ORIGINAL ARTICLE



Retrospective analysis of complications in 190 mandibular resections and simultaneous reconstructions with free fibula flap, iliac crest flap or reconstruction plate: a comparative single centre study

Lucas M. Ritschl¹ · Thomas Mücke² · Diandra Hart¹ · Tobias Unterhuber¹ · Victoria Kehl³ · Klaus-Dietrich Wolff¹ · Andreas M. Fichter¹

Received: 19 May 2020 / Accepted: 25 September 2020 \odot The Author(s) 2020

Abstract

Objectives The purpose of this study was to evaluate the incidence of complications following mandibular reconstruction and to analyse possible contributing factors.

Materials and methods Clinical data and computed tomography scans of all patients who needed a mandibular reconstruction with a reconstruction plate, free fibula flap (FFF) or iliac crest (DCIA) flap between August 2010 and August 2015 were retrospectively analysed.

Results One hundred and ninety patients were enrolled, encompassing 77 reconstructions with reconstruction plate, 89 reconstructions with FFF and 24 reconstructions with DCIA flaps. Cutaneous perforation was most frequently detected in the plate subgroup within the early interval and overall (each p = 0.004). Low body mass index (BMI) and total radiation dosage were the most relevant risk factors for the development of analysed complications.

Conclusions Microvascular bone flaps have overall less skin perforation than reconstruction plates. BMI and expected total radiation dosage have to be respected in choice of reconstructive technique.

Clinical relevance A treatment algorithm for mandibular reconstructions on the basis of our results is presented.

Keywords Mandibular reconstruction · Free bone flaps · Free fibula flap · Complications · Radiation

Introduction

Mandibular reconstruction after partial or continuity resection is a complex procedure with non-negligible sequela on the

Electronic supplementary material The online version of this article (https://doi.org/10.1007/s00784-020-03607-8) contains supplementary material, which is available to authorized users.

Lucas M. Ritschl lucas.ritschl@tum.de

- ¹ Department of Oral and Maxillofacial Surgery, School of Medicine, Technical University of Munich, Klinikum rechts der Isar, Ismaninger Straße 22, 81675 Munich, Germany
- ² Department of Oral and Maxillofacial Surgery, Malteser Kliniken Rhein-Ruhr, Kurfürstenstraße 69, 47829 Krefeld-Uerdingen, Germany
- ³ School of Medicine, Institute of Medical Informatics, Statistics and Epidemiology, Technical University of Munich, Grillparzerstraße 18, 81675 Munich, Germany

oro-pharyngeal integrity and function, aesthetics and quality of life [1–3]. With continuous technical and surgical improvements, the restoration of mandibular continuity has become a highly standardized procedure in high-volume centres over the last decades. Microvascular tissue transplantation has evolved to a gold standard procedure in reconstructive cranio-maxillofacial surgery, especially since the introduction of the free fibula flap (FFF) for mandibular reconstruction by Hidalgo [4, 5]. Another milestone in the continuous development and improvement of mandibular reconstruction was the introduction of computer-aided design and computer-aided manufacturing (CAD/CAM) in this field. The application of the CAD/CAM technique increased the quality, function, safety and symmetry of this surgical procedure and reduced the operating time significantly [2, 6–8].

Microvascular bone transplants like the FFF or iliac crest (DCIA) flap are reported to be safe reconstructive possibilities with transplant-specific advantages and disadvantages. In general, they are associated with good long-term results in specialized centres and have a reliable consistency of vertical bone dimension in conventional panoramic radiographs [9, 10].

Nevertheless, mandibular reconstruction is also associated with complications, which might vary according to the reconstructive technique applied and the demanding patient cohort itself [11]. Postoperative complications after reconstructive microsurgery are generally associated with prolonged inhospital stay, increased costs and prolonged treatment in the outpatient setting. This potentially reduces the quality of life in the synopsis.

The purpose of this retrospective study was to evaluate short- and intermediate-term complications and to describe possible factors that might contribute to the unwanted clinical course.

Materials and methods

Ethical statement and patient recruitment

All clinical investigations and procedures were conducted according to the principles expressed in the Declaration of Helsinki. The retrospective analysis was approved by the Ethical Committee of the local government based on this informed consent to participate in this study and on the willingness to undergo the required medical care (Approval No. 87/ 17 S).

Any patient requiring a mandibulectomy and simultaneous reconstruction with a reconstruction plate (intraoperatively or preoperatively bend), a FFF (conventional free hand or CAD/CAM-assisted) or a DCIA flap because of a benign or malign process between August 2010 and August 2015 at our department was included in this retrospective study. Patients with a history of previous microvascular flap loss or reconstruction with a scapula free flap and patients receiving more than one flap (double free or sandwich flap reconstruction) in the same operation were excluded [12, 13]. No further exclusion criteria were applied.

Patient collective and study analyses

Patient records and datasets of all patients operated with a standardized scheme for mandibular continuity resection between August 2010 and August 2015 were screened for postoperative complications including revision of microvascular anastomosis and (partial) flap loss in microvascular free flap cases. Postoperative fistula, dehiscence, intraoral bone exposure, cutaneous perforation, screw loosening and plate fracture were registered in all enrolled cases. Fistula was defined as an extraoral, round wound with a diameter < 10 mm, not necessarily adjacent to the incision line/scar communicating with the oral cavity or with an abscess formation. Dehiscence was defined as an extraoral wound adjacent to the former incision line/scar with a longitudinal extent of > 10 mm.

The incidence of any complication was registered, including the time interval between operation and date of recorded incidence. We defined three clinical observation intervals: early (≤ 100 days) and late complication (> 100 days) as well as overall complication.

Potential contributing factors for the registered complications like age, gender, body mass index (BMI), ASA status, history of smoking, diabetes mellitus and history of chemoand radiation therapy were additionally recorded.

Statistical methods

The occurrence of complications was described by treatment group using absolute and relative frequencies. Hypothesis testing of differences between subgroups was performed by χ^2 or Fisher's exact test as necessary. Continuous variables were summarized by reconstruction type with mean and standard deviation and compared between groups using the Mann-Whitney U test.

All statistical tests were performed on an exploratory twosided 5% significance level. No adjustment for multiple testing was incorporated. Analysis was done with IBM SPSS 24 for Windows software (IBM Corp, Armonk, NY).

Results

Clinical outcome and descriptive results (Tables 1, 2 and Figure 1)

One hundred and ninety patients were enrolled for final analysis. The overall distribution of gender, age, indication, history of surgical intervention and radiation for the reconstructive subgroups is shown in Table 1. Mandibles were reconstructed with reconstruction plates in 77 cases, FFF in 89 cases and DCIA flap in 24 cases after continuity resection. In cases with reconstruction plates, 37 plates were bent intraoperatively, and 40 plates were individually pre-bend using an in-house printed mandible (ProJet 160 printer, VisiJet PXLCore Eco and Binder; 4D Concepts GmbH, Gross-Gerau, Germany) as described by others [14]. In the case of FFF, 63 cases were reconstructed conventionally by free hand, and 26 reconstructions were performed using the CAD/CAM technique as described elsewhere [15, 16].

Median overall follow-up time was 1728 days (8–3016) for reconstructions plate, 1794 days (23–3060) for FFF and 2017 days (1070–2792) for DCIA. More precisely it was 1728 days (359–3016) for intraoperatively bend and 1726 days (8–2973) pre-bend plates and 2039 days (23–3060) for conventional FFF and 1407 days (181–2015) for CAD/CAM FFF.

	Rec. plate intraop. bend ($n = 37$)	Rec. plate pre-bend (<i>n</i> = 40)	FFF conventional ($n = 63$)	FFF CAD/CAM ($n = 26$)	DCIA flap (<i>n</i> = 24)
Gender male (%)	21 (56.8)	25 (62.5)	41 (65.1)	18 (69.2)	13 (54.2)
Age median (range)	62.5 (43-86)	66.0 (14–93)	60.0 (27-76)	56.0 (18-76)	53.5 (24–71)
Malignancy yes (%)	29 (78.4)	29 (72.5)	27 (42.9)	10 (38.5)	2 (8.3)
1 st surgical intervention Yes (%)	25 (67.6)	30 (75.0)	33 (52.4)	11 (42.3)	15 (62.5)
History of radiation Yes (%)	13 (35.1)	9 (22.5)	40 (63.5)	13 (50.0)	2 (8.3)
Adjuvant radiation Yes (%)	18 (48.6)	17 (42.5)	13 (20.6)	5 (19.2)	2 (8.3)

 Table 1
 Overview of enrolled patients and their reconstruction technique with regard to gender, indication, history of surgical therapy and radiation therapy

Rec. reconstruction, *intraop.* intraoperatively, *FFF* free fibula flap, *CAD/CAM* computer-aided design and computer-aided manufacturing, *DCIA* deep circumflex iliac artery

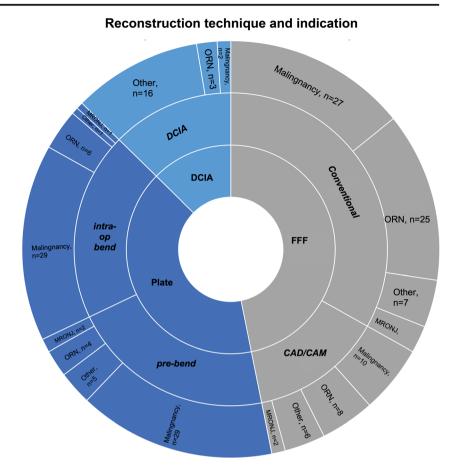
Incidence and distribution of all registered complications are listed in Table 2. Ten anastomoses had to be revised in the FFF reconstructed cases (six cases conventional FFF and four CAD/CAM FFF), which resulted in four total flap losses (two cases each in the conventional and CAD/CAM FFF groups) and four partial losses of the skin paddle (three cases conventional FFF and one CAD/CAM FFF) in total. Overall, the median time of revision of failing FFF was 5 days (range 1–

Complication	Rec. plate intraop. bend n (%)	Rec. plate pre-bend n (%)	FFF conventional <i>n</i> (%)	FFF CAD/CAM n (%)	DCIA flap <i>n</i> (%)	<i>p</i> value Fisher exact test		
Revision anastomosis \leq 30 d	/	/	6 (9.5)	4 (15.4)	0 (0.0)	0.137		
Flap failure ≤ 30 d	/	/	2 (3.2)	2 (7.7)	0 (0.0)	0.502		
Partial flap loss ≤ 30 d	/	/	3 (4.8)	1 (3.8)	0 (0.0)	1.000		
Fistula overall	6 (16.2)	6 (15.0)	14 (22.2)	4 (15.4)	4 (16.7)	0.920		
Fistula ≤ 100 th d	3 (8.1)	4 (10.0)	3 (4.8)	1 (3.8)	2 (8.3)	0.791		
Fistula > 100th d	3 (8.1)	2 (5.0)	11 (17.5)	3 (11.5)	2 (8.3)	0.401		
Dehiscence overall	10 (27.0)	8 (20.0)	13 (20.6)	3 (11.5)	2 (8.3)	0.378		
Dehiscence ≤ 100 th d	8 (21.6)	5 (12.5)	9 (14.3)	2 (7.7)	0 (0.0)	0.121		
Dehiscence > 100th d	2 (5.4)	1 (2.5)	2 (3.2)	1 (3.8)	1 (4.2)	0.955		
Io bone exposure overall	3 (8.1)	2 (5.0)	5 (7.9)	1 (3.8)	1 (4.2)	0.938		
Io bone exposure ≤ 100 th d	1 (2.7)	0 (0.0)	2 (3.2)	1 (3.8)	0 (0.0)	0.818		
Io bone exposure > 100 th d	2 (5.4)	2 (5.0)	4 (6.3)	0 (0.0)	1 (4.2)	0.854		
Cutaneous perforation overall	11 (29.7)	3 (7.5)	4 (6.3)	1 (3.8)	1 (4.2)	0.004*		
Cutaneous perforation \leq 100th d	5 (13.5)	1 (2.5)	0 (0.0)	0 (0.0)	0 (0.0)	0.004*		
Cutaneous perforation > 100th d	5 (13.5)	2 (5.0)	4 (6.3)	1 (3.8)	1 (4.2)	0.617		
Screw loosening overall	2 (5.4)	2 (5.0)	2 (3.2)	2 (7.7)	2 (8.3)	0.566		
Screw loosening ≤ 100 th d	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	1 (4.2)	0.004*		
Screw loosening > 100th d	2 (5.4)	2 (5.0)	2 (3.2)	2 (7.7)	1 (4.2)	0.972		
Plate fracture overall	0 (0.0)	2 (5.0)	3 (4.8)	4 (15.4)	0 (0.0)	0.067		
Plate fracture ≤ 100 th d	0 (0.0)	0 (0.0)	1 (1.6)	2 (7.7)	0 (0.0)	0.172		
Plate fracture > 100th d	0 (0.0)	2 (5.0)	2 (3.2)	2 (7.7)	0 (0.0)	0.371		

d days, Rec. reconstruction, intraoperatively, FFF free fibula flap, DCIA flap deep circumflex iliac artery, CAD/CAM computer-aided design/ computer-aided manufacturing, Io intraoral

*Significant at the 5% level

Fig. 1 Sunburst diagram with the distribution of reconstruction techniques according to the underlying indication for mandibular reconstruction (FFF, free fibula flap; DCIA, deep circumflex iliac artery; CAD/ CAM, computer-aided design and computer-aided manufacturing; ORN, osteoradionecrosis; MRONJ, medication-related necrosis of the jaw)



21), and final flap loss was seen at 17 days (6–29). No DCIA flap required revision and no flap failure occurred within the retrospective observation time.

Postoperative fistula occurred four times in the reconstruction plate group, seven time in the plate, four times in the FFF and twice in the DCIA reconstructed cases within 100 days (p = 0.791). Within the late interval (> 100 days), fistula occurred five times in the plate group, 14 times the FFF group and twice in the DCIA group (p = 0.401). Dehiscence was registered 13 times in the plate group, eleven times in the FFF group and never in the DCIA group within 100 days (p = 0.121). Within the late interval, dehiscence occurred three times in the plate group, three times in the FFF group and once in the DCIA group (p =0.955). Intraoral bone exposure occurred once each in the plate group and four times in the FFF group within 100 days (p = 0.818) and four times in the plate group and four times in the FFF group in the late interval (p = 0.854). Cutaneous perforation within 100 days was only seen in the plate group six times (p = 0.004). In the late interval, it was registered seven in the plate group, five times in the FFF group and once in the DCIA group (p = 0.617). Screw loosening was only seen once in the DCIA group within 100 days (p = 0.004). In the late interval, it became evident each four times in the plate and FFF groups and once in the DCIA group (p = 0.972). Plate fracture was only seen in the FFF group three times within 100 postoperative days (p = 0.172). In the late interval, it was registered twice in the plate group, four times in the FFF group and never in the DCIA group (p = 0.371).

Detailed results for the incidences of postoperative fistula, dehiscence and cutaneous perforation formation within the subgroups with regard to all analysed possible confounding factors are shown in the Supplementary Tables S1–3.

The incidences of postoperative fistula, dehiscence and cutaneous perforation within the subgroups with regard to the most relevant possible confounding factors on the basis of the Supplementary Tables S1–3 are summarized in Table 3 (age, BMI and history of radiation represented as "TotalGy"). Overall, history of radiation with the total amount of gray (Gy) was significantly associated with all complications. Within the differentiation of the subgroups, the mandibular reconstruction technique with intraoperatively bend reconstruction plates showed a strong association between the history of radiation and the incidence of fistula formation.

A low BMI significantly more often led to postoperative dehiscence when intraoperatively bend reconstruction plates were used ($18.1 \pm 2.5 \text{ kg/m}^2$; p = 0.006). A low BMI was also associated with cutaneous perforation within the CAD/CAM FFF subgroup ($18.2 \pm 1.5 \text{ kg/m}^2$; p = 0.004).

			Postoperative fistula					Postoperative dehiscence					Postoperative cutaneous perforation				
			No		Yes			No		Yes			No		Yes		
			Mean	± SD	Mean	± SD	<i>p</i> value	Mean	± SD	Mean	±SD	<i>p</i> value	Mean	± SD	Mean	± SD	<i>p</i> value
Reconstruction technique	Rec. plate intraop. bend	Age	63	12	68	10	0.408	64	12	59	10	0.363	65	12	59	11	0.247
		BMI	22.7	5.5	23.6	3.4	0.733	23.8	5.0	18.1	2.9	0.006	23.3	5.5	22.3	4.0	0.615
		TotalGy	46.1	30.8	89.5	46.3	0.058	51.8	37.8	60.8	34.0	0.664	49.0	39.5	63.7	29.4	0.255
	Rec. plate pre-bend	Age	65	15	68	8	0.676	65	15	66	9	0.764	65	14	65	10	0.712
		BMI	25.1	5.7	21.8	2.5	0.110	24.2	5.5	25.6	5.2	0.536	24.4	5.4	25.0	5.8	1.000
		TotalGy	36.8	34.8	45.3	31.5	0.326	35.2	35.4	44.5	31.0	0.245	38.9	34.1	32.7	37.7	1.000
	FFF conv.	Age	60	11	57	12	0.339	60	11	56	11	0.174	59	11	56	4	0.253
		BM <i>I</i>	23.1	5.1	22.1	2.5	0.528	22.7	4.8	23.5	3.8	0.356	23.2	4.6	18.2	1.5	0.004
		TotalGy	45.9	32.0	57.1	28.5	0.554	46.8	30.4	57.9	35.3	0.244	47.6	32.1	65.1	4.3	0.118
	FFF CAD/CAM	Age	53	14	55	1	0.886	53	14	50	14	0.490	52	13	65		
		BMI	22.8	2.7	27.6	10.0	0.745	23.0	3.7	23.9	2.8	0.490	23.0	3.6	25.3		
		TotalGy	42.9	31.3	66.2	3.1	0.345	44.8	30.7	43.8	38.0	1.000	43.9	31.1	64.0		
	DCIA	Age	50	16	56	12	0.627	51	15	52			51	15	52		
		BMI	26.1	5.1	28.0	3.6	0.477	26.6	4.8	22.1			26.6	4.8	22.1		
		TotalGy	6.7	20.6	16.0	32.0	0.682	5.8	19.3	64.0			5.8	19.3	64.0		
	Total	Age	59	14	60	12	0.630	59	14	59	11	0.399	59	14	60	9	0.589
		BMI	23.8	5.1	23.3	3.8	0.503	23.8	5.0	23.2	4.7	0.504	23.9	4.9	22.2	4.4	0.082
		TotalGy	38.4	33.3	56.1	36.8	0.043	39.3	34.6	53.0	32.4	0.043	39.7	34.8	57.5	28.2	0.018

Table 3 Mean and standard deviation (SD) of postoperative fistula, dehiscence and cutaneous perforation

Rec. reconstruction, intraop. intraoperatively, FFF free fibula flap, conv. conventional, DCIA flap deep circumflex iliac artery, CAD/CAM computeraided design/computer-aided manufacturing, BMI body mass index, Gy gray

Distinguishing between the timing of radiation therapy, a positive history of radiation therapy was significantly associated with wound healing disturbances like postoperative fistula or dehiscence formation (p = 0.009, Table S1, and p = 0.027, Table S2). Cutaneous perforation occurred more frequently in cases with intraoperatively bend reconstruction plates and conventional FFF reconstructions compared with pre-bend plates and CAD/CAM FFF in the Fisher exact tests for difference in frequency of this complication between reconstruction type within the subgroups (p = 0.055, Table S3) in cases with a positive history of radiation therapy.

Discussion

This retrospective study investigated the most common postoperative complications after mandibulectomy and simultaneous reconstruction using only reconstruction plates, free fibula flaps or DCIA flaps in 190 cases. Potential risk factors for the occurrence of postoperative complications were analysed.

Frequency and nature of complications observed

The overall incidence of complications after mandibular reconstruction varies between 8 and 57% in the literature [11, 17-19]. Lee et al. described fistula formation (8.4%), hardware plate exposure (7.1%) and flap wound infections (6.5%)to be the most common complications. Our complication rates are higher for all mentioned complications, except for flap wound infection. This might be attributed to somewhat different definitions of complications or a differing complexity of the studied cohorts. As stated earlier, our understanding and precise definition of fistula and dehiscence might be different to the definitions authors have used to describe their woundhealing disorders [18, 20, 21]. Infection of the flap (wound) is rarely seen in daily routine at our department, which might be attributed to an established protocol for the intra- and postoperative administration of intravenous antibiotic therapy and the common standards of hygiene [22].

Plate fracture after mandibular reconstruction has been reported in the literature. Rendenbach et al. analysed the stability and strength of miniplates, common reconstruction plates and CAD/CAM plates in an in vitro study. They demonstrated no significant difference up to a load of 300 N [23]. There exists a diversity of results regarding stability, associated complications and costs of used fixation plates for the FFF and DCIA flap [24–30]. The bottom line of most manuscripts on this topic is that miniplate osteosynthesis remains the fixation system of choice for the FFF due to a low rate of associated complications and its good clinical handling. Cost-effective analyses and long-term results with an adequate scientific quality of patient-specific, laser-sintered titanium plates need to be conducted and are expected in the future [31].

Impact of the choice of reconstructive technique on the occurrence of complications

Markiewicz et al. described an overall high success rate of 94.8% of free flap reconstruction for the mandible using the FFF, the DCIA flap, the osteocutaneous radial forearm flap and the scapula flap. No significant difference in failure was found comparing FFF and DCIA in their meta-analysis encompassing 17 studies [1].

On the other hand, Lodders et al. described, for example, a higher complication rate in FFF compared with other free flaps [18]. This is in concordance with our results, where we registered more fistula and dehiscence formation in comparison with the FFF group.

Further, the extent of mandibular reconstruction was also identified as a relative risk factor in a literature review by Sadr-Eshkevari and colleagues. In their study, crossing the midline seemed to favour plate exposure, a phenomenon that has also been described by others [32–34]. Plate exposure is a severe complication that necessitates a critical review of the treatment plan, and a new operation including plate removal and secondary reconstruction [35]. Wei et al. reported a plate exposure rate of 46.15% in a retrospective analysis of patients who had been reconstructed with a reconstruction plate and soft tissue free flap [36]. Their high incidence of plate exposure was probably due to patient selection, including only patients with recurrent carcinomas and stage IV tumours. In the authors' experience, in cases of mandibular reconstruction (Fig. 1) using the reconstruction plate, the refixation of the temporarily released supramyohyoidal muscles is important from the perspective of soft tissue in addition to functional considerations. Further, the plate design with a suggested retrognathic position and a counterbored plate position on the remaining mandible might release some of the soft tissue tension over the reconstruction plate and reduce the dead space below the plate [33]. Pre-bending the plate reduces operation time and increases the accuracy of the reconstructive result, as shown by others [37, 38]. Despite following all these known and described suggestions, both plate groups (pre-bend plate and intraoperatively bend plate) had high rates of complications, with the lower incidence in the pre-bend plate subgroup.

Impact of total radiation dosage on the occurrence of complications

The highest overall recipient complication rate, though, was observed after the use of reconstruction plates. However, this observation might also be attributable to selection bias in this group, as seen in the study by Wei et al. [36]. The plate group had the highest number of malignancies and, if required in accordance with the current guidelines [39], consecutive adjuvant radiation therapy. Our FFF group had the highest number of cases of osteoradionecrosis (ORN). Both underlying indication and clinical situation favour wound-healing disturbances in the head and neck area as reported in the literature [40, 41]. In our resent study, postoperative fistula formation and development of dehiscence were more frequently seen in patients with a positive history of radiation therapy (p = 0.009and p = 0.027, respectively), rather than in the study population with adjuvant radiation therapy (p = 0.441 and p = 0.670, respectively) (Tables S1 and 2). On the other hand, cutaneous perforation was not significantly influenced with regard to the timing of radiation therapy and reconstruction technique (Table S3). In this context, a positive history of radiation therapy is a known risk factor for complications [42, 43] that has led us to adapt the concept of double free flap reconstructions [12, 44]. In a recent systematic review and meta-analysis, Mijiti et al. demonstrated an odds ratio of 1.90 that suggests a significantly increased rate for fistula formation [45]. On the other hand, the development of wound dehiscence or plate exposure was not significantly associated with a positive history of radiation. But only a minority of included studies have addressed these complications.

Overall, these unfavourable clinical situations and above-mentioned complications can be caused by secondary tissue and vascular fibrosis, altered extracellular matrix remodelling and changes in vasculature and local host defence peptides [46–48]. In accordance with this assumption, Lonie et al. described a higher recipient site complication rate for the DCIA flap compared with the FFF in a comparative literature review [49]. In their included studies, the incidence of radiotherapy was higher in the DCIA group, which is the opposite of our cohort, where the DCIA flap was almost exclusively applied for mandibular reconstructions of benign processes (Table 1).

This might lead to a key feature that has to be considered when it comes to reconstruction in irradiated head and neck areas: tensionless wound closure with robust free flap selection. An estimation of potential risk factors that favour the incidence of complications, such as fistula/ dehiscence formation or cutaneous perforation, must be addressed in a thorough preoperative, clinical examination of the patient.

Patient constitution and behaviour-related risk factors

The analysis of risk factors that contribute to the incidence of postoperative complications is difficult. Most studies analysed a heterogeneous cohort with a varying number of cases and microvascular free flaps. Overall, several risk factors have already been assumed for microvascular free flap transfer, including age, tobacco use, comorbidity, previous flap loss, ASA status and operating time [50–53].

In this study, low BMI and total amount of radiation dosage (Gy) emerged to be the most relevant risk factors (Table 3). Lo et al. also described an increased recipient site complication rate in patients with low BMI [19]. This is probably due to a decreased amount of subcutaneous fat tissue and muscle layers that could buffer the permanent mechanical stress during mandibular movements. An increased infection rate in obese patients with high BMI could not be described by Khan et al. [54].

ASA status showed a significant influence only on fistula formation (p = 0.010; Supplementary Table S1). As described previously and by others, ASA status is also a risk factor for flap loss [53]. However, this finding must be interpreted with caution, since this parameter reflects the constitution of the patients and therefore is biased by other underlying diseases that might also contribute to flap loss or complications.

Smoking and diabetes mellitus had no significant influence on fistula or dehiscence formation (Supplementary Tables S1– 3). Smoking, however, had a strong trend towards the development of postoperative cutaneous perforation (Supplementary table S3). Within the limits of this study, one could again argue that a potential selection bias has contributed to the coincidence of cutaneous perforation in the plate and FFF reconstruction subgroups, as smoking remains one of the main risk factors for the development of oral squamous cell carcinoma (OSCC).

Limitations

The presented study is a retrospective study with limitations that go along with this kind of study type. The provided clinical data has a potential for variability in reports. The authors attempted to minimize bias through definitions and data collection by a single researcher (DH). Secondly, the enrolled patients might not be representative of the entire population, although the study cohort reflects the regular distribution as reported in the literature. Thirdly, this study focused on the most common complications in the head and neck area after mandibular reconstruction and excluded donor site comorbidity, which might be associated with flap harvesting, as reported by others.

Treatment algorithm

Despite their different complication profiles, all techniques used in this study have a scope of application, and the choice

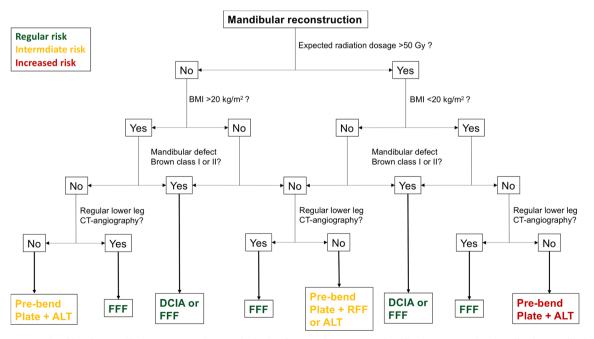


Fig. 2 Treatment algorithm for mandibular reconstructions on the basis of our results. Brown's classification was used to describe the mandibular defect situation [55]

of reconstruction method should be made for each individual patient. Within the limitations of this retrospective study and the experience of the authors, the different complication profiles and potential risk factors have helped us develop a treatment algorithm that can help in the decision-making process, which is illustrated in Fig. 2. This algorithm incorporates the most relevant risk factors (radiation dosage and patient BMI) and the expected mandibular defect as defined by Brown et al. [55].

Conclusions

Mandibular reconstruction is a safe procedure within the limits of underlying indication and applied reconstructive technique. Age, low BMI and total amount of radiation dosage (Gy) seem to have a significant impact on postoperative fistula, dehiscence and cutaneous perforation formation. These risk factors should be kept in mind in reconstructive technique selection. CAD/CAM FFF and preoperatively bend reconstruction plates had fewer complications than the corresponding conventional reconstructive technique. Free bone transplants offer the best functional and anatomical reconstruction option and have a lower risk of recipient complications than reconstruction plates. This reconstructive technique should therefore be applied whenever possible.

Funding Open Access funding enabled and organized by Projekt DEAL.

Compliance with ethical standards

Conflict of interest Author LMR declares that he has no conflict of interest. Author TM declares that he has no conflict of interest. Author DH declares that he has no conflict of interest. Author TU declares that he has no conflict of interest. Author VK declares that he has no conflict of interest. Author KDW declares that he has no conflict of interest. Author AMF declares that he has no conflict of interest.

Ethical approval All procedures performed in studies involving human participants were in accordance with the ethical standards of the institutional and national research committee and with the 1964 Helsinki declaration and its later amendments or comparable ethical standards.

Informed consent For this type of study, formal consent is not required.

Abbreviations ASA, American Society of Anaesthesiologists status; BMI, body mass index; CAD/CAM, computer-aided design and computer-aided manufacturing; CT, computed tomography; DCIA, free iliac crest flap; FFF, free fibula flap; Gy, gray; MRONJ, medicationrelated necrosis of the jaw; OSCC, oral squamous cell carcinoma; ORN, osteoradionecrosis

Open Access This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit http://creativecommons.org/licenses/by/4.0/.

References

- Markiewicz MR, Bell RB, Bui TG, Dierks EJ, Ruiz R, Gelesko S, Pirgousis P, Fernandes R (2015) Survival of microvascular free flaps in mandibular reconstruction: a systematic review and metaanalysis. Microsurgery 35:576–587. https://doi.org/10.1002/micr. 22471
- Ritschl LM, Mücke T, Fichter AM, Roth M, Kaltenhauser C, Pho Duc JM, Kesting MR, Wolff K-D, Loeffelbein DJ (2017b) Axiographic results of CAD/CAM-assisted microvascular, fibular free flap reconstruction of the mandible: a prospective study of 21 consecutive cases. J Craniomaxillofac Surg 45:113–119. https:// doi.org/10.1016/j.jcms.2016.11.001
- Wetzels JW, Merkx MA, de Haan AF, Koole R, Speksnijder CM (2014) Maximum mouth opening and trismus in 143 patients treated for oral cancer: a 1-year prospective study. Head Neck 36:1754– 1762. https://doi.org/10.1002/hed.23534
- 4. Hidalgo DA (1989) Fibula free flap: a new method of mandible reconstruction. Plast Reconstr Surg 84:71–79
- Disa JJ, Cordeiro PG (2000) Mandible reconstruction with microvascular surgery. Semin Surg Oncol 19:226–234
- Tarsitano A, Ciocca L, Scotti R, Marchetti C (2016) Morphological results of customized microvascular mandibular reconstruction: a comparative study. J Craniomaxillofac Surg 44:697–702. https:// doi.org/10.1016/j.jcms.2016.03.007
- Weitz J, Bauer FJ, Hapfelmeier A, Rohleder NH, Wolff KD, Kesting MR (2016) Accuracy of mandibular reconstruction by three-dimensional guided vascularised fibular free flap after segmental mandibulectomy. Br J Oral Maxillofac Surg 54:506–510. https://doi.org/10.1016/j.bjoms.2016.01.029
- Sieira Gil R, Roig AM, Obispo CA, Morla A, Pages CM, Perez JL (2015) Surgical planning and microvascular reconstruction of the mandible with a fibular flap using computer-aided design, rapid prototype modelling, and precontoured titanium reconstruction plates: a prospective study. Br J Oral Maxillofac Surg 53:49–53. https://doi.org/10.1016/j.bjoms.2014.09.015
- Hölzle F, Watola A, Kesting MR, Nolte D, Wolff K-D (2007) Atrophy of free fibular grafts after mandibular reconstruction. Plast Reconstr Surg 119:151–156. https://doi.org/10.1097/01.prs. 0000240703.02620.24
- Wilkman T, Apajalahti S, Wilkman E, Tornwall J, Lassus P (2017) A comparison of bone resorption over time: an analysis of the free scapular, iliac crest, and fibular microvascular flaps in mandibular reconstruction. J Oral Maxillofac Surg 75:616–621. https://doi.org/ 10.1016/j.joms.2016.09.009
- Lee M, Chin RY, Eslick GD, Sritharan N, Paramaesvaran S (2015) Outcomes of microvascular free flap reconstruction for mandibular osteoradionecrosis: a systematic review. J Craniomaxillofac Surg 43:2026–2033. https://doi.org/10.1016/j.jcms.2015.03.006
- Weitz J, Kreutzer K, Bauer FJ, Wolff KD, Nobis CP, Kesting MR (2015) Sandwich flaps as a feasible solution for the management of huge mandibular composite tissue defects. J Craniomaxillofac Surg 43:1769–1775. https://doi.org/10.1016/j.jcms.2015.07.038
- 13. Wei FC, Demirkan F, Chen HC, Chen IH (1999) Double free flaps in reconstruction of extensive composite mandibular defects in head

and neck cancer. Plast Reconstr Surg 103:39–47. https://doi.org/10. 1097/00006534-199901000-00008

- Lethaus B, Poort L, Bockmann R, Smeets R, Tolba R, Kessler P (2012) Additive manufacturing for microvascular reconstruction of the mandible in 20 patients. J Craniomaxillofac Surg 40:43–46. https://doi.org/10.1016/j.jcms.2011.01.007
- Wolff KD, Hölzle F, Kolk A, Hohlweg-Majert B, Steiner T, Kesting MR (2011) Raising the osteocutaneous fibular flap for oral reconstruction with reduced tissue alteration. J Oral Maxillofac Surg 69:e260–e267. https://doi.org/10.1016/j.joms.2010.11.040
- Tarsitano A, Del Corso G, Ciocca L, Scotti R, Marchetti C (2015) Mandibular reconstructions using computer-aided design/ computer-aided manufacturing: a systematic review of a defectbased reconstructive algorithm. J Craniomaxillofac Surg 43:1785– 1791. https://doi.org/10.1016/j.jcms.2015.08.006
- Wu CC, Lin PY, Chew KY, Kuo YR (2014) Free tissue transfers in head and neck reconstruction: complications, outcomes and strategies for management of flap failure: analysis of 2019 flaps in single institute. Microsurgery 34:339–344. https://doi.org/10.1002/micr. 22212
- Lodders JN, Schulten EA, de Visscher JG, Forouzanfar T, Karagozoglu KH (2016) Complications and risk after mandibular reconstruction with fibular free flaps in patients with oral squamous cell carcinoma: a retrospective cohort study. J Reconstr Microsurg 32:455–463. https://doi.org/10.1055/s-0036-1571794
- Lo SL, Yen YH, Lee PJ, Liu CC, Pu CM (2017) Factors influencing postoperative complications in reconstructive microsurgery for head and neck cancer. J Oral Maxillofac Surg 75:867–873. https://doi.org/10.1016/j.joms.2016.09.025
- Ritschl LM, Mücke T, Fichter A, Güll FD, Schmid C, Duc JMP, Kesting MR, Wolff K-D, Loeffelbein DJ (2017a) Functional outcome of CAD/CAM-assisted versus conventional microvascular, fibular free flap reconstruction of the mandible: a retrospective study of 30 cases. J Reconstr Microsurg 33:281–291. https://doi. org/10.1055/s-0036-1597823
- Chen SH, Chen HC, Horng SY, Tai HC, Hsieh JH, Yeong EK, Cheng NC, Hsieh TM, Chien HF, Tang YB (2014) Reconstruction for osteoradionecrosis of the mandible: superiority of free iliac bone flap to fibula flap in postoperative infection and healing. Ann Plast Surg 73(Suppl 1):S18–S26. https://doi.org/10. 1097/SAP.00000000000270
- Mücke T, Rohleder NH, Rau A, Ritschl LM, Kesting M, Wolff K-D, Mitchell DA, Loeffelbein DJ (2015) The value of perioperative antibiotics on the success of oral free flap reconstructions. Microsurgery 35:507–511. https://doi.org/10.1002/micr.22470
- Rendenbach C, Sellenschloh K, Gerbig L, Morlock MM, Beck-Broichsitter B, Smeets R, Heiland M, Huber G, Hanken H (2017) CAD-CAM plates versus conventional fixation plates for primary mandibular reconstruction: a biomechanical in vitro analysis. J Craniomaxillofac Surg 45:1878–1883. https://doi.org/10.1016/j. jcms.2017.08.024
- Fontana SC, Smith RB, Nazir N, Andrews BT (2016) Biomechanical assessment of fixation methods for segmental mandible reconstruction with fibula in the polyurethane model. Microsurgery 36:330–333. https://doi.org/10.1002/micr.30052
- Kim NK, Nam W, Kim HJ (2015) Comparison of miniplates and biodegradable plates in reconstruction of the mandible with a fibular free flap. Br J Oral Maxillofac Surg 53:223–229. https://doi.org/ 10.1016/j.bjoms.2014.11.010
- Al-Bustani S, Austin GK, Ambrose EC, Miller J, Hackman TG, Halvorson EG (2016) Miniplates versus reconstruction bars for oncologic free fibula flap mandible reconstruction. Ann Plast Surg 77:314–317. https://doi.org/10.1097/SAP. 0000000000000497
- Robey AB, Spann ML, McAuliff TM, Meza JL, Hollins RR, Johnson PJ (2008) Comparison of miniplates and reconstruction

plates in fibular flap reconstruction of the mandible. Plast Reconstr Surg 122:1733–1738. https://doi.org/10.1097/PRS. 0b013e31818a9ac5

- Zavattero E, Fasolis M, Garzino-Demo P, Berrone S, Ramieri GA (2014) Evaluation of plate-related complications and efficacy in fibula free flap mandibular reconstruction. J Craniofac Surg 25: 397–399. https://doi.org/10.1097/SCS.00000000000656
- Grohmann I, Raith S, Mucke T, Stimmer H, Rohleder N, Kesting MR, Holzle F, Steiner T (2015) Biomechanical loading test on reconstructed mandibles with fibular, iliac crest or scapula graft: a comparative study. Br J Oral Maxillofac Surg 53:741–747. https:// doi.org/10.1016/j.bjoms.2015.05.022
- Pietsch AP, Raith S, Ode JE, Teichmann J, Lethaus B, Mohlhenrich SC, Holzle F, Duda GN, Steiner T (2016) Biomechanical competence of six different bone screws for reconstructive surgery in three different transplants: fibular, iliac crest, scapular and artificial bone. J Craniomaxillofac Surg 44:689–696. https://doi.org/10.1016/j. jcms.2016.03.008
- Goodson AM, Kittur MA, Evans PL, Williams EM (2019) Patientspecific, printed titanium implants for reconstruction of mandibular continuity defects: a systematic review of the evidence. J Craniomaxillofac Surg 47:968–976. https://doi.org/10.1016/j. jcms.2019.02.010
- 32. Sadr-Eshkevari P, Rashad A, Vahdati SA, Garajei A, Bohluli B, Maurer P (2013) Alloplastic mandibular reconstruction: a systematic review and meta-analysis of the current century case series. Plast Reconstr Surg 132:413e–427e. https://doi.org/10.1097/PRS. 0b013e31829ad0d9
- Onoda S, Kimata Y, Yamada K, Sugiyama N, Onoda T, Eguchi M, Mizukawa N (2012) Prevention points for plate exposure in the mandibular reconstruction. J Craniomaxillofac Surg 40:e310– e314. https://doi.org/10.1016/j.jcms.2012.01.013
- Mariani PB, Kowalski LP, Magrin J (2006) Reconstruction of large defects postmandibulectomy for oral cancer using plates and myocutaneous flaps: a long-term follow-up. Int J Oral Maxillofac Surg 35:427–432. https://doi.org/10.1016/j.ijom.2005.10.008
- Wood CB, Shinn JR, Amin SN, Rohde SL, Sinard RJ (2018) Risk of plate removal in free flap reconstruction of the mandible. Oral Oncol 83:91–95. https://doi.org/10.1016/j.oraloncology.2018.06. 008
- Wei FC, Celik N, Yang WG, Chen IH, Chang YM, Chen HC (2003) Complications after reconstruction by plate and soft-tissue free flap in composite mandibular defects and secondary salvage reconstruction with osteocutaneous flap. Plast Reconstr Surg 112: 37–42. https://doi.org/10.1097/01.PRS.0000065911.00623.BD
- 37. Naros A, Weise H, Tilsen F, Hoefert S, Naros G, Krimmel M, Reinert S, Polligkeit J (2018) Three-dimensional accuracy of mandibular reconstruction by patient-specific pre-bent reconstruction plates using an "in-house" 3D-printer. J Craniomaxillofac Surg 46:1645–1651. https://doi.org/10.1016/j.jcms.2018.05.047
- Wurm MC, Hagen J, Nkenke E, Neukam FW, Schlittenbauer T (2019) The fitting accuracy of pre-bend reconstruction plates and their impact on the temporomandibular joint. J Craniomaxillofac Surg 47:53–59. https://doi.org/10.1016/j.jcms.2018.05.033
- Wolff KD, Follmann M, Nast A (2012) The diagnosis and treatment of oral cavity cancer. Dtsch Arztebl Int 109:829–835. https:// doi.org/10.3238/arztebl.2012.0829
- Mücke T, Rau A, Weitz J, Ljubic A, Rohleder N, Wolff K-D, Mitchell DA, Kesting MR (2012) Influence of irradiation and oncologic surgery on head and neck microsurgical reconstructions. Oral Oncol 48:367–371. https://doi.org/10.1016/j.oraloncology. 2011.11.013
- Thankappan K (2010) Microvascular free tissue transfer after prior radiotherapy in head and neck reconstruction - a review. Surg Oncol 19:227–234. https://doi.org/10.1016/j.suronc.2009.06.001

- 42. Herle P, Shukla L, Morrison WA, Shayan R (2015) Preoperative radiation and free flap outcomes for head and neck reconstruction: a systematic review and meta-analysis. ANZ J Surg 85:121–127. https://doi.org/10.1111/ans.12888
- Benatar MJ, Dassonville O, Chamorey E, Poissonnet G, Ettaiche M, Pierre CS, Benezery K, Hechema R, Demard F, Santini J, Bozec A (2013) Impact of preoperative radiotherapy on head and neck free flap reconstruction: a report on 429 cases. J Plast Reconstr Aesthet Surg : JPRAS 66:478–482. https://doi.org/10.1016/j.bjps.2012.12.019
- Sokoya M, Bahrami A, Vincent A, Kadakia S, Inman J, Saman M, Ducic Y (2018) Preoperative radiation and complication rates after double free flap reconstruction of head and neck cancer. Am J Otolaryngol 39:558–560. https://doi.org/10.1016/j.amjoto.2018. 06.015
- 45. Mijiti A, Kuerbantayi N, Zhang ZQ, Su MY, Zhang XH, Huojia M (2020) Influence of preoperative radiotherapy on head and neck free-flap reconstruction: systematic review and meta-analysis. Head Neck 42:2165–2180. https://doi.org/10.1002/hed.26136
- 46. Koerdt S, Steinstraesser L, Stoeckelhuber M, Wales CJ, Rohleder NH, Babaryka G, Steiner T, Wolff K-D, Loeffelbein DJ, Mücke T, Nieberler M, Kesting MR (2016) Radiotherapy for oral cancer decreases the cutaneous expression of host defence peptides. J Craniomaxillofac Surg 44:882–889. https://doi.org/10.1016/j.jcms.2016.04.014
- 47. Rohleder NH, Flensberg S, Bauer F, Wagenpfeil S, Wales CJ, Koerdt S, Wolff K-D, Hölzle F, Steiner T, Kesting MR (2014) Can tissue spectrophotometry and laser Doppler flowmetry help to identify patients at risk for wound healing disorders after neck dissection? Oral Surg Oral Med Oral Pathol Oral Radiol 117:302– 311. https://doi.org/10.1016/j.oooo.2013.11.497
- Lee S, Thiele C (2010) Factors associated with free flap complications after head and neck reconstruction and the molecular basis of fibrotic tissue rearrangement in preirradiated soft tissue. J Oral Maxillofac Surg 68:2169–2178. https://doi.org/10.1016/j.joms. 2009.08.026

- Lonie S, Herle P, Paddle A, Pradhan N, Birch T, Shayan R (2016) Mandibular reconstruction: meta-analysis of iliac- versus fibulafree flaps. ANZ J Surg 86:337–342. https://doi.org/10.1111/ans. 13274
- Ferrier MB, Spuesens EB, Le Cessie S, Baatenburg de Jong RJ (2005) Comorbidity as a major risk factor for mortality and complications in head and neck surgery. Arch Otolaryngol Head Neck Surg 131:27–32. https://doi.org/10.1001/archotol.131.1.27
- McMahon JD, MacIver C, Smith M, Stathopoulos P, Wales C, McNulty R, Handley TP, Devine JC (2013) Postoperative complications after major head and neck surgery with free flap repair– prevalence, patterns, and determinants: a prospective cohort study. Br J Oral Maxillofac Surg 51:689–695. https://doi.org/10.1016/j. bjoms.2013.04.015
- Haughey BH, Wilson E, Kluwe L, Piccirillo J, Fredrickson J, Sessions D, Spector G (2001) Free flap reconstruction of the head and neck: analysis of 241 cases. Otolaryngol Head Neck Surg 125: 10–17. https://doi.org/10.1067/mhn.2001.116788
- 53. Mücke T, Ritschl LM, Roth M, Güll FD, Rau A, Grill S, Kesting MR, Wolff K-D, Loeffelbein DJ (2016) Predictors of free flap loss in the head and neck region: s four-year retrospective study with 451 microvascular transplants at a single centre. J Craniomaxillofac Surg 44:1292–1298. https://doi.org/10.1016/j.jcms.2016.04.029
- 54. Khan MN, Russo J, Spivack J, Pool C, Likhterov I, Teng M, Genden EM, Miles BA (2017) Association of body mass index with infectious complications in free tissue transfer for head and neck reconstructive surgery. JAMA Otolaryngol Head Neck Surg 143:574–579. https://doi.org/10.1001/jamaoto.2016.4304
- Brown JS, Barry C, Ho M, Shaw R (2016) A new classification for mandibular defects after oncological resection. Lancet Oncol 17: e23–e30. https://doi.org/10.1016/S1470-2045(15)00310-1

Publisher's note Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.