BEST-EVIDENCE CONSENSUS



The significance of surgically modifying soft tissue phenotype around fixed dental prostheses: An American Academy of Periodontology best evidence review

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Abstract

Background: This systematic review endeavored to investigate the effect of soft tissue phenotype modification therapy (PhMT-s) at sites with a tooth or an implant supported fixed dental prosthesis.

Methods: A comprehensive literature search was conducted by two independent examiners to identify relevant studies reporting differences in clinical, esthetic, or radiographic outcomes of interest between sites underwent PhMT-s and sites that remained untreated. Risk of bias assessment was calculated for all included studies. Meta-analyses involving endpoints of interest were performed when feasible.

Results: No controlled studies pertaining to tooth sites were identified. A total of six articles reporting on the outcomes of buccal soft tissue phenotype modification around implants were selected, of which, five were included in the meta-analyses. Quantitative analyses showed a weighted mean difference (WMD) of 0.98 mm (95% CI = 0.25 to 1.72 mm, P = 0.009) for change of tissue thickness; a WMD of -4.87% (95% CI = -34.27 to 24.53%, P = 0.75) for bleeding on probing (BOP); a WMD of 0.36 mm (95% CI = 0.12 to 0.59 mm, P = 0.003) for mucosal recession (MR); a WMD of 0.13 mm (95% CI = -0.11 to 0.36 mm, P = 0.30 for probing depth (PD); a WMD of 1.08 (95% CI = -0.39 to 2.55, P = 0.15) for pink esthetic score (PES), and a WMD of 0.40 mm (95% CI = -0.34 to 1.14 mm, P = 0.28) for marginal bone loss (MBL).

Conclusions: Surgical modification of peri-implant soft tissue phenotype via PhMT-s may decrease the amount of MR. Future clinical trials are needed to warrant the clinical benefits of modifying soft tissue phenotype around tooth-supported restorations.

KEYWORDS

dental implants, dental implantation, esthetics, gingival recession, peri-implantitis, systematic review

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1 | INTRODUCTION

Phenotype can be defined as the "appearance of an organ based on a multifactorial combination of genetic traits and environmental factors." The term "periodontal phenotype," which encompasses both the gingival phenotype (threedimensional gingival volume) and the thickness of the buccal bone plate (dentoalveolar bone morphotype), was recently adopted by the specialty of Periodontics¹ to replace the largely misused term "biotype." Historically, two main gingival phenotypes, thick-flat and thin-scalloped, have been widely employed to describe soft tissue appearance around teeth. Sites presenting a "thick-flat" phenotype are typically associated with squared tooth crown forms and wider contact areas between the teeth.² Additionally, the contact point is more apically positioned, often resulting in shorter interdental papillae. On the contrary, sites exhibiting a "thin-scalloped" phenotype normally present with tapered crown forms and shorter contact areas between the teeth. Because the contact point is usually located more coronally, the interdental papilla is often more volume.²

Unlike the dentate situation, the phenotypical categorization should be used with caution regarding implant sites because of wide variations resulting from site development procedures, implant placement, relative ridge positioning, and restorative design.³ Although the relationship of the papilla to the restoration is changed,⁴ much of the marginal inflammation and bone loss around peri-implant tissues may be related to the tissue phenotype.

The results of 2014 American Academy of Periodontology Regeneration Workshop provide us with a variety of strategies for phenotype modification of thin to thick phenotype.⁵ Decades of clinical experience indicate that this is "best practice" strategy for preventing gingival recession and future loss of attachment.⁵ Several methods have been proposed to categorize soft tissue phenotype around teeth and dental implants. Among all of them, visual assessment is arguably the most popular method because of its simplicity and noninvasiveness.6 This method defines a thin periodontal phenotype if the outline of the probe can be visualized through the marginal soft tissue and a thick phenotype if the outline of the probe cannot be seen. This classification for determining thin versus thick phenotype has been widely used⁶⁻⁸ and is reported to be a reliable alternative to other measurements. Because of its subjective nature, it is difficult to have an objective standard for comparison among studies. Alternatively, other proposed methods include direct clinical, 9 radiographic 10 or ultrasonic measurements 11 that provide objective measures for research comparisons. With the probe transparency method, a recent study⁷ has shown that the tissue thickness was consistently qualified as thin if the thickness was 0.6 mm or less, and thick if this value was >1.2 mm. For thickness between 0.7 and 1.2 mm, the frequency distributions showed a descending trend in thin phenotype and an ascending trend in thick phenotype.⁷

A thin periodontal phenotype may predispose the initiation or progression of recession defects. 12,13 Olsson and Lindhe analyzed the characteristics of maxillary central incisors in a cohort of 113 subjects and showed that long-narrow teeth presented more buccal marginal tissue recession than those with a short-wide tooth form. 14 In addition, a native thick tissue phenotype has been associated with more favorable clinical outcomes following corrective periodontal procedures. such as root coverage¹⁵ and periodontal regeneration. ¹⁶ Similarly, evidence supports that thin buccal peri-implant soft tissues are associated with an increased risk of future mucosal recession. 17,18 However, the decision of surgically modifying a thin to a thick phenotype using soft tissue grafting procedures (soft tissue phenotype modification therapy, PhMT-s) with the ultimate goal of achieving satisfactory long-term outcomes remains a controversial topic. 19 The aim of this systematic review was to investigate the effect of modifying a thin to a thick buccal soft tissue phenotype via PhMT-s around toothand implant-supported fixed prostheses in the function of relevant clinical, esthetic, and radiographic endpoints.

2 | MATERIALS AND METHODS

This systematic review adhered to the PRISMA (Preferred Reporting Items for Systematic Review and Meta-Analyses) statement guidelines.²⁰

2.1 | Focused question

What is the effect of surgically modifying a thin to a thick buccal soft tissue phenotype via PhMT-s around tooth- and implant-supported fixed prostheses in the function of relevant endpoints, for example, change in clinical, radiographic, and esthetic parameters?

Population: Adult individuals presenting intraoral sites with fixed tooth- or implant-supported prostheses

Intervention: Surgical augmentation procedures (PhMT-s) to modify the buccal soft tissue phenotype after restoration

Comparison: No surgical augmentation procedures to modify the buccal soft tissue phenotype

Outcomes: Changes in clinical, radiographic, or esthetic parameters with at least a 6-month follow-up

2.2 | Article eligibility criteria

The included articles had to fulfill all the following criteria:

- 1. Randomized controlled trials (RCTs), non-RCTs, cohort studies, case-control studies, or case series
- 2. A minimum of 10 treatment sites per group

- 3. Report at least one of the aforementioned outcomes of interest
- 4. Published in English

2.3 | Information sources and literature search strategy

An electronic search of Ovid MEDLINE, EMBASE, Web of Science and Cochrane Central was conducted on October 23, 2018, to identify relevant studies.

The search terms used for Ovid MEDLINE, where mh represented the Medical Subject Headings (MeSH), were: ("Gingival recession"[mh] OR "gingival recession"[all] OR "peri-implantitis"[mh] OR "peri-implantitis"[all] OR "dental implants"[mh] OR "dental implants"[all] OR "esthetics, dental"[mh] OR "esthetics"[all] OR "papilla"[all] OR "complications"[all] OR "complications"[all]) AND ("dental"[all] or "dentistry"[all]) AND ("phenotype"[all]) OR "biotype"[all])

The search terms used for EMBASE, where exp represented the explosion in the search strategy, were:

("gingiva disease"/exp OR "gingiva disease" OR "periimplantitis"/exp OR periimplantitis OR "tooth implantation"/exp OR "tooth implantation" OR "dental procedure"/exp OR "dental procedure" OR "esthetics"/exp OR esthetics OR "papilla"/exp OR papilla OR "complication"/exp OR complication OR "complications"/exp OR complications) AND ("dental"/exp OR dental OR "dentistry"/exp OR dentistry) AND ("biotype"/exp OR biotype OR "phenotype"/exp OR phenotype)

The search terms used for Web of Science were: (perimplantitis or perimplantitis or "dental implants" or "gingival recession" or papilla or complication or complications or esthetics) AND (dental or dentistry) AND (biotype or phenotype)

The search terms used for Cochrane Central were a combination of different keywords, including peri-implantitis, biotype, phenotype, dental implants, etc.

A hand search was also carried out in dental and implantrelated journals from January 2018 to October 2018, including Journal of Periodontology, Journal of Clinical Periodontology, Clinical Implant Dentistry and Related Research,
International Journal of Oral and Maxillofacial Implants,
Clinical Oral Implants Research, Journal of Dental Research,
Journal of Prosthetic Dentistry, International Journal of
Prosthodontics, Journal of Oral Implantology, and International Journal of Periodontics and Restorative Dentistry.
Additionally, a hand search of the references in the included
papers and review articles was conducted for relevant publications. For the search of grey literatures, Google Scholar
was used to identify any articles not included in the aforementioned databases.

2.4 | Literature selection

The initial screening of titles and abstracts was performed independently by two reviewers (GL and DC). Potential articles were examined in full-text after the initial screening and their eligibility for this review was confirmed after discussion. Agreement between the reviewers regarding study inclusion was calculated using kappa statistics.

2.5 | Data extraction

Data pertaining the pre-established outcomes of interest were extracted from the included studies by two independent reviewers (GL and DC) for subsequent qualitative and quantitative analyses. Data collected from each study included authors' names, year of publication, study design, sample size, demographic information of the participants (age, gender and smoking status), tooth/implant location, type of surgical approach, and follow-up period. Outcomes that were considered for the analyses included soft tissue dimensional changes, bleeding on probing (BOP), plaque index (PI), papillary fill index,²¹ keratinized tissue width (KTW), midbuccal recession (MR), probing depth (PD), pink esthetic score (PES),²² and marginal bone level (MBL). Corresponding authors of reviewed citations were contacted if further clarification regarding study methods and/or a more detailed data were needed.

2.6 | Risk of bias assessment

The Randomized Clinical Trial Checklist of the Cochrane Center²³ criteria were applied to evaluate the following methodological aspects of included RCTs: random sequence generation, allocation concealment method, blinding of participants and personnel, blinding of outcome assessment, incomplete outcome data addressed, selective reporting and other bias (Table 1). The degree of bias was categorized as: low, high, or uncertain risk.²³ Meanwhile, the included non-RCTs were assessed using the Newcastle–Ottawa Scale (Table 2).²⁴ Each non-RCT study was evaluated and rated from a maximum of nine stars to a minimum of no stars. Two reviewers (GL and DC) assessed all the included articles independently.

2.7 | Data synthesis

The primary outcome was the difference in the recorded parameters when comparing the sites with and without soft tissue grafting procedures to modify the tissue phenotype. For each parameter, the pooled weighted mean difference (WMD) between the grafted and non-grafted sites was

TABLE 1 Risk assessment of publication bias for the included RCTs

Study	Random sequence generation	Allocation concealment	Blinding of participants and personnel	Blinding of outcome assessment	Incomplete outcome data addressed	Selective reporting	Other bias
Migliorati et al. (2015) ⁵²	Low	Uncertain	Low	Low	Low	Low	Low
Wiesner et al. (2010) ⁵³	Low	Uncertain	Low	High	Low	Low	Low
Yoshino et al. (2014) ⁵⁴	Low	Low	Low	Uncertain	Low	Low	Low
Zuiderveld et al. (2018) ⁵⁵	Low	Low	Low	Low	Low	Low	Low

TABLE 2 Newcastle-Ottawa Quality Assessment Scale of included non-RCTs

Study	Selection	Comparability	Outcome
Bienz et al. (2017) ⁵⁰	***	*	**
Fenner et al. (2016) ⁵¹	**	*	***

estimated with computer software.* The contribution of each article was weighed based on sample size. Forest plots were produced to graphically represent outcome differences between the grafted and non-grafted groups using the number of sites as the unit of analysis. In addition, funnel plots (see supplementary Figures S1A to F in online *Journal of Periodontology*) were generated to assess the presence of publication bias. A P value = 0.05 was used as the level of significance. Heterogeneity was assessed with a chi-square test and I^2 test, which ranges between 0% and 100% with lower values indicating less heterogeneity. Random-effects meta-analyses of the selected studies were applied if the I^2 test showed a value of more than 50%; fixed-effects meta-analyses were applied if the I^2 test presented a value less than 50%.

3 | RESULTS

The screening process is shown in Figure 1. Electronic and hand searches yielded 1831 entries. After screening titles and abstracts, 32 articles were selected for full-text evaluation. Twenty-six articles were further excluded from the qualitative and quantitative analyses^{9,25–49}; the reasons for exclusion are listed in Table 3. After full-text review, no literature regarding tooth-supported prostheses was identified. Therefore, this specific aim could not be assessed because of lack of evidence. For implant-supported prostheses, six articles^{50–55} were included for qualitative/quantitative analyses. The kappa value for inter-reviewer agreement was 0.91 for identified titles and abstracts and 0.92 for full-text articles, indicating an "almost perfect" agreement between the two reviewers.⁵⁶

The main features and conclusions of the included studies were summarized in Table 4. The features and outcomes of the studies that included a secondary outcome analysis of tissue phenotype are displayed in supplementary Table S1 (around implants)^{3,37–40,43,57–75} and supplementary Table S2 (around teeth)^{48,76} in online *Journal of Periodontology* where the influence of phenotype on clinical outcomes is identified.

3.1 | Features of the included studies (implant-supported restorations)

3.1.1 | Study design and participant features

Four RCTs, ^{52–55} one cohort study, ⁵¹ and one case-control study ⁵⁰ were included. The age of the participants ranged from 19 ⁵⁵ to 87 ⁵⁴ years of age. Three studies ^{53–55} excluded smokers from participating their studies. All included studies had one study arm using subepithelial connective tissue graft (SCTG) for PhMT-s to thicken the buccal soft tissue phenotype and another study arm without using SCTG to serve as a control.

3.1.2 | Assessment method of tissue volumetric change

In terms of the methods to measure the phenotype change, one article⁵⁰ used stereolithographic files to assess the volumetric change digitally. Two studies^{54,55} determined the phenotype based on the transparency of a periodontal probe. Another two studies^{52,53} used endodontic reamers to assess the volumetric change. One article⁵¹ did not specify the method of assessing phenotype.

3.1.3 | Anatomic location of study sites

In four of the six included studies, ^{50,52,54,55} all fixtures were placed in either the anterior or premolar regions of the maxillary arch. One study reported placement of the implant fixtures only in the premolar or molar sites of the maxillary or mandibular arch. ⁵³ One study did not specify the implant location. ⁵¹

^{*} Review Manager (RevMan) (Version 5.0). The Nordic Cochrane Centre, The Cochrane Collaboration, 2008, Copenhagen, Denmark.

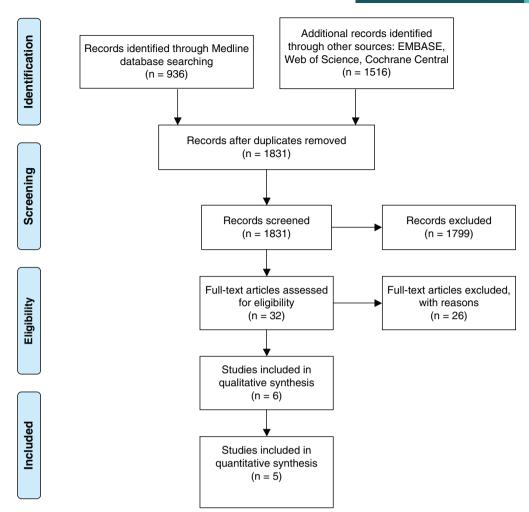


FIGURE 1 Flow chart illustrating the publication selection process

3.1.4 □ Bone grafts and membranes

In addition to SCTG procedure, xenogeneic bone grafting materials and collagen membranes were used in one study⁵¹ in which buccal augmentation via guided bone regeneration (GBR) was applied. The other studies did not perform GBR procedures in the study sites.

3.1.5 | Immediate implant placement and provisionalization

In addition to SCTG procedure, immediate implant placement and provisionalization (IIPP) protocol was used in two studies.^{54,55} One study⁵⁴ used xenogeneic graft material to fill the gap between the implant and buccal plate, whereas another study⁵⁵ used a combination of autogenous and xenogeneic graft to fill the gap.

3.2 | Results of meta-analyses (implant-supported restorations)

The meta-analysis conducted in the current study only included cohort studies and RCTs with data comparing

the clinical parameters between groups with and without SCTG. One case-control study⁵⁰ was excluded from the metaanalyses because the study performed soft tissue grafting procedures only in sites with a volume deficit on the buccal aspect of the implants, and therefore posed a risk of bias in baseline conditions between the grafted and non-grafted groups. The information of this case-control study is still shown in Table 4 for further reference.

Two RCTs^{52,53} evaluated change in tissue thickness. The results presented a WMD of 0.98 mm (95% CI = 0.25 to 1.72 mm, P = 0.009, Figure 2A), favoring the SCTG group. The comparison presented a high heterogeneity between the pooled studies ($I^2 = 80\%$).

Two articles^{51,52} evaluated BOP reduction. The results indicated a WMD of -4.87% (95% CI = -34.27 to 24.53%, P = 0.75). No statistical significance was found (Figure 2B) between groups. The comparison presented a high heterogeneity between the pooled studies ($I^2 = 77\%$).

Three articles^{51,54,55} evaluated MR. The results indicated a WMD of 0.36 mm (95% CI = 0.12 to 0.59 mm, P = 0.003). A statistically significant difference was detected (Figure 2C),

TABLE 3 Summary of the excluded articles

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Reason for exclusion	Author (year)
No data on comparing groups with and without soft tissue grafting procedures	Aguirre-Zorzano et al. 2013 ²⁵ Ahmed et al. 2018 ²⁶ Akcali et al. 2017 ²⁷ An et al. 2009 ²⁸ Bhat et al. 2015 ³¹ Cosyn et al. 2013 ³³ Kan et al. 2003 ³⁸ Kan et al. 2013 ³⁷ Kim et al. 2016 ³⁹ Nisapakultorn et al. 2010 ⁴⁰ Paniz et al. 2016 ⁴¹ Patil et al. 2013 ⁴² Ross et al. 2014 ⁴³ Spinato et al. 2012 ⁴⁶ Studer et al. 2010 ⁴⁸ Yilmaz and Tözüm 2012 ⁴⁹
No control group	Anderson et al. 2014 ²⁹ Batista et al. 2001 ³⁰ De Bruyckere et al. 2015 ³⁵ Hutton et al. 2018 ⁹ Schneider et al. 2011 ⁴⁴ Speroni et al. 2010 ⁴⁵
Inadequate data to be analyzed	Bianchi and Sanfilippo 2004 ³² Jyothi et al. 2013 ³⁶

favoring the SCTG group. There was a low ($I^2 = 31\%$) heterogeneity among compared studies.

Regarding PD reduction, three articles^{51,52,55} were analyzed. The results indicated a WMD of 0.13 mm (95% CI = -0.11 to 0.36 mm, P = 0.30). No statistical significance was found (Figure 2D) between groups. The comparison presented a low heterogeneity among the pooled studies (I2 = 0%).

In terms of PES, three studies 52,53,55 were analyzed. The results indicated a WMD of 1.08 (95% CI = -0.39 to 2.55, P = 0.15). No statistical significance was found (Figure 2E) between groups. The comparison presented a high heterogeneity among the pooled studies ($I^2 = 90\%$).

Four RCTs^{52–55} were pooled to evaluate MBL. The results presented a WMD of 0.40 mm (95% CI = -0.34 to 1.14 mm, P = 0.28). No statistical significance was found (Figure 2F). The comparison presented a high heterogeneity ($I^2 = 77\%$) among the studies.

Because of the lack of sufficient data, a meta-analysis could not be completed on PI, KTW, and papillary index. One cohort study⁵¹ reported the outcome of PI and did not detect a statistically significant difference between the grafted and non-grafted groups (P = 0.118). Only one study⁵² reported the change of KTW after grafting and did not find a significant difference. One RCT⁵⁴ reported the outcome

of papillary index and did not find a statistically significant difference between the grafted and non-grafted groups (P = 0.47 for mesial papilla and P = 0.35 for distal papilla, respectively). The findings for PI and papillary fill index were also reported in a case-control study,⁵⁰ where no difference in these parameters was identified between groups with and without SCTG.

3.3 | Risk of bias assessment

The risk of bias evaluation for RCTs were summarized in Table 1. Of the four included RCTs, one study⁵⁵ was ranked low for risk of bias in every category. Two studies^{52,54} were considered to have a category with an uncertain risk of bias. One study⁵³ was identified with an uncertain risk of bias in one area and a high risk of bias in a second category.

The risk of bias assessment for non-RCTs were summarized in Table 2. The two studies^{50,51} were scored six stars out of nine stars according to the Newcastle–Ottawa Scale,²⁴ and therefore were determined to have a considerable risk of bias.

4 | DISCUSSION

The significance of KTW around teeth with restorations 77,78 or dental implants⁷⁹ has been evaluated whereas the importance of soft tissue phenotype has not been widely analyzed. Therefore, the current review aimed to identify the potential benefit of modifying thin phenotype to thick phenotype through PhMT-s. From this review process, only two articles^{48,76} (see supplementary Table S2 in online *Journal* of Periodontology) were identified that contained secondary data analyses related to tooth-borne restorations. One in vitro study⁷⁶ concluded that a thick gingival phenotype could prevent tissue color change caused by the materials. Another study⁴⁸ concluded that crowns with a thick gingival phenotype resulted in significantly less recession than those with a thin phenotype when using metal-ceramic crowns. Because of the scarcity of clinical trials, future studies are warranted to evaluate the clinical benefits of surgically augmenting a thin gingival phenotype to a thick phenotype around a tooth-borne restoration.

Several studies^{11,80,81} have reported a positive correlation between the gingival phenotype and the buccal plate thickness. Therefore, when encountering a site with a thin gingival phenotype, clinicians should be aware of a possible thin underlying buccal plate for future implant placement. Interestingly, one study³ reported that there was no significant correlation between the gingival phenotype before tooth extraction and the peri-implant tissue phenotype after implant placement. This lack of correlation may result from several factors, including tissue remodeling processes, implant type, implant

TABLE 4 Summary of the articles analyzing the soft tissue outcomes between sites with and without soft tissue grafting procedures

	Main conclusions	Implant sites with and without soft tissue grafting showed only minimal changes and stability of peri-implant parameters over 5 years.	No statistically significant difference in the clinical parameters was observed between the sites with or without soft tissue grafting procedures.	None of the clinical parameters showed statistically significant difference between groups except for PES. Sites that received SCTG presented with more favorable PES than sites without SCTG.	Based on the outcomes of this split-mouth RCT, sites that received SCTG presented more favorable PES outcome when compared to sites without SCTG.	Subjects who underwent IIPP+SCTG experienced less facial gingival level change than those who did not receive a SCTG.	Zuiderveld RCT 60 19.5-82.2 29 12 All max -0.05 NA NA
(m	၁	3.65 (1.38)	3.97 (1.07)	3.2 (0.5)	NA	NA A	2.68
РД (тт)	T	3.67 (0.79)	4.09	3.4 (0.5)	X Y	Z Z	2.55 (0.95)
	C	NA V	NA A	6.39	8.45 (1.46)	Z A	6.8 (1.5)
PES	T	NA	NA A	(1.57) (0.83)	11.32 (1.63)	Z Y	6.4
KTW (mm)	C	NA	N A	-0.7 (1.57)	NA	X A	NA
KTW	T	NA	NA	-0.4 (1.66)	NA	NA	NA S
ive = ion)	C	-0.35 (0.30)	-0.06 NA (0.47)	NA	₹ Z	-0.70 NA (0.48)	(1.1)
MR (mm) (negative value = recession)	T	-0.47	0.03 (0.66)	NA	NA	-0.25 (0.35)	0.1 (0.8)
ary	C	1.55 (0.70)	NA	NA	NA	2.15 (0.75)	NA
Papillary index	T	1.38 (1.04)	NA	NA	NA	2.10 (1.02)	NA .
3 OP (%)	D	i); PI 14 (17); BOP 38 (29)	D; PI 28 (28); NA BOP 46 (24)	BOP 40 (40)	Z Z	NA	N A
PI (%)/ BOP (%)	T	1 Pt 10 (15);) BOP 38 (26)	PI 15 (32); BOP 56 (32)) BOP 20	NA V	NA	NA Y
e ness	C	(0.41)	^K Z	0.20 (0.70)	-0.15	N A	Ž Ž
Tissue thickness (mm)	T	-0.50	NA	0.40 (0.97)	1.20 (0.63)	N A	NA .
MBL (mm) (negative value = bone loss)	C	NA A	NA A	-0.17	-1.06	-0.14	-0.05
MBL (m (negative value = loss)	T	NA	NA	-0.06	-1.14	(0.27)	-0.03
	Follow- up months Location	ant or premolars	e-	All max ant or premolars	All –1.14 premolars (0.29) or molars	All max ant or premolars	All max ant or premolars
	Follow- up months	Average 60.5 53-152	Average 86.4 63.6- 111.6	24	12	12	17
ıts	် င	8 10 Avera GBR+ GBR 60.5 SCTG 53-15	22	23	10	10 IIPP	IIPP
Implants	T S	8 GBR+ SCTG	14 SCTG	24 SCTG	10 SCTG	10 IIPP+ SCTG	29 IIPP+ SCTG
Individuals	Age (gender)	18 27-76.6 11f7m	8 27-82 13f15m	48 22-70 25f23m	10 25-60 7f 3m	20 27-87 13f7m	60 19.5-82.2 29 32f.28m IIP SC
А	Design N	Case- 13	Cohort 28 27-82 13f 15		RCT 10	RCT 24	RCT 6
	Authors (year)	Bienz et al. (2017) ⁵⁰	Fenner et al. (2016) ⁵¹	Migliorati RCT et al. (2015) ⁵²	Wiesner et al. (2010) ⁵³	Yoshino et al. (2014) ⁵⁴	Zuiderveld RCT et al. (2018) ⁵⁵

RCT, randomized controlled trial; N, number; T, test group (with soft tissue grafting procedures); C, control group (without soft tissue grafting procedures); f, females; max, maxillary; mand, mandibular; ant, anterior teeth; MBL, marginal bone loss; PI, plaque index; BOP, bleeding on probing; MR, mucosal recession; KTW, keratinized tissue width; PES, pink esthetic score; PD, probing depth; IIPP, immediate implant placement and provisionalization; GBR, guided bone regeneration; SCTG, subepithelial connective tissue graft; NA, not available. Data in parentheses represent standard deviation.

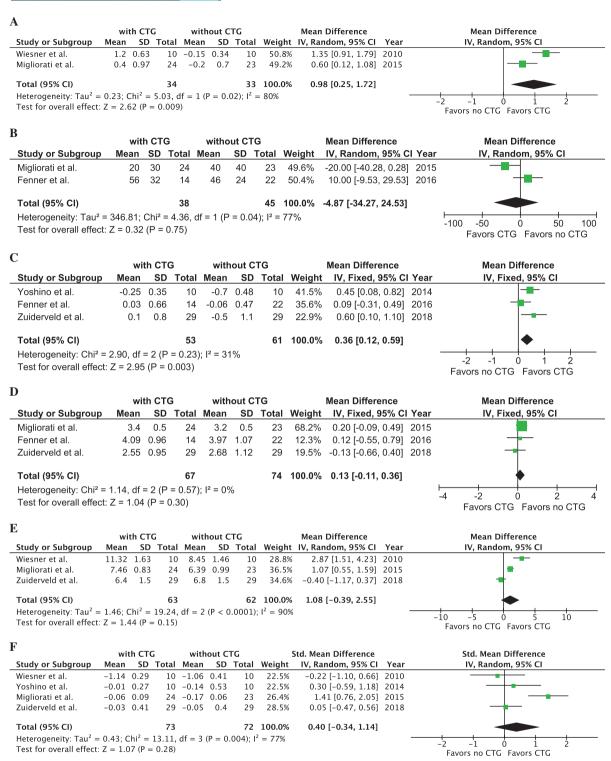


FIGURE 2 A) The result of meta-analysis for the change of peri-implant tissue thickness presented a WMD of 0.98 mm (95% CI = 0.25 to 1.72 mm, P = 0.009), favoring the SCTG group. The comparison presented a high heterogeneity ($I^2 = 80\%$). **B**) The result of meta-analysis for BOP reduction at implant sites presented a WMD of -4.87% (95% CI = -34.27 to 24.53%, P = 0.75). No statistical significance was found. The comparison presented a high heterogeneity ($I^2 = 77\%$). **C**) The result of meta-analysis for MR at implant sites presented a WMD of 0.36 mm (95% CI = 0.12 to 0.59 mm, P = 0.003), favoring the SCTG group. The comparison presented a low heterogeneity ($I^2 = 31\%$). **D**) The result of meta-analysis for PD reduction at implant sites presented a WMD of 0.13 mm (0.13 mm (0.13 mm (0.13 mm (0.13 mm). No statistical significance was found. The comparison presented a low heterogeneity (0.13 mm) (

orientation/position, and possible grafting procedures.^{3,82} Clinicians are advised to consider soft tissue grafting procedures when an undesirable implant outcome is foreseen.⁸³

PhMT-s has been widely used to successfully modify a thin tissue phenotype to a thick tissue phenotype around dental implants. 55,84 Our study confirmed the efficacy of PhMT-s and found that approximately a 1 mm gain of tissue thickness can be expected from this approach based on the metaanalysis. Therefore, a gain of 1 mm tissue thickness should be considered an endpoint for PhMT-s using SCTG aiming to thicken tissue phenotype. In a recent study,⁵² it was reported that sites with SCTG gained 34.3% tissue thickness after two years of follow-up, whereas sites without SCTG lost 9.9% tissue thickness. In addition, when performing IIPP procedure, the use of SCTG procedure has been shown to result in a more favorable peri-implant tissue thickness than the one without SCTG procedure. 85 Therefore, performing soft tissue grafting procedures to change tissue phenotype seems to be an enduring and predictable approach.

Increasing the soft tissue thickness provides the advantages of decreasing the soft tissue discoloration and show-through when a patient has a thin tissue phenotype and the implant or abutment is visible through the tissue. The thickened tissue also provides the restorative dentist more tissue volume by which to develop more idealized crown contours, which has both esthetic and biologic advantages. When the soft tissue phenotype is thin, ridge lapping is often necessary which limits access for cleaning and is not stable esthetically. By thickening the patient's soft tissue phenotype, it is easier to avoid the ridge-lap of crown restorations and develop a crown emergence profile that is more esthetic and biologically stable to facilitate patient's oral hygiene and tissue health.

In terms of peri-implant parameters, our results did not detect a difference in BOP between the sites with and without PhMT-s, which is in agreement with previous studies. 54,83 This indicates that BOP around implants depends on the health of the peri-implant tissue instead of the tissue phenotype. If the tissue presents healthy, BOP should not be a common finding on examination.⁹¹ However, soft tissue grafting procedures have been widely performed as one of the treatment modalities to manage peri-implantitis. 72 With the modification of prosthetic designs, soft tissue grafting procedures have also been introduced to manage mal-positioned implant fixtures. 92 In addition, evidence supports that PhMT-s can increase KTW and further improve patient comfort and compliance during oral hygiene. 93 Therefore, the need for these procedures should be based on the health status of the periimplant tissue and is determined on a case-by-case basis.

Results indicated a significantly less MR at sites with PhMT-s via SCTG than those without. Although the WMD was only 0.36 mm, two^{51,55} out of the three pooled studies reported a decrease in MR in the SCTG group. In contrast, the group without SCTG exhibited increased MR. This finding is

consistent with other studies, ^{54,61} and thus support the concept that modification of a thin to thick tissue phenotype by soft tissue augmentation could potentially reduce the amount of MR. With the use SCTG, creeping attachment may occur around natural teeth⁹⁴ or dental implants, ⁹⁵ which could further reduce the amount of MR. Therefore, clinicians should consider developing a thick tissue phenotype through grafting procedures whenever possible if the site presents with a high risk of future recession.

Our review shows that there is no statistically significant difference in change of PD when comparing sites with SCTG to the ones without SCTG. Previously published studies^{96,97} have shown that the healing after SCTG procedure is mediated by a combination of epithelial down growth and connective tissue attachment. Therefore, the difference in change of PD between the sites with and without SCTG was expected to be minimal, and the use of SCTG procedure will not result in deeper PD.

There was no difference in papillary fill reported between groups having or not having soft tissue grafting procedures that were performed to thicken the phenotype. 50,54 Although a recent study²⁶ reported that the phenotype may impact the heights and fill of interdental papilla by affecting papilla proportion and distances between the facial and palatal papilla, most studies^{4,98} showed that the papillary fill depends on the distance between the adjacent bone level and the contact point of the crowns. Currently there is insufficient evidence to support the rationale for modifying tissue phenotype to enhance papillary fill.

It remains controversial whether thickening the perimplant phenotype could result in an improved PES.⁵³ Although some evidence^{52,53} suggested a potential benefit of improved esthetics, the meta-analysis did not detect a significant improvement in PES with SCTG procedure. Among the three studies^{52,53,55} pooled in the meta-analysis, two studies^{52,53} reported a significant improvement in PES after thickening the tissue phenotype whereas a third study⁵⁵ reported no significant change in PES after surgically thickening the phenotype.

Based on the results of the meta-analysis, peri-implant sites, which are surgically modified to a thick soft tissue phenotype, do not exhibit a reduced amount of MBL compared to sites with a thin phenotype. This is consistent with several published reports. Whether a peri-implant site is with a thick or thin tissue phenotype, bone remodeling is an unavoidable process that occurs after tooth extraction therefore, other surgical modalities such as bone augmentation to thicken the peri-implant soft tissue phenotype may minimize but not prevent future bone loss.

Recent studies^{53,55} have also investigated the influence of PhMT-s to increase the amount of KTW using soft tissue grafting procedures. A systematic review by Thoma et al.⁸³

concluded that PhMT may result in more favorable periimplant tissue health such as a gain of KTW, an improvement of bleeding indices, and a higher marginal bone levels. Based on this review, higher bone level was noted in sites with apically positioned flap (APF) plus autogenous grafts versus all control treatments, including APF or vestibuloplasty procedure alone, APF with the use of collagen matrix, no treatment with or without residual keratinized tissue. Therefore, increasing soft tissue thickness and the amount of KTW via PhMT-s may be beneficial for providing more favorable periimplant tissue health. In addition, despite a lack of strong evidence, PhMT-s should be considered to achieve a wide band of KTW around tooth-borne restorations with a subgingival margin to facilitate gingival health. 77,78 Whenever a gain of KTW is needed, APF plus autogenous grafts is considered as the gold standard among all available treatment modalities.⁸³

All the studies pertaining to peri-implant mucosa thickening included in this systematic review involved a PhMT-s using an autologous SCTG after delivering the final implant-supported restoration. Interestingly, a recently published RCT¹⁰⁰ investigated the effect on MBL of periimplant soft tissue phenotype modification via CTG at the time of implant placement in a submerged approach (test), as compared to conventional implant placement (control). At implant uncovering, test sites presented less MBL compared to controls. However, this finding was only significant in sites with thin peri-implant soft tissue (≤ 2.5 mm) at baseline, but not in sites that presented thick tissue (>2.5 mm). This study also concluded that interim soft tissue modification before crown delivery did not significantly increase KTW. Therefore, if the peri-implant soft tissue thickness is ≤ 2.5 mm at baseline, it may be beneficial to perform PhMT-s to thicken the tissue simultaneously with implant placement with the purpose of minimizing MBL.

The limitations of this systematic review include (1) only five papers with comparable data were identified and pooled in the meta-analyses; (2) relatively short follow-up period of the included articles was noted; (3) considerable risk of bias was identified in non-RCTs; (4) four out of six reported meta-analyses had a high heterogeneity; (5) large variations in the study designs, implant placement protocols, outcome assessment methods, and reported parameters. Therefore, clinicians should interpret the results of this study cautiously after considering all the aforementioned limitations.

5 | CONCLUSIONS

On the basis of the evidence included in this systematic review, it was observed that surgical modification of perimplant soft tissue phenotype (PhMT-s) may decrease the amount of MR (WMD = 0.36 mm based on the meta-analysis) around implants. However, it remains inconclusive whether

thickening the peri-implant soft tissue positively influences PD, BOP and esthetic parameters, such as papillary fill and PES. In addition, clinical trials are needed to explore the effect of soft tissue phenotype modification around tooth-supported fixed dental prostheses.

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SUPPORTING INFORMATION

Additional supporting information may be found online in the Supporting Information section at the end of the article.

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