

Effect of Laser and Energy-based Device Therapies to Minimize Surgical Scar Formation: A Systematic Review and Network Metaanalysis

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Utilization of lasers and energy-based devices for surgical scar minimization has been substantially evaluated in placebo-controlled trials. The aim of this study was to compare reported measures of efficacy of lasers and energy-based devices in clinical trials in preventing surgical scar formation in a systematic review and network meta-analyses. Five electronic databases, PubMed, Scopus, Embase, ClinicalTrials.gov, and the Cochrane Library, were searched to retrieve relevant articles. The search was limited to randomized controlled trials that reported on clinical outcomes of surgical scars with treatment initiation no later than 6 months after surgery and a follow-up period of at least 3 months. A total of 18 randomized controlled trials involving 482 participants and 671 postsurgical wounds were included in the network meta-analyses. The results showed that the most efficacious treatments were achieved using low-level laser therapy) (weighted mean difference -3.78; 95% confidence interval (95% CI) -6.32, -1.24) and pulsed dye laser (weighted mean difference -2.46; 95% CI -4.53, -0.38). Nevertheless, low-level laser therapy and pulsed dye laser demonstrated comparable outcomes in surgical scar minimization (weighted mean difference -1.32, 95% CI -3.53, 0.89). The findings of this network meta-analyses suggest that low-level laser therapy and pulsed dye laser are both effective treatments for minimization of scar formation following primary closure of surgical wounds with comparable treatment outcomes.

Key words: surgical scar; laser; energy-based device; prevention; minimization; mitigation.

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SIGNIFICANCE

Various types of lasers and energy-based devices have been used to prevent scar formation following primary closure of surgical wounds. Although data are promising, they are insufficient to draw firm conclusions about the relative efficacies of these treatments. In this systematic review and network meta-analysis of 18 randomized clinical trials including 482 participants and 671 postsurgical scars, lowlevel laser and pulsed dye laser treatment were associated with significant reductions in Vancouver Scar Scale score than the untreated control. This network meta-analysis may help clinicians and patients to make informed treatment choices for the prophylaxis of surgical scar formation.

Wound healing is a complex tissue-response to injury that leads to skin restoration. Cutaneous aberrant scarring is characterized by an imbalance between cell growth and excessive deposition of extracellular matrix and is the consequence of skin injury. Alterations in the wound healing process can result in hypertrophic or keloid scarring causing functional impairment and psychological distress, especially when the scars are on a conspicuous part of the body (1). Therefore, effective interventions to minimize postoperative and/or posttraumatic scar formation are essential.

While lasers and energy-based devices (EBDs) have been generally accepted as a prophylaxis against scarring, there is no consensus on the appropriate timing for initiation of treatment for best outcomes. Laser treatments for scars have traditionally been performed at a minimum of 2–3 months after surgery because of scar stabilization and the disappearance of erythema at the operation site (2). However, increasing evidence emphasizes the importance of earlier initiation of treatment in scar minimization (3–5). The sparsity of head-to-head trials for scar minimization treatment modalities makes direct comparisons of their efficacy difficult. Few meta-analyses have been conducted, with 1 meta-analysis demonstrating that laser treatments

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result in scar prevention (4). A systematic review and a network meta-analysis (NMA) were conducted to assess and compare the efficacy of laser and EBD therapies in patients who had undergone intervention for scar prevention after primary closure of surgical wounds.

MATERIALS AND METHODS

Protocol and registration

A systematic review and NMA were performed and reported according to the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) extension statement for NMA (6). This study was registered with the trial registration number CRD42022381360 under the international prospective register of systematic reviews (PROSPERO: www.crd.york.ac.uk/ PROSPERO).

Eligibility and exclusion criteria for considering studies for this review

The analysis included randomized controlled trials (RCTs) that treated participants with lasers, EBDs or silicone gels and compared them with conventional postsurgical wound care to prevent postoperative scars. The primary outcome was differences in the mean Vancouver Scar Scale (VSS) score between the baseline and at the latest follow-up time-point. The secondary outcome was the mean difference of Observer Scar Assessment Scale (OSAS) between baseline and the latest follow-up visit. The study protocol only included interventions that were performed within 6 months after surgery and those with follow-up periods of more than, or equal to, 3 months after primary closure of the wounds in order to evaluate their effects on surgical scarring. RCTs that lacked sufficient information, such as standard deviation (SD), or studies that did not perform RCTs were not included in the current analysis.

Information sources and search strategy

Searches were performed for relevant published articles from 5 electronic databases (PubMed, Scopus, Embase, ClinicalTrials.gov, and the Cochrane Library) up until 24 January 2023, and medical subject headings (MeSH) were applied to each database, as applicable. The keywords included "pulsed dye laser", "fractional laser", "Nd:YAG laser", "picosecond laser", "erbium:YAG (Er:YAG) laser", "diode laser", "KTP laser", "fractional carbon dioxide (CO₂) laser", "fractional erbium glass (Er:glass) laser", "fractional erbium YAG (Er:YAG) laser", "intense pulsed light (IPL)", "low-level light", "radiofrequency device", "surgical wound", "surgical scar". Bibliographic lists of related articles were also explored. The complete search strategy is provided in Appendix S1; Table SI.

Study selection

Two investigators (SY and PA) independently evaluated the titles and abstracts, to identify potential eligibility from the searches, and relevant full-text articles were retrieved. The full-text articles were then assessed for final eligibility by the same individuals. Only English articles were included in the evaluation. Conflicts arising from the full-text articles were resolved through discussion or consultation with a team of experts.

Data extraction and study appraisal

The 2 investigators evaluated all potentially pertinent papers in a full-text search against the qualifying standards before selecting the paper in a data-extraction procedure. Data were extracted,

including the location of study, study design, intervention details (such as the regimen, treatment parameter, and duration of treatment), the study size (number of patients and number of scars); population characteristics (age, Fitzpatrick skin type, anatomical area of the scar) and treatment outcomes (i.e. the reported mean and/or standard deviation (SD) values of VSS score and OSAS score at the last follow-up), which were the indicative measures of the effects of the interventions. When mean and/or SD were not reported, continuous outcomes were estimated by using the reported statistics (e.g. median, interquartile range, etc.) (7). Study authors were contacted to obtain the missing outcomes of the relevant studies. If the authors did not respond within 1 month, the study was excluded from the analysis.

Quality assessments

Using the updated Cochrane risk-of-bias tool for randomized trials (RoB 2.0, London, UK), 2 researchers independently evaluated each study's risk of bias (8). This assessment addressed specific bias domains, including methods for generating the random sequence; allocation concealment; blinding of participants and investigators; blinding of the outcome assessment; incompleteness of the outcome data; and selective outcome reporting. The outcomes of each study's assessment of each item were displayed in the risk-of-bias summary graph and the risk-of-bias summary itself. Adjudication of the risk of bias was completed by answering pre-specified questions concerning the methodologies reported by each study in relation to the risk domain. The results were either a low risk of bias, an uncertain risk of bias, or a high risk of bias. Any differences between the 2 researchers were settled through consensus or by consulting an expert.

Synthesis and statistical analysis

Pairwise meta-analyses were performed using the DerSimonian and Laird random effects model to estimate outcomes (9). The outcomes were then reported as weighted mean differences and 95% confidence intervals (95% CIs). The I-squared statistic and the χ^2 test were used to evaluate the statistical heterogeneity in each pairwise comparison. When the *p*-value was less than 0.1, heterogeneity was considered to exist. Also, we used the network command in Stata Statistical Software: Version 16 (StataCorp LP, College Station, TX, USA) and the random-effects NMA techniques outlined by Lu and Ades to integrate direct and indirect evidence of all relative alternative effects (10). To rank the options, hierarchy of competing for intervention in the NMA, the rankogram; the surface under the cumulative ranking (SUCRA) curves; the mean ranks; and the league tables were used (11). A global inconsistency test was used to evaluate the network consistency between direct and indirect evidence (*p*-value ≥ 0.05 indicated consistency). A comparison-adjusted funnel plot was used to determine any publication bias and small-study effects.

Sensitivity analyses, concentrating on the aforementioned effects of laser treatments, were conducted to ascertain whether the results were impacted by the variability in the studies' characteristics. To examine the robustness of the results, several sensitivity analyses were carried out. These were based on: (*i*) the subgroup of the initial time-points after surgery to create highly effective prevention of scars; and (*ii*) the anatomical locations. To demonstrate the statistical significance, 2-sided statistical testing with a *p*-value of 0.05 was utilized.

RESULTS

A total of 1,967 articles were identified from PubMed, Scopus, Embase, ClinicalTrials.gov, and the Cochrane



Fig. 1. Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) flow diagram summary of the study selection process.

Library. Of these, 138 duplicated articles were removed. The full texts of 63 publications were assessed, and 1,904 studies were excluded for the reasons shown in Fig. 1. Ultimately, 18 eligible articles were obtained. Thirteen RCTs focused on the effects of the interventions to prevent surgical scars and reported VSS measurements (3, 5, 12–22). The other 7 RCTs were conducted on effects of the interventions to prevent surgical scars and depicted the OSAS measurements (3, 17, 23-27). Two RCTs reported results in both VSS and OSAS score (3, 17) (**Table I**). The study selection process flow is summarized in a PRISMA flow diagram (Fig. 1). The eligible trials were published between 2003 and 2023. A total of 482 randomized patients with 671 scars were included in the studies, of which 11 trials performed the split-scar technique. Nine trials were conducted in Asia, 3 trials recruited participants from Europe, 2 trials recruited participants from South America, and 4 trials were from North America. Nine studies obtained patients' age \geq 20 years. The mean last follow-up time reported for participants was 5.5 months. Thirteen studies excluded patients who had a history of hypertrophic or keloid scars.

Risk of bias

Appendix S1 provides an overall risk of bias of the RCTs included in the current study. Four studies reported a low

risk of bias. Eleven studies had some concerns for risk of bias. In addition, 3 studies showed a high risk of bias, with 2 of them having a high risk of bias for missing outcome data. Complete study characteristics and extracted outcomes data are found at Appendix S1; Figs. S1 and S2.

For network meta-analysis in Vancouver Scar Scale

Thirteen studies, which reported VSS as the primary outcome, were included to the NMA. Of these, the fractional CO₂ laser was the most frequently used comparator, studied in 5 out of 13 trials (5/13, 38.4%). There were a total of 9 treatment arms, including 7 lasers and EBDs. Low level laser therapy (LLLT), pulsed dye laser (PDL), fractional erbium: YAG (Er: YAG) laser, intense pulsed light (IPL), IPL+fractional erbium: YAG laser (IPL+ fractional Er: YAG laser), fractional ablative CO₂ laser (FACL), fractional erbium glass (Er:glass) laser, silicone gel, and untreated control were studied. Three papers compared FACL with untreated control and PDL to untreated control. LLLT to untreated control was investigated in 2 publications. Fractional Er:glass laser,

fractional Er:YAG laser, IPL+ fractional Er:YAG and IPL were compared with untreated control in 1 comparison. FACL was compared with silicone in 1 comparison. In addition, there were comparisons between EBD and laser as IPL+ fractional Er:YAG laser with fractional Er:YAG laser, laser to laser as PDL to FACL and FACL to silicone-based preparation (**Fig. 2**). The network metaanalysis result of OSAS is illustrated in Appendix S1; Table SV and Fig. S3.

Fig. 3 shows the results based on NMA combining direct and indirect comparisons. The treatment associated with the highest reduction in VSS (lower VSS scores imply better scar appearance) was LLLT and PDL-3.78 (95% CI-6.32, -1.24) and -2.46 (95% CI-4.53, -0.38), respectively. Following LLLT and PDL, based on the SURCA value, the results depicted silicone gel -1.75(95% CI -3.63, 0.14), fractional Er:YAG laser -0.83 (95% CI -2.43, 0.77), FACL -0.66 (95% CI -2.81, 1.49), IPL+ fractional Er:YAG laser -0.27 (95% CI -3.03, 2.49), fractional Er:glass -0.16 (95% CI -1.87, 1.56) and IPL 0.72 (95% CI -2.00, 3.44). Only LLLT and PDL were correlated with a significantly lower weighted mean difference of VSS score for surgical scar prevention compared with other interventions (Appendix S1; Table SIV).

However, there was no significant statistical difference between LLLT and PDL, with -1.32 (95% CI -3.53,

Table I. Char	acteristics of randomized										
Source	Study design	Location	Age, mean (SD), years	Fitzpatrick skin type	Anatomical site	Assessment	Intervention	Number of patients	Number of scars	Scar age at start	Duration of follow-up
Chi et al,	Observer-blinded, randomizeo	l China	NR	NR	Face	VSS	FACL	10	10	1 month	4 months
(14) 2023	controlled trial						FACL	10	10	3 months	6 months
							FACL	10	10	6 months	9 months
							Conventional conservative	12	12	NR	NR
Kang et al,	Observer-blinded, randomized	I USA	62.5 (13.2)	V-I	Head, neck	POSAS and OSAS	u edunenu (NS) 595nm PDL + 1550nm	28	28	2-8 weeks	5 months
(25) 2022	controlled trial			1			fractional Er:glass laser	1			
							Untreated control	24	24		
Kim et al, (18) 2022	Double-blinded, randomized	Korea	42 (7.08)	IV	Neck	VSS	Home-based 830nm LED	21	21	Not more than	6 months
Cheon et al		1	46.9 (10.74)		-			77	77		-
(24) 2022	Observer-blinded, randomized controlled trial	l Korea	48.0 (1.3)	NK	Neck	POSAS and OSAS	1550nm fractional Er:glass laser + ILS1 + Silicone gel (daytime) + Silicone sheet (nichttime)	32	32	3 weeks	6 months
			50.3 (13.3)				Silicone gel (daytime) + Silicone sheet (ninhttime)	32	32		
Kim et al, (17) 2021	Observer-blinded, split-scar, randomized controlled trial	Korea	62.13 (11.00)	VI-III	Abdomen	OSAS and VSS	555–950nm IPL + 2940nm fractional Er:YAG laser	17	17	1 week	5 months
							2940nm fractional Er:YAG laser		17		
Su et al, (22)	Ohserver-hlinded solit-scar	China	30.2 (6.74)	111-1/	Various sites	755		σ	ζđ	2 weeks	4 5 months
2021	randomized controlled trial	5		•)	Untreated control	'n	ით		
Shin et al,	Observer-blinded, split-scar,	Korea	43.4 (7.4)	NR	Breast	VSS and VAS	FACL	15	15	2-3 weeks	6 months
(20) 2021	randomized controlled trial						Untreated control		15		
Safra et al, (27) 2019	Observer-blinded, split-scar, randomized controlled trial	Israel	51.3 (11.3)	II-IV	Breast	POSAS and OSAS	595nm PDL + FACL+ Moisturizer + Silicone gel	18	18	2-6 weeks	9 months
							Moisturizer + Silicone gel		18		
Pongcharoen	Observer-blinded, split-scar,	Thailand	66.3 (8.92)	V-III	Knee	VSS	595nm PDL	39	40	2 weeks	6 months
er al, (12) 2013	randomized controlled trial						Untreated control		40		
Karmisnoit et al, (3) 2018_1	 Observer-blinded, split-scar, randomized controlled trial 	Denmark	65.5 (7.88)	111-1	Various sites	OSAS and VSS	1540nm fractional Er:glass laser	30	30	Immediately before excision	4.5 months
							Untreated control		30		
Karmisholt et al, (26) 2018_2	 Observer-blinded, randomizec controlled, intraindividual trial 	l Denmark	25.5 (1.56)	111-11	Buttock	OSAS and VAS	1540nm fractional Er:glass laser	16	144	1 day before excision	3 months
Alharti at al	-					000	Untreated control	0	16	-	-
(12) 2017	Double-blinded, randomized controlled trial	Brazil	38.2 (9.91)	I-IV	Breast, abdomen	VSS	FACL + Silicone gel Silicone gel	20 21	20 21	3 weeks	6 months
Buelens et al,	Observer-blinded, split-scar,	Belgium	45.46 (10.2)	I-IV	Head, neck	PhGA/PGA and	FACL	6	6	Within 3 month	is 6 months
Sobanko et al.	Observer-blinded solit-scar	VOI1	63 E (0 7E)	1_1/1	Earo		Untreated control	00	ь с	diref surgery	3 months
(21) 2015	randomized controlled trial	5			-		Untreated control	0	20		
Kim et al, (16) 2014	Observer-blinded, split-scar, randomized controlled trial	Korea	61 (17.04)	V-III	Face, flank	VSS	FACL 595nm PDI	14	14 14	2 weeks	3 months
Carvalho et al,	Observer-blinded. randomized	Brazil	47.07 (13.02)	NR	Groin	VSS and VAS and	LLLT (830nm diode laser)	28	14	1 dav	6 months
(13) 2010	controlled trial	5	47.14 (16.57)			scar thickness measurement	Untreated control	2	14	(m	
Conolouge et al, (15) 2006	Observer-blinded, split-scar, randomized controlled trial	NSA	59 (13.45)	I-IV	Various sites	VSS	595nm PDL Untreated control	13	13 13	Immediately after suture	3-5 months
Nouri et al.,	Observer-blinded, split-scar,	NSA	55 (11.17)	I-IV	Various sites	VSS	585nm PDL	11	12	removal Immediately	3-6 months
(5) 2003	randomized controlled trial			1			Intrested control	1	5	after suture	
							טוונו בפרבת כטוונו טו		77	removal	
SD: standard de Observer compo injection; IPL: in	viation; y: year; NR: not reports nent of the Patient and Observe tense pulse light; fractional Er:Y	ed; VSS: Vano r Scar Assess 'AG laser: fra	couver Scar Scale ment Scale; PDL: ctional erbium:YA	e; NS: not spec : pulsed dye las NG laser; VAS: *	ified; FACL: fract er; +: combined visual analogue :	tional ablative CO ₂ I d with; fractional Er scale: LLLT: low-leve	aser; POSAS: Patient component :glass laser: fractional erbium:gla el laser therapy: PhGA/PGA: Phys	of the Patient all Iss laser; LED: li dician/Patient Glo	nd Observer ght-emitting	Scar Assessment J diode; ILSI: intr Jent: CVAS: rosm	Scale; OSAS: alesional steroid atic visual

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Fig. 2. Networks of all options comparisons for reducton in Vancouver Scar Scale (VSS) score.

0.89). Six RCTs were used to conduct NMA in OSAS, (3, 17, 23, 24, 26, 27) and the result is shown in Appendix S1; Table SV, Fig. S3. The SURCA rank-bar chart, which illustrates SURCA cumulative probabilities of all outcomes associated with the prevention of surgical scars, is shown in **Fig. 4**.

Optimal period for treatment initiation

Meta-analysis was conducted between the laser/EBD group and non-laser group (consisting of silicone and untreated controls) in both VSS and OSAS (Appendix S1; Figs. S6–S16). Initial treatment times between the laser/EBD groups and the non-laser group were compared in meta-analysis. The forest plot depicted a greater overall decrease in VSS in the group treated within 1 week after surgery than in the group initiating treatment later than 1 week after surgery, with weighted mean difference of -1.51 (-2.43, -0.59) (I2 = 84%) and -0.76 (-1.45, -0.06) (I2 = 76%), respectively (Appendix S1; Fig. S10). Subgroup analysis in trials that performed the interven-

tions within 1 week showed that LLLT had the highest VSS reduction compared with others and untreated control. (Appendix S1; Fig. S17 and Table SVI). Other subgroup and sensitivity analyses were shown in Appendix S1; Figs. S18–S23, Table SVII–SXII.

Adverse effects

Regarding adverse effects, out of the 17 trials included in the study, only minor adverse effects were reported following the interventions. In the PDL treatment group, only minor purpuras were reported and all of them resolved spontaneously (5, 16, 19). No adverse event or significant pain based on the visual analogue score were reported in LLLT group (13, 18). In ablative laser interventions, crusting occurred in 2 patients and resolved in 1 week (17, 20).

Network consistency and small study effects

There was no evidence of any inconsistency in the results of the network meta-analysis in VSS. The comparisonadjusted funnel plots revealed no evidence of small study effects for the VSS score. However, there is an inconsistency in OSAS score as reported in Appendix S1; Table SXIII.

DISCUSSION

Cutaneous scarring is often a major concern for patients following primary closure of surgical wounds. The results of the current study expand on existing systematic reviews (28, 29) and meta-analysis (4), by comparing 8 different laser and EBD treatment interventions and untreated control in a single framework to investigate the benefits of treatment for reducing surgical scar formation. The results confirmed that laser and EBD interventions have the potential to improve the appearance of surgical scars, consistent with previous studies (4, 26). NMA suggests 830nm LLLT and 585nm and 595nm

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Fig. 3. The summarized results of using laser and energy-based device (EBD) treatment for the reduction in Vancouver Scar Scale (VSS) score. Silicone sheet/gel are also evaluated. WMD: weighted mean difference.



Fig. 4. Rank-bar chart with surface under the cumulative ranking (SUCRA) values for outcomes associated with lasers and energy-based devices used in patients with postsurgical scars.

PDL are the most effective interventions among all analysed trials in reducing VSS compared with untreated control and provide highest reduction of VSS of the scars at the last follow-up visits. Although the weighted mean difference between the 2 interventions was not significant, the efficacy of LLLT and PDL was determined to be equally effective. The advantages of LLLT and PDL are related to their non-ablative wounding characteristics and low degree of treatment discomfort, making patients comfortable to start the treatment on the early stage of the scar.

While LLLT and PDL exhibited similar efficacy in reducing scar formation following primary closure of surgical wounds, it is noteworthy that these 2 light sources operate via distinctive mechanisms. LLLT is postulated to modulate the inflammatory and proliferative phases of wound healing through a process known as photobiomodulation (18). In contrast, PDL relies on selective photothermolysis to target the scar's microvasculature, inducing localized hypoxaemia and consequently influencing collagen production (25). Our understanding suggests that the mechanisms of action employed by LLLT and PDL in the treatment of mature scars are fundamentally rooted in the principles applicable to early scar interventions. Commencing scar treatment during its early stages may potentially mitigate scar formation, particularly when the pathophysiology is immature and amenable to intervention, leading to more substantial

enhancements in scar appearance compared with interventions initiated at a later stage.

Although evidence from previous systematic reviews support early intervention as a key to control hyperplastic response leading to hypertrophic scars and keloids, the optimal time to initiate treatment remains controversial. A previous systematic review by Karmisholt et al. (29) reported a wide range of timing for treatment initiation, including the initiation in the inflammatory, proliferative and remodelling phases being effective for significant scar improvement. In contrast, a systematic review and meta-analysis by Kent at al.(4) indicates that laser treatment for minimizing primarily closed surgical scar should be initiated no later than 1 month after operation. However, the most recent systematic review by Behrouz-Pirnia and colleagues (28) could not conclude whether early laser treatment can reduce scar formation, due to the considerable heterogeneity (I^2 statistic=86%) of the currently available evidence. The current analysis further bolsters the trend that LLLT and PDL treatments initiated within 1 week after surgery are associated with a greater decrease in an overall effect on VSS.

Although the current systematic review and NMA focused primarily on closed full-thickness surgical scars for homogeneity, the assumptions can be extrapolated to the other types of scars, including acne, traumatic, and burn scars. The current study only included interventions initiated within 6 months after surgery because it aimed to assess the outcome of early initiation of treatment within the time period that the scar tissue has not yet reached the maturation phase (1). The current study only assessed studies with a follow-up period of at least 3 months postoperatively, in order to ensure that the interventions provided a certain duration of scar mitigation effect.

Silicone-based preparations in the form of sheets and gels are universally considered prophylactic options for hypertrophic scars and keloids. Based on the SURCA value, this NMA showed that silicone-based treatment improves scar appearance to -1.75 (-3.63, 0.14) compared with the untreated control. The mode of action of silicone-based products on scar tissue is unknown, but many agree that it acts at the stratum corneum by reducing evaporation and restores homeostasis, thereby reducing mast cell activity, oedema, vasodilation, and excessive extracellular matrix formation. The authors did not include silicone-based and other topical preparations used as a monotherapy for scar treatment in the current analysis because this systematic review and NMA focused on the use of lasers and EBDs for minimization of surgical scars. Although many RCTs have demonstrated ActaDV

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promising outcome of silicone-based preparations in prevention of hypertrophic scar development (30, 31), the effectiveness of silicone in preventing scar formation remains controversial. A systematic review by O'Brien & Jones (32) showed that silicone gel sheeting reduces the incidence of hypertrophic scarring in people prone to scarring compared with untreated control (risk ratio (RR) 0.46, 95% CI 0.21–0.98). However, the studies analysed in this review had a high susceptibility to bias. In contrast, a recent systemic review and meta-analysis evaluating 6 RCTs with a total of 375 patients demonstrated that topical silicone gel significantly reduced pigmentation, height, and pliability scores postoperatively compared with placebos or no treatment (33).

The majority of the RCTs analysed in this systematic review and NMA were monotherapy (using a single laser or EBD). However, in current practice, state-of-the-art treatment for scars is to apply multimodal treatments to maximize the therapeutic outcome. Although not investigated in large RCTs, evidence suggests that combining lasers targeting distinct chromophores (i.e. vascular lasers and non-ablative/ablative fractional lasers) (34), with intralesional and/or topically applied (laser-assisted drug delivery) triamcinolone acetonide (35, 36) and/or 5-fluorouracil (37), can yield superior results.

Study limitations

This NMA has several limitations. First, the heterogeneity of studies may limit the strength of the results. For instance, some studies adjusted for different sets of covariates, and not all studies provided fully adjusted effect estimates; hence the raw data for the meta-analysis was pooled. Secondly, treatment comparison arms were based on a single trial or trials with small sample sizes. Thus, the evidence certainty of the current analysis is limited by the inherent limitations of individual included trials. Thirdly, despite a comprehensive search approach, potential small-study effects exist and might contribute to some of the network effect estimates we observed. Fourthly, several of the studies included in this analysis assessed scar improvement without inclusion of untreated control groups. It is important to acknowledge the possibility that scars may improve over time even in the absence of treatment. Finally, we acknowledge the limited interconnections in the results of the network estimates.

Conclusion

Overall, the results of this systematic review and NMA suggest that low-level laser and pulsed dye laser are similarly effective treatments for minimizing surgical scar formation. There is a trend toward a better therapeutic outcome when treatment is initiated as early as 1 week postoperatively.

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