s disease, predominantly affects the gastrointestinal tract, although a direct influence on the oral mucosa can also occur. The autoimmune inflammatory reactions are triggered by antigen-antibody complexes and this is why immunosuppressive and anti-inflammatory drugs are part of the treatment spectrum.

Pertaining to the therapy of patients with dental implants and Crohn’s disease, a correlation was observed between implant loss and Crohn’s disease in studies investigating early implant failure [2, 3, 30]. These results were statistically supported by further studies performed in 2007 and 2008. The basis for the cumulative incidence of early implant loss in Crohn’s disease patients is controversially debated in the scientific community. Thus far, it has been shown that the osseointegration of dental implants can be influenced by antigen-antibody complexes due to autoimmune reactions in the area of bone-implant contact [26]. Moreover, malnourishment, which frequently occurs in the course of Crohn’s disease, may also lead to inadequate bone healing around dental implants [9].

**HIV**

In the last 30 years, the HIV infection and the subsequent Acquired Immunodeficiency Syndrome (AIDS) has developed from being an epidemic with a devastating deterioration of patient’s health, to being a stable, chronic disease thanks to current therapeutic management. Consequently, there is an increasing number of patients at different stages of the disease who wish for implant-based dental rehabilitation.

Compared to healthy patients, both HIV-seropositive patients with a CD4 cell count > 200 cells/μl and severely immunocompromised patients with a CD4 count of less than 200 cells/μl showed no significant differences after implant insertion with regard to the healing reaction, infection rate or postoperative complications. In addition, higher rates of implant loss were not observed [6, 16, 18]. Consequently, there was no evidence of a direct relationship between the risk of postoperative infections after implant treatment and the CD4 count [10, 20, 29].

In 75 % of the studies which were analyzed, different forms of anti-biotic therapy were used for implant surgery. The prophylactic administration of antibiotics had no influence on the risk of postoperative infections, but it was shown to reduce the risk of implant failure [5, 15]. In summary, in a systematic review, antibiotic therapy was identified as one of the key factors which influences the analysis of dental implant osseointegration in HIV-positive patients [4].

**Chemotherapy**

Chemotherapy still constitutes one of the main pillars of modern cancer treatment. To date, the number of approved antineoplastic drugs is continuously growing due to the diversification of drugs which target specific types of cancer. Therefore, it is difficult to thoroughly investigate the mechanisms by which chemotherapeutic agents exert their biological effects on dental implants. For this reason, few studies have explored the potential effect of chemotherapeutic agents on osseointegration in spite of the fact that it is one of the most important parameters for successful implant therapy.

For already existing implants, there is evidence to suggest that chemotherapy can have a variety of negative effects such as mucositis, painful peri-implant infections as well as systemic effects such as fever and sepsis. However, these reported side effects have been observed primarily in blade and subperiosteal implants, which are hardly used nowadays [14]. Due to the fact that chemotherapy is associated with serious underlying diseases, there is a need for rigorous risk stratification involving an interdisciplinary approach; thus, close cooperation with the oncologist in charge of therapy is recommended.

**Immunosuppression after organ transplantation**

The number of organ transplant recipients is increasing in tandem with medical progress. As a general rule, oral and maxillofacial infectious foci should be surgically operated prior to organ transplantation in order to reduce the rate of postoperative infection. After successful transplantation, patients who have had multiple tooth extractions before transplantation often require the functional rehabilitation of their masticatory functions [24, 27]. It is not uncommon for dental practitioners and oral and maxillofacial surgeons who are performing the treatment to be addressed with the patient’s expressed wish for implant-supported prosthetic restorations.

In numerous studies, no demonstrable effect of the various post-transplantation protocols and their respective immunosuppression regimes, which use steroid and immunomodulating drugs (tacrolimus, sirolimus, cyclosporine and mycophenolate), could be shown on the implant survival rate.

The lack of randomized controlled trials limits the ability to draw conclusions. Nevertheless, no constraints for treatment with dental implants could be identified based on the results of the abovementioned studies.

Implant surgery should only be performed after consultation with the transplant physician in charge.
especially with regard to prophylactic/prolonged antibiotic therapy.

The practice-oriented treatment recommendations and checklist for risk stratification for indicating and managing dental implant treatment in immunocompromised patients is based on the current S3-guideline „Dental implants in patients with immunodeficiency“.

**Practice-oriented treatment recommendations**

1. **Indications**
The medical status of the patient at the time of indication has a significant influence on the success rate of implant treatment. Conversely, it must also be ensured that the planned implant therapy does not endanger the patient’s health [12]. Thus, before indicating implant treatment, the individual risk for implant loss and complications should first be assessed in all patients displaying an autoimmune disease, immunodeficiency, or who are under immunosuppression.

   Primarily, an acute status of the underlying disease as well as any local or systemic contraindications should first be excluded. Further treatment planning should ideally include interdisciplinary cooperation (internists, rheumatologists, or other specialist disciplines).

2. **Preoperative pre-treatment and diagnostics**
In order to reduce the risk of infection and eliminate existing foci of infection, any necessary surgical interventions should be performed before implant surgery. Wound healing can provide an initial insight into the function of the immune system.

   In addition to radiological diagnostics, the reported clinical findings should be included for the purpose of risk assessment, as they may provide possible clues regarding compromised soft tissue healing, bone remodeling or rates of bone regeneration. The prosthetic evaluation (prognosis of the remaining dentition, benefits of abutment augmentation or mucosal load reduction) is performed in the same manner as for healthy patients.

3. **Implant prognosis**
   Literature data indicates that there are no relevant differences between patients with and without immunosuppression based on a follow-up period of at least 24 months. An exception to this are patients suffering from Crohn’s disease.

4. **Necessity of augmentation**
   Jaw bone augmentation is accompanied by increased demands on the bony recipient tissue. In cases of immunosuppression or immunodeficiency, it can be assumed that there is a less than adequate systemic immune response of the recipient tissue during wound healing [11, 21, 23].

5. **Informing the patient**
   As part of routine patient education before planning any type of implant-based treatment, immunocompromised patients should be informed about the individual risks of disease-related complications (e.g. poorer implant prognosis in patients with Crohn’s disease) and implant loss. Additionally, patients should be well-informed about the importance of follow-up care, which is adapted and structured according to their individual risk, and any potential follow-up costs.

6. **Perioperative management**
   The low complication rates/implant loss risks observed in the studies were all attained from patients undergoing perioperative systemic antibiotic prophylaxis.

7. **Implant insertion**
   Submucosal and transmucosal healing are both possible. A recommendation regarding which type of healing is preferable cannot be derived from literature. Since the rates of bone remodeling and bone regeneration are reduced under immunosuppressive therapy, immediate or early loading must be critically appraised; this is also true for immediate implant insertion.

8. **Prosthetic treatment**
   With regard to prosthetic treatment adapted to immunocompromised or immunosuppressed patients, no reliable data can be found in literature. Given the increased demands on the peri-implant soft tissue because of an inadequately functioning immune system, prosthetic designs which facilitate good oral hygiene and, if necessary, mucosal load relief should be pursued.

9. **Follow-up care**
   An essential aspect for successful long-term implant therapy is regular follow-up care. In patients with immunodeficiency, the required follow-up care should be individually determined and consider the underlying disease, while being performed on a regular basis. It is advisable to provide patients with additional follow-up care during acute phases of immunodeficiency.

**Checklist for risk stratification (low risk profile):**
- phase of the underlying disease (chronic or inactive)
- stably adjusted immunosuppressive medication
- adequate oral hygiene
- focal infections are treated before implant surgery
- clinical and radiological examination reveals normal healing of hard and soft tissues
- bone augmentation is not necessary before implant surgery
- perioperative systemic antibiotic prophylaxis
- conventional loading (> 2 months healing period)
- features that facilitate good oral hygiene are integrated into prosthetic design
- follow-up care is adapted and structured according to the patient’s specific risk

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