

REVIEW

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Free versus pedicled flaps for reconstruction of head and neck cancer defects: a systematic review

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Abstract

Objective: The present review focuses on comparative studies of reconstruction with free flaps (FF) versus pedicled flaps (PF) after oncologic resection.

Method: A systematic review was developed in compliance with PRISMA guidelines and performed using the Pubmed, Medline, EMBASE, Amed and Biosis databases.

Results: A total of 30 articles were included. FF are associated with a longer operative time, a higher cost and a higher incidence of postoperative revisions compared to PF. FF are associated with a longer stay at the intensive care unit than the supraclavicular artery island flap (SCAIF) and with a more extended hospital stay compared to the submental island flap (SMIF). FF are associated with fewer infections and necrosis compared to the pectoralis major myocutaneous flap (PMMF).

Conclusion: The comparison of both type of flaps is limited by the inherent design of the studies included. In sum, FF seem superior to the PMMF for several outcomes. SMIF and SCAIF compare favorably to FF for some specific indications achieving similar outcomes at a lower cost.

Keywords: Flaps, Oncology, Reconstruction, Surgery, Outcomes

Introduction

Head and neck reconstruction surgery has considerably evolved over the past decades, along with the trend of using either a free or a pedicled flap for the reconstruction of oncologic defects. Tracing back the history of flaps, the first pedicled flap (PF) was described by Susruta in 800 BC and consisted of a forehead flap [1]. It was later popularized by McGregor in 1963 and marked a turning point in reconstructive surgery, being the first ever reliable transposition flap [2]. A decade later, the pectoralis major myocutaneous

flap (PMMF), supplied by the pectoral branch of the thoracoacromial artery, was introduced by Ariyan in 1979 [3]. The PMMF became the flap of choice for head and neck reconstruction in many centers and was extensively studied. However, concerns regarding the reliability of this flap for some defects resulted in the emergence of free flaps and other regional pedicled flaps, such as the supraclavicular artery island flap (SCAIF) and the submental island flap (SMIF).

With the advent of microvascular surgery in the 1970s, harvesting free flaps became popular in head and neck reconstruction surgery. Free tissue transfer was described by various authors, such as Daniel and Taylor who described the first cutaneous free flap in 1973 [4]. Free flap (FF) reconstruction slowly gained popularity over time to become the standard of care for large head & neck defects.

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Free flaps require the expertise of microvascular surgery and longer operative times, but they show more versatility and robustness than PF for some defects [5, 6]. Pedicled flaps are accessible to both academic and community surgeons and considered more reliable in specific settings but are not suitable for every defect [7, 8].

Flap selection is a complex process, with FF and PF having both their respective pros and cons. More importantly, patients pre-operative conditions, the nature of the disease, and the available resources are significant factors to consider when choosing the appropriate reconstructive technique.

Favoring one type over the other to obtain the best outcomes is a challenge and a source of debate in the literature. The purpose of this study is to review all articles explicitly comparing FF to PF for head and neck defects reconstruction regarding demographic parameters, risk factors, tumor staging, operative time, hospitalization length, cost, post-operative complications, and outcomes, in order to better characterize the benefits and disadvantages of these flaps. Regarding post-operative complications, donor and recipient sites morbidity, as well as the impact of either FF or PF reconstruction techniques on patients' quality of life, was evaluated to facilitate the choice for clinicians in the future.

Materials and methods

Literature review

The systematic review was performed in accordance with PRISMA guidelines, and a formal PROSPERO protocol was published according to the NHS International Prospective Register of Systematic Review (PROSPERO #42017055252). The Pubmed, Ovid-MEDLINE, EMBASE, Amed and Biosis databases were used to perform a literature review of English-language publication dating from 1948 to February 2017. Keyword combination included: free flaps AND pedicled flaps AND head and neck AND reconstruction surgery. The comparative study option was used as a limit to refine the search. Additionally, references in all articles were manually searched to identify other articles.

Selection criteria

Prospective and retrospective articles explicitly comparing the use of free flap versus pedicled flaps for head and neck oncologic defects were included. The data compared had to include one of the following parameters: demographic characteristics, risk factors, radiation or chemotherapy use, operative time, length of stay, total cost, post-operative complications and outcomes

concerning survival and quality of life. The paediatric population was excluded. Articles describing only revision surgery were also excluded.

Titles and abstracts were initially screened by two investigators (F.G.F and P.T) to discard irrelevant studies. All reference lists of identified studies were then further analyzed to include any additional articles of interest (Fig. 1). Selection of relevant studies was determined independently based on inclusion criteria. Any disagreement between reviewers was solved by discussions among the authors to reach consensus or by a third party (T.A), if necessary. The selection process was conducted per PRISMA guidelines. (Fig. 1).

Quality assessment

The methodological quality of evidence and the risks of bias of the included studies were assessed with the MINORS criteria (Methodological Index for Non-randomized Studies) [9]. Twelve criteria are used to evaluate the level of evidence of comparative studies. Criteria are graded from 0 to 2 (0: not reported; 1: reported but inadequate; 2: reported and adequate), for a global ideal score of 24. Studies with MINORS score > 18 were considered to have low risk bias. Quality assessment was conducted independently by two investigators (F.G.F and P.T) and discrepancies were resolved through a mutual re-review.

Results

A total of 30 articles were included for qualitative analysis after selection process (Fig. 1). All studies were retrospective except for one.

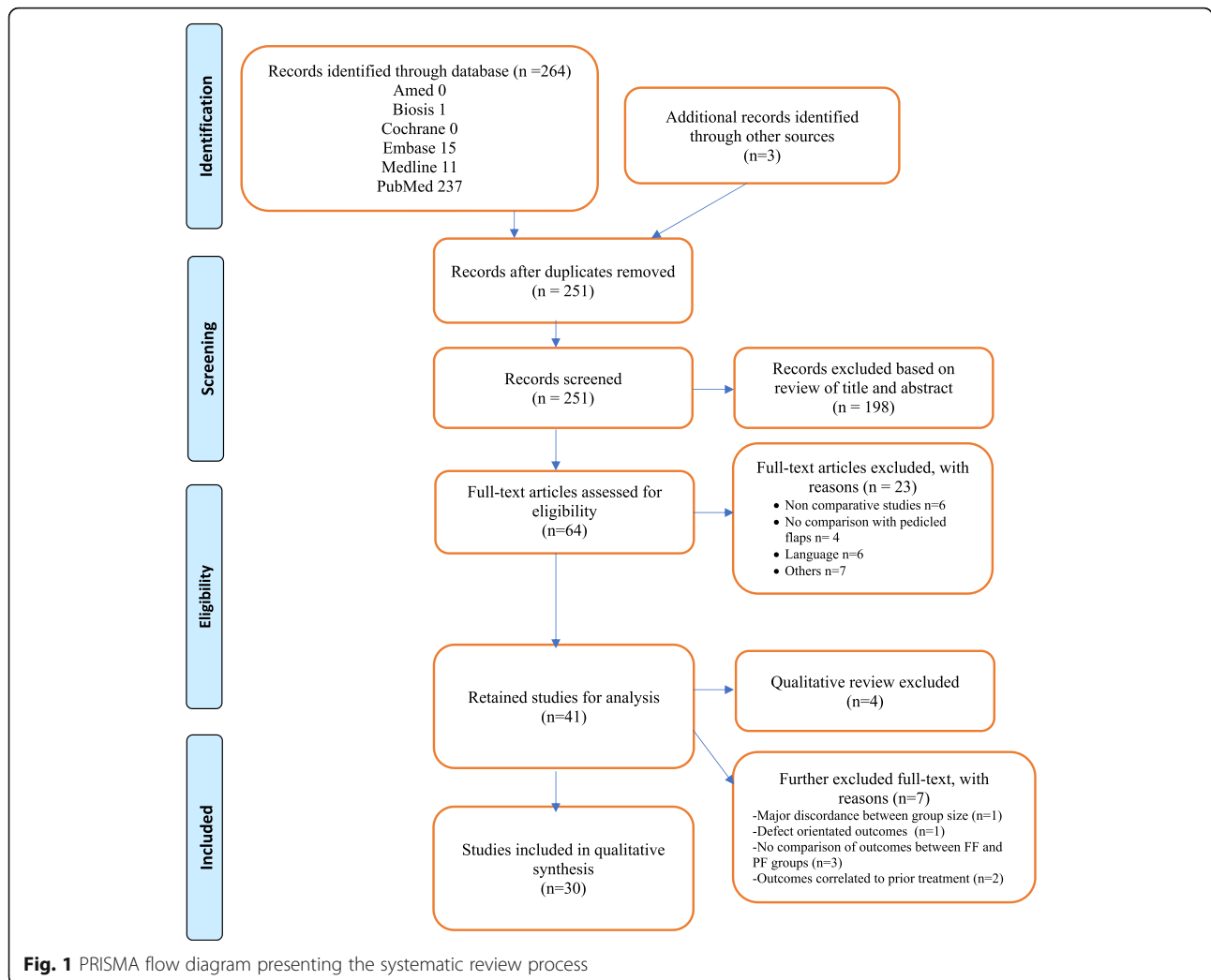
Types of flaps

Of the included studies, 53.3% ($n = 16$) compared FF to PMMF. Ten percent compared FF to supraclavicular artery island flap (SCAIF) ($n = 3$) and 10% compared FF to submental artery island flap (SMIF) ($n = 3$). The other studies compared FF to an array of different pedicled flaps or unspecified pedicled flaps. Types of flaps of all included studies are detailed in Table 1.

None of the included studies compared osseous free flaps to osseous pedicled flaps. All studies compared myocutaneous or fasciocutaneous flaps with the exception of two articles comparing fibular free flaps, along with other free flaps, to a variety of myocutaneous PF.

Quality of studies

The methodological quality of each study was evaluated with the MINORS criteria [9]. (Table 1 and details in Appendix). The studies scores ranged from 6 to 20. The mean and median scores were 18.2 and 18.5 respectively. Eight studies had a MINORS score



of > 20 and were considered to have low risk bias. Most studies were retrospective reviews and were deficient in categories of blinded evaluations, power calculation, and adequacy of group control. All the studies had clearly stated aims and end points were appropriate to the aim of the studies. One study had a low MINORS score of 6, but it was not excluded considering we had not established a minimum threshold for inclusion.

Defect location

Table 1 summarizes the defects location. Twenty-three studies over the 30 included mentioned the reconstruction site, with 18 studies describing defects location and 5 studies describing tumors location. The grouping of these sites differed among the studies and variable sub-site divisions were used. Both categories of flaps were not used equally to reconstruct a specific defect location within the studies, except for matching cohort studies. (Table 1 and details in Appendix).

Demographic parameters

Mean, or median age of patients was mentioned in 19 articles. Groups were comparable in 14 studies. Five articles showed that patients were significantly younger in the FF group compared to PF ($p < 0.05$) [10–14]. Gender representation was similar between both groups in the 20 studies reporting gender, as one study [15] showed fewer males in the FF group (70.8% vs 100%, $p < 0.05$) and another [16] showed fewer males in the PF group (88% vs 32%, $p < 0.05$). (Table 2).

Preoperative risk factors

Table 2 shows the preoperative risk factors for the FF and PF groups. Both were comparable for the incidence of smoking, chronic obstructive pulmonary disease (COPD), hypertension (HTA), cardiac atherosclerosis disease (CAD), dyslipidemia (DLP), chronic heart failure (CHF), alcoholism and the incidence of other cancer.

Table 1 Overview of the included studies

Article	Ref	Study type	MINORS Score ^a	N total	n FF	n PF	Type of free flaps	Type of pedicled flaps	Defect / tumor location
Sinha, 2017 [14]	A	Retrospective review	20	517	384	133	ALT, FibF, RFFF	PMMF, SCAIF, SMIF	N/A
Goyal, 2017 [10]	B	Retrospective review	18	797	589	208	NA	PMMF, SCAIF, SMIF, TEMP, TRPF, DELT	Cutaneous/ skull base Oral cavity Oropharynx Larynx / hypopharynx Mandibular Sinonasal Composite/multiple sites
Li, 2016 [15]	C	Retrospective review	19	41	24	17	RFFF	PMMF	Oral cavity
Kozin, 2016 [5]	D	Retrospective review	20	72	28	45	RFFF, ALT	SCAIF	Cutaneous defect Parotid/temporal bone
Howard, 2016 [21]	E	Retrospective review	18	31	9	16	ALT 6	SMIF PLDF	Lateral skull base
Geiger, 2016 [17]	F	Retrospective review	20	105	50	55	39 RFFF, 3 ALT, 4 FibF, 2 theLDF, 2 SFF, 2 ORFF	51 PMMF, 2 DIEP, 2 TPF	N/A
Gao, 2016 [49]	G	Retrospective review	16	60	34	26	RFFF	PMMF	N/A
Forner, 2016 [27]	H	Retrospective review	20	21	12	9	RFFF	SMIF	Oral cavity Oropharynx
Zhang S, 2015 [32]	I	Retrospective review	19	37	15	12	RFFF	SCAIF	Oral cavity
Zhang X, 2014 [16]	J	Retrospective review	19	110	79	31	ALT	PMMF	Oral cavity Oropharynx
Jing, 2014 [22]	K	Retrospective review	17	49	22	27	GFMF	PMMF	Larynx
Granzow, 2013 [7]	L	Retrospective review	18	34	16	18	FFF	SCAIF	Larynx / hypopharynx Parotid Oral cavity Esophagus
Deganello, 2013 [11]	M	Retrospective review	18	36	16	20	RFFF	10 TEMP, 10 PMMF	Oral cavity Oropharynx
Fang, 2013 [12]	N	Retrospective review	18	56	20	24	RFF 12 ALT	PLTM	Oral cavity
Paydarfar, 2011 [23]	O	Retrospective review	20	60	33	27	RFF	SMIF	Oral cavity
Hsing, 2011 [25]	P	Retrospective review	18	100	42	58	N/A	PMMF	Oral cavity
Chan Y, 2011 [35]	Q	Prospective review	18	202	24	92	ALT 86 FJF	PMMF	Hypopharynx
Demirtas, 2010 [18]	R	Retrospective review	20	20	12	8	10 ALT, 2 LDF	PLDF	N/A
O'Neil, 2010 [30]	S	Retrospective review	20	114	77	37	RFFF	PMMF	N/A
Mallet, 2009 [6]	T	Retrospective review	18	70	25	45	18 RFFF, 3 LD, 3 ALT, 1 PCFF, 1 FJF	PMMF	Oral cavity Oropharynx

Table 1 Overview of the included studies (Continued)

Article	Ref	Study type	MINORS Score ^a	N total	n FF	n PF	Type of free flaps	Type of pedicled flaps	Defect / tumor location
de Bree, 2007 [20]	U	Retrospective review – Matched cohort	19	80	40	40	RFFF	PMMF	Oral cavity Oropharynx
Smeele, 2006 [29]	V	Retrospective – Matched cohort	20	64	32	32	N/A	PMMF	Oral cavity Oropharynx
Chien, 2005 [26]	W	Case series	17	27	11	16	RFFF	PBFF	Oral cavity
Chepeha, 2004 [8]	X	Retrospective review	19	179	71	108	N/A	PMMF	Oral cavity Oropharynx Hypopharynx Neck Others
Funk, 2002 [19]	Y	Case-control, matched pairs study	20	42	21	21	FTT	N/A	Oral cavity Oropharynx Hypopharynx Larynx
Petruzzelli, 2002 [31]	Z	Retrospective review	18	39	24	15	FTT	N/A	N/A
Amarante, 2000 [34]	Aa	Case-series	6	117	49	68	23 RFFF, 3 ORFF, 2 CFF, 5 LDF, 5 RAFF, 3 MSA, 2 PSFF, 2 ICC, 1 GOM, 3 FJF	47 PMMF, 7 PLDF, 2 PLTM, 9 TEMP, 3 MEC	Orbit Parotid Skull base Oropharynx Larynx / hypopharynx Mandibular Neck
Tsue, 1997 [24]	Ab	Retrospective review	19	53	29	24	N/A	PMMF	Oral cavity Oropharynx
Kroll, 1997 [13]	Ac	Retrospective review	17	178	145	33	89 RFFF, 56 RAFF	PMMF	Oral cavity Oropharynx
Kroll 1992 [33]	Ad	Retrospective review	18	69	30	39	RAFF	PMMF	N/A

Ref: Reference in subsequent tables

^aMINORS score ranges from 0 to 24. A value ≥20 indicates low risk of bias

Flaps abbreviation: ALT Anterolateral thigh, CFF Cubital forearm flap, DELT Deltopectoralis, DIEP Deep inferior epigastric perforator flap, FibF Fibular free flap, FFF free fasciocutaneous flap, FTT Microvascular free tissue transfer, FJF Free jejunal flap, GFMF Gracilis free muscle flap, GOM Greater Omentum, ICC Iliac crest, LDF Latissimus dorsi free flap, MEC musculocutaneous sternocleidomastoid flap, MSA Muscular serratus anterior, N/A Not available, ORFF Osteocutaneous Radial free flap, PBFF pedicled buccal fat flap, PCFF Pectoralis major free flap, PLTM Platysma myocutaneous island flap, PLDF Pedicled latissimus mucocutaneous dorsi flap, PMMF Pectoralis major pedicled flap, PSFF Parascapular free flap, RFFF Radial forearm free flap, RAFF Rectus abdominis free flap, SCAIF Supraclavicular artery island flap, SMIF Submental Island flap, TEMP Temporal flap, TRPF Trapezius flap

Prior head and neck surgery was reported in 4 studies. Two studies [10, 17] found there was a lower proportion of patients who had prior head and neck surgery in the FF group compared to the PF group (32% vs 59 and 14% vs 31%, $p < 0.05$). One study [12] reporting the incidence of preoperative systemic disease showed a lower regrouped incidence of diabetes, cardiovascular disease and hypertension in the FF group (25% vs 83%, $p < 0.05$). In contrast, another study [7] demonstrated a higher incidence of diabetes mellitus in the FF group compared to PF (31% vs 0, $p < 0.05$). The only study reporting the incidence of atrial fibrillation [14] showed less patient with atrial fibrillation in the FF group compared to PF (7% vs 15%, $p < 0.05$).

American Society of Anesthesiologists (ASA) classification

Six studies reported ASA class. Among them, four showed similar ASA classes [6, 7, 18, 19]. Two studies showed a significant difference in ASA class between FF and PF groups with contrasting results. Goyal et al. [10] showed a higher proportion of ASA class I-II in the FF group (41.6% vs 26.9%, $p < 0.05$) and a lower proportion of ASA class III-IV in the FF group (58.2% vs 73.1%, $p < 0.05$) compared to PF.

In contrast, de Bree et al. [20] demonstrated a lower proportion of patients with ASA class I-II in the FF group (80% vs 92.5%, $p < 0.05$) and a higher proportion of ASA class III-IV in the FF group (20% vs 8%, $p < 0.05$). (Table 2.)

Table 2 Demographic data, preoperative risk factors and ASA class

	Total # of articles reporting (total = n)	# of articles reporting differences (total n)	Articles reporting differences	FF	PF	p-value
Age, mean ± SD or mean (range)	16 ^{B, D, H, K, L, M, N, O, P, R, T, W, X, Z, Aa, Ac} (n = 1855)	4 (n = 1067)	Goyal, 2017 Deganello, 2013 Fang, 2013	64.0 ± 12.0 58.2 ± 6.32 58.0 (25–78)	66.5 ± 12.9 69.6 ± 6.8 72.4 (55–80)	0.017 < 0.01 < 0.001
				57.2 (46–72)		< 0.001
			Kroll, 1997	56 ± 13	62 ± 12	0.0046
Age (median)	3 ^{A, E, Ab} (n = 601)	1 (n = 517)	Sinha, 2017	65.9 (57.7–74.2)	67.9 (60.3–76.8)	0.037
Age > 50 years	4 ^{C, F, I, J} (n = 293)	0				
Male	20 ^{A, D, C, F, E, H, I, J, K, L, M, N, O, P, R, T, W, X, Aa, Ab} (n = 1609)	2 (n = 151)	Li, 2016 Zhang X, 2014	17(70.83%) 66 (88%)	17(100%) 31 (32.0%)	0.043 0.018
Smoking	5 ^{F, K, L, T, Y} (n = 300)	0		62.0%	61.8%	0.985
Prior head and neck surgery	4 ^{B, F, X, Ab} (n = 1134)	2 (n = 902)	Goyal, 2017 Geiger, 2016 ^a	189 (32.1%) 14.0%	122 (58.7%) 30.9%	< 0.001 0.039
Systemic diseases (CVD, HTA, DM)	1 ^N (n = 56)	1 (n = 56)	Fang, 2013	4 (20%) 3 (25%)	20 (83.33%)	< 0.001 < 0.001
COPD	2 ^{A, T} (n = 587)	0				
DM	3 ^{A, L, P} (n = 651)	1 (n = 34)	Granzow, 2013	5 (31%)	0	0.02
HTA	2 ^{F, L} (n = 139)	0				
CAD	3 ^{A, L, T} (n = 621)	0				
DLP	1 ^F (n = 105)	0				
CHF	1 ^A (n = 517)	0				
aFIB	1 ^A (n = 517)	1 (n = 517)	Sinha, 2017	7.0%	15.0%	0.0083
Alcoholism	1 ^T (n = 70)	0				
Other cancer	1 ^T (n = 70)	0				
ASA Class I-II, n (%)	6 ^{B, L, R, T, U, Y} (n = 1043)	2 (n = 877)	Goyal, 2017 de Bree, 2007	245 (41.6%) 32 (80%)	56 (26.9%) 37 (92.5%)	0.001 0.028
ASA Class III-IV, n (%)	6 ^{B, L, R, T, U, Y} (n = 1043)	2 (n = 877)	Goyal, 2017 de Bree, 2007	343 (58.2%) 8 (20%)	152 (73.1%) 3 (8%)	0.001 0.028
ASA mean factor (mean ± SD)	4 ^{A, H, R, Ac} (n = 736)	2 (n = 538)	Sinha, 2017 Forner, 2016	2.6 ± 0.03 2.3	2.8 ± 0.05 2.4	0.0007 0.05

aFIB atrial fibrillation, ASA risk factor: scored using the American Society of Anesthesiology Scale (ASA), CAD Cardiac artery disease, CHF Chronic heart failure, COPD Chronic obstructive pulmonary disease, CVD Cardiovascular disease, DM Diabetes mellitus, DLP Dyslipidemia, HTA Hypertension

Prior radiation or chemotherapy

Exposure to prior head and neck radiation therapy was mentioned in 11 articles, and was comparable between the FF and PF groups in 9 of them [5–8, 13, 17, 21–23]. (Table 3) Two articles [10, 24] showed a significantly

lower proportion of prior radiotherapy with FF reconstruction compared to PF (46% vs 62 and 28% vs 54% *p* < 0.05). No difference was seen in the incidence of prior chemotherapy between FF and PF, as it was reported in five studies [5, 7, 17, 23, 24]. The incidence of adjuvant

Table 3 Staging and treatment data

	Total # of articles reporting (total = n)	# of articles reporting differences (total n)	Articles reporting difference	FF	PF	p-value
Prior radiation	11 ^{B, D, E, F, K, L, O, T, X, Ab, Ac} (n = 1628)	2 (n = 850)	Goyal, 2017 Tsue, 1997	272 (46.2%) 8 (28%)	130 (62.5%) 13 (54%)	< 0.001 0.05
Prior chemotherapy	5 ^{D, K, L, O, Ab} (n = 268)	0				
T1	8 ^{J, K, M, N, O, P, W, Y} (n = 480)	1 (n = 36)	Deganello, 2013	0	4 (20%) [#]	< 0.01
T2	8 ^{J, K, M, N, O, P, W, Y} (n = 480)	1 (n = 36)	Deganello, 2013	7 (43.8%) [#]	5 (25%) [#]	< 0.01
T3	8 ^{J, K, M, N, O, P, W, Y} (n = 480)	1 (n = 36)	Deganello, 2013	8 (50%)	8 (40%)	< 0.01
T4	8 ^{J, K, M, N, O, P, W, Y} (n = 480)	1 (n = 36)	Deganello, 2013	1 (6.25%)	3 (15%)	< 0.01
T1-T2	4 ^{C, H, T, Ab} (n = 185)	0				
T3-T4	4 ^{C, H, T, Ab} (n = 185)	0				
Stage I-II	1 ^X (n = 179)	0				
Stage III-IV	1 ^X (n = 179)	0				
Surg + chemoradio	6 ^{C, N, O, P, U} (n = 516)	1 (n = 60)	Paydarfar, 2011	16 (48.48%) [#]	12 (44.44%) [#]	0.03
Surg + radio	4 ^{J, O, P, X} (n = 449)	0				
Tumor stage	1 ^{Ac} (n = 178)	0				
Tumor recurrence	1 ^{Ac} (n = 178)	0				

Surg + chemoradio: Surgical resection and adjuvant chemoradiotherapy

Surg + radio: Surgical resection and adjuvant radiotherapy

Percentage calculated relying on the data presented. Percentage not provided by the article

Bold = Statistically significant, p-value ≤ 0.05

chemoradiotherapy after surgery was higher in the FF group in one study (48% vs 44%, $p < 0.05$) [23].

Tumor staging

Tumor stages were compared in 13 studies; T stage or global staging was reported (Table 3). No significant difference in cancer staging was found in the other 12 studies [6, 11, 12, 15, 16, 19, 22–27]. A study comparing RFFF to PMMF and temporalis flap [11] showed a lower proportion of T1 and T4 in the FF group (T1: 0% vs 20%; T4: 6.25% vs 15%, $p < 0.05$) as well as a higher proportion of T2 and T3 in the FF group (T2: 43.8% vs 25%; T3: 50% vs 40%, $p < 0.05$) when compared to PF.

Operative time

Nineteen studies compared the operative time between both reconstruction techniques. All showed that FF was associated with a longer operating time than PF. This difference was statistically significant in 14 studies (Table 4) [5, 7, 10, 13–16, 20, 21, 23, 24, 27–29].

Hospitalization and ICU length of stay

Seventeen studies compared the duration of hospital stay. (Table 4). Ten studies showed a similar hospitalization stay with FF compared to PF [5–7, 11, 18, 24, 27, 29–31]. However, when FF were compared to SMIF and SCAIF specifically the results differed. FF patients had a longer hospitalization stay than SMIF patients for skull base (9.8 days vs 4.75, $p < 0.05$) [21] and oral cavity (14.0 days vs 10.6, $p < 0.05$) defects [23]. SCAIF patients had a shorter length of stay than FF patients for oral cavity defects (12 ± 1.7 vs 17 ± 2.5 , $p < 0.05$) [32]. On the other hand, four studies [8, 13, 20, 33] showed a shorter hospitalization stay with FF compared to PMMF ($p < 0.05$).

Four studies assessed the ICU length of stay [7, 24, 27, 29]. One study comparing FF to SCAIF for larynx/pharynx reconstruction [7] concluded in a longer stay for FF reconstruction ($p < 0.05$).

Cost

Nine studies compared hospital costs between FF and PF [5, 11, 13, 18, 24, 27, 29, 31]. As reported by three studies, FF was associated with a

Table 4 OR time, Hospital and ICU length and hospital cost

	Total # of articles reporting (total = n)	# of articles reporting differences (total n)	Articles reporting difference	FF	PF	p-value			
OR time, min (mean ± SD)	12 ^{A, B, C, E, H, I, L, M, O, R, U, Ab} (n = 1727)	9 (n = 1634)	Sinha, 2017	421.4 ± 4.4	332.7 ± 10.7	0.0001			
			Goyal, 2017	427.2 ± 92.3	310.8 ± 125.0	0.001			
			Li, 2016	405 ± 107	365 ± 48	< 0.05			
			Howard, 2016	683 (575–979)	544 (396–700)	0.00817			
			Forner, 2016	552	347	< 0.05			
			Granzow, 2013	816.3 ± 148.9	587.9 ± 130.5	0.0002			
			Paydarfar, 2011	780	506.4	0.001			
			de Bree, 2007	692	462	< 0.005			
			Tsue, 1997	684 ± 16	666 ± 20	0.003			
			OR time, hour (mean ± SD)	6 ^{D, N, T, V, Ac, Ad} (n = 474)	4 (n = 384)	Kozin, 2016	8.1	6.7	0.002
Mallet, 2009	7.01 ± 1.19)	4.19 ± 0.57				< 0.001			
Smeele, 2006	12.5 ± 1.9	9.9 ± 1.5				< 0.0001			
Kroll, 1997	10.49 ± 2.06	9.39 ± 2.59				0.029			
Zhang X, 2014	59 (74.68%)	3 (9.68%)				0.001			
OR time of > 600 min	1 ^J (n = 110)	1 (n = 110)	Zhang X, 2014	59 (74.68%)	3 (9.68%)	0.001			
			Hospital length, days (mean ± SD)	17 ^{D, E, H, I, L, M, O, R, S, T, U, V, X, Z, Ab, Ac, Ad} (n = 1104)	7 (n = 634)	Howard, 2016	9.8 (7–22)	4.75 (2–14)	0.004
						Zhang S, 2015	17 ± 2.5	12 ± 1.7	< 0.05
						Paydarfar, 2011	14.0	10.6	0.008
						de Bree, 2007	24	28	0.005
						Chepeha, 2004	12	14	0.006
						Kroll, 1997	13.2 ± 5.4	19.8 ± 11.5	0.003
ICU length, days (mean ± SD)	4 ^{H, L, V, Ab} (n = 172)	1 (n = 34)	Kroll, 1992	11.3	21.2	0.003			
			Granzow, 2013	5.6 (4–9)	1.8 (0–5)	0.0001			
			Hospital cost	9 ^{D, H, M, R, U, V, Z, Ab, Ac} (n = 563)	4 (n = 339)	Kozin, 2016	SCAIF 32% less expensive than FTT	0.0001	
Deganello, 2013	22,924	19,872				0.043			
Tsue, 1997	50,026 ± 4340	38,246 ± 1440				0.003			
Kroll, 1997	28,460 ± 8435	40,992 ± 1958				0.001			

significantly higher cost compared to PF. Of these, Kozin et al. [5] showed that SCAIF was 32% less expensive than FF for total laryngectomy, parotid/temporal bone, and cutaneous defect reconstruction. Reconstruction for oral cavity and oropharynx were also less expensive with PMMF compared to FF. (38,246\$ vs 50,026\$, $p < 0.05$) [24]. A study comparing temporal flap (TEMP) and PMMF reconstruction to RFFF for oral cavity and oropharyngeal defects to RFFF led to similar findings (19,872\$ vs 22,924\$, $p < 0.05$) [11]. In contrast, one study [13] showed a lower hospital cost with FF compared to PMMF for the reconstruction of oral and oropharyngeal defects. (28,460 ± 8435 vs 40,992 ± 1958, $p < 0.05$) (Table 4).

Post-operative complications

Articles differed in the definitions of their studied complications. (Table 5). For example, some studies grouped recipient and donor site complications, as other separated them. Some studies were less specific and only reported the incidence of any complications. Others were selectively reporting the incidence of infection, fistula, abscess, dehiscence, hematoma, and others. Articles and results were grouped according to the definition of their studied complications.

Seven articles reported the incidence of “any complications” [7, 16, 17, 22, 23, 33, 34]. One article [17] showed that FF was associated with a higher incidence of any complication (68.0% vs 36.4%, $p < 0.05$). This article included various types

Table 5 Post-operative complications and outcomes

	Total # of articles reporting (total = n)	# of articles reporting differences (total n)	Articles reporting difference	FF	PF	p-value
Any complications	7 ^{F, J, K, L, O, Aa, Ad} (n = 544)	3 (n = 284)	Geiger, 2016	68.0%	36.4%	0.001
			Zhang X, 2014	13 (16.46%)	14 (45.16%)	0.002
			Kroll, 1992	4 (13%)	17 (44%)	0.0145
Infection	3 ^{F, L, T} (n = 209)	0				
Recipient site infection	6 ^{B, D, O, S, T, X} (n = 1292)	1 (n = 179)	Chepeha, 2004	2 (3%)	18 (17%)	< 0.004
Donor site infection	3 ^{B, D, S} (n = 983)	0				
Donor site morbidity	1 ^Q (n = 202)	0				
Fistula	11 ^{B, F, K, O, Q, S, R, T, V, X, Aa} (n = 1777)	2 (n = 902)	Goyal, 2017	18 (3.1%)	17 (8.2%)	0.005
			Geiger, 2016	22.0%	7.3%	0.039
Abscess	1 ^F (n = 105)	0				
Dehiscence recipient or donor site	4 ^{F, K, S, V} (n = 332)	1 (n = 105)	Geiger, 2016	44.0%	23.6%	0.029
Dehiscence recipient site	5 ^{D, L, O, V, X} (n = 409)	1 (n = 179)	Chepeha, 2004	0	11 (10%)	< 0.008
Dehiscence donor site	8 ^{D, I, L, O, V, K, L, R} (n = 370)	0				
Hematoma	2 ^{X, S} (n = 293)	0				
Hematoma Donor site	2 ^{E, V} (n = 95)	0				
Hematoma recipient site	2 ^{O, V} (n = 124)	0				
Partial flap necrosis	6 ^{I, O, V, X, Aa, Ad} (n = 526)	1 (n = 179)	Chepeha, 2004	2 (2.82%) [#]	12 (11%) [#]	< 0.006
Total flap necrosis	2 ^{V, Aa} (n = 181)	0				
Partial or total flap necrosis	1 ^T (n = 70)	1 (n = 70)	Mallet, 2009	1 (4%)	14 (31%)	0.02
Osteonecrosis	2 ^{B, F} (n = 902)	1 (n = 105)	Geiger, 2016	24.0%	3.6%	0.007
Deep Vein Thrombosis (Inferious member)	2 ^{A, S} (n = 631)	0				
Venous obstruction (At site)	1 ^O (n = 60)	0				
Late anastomotic stricture	1 ^Q (n = 202)	0				
Operative revision surgery	8 ^{E, F, L, O, S, R, X, Aa} (n = 660)	2 (n = 136)	Howard, 2016	1.6 (1–3)	0.6 (0–1)	< 0.00001
			Geiger, 2016	34%	9.1%	0.003
Flap failure	8 ^{E, I, K, L, O, T, U, V} (n = 425)	1 (n = 70)	Mallet, 2009	1 (4%)	14 (31%)	0.02
Mortality at 30 days	2 (n = 228) ^{K, X}					
Mortality at 1-year	2 (n = 76) ^{L, Y}					
Mortality at 2-year	1 (n = 80) ^U					

Percentage not provided by the original article, calculated by the authors from the data presented

of flaps in both FF and PF groups. Two studies [16, 33] showed the opposite with a lower incidence of “any complication” in the FF group compared to PF. Of those, Zhang X et al. [16] showed a significantly lower rate of complications in the FF group compared to PMMF. (16.5% vs 45.2%, $p < 0.05$).

Infections at large, recipient site infection and donor site infection were reported in some studies. In one study [8], the rate of infection at the recipient site was lower in the FF group compared to the PMMF group (3% vs 17%, $p < 0.05$).

Two studies showed significant differences in the incidence of fistula. Goyal et al. [10] showed a lower rate of fistula in the FF group compared to PF (3.1% vs 8.2%, $p < 0.05$). The exact defect location was not specified in this study including multiple reconstruction sites, i.e. skull base, sinonasal cavities, oral cavity, and larynx. However, in a study focusing on intraoperative brachytherapy [17], the rate of fistula was higher in the FF group (22% vs 7.3%, $p < 0.05$). Neither the defect location nor the exact type of flaps was mentioned in this study.

The incidence of dehiscence either at the recipient, donor, or recipient and/or donor sites was reported by several studies. Dehiscence at “recipient and/or donor” site was higher with FF reconstruction compared to PF, according to Geiger et al. (44% vs 23.6%, $p < 0.05$) [17].

Dehiscence at recipient site was lower in FF group compared to PMMF in one study (0 vs 10%, $p < 0.05$) [8]. As for dehiscence at the donor site, no significant difference was observed between FF and PF in the eight studies reporting this complication [5, 7, 18, 22, 23, 29, 32].

The incidence of hematomas either at the recipient, donor, or recipient and/or donor site was analyzed by very few studies [8, 21, 23, 29, 30], without statistically significant differences between both techniques.

One study [8] showed a lower incidence of partial flap necrosis with FF reconstruction compared to PMMF (2.8% vs 11%, $p < 0.05$) for various defect locations (oral cavity, oropharynx, hypopharynx, neck, and others). The same was revealed by the Mallet et al. [6] study where “partial or total flap” necrosis was higher with PMMF for oral tongue and base of tongue reconstruction (4% vs 31%, $p < 0.05$).

Post-operative outcomes

Operative revision surgery was significantly higher in the FF group in two studies [21] (Table 5). One compared FF to SMIF (1.6 vs 0.6, $p < 0.05$) for lateral skull base defects [21] and the other compared various FF to PF (34% vs 9.1%, $p < 0.05$) without specifying the defect location [17]. Although not being statistically significant, six other studies also showed a

higher occurrence of revision with FF reconstruction [7, 8, 18, 23, 29, 30, 34]. One study [6] showed that flap failure was more frequent with PMMF compared to FF (4% vs 31%, $p < 0.05$) for oral tongue and base of tongue reconstruction. No difference between both groups was reported for mortality at 30 days [8, 22], at 1 year [7, 19] and at 2 years [20].

Quality of life

Table 6 shows the quality of life of patients after surgical reconstruction with either FF or PF. The University of Washington Quality of Life Questionnaire (UW-QOL), including 14 items, was used by three studies [15, 16, 25] to measure the quality of life after surgical reconstruction with either FF or PMMF. Differences were seen in speech. Zhang X. et al. [16] showed a lower quality of speech with FF (57.5 ± 20.1 vs 76.1 ± 13.3 , $p < 0.05$) with a mean follow-up of 5.9 years. Hsing et al. showed a better quality of speech with FF compared to PMMF (66.7 ± 27.2 vs 44.7 ± 35.0 , $p < 0.05$) from data of patients operated 2 to > 10 years earlier.

Speech quality was also specifically assessed by two other studies not using the UW-QOL. O’Neil et al. [30] found a difference in speech quality ($p < 0.05$), with RFFF patients being more often “always understandable” than PMMF patients (53.1% vs 22.2%, follow-up period not mentioned). Additionally, Zhang S. et al. [32] graded the speech quality as excellent, good or poor, and found no difference in the speech quality of reconstruction with either FF or SCAIF flaps 6 months after the surgery.

Shoulder function, evaluated with UW-QOL, was significantly better in the FF group compared to the PMMF group in all three studies [15, 16, 25]. Follow-up time was ranging from 1 to over 10 years. One study [25] showed that FF was associated with a better mood compared to PMMF (76.2 ± 24.7 vs 60.8 ± 32.8 , $p < 0.05$).

In addition, looking at studies using the UW-QOL, FF and PMMF scored similarly on global quality of life, pain, swallowing, chewing, speech, activity, recreation, taste, saliva, anxiety and composite score [15, 16, 25].

Recovery to a normal diet was reported in five studies [23, 24, 30, 32, 35]. According to one study [24], the incidence was higher in the FF group compared to the PMMF for reconstruction of oral or base of tongue defects (34% vs 17%, $p < 0.05$).

Preoperative and postoperative mouth opening were reported by two studies, one comparing RFFF and ALT to platysma myocutaneous island flap (PMIF) [12] and the other comparing RFFF to pedicled buccal fat pad flap [26]. Mouth opening was similar between FF and PF groups.

Table 6 Quality of Life data

	Article	FF	PF	<i>p</i> -value
UW-QOL Global	Li, 2016 [#]	55.14 ± 9.24	54.36 ± 8.13	0.965
	Zhang X, 2014 [§]	70.5 ± 16.7	67.3 ± 12.9	0.860
	Hsing, 2011 ^{&}	66.0 ± 18.5	57.8 ± 18.2	0.090
UW-QOL: Pain	Li, 2016	71.63 ± 9.91	72.94 ± 11.13	0.751
	Zhang X, 2014	86.2 ± 10.8	89.9 ± 11.4	0.425
	Hsing, 2011	76.8 ± 23.0	68.1 ± 27.2	0.138
UW-QOL: Swallowing	Li, 2016	44.00 ± 16.27	43.78 ± 4.95	0.741
	Zhang X, 2014	49.4 ± 14.7	51.3 ± 21.7	0.840
	Hsing, 2011	49.3 ± 37.2	48.6 ± 32.7	0.962
UW-QOL: Chewing	Li, 2016	42.45 ± 6.15	43.43 ± 12.37	0.817
	Zhang X, 2014	52.6 ± 17.1	59.4 ± 12.9	0.498
	Hsing, 2011	34.5 ± 39.0	33.6 ± 36.7	0.973
UW-QOL: Speech	Li, 2016	51.27 ± 11.24	52.63 ± 12.43	0.461
	Zhang X, 2014	57.5 ± 20.1	76.1 ± 13.3	0.017
	Hsing, 2011	66.7 ± 27.2	44.7 ± 35.0	0.002
UW-QOL: Appearance	Li, 2016	57.47 ± 11.44	68.54 ± 13.24	0.0001
	Zhang X, 2014	76.4 ± 18.6	70.3 ± 17.1	0.308
	Hsing, 2011	67.3 ± 25.0	69.8 ± 25.5	0.535
UW-QOL: Activity	Li, 2016	64.23 ± 9.52	63.73 ± 8.41	0.641
	Zhang X, 2014	71.9 ± 11.5	74.8 ± 10.2	0.710
	Hsing, 2011	67.9 ± 24.2	66.8 ± 27.9	0.760
UW-QOL: Recreation	Li, 2016	66.59 ± 11.62	67.26 ± 9.23	0.445
	Zhang X, 2014	72.1 ± 10.2	78.9 ± 11.2	0.590
	Hsing, 2011	69.1 ± 32.6	62.5 ± 32.2	0.221
UW-QOL: Shoulder	Li, 2016	61.52 ± 7.83	54.65 ± 11.24	0.0001
	Zhang X, 2014	87.1 ± 14.4	65.6 ± 20.0	< 0.001
	Hsing, 2011	81.4 ± 14.7	50.5 ± 29.8	< 0.001
UW-QOL: Taste	Li, 2016	50.91 ± 10.64	51.24 ± 11.23	0.673
	Zhang X, 2014	48.4 (18.3)	52.9 (19.6)	0.713
	Hsing, 2011	55.0 ± 43.2	45.9 ± 39.6	0.226
UW-QOL: Salive	Li, 2016	45.48 ± 16.92	44.17 ± 12.78	0.723
	Zhang X, 2014	70.9 ± 9.5	72.3 ± 23.1	0.813
	Hsing, 2011	71.7 ± 34.8	73.8 ± 28.1	0.964
UW-QOL: Mood	Li, 2016	69.94 ± 9.51	68.31 ± 14.72	0.474
	Zhang X, 2014	76.0 ± 14.7	71.6 ± 18.8	0.114
	Hsing, 2011	76.2 ± 24.7	60.8 ± 32.8	0.022
UW-QOL: Anxiety	Li, 2016	70.57 ± 15.11	72.55 ± 15.19	0.219
	Zhang X, 2014	78.5 ± 9.64	86.4 ± 17.5	0.775
	Hsing, 2011	75.9 ± 26.3	68.9 ± 33.9	0.423
UW-QOL: Composite score	Hsing, 2011	66.0 ± 18.5	57.8 ± 18.2	0.090

Table 6 Quality of Life data (Continued)

		Article	FF	PF	p-value
Speech	Excellent	Zhang S, 2015 [§]	12 (80.0%) [#]	11 (91.7%) [#]	0.62
	Good		3 (20%)	1 (8.3%)	
	Poor		0	0	
	Always understandable	O'Neil, 201	17 (53.1)	4 (22.2)	0.014
	Usually understandable		14 (43.8)	9 (50.0)	
	Difficult to understand		1 (3.1)	5 (27.8)	
Swallowing full/regular diet at follow-up (vs soft, liquid) n (%)		Zhang S, 2015 [§]	13 (86.7%) [#]	10 (83.3%) [#]	1.00
		Paydarfar, 2011 [¶]	19	20	0.60
		Chan Y, 2011*	8 (38.2%)	24 (35.8%)	ND
			52 (61.9%)		ND
		O'Neil, 2010**	17 (59.4%)	6 (33.3%)	0.202
		Tsue, 1997***	8 (34%)	4 (17%)	0.02
Preoperative mouth-open width distance (mean) cm		Fang, 2013	1.5–6.2 (4.6)	1.2–6.2 (4.8)	ND
			0.9–6.0 (3.5)		ND
		Chien, 2005	6.3–3.5 (5.7)	6.1–2.5 (5.1)	0.384
Postoperative mouth-open width		Fang, 2013	1.4–5.8 (4.3)	1.1–4.7 (3.2)	ND
			0.8–5.8 (3.3)		ND
		Chien, 2005	5.9–3.2 (5.2)	5.6–1.6 (3.6)	0.384
Mouth-open width change (%)		Fang, 2013	4.0–9.1%	8.3–47.5%	< 0.001
			3.3–11.1%		< 0.001
		Chien, 2005	4.8–9.8%	5–45.5%	< 0.001
G-tube at 6 months postoperatively		Smeele, 2006	21.8%	34.3%	NS
G-tube dependence, n (%)		Chepeha, 2004	10 (16%)	40 (42%)	0.001
Feeding tube for >21 days		Mallet, 2009	8 (36%)	17 (42%)	0.84
Feeding tube at discharge		Tsue, 1997	20 (69%)	20 (83%)	NS
Feeding tube at follow up ^e			11 (39%)	17 (85%)	0.002

Follow up ranging from 13 to 108 months

\$ Mean-follow up = 5.9 years

& Follow up ranging from 2 to >10 years

§ Follow-up = 6 months

% at most recent follow-up

* Regular PO follow-up, median follow-up period was 82 months

** Follow-up period not mention

*** Median follow-up was 298 days

Percentage calculated relying on the data presented. Percentage not provided by the article

Bold = Statistically significant, p-value ≤ 0.05

The incidence of feeding tube dependence was reported by some studies and different postoperative time-points were evaluated. One study [8] showed a lower incidence of feeding tube dependence in the FF group compared to PMMF (16% vs 42%, $p < 0.05$) for reconstruction of various defects. The FF group was also associated with a lower rate of incidence of feeding tube at follow up, with a median follow-up of 298 days, compared to PMMF (39% vs 85%, $p < 0.05$) for oral cavity and oropharynx reconstruction [24]. Feeding tube dependence at 21 days [6] and at discharge [24] was similar between the PF and FF groups in two studies.

Discussion

Choosing between FF and PF in head and neck reconstruction is a challenge for some defects, especially with the recent resurgence of PF and their expanding indications. In this era of economic awareness in the healthcare system, use of microvascular reconstruction needs to be justified if other comparable and less expensive alternatives are available. The present study aimed to review the literature comparing FF to PF for reconstruction of oncologic head and neck defects and determine the relative benefits and drawbacks of both flap types.

To our knowledge, this is the first systematic review of studies comparing the postoperative complications and outcomes of FF and PF for head and neck reconstruction of oncologic defects.

The major findings of the present study are that: (a) FF was associated with a longer operating time and, in general, a higher cost compared to PF, including compared to SCAIF. (b) FF was associated with a lower hospitalization stay compared to PMMF, but a higher hospitalization stays when compared to SCAIF and SMIF. (c) Recipient site morbidity was lower with FF reconstruction compared to PMMF, including a lower incidence of infection, dehiscence, and necrosis. The incidence of hematoma and fistula were equivocal. (d) Donor site morbidity was equivocal between FF and PF reconstruction, with no distinction in the rate of infection, dehiscence, and hematoma. (e) Revision surgery was higher with FF reconstruction compared to PF and SMIF. (f) Speech quality was better with FF than with PMMF for oral cavity defects, and FF and PMMF scored similarly on global quality of life, pain, swallowing, chewing, speech, activity, recreation, taste, saliva, anxiety and composite score.

Those conclusions are drawn from retrospective studies lacking methodological homogeneity, thus limiting a truly valid comparison between FF and PF reconstruction. The main issues that need to be further address are the inherent differences among the studied groups in term of patients' preoperative characteristics and defect locations. The findings of the studies included in this review can result from surgeon's bias itself opting for either a FF or a PF based on patient's characteristics and considering it more suitable for a certain location. In fact, patients in the FF group were younger than patients in the PF group with more than a 10-year age difference noted in some studies [11, 12]. Distal extremity reconstruction donor sites are thought to be affected by the patients' health status and age in relation to the condition of peripheral vessels. However, according to several studies age is not considered a risk factor for FF failure [36]. FF reconstruction was also considered with favorable long-term outcomes in patients of 90 years old in a study by Wester et al. [37].

Overall, only a minority of studies showed significant differences in the preoperative characteristics of FF and PF groups. The patients characteristics and T stage were similar between FF and PF groups in most of the studies with a few exceptions. In those, some even showed opposite findings, as it is the case for the ASA class and the incidence of diabetes mellitus [7, 10, 12, 20]. Thus, in front of a

majority of studies with similar baseline characteristics between the PF and FF groups, we could extrapolate with caution that the intrinsic flaps characteristics have an essential contribution to the surgical outcomes depicted in these studies.

A unanimous finding among all studies in this review was the longer operative (OR) time necessary for FF reconstruction which was frequently explained by the microvascular anastomosis. Interestingly, four distinct articles mentioned longer hospitalization time for PF when compared to PMMF [8, 13, 20, 33]. The higher complication rate in PMMF and the poorer patients' preoperative health status in two of those studies may explain this finding [25, 33]. In contrast, SMIF and SCAIF showed a shorter hospitalization and ICU length of stay when compared to FF in similar patients groups [7, 21, 23, 32].

Cost analysis favour PF over FF in a study focusing on the SCAIF [5]. The study by Forner et al. also showed a favorable cost-analysis for SMIF over RFFF but no statistical analysis was provided to be able to conclude on a significant difference [27]. Conclusions on the relative cost of PMMF are harder to draw because studies are showing divergent results [11, 13, 24]. The differences in costs for the PMMF between studies can be explained by the different indications for the use of this flap by the authors. In an era of limited resources and increased attention to health economics, cost analysis studies should be encouraged.

Studies demonstrating significant differences in complication rates were all specifically comparing PMMF to FF. In fact, there was a higher incidence of overall complications, recipient site infection, dehiscence of recipient site, necrosis and flap failure with PMMF reconstruction in all articles except one. Geiger et al. [17] presented different results with regards to fistula, dehiscence and osteonecrosis rates. It is important to note, however, that the authors compared RFFF to PMMF only in the presence of intraoperative brachytherapy implants. These implants, which supplied high doses of radiation, may have led to direct tissue damage in the thinner free flaps, subsequently leading to a higher risk of fistula and dehiscence. The authors themselves associated the lower complication rate of the PMMF group to their increased bulk.

When comparing ALT to SMIF, Howard et al. [21] showed a higher complication rate as well as higher operative revision rates when using the ALT. Similarly, Zhang et al. [23] demonstrated higher rates of donor site complications in the RFFF when compared to SCAIF. Paydarfar et al. [23] demonstrated

higher recipient and donor site complications when comparing RFFF to SMIF (no *p*-values were available); this latter flap has previously been cited as having a low donor site morbidity in another study [38].

PMMF was associated with poorer QoL outcomes when compared to FF [15, 16, 25, 32]. Tsue and al [24], even demonstrated a lower capacity to progress to a regular diet following oral cavity and oropharyngeal reconstruction. This was corroborated by Chepeha et al. [8] who showed a higher incidence of gastrostomy tube dependence after PMMF which they attributed to the flap's downward pull, small size, a limited axis of rotation and inability to fold.

Thereby, PMMF seem to be inferior to FF or other pedicled flaps on many different levels. However, we must remember that higher ASA classes were more common in the PMMF groups representing a considerable bias in the literature. It is our opinion that PMMF should still be considered a reliable and useful flap, especially in a salvage surgery setting.

The present review did not allow us to find any comparative study between osseous or composite FF and osseous or composite PF. Composite head and neck defects have previously been reconstructed with PF including a bony component such as the pectoralis major osteomyocutaneous flap with rib or sternum, the sternocleidomastoid flap with part of the clavicle, or the trapezius flap with the scapular spine. These flaps did not withstand the test of time because of their lack of robustness, reliability and versatility in comparison to their homologue free flaps [39–43]. The more recent SMIF has also been used as a composite flap for mandible, maxilla and orbital defects reconstruction [10]. Yet, its role in osseous reconstruction remains to be defined in an era dominated by FF. Despite the lack of comparative studies, we can safely state that FF are superior to PF for bony reconstructions, especially in radiated patients.

This review suggests that SMIF and SCAIF can be considered reasonable alternatives to free flaps for the reconstruction of head and neck tissue defects given the similar functional outcomes and better performance in OR time, hospitalization/ICU length, and cost. Some articles described their use more frequently in higher ASA classes which further highlights their utility. They can usually be closed primarily and do not typically require skin grafting [38]. Furthermore, SMIF has also been cited as having superior color matching for cervicofacial skin defects [44]. Nonetheless, SMIF and SCAIF are not

suitable for all head and neck defects. Patients with previous history of radiation or ipsilateral neck dissection are not optimal candidates [21]. Additionally, reconstruction of the midface or upper face can sometimes be limited by the length of their respective pedicles. Finally, the use of SMIF in patients requiring level I neck dissection is still debated [21], as its oncological safety and the potential risk to transfer cervical neoplastic cells to the recipient site is controversial in the literature [45]. However, recurrence is thought to be due to the aggressiveness of the resected tumor than the flap itself [46]. A careful flap dissection, at the subplatysmal plane, after completing the neck dissection, helps minimize the risk of tumor spread [47]. The SMIF is a reliable reconstruction technique if level I, A and B, nodes are thoroughly removed, as supported by Howard et al. 11-years case-series study, where no recurrences related to the SMIF transfer of metastatic tissue were noted [21]. Still, SMIF should not be performed in the presence of clinical or radiographic evidence of level 1 cervical lymph node disease [48].

Limitation

Of the thirty studies reviewed, some do not specify the primary tumor location and consequently the defect site. Many articles did not mention the specific flaps used and lacked standard definitions for post-operative complications and outcomes. Furthermore, retained articles were for the majority retrospective studies and comprised risk bias, as assessed by the MINORS criteria. These factors limited the authors ability to analyze specific discordant results between articles and to draw robust conclusions from this systematic review.

Conclusion

The articles included in this review are lacking of methodological homogeneity. Their retrospective nature and the inherent disparities in term of pre-operative characteristics between the groups in some studies are limiting. Although the conclusions should be interpreted with caution, it is safe to assume that free flaps are an excellent choice for reconstruction in relatively healthy subjects with low ASA classes. It appears that FF are superior to the PMMF for several postoperative outcomes. However, other pedicled flaps such as the SMIF and SCAIF compare favorably to FF for some specific indications achieving similar outcomes at a lower cost.

Appendix
Table 7 Methodological Quality of Included Studies by MINORS Criteria

Study, Year	Type of Study	Score	Methodological items for non-randomized studies										Additional criteria in the case of comparative study				
			Clearly Stated Aim	Inclusion of Consecutive Patients	Prospective Collection of Data	Endpoints Appropriate to Aim of Study	Unbiased Assessment of Study Endpoint	Appropriate Follow-up Period to Aim of Study	Loss to Follow-up <5%	Prospective Calculation of Study Size	An adequate control group	Contemporary groups	Baseline equivalence of groups	Adequate statistical analyses			
Sinha, 2017	Retrospective review	20	2	2	2	2	0	2	2	2	2	2	2	2	0	0	2
Goyal, 2017	Retrospective review	18	2	2	2	2	0	0	2	2	2	0	0	2	0	0	2
Li, 2016	Retrospective review	19	2	2	2	2	0	0	2	2	2	0	0	2	1	1	2
Kozin, 2016	Retrospective review	20	2	2	2	2	0	0	2	2	2	0	0	2	2	2	2
Howard, 2016	Retrospective review	18	2	2	2	2	0	0	2	2	2	0	0	2	0	0	2
Geiger, 2016	Retrospective review	20	2	2	2	2	0	0	2	2	2	0	0	2	2	2	2
Gao, 2016	Retrospective review	16	2	0	2	2	0	0	2	2	2	0	0	2	2	0	2
Fomer, 2016	Retrospective review	20	2	2	2	2	0	0	2	2	2	0	0	2	2	2	2
Zhang S, 2015	Retrospective review	19	2	2	2	2	0	0	2	2	2	0	0	2	1	1	2
Zhang X, 2014	Retrospective review	19	2	2	2	2	0	0	2	2	2	1	0	2	2	2	2
Jing, 2014	Retrospective review	17	1	2	2	2	0	0	2	2	2	0	0	2	2	0	2
Granzow, 2013	Retrospective review	18	1	2	2	2	0	0	2	2	2	0	0	2	1	1	2
Deqanello, 2013	Retrospective review	18	2	2	2	2	0	0	2	2	2	0	0	2	0	0	2
Fang, 2013	Retrospective review	18	2	2	2	2	0	0	2	2	2	0	0	2	0	0	2
Paydarfar, 2011	Retrospective review	20	2	2	2	2	0	0	2	2	2	0	0	2	2	2	2
Hsing, 2011	Retrospective review	18	2	2	2	2	0	0	2	2	2	0	0	2	0	0	2
Chan Y, 2011	Prospective	18	2	2	2	2	0	0	2	2	2	0	0	2	0	0	2
Dermirtas, 2010	Retrospective review	20	2	2	2	2	0	0	2	2	2	0	0	2	2	2	2

Table 7 Methodological Quality of Included Studies by MINORS Criteria (Continued)

Study, Year	Type of Study	Score	Methodological items for non-randomized studies										Additional criteria in the case of comparative study			
			Clearly Stated Aim	Inclusion of Consecutive Patients	Prospective Collection of Data	Appropriate Endpoints to Aim of Study	Unbiased Assessment of Study Endpoint	Appropriate Follow-up Period to Aim of Study	Loss to Follow-up <5%	Prospective Calculation of Study Size	An adequate control group	Contemporary groups	Baseline equivalence of groups	Adequate statistical analyses		
O'Neil, 2010	Retrospective review	20	2	2	2	2	0	1	2	2	2	2	2	2	1	2
Mallet, 2009	Retrospective review	18	2	2	2	2	0	1	2	0	0	2	2	2	1	2
de Bree, 2007	Retrospective review – Matched cohort	19	2	2	2	2	0	2	1	0	0	2	2	2	2	2
Smeele, 2006	Retrospective review – Matched cohort	20	2	2	2	2	0	2	2	0	0	2	2	2	2	2
Chien, 2005	Case series	17	2	2	2	2	0	1	2	0	0	2	2	0	0	2
Chepeha, 2004	Retrospective review	19	2	2	2	2	0	1	2	0	0	2	2	2	2	2
Funk, 2002	Case-control study	20	2	2	2	2	0	2	2	0	0	2	2	2	2	2
Petrizzelli, 2002	Retrospective review	18	2	2	2	2	0	2	2	0	0	2	2	2	0	2
Amarante, 2000	Case-series	6	1	0	2	1	0	0	0	0	0	2	0	0	0	0
Tsue, 1997	Retrospective review	19	2	2	2	2	0	2	2	0	0	2	2	2	1	2
Kroll, 1997	Retrospective review	17	2	2	2	2	0	2	2	0	0	2	2	2	1	0
Kroll 1992	Retrospective review	18	2	2	2	2	0	2	2	0	0	2	2	2	0	2

The items are scored 0 (not reported), 1 (reported but inadequate) or 2 (reported and adequate). The global ideal score being 16 for non-randomized studies and 24 for comparative studies

Table 8 Defect location

Articles	Defect / tumor location	FF	PF	P-value
Goyal, 2017	Cutaneous/lateral skull base	60 (10.2%)	56 (26.9%)	< 0.001
	Oral cavity	154 (26.2%)	32 (15.4%)	
	Oropharynx	42 (7.1%)	4 (1.9%)	
	Laryngectomy/pharyngectomy	109 (18.5%)	58 (27.9%)	
	Mandibular	96 (16.3%)	9 (4.3%)	
	Sinonasal	49 (8.3%)	9 (4.3%)	
	Composite/multiple sites	79 (13.4%)	40 (19.2%)	
	Cutaneous defect	6 (21.4%)	15 (33.3%)	
	Parotid/temporal bone	9 (32.1%)	16 (35.6%)	
	Total auriculectomy	4 (44%)	8 (50%)	
Kozin, 2016	Lateral TBR	7 (78%)	14 (88%)	0.367
	Subtotal TBR	2 (22%)	5 (83%)	0.071
	Facial nerve sacrifice	6 (67%)	14 (88%)	0.740
Howard, 2016	Tongue	18(75.00%)	12(70.59%)	0.016
	Tongue and FOM	7 (46.7%) ^b	7 (58.3%) ^b	
	Tongue	8 (53.3%) ^b	5 (41.7) ^b	
	FOM	43 (65.2%)	23 (34.8%)	
	Gum	21 (80.8%)	5 (19.2%)	
	Buccal	9 (90%)	1 (10%)	
	Palate	4 (66.7%)	2 (33.3%)	
	Total laryngectomy	2 (100%)	0 (0)	
	Oral cavity	22 (100%) ^b	27 (100%) ^b	
	Oropharynx	9 (56.25%) ^b	11 (55%) ^b	
Li, 2016 ^a	Buccal defects	7 (43.75%) ^b	9 (45.0%) ^b	ND
		20	24	
		12		
		35 (28.9%)	86 (71.1%)	
Zhang S, 2015	Tongue	7 (53.8%)	6 (46.2%)	< 0.001
	FOM	17 (94.4%)	1 (5.6%)	
	Lip			
Zhang X, 2014 ^a				0.331
Jing, 2014				ND
Deganello, 2013				ND
Fang, 2013				ND
Hsing, 2011 ^a				< 0.001

Table 8 Defect location (Continued)

Articles	Defect / tumor location	FF	PF	p-value
Paydarfar, 2011	Gum	15 (34.9%)	28 (65.1%)	
	Buccal	101 (36.9%)	173 (63.1%)	
	Palate	7 (58.1%)	5 (41.7%)	
	Retromolar trigone	4 (40%)	6 (60%)	
	Tongue	14 (42.4%) ^b	12 (44.4%) ^b	0.90
	Tongue and FOM	10 (30.3%) ^b	9 (33.3%) ^b	ND
	FOM	9 (27.3%) ^b	6 (22.2%) ^b	0.90
	Hypopharynx	24 (100%) ^b	92 (100%) ^b	ND
		86 (100%) ^b		
		16 (64%)	20 (44%)	0.17
Mallet, 2009	Oral tongue (OT)			
	Base of tongue	2 (8%)	11 (24%)	
	Lateral tongue	8 (20%)	8 (20%)	ND
	FOM	10 (25%)	10 (25%)	
	Base of tongue	3 (8%)	3 (8%)	
de Bree, 2007 ^a	Buccal mucosal	11 (100%)	16 (100%)	ND
Chien, 2005	Oral cavity	38 (54%)	52 (48%)	NS
Chepeha, 2004	Oropharynx	16 (23%)	45 (42%)	
	Hypopharynx	10 (14%)	11 (10%)	
	Neck	2 (3%)	2 (2%)	
	Other	6 (10%)	10 (9%)	
Petruzzelli, 2002	Upper aerodigestive tract + tracheotomy	24 (100%)	15 (100%)	ND
Amarante, 2000	Orbital	-	6 (8.8%) ^b	ND
	Parotid	-	3 (4.4%) ^b	
	Middle 1/3 skull base	11 (22.4%) ^b	-	
	Oropharynx	-	4 (5.9%) ^b	
	Pharyngo-laryngeal	-	34 (50%) ^b	
	Pharynx cer. Esoph.	3 (6.1%) ^b	-	
	Mandibular	-	3 (4.4%) ^b	
	Cervical	-	1 (1.5%) ^b	
	Brachial plexus	1 (2.0%) ^b	-	

Table 8 Defect location (Continued)

Articles	Defect / tumor location	FF	PF	p-value
Tsue, 1997 ^a	Tongue	1 (20%)	1 (80%)	ND
	FOM	1 (20%)	4 (80%)	
	Lateral and posterior oropharynx	1 (25%)	3 (75%)	
	Base of tongue	15 (76%)	4 (24%)	
	Retromolar trigone or Alveolus	10 (71%)	4 (28%)	
	Tonsil	3 (27%)	8 (73%)	

TBR Temporal bone resection, FOM Floor of the mouth

^a The article describes the tumor location instead of the defect location

^b Percentage calculated relying on the data presented. Percentage not provided by the article

Bold = Statistically significant, p-value ≤ 0.05

Table 9 Demographic data

	Article	FF	PF	p-value	
Age, mean ± SD or mean (range)	Goyal, 2017	64.0 ± 12.0	66.5 ± 12.9	0.017	
	Kozin, 2016	64 ± 10.1	67.2 ± 10	0.1876	
	Forner, 2016	65	63	NS	
	Jing, 2014	64.8 (50–84)	67.4 (44–89)	ND	
	Granzow, 2013	58.75 (44–77)	55.61 (34–83)	0.39	
	Deganello, 2013	58.2 ± 6.32	69.6 ± 6.8	< 0.01	
	Fang, 2013	58.0 (25–78)	72.4 (55–80)	< 0.001	
		57.2 (46–72)		< 0.001	
	Paydarfar, 2011	54.7 (38–75)	58.1 (34–82)	0.24	
	Hsing, 2011	54.1 ± 9.6	54.5 ± 12.5	0.839	
	Demirtas, 2010	60 (38–73)	65.3 (49–82)	ND	
			58.5 (17.3%)	ND	
	Mallet, 2009	52.7 ± 9.3	56.8 ± 8.7	0.07	
	Chien, 2005	53.5 (40–77)	50.1(38–66)	ND	
	Chepeha, 2004	58	58	NS	
	Petruzzelli, 2002	58.2 (9.8)	64 (12.8)	ND	
	Amarante, 2000	15–81	37–71	ND	
	Kroll, 1997	56 ± 13	62 ± 12	0.0046	
	Age (median)	Sinha, 2017	65.9 (57.7–74.2)	67.9 (60.3–76.8)	0.037
		Howard, 2016	68.3	76.9	0.172 ^a
			73.3		
Age > 50 years	Tsue, 1997	63 (62 ± 2)	64 ^a (64 ± 2)	ND	
	Li, 2016	9(27.50%)	3(17.65%)	0.304	
	Gieger, 2016	82%	85.4%	0.661	
	Zhang S, 2015	12 (80%) ^a	10 (83.33%) ^a	1.00	
	Zhang X, 2014	19 (76%)	6 (24%)	0.597	
Male n (%)	Sinha, 2017	63.0%	71.40%	0.0794	
	Kozin, 2016	78.6%	68.9%	0.43	
	Li, 2016	17(70.83%)	17(100%)	0.043	
	Gieger, 2016	68.0%	76.4%	0.338	
	Howard, 2016	7 (77.8)	15 (93.8)	0.240	
		7 (77.8)	6 (100)		
	Forner, 2016	75%	43%	NS	
	Zhang S, 2015	10 (66.7%) ^a	9 (75%) ^a	0.69	
	Zhang X, 2014	66 (88%)	31 (32.0%)	0.018	
	Jing, 2014	19 (86.4%) ^a	26 (96.3%) ^a	ND	
	Granzow, 2013	11 (69%)	13 (72%)	1.0	
	Deganello, 2013	12 (75)	16 (80)	0.88	
	Fang, 2013	9 (45%)	11 (45.83%)	ND	
		8 (66.67%)		ND	
	Paydarfar, 2011	22 (66.7%) ^a	20 (90.1%) ^a	0.38	
	Hsing, 2011	39 (40.2%)	58 (59.8%)	0.745	
	Demirtas, 2010	7 (58.33)	8 (100%)	ND	
			7 (100%)	ND	

Table 9 Demographic data (Continued)

	Article	FF	PF	p-value
	Mallet, 2009	19 (76%)	38 (84.4%)	0.58
	Chien, 2005	11(100%)	14 (87.5%)	ND
	Chepeha, 2004	54 (76%)	84 (78%)	NS
	Amarante, 2000	36 (73.47%)	49 (72.06%)	ND
	Tsue, 1997	16 (67%)	8 (27.6%)	ND

^aPercentage calculated relying on the data presented. Percentage not provided by the article

Bold = Statistically significant, p-value ≤ 0.05

Table 10 Preoperative risk factors

	Article	FF	PF	p-value
Smoking	Gieger, 2016	62.0%	61.8%	0.985
	Jing, 2014	4 (18.18%) ^a	1 (3.70%) ^a	ND
	Granzow, 2013	9 (56%)	13 (72%)	0.5
	Mallet, 2009	23 (92%)	43 (98%)	ND
	Funk, 2002	13 (61.9%)	18 (85.7%)	0.159
Prior head and neck surgery	Goyal, 2017	189 (32.1%)	122 (58.7%)	< 0.001
	Gieger, 2016 ^a	14.0%	30.9%	0.039
	Chepeha, 2004	25 (35%)	31 (29%)	ND
	Tsue, 1997	5 (17%)	11 (46%)	0.05
Systemic diseases (CVD, HTA, DM)	Fang, 2013	4 (20%)	20 (83.33%)	< 0.001
		3 (25%)		< 0.001
COPD	Sinha, 2017	12.2%	15%	0.4538
	Mallet, 2009	6 (24%)	10 (22%)	0.96
DM	Sinha, 2017	17.7%	15.8%	0.6900
	Granzow, 2013	5 (31%)	0	0.02
	Hsing, 2011	2 (33.3%)	4 (66.7%)	0.986
HTA	Gieger, 2016	56.0%	58.2%	0.821
	Granzow, 2013	6 (38)	5 (28)	0.7
CAD	Sinha, 2017	15.6%	18.8%	0.4162
	Granzow, 2013	1 (6%)	1 (6%)	1.0
	Mallet, 2009	6 (24%)	16 (36%)	0.46
DLP	Gieger, 2016	34.0%	34.6%	0.953
CHF	Sinha, 2017	3.4%	4.5%	0.5939
aFIB	Sinha, 2017	7.0%	15.0%	0.0083
Alcoolism	Mallet, 2009	24 (96%)	41 (95%)	ND
Other cancer	Mallet, 2009	3 (12%)	10 (22%)	0.46
Kaplan-Feinstein grade (severe comorbidity)	Funk, 2002	1 (4.8%)	5 (23.8%)	0.196

aFIB atrial fibrillation, *CAD* Cardiac artery disease, *CHF* Chronic heart failure, *COPD* Chronic obstructive pulmonary disease, *CVD* Cardiovascular disease, *DM* Diabetes mellitus, *DLP* Dyslipidemia, *HTA* Hypertension

Bold = Statistically significant, p-value ≤ 0.05

Table 11 ASA class and risk

Article	FF	PF	p-value	
ASA class I-II n, (%)	Goyal, 2017	245 (41.6%)	56 (26.9%)	0.001
	Granzow, 2013	3 (18.75%) ^a	3 (16.67%) ^a	0.5
	Demirtas, 2010	9 (75%) ^a	5 (62.5%) ^a	ND
			5 (71.4%) ^a	ND
	Mallet, 2009	18 (75%)	33 (73.33%)	1.00
	de Bree, 2007	32 (80%)	37 (92.5%)	0.028
	Funk, 2002	10 (47.62%)	12 (57.14%)	0.226
ASA class III-IV n, (%)	Goyal, 2017	343 (58.2%)	152 (73.1%)	0.001
	Granzow, 2013	13 (81.25%) ^a	15 (83.33%) ^a	0.5
	Demirtas, 2010	3 (25%) ^a	3 (37.5%) ^a	ND
			2 (28.57%) ^a	ND
	Mallet, 2009	6 (25%)	12 (27%)	1.00
	de Bree, 2007	8 (20%)	3 (8%)	0.028
	Funk, 2002	11 (52.38%)	9 (42.86%)	0.226
ASA risk factor (mean ± SD)	Sinha, 2017	2.6 ± 0.03	2.8 ± 0.05	0.0007
	Forner, 2016	2.3	2.4	0.05
	Demirtas, 2010	2.25 ± 0.4	2.37 ± 0.5	ND
			2.28 ± 0.5	ND
	Kroll, 1997	2.74 ± 0.61	2.80 ± 0.68	0.8011

ASA risk factor scored using the American Society of Anesthesiology Scale (ASA)

^aPercentage calculated relying on the data presented. Percentage not provided by the article

Bold = Statistically significant, p-value ≤ 0.05

Table 12 Staging and treatment data

Article	FF	PF	p-value		
Prior radiation	Goyal, 2017	272 (46.2%)	130 (62.5%)	< 0.001	
	Kozin, 2016	12 (46.2%)	22 (48.9%)	0.824	
	Howard, 2016	6 (67%)	7 (44%)	0.071	
			3 (50%)		
	Gieger, 2016	98.0%	94.6%	0.356	
	Jing, 2014	15 (68.2%) ^a	23 (85.2%) ^a	ND	
	Granzow, 2013	4 (25%)	7 (39%)	0.5	
	Paydarfar, 2011	4 (12%) ^a	0	0.14	
	Mallet, 2009	3 (12%)	15 (33%)	0.09	
	Chepeha, 2004	37 (52%)	56 (52%)	NS	
	Tsue, 1997	8 (28%)	13 (54%)	0.05	
	Kroll, 1997	49 (33.8%)	9 (28.1%)	0.5360	
	Prior chemotherapy	Kozin, 2016	7 (26.9%)	11 (25%)	0.859
		Jing, 2014	7 (31.8%) ^a	4 (14.8%) ^a	ND
Granzow, 2013		1 (6%)	6 (33%)	0.09	
Paydarfar, 2011		1 (3%) ^a	1 (3.7%) ^a	0.14	
Tsue, 1997		0	1 (4%)	ND	
T1	Zhang X, 2014	1 (50%)	1 (50%)	0.369	
	Jing, 2014	7 (31.3%) ^a	8 (29.6%) ^a	ND	
	Deganello, 2013	0	4 (20%) ^a	< 0.01	

Table 12 Staging and treatment data (Continued)

	Article	FF	PF	p-value	
T2	Fang, 2013	0	3 (15%) ^a	ND	
		0		ND	
	Paydarfar, 2011	1 (3.0%) ^a	2 (7.4%) ^a	0.38	
	Hsing, 2011	46 (42.2%)	63 (57.8%)	0.904	
	Mallet, 2009	32% (8/25)	44% (20/45)	0.44	
	Chien, 2005	0	9 (56.25%) ^a	NS	
	Funk, 2002	1 (4.8%)	1 (4.8%)	NS	
	Tsue, 1997	11 (37.93%) ^a	6 (25%) ^a	ND	
	Zhang X, 2014	10 (76.9%)	3 (23.1%)	0.369	
	Jing, 2014	6 (27.3%) ^a	10 (37.0%) ^a	ND	
	Deganello, 2013	7 (43.8%) ^a	5 (25%) ^a	< 0.01	
	Fang, 2013	7 (35%) ^a	12 (50%) ^a	ND	
	T3		4 (33.33%) ^a		
Paydarfar, 2011		15 (45.45%) ^a	17 (62.96%) ^a	0.38	
Hsing, 2011		90 (35.3%)	165 (64.7%)	0.621	
Chien, 2005		6 (54.5%) ^a	7 (43.75%) ^a	NS	
Funk, 2002		1 (4.8%)	0	NS	
Zhang X, 2014		37 (66.1%)	19 (33.9%)	0.369	
Jing, 2014		7 (31.8%) ^a	2 (7.4%) ^a	ND	
Deganello, 2013		8 (50%)	8 (40%)	< 0.01	
Fang, 2013		3 (15%)	6 (25%)	ND	
		2 (16.7%)			
Paydarfar, 2011		11 (33.33%) ^a	6 (22.22%) ^a	0.38	
Hsing, 2011		38 (40.0%)	57 (60.0%)	0.621	
T4		Chien, 2005	5 (45.45%) ^a	0	NS
	Funk, 2002	5 (23.8%)	5 (23.8%)	NS	
	Zhang X, 2014	31 (79.5%)	8 (20.5%)	0.369	
	Jing, 2014	1 (4.5%) ^a	7 (25.9%) ^a	ND	
	Deganello, 2013	1 (6.25%)	3 (15%)	< 0.01	
	Fang, 2013	9 (45%)	12.5%	ND	
		6 (50%)			
	Paydarfar, 2011	6 (18.18%) ^a	2 (7.4%) ^a	0.38	
	Hsing, 2011	12 (37.5%)	20 (62.5%)	0.621	
	Chien, 2005	0	0	NS	
	Funk, 2002	14 (66.6%)	15 (71.4%)	NS	
	T1-T2	Li, 2016	10 (41.67%)	8 (47.06%)	0.981
		Forner, 2016	5 (41.7%) ^a	6 (66.6%) ^a	ND
Mallet, 2009		8 (32%)	20 (44%)	0.44	
Tsue, 1997		11 (37.93%) ^a	6 (25%) ^a	ND	
T3-T4		Li, 2016	14(58.33%)	9(52.94%)	0.981
		Forner, 2016	7 (58.3%) ^a	3 (33.3%) ^a	ND
		Mallet, 2009	17 (68%)	25 (56%)	0.44
	Tsue, 1997	17 (58.6%) ^a	18 (75%) ^a	ND	
Stage I-II	Chepeha, 2004	16 (23%)	23 (21%)	NS	

Table 12 Staging and treatment data (Continued)

	Article	FF	PF	p-value
Stage III-IV	Chepeha, 2004	85 (79%)	55 (77%)	NS
Surg +chemoradio	Li, 2016	8(33.33%)	7(41.18%)	0.854
	Fang, 2013	4 (20%)	5 (20.83%)	ND
		2 (16.67%)		ND
	Hsing, 2011	25 (39.7%)	38 (60.3%)	0.687
	de Bree, 2007	40 (100%)	37 (92.5%)	ND
	Chepeha, 2004	27 (38%)	46 (43%)	NS
	Paydarfar, 2011	16 (48.48%) ^a	12 (44.44%) ^a	0.03
Surg + radio	Zhang X, 2014	41 (51.9%)	17 (54.84%)	0.781
	Paydarfar, 2011	13 (39.39%) ^a	5 (18.5%) ^a	0.3
	Hsing, 2011	25 (39.7%)	38 (60.3%)	0.687
	Chepeha, 2004	27 (38%)	46 (43%)	NS
	Tumor stage	Kroll, 1997	2.80 ± 0.94	3.14 ± 0.83
Tumor recurrence	Kroll, 1997	44(32.6%)	11 (33.3%)	0.9350

Surg + chemoradio: Surgical resection and adjuvant chemoradiotherapy

Surg + radio: Surgical resection and adjuvant radiotherapy

^aPercentage calculated relying on the data presented. Percentage not provided by the article

Bold = Statistically significant, p-value ≤ 0.05

Table 13 OR time, Hospital and ICU length and hospital cost

	Article	FF	PF	p-value	
OR time, min (mean ± SD)	Sinha, 2017	421.4 ± 4.4	332.7 ± 10.7 min	0.0001	
	Goyal, 2017	427.2 ± 92.3	310.8 ± 125.0	0.001	
	Li, 2016	405 ± 107	365 ± 48	< 0.05	
	Howard, 2016	683 (575–979)	544 (396–700)	0.00817	
			601 (474–841)	0.2488	
	Forner, 2016	552	347	< 0.05	
	Zhang S, 2015	81 ± 8	55 ± 7	0.05	
	Granzow, 2013	816.3 ± 148.9	587.9 ± 130.5	0.0002	
	Deganello, 2013	570 ± 96	444 ± 54	0.14	
	Paydarfar, 2011	780	506.4	0.001	
	Demirtas, 2010	204 ± 52	111 ± 18	ND	
			153 ± 18	ND	
	de Bree, 2007	692	462	< 0.005	
	Tsue, 1997	684 ± 16	666 ± 20	0.003	
	OR time, hour (mean ± SD)	Kozin, 2016	8.1	6.7	0.002
Fang, 2013		8.2 ± 2.7	5.4 ± 1.0	NS	
		8.4 ± 1.7		NS	
Mallet, 2009		7.01 ± 1.19)	4.19 ± 0.57	< 0.001	
Smeele, 2006		12.5 ± 1.9	9.9 ± 1.5	< 0.0001	
Kroll, 1997		10.49 ± 2.06	9.39 ± 2.59	0.029	
Kroll, 1992		8.7	8.3	NS	
OR time of > 600 min		Zhang X, 2014	59 (74.68%)	3 (9.68%)	0.001
Hospit length, days (mean ± SD)		Kozin, 2016	11	8.8	0.12
		Howard, 2016	9.8 (7–22)	4.75 (2–14)	0.00424
6.5 (6–7)	0.15886				

Table 13 OR time, Hospital and ICU length and hospital cost (Continued)

	Article	FF	PF	p-value
	Forner, 2016	15.4	12.4	> 0.05
	Zhang S, 2015	17 ± 2.5	12 ± 1.7	< 0.05
	Granzow, 2013	18.50 (9–59)	16.4 (5–57)	0.4
	Deganello, 2013	23.2 ± 7.5	26.5 ± 9.9	0.63
	Paydarfar, 2011	14.0	10.6	0.008
	Demirtas, 2010	16.0 ± 8.7	15.2 ± 8.7	ND
			17.0 ± 4.7	ND
	O'Neil, 2010	34.298	29.649	0.184
	Mallet, 2009	18.1 ± 6.7	23.2 ± 14.1	0.10
	de Bree, 2007	24	28	0.005
	Smeele, 2006	22.9 ± 11.5	22.8 ± 15.7	NS
	Chepeha, 2004	12	14	0.006
	Petruzzelli, 2002	6.9 ± 1.9	7.5 ± 4.3	ND
	Tsue, 1997	13 ± 1.4	13 ± 1.1	NS
	Kroll, 1997	13.2 ± 5.4	19.8 ± 11.5	0.003
	Kroll, 1992	11.3	21.2	0.003
ICU length, days (mean ± SD)	Forner, 2016	4.7	0.14	ND
	Granzow, 2013	5.6 (4–9)	1.8 (0–5)	0.0001
	Smeele, 2006	0.1 ± 0.5	0.28 ± 0.81	NS
	Tsue, 1997	3 ± 0.8	2 ± 0.3	NS
Hospital cost \$	Kozin, 2016	SCAIF 32% less expensive than FTT		0.0001
	Forner, 2016	43,617.60	18,158.40	ND
	Deganello, 2013	22,924	19,872	0.043
	Demirtas, 2010 ^a	2963 ± 1715 ^a	2053 ± 687 ^a	ND
			2924 ± 100 ^a	ND
	de Bree, 2007	48,097	51,963	ND
	Smeele, 2006	23,600 ^b	20,400 ^b	ND
	Petruzzelli, 2002	22,821.04 ± 5062.81 ^a	17,648.20 ± 7817.86 ^a	ND
	Tsue, 1997	50,026 ± 4340	38,246 ± 1440	0.003
	Kroll, 1997	28,460 ± 8435	40,992 ± 1958	0.001

^aUS\$^bCAN\$

Bold = Statistically significant, p-value ≤ 0.05

Table 14 Post-operative complications

	Article	FF	PF	p-value	
Any complications	Gieger, 2016	68.0%	36.4%	0.001	
	Zhang X, 2014	13 (16.46%)	14 (45.16%)	0.002	
	Jing, 2014	8 (36.36%)	12 (44.44%)	ND	
	Granzow, 2013	7 (44%)	7 (39%)	1.0	
	Paydarfar, 2011	25 (75.75%) ^a	10 (37.04%) ^a	ND	
	Amarante, 2000	31%	0		
	Kroll, 1992	4 (13%)	17 (44%)	0.0145	
Infection	Gieger, 2016	16.0%	7.3%	0.170	
	Granzow, 2013	0	1 (6.25%)	1.0	
	Mallet, 2009	4 (16%)	6 (13%)	1.0	
Recipient site infection	Goyal, 2017	84 (14.3%)	19 (9.1%)	0.076	
	Kozin, 2016	2 (7.14%) ^a	2 (4.44%) ^a	0.106	
	Paydarfar, 2011	2 (6.06%) ^a	2 (7.41%) ^a	ND	
	O'Neil, 2010	4 (5.2%)	2 (5.4%)	0.962	
	Mallet, 2009	16% (4/25)	13% (6/45)	1.00	
	Chepeha, 2004	2 (3%)	18 (17%)	< 0.004	
Donor site infection	Goyal, 2017	23 (3.9%)	5 (2.4%)	0.43	
	Kozin, 2016	3 (10.71%) ^a	2 (4.44%) ^a	0.106	
	O'Neil, 2010	1 (1.3%)	1 (2.7%)	0.593	
Donor site morbidity	Chan Y, 2011	1(4.2%)	7(7.6%)	ND	
		2(2.3%)		ND	
Fistula	Goyal, 2017	18 (3.1%)	17 (8.2%)	0.005	
	Geiger, 2016	22.0%	7.3%	0.039	
	Jing, 2014	5 (22.72%) ^a	6 (22.22%) ^a	ND	
	Paydarfar, 2011	5 (15.15%) ^a	0	ND	
	Chan Y, 2011	3 (12.5%)	22 (23.9%)	ND	
		4 (4.6%)		ND	
	O'Neil, 2010	2/77 (2.6)	3/37 (8.1)	0.179	
	Demirtas, 2010	0	1 (12.5%) ^a	ND	
	Mallet, 2009	2 (8%)	4 (9%)	1.00	
	Smeele, 2006	3 (0.09%) ^a	1 (0.03%) ^a	ND	
	Chepeha, 2004	4 (5%)	5 (5%)	0.74	
	Amarante, 2000	0	14 (20.59%) ^a	ND	
	Abscess	Gieger, 2016	4.0%	7.3%	0.477
	Dehiscence recipient or donor site	Gieger, 2016	44.0%	23.6%	0.029
Jing, 2014		1 (4.55%) ^a	0	ND	
O'Neil, 2010		2 (2.6%)	1 (2.7%)	0.974	
Smeele, 2006		4 (12.5%) ^a	1 (1.47%) ^a	ND	
Dehiscence recipient site	Kozin, 2016	4 (14.29%) ^a	6 (13.33%) ^a	NS	
	Granzow, 2013	0	1 (5.55%) ^a	1.0	
	Paydarfar, 2011	5 (15.15%) ^a	0	ND	
	Smeele, 2006	1 (3.13%) ^a	1 (3.13%) ^a	ND	
	Chepeha, 2004	0	11 (10%)	< 0.008	

Table 14 Post-operative complications (*Continued*)

	Article	FF	PF	p-value
Dehiscence donor site	Kozin, 2016	0	4 (8.89%) ^a	NS
	Zhang S, 2015	1 (6.67%) ^a	0	ND
	Granzow, 2013	3 (18.75%) ^a	2 (11.11%) ^a	0.6
	Paydarfar, 2011	0	2 (9.09%) ^a	ND
	Smeele, 2006	3 (9.38%) ^a	0	ND
	Kozin, 2016	3 (10.71%) ^a	0	NS
	Jing, 2014	0	4 (14.81%) ^a	ND
	Granzow, 2013	0	0	1.0
	Demirtas, 2010	1 (8.33%) ^a	0	ND
	Hematoma	Chepeha, 2004	4 (6%)	4 (4%)
O'Neil, 2010		1 (1.3%)	1 (2.7%)	0.593
Hematoma Donor site	Howard, 2016	1 (11.11%) ^a	1 (6.25%) ^a	ND
			0	ND
	Smeele, 2006	0	1 (3.13%) ^a	ND
Hematoma recipient site	Paydarfar, 2011	2 (6.06%) ^a	2 (9.09%) ^a	ND
	Smeele, 2006	1 (3.13%) ^a	1 (3.13%) ^a	ND
Partial flap necrosis	Zhang S, 2015	0	2 (16.67%) ^a	ND
	Paydarfar, 2011	0	3 (13.64%) ^a	ND
	Smeele, 2006	1 (3.125%) ^a	2 (6.25%) ^a	NS
	Chepeha, 2004	2 (2.82%) ^a	12 (11%) ^a	< 0.006
	Amarante, 2000	0	11 (16.2%) ^a	ND
	Kroll, 1992	0	4 (10%)	0.1979
Total flap necrosis	Smeele, 2006	2 (6.25%) ^a	1 (3.125%) ^a	
	Amarante, 2000	3 (6.12%) ^a	1 (1.5%) ^a	ND
Partial or total flap necrosis	Mallet, 2009	1 (4%)	14 (31%)	0.02
Osteonecrosis	Goyal, 2017	36 (6.1%)	17 (8.2%)	0.33
	Gieger, 2016	24.0%	3.6%	0.007
Deep Vein Thrombosis (Inferious member)	Sinha, 2017	1.6%	2.2%	0.2775
	O'Neil, 2010	3 (3.9%)	1 (2.7%)	0.746
Venous obstruction (At site)	Paydarfar, 2011	1 (3.03%) ^a	0	ND
Late anastomotic stricture	Chan Y, 2011	3 (12.5%)	25 (27.2%)	ND
		2 (2.3%)		ND
Blood transfusion	Deganello, 2013	3 (19%)	4 (20%)	0.96

^aPercentage calculated relying on the data presented. Percentage not provided by the article

Bold = Statistically significant, p-value ≤ 0.05

Table 15 Post-operative outcomes

	Article	FF	PF	p-value
Operation revision	Howard, 2016	1.6 (1–3)	0.6 (0–1)	< 0.00001
			1.3 (1–2)	NS
	Gieger, 2016	34%	9.1%	0.003
	Granzow, 2013	4 (25%)	3 (17%)	0.7
	Paydarfar, 2011	2	1	ND
	O'Neil, 2010	13 (16.9%)	3 (8.1%)	0.207
	Demirtas, 2010	3	0	ND
	Chepeha, 2004	6 (8%)	13 (12%)	0.60
	Amarante, 2000	7	0	ND
	Flap failure	Howard, 2016	0	0
			0	NS
Zhang S, 2015		1 (6.7%)	0	ND
Jing, 2014		1	0	ND
Granzow, 2013		1	0	0.5
Paydarfar, 2011		1	0	ND
Mallet, 2009		1 (4%)	14 (31%)	0.02
de Bree, 2007		1	0	ND
Smeele, 2006		2	1	ND
Mortality at 30 days		Jing, 2014	0	1
	Chepeha, 2004	0	6 (6%)	0.08
Mortality at 1-year	Granzow, 2013	0	0	1.0
	Funk, 2002	2 (20%)	6 (28.6%)	ND
Mortality at 2-year	de Bree, 2007	13 (33%)	16 (40%)	0.602

Bold = Statistically significant, p-value ≤ 0.05

Table 16 Quality of Life data

	Article	FF	PF	p-value	
UW-QOL: global	Li, 2016	55.14 ± 9.24	54.36 ± 8.13	0.965	
	Zhang X, 2014	70.5 ± 16.7	67.3 ± 12.9	0.860	
	Hsing, 2011	66.0 ± 18.5	57.8 ± 18.2	0.090	
UW-QOL: Pain	Li, 2016	71.63 ± 9.91	72.94 ± 11.13	0.751	
	Zhang X, 2014	86.2 ± 10.8	89.9 ± 11.4	0.425	
	Hsing, 2011	76.8 ± 23.0	68.1 ± 27.2	0.138	
UW-QOL: Swallowing	Li, 2016	44.00 ± 16.27	43.78 ± 4.95	0.741	
	Zhang X, 2014	49.4 ± 14.7	51.3 ± 21.7	0.840	
	Hsing, 2011	49.3 ± 37.2	48.6 ± 32.7	0.962	
UW-QOL: Chewing	Li, 2016	42.45 ± 6.15	43.43 ± 12.37	0.817	
	Zhang X, 2014	52.6 ± 17.1	59.4 ± 12.9	0.498	
	Hsing, 2011	34.5 ± 39.0	33.6 ± 36.7	0.973	
UW-QOL: Speech	Li, 2016	51.27 ± 11.24	52.63 ± 12.43	0.461	
	Zhang X, 2014	57.5 ± 20.1	76.1 ± 13.3	0.017	
	Hsing, 2011	66.7 ± 27.2	44.7 ± 35.0	0.002	
UW-QOL: Appearance	Li, 2016	57.47 ± 11.44	68.54 ± 13.24	0.0001	
	Zhang X, 2014	76.4 ± 18.6	70.3 ± 17.1	0.308	
	Hsing, 2011	67.3 ± 25.0	69.8 ± 25.5	0.535	
UW-QOL: Activity	Li, 2016	64.23 ± 9.52	63.73 ± 8.41	0.641	
	Zhang X, 2014	71.9 ± 11.5	74.8 ± 10.2	0.710	
	Hsing, 2011	67.9 ± 24.2	66.8 ± 27.9	0.760	
UW-QOL: Recreation	Li, 2016	66.59 ± 11.62	67.26 ± 9.23	0.445	
	Zhang X, 2014	72.1 ± 10.2	78.9 ± 11.2	0.590	
	Hsing, 2011	69.1 ± 32.6	62.5 ± 32.2	0.221	
UW-QOL: shoulder	Li, 2016	61.52 ± 7.83	54.65 ± 11.24	0.0001	
	Zhang X, 2014	87.1 ± 14.4	65.6 ± 20.0	< 0.001	
	Hsing, 2011	81.4 ± 14.7	50.5 ± 29.8	< 0.001	
UW-QOL: Taste	Li, 2016	50.91 ± 10.64	51.24 ± 11.23	0.673	
	Zhang X, 2014	48.4 (18.3)	52.9 (19.6)	0.713	
	Hsing, 2011	55.0 ± 43.2	45.9 ± 39.6	0.226	
UW-QOL: Salive	Li, 2016	45.48 ± 16.92	44.17 ± 12.78	0.723	
	Zhang X, 2014	70.9 ± 9.5	72.3 ± 23.1	0.813	
	Hsing, 2011	71.7 ± 34.8	73.8 ± 28.1	0.964	
UW-QOL: Mood	Li, 2016	69.94 ± 9.51	68.31 ± 14.72	0.474	
	Zhang X, 2014	76.0 ± 14.7	71.6 ± 18.8	0.114	
	Hsing, 2011	76.2 ± 24.7	60.8 ± 32.8	0.022	
UW-QOL: Anxiety	Li, 2016	70.57 ± 15.11	72.55 ± 15.19	0.219	
	Zhang X, 2014	78.5 ± 9.64	86.4 ± 17.5	0.775	
	Hsing, 2011	75.9 ± 26.3	68.9 ± 33.9	0.423	
UW-QOL: Composite score	Hsing, 2011	66.0 ± 18.5	57.8 ± 18.2	0.090	
Speech	Excellent	Zhang S, 2015	12 (80.0%) [#]	11 (91.7%) [#]	0.62
	Good		3 (20%)	1 (8.3%)	
	Poor		0	0	
	Always understandable	O'Neil, 201	17 (53.1)	4 (22.2)	0.014
	Usually understandable		14 (43.8)	9 (50.0)	
	Difficult to understand		1 (3.1)	5 (27.8)	

Table 16 Quality of Life data (Continued)

	Article	FF	PF	<i>p</i> -value
Swallowing full/regular diet at follow-up (vs soft, liquid) n (%)	Zhang S, 2015 ^S	13 (86.7%) [#]	10 (83.3%) [#]	1.00
	Paydarfar, 2011 [%]	19	20	0.60
	Chan Y, 2011 [*]	8 (38.2%)	24 (35.8%)	ND
		52 (61.9%)		ND
	O'Neil, 2010 ^{**}	17 (59.4%)	6 (33.3%)	0.202
Tsue, 1997 ^{***}	8 (34%)	4 (17%)	0.02	
Preoperative mouth-open width distance (mean) cm	Fang, 2013	1.5–6.2 (4.6)	1.2–6.2 (4.8)	ND
		0.9–6.0 (3.5)		ND
	Chien, 2005	6.3–3.5 (5.7)	6.1–2.5 (5.1)	0.384
Postoperative mouth-open width	Fang, 2013	1.4–5.8 (4.3)	1.1–4.7 (3.2)	ND
		0.8–5.8 (3.3)		ND
	Chien, 2005	5.9–3.2 (5.2)	5.6–1.6 (3.6)	0.384
Mouth-open width change (%)	Fang, 2013	4.0–9.1%	8.3–47.5%	< 0.001
		3.3–11.1%		< 0.001
	Chien, 2005	4.8–9.8%	5–45.5%	< 0.001
G-tube dependence, n (%)	Smeele, 2006	21.8%	34.3%	NS
	Chepeha, 2004	10 (16%)	40 (42%)	0.001
	Mallet, 2009	8 (36%)	17 (42%)	0.84
Feeding tube for > 21 days	Tsue, 1997	20 (69%)	20 (83%)	NS
Feeding tube at follow up ^e		11 (39%)	17 (85%)	0.002

§ Follow-up = 6 months

% at most recent follow-up

* Regular PO follow-up, median follow-up period was 82 months

** Follow-up period, time not mention

*** Median follow-up was 298 days

Percentage calculated relying on the data presented. Percentage not provided by the article

Bold = Statistically significant, *p*-value ≤ 0.05

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Availability of data and materials

This review is using data from published articles to portrayed the current literature on head and neck reconstruction surgery with either a free or a pedicled flap. All the references are properly listed as they are being used in the manuscript.

Authors' contributions

TA designed and directed the study. FGF and PT wrote the manuscript with support from TA. AR, EB, AC and TA were involved in the critical revision of the manuscript. All authors read and approved the final manuscript.

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