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Review

A systematic review and meta-analysis of removable and fixed implantsupported prostheses in edentulous jaws: post-loading implant loss

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[Corrections added after initial online publication on 9 February, 2015: the word five studies has been changed as 54 studies in the Results section in Abstract.]

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Kern J-S, Kern T, Wolfart S, Heussen N. A systematic review and meta-analysis of removable and fixed implant-supported prostheses in edentulous jaws: post-loading implant loss. *Clin. Oral Impl. Res.* 27, 2016, 174–195 doi: 10.1111/clr.12531 Key words: edentulous mandible, edentulous maxilla, implant-supported prosthesis, meta-analysis, systematic review

Abstract

Objectives: The aim of this systematic review was to analyze post-loading implant loss for implantsupported prostheses in edentulous jaws, regarding a potential impact of implant location (maxilla vs. mandible), implant number per patient, type of prosthesis (removable vs. fixed), and type of attachment system (screw-retained, ball vs. bar vs. telescopic crown).

Material and methods: A systematic literature search for randomized-controlled trials (RCTs) or prospective studies was conducted within PubMed, Cochrane Library, and Embase. Quality assessment of the included studies was carried out, and the review was structured according to PRISMA. Implant loss and corresponding 3- and 5-year survival rates were estimated by means of a Poisson regression model with total exposure time as offset.

Results: After title, abstract, and full-text screening, 54 studies were included for qualitative analyses. Estimated 5-year survival rates of implants were 97.9% [95% CI 97.4; 98.4] in the maxilla and 98.9% [95% CI 98.7; 99.1] in the mandible. Corresponding implant loss rates per 100 implant years were significantly higher in the maxilla (0.42 [95% CI 0.33; 0.53] vs. 0.22 [95% CI 0.17; 0.27]; P = 0.0001). Implant loss rates for fixed restorations were significantly lower compared to removable restorations (0.23 [95% CI 0.18; 0.29] vs. 0.35 [95% CI 0.28; 0.44]; P = 0.0148). Four implants and a fixed restoration in the mandible resulted in significantly higher implant loss rates compared to five or more implants with a fixed restoration. The analysis of one implant and a mandibular overdenture also revealed higher implant loss rates than an overdenture on two implants. The same (lower implant number = higher implant loss rates for maxillary overdentures on <4 implants were significantly higher than for four implants (7.22 [95% CI 5.41; 9.64] vs. 2.31 [1.56; 3.42]; P < 0.0001).

Conclusions: Implant location, type of restoration, and implant number do have an influence on the estimated implant loss rate. Consistent reporting of clinical studies is necessary and high-quality studies are needed to confirm the present results.

Introduction and rationale

Complete edentulism still is a common health problem. Although oral health studies illustrated a decrease of individuals suffering from an edentate status, in Germany still 22.6% of 65- to 70-year olds were completely edentulous in the year 2005 (Micheelis & Schiffner 2006).

A complete denture is the classic therapy of full edentulism. Nowadays, this kind of rehabilitation might not be considered as the standard therapy for the lower edentulous jaw any longer. The stabilization of the lower denture with at least two endosseous implants is applied for more than 20 years and was recommended by Feine and co-workers in the McGill consensus statement as standard therapy in 2002 already (Feine et al. 2002a,b,c).

The diversity of problems caused by complete dentures is not a modern issue. Patients do not only complain about insufficient chewing abilities and articulation problems, but also experience psychic strain and social impairment (Albaker 2013). On the contrary, clinical studies investigating the potential impact of implant-supported prostheses on

© 2015 The Authors. Clinical Oral Implants Research published by John Wiley & Sons Ltd. This is an open access article under the terms of the Creative Commons Attribution-NonCommercial-NoDerivs License, which permits use and distribution in any medium, provided the original work is properly cited, the use is non-commercial and no modifications or adaptations are made. the oral health-related quality of life were able to show clear improvement after implants had been inserted (Zitzmann & Marinello 2000a; Allen & McMillan 2003; Scala et al. 2012; Zembic & Wismeijer 2014). It is worth mentioning that clear evidence of benefits for the patient is merely available for the edentulous lower jaw with two interforaminal implants and an overdenture compared to a complete denture. The few studies concerning patient-centered outcome for implant-supported prostheses in the maxilla indicate advantages for the patient. However, considering daily practice, it has to be assumed that the majority of patients with a maxillary complete denture do not articulate major problems.

This systematic review is an update of our own (Schley & Wolfart 2011) and other previously published reviews on the edentulous jaw. As a result of clinical diversity reasons, usually, only a limited number of studies were included in these reviews. Moreover, probably due to a lack of high-quality studies, most of them also included retrospective studies (Lambert et al. 2009; Slot et al. 2009; Heydecke et al. 2012), which are known to have a lower level of evidence. Furthermore, they either included the edentulous maxilla (Slot et al. 2009) or mandible (Payne & Solomons 2000; Roccuzzo et al. 2012; Papaspyridakos et al. 2013) or pooled the results for both jaws (Papaspyridakos et al. 2012). Two very interesting systematic reviews with meta-analysis were recently published (Papaspyridakos et al. 2012, 2013). They focused on biologic and technical complications of fixed implant restorations in edentulous mandibles and implant and prosthodontic survival rates of both jaws and reported an implant survival rate of 97.3% after 10 years.

There is still a large variety of opinions on the best rehabilitation of an edentulous patient. The patient's wish and his or her individual circumstances, which also include financial capacities, have first priority in the decision-making. The anatomic situation and the dentist's knowledge, that is his or her internal evidence, determine the further procedure. Nowadays, the insertion and/or restoration of dental implants in edentulous jaws can considered to be one of the basic treatment modalities in a dentist's everyday practice. Therefore, it seems to be essential to define reproducible treatment protocols that support the individual's expertise and help to establish clear concepts in the sense of an evidence-based dentistry.

The "optimal" number of implants for edentulous jaws still seems to be debatable.

Different reviews tried to address this question (Lambert et al. 2009; Slot et al. 2009; Roccuzzo et al. 2012) and a recently published clinical guideline at least provided key recommendations concerning number of implants and type of implant prosthesis for the edentulous maxilla (Schley & Wolfart 2011; Schley et al. 2013).

To the authors' best knowledge, the potential influence of several factors (not only implant number) on the outcome of dental implants in edentulous patients has not been systematically elaborated, statistically analyzed and compared for both fixed and removable restorations for maxilla *and* mandible in *one* review.

Thus, the aim of this systematic review was to address the following focused question:

Is there an impact of implant location (maxilla vs. mandible), implant number, type of prosthesis (fixed vs. removable) and/or different anchorage systems on the implant loss rate concerning the implant-prosthodontic rehabilitation of edentulous patients?

Material and methods

Protocol

Prior to the systematic literature search, a review protocol was determined with the software Review Manager, version 5.2.

Structure of the review

The systematic review was edited according to the "Preferred Reporting Items for Systematic Reviews and Meta-Analyses" (PRISMA) (Moher et al. 2009).

Eligibility criteria

The focused question was formulated according to the PICOS format, as suggested by the Center for Evidence-Based Medicine and served as a basis for the systematic literature search (Askig Focused Questions 2014):

Patients: edentulous patients (both jaws or either upper or lower jaw) with an implant-retained fixed or removable prosthesis;

Interventions: insertion of either machined or rough-surfaced endosseous titanium implants with a root-like or cylindrical form, irrespective of implant number, length, diameter, position, or angulation, into either local or augmented bone, prosthodontic rehabilitation with a fixed full-arch bridge, segmented reconstructions or a removable overdenture according to an immediate, early or conventional loading protocol.

Comparisons: comparison of different types of prostheses (fixed vs. removable) and/or anchorage systems (ball/locator, bar, telescopic crowns) or fixation mode (screw-retained/cemented) with different implant numbers, in one or between both jaws.

Outcomes: implant survival rate or number of implant losses after prosthetic loading after an observation period of at least 3 years.

Study design: randomized-controlled trials (RCTs) or prospective clinical studies as reported by the authors

Definitions: A prosthesis not being detachable by the patient himself was defined as "fixed prosthesis," that is, screw-retained or cemented fixed full-arch or segmented prostheses. An overdenture retained by different anchorage systems (bar, ball/locator or telescopic crown), and accordingly being removable by the patient, was defined as "removable prosthesis." Regarding different implant surfaces merely a simple distinction between machined and so-called rough implant surfaces was made. A further differentiation of roughening methods or surface modifications, respectively, was not applicable. The loading protocols were defined according to Esposito et al. (2007), that is, an immediate loading was considered to be within 1 week after implant insertion, an early loading between 1 weeks and 2 months, and a conventional loading after a healing period of more than 2 months.

An implant being still *in situ* with a bony anchorage after the observation period was defined as "implant survival," irrespective of hard or soft tissue condition around the implant. Prosthetic loading (immediate or after a conventional healing period) was defined as baseline, meaning, that so-called early losses, that is losses before prosthetic loading, were noted but not statistically evaluated.

Exclusion criteria: no clinical study, retrospective studies, observation period of <3 years, no mean observation period or detailed information on time of implant loss/dropout, no separate reporting of maxilla and mandible or fixed and removable prostheses, provisional implants, ceramic implants, or implants placed into the pterygomaxillary, zygomatic or palatal region, transmandibular implants, studies reporting on the same patient cohort more than once.

Information sources

The electronic databases of Medline (Pub-Med), Cochrane Library, and Embase were searched. A supplementary manual search in different German dental journals (Deutsche Zahnärztliche Zeitschrift, Implantologie, Quintessenz, Zeitschrift für Zahnärztliche Implantologie), reference lists of available publications, and private databases (End Note libraries) was conducted. Authors of available studies were contacted per mail in case of unclear data.

May 7, 2014 was the last date of search. (Table 1).

Search strategy

The search strategy is described in Table 1. The PubMed search complied with the PICOS question addressing Patients, Intervention, Comparison, Outcome and Study design.

Study selection

The resulting initial hits of the above-mentioned search were screened, and a first preselection by title was undertaken. Titles were sequentially excluded if they indicated a nonrelevant content (e.g., no dental implants, animal or *in vitro* study). In case of any uncertainty, an additional abstract reading was performed. Abstracts of the selected titles were inspected for relevance resulting in a choice of possibly eligible full texts. If studies were published by the same author or institution several times, the according manuscripts were thoroughly read and compared to avoid the inclusion of duplicate data. After full-text selection and data extraction, it was decided whether the publication was adequate for the intended systematic review.

Study selection and data extraction were performed independently by two reviewers (JSK, TK), and any disagreement was solved by discussion. To assess consistency among the reviewers, the interreviewer reliability using Cohen's Kappa statistic (κ) was analyzed.

Data collection and data items

Extracted data were filled into pre-defined forms and included the following parameters: author, year, total number of patients/prostheses investigated, observation period, total number of implants, number and time of dropouts on implant level, number of implants per patient, type of implant prosthesis, type of anchorage system, implant survival and implant losses before and after loading. Moreover, implant system, implant surface, loading protocol, and bone augmentation procedures were noted. All variables were pre-determined and no additional variables were added after the reviewing had started.

Risk of bias within and across studies

A potential risk of bias within the included studies was assessed using the methodology checklists provided by the Scottish Intercollegiate Guidelines Network (SIGN). These lists comprise the critical appraisal of the selection of subjects, the applied assessment, potential confounders, and the statistical analysis, and finally, the overall assessment of the methodological quality of the study:

- High quality: (++) Majority of criteria met. Little or no risk of bias. Results unlikely to be changed by further research.
- Acceptable quality: (+) Most criteria met. Some flaws in the study with an associated risk of bias, Conclusions may change in the light of further studies.
- Low quality: (-) Either most criteria not met, or significant flaws relating to key aspects of study design. Conclusions likely to change in the light of further studies.

Further explanations are shown as footnote of Table 2.

Information sources	Electronic databases	PubMed, EMBASE, The Cochrane Library
	Additional sources	German dental journals (not Medline listed), Private databases, reference lists
Search strategy	Population	#1: (("mouth, edentulous"[MeSH Terms] OR ("mouth"[All Fields] AND
PubMed		"edentulous" [All Fields]) OR "edentulous mouth" [All Fields] OR "edentulous"
		[All Fields]) OR (completely[All Fields] AND ("mouth, edentulous"[MeSH Terms] OR
		("mouth"[All Fields] AND "edentulous"[All Fields]) OR "edentulous mouth"[All Fields]
		OR "edentulous" [All Fields]))) AND ("maxilla" [MeSH Terms] OR "maxilla" [All Fields]
		OR ("mandible"[MeSH Terms] OR "mandible"[All Fields])) AND ("dental implants"
		[MeSH Terms] OR ("dental"[All Fields] AND "implants"[All Fields]) OR
		"dental implants"[All Fields] OR ("dental"[All Fields] AND "implant"
		[All Fields]) OR "dental implant"[All Fields])
	Intervention	#2: #1 AND ((implant[All Fields] AND ("denture, overlay"[MeSH Terms] OR
		("denture"[All Fields] AND "overlay"[All Fields]) OR "overlay denture"[All Fields]
		OR "overdenture" [All Fields])) OR (complete [All Fields] AND implant [All Fields]
		AND removable[All Fields] AND ("dental prosthesis" [MeSH Terms] OR ("dental"
		[All Fields] AND "prosthesis"[All Fields]) OR "dental prosthesis"[All Fields]))) OR
		(complete[All Fields] AND fixed[All Fields] AND ("dental prosthesis"[MeSH Terms]
		OR ("dental"[All Fields] AND "prosthesis"[All Fields]) OR "dental prosthesis"
		[All Fields]))) OR (full-arch[All Fields] AND restoration[All Fields])) OR
		("dental prosthesis, implant-supported"[MeSH Terms] OR ("dental"
		[All Fields] AND "prosthesis"[All Fields] AND "implant-supported"
		[All Fields]) OR "implant-supported dental prosthesis"[All Fields] OR ("implant"[All Fields] AND "supported"[All Fields] AND "dental"
		[All Fields] AND "prosthesis"[All Fields]) OR "implant-supported
		dental prosthesis" [All Fields]))
	Comparison	Covered by Population, Intervention and Outcome search
	Outcome	#3: #1 AND ("survival rate" [MeSH Terms] OR ("survival" [All Fields] AND
	outcome	"rate" [All Fields]) OR "survival rate" [All Fields])
	Limits (filters)	Clinical trial, humans
	Period	No time restriction
Search strategy		"dental implant"/exp OR "dental implant" AND "edentulous"
EMBASE	Limits (filters)	No
Search strategy	(,	"dental implant AND edentulous"
Cochrane Central Register of Controlled Trials	Limits (filters)	Trials

A special assessment of possible publication bias or selective reporting was not performed. There were no clues indicating that data within studies were missing. Several studies were industrially sponsored.

Summary measures and synthesis of results

In the majority of included studies, the investigated patients were subdivided into different groups, for example, to compare different loading protocols, anchorage systems, implant numbers or implant types. Whenever possible, data of these groups were recorded separately so that the statistical analysis incorporated more study populations than indicated by the number of included studies.

The primary outcome of the meta-analysis was the estimated implant loss rate per 100 implant years in the edentulous maxilla and mandible depending on type of prosthesis (fixed or removable), type of attachment (bar/ball/telescopic crowns, screw-retained/ cemented), and implant number. This rate describes, for example, the risk of an implant loss regarding 100 implants over the course of 1 year or the risk of an implant loss regarding 10 implants over 10 years.

Based on these implant loss rates, 3- and 5year implant survival rates were estimated.

For simplification, implant numbers were categorized for both jaws. For the mandible, these categories were as follows: one implant, two implants, four implants, and ≥ 5 implants. For the maxilla, a subdivision was chosen as follows: <4 implants, four implants, and ≥ 6 implants. Whenever information on the exact implant number per patient could not be extracted, further subcategories were chosen (2-4 implants and 4-6 in the mandible, and 5-6 implants in the maxilla). Data of these overlapping categories were used to strengthen the overall analysis, but were not included for any comparisons. The same applies to missing or not extractable information of other categories (e.g., loading protocol or implant surface, declared as "not applicable"). Tables 3 and 4 illustrate in detail which particular category was "not applicable". The number of included study populations for each analysis is shown in the Tables, as well.

Ball and locator attachments were summarized in one category ("ball"). The category "bar" included all types of bars. The category "telescopic crowns" included all types of double crowns.

Additional subgroup analyses were carried out to calculate the estimated implant loss rates per 100 implant years with regard to loading protocol (immediate vs. conventional) and implant surface (rough vs. machined).

According to Pjetursson et al. (2007) implant loss rates were calculated by dividing the number of events (loss after loading) by the total exposure time of the implants. The total exposure time consisted of a) the exposure time of the implants being followed for the complete observation period, b) the exposure time of the implants until loss, and c) the exposure time until an implant dropout had occurred (withdrawal for different reasons, patient's death/illness, patient missed recall or moved). If the explicit information on an implant was not provided, that is time of dropout or loss, the total exposure time was calculated by multiplying the number of initially inserted implants (minus losses before loading) by the mean follow-up time. Implant loss rates were calculated for every study population by dividing the number of events (post-loading losses) by the total implant exposure time in years.

A Poisson regression models with a logarithmic link function and the logarithm of total exposure time as an offset variable were fitted to the data to obtain a cumulative estimate for the appropriate implant loss rate and a corresponding 95% confidence interval. 3- and 5-year implant survival rates and related 95% confidence limits were derived from the equation $S(t) = e^{\lambda t}$ where t denotes the time and λ the implant loss rate by assuming constant event rates over time. Comparison of loss rates in different subgroups were contrasted by descriptive P-values resulting from the correspondent Poisson regression model. Factors, which showed significant influence on implant loss in the unianalysis, were simultaneously variate analyzed in a multivariate Poisson regression model. To explore possible effect modifiers, all two-way interactions between factors were evaluated within this model. The final Poisson regression model included all main effects and significant two-way interactions. P-values less than or equal to 0.05 were regarded as statistically meaningful. Due to the explorative nature of the study, no adjustment to the significance level was made. All statistical analyses were performed using the software SAS (SAS Institute Inc., Cary, NC, USA, Version 9.3).

Results

Literature search

The search strategy, as described in Figure 1 and Table 1, resulted in an initial number of

4317 titles. 3823 titles could be excluded after screening. The manual search revealed 80 further abstracts.

After filtering the abstracts and excluding the duplicates, the reviewers decided to conduct a full-text analysis of 210 publications. Fifty-six publications, describing 54 studies, could be considered for a quantitative analysis. The interreviewer agreement was found to be $\kappa = 0.9$ (SD 0.098) concerning final study selection.

Study characteristics

The included clinical trials were published within an almost 20-year period (1996–2013). Ten of them investigated the edentulous maxilla, 36 the edentulous mandible, and eight investigated both jaws. Four studies were RCTs, and the rest were prospective clinical studies, sometimes described as "prospective, randomized" or "prospective, controlled" (Table 2).

In the majority of studies, observation periods between 3 and 10 years were stated, and in four studies, 11 or more years of follow-up were reported (Table 3). Within the 54 included clinical trials, altogether 81 study populations have been investigated. Whenever subgroups were described in a study, this information is shown in Tables 4 and 5. In 30 study populations, patients were restored with fixed full-arch prostheses, and in the residual 51 study populations, patients received removable overdentures. All of the fixed, definitive prostheses had a metal framework (Au, CoCr, or Ti), veneered with acrylic resin or ceramic and were screwretained. None of the studies reported on cemented or adhesively fixed prostheses. The removable prostheses were generally fabricated out of acrylic resin, reinforced with a metal framework or reinforcement (CoCr) and attached by different anchorage systems (ball, locator, telescopic crown as un-splinted retention elements and different bars enabling a primary splinting).

Altogether 2368 patients received 9267 implants. Various implant types with differsurface modifications were ent used (Table 3). All implants were titanium implants with different lengths and diameters. Implant numbers per patient varied between 1 and 6 implants in the mandible and 2 and 10 in the maxilla. The interforaminal area was the preferred area for implant positioning in the mandible. If only one implant was inserted in the edentulous lower jaw, it was located in the midline symphysis, representing the absolute minimal treatment concept. In the maxilla, implant positions

Table 2. Risk of bias within studies

		Overall		
Studies in alphabetical		assessment	Level of	
order	Study design	of the study*	evidence†	Sponsoring/support as reported by the authors
Agliardi et al. (2012)	Prospective	+	2+	n.r.
Akca et al. (2010)	Prospective	+	2+	Partly supported by State Planning Organization, Prime Ministry, Republic of Turkey
Akoglu et al. (2011)	Prospective	++	2++	n.r.
Arvidson et al. (1998)	Prospective	+	2+	n.r.
Arvidson et al. (2008)	Prospective	++	2++	Supported and sponsored by Institut Straumann AG, Basel, Switzerland
Behneke et al. (2002)	Prospective	+	2+	n.r.
Bergendal & Engquist (1998)	Prospective	+	2+	n.r.
Cehreli et al. (2010)	RCT	+	1+	Partly supported by the State Planning Organization, Prime Ministry, Republic of Turkey
Chiapasco & Gatti (2003)	Prospective	+	2+	n.r.
Collaert & De Bruyn (2008)	Prospective	+	2+	n.r.
Cooper et al. (2008)	Prospective	++	2++	n.r.
Cordioli et al. (1997)	Prospective	+	2+	n.r.
Covani et al. (2012)	Prospective	+	2+	n.r.
Crespi et al. (2012)	Prospective	+	2+	n.r.
De Bruyn et al. (2008)	Prospective	+	2+	n.r.
De Santis et al. (2012)	Prospective	+	2+	n.r.
Degidi et al. (2010) Ekelund et al. (2003)/	Prospective	++++	2+ 2+	n.r.
Lindquist et al. (1996)	Prospective	Ŧ	ZT	n.r.
Eliasson et al. (2010)	Prospective, randomized	+	2+	Supported by a grant from Public Dental Health, Örebro County Council, Sweden
Elsyad et al. (2012)	RCT	+	1+	n.r.
Engquist et al. (2005)	Prospective, controlled	++	2++	n.r.
Fischer & Stenberg (2012)	Prospective	+	2+	Institut Straumann AG, Basel, Switzerland assisted to prepare the manuscript
Gotfredsen & Holm (2000)	Prospective, randomized	+	2+	Astra Tech, Sweden supplied implants and implant components
Harder et al. (2011)	Prospective	+	2+	Supported by a grant from Camlog Biotechnologies AG, Basel, Switzerland
Heijdenrijk et al. (2006)	Prospective, randomized	+	2+	n.r.
Heschl et al. (2013)	Prospective	+	2+	n.r.
Jemt et al. (1996)/ Watson et al. (1997)	Prospective	+	2+	Nobelpharma AB, Göteborg, Sweden supported with components
Krennmair et al. (2008)	Prospective, randomized	++	2++	Study was self-funded by the authors and their institution
Krennmair et al. (2011)	Prospective, randomized	++	2++	n.r.
Krennmair et al. (2012)	Prospective, randomized	++	2++	Study was self-funded by the authors and their institution
Lethaus et al. (2011)	Prospective	+	2+	Supported and sponsored by Institut Straumann, Basel, Switzerland
Liddelow & Henry (2010)	Prospective	+	2+	n.r.
Lorenzoni et al. (2013)	Prospective	+	2+	n.r.
Meijer et al. (2004)	RCT	+	1+	n.r.
Meijer et al. (2009a)	Prospective	+	2+	n.r.
Meijer et al. (2009b) Mertens et al. (2012)	Prospective Prospective	+ +	2+ 2+	n.r. Supported by Astra Tech AB, Sweden and Bioscientia, Germany, provided kits for IL-1 composite genotype tests
Murphy et al. (2002)	Prospective	+	2+	n.r.
Naert et al. (1998)	Prospective	+	2+	Ceka NV, Belgium, support with prosthesis components
Nyström et al. (2009a,b)	Prospective	+	2+	n.r.
Nyström et al. (2009a,b)	Prospective	+	2+	n.r.
Örtorp & Jemt (2012)	Prospective	+	2+	Supported by Wilhelm and Martina Lundgren Research Foundation
Rasmusson et al. (2005)	Prospective	+	2+	n.r.
Richter & Knapp (2010)	Prospective	+	2+	Biomet 3i supported the study with implants and implant components
Romeo et al. (2004)	Prospective	+	2+	n.r.
Schwarz et al. (2010)	Prospective	+	2+	Financially supported by FRIADENT GmbH, Germany
Sjöström et al. (2007)	Prospective	+	2+	n.r.
Stoker et al. (2012)	RCT	+	1+	Supported by a grant (188/2000) from the ITI Foundation for the Promotion of Oral Implantology, Switzerland
Testori et al. (2004)	Prospective	+	2+	n.r.
Van de Velde et al. (2007)	Prospective	+	2+	n.r.
Vroom et al. (2009)	Prospective	+	2+	Partly supported by Astra Tech AB, Sweden
Weinländer et al. (2010)	Prospective	+	2+	Study was self-funded by the authors and their institution
Zitzmann & Marinello (2000a,b)	Prospective	+	2+	n.r.

Table 2. (continued)

Studies in alphabetical order	Study design	Overall assessment of the study*	Level of evidence†	Sponsoring/support as reported by the authors
Zou et al. (2013)	Prospective	÷	2+	Funded by Combined Engineering and Medical Project of Shanghai Jiao Tong University the National Natural Science Foundation of (YG2010MS56), Science and Technology Commission of Shanghai Municipality (13ZR1424000), China (81100788, 31370983, 81371190), the Key Project of Chinese Ministry of Education (212080), Grants for Scientific Research of BSKY (XJ201109), and the Young Top-notch Talent Support Scheme from Anhui Medical University
†Level of evidence accord reviews of case-control or the relationship is causal.	ing to SIGN: 1+ = well-co cohort studies, high-qua 2+ = well-conducted case	nducted meta-analyse lity case–control or co e–control or cohort st	es, systemation whort studies tudies with a	-) High quality (+) Acceptable (-) Low quality. c reviews, or RCTs with a low risk of bias; 2++ = high-quality systematic with a very low risk of confounding or bias and a high probability that a low risk of confounding or bias and a moderate probability that the unding or bias and a significant risk that the relationship is not causal;

were often not described precisely. Only the following authors described the area of implant placement in more detail: Fischer & Stenberg (2012, 2013) located 5–6 implants from second premolar to second premolar. Agliardi et al. (2012) and Degidi et al. (2010) placed implants in the anterior area and (tilted) implants in the regions of the anterior and posterior sinus wall. De Santis et al. (2012) inserted 6–10 implants in the positions of former incisors, canines, premolars, and molars.

3 = Non-analytic studies, for example case reports, case series.

The results for fixed prostheses presented by Romeo et al. (2004) could not be considered, because only three patients had been provided with a fixed prosthesis. In another trial, the observation period was too short, and therefore, the "removable cases" had to be excluded (Zitzmann & Marinello 2000a, b). Covani et al. (2012) merely included six patients with an edentulous lower jaw, and hence, these cases were not regarded in this review. Some authors observed the same study population but reported on different clinical outcomes in different publications (surgical, periodontal, prosthetic) (Jemt et al. 1996; Watson et al. 1997; Fischer & Stenberg 2012, 2013). Their results were summarized.

Generally, criteria for the inclusion or exclusion of patients were pre-defined. For obvious reasons, these criteria were not consistent among the studies. Mostly, patients with severe diseases or uncontrolled diabetes, psychological problems, and heavy smokers were excluded. In general, the average age of the patients was between 50 and 60 years, although it is worth mentioning that mean ages were not always provided or sometimes not for all indications being investigated in one particular study (e.g., maxilla or mandible, edentulous or partially edentulous).

In the majority of studies, a 2-stage surgical procedure and a conventional loading protocol were carried out, but non-submerged healing (1-stage surgery) followed by immediate prosthetic loading was applied, as well (Table 3). Pre-implantological or simultaneous bone augmentation was reported in six studies and ranged from rather simple procedures (e.g., filling of post-extraction sites (Agliardi et al. 2012; Zou et al. 2013) to complex reconstructions such as Le Fort I osteotomies with interpositional bone grafts (Nyström et al. 2009b; De Santis et al. 2012) or onlay osteoplastics (Nyström et al. 2009a). Sjöström et al. (2007) either applied inlay, onlay, or interpositional grafting with free iliac grafts. Covani et al. (2012) partly carried out simultaneous sinus floor elevation with the ostetome technique. Richter & Knapp (2010) performed either bone splitting or bone spreading but no augmentation in case of heavy bone resorption. Three other studies (De Bruyn et al. 2008; Heschl et al. 2013; Lorenzoni et al. 2013) reported not to have applied augmentative or regenerative procedures. The rest of the studies cannot be commented as the authors did not make any statements about bone augmentation.

The examination of patients usually comprised the recording of several indices, that is, plaque indices, bleeding indices, and pocket depth. Implant stability was checked, sometimes by means of radio-frequency analysis or "damping capacity assessment" (Heschl et al. 2012). In the majority of the included studies, a radiographic examination was performed to measure marginal bone level changes. Several techniques were used for this, for example, standardized radiographic holders to achieve the highest possible reproducibility. In many cases, merely panoramic radiographs were compared.

Overall implant survival and loss

Results of individual patient groups

Estimated implant survival rates after 5 years ranged from 89.0% to 100% for fixed prostheses concerning both jaws (Tables 4 and 5). For removable prostheses, estimated survival rates of 24.9% up to 100% were calculated. The very low survival rate of 24.8%, with an associated annual implant loss rate of 27.8 per 100 implant years, is related to a very small patient group (n = 7) that was restored with merely 2 diameter-reduced implants and an overdenture in the edentulous maxilla (Richter & Knapp 2010).

Synthesis of results

Comparing the overall implant loss rate per 100 implant years for fixed vs. removable prostheses, a statistically significant difference could be assessed (P < 0.0001) if the category <4 implants (maxilla) was included (Tables 6 and 7). Excluding this latter category, there was also a significantly higher implant loss rate per 100 implant years comparing fixed and removable restorations (0.23 [95% CI 0.18; 0.29] vs. 0.35 [95% CI 0.28; 0.44]; P = 0.0148).

Regarding different attachment types for overdentures in both jaws, no significant differences could be detected for ball vs. bar anchorage. The estimated implant loss rate per 100 implant years was similar (0.34 [95% CI 0.16; 0.72] vs. 0.35 [95% CI 0.27; 0.46] per 100 implant years; P = 0.9607). The comparison of bar vs. telescopic crown and ball vs. telescopic crown was not possible (no implant losses, merely three study populations included (not regarding the study of Richter & Knapp (2010), as it belonged to the group <4 implants, see below).

Study (Year of publication)	Study design	Jaw	Type of prosthesis	Type of anchorage	Implant system (as reported by the authors)	Loading protocol	Total number of implants	Total number of prostheses	Follow-up period (years)
Agliardi et al. (2012)	Prospective	Maxilla	Fixed	Screw- retained	Brånemark, Nobel Speedy (Nobel	Immediate	192	32	4.6
Collaert & De Bruyn (2008)	Prospective	Maxilla	Fixed	Screw- retained	Biocare) TiOblast Astra Tech	Immediate	195	25	m
De Santis et al.	Prospective	Maxilla	Fixed	Screw-	(Dentspiy) n.a.	Conventional	154	20	4.3
(2012) Degidi et al. (2010)	Prospective	Maxilla	Fixed	Screw- ratained	Xive (Dentroliv)	Immediate	210	30	ĸ
(2010) Fischer & Stenberg (2012)	Prospective	Maxilla	Fixed	Screw- retained	Esthetic Plus SLA	lmmediate/ conventional	142	24	10
Mertens et al.	Prospective	Maxilla	Fixed	Screw-	Astra Tech,	Conventional	106	17	11.3
vzu z) Naert et al. (1998)	Prospective	Maxilla	Fixed	Screw- retained	Brånemark (Nobel Riocare)	Conventional	53	1	m
Nyström et al. (2009b)	Prospective	Maxilla	Fixed	Screw- retained	Brånemark (Nobel Riocare)	Conventional	167	26	13
Nyström et al. (2009a)	Prospective	Maxilla	Fixed	Screw- retained	Brånemark (Nobel Riocare)	Conventional	334	44	11
Richter & Knapp (2010)	Prospective	Maxilla	Remov- able	Telescopic crown, locator	Osseotite (Biomet 3i)	Conventional	44	27	Ŋ
Sjöström et al. (2007)	Prospective	Maxilla	Fixed	Screw- retained	Brånemark (Nobel Biocare)	Conventional	222	29	m
Zitzmann & Marinello (2000b)	Prospective	Maxilla	Fixed	Screw- retained	Brånemark (Nobel Riocare)	Conventional	84	10	с. С
Zou et al. (2013)	Prospective	Maxilla	Remov- able	Telescopic crown, bar, locator	ITI (Straumann)	Conventional	120	30	m
Akoglu et al. (2011)	Prospective	Mandible	Remov- able	Ball	ITI (Straumann), Swiss Plus (Zimmer Dental), Astra Tech (Dentralv)	Conventional	72	98	Ŋ
Arvidson et al.	Prospective	Mandible	Fixed	Screw- retained	Astra Tech (Dentsolv)	Conventional	618	107	5
Arvidson et al. (2008)	Prospective, multicenter	Mandible	Fixed	Screw- retained	ITI Monotype SLA (Straimann)	Early	250	61	m
Behneke et al. (2002)	Prospective	Mandible	Remov- able	Bar	(Straumann)	Conventional	340	100	5.8

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Table 3. (continued)									
Study (Year of publication)	Study design	Jaw	Type of prosthesis	Type of anchorage	Implant system (as reported by the authors)	Loading protocol	Total number of implants	Total number of prostheses	Follow-up period (vears)
Cehreli et al. (2010)	ي لاط	Mandible	Remov- able	Ball	SLA (Straumann), Brånemark TiUnite (Nobel	Early	56	5 8	ц
Chiapasco & Gatti (2003)	Prospective	Mandible	Remov- able	Bar	Biocore Biocore Ha-Ti (Mathys Dental), ITI (Straumann), Bränemark Conical (Nobel Biocore), Biocore),	Immediate	328	82	5.2
Cooper et al.	Prospective	Mandible	Remov-	Ball	(Dentsply) Friatec	Conventional	118	59	5
(2008) Cordioli et al.	Prospective	Mandible	able Remov	Ball	(Dentsply) Biomet 3i	Conventional	21	21	5
De Bruyn et al. (2008)	Prospective	Mandible	Fixed	Screw- retained	TiOblast Astra Tech	Immediate	125	25	m
Ekelund et al. (2003)/Lindquist et al. (1996)	Prospective	Mandible	Fixed	Screw- retained	Standard Brånemark (Nobel Biocare)	Conventional	273	47	21.5
Eliasson et al. (2010)	Prospective, randomized	Mandible	Fixed	Screw- retained	Paragon TPS (Zimmer	Conventional	168	29	Ŋ
Elsyad et al. (2012)	RCI	Mandible	Remov- able	Ball	Dental) Spectra System Screw Plant (Implant	Immediate/ conventional	72	36	m
Engquist et al. (2005)	Prospective, controlled	Mandible	Fixed	Screw- retained	Direct LLC) Standard Bränemark, Bränemark conical 1- piece, Bränemark Mk II (Nobel	Early/ conventional	432	108	m
Gotfredsen & Holm (2000) Harder et al. (2011)	Prospective, randomized Prospective	Mandible Mandible	Remov- able Remov- able	Bar, ball Ball	Biocare) Astra Tech (Dentsply) Camlog Promote screw line	Conventional Conventional	52 11	26	3.6 3.6
Heijdenrijk et al. (2006)	Prospective, randomized	Mandible	Remov- able	Bar	(Camlog) IMZ TPS, solid screw TPS	Conventional	120	60	2
Heschl et al. (2013)	Prospective	Mandible	Remov- able	Bar	(Juraumann) Xive S plus (Dentsply)	Conventional	156	39	Ŋ

iable 3. (continued)					Implant				
Study (Year of publication)	Study design	Jaw	Type of prosthesis	Type of anchorage	system (as reported by the authors)	Loading protocol	Total number of implants	Total number of prostheses	Follow-up period (years)
Krennmair et al. (2008)	Prospective, randomized	Mandible	Remov- able	Bar	IMZ (Dentsply), Frialoy, (Dentsply), Camlog root	Conventional	204	51	Ω
Krennmair et al. (2011)	Prospective, randomized	Mandible	Remov- able	Ball, telescopic	line (Camlog) Camlog root line (Camlog)	Conventional	50	25	5
Krennmair et al. (2012)	Prospective, randomized	Mandible	Remov- able	crown Bar, telescopic crown	Camlog root line/screw	Conventional	204	51	m
Lethaus et al.	Prospective	Mandible	Remov- able	Bar	IINE (Lamiog) SLA (Straiimann)	Early	70	14	5
Liddelow & Henry (2010)	Prospective	Mandible	able	Ball	Brånemark Mk III TiUnite,	Immediate	32	32	m
					Branemark Mk III machined (Nobel				
Lorenzoni et al.	Prospective	Mandible	Remov- abla	Bar	Blocare) Xive S Plus (Dentsniv)	Immediate/ conventional	160	40	5
Meijer et al. (2004)	RCT	Mandible	Remov- able	Bar	(Dentsply), (Dentsply), Brånemark (Nobel	Conventional	122	61	10
Meijer et al. (2009b)	Prospective	Mandible	Remov- able	Bar	Brocare) IMZ TPS (Dentsply), Bränemark (Nobel Biocare), ITI solid screw TPS	Conventional	180	06	9
Meijer et al. (2009a)	Prospective	Mandible	Remov- ahle	Bar	(Straumann) IMZ TPS (Dentsolv)	Conventional	180	60	10
Murphy et al. (2002)	Prospective	Mandible	Fixed	Screw- retained	Astra Tech (Dentsplv)	Conventional	131	26	5
Schwarz et al. (2010) Stoker et al. (2012)	Prospective RCT	Mandible Mandible	Fixed Remov-	Screw- retained n.a.	Frialoc (Dentsply) 1-stage TPS	Early Conventional	158 296	37 110	4.5 8.3
Testori et al. (2004)	Prospective	Mandible	able Fixed	Screw- retained	Bonefit (Straumann) Osseotite, dual acid-	Immediate	116	19	3.2
					etched, cylindrical, screw-shaped (3i)				

Table 3. (continued)									
					Implant system (as		Total		
Study (Year of publication)	Study design	Jaw	Type of prosthesis	Type of anchorage	reported by the authors)	Loading protocol	number of implants	Total number of prostheses	Follow-up period (years)
Van de Velde et al. (2007)	Prospective	Mandible	Fixed	Screw- retained	Brånemark Mk III/Mk IV	Immediate	91	18	3.8
					(Nobel				
Vroom et al.	Prospective	Mandible	Remov-	Bar	Astra Tech	Conventional	80	20	12
(2009)			able		turned/ +ichloctod				
					uopiasted (Dentsply)				
Weinländer et al.	Prospective	Mandible	Remov-	Bar	IMZ cylindric,	Conventional	252	76	5
(2010)			able		Frialoc (Denctolv)				
					روانطارداتعکار Camlog				
					screw line (Camloo)				
Akca et al. (2010)	Prospective	Both jaws	Remov-	Bar	ITI SLA/TPS	Conventional	124	35	4.9
			able		(Straumann)				
Bergendal &	Prospective	Both jaws	Remov-	Bar, ball	Brånemark	Conventional	115	50	5.2
Engquist (1998)			able		(Nobel Biorare)				
Covani et al.	Prospective	Both jaws.	Fixed	Screw-	Ossean (Intra	Immediate	128	16	3.6
(2012)		only maxilla included		retained	Lock Int)				
Crespi et al. (2012)	Prospective	Both jaws	Fixed	Screw-	PAD system	Immediate	176	44	c
				retained	(Sweden- Martina)				
Jemt et al. (1996)/	Prospective	Both jaws	Remov-	Bar	Brånemark	Conventional	315	133	5
Watson et al. (1997)			able		(Nobel Biocare)				
Örtorp & Jemt	Prospective	Both jaws	Fixed	Screw-	Brånemark	Conventional	728	129	10
(2012)				retained	(Nobel Biorare)				
Rasmusson et al	Prosnertive	Roth jawe	Fixed	Screw-	TiOhlact Actra	Conventional	199	36	10
(2005)				retained	Tech)) -	2	2
Domao at al	Drospotivo	Both issue	Bemoushle	c s	(Denstply)	[control	361	27	0
(2004)	LIOSpective	both Jaws	Velliovable	n.a.	(Straumann)	COLIVERIDORIAL	071	10	v.c

Study	Subgroups within study	Number of implants per patient	Total number of implants	Type of prosthesis and anchorage	Number of post- loading implant losses	Total implant exposure time (implant years)	Estimated implant loss (per 100 implant years)	Estimated implant survival after 5 years (%)
Richter & Knapp	Locator	<4*	14	Removable, BL	12	43.1	27.8	24.6
	Telescopic	<4*	30	Removable, TC	18	119.5	15.1	47.1
	crowns							
Bergendal & Engquist	Bar	<4*	29	Removable, BR	9	140.3	4.3	80.7
	Ball	<4*	18	Removable, BL	7	88.0	8.0	67.2
Romeo et al.	n.a.	<4*	42	Removable		246.0	1.2	94.1
Naert et al.	n.a.	4	53	Removable, BR	m	135.9	2.21	89.4
Akca et al.	n.a.	4	44	Removable, BR	-	210.7	0.5	97.6
Crespi et al.	n.a.	4	96	Fixed, SR	-	285.3	0.35	98.3
Jemt/Watson et al.	n.a.	4	117	Removable, BR	21	375.5	5.6	75.0
Zou et al.	Telescopic	4	40	Removable, TC	0	120.0	0	100
	crowns							
	Bar	4	40	Removable, BR	0	120.0	0	100
	Locator	4	40	Removable, BL	0	120.0	0	100
Agliardi et al.	n.a.	9<⊓	192	Fixed, SC	0	576.0	0	100
Fischer & Stenberg	n.a.	5-6	142	Fixed, SC	4	1095.0	0.4	98.2
Rasmusson et al.	n.a.	5-6	91	Fixed, SC	0	787.0	0	100
Collaert & De Bruyn	n.a.	9<	195	Fixed, SC	0	529.0	0	100
De Santis et al.	n.a.	9<	154	Fixed, SC	0	780.0	0	100
Degidi et al.	n.a.	9~1	210	Fixed, SC	-	630.1	0.2	99.1
Mertens et al.	n.a.	9<	106	Fixed, SC	2	1050.9	0.2	99.1
Nytröm et al.	n.a.	9~	167	Fixed, SC	5	2132.0	0.2	98.8
Nyström et al.	n.a.	9⊂	334	Fixed, SC	4	3674.0	0.1	99.5
Sjöström et al.	n.a.	9⊂	222	Fixed, SC	8	197.5	4.1	81.3
Zitzmann & Marinello	n.a.	9⊂	84	Fixed, SC	0	820.0	0	100
Covani et al.	n.a.	9~	128	Fixed, SC	0	460.8	0	100
Örtorp et al.	n.a.	9<	355	Fixed, SC	17	2042.5	0.8	95.9

Implant survival and loss in the maxilla

Results of individual patient groups

Concerning the estimated 3- and 5-year implant survival rates of both, removable and fixed implant-supported prostheses, these were higher than 95% for the majority of study populations (Table 4). For five of the investigated groups, the estimated implant survival rate was <90% (67.2–89.4%) and for two groups even <50% (24.8% and 47.2%) after 5 years. The low survival rates were associated with an implant number of <4, and removable overdentures and corresponding annual implant loss rates were between 8.0 and 27.8.

Synthesis of results and subgroup analyses

If fixed and removable implant prostheses were compared, the removable prostheses had a significantly higher implant loss rate $(0.28 \ [95\% \ CI \ 0.21; \ 0.38] \ vs. \ 2.31 \ [95\% \ CI \ 1.56; \ 3.42]; \ P < 0.0001)$ (Table 8). Comparing the implant numbers <4 vs. 4 implants in the "removable group", the risk of implant loss is more than three times higher with <4 implants (7.22 \ [95% \ CI \ 5.41; \ 9.64] \ vs. \ 2.31 [95% \ CI \ 1.56; \ 3.42]; \ P < 0.0001). Therefore, this category (<4 implants in the maxilla) was excluded from further statistical analysis.

Fixed restorations with six or more implants demonstrated an implant loss rate of 0.28 [95% CI 0.20; 0.39] per 100 implant years. A comparison of this latter category with lower implant numbers was not possible due to a lack of studies.

Implant survival and loss in the mandible

Results of individual patient groups

Estimated implant survival rates after 3 and 5 years for fixed restorations were generally very high (95–100%) (Table 5). For one patient group, implant survival was <90% (88.8%). Also the results for removable prostheses revealed high survival rates. Here, a small study population being treated with one machined implant and an overdenture stands out in a negative sense with an implant loss rate of 24.2 per 100 implant years and a corresponding 5-year survival estimation of 25.1%.

Synthesis of results

*This category was excluded from further statistical analysis.

Comparing the estimated implant loss rates per 100 implant years in the mandible for fixed and removable prostheses, no significant difference could be detected (0.19 [95% CI 0.13; 0.27] vs. 0.24 [95% CI 0.18; 0.32];

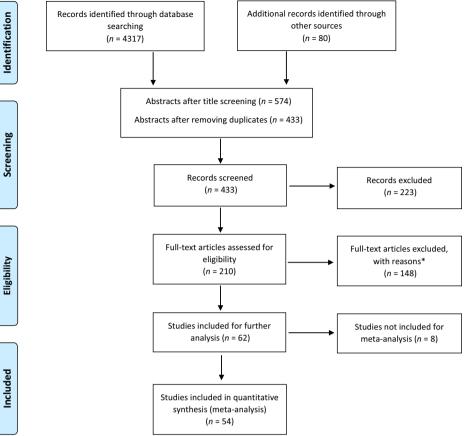


Fig. 1. Modified PRISMA flow chart: study selection process. *zygoma, pterygomaxillary, transmandibular or other region, no clinical examination/no regular follow-up, observation <3 year, no titanium implant, no clinical study, not edentulous, retrospective/study design unclear, no separate reporting of edentulous jaws/removable or fixed prostheses, same patient cohort at earlier stage, case report, no survival rate nor implant loss, different emphasis,

study being judged with "2-" according to SIGN (high risk of bias)

P = 0.2980) (Tables 9 and 10). Comparing different implant numbers, less implants always resulted in significantly higher implant loss estimations (1 vs. 2, 2 vs. 4 (removable prostheses) 4 vs. ≥5 implants (fixed prostheses)). No statistically significant differences were revealed regarding ball vs. bar attachment (0.34 [95% CI 0.16; 0.72] vs. 0.20 [95% CI 0.14; 0.28]; P = 0.1499). Four implants with a fixed restoration resulted in a significantly higher (P < 0.0001) estimated implant loss rate (0.79 [95% CI 0.49; 1.30]) than with a removable restoration [0.11 [95% CI 0.06; 0.23]).

Further subgroup analyses

Implant surface

Machined implants were more prone to implant loss than rough-surfaced implants, in almost every subgroup (Table 11). Concerning fixed restorations, no significant difference in *post-loading* implant loss could be demonstrated comparing machined vs. rough implant surfaces, although machined implant surfaces tend to result in higher loss rates (0.28 [95% CI 0.21; 0.37] vs. 0.19 [95% CI 0.13; 0.28]; P = 0.1177). Furthermore, in the edentulous mandible, no difference in estimated implant loss per 100 implant years between machined and rough implants could be shown (0.25 [95% CI 0.17; 0.36] vs. 0.21 [95% CI 0.16; 0.27]; P = 0.4518).

Loading protocols

Both loading protocols, immediate and conventional, exhibited low implant loss rates per 100 implant years, and no statistically significant differences could be shown concerning fixed restorations in both jaws (0.27 [95% CI 0.15; 0.50] vs. 0.17 [0.12; 0.23]; P = 0.1652) (Tables 12 a, b). However, there was a significantly lower risk of implant loss with a conventional loading protocol concerning the overall analysis, removable prostheses, and the edentulous mandible. Merely for an immediate loading in the maxilla, a significantly lower implant loss rate was shown (0.08 [95% CI 0.02; 0.32] vs. 0.49 [95% CI 0.38; 0.62]; P = 0.0125).

Multivariate analysis

To explore the independent effects and interrelation between factors influencing the estimated implant loss rate, a multivariate Poisson regression model was fitted to the data of univariate meaningful factors. The first model included the location of implants, the type of prosthesis, the surface of implants, the loading protocol, and the number of implants per patient and all two-way interaction terms. Due to the sparse distribution of number of implants across the remaining factors, the Poisson regression model did not converge. Thus, the final model was reduced to the location of implants, the type of prosthesis, the surface of implants, and the loading protocol as main effects. Additionally, the significant two-way interaction between location and loading protocol remained in the model. Within this model, type of prosthesis (P < 0.0001 fixed vs. removable), surface of implants (P = 0.0001 machined vs. rough), and the interaction term between jaw and loading protocol (P = 0.0006) demonstrated significant influence on the estimated implant loss rate. From the significant interaction between jaw and loading protocol, a significant difference between conventional and immediate loading in the mandible (P < 0.0001) and between mandible and maxilla in the conventional loading protocol (P <0.0001) followed. The comparisons between conventional and immediate loading in the maxilla (P = 0.1745) and between mandible and maxilla in the immediate loading protocol (P = 0.9886) showed no significant differences (Table 13).

Bone augmentation

The analysis of a potential impact of bone augmentation on implant loss or survival was not a part of the focused question and serves as additional information.

Studies reporting on complex augmentative procedures (e.g., Le Fort I, onlay osteoplastic with iliac graft) were already described. Bone augmentation was reported for the maxilla, exclusively. Assuming that complex procedures would have been reported if executed, a comparison of post-loading implant loss per 100 implant years revealed a significantly higher rate for non-augmented (0.93 [95% CI 0.76152; 1.14; 22 study populations) vs. augmented (0.25 [95% CI 0.16; 0.40]; 4 study populations) edentulous maxillae (P < 0.0001). Corresponding 5-year implant survival estimations were 95.45% [95% CI 94.47; 96.26] for non-augmented and 98.75% [95% CI 98.00; 99.22] for augmented

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	Subaroups	Number of implants per	Total number	Tvpe of prosthesis	Number of post- loading implant	Total implant exposure time	Estimated implant loss (per 100	Estimated implant
Study	within study	patient	of implants*	and anchorage	losses	(implant years)	implant years)	survival after 5 years (%)
Cordioli et al.	n.a.	1	21	Removable, BL	0	95.0	0	100
Harder et al.	n.a.	-	11	Removable, BL	0	38.9	0	100
Liddelow et al.	Machined	, ,	7		с (12.4	24.2 ĉ	25.1
	Kough	(۲7 ۲		0 0	36.5 260.0	0 0	100
Akogiu et al. Cobroli of al	n.a.	ע ר	12	Removable, BL		0.005		100
Conner et al	.e.	40	118 118			557.0		100
Elsvad et al.	Immed. load.	1 0	36		0	0.96	0 0	100
	Convent. load.	- 7	36		4	92.3	ç 4.3	80.1
Gotfredsen & Holm	Bar	2	22		0	110.0	0	100
	Ball	2	30		0	148.0	0	100
Heijdenrijk et al.	1-stage	2	40		0	381.1	0	100
-	2-stage	2	80		2	192.0	1.0	94.9
Krennmair et al.	Talore crowne	א ר	97 VC	Removable, BL		122.0	5 0	100
Meijer et al		4 0	1 85		o	560.0	0 C U	99.1
	Brå	- 7	64	Removable, BR	4	536.0	0.7	96.3
Meijer et al.	Brå	2	60	Removable, BR	0	546.0	0	100
	IMZ	2	60	Removable, BR	ε	570.0	0.5	97.4
	ITI Str	2	60		0	562.0	0	100
Meijer et al.	2 implants	2	60	Removable, BR	m	1008.5	0.3	98.5
Weinländer et al.	2 implants	2	48	Removable, BR	0	216.0	0	100
Jemt et al.Watson et al.	n.a.	2 - 2	198	Removable, BR	7	880.0	0.8	96.1
Behneke et al.	n.a.	2-4	340		0 0	1530.0	0	100
Stoker et al.	n.a.	2-4	296	Removable, n.a.	л т	2124.8	0.4	9.79
Akca et al.	n.a.	2-4	80	Removable, BR	- 0	392.0 197.8	0.3	98. / 100
bergenaal & Engquist	Bar Ball	2-4 7-0	20	Removable, BK Removable BI		140.6		100
Romeo et al		P-C	84	Removable, br	о С	472.0	0.4	97.9
Chiapasco et al.	n.a.	4	328			1704.0	0.4	98.0
Engquist et al.	1-stage Brå/Str	. 4	120		× m	331.0	6.0	95.6
-	2-stage Brå/Str	4	120	Fixed, SC	0	347.0	0	100
	1-stage Brå 1-piece	4	88	Fixed, SC	-	243.0	0.4	98.0
	1-stage Brå Mk III	4	104		7	299.0	2.3	88.8
Heschl et al.	n.a.	4	156		,	745.0	0.1	99.3
Krennmair et al.	n.a.	4 •	204		0 0	768.0	0 0	100
Krennmair et al.	Talace crowing	4 <	104	Removable, BK	0 0	0.962	5 6	100
l oranzoni at al	Convent load	4 4	80			400.0		100
	Immed. load.	4	80		0 0	400.0	0 0	100
Meijer et al.	n.a.	4	120		0	586.3	0	100
Vroom et al.	n.a.	4	80	Removable, BR	0	852.0	0	100
Weinländer et al.	n.a.	4	204	e,	0	944.0	0	100
Crespi et al.	n.a.	4	80	Fixed, SC	2	273.6	0.7	96.4
Arvidson et al.	n.a.	4-6	618	Fixed, SC	0	2809.0	0	100
Arvidson et al.	n.a.	4	250	Fixed, SC	ε Ω	511.3	0.6	97.1
De Bruyn et al.	n.a.	-\^ 5	125	Fixed, SC	0	350.0	0	100
Ekelund et al./	n.a.	1>5	273	Fixed, SC	-	4835.0	0.02	99.9
Lindquist et al.		L		-	c		c	000
Eliasson et al.	n.a.	2 I	168 	Fixed, SC	0 0	756.0	0 0	100
Lethaus et al.	n.a.	U I	70	Removable, BR	0 0	286.0	0 0	100
Nuurpny et al. Schwarz et al	П.а. П.а.	Ω Υ	151	Fixed SC	0 -	0.020.0	0	95.1
Junivalz et al. Tectori et al		ר ני ער א	116	Fived SC	- 0	C.C.C.D.	0	1.00 8 No
restorr et al. Van de Velde et al.	n.a.	c/ −4	91	Fixed, SC	n m	302.4	- 1	95.1 95.1
		2		12/20/01	n	1.100	2	

Table 5. (continued)	nued)									
Study	Subgroups within study	ips tudy	Number of implants per patient	Total number of implants*	Type of prosthesis and anchorage	Number of post- loading implant losses	Total implant exposure time (implant years)	Estimated implant loss (per 100 implant years)	Estimated implant survival after 5 years (%)	ant years (%)
Örtorp et al. Rasmusson et al.	n.a. al. n.a.		∨ ∨ 5	373 108	Fixed, SC Fixed, SC	0 0	2200.0 1054.0	0 0	100 100	
BL, ball; BR, b	BL, ball; BR, bar; TC, telescopic crown; SC, screw-retained; n.a., not applicable.	; SC, screw-reta	ined; n.a., not a	pplicable.						
Table 6. Overall comparison Number o	ll comparison Number of study	Number of	Number of	Total number of post-loading	Total exposure time (implant	Estimated loss rate per 100 implant			Estimated 5-year implant	
	populations	patients	implants	implant losses	years)	years [95% Cl]	[95% CI]	survival (%) [95% Cl]	[95% CI]	<i>P</i> -value
Maxilla vs. mandible Maxilla 25 ³ Maxilla 20 [†] Mandible 56	andible 25* 20† 56	496 457 1872	2850 2717 6417	113 67 77	16779.08 16142.18 35486.20	0.67 [0.55; 0.80] 0.42 [0.33; 0.53] 0.22 [0.17; 0.27]	97.99 [97.60; 98.33] 98.76 [98.43; 99.02] 99.35 [99.19; 99.48]	.33 96.68 [96.03;97.24] .02 97.95 [97.40; 98.38] .48 98.93 [98.66; 99.14]		<0.0001 0.0001

	Number of study populations	Number of patients	Number of implants	I otal number of post-loading implant losses	l otal exposure time (implant years)	estimated loss rate per 100 implant years [95% Cl]	Estimated 3-year implant survival (%) [95% CI]	Estimated 5-year im survival (%) [95% CI
Maxilla vs. mandible	Idible							
Maxilla	25*	496	2850	113	16779.08	0.67 [0.55; 0.80]	97.99 [97.60; 98.33]	96.68 [96.03;97.24]
Maxilla	20†	457	2717	67	16142.18	0.42 [0.33; 0.53]	98.76 [98.43; 99.02]	97.95 [97.40; 98.38]
Mandible	56	1872	6417	77	35486.20	0.22 [0.17; 0.27]	99.35 [99.19; 99.48]	98.93 [98.66; 99.14]
Fixed vs. remov	Fixed vs. removable prostheses							
removable	51*	1383	3901	118	21281.16	0.55 [0.46; 0.66]	98.35 [98.03; 98.62]	97.27 [96.73; 97.71]
removable	46†	1354	3768	72	20644.26	0.35 [0.28; 0.44]	98.96 [98.69; 99.17]	98.27 [97.82; 98.63]
fixed	30	955	5306	72	30984.12	0.23 [0.18; 0.29]	99.31 [99.13; 99.45]	98.84 [98.55; 99.08]
*Category <4 i †Category <4 i	*Category <4 implants (maxilla) included. †Category <4 implants (maxilla) excluded.	ded. ded.						

0.0148 ≤0.0001 maxillae. Both groups were pooled for further analyses.

Risk of bias within and across studies

Table 2 shows the risk of bias for each study as identified by the respective SIGN checklist. According to the terms of SIGN, most of the included clinical cohort studies or RCTs were of an acceptable or high quality, meaning "some flaws in the study with an associated risk of bias" or little to no risk of bias. Selective reporting or publication bias cannot be completely ruled out, especially, as some of the studies were sponsored by dental companies or a foundation being associated with a dental company.

Discussion

Summary of evidence

The objective of this systematic review and meta-analysis was to attend to the focused question: Is there an impact of implant location (maxilla vs. mandible), implant number, type of prosthesis (fixed vs. removable) and/ or different anchorage systems on the implant loss rate concerning the implantprosthodontic rehabilitation of edentulous patients?

Furthermore, additional analyses were performed to reveal a potential influence of implant surface and loading protocols on the implant loss rate for edentulous jaws.

In summary, the data situation or rather availability of literature concerning the edentulous jaw is comparatively satisfactory. Altogether 54 studies could be included for statistical analysis, although admittedly the majority of clinical studies investigated the edentulous mandible. This fact has also been observed by other authors of systematic reviews over the last 7 years (Sadowsky 2007; Slot et al. 2009; Roccuzzo et al. 2012).

To attain a reasonable level of evidence, retrospective studies were excluded. The overall evidence for the included randomizedcontrolled and prospective studies can be rated acceptable. The majority of included studies had an evidence level of 2+, although it must be noted that the little number of included RCTs did not always directly address the focused question, meaning that a high level of evidence can be assumed for certain investigations, exclusively (e.g., different implant types were randomized). (Table 2).

In contrast to a previously published review regarding optimal implant numbers

Table 7. Overall con	Table 7. Overall comparison bar vs. ball vs. telescopic crown (category <4 implants excluded)	/s. telescopic cro	own (category	<4 implants exclude	(p				
	Number of study populations	Number of patients	Number of implants	Total number of post-loading implant losses	Total exposure time (implant years)	Estimated loss rate per 100 implant years [95% CI]	Estimated 3-year implant survival (%) [95% Cl]	Estimated 5-year implant survival (%) [95% CI]	<i>P</i> -value
Ball Bar	13 28	267 928	507 2777	7 54	2048.73 15494.73	0.34 [0.16; 0.72] 0.35 [0.27; 0.46]	98.98 [97.87; 99.51] 98.96 [98.64; 99.20]	98.31 [96.46; 99.20] 98.27 [97.73; 98.66]	0.9607
Telescopic crown Bar	3 28	47 928	164 2777	0 54	504.0 15494.73	Not estimable 0.35 [0.27; 0.46]	98.96 [98.64; 99.20]	98.27 [97.73; 98.66]	
Ball Telescopic crown	0 m	267 47	507 164	7 0	2048.73 504.0	0.34 [0.16; 0.72] Not estimable	98.98 [97.87; 99.51]	98.31 [96.46; 99.20]	
<i>Table 8</i> . Comparison in the maxilla	in the maxilla								
				+ 	Total among and a			Patients	

Table 8. Compai	Table 8. Comparison in the maxilla								
	Number of study populations	Number of patients	Number of implants	Number of post-loading implant losses	Total exposure time (implant years)	Estimated loss rate per 100 implant years [95% CI]	Estimated 3-year implant survival (%) [95% Cl]	Estimated 5-year implant survival (%) [95% CI]	<i>P</i> -value
Fixed vs. remov	Fixed vs. removable prostheses (category <4 implants excluded)	/ <4 implants exc	luded)						
removable	9	84	334	25	1082.10	2.31 [1.56; 3.42]	93.30 [98.91; 99.41]	89.09 [84.28; 92.50]	
fixed	14	363	2383	42	15060.08	0.28 [0.21; 0.38]	99.17 [98.87; 99.38]	98.62 [98.13; 98.97]	<0.0001
Removable: <4	Removable: <4 implants vs. 4 implants								
4	5	49	133	46	636.90	7.22 [5.41; 9.64]	80.52 [74.88; 85.02]	69.70 [61.75; 76.30]	
4	9	84	334	25	1082.10	2.31 [1.56; 3.42]	93.30 [90.25; 95.42]	89.09 [84.29; 92.50]	<0.0001
Fixed: ≥6 (no co	Fixed: 26 (no comparison feasible)								
9/1	11	303	2057	37	13099.78	0.28 [0.20; 0.39]	99.16 [98.84; 99.39]	98.60 [98.07; 98.98]	

for the completely edentulous maxilla (Schley & Wolfart 2011), the authors decided to perform a statistical analysis. Analyzing non-randomized, non-controlled studies raises a complex of problems and does not allow for a classical analysis in form of a forest plot that always intents to compare different intervention groups, that is, randomized-controlled trials. Furthermore, the inconsistent reporting of results among the studies complicates a meaningful analysis. The absence of exact information on implant/prosthesis loss or dropout and/or the absence of a mean observation period led to the exclusion of several articles. Hence, the authors adopted a frequently applied statistical method, suggested by Pjetursson et al. (2007) and Sailer et al. (2007) using the "total exposure time" of the investigated objects and estimating failure (or loss) and survival rates by Poisson regression. Recently, Pjetursson et al. (2014) applied the same method to describe the implant failure and the survival in a systematic review. Also, the present calculation of the "implant loss rate per 100 implant years" is based on the assumption of a constant event rate over time. The resulting "data distortion" is mainly caused by those studies with a very long or short observation period leading to an extrapolation or adaption of the available data, respectively. From a clinical point of view, this assumption is debatable; however, in the authors' opinion, currently, it is the best method to compare the results of the different clinical studies with each other. To provide full information, the actual implant losses and observation periods are given in Tables 3-5.

Considering the focused question, it can be stated that all of the mentioned factors (jaw, implant number, type of prosthesis and anchorage system) seem to have an impact on implant survival and implant loss. Generally, estimated implant survival was satisfactory for both, fixed and removable rehabilitation concepts.

The risk for implant loss per 100 implant years in the edentulous mandible is significantly lower than in the maxilla (0.22 [95% CI 0.17; 0.27] vs. 0.41 [95% CI 0.32; 0.52]; P = 0.0001). Regarding the direct comparisons of implant numbers in the mandible, higher numbers showed a clear tendency of resulting in lower implant loss rates. The therapeutic concept of one implant inserted into the midline symphysis in the edentulous lower jaw is an ongoing and intensively discussed topic. The present data of this concept are based on merely three studies and revealed a 5-year survival estimation of 92.1%, which is

<i>Table 9.</i> Compa	Table 9. Comparison in the mandible								
	Number of study populations	Number of patients	Number of implants	Total number of post-loading implant losses	Total exposure time (implant years)	Estimated loss rate per 100 implant years [95% CI]	Estimated 3-year implant survival (%) [95% Cl]	Estimated 5-year implant survival (%) [95% CI]	<i>P</i> -value
Fixed vs. remo	Fixed vs. removable prostheses	0007	1010	ţ					
Kemovable Fixed	40 16	1280 592	3494 2923	4/ 30	19562.16 15924.04	0.24 [0.18; 0.32] 0.19 [0.13; 0.27]	99.28 [99.05; 99.46] 99.44 [99.20; 99.61]	98.81 [98.41; 99.10] 99.06 [98.66; 99.34]	0.2980
Fixed: 4 implar	Fixed: 4 implants vs. 25implants								
4	. 9	189	762	16	2004.85	0.80 [0.49;1.30]	97.63 [96.17; 98.54]	96.10 [93.69; 97.59]	
5	ø	278	1452	11	10807.77	0.10 [0.06;0.18]	99.70 [99.45; 99.83]	99.49 [99.10; 99.72]	<0.0001
Removable: 1	Removable: 1 implant vs. 2 implants								
-	4	66	66	c	182.81	1.64 [0.53; 5.09]	95.20 [85.84; 98.42]	92.13 [77.53; 97.38]	
2	19	557	1134	24	7274.90	0.33 [0.22; 0.49]	99.02 [98.53; 99.34]	98.36 [97.58; 98.91]	0.0088
Removable: 2	Removable: 2 implants vs. 4 implants								
2	19	557	1134	24	7274.90	0.33 [0.22; 0.49]	99.02 [98.53; 99.34]	98.36 [97.58; 98.91]	
4	10	365	1366	8	6971.25	0.11 [0.06; 0.23]	99.66 [99.31; 99.82]	99.45 [98.86; 99.70]	0.0097
4 Implants: fixe	4 Implants: fixed vs. removable								
removable	10	365	1366	80	6971.25	0.11 [0.06; 0.23]	99.66 [99.31; 99.82]	99.43 [98.86; 99.70]	
fixed	9	189	762	16	2004.85	0.79 [0.49; 1.30]	97.63 [96.17; 98.54]	96.10 [93.69; 97.59]	<0.0001
Table 10. Comp	Table 10. Comparison bar vs. ball vs. telescopic crown for mandible	telescopic crov	n for mandible						
		-							
	Number of study populations	Number of patients	Number of implants	Total number of post-loading implant losses	Total exposure time (implant vears)	Estimated loss rate per 100 implant vears [95% CI]	Estimated 3-year implant survival (%) [95% CI]	Estimated 5-year implant survival (%) [95% CI]	P-value
					((
Ball Bar	13 24	257 864	467 2523	7 29	2048.73 14652.63	0.34 [0.16; 0.72] 0.20 [0.14; 0.28]	98.98 [97.87; 99.51] 99.41 [99.15; 99.59]	98.31 [96.46; 99.20] 99.01 [98.61; 99.30]	0.1499

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satisfactory. Nevertheless, implant loss rates for two and four implants with an overdenture were significantly lower, and data were predicated on 19 and 10 patient groups, respectively. The "gold-standard concept" of two implants with an overdenture seems to be consolidated by the analyses of this systematic review, regarding post-loading implant survival, exclusively.

[96.46; 99.20] [98.61; 99.30]

[97.87;

0.34 [0.16; 0.72] Not estimable

2048.73 384.0 14652.63

~ 0

<u>~</u> 24

Telescopic

crown

0.34

99.01 98.31

99.59] 99.51]

5;

、 . 66]

99.41 98.98

0.20 [0.14; 0.28] Not estimable

384.0

C 29

124 2523 467 124

37 364 257 37

2

⁻elescopic

crown

Only 21 studies could be included regarding the edentulous upper jaw, rendering extensive statistical comparisons difficult. However, the present analyses clearly indicate that at least four implants are needed in the edentulous maxilla, irrespective of the type of restoration. Less than four implants have been suggested not to be feasible for the edentulous maxilla in an experts' consensus conference and is not recommendable at the time being (Schley et al. 2013). This fact was proven by the present analysis that revealed unacceptable survival estimations after 5 years (69.7% [95% CI 61.75; 76.30]) and significantly higher implant loss rates per 100 implant years when compared to implant numbers of four and more (7.22 [95% CI 5.41; 9.64] vs. 2.31 [95% CI 1.56; 3.42]; P < 0.0001). Therefore, it was decided to merely include this group for an overall survival analysis, but to exclude it from further statistical evaluations and comparisons. No statistically significant differences for postloading implant loss could be assessed when comparing bar or ball anchorage. Estimated implant survival was very high for both attachment types (ball: 98.31% [95% CI 96.46; 99.20]; bar: 98.27 [95% CI 97.73; 98.66]). Telescopic crowns could not be evaluated, as the included number of studies was to low, and no implant losses had occurred after observation periods of 3 years. Furthermore, no statements can be made regarding cemented or adhesively luted fixed restorations as the systematic literature review did not reveal such studies. Considering the socalled all-on-4 concept, meaning four implants being restored with a fixed prosthesis, the existing literature provides sufficient evidence for the edentulous mandible. Crespi et al. (2012) also implemented this concept for the edentulous maxilla and reported an implant survival of 98.96% after 3 years. For obvious reasons, this one study could not be used for statistical comparisons. However, retrospective clinical studies demonstrate comparable results (Malo et al. 2011, 2012).

Additional subgroup analyses were conducted regarding the aspects implants surface (machined vs. rough) and different loading protocols. Different surface roughness values

	Number of study Number of populations patients	Number of patients	Number of implants	Total number of post- loading implant losses	Total exposure time (implant years)	Estimated loss rate per 100 implant years [95% CI]	Estimated 3-year implant survival (%) [95% CI]	Estimated 5-year implant survival (%)	<i>P</i> -value
Overall comparison	arison								
Machined	21	657	2888	87	21089.63	0.41 [0.33; 0.51]	98.77 [98.48; 99.00]	97.63 [97.09; 98.02]	
Rough	53	1662	6246	57	29123.75	0.20 [0.15; 0.25]	99.41 [99.24; 99.55]	99.03 [98.74; 99.25]	<0.0001
Fixed (maxilla	Fixed (maxilla and mandible)								
Machined	12	393	2240	49	17423.42	0.28 [0.21; 0.37]	99.16 [98.89; 99.36]	98.17 [97.67; 98.56]	
Rough	16	536	1714	23	12145.70	0.19 [0.13; 0.28]	99.43 [99.15; 99.62]	99.06 [98.59; 99.37]	0.1177
Removable (m	Removable (maxilla and mandible)								
Machined	6	264	648	38	3666.21	1.04 [0.75; 1.42]	96.94 [95.82; 97.76]	94.93 [93.15; 96.32]	
Rough	37	1059	3180	34	16978.05	0.20 [0.14; 0.28]	99.40 [99.16; 99.57]	99.01 [98.61; 99.30]	<0.0001
Maxilla (fixed	Maxilla (fixed and removable)								
Machined	7	163	1162	58	9377.40	0.62 [0.48; 0.80]	98.16 [97.63; 98.58]	96.95 [96.08; 97.64]	
Rough	12	241	1385	6	5984.78	0.15 [0.08; 0.29]	99.55 [99.14; 99.77]	99.25 [98.57; 99.61]	<0.0001
Mandible (fixe	Mandible (fixed and removable)								
Machined	13	451	1556	29	11712.23	0.25 [0.17; 0.36]	99.26 [98.94; 99.49]	98.76 [98.22; 99.15]	
Rough	41	1395	4730	48	23138.97	0.21 [0.16; 0.27]	99.38 [99.17; 99.53]	98.96 [98.63; 99.22]	0.4518

could not be distinguished, and any type of surface modification was summarized under "rough" implant surface.

Comparing immediate vs. conventional loading in general, the estimated implant loss rate was slightly, but still significantly higher for an immediate protocol (P = 0.0151). Regarding fixed restorations, exclusively, implant loss rates did not significantly differ concerning immediate vs. conventional loading (P = 0.1652). This is in accordance with Papaspyridakos et al. (2014) who recently reported estimated survival rates between 99.10% and 99.90% for immediate and conventional loading for fixed prostheses in the edentulous maxilla and mandible in a systematic review. Surprisingly, the present results for fixed reconstructions in the edentulous mandible indicate a better outcome for а conventional loading protocol (P < 0.0001) and a better outcome for an immediate loading protocol for maxilla in general (P = 0.0108). If removable overdentures are planned, a conventional loading protocol still seems to result in a superior outcome concerning post-loading implant losses per 100 implant years (0.32 [95% CI 0.25; 0.42] vs. 0.62 [95%CI 0.37; 1.05]; P = 0.0282). Also Schimmel et al. (2014) concluded in their systematic review that implant-supported overdentures tend to have lower 1-year implant failure rates after application of a conventional loading protocol when compared to an immediate loading protocol. Moreover, they stated a necessity for "well-designed research protocols", because they partly experienced contradicting findings in their review.

The superior results for rough implant surfaces in almost all of the subcategories were not surprising. Better osseointegration capabilities of rough-surfaced implants have been shown in the past (Cordioli et al. 2000; Wennerberg & Albrektsson 2010).

Regarding post-loading implant loss, the classical implant-prosthodontic rehabilitation concepts, that is bar- or ball-retained overdentures and screw-retained full-arch reconstructions, have shown an excellent outcome according to the present analyses. A certain number of implants seem to ensure a reliable outcome for implants with a fixed or a removable restoration. However, prosthesisrelated technical complications need to be taken into consideration, as well. Therefore, we plan to analyze technical complications and correlated complication-free rates for implant-supported prostheses, related to implant location and certain implant numbers, in another systematic review.

Limitations

The presented results have to be interpreted with the following limitations:

The estimated implant loss rates and survival estimations were mostly derived from non-comparative studies. Due to a lack of high-quality studies (i.e., RCTs), the currently best option of receiving meaningful results is to analyze the best available evidence (mostly single arm cohort studies). Our focus was on potentially influencing aspects such as implant number, loading protocol and different prosthodontic treatment options. Due to the high degree of separation, a statistical analysis considering all of the potential influencing factors simultaneously was not feasible. However, a multivariate Poisson regression model concerning the location of implants, the type of prosthesis, the surface of implants, and the loading protocol as main effects, was fitted to the data of univariate meaningful factors.

Due to the observational nature of the included studies, confounding of observable, as well as unobservable factors is an intrinsic limitation of our derived results. Of course, a future aim is to analyze which combination of the above-mentioned factors is decisive. and therefore, more well-designed RCTs are needed. However, in dentistry and especially in the field of implant dentistry, several aspects such as high treatment costs, long duration of treatment, and limiting inclusion criteria (edentulous patients not being satisfied with complete overdentures) render RCTs difficult at best. It has to be recognized that CONSORT and consequently PRISMA statements or the "Cochrane Handbook" are mainly intended for medical studies and do perfectly fit for study concepts such as placebo vs. active agent. If our analysis strictly adhered to these protocols, merely a few studies would have been included thus setting a limitation, as well. In the authors' opinion, the inclusion of 54 studies with 9267 implants should inherently partly compensate the methodological handicap and thus could represent the best available "compromise".

The primary outcome was post-loading implant loss and not implant failure or success, which clearly would be the more accurate approach. However, the inhomogeneous reporting of success and failure among the studies (if reported at all) did not permit a statistical evaluation of success or failure rates.

As a matter of fact, several important aspects could not be considered in the present analyses, setting a limitation to this review. For example, different implant parameters such as length, diameter, form Table 12. (a) Conventional loading vs. immediate loading (category <4 implants excluded). (b) Comparison of immediate loading vs. conventional loading for mandible/fixed and mandible/ removable

	Number of study	Number of	Number of	Total number of post-loading	Total exposure	Estimated loss rate	Estimated 3-vear implant	Estimated 5-vear implant	
	populations	patients	implants	implant losses	years)	years [95% CI]	survival (%) [95% CI]	survival (%) [95% CI]	<i>P</i> -value
(a)									
Overall comparison	F								
Conventional	56	1773	6835	66	42576.93	0.24 [0.19; 0.28]	99.30 [99.15; 99.43]	98.66 [98.41; 98.91]	
Immediate	14	361	1532	24	5936.87	0.40 [0.27; 0.60]	98.79 [98.21; 99.19]	97.99 [97.03; 98.65]	0.0151
Fixed (maxilla and mandible)	i mandible)								
Conventional	17	598	3512	41	24693.90	0.17 [0.12; 0.23]	99.50 [99.33; 99.63]	98.86 [98.51; 99.10]	
Immediate	6	209	1143	10	3691.64	0.27 [0.15; 0.50]	99.19 [98.50; 99.56]	98.65 [97.51; 99.27]	0.1652
Removable (maxilla and mandible)	a and mandible)								
Conventional	39	1175	3323	58	17883.03	0.32 [0.25; 0.42]	99.03 [98.75; 99.25]	98.41 [97.92; 98.76]	
Immediate	5	152	389	14	2245.23	0.62 [0.37; 1.05]	98.15 [96.89; 98.90]	96.95 [94.89; 98.17]	0.0282
Maxilla (fixed and removable)	removable)								
Conventional	14	296	1847	61	12566.00	0.49 [0.38; 0.62]	98.55 [98.15; 98.87]	97.60 [96.93; 98.13]	
Immediate	5	127	731	2	2688.18	0.08 [0.02; 0.32]	99.75 [99.04; 99.94]	99.60 [98.40; 99.90]	0.0125
Mandible (fixed and removable)	nd removable)								
Conventional	42	1477	4988	38	30010.93	0.13 [0.09; 0.17]	99.62 [99.48; 99.72]	99.35 [99.15; 99.55]	
Immediate	6	234	801	22	3455.69	0.64 [0.42; 0.97]	98.12 [97.14; 98.75]	96.87 [95.28; 97.93]	<0.0001
(q)									
Mandible/fixed									
Conventional	6	386	1999	5	13210.00	0.04 [0.02; 0.09]	99.89 [99.73; 99.95]	99.80 [99.55; 99.90]	
Immediate	4	82	412	4	1210.46	0.66 [0.33; 1.32]	98.04 [96.11; 99.01]	96.75 [93.61; 98.36]	<0.0001
Mandible/removable	ble								
Conventional	33	1091	2989	33	16800.93	0.20 [0.14; 0.28]	99.41 [99.17; 99.58]	99.01 [98.61; 99.30]	
Immediate	5	152	389	14	2245.23	0.62 [0.37; 1.05]	98.14 [96.89; 98.90]	96.93 [94.87; 98.17]	0.0003

(cylindric, root-like), implant-abutment connection, bone-to-implant interface, or the difference of one- or two-piece implants could not be assessed. Furthermore, studies investigating implants in either local or augmented bone (four studies) were pooled. However, the analysis of augmented bone in the maxilla did not reveal negative results concerning estimated implant loss and corresponding survival rates. The duration of edentulism as a potential confounder could not be regarded either, but, in most studies, the "typical" completely edentate patient was subject of the investigation.

The analysis of biologic complications was not part of our focused question and explains why these complications were not evaluated in detail.

It is self-evident that the "best" choice of an implant-prosthdontic restoration, cannot simply be based on the analyzed and aforementioned aspects. Individual, patient-based circumstances determine any surgical or prosthodontic procedures. In this context, it was not possible to regard important facts such as patients' preferences, esthetic complexity, maxillomandibular relationship, bone quality and quantity, soft tissue conditions, condition or type of restoration of the opposing jaw, or differences of treatment/manufacturing costs. Even though several authors gave information on the type of restoration in the opposite jaw (full denture, fixed or removable prosthesis), a conclusion, if implant outcome is affected by this factor, could not be evaluated.

Moreover, oral health-related quality of life (OHRQoL) is an omnipresent topic, and especially, the rehabilitation of the edentulous jaw by means of implant-prosthodontic procedures can offer a great potential of improving patients' quality of life (Turkyilmaz et al. 2010). For the edentulous maxilla, in particular, there is a huge backlog demand for studies on OHRQoL. In this respect, Zembic & Wismeijer (2014) recently published an interesting approach. Patients received conventional complete dentures in a first step, and 2 months later, two implants were inserted the implant-retained overdentures "provided some significant short-term improvements over conventional dentures in oral- and health-related quality of life".

Many of the aforementioned parameters demand for a consolidated internal evidence, meaning the dentist's experience, which serves as an important component of evidence-based medicine/dentistry. In combination with the external evidence (current state of science) and the patient's values and

Table 13. Estimates with corresponding standard errors and *P*-values resulting from the multivariate Poisson regression model

Factor/Interaction	Estimate	Standard Error	P-value
Intercept	-4.7007	0.7327	< 0.0001
Jaw	0.0108	0.7610	0.9886
Type of prosthesis	-2.4227	0.1921	< 0.0001
Surface of implant	0.7254	0.1913	0.0001
Loading protocol	1.0085	0.7427	0.1745
Jaw-loading protocol	-2.6806	0.7790	0.0006
Conventional vs. immediate (mandible)	-1.6721	0.2748	< 0.0001
Conventional vs. immediate (maxilla)	1.0085	0.7427	0.1745
Mandible vs. maxilla (conventional)	-2.6698	0.2048	< 0.0001
Mandible vs. maxilla (immediate)	-0.9976	0.2677	0.9886

wishes, a participatory decision-making process can be developed (Türp & Antes 2013). This procedure provides a reasonable degree of safety for both patient and dentist.

Conclusions

Considering the above-mentioned limitations, the following conclusions can be drawn:

- Only four of the included studies report on observation periods of more than 10 years.
- The current evaluations show a successful outcome for screw-retained fixed restorations and bar- or ball-retained overdentures in the completely edentulous jaw. Disregarding more than the included potential confounders (such as anatomic situation, bone quality, jaw relation, implant-related components) and relating to the estimated post-loading implant loss, exclusively, the following statements can be made:
- Maxilla:
 - (a) The insertion of six or more implants for a fixed reconstruction in the maxilla reveals favorable results. Considering the "all-on-4" concept for the maxilla, one study (Crespi et al. 2012) with an acceptable level of evidence was found, revealing a satisfactory outcome. For obvious reasons, this one study could not be used for a meaningful statistical comparison.
 - (b) The insertion of four implants for a removable overdenture in the maxilla reveals satisfying results. Data on minimal concepts with <4 implants in the maxilla is scarce and demonstrated significantly worse results,

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Askig Focused Questions (2014) CEBM Center for Evidence Based Medicine. www.cebm.net. Oxford: University of Oxford. calling for a cautious and controlled application of these therapeutic options.

- Mandible:
 - (a) The insertion of four implants for a fixed restoration in the edentulous mandible reveals satisfying results. However, it has to be noticed that five or more implants showed a slightly better outcome.
 - (b)The insertion of two implants for a removable overdenture in the mandible shows favorable results. However, it has to be noticed that four implants revealed a slightly better outcome. Furthermore, four implants with a removable prosthesis had a better outcome than four implants with a fixed prosthesis in the mandible. Data on the minimal concept with only 1 implant is scarce and shows promising results. However, the results are negatively influenced when using machined-surfaced implants and an immediate loading protocol (Liddelow & Henry 2010). The application of this therapeutic option can only be recommended, when the insertion of 2 or more implants is not feasible, e.g. due to economic reasons.
- In general:
- (a) Implants with fixed prostheses show slightly but significantly better results than removable prostheses regarding both jaws.
- (b) Rough-surfaced implants demonstrated favorable results compared to machined implants.

(c) In general, conventional loading tended to result in fewer implant losses. However, the implant loss rate for fixed prostheses in maxilla and mandible did not significantly differ concerning immediate and conventional loading. It has to be noted, though, that immediate loading was generally attached to strict conditions (e.g., a pre-defined insertion torque).

Future research

Consequential suggestions for future research: Future RCTs should investigate different attachment systems with different implant numbers, especially for 1 vs. 2 implants in the mandible and <4 implants in the maxilla. Furthermore, the comparison of 4 implants vs. >4 implants with a fixed prosthesis in the maxilla and mandible would be desirable.

General suggestions for future research: Clinical studies should not only concentrate on implant success rates but also on the patients' benefit with regard to quality of life, improvement of mastication abilities, hygiene capability, psychological aspects, and financial considerations.

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Conflict of interest

The authors declare that they have no conflict of interest related to this article.

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Supporting Information

Additional Supporting Information may be found in the online version of this article:

Appendix S1. PRISMA 2009 Checklist.