

Evaluating evidence for atrophic scarring treatment modalities

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Summary

Introduction: Atrophic scars cause significant patient morbidity. Whilst there is evidence to guide treatment, there does not appear to be a systematic review to analyse the efficacy of treatment options.

Objectives: To retrieve all evidence relating to atrophic scar treatment and evaluate using the Clinical Evidence GRADE score in order to allow clinicians to make evidence-based treatment choices.

Method: Searches were performed in Medline, EMBASE, CINHL and Cochrane to identify all English studies published evaluating treatment of atrophic scars on adults excluding journal letters. Each study was allocated a GRADE score based on type of study, quality, dose response, consistency of results and significance of results. The end score allowed categorisation of evidence into high, moderate, low or very low quality.

Results: A total of 41 studies were retrieved from searches including randomised controlled trials, observational studies, retrospective analyses and case reports of which 7% were allocated a high-quality score, 10% a moderate score, 7% a low score and 75% a very low score. Treatment modalities included ablative laser therapy, non-ablative laser therapy, autologous fat transfer, dermabrasion, chemical peels, injectables, subcision, tretinoin iontophoresis and combination therapy.

Conclusion: There is a paucity of good-quality clinical evidence evaluating treatment modalities for atrophic scarring. Evidence supports efficacy of laser, surgery and peel therapy. Further biomolecular research is required to identify targeted treatment options and more randomised controlled trials would make the evidence base for atrophic scar treatment more robust.

Keywords

atrophic scarring, evidence, GRADE score, treatment

Introduction

Scars are the end outcome of the natural healing and reparative process as a result of dermal fibrotic scar formation postinflammation. They have been categorised according to clinical and histological appearance into various categories: keloid, hypertrophic, stretched and atrophic scars. Atrophic scars are broadly described as exhibiting generalised cutaneous atrophy resulting in loss of cutaneous cells in the epidermis although appear clinically as a loss of normal dermis. Clinically, atrophic scars classically appear as depressions of the skin and commonly occur post acne amongst other causes listed in Table 1.

There are various atrophic scar treatment modalities in this clinical area of interest. This article aims to describe and evaluate the evidence in published literature including randomised controlled trials (RCTs) for atrophic scarring. The method of evaluation of studies will be via the Clinical Evidence GRADE score, which assigns a score to categorise all interventions according to their likely effectiveness based on type of study, quality, dose response, consistency of results and significance of results.

Methods

We conducted a Medline, EMBASE, Cochrane and CINHL search to retrieve any studies including RCTs evaluating treatment modalities for atrophic scarring. Key words used in each search engine included (with wildcard truncation used as*) atrophic, scar*, therapeutics (mapped to thesaurus to include treatment, intervention), drug treatment, conservative treatment, surgery, laser, treatment outcome and treatment response. We excluded journal letters and interventions on paediatric patients. We included all English publications published in any year analysing the effectiveness of treatment modalities for atrophic scarring. A total of 45 studies were identified which included RCTs, observational studies, retrospective analyses and case reports. No review articles were identified.

All abstracts of studies found were assessed by two independent clinicians against criteria for inclusion to ensure applicability of study within the systematic review. Randomised control trials were included regardless of the length of follow-up period or patient drop-out rate and those that were at least single blinded. A final number of 41 studies were identified.

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Table	Ι.	Main	causes	and	risk	factors	for o	devel	opin	g atrop	phic
scars.											

	Cause/risk factor
Inflammatory	Acne
	Cyst
	Discoid lupus erythematosus
Infective	Postvaricella
Trauma	Injury
	Burn
	latrogenic – surgery
Patient factors	Tendency toward atrophic scarring
	Previous atrophic scars
	Ehlers–Danlos syndrome
	Primary anetoderma

Having retrieved the full articles of all studies identified, each study was evaluated with the GRADE Clinical Evidence score as per parameters shown in Table 2.¹ The scoring system aimed to assess the type of study, quality, consistency, directness and effect size allowing the overall score to rate the study as high, moderate, low or very low in terms of intervention effectiveness for atrophic scarring.

Results

Forty-one studies reporting treatments for atrophic scars were identified including randomised control trials (8), retrospective cohort study analysis (8), prospective cohort analysis (21) and case studies (4) as demonstrated in Figure 4.

The available treatment modalities for atrophic scarring available in literature are ablative fractional laser therapy (6), non-ablative laser therapy (16), dermabrasion (2), chemical peel therapy (5), surgical techniques such as subcision, autologous fat transfer and injectables (5) and combination therapies (7) highlighted in Table 3.

Of these studies, three were allocated a high GRADE score (7%), four studies scored as moderate (10%), three scored as low (7%) and the majority of 31 studies scored very low (75%), demonstrated in Figure 5.

Of the eight RCTs evaluated, five were regarded as high- or moderate-quality studies. Studies were deducted points due to low sample size and short-term follow-up although all RCTs showed statistically significant improvement of scars with treatment. Efficacious treatment modalities were CO_2 ablative therapy and non-ablative laser therapy. Of the 21 observational studies, identified scores ranged from moderate- to very-low-quality studies. Again score variances identify issues of interstudy heterogeneity with regard to sample sizes, methodology, treatment outcome measures used, and methods of evaluating scar improvement. Lastly, most of the retrospective analysis and case report studies scored as low or very low quality due to the limitations of sample size and generalisability of results.

Discussion

Several key papers that discuss atrophic scarring focus on acne. Community-based studies report acne is prevalent in 90% of adolescent patients and can occur in any regions where there is an abundance of pilosebaceous glands such as the face, shoulders, back and chest. The process of acne is caused by various factors which increase sebum production of pilosebaceous glands such as increased systemic production of androgens and concomitant Propionibacterium acnes proliferation within follicles. The resultant infrainfundibular inflammatory process which ensues can cause follicular rupture or abscess formation. The wound healing process which then occurs can cause atrophic scars. In the adult population, 1% of patients are reported to have persistent acne scarring from adolescence.⁴³ The risk factors for developing scarring are multifactorial including a genetic predisposition to scarring, and a delay in acne treatment. The extent of scarring in acne can be reduced by early treatment during the inflammatory phase.

An observational study of normal scarring shows that histologically the scar maturation process occurs over a year with fibroblastic changes in dermal layers of skin; however, clinically the appearance remains unchanged except from diminishing erythema as angiogenesis ceases.⁴⁴ In addition, atrophic scarring is reported to worsen with age due to the natural lipoatrophy which further accentuates the scars.⁴⁵

Atrophic scars are defined histologically as scars showing a loss of collagen. They have been subclassified into the icepick scars which are the most prevalent subcategory at 60-70%, boxcar scars at 20-30% and rolling scars at 15-25%.⁴⁶ Icepick scars are described as a 'V shaped' extension scar into dermis whereas boxcar and rolling scars are more superficial with a wider base (Figure 1).

However, boxcar and rolling scars can be deep and all three subcategories can co-exist making clinical identification of type difficult. Histologically,

Parameter	Areas examined within parameter for each study	Score	Score explained
Type of evidence	RCTs/SR of RCTs	+4	
	Observational evidence (e.g. cohort, case–control)	+2	
Quality	Blinding and allocation process	0	No problems
	Follow-up	I	Problem with I element
	Withdrawal of participants	2	Problem with 2 elements
	Sparsity of data	3	Problem with 3 elements
Consistency	Degree of consistency of effect between or within studies	+1	Evidence of dose response across or within studies (or inconsistency across studies is explained by a dose response); also I point added if adjustment for confounders would have increased the effect size
		0	All/most studies show similar results
		I	Lack of agreement between studies (e.g. statistical heterogeneity between RCTs, conflicting results)
Directness	The generalisability of population and outcomes from each study to population of interest	0	Population and outcomes broadly generalisable
		I	Problem with I element
		2	Problem with 2 or more elements
Effect size	The reported OR/RR/HR for comparison	0	Not all effect sizes >2 or <0.5 and significant; or if OR/RR/HR not significant
		+1	Effect size >2 or <0.5 for all studies/meta- analyses included in comparison and significant
		+2	Effect size >5 or <0.2 for all studies/meta- analyses included in comparison and significant

Table 2. Clinical evidence GRADE score components (adapted from Clinical Evidence¹).

Final score (quality of evidence) High = 4 points overall, Moderate = 3 points, Low = 2 points, Very low = 1 point or less. RCT: randomised controlled trial.

atrophic scars exhibit thinning of the skin with a loss of collagen, elastin and deep dermal fat which cause a downward traction pull of the epidermis (Figure 2). Atrophic scar formation over time can be demonstrated in Figure 3.

The histological changes noted and described arise due to the inflammatory and reparative process of wound healing. The reparative phase is subdivided into (a) inflammation, (b) granulation and (c) matrix remodelling phases.

The main causes for patient distress from acne are facial disfigurement from inflammation, pigmentation and scarring. The psychosocial consequences resulting from acne were first recognised by Sulzberger and Zaidens.⁴⁹ A study examining the quality of life acne patients reported showed they experienced psychological and emotional morbidity comparable to chronic, disabling conditions such as epilepsy, diabetes and arthritic pain.⁵⁰ In addition, severe acne has been correlated with depression and suicidal ideation.⁵¹ Patients can experience body dysmorphia due to the visible acne; however, the degree of psychological distress may not always correlate with the severity of acne and can be present in

Authors	Treatment modality	Score	Study design	Outcome
CO ₂ ablative laser				
Hedelund et al. ²	CO ₂ laser re-surfa- cing vs. placebo	+5	RCT (single blinded): 12 patients treated with randomised split face treatment three times in 4/5-week intervals. Followed up till six months.	Objective, statistically assessed increase in scar smoothness.
Walia and Alster ³	CO ₂ laser re- surfacing	+3	Observational: 60 patients with acne atrophic scars treated with laser re- surfacing and assessed between 1 and 18 months with biopsies.	Subject investigator improve- ment scores and positive histological evidence of new collagen deposition.
Weiss <i>et al.</i> ⁴	CO ₂ ablative frac- tional therapy	-3	Observational: 19 non-acne atro- phic scars treated with three treat- ments of ablative fractional therapy at 1–4-month intervals and followed up for six months.	Subjective patient and inves- tigator improvement scores and non-significant objective topographical analysis improvement.
Cho et al. ⁵	CO ₂ ablative frac- tional therapy	-3	Observational: 20 acne atrophic scars treated with ablative fractional therapy and followed up for three months.	Subjective patient and inves- tigator improvement scores
Kim ⁶	CO ₂ laser	-3	Observational: 35 acne atrophic scar patients treated with five sessions at 2–3-week intervals of pinpoint CO_2 laser, nil long-term follow-up.	Subjective patient and inves- tigator improvement scores.
Manuskiatti et <i>al.⁷</i>	CO ₂ ablative therapy	-1	Observational: 13 acne atrophic scar patients treated with three seven-weekly laser sessions and fol- lowed up till six months.	Objective and subjective patient and investigator improvement scores
Non-ablative laser				
Tanzi and Alster ⁸	1450 nm diode laser vs. 1320 nm Nd:YAG laser	+5	RCT (single blinded): 20 atrophic scar patients received split face laser treatments at three-week intervals and followed up for 12 months.	Significant increased histo- logical deposition of collagen for both and improved objective photographic scar quality for both.
Min et al. ⁹	Laser: long-pulse vs. combined 585/ 1064 nm	+4	RCT (single blinded): 19 patients with acne atrophic scars received split face long laser and combined at two-week intervals and followed up for 14 weeks.	Objective investigator and patient improvement scores for both treatments but nil significant difference between treatments. Significantly increased histological colla- gen deposition for both but nil significant difference between treatments.
Wanitphakdeedec- ha et <i>al</i> . ¹⁰	Er YAG laser: short pulse vs. extra long pulse	+3	RCT (single blinded): 22 patients with atrophic acne scars rando- mised to treatment with SP or ELP	Objective investigator improvement scores.
				(continued)

Table 3. Atrophic scar to	reatment modality and	GRADE scoring.
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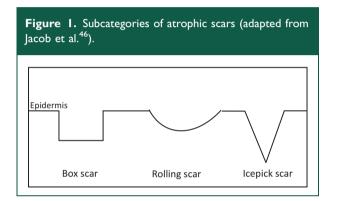
Authors	Treatment modality	Score	Study design	Outcome
			for two sessions monthly. Followed up till four months.	
Hedelund et al. ¹¹	Fractional non- ablative laser, 1540nm laser vs. control	+2	RCT (single blinded): 10 patients randomised to 1540 nm laser vs. no therapy. Laser therapy given four weekly three times. Followed up till 12 weeks after last laser session.	Objective investigator and patient improvement scores
Chan et al. ¹²	Non-ablative laser	+2	Retrospective analysis: 47 acne atrophic patients who received non- ablative laser treatment between December 2005 and February 2009.	Objective improvement in scar texture, pigmentation and extent
Chua et al. ¹³	Non-ablative I 450 nm diode laser	+1	Retrospective analysis: 57 acne atrophic patients who received laser treatment from May 2002 to December 2003.	Objective patient and investi- gator improvement scores.
Sadick and Schecter ¹⁴	Laser: Nd YAG I 320 nm	-1	Observational: eight patients with atrophic acne scars given \times 3 treatments of laser. Followed up for six months to one year.	Objective patient and investi- gator improvement scores.
Chan et al. ¹⁵	Laser re-surfacing	-1	Observational: 27 acne atrophic patients treated for six months with laser and followed up till 18 months.	Subjective increase in investi- gator and patient scores, objective histological increase in collagen and objective improvement in scar viscoelasticity.
Rogachefsky et al. ¹⁶	Laser re-surfacing	-1	Observational: 12 patients with acne atrophic scars treated with \times 3 laser treatment at one-monthly intervals and followed up till six months.	Objective patient and investi- gator improvement scores.
Badawi et <i>al</i> . ¹⁷	Non-ablative 1064 laser	-1	Retrospective analysis: 22 acne atrophic patients who received six months of laser treatment between February and July 08.	Objective improvement in scar improvement, texture and postinflammatory hyperpigmentation.
Jih et al. ¹⁸	1450 nm diode laser	-3	Case study analysis	Subjective investigator and patient improvement scores
Tanzi and Alster ¹⁹	2940 nm Er YAG laser	-3	Observational: 25 patients treated with laser and followed up till 12 months.	Subjective investigator improvement scores
Park et <i>al.</i> ²⁰	Fractional photothermolysis	-3	Observational: 59 patients with atrophic scars treated with photo- thermolysis at 3–4-week sessions for three weeks.	Subjective investigator and patient improvement scores and increased collagen on biopsy
Cho et al. ²¹	Fractional photothermolysis	-3	Observational: 12 acne atrophic scar patients treated with \times 3 1550 nm Erb laser at monthly	Subjective investigator and photographic improvement scores

Authors	Treatment modality	Score	Study design	Outcome
			intervals and followed up till four months.	
Deng et al. ²²	Fractional photo- thermolysis laser	-3	Observational: 26 acne atrophic scar patients treated with laser, follow-up time not specified.	Subjective patient and inves- tigator improvement scores
Koo et al. ²³	Laser punch out	-3	Observational: 71 patients with acne atrophic scars treated with laser punch out and followed up till 12 months.	Subjective patient and inves- tigator improvement scores
Autologous fat transfe	r			
Roh et al. ²⁴	Autologous fat transfer	-3	Retrospective study: 20 sclero- derma atrophic scar patients treated with three-weekly intervals of fat transfer identified from lower abdomen or buttocks. Followed up till 12 months.	Subjective investigator improvement scores
Lapiere et al. ²⁵	Autologous fat transfer	-3	Two case studies of atrophic scar patients treated with fat transfer.	Subjective investigator and patient improvement scores
Dermabrasion				
Bagatin et al. ²⁶	Dermabrasion	-3	Observational: seven patients on oral isotretinoin for acne atrophic scarring treated with manual derm- abrasion and followed up till 180 days.	Subjective photographic evi- dence improvement
Majid ²⁷	Microneedling	-3	Observational: 36 patients with facial atrophic scars of various aeti- ology treated with maximum of four months derma roller treatment and followed up till two months.	Subjective patient and inves- tigator improvement scores.
Chemical peels				
Erbagci and Akcah ²⁸	Glycolic acid peel: daily application vs. biweekly	+3	RCT (Single blinded): 48 patients randomised to (a) biweekly peels (b) daily peels (c) control and followed up till 24 weeks	Objective patient and investi- gator improvement scores with biweekly peels vs. daily peels.
Barikbin et al. ²⁹	TCA peel	I	Observational: 100 varicella atro- phic scar patients received 70% TCA and followed up in 12 weeks.	Subjective improvement in investigator and patient scores.
Lee et al. ³⁰	TCA via CROSS method	-2	Retrospective analysis of 58 acne atrophic patients treated with 65% or 100% TCA via CROSS method.	Subjective patients and inves- tigator improvement scores, better scores with higher concentration.
Fabbrocini et al. ³¹	CROSS technique: 50% TCA	-3	Observational: five patients with acne atrophic scars treated with three TCA treatments at four-	Subjective patient and inves- tigator improvement scores

Authors	Treatment modality	Score	Study design	Outcome
			weekly intervals. Followed up at end of last treatment.	and increased collagen deposition histologically
Khunger et al. ³²	CROSS technique: 100% TCA	-3	Observational: 30 acne atrophic scar patients treated with four two- weekly sessions of 100% TCA and followed up at three months.	Subjective patient and inves- tigator improvement scores
Injectables				
Sadove ³³	Injectable poly-L- lactic acid	-3	Two case study analyses of PLLA treated atrophic acne scars.	Subjective investigator improvement scores
Richards and Rashid ³⁴	Hyaluronic acid filler	-3	Case report of atrophic treated with hyaluronic acid filler.	Subjective patient improvement
Subcision				
Harandi et al. ³⁵	Subcision – suction combination	+2	Observational: 58 acne atrophic patients treated with subcision and then suction therapy. Followed up till six months.	Objective investigator and patient improvement scores
Other				
Leheta et al. ³⁶	Percutaneous colla- gen induction vs. 100% TCA	+3	Randomised (single blinded) study: 30 patients randomised to four sessions of TCA/PCI at four-weekly intervals.	Objective patient and investi- gator improvement scores.
Kim and Cho ³⁷	Ablative fractional laser vs. combined ablative and non- ablative laser	0	RCT (single blinded): 20 patients randomly received split face ablative fractional therapy on one half and ablative fractional therapy plus non- ablative laser on the other. Followed up at four weekly intervals.	Objective patient and investi- gator improvement scores, better results with combin- ation therapy.
Sage et al. ³⁸	Subcuticular inci- sion vs. porcine collagen	0	RCT (single blinded): nine patients underwent random split face treat- ment of incision on one half and collagen on other. Followed up till six months.	Objective patient improve- ment scores of incision vs. collagen, nil significant inves- tigator improvement between both.
Epstein and Spencer ³⁹	Subcision and arte- fill filler	-3	Observational: 14 patients treated with subcision and then artefill. Followed up till eight months.	Subjective patient and inves- tigator improvement scores.
Kang et al. ⁴⁰	Dot peeling and subcision and frac- tional laser	-3	Observational: 10 patients with atrophic acne scars each received 3–4 treatments of laser monthly. Then two weeks after, received dot and subcision treatment together. Total treatment for one year and followed up at three months after all combined.	Subjective investigator improvement scores

Authors	Treatment modality	Score	Study design	Outcome
Carniol et al. ⁴¹	1450 nm laser and 30% TCA com- bined therapy	-3	Observational: nine atrophic scar patients received four-monthly laser treatment followed by bimonthly 30% TCA peels.	Subjective increased investi- gator and patient improve- ment scores.
Schmidt et <i>al</i> . ⁴²	Tretinoin iontophoresis	-3	Observational: 32 patients treated with tretinoin iontophoresis and followed up.	Subjective investigator and patient improvement scores. Nil significant histological collagen deposition.

RCT: randomised controlled trial; TCA: trichloroacetic acid.



clinically mild acne causing low self-esteem and a reduced confidence to socialise.⁵² Due to these recognised psychological comorbidities acne patients may have, it is important to offer holistic management and recognise psychological symptoms early.

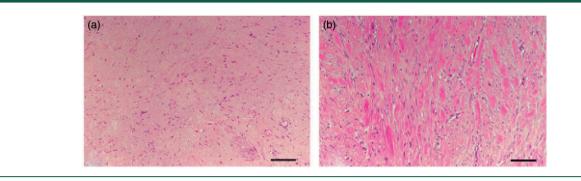
Given the significant patient morbidity, our study highlights a lack of robust, good-quality evidence evaluating treatments. Overall, there is a lack of evidence evaluating treatment of this particular scar subcategory in comparison to keloid or hypertrophic scarring. In addition, there appears to be a lack of concrete pathophysiological knowledge of how atrophic scars occur and develop which could explain the paucity of treatment evidence as further biomolecular and pathological research examining the nature of atrophic scars needs to be conducted to further identify targets for treatment.

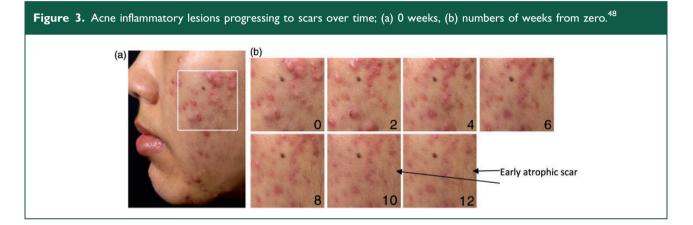
We chose the Clinical Evidence GRADE Score system to evaluate the quality of studies in our review over other scoring systems as we aimed to focus on outcome evidence of scar severity improvement, which clinicians could use to make judgements about efficacy of treatment for individual patients. Hence, the end scores assigned to each study do not take into account detailed analysis of methodology such as single or double blinding as well as wider practical issues such as treatment cost effectiveness. Two independent clinicians assigned each final score and crosschecked for congruence and in cases of discrepancy the paper was re-examined and the end score was achieved together. Despite this, the main disadvantage of the GRADE scoring system is that it is prone to being more subjective than other scores as it relies on the assessor's own judgement about certain criteria and where studies have very different methodologies comparison for scoring purposes can be difficult.

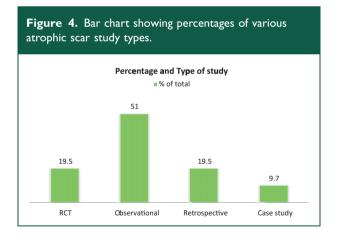
As is evident, there is a scarcity of RCTs and all are single blinded due to the nature of treatments subjected to trial; however, they provide a basis for further large-scale studies particularly with regard to the injectable therapies and the CROSS method of trichloroacetic acid (TCA) chemical peel therapy. Clearly, further RCTs with standardised methodology would need to be conducted before a metaanalysis of results can be compiled.

Table 4 is a visual summation of the results identified. In practice, it is very difficult to give clear guidelines of which treatment modality is best, as the choice of treatment will depend on individual patient characteristics such as skin tone, original scar location, previous treatment modalities experienced, overall treatment therapy duration, treatment downtime, side effects and their visibility, patient expectations of treatment and willingness to trial combination therapy. However, the evidence examined for this review does suggest a hierarchy of treatment in terms of efficacy.

The main limitation of our study is the lack of total number of articles retrieved from which to draw a sound conclusion. However, we did not search for articles in other languages or nonpublished data rendering the results subject to publication bias. In addition, the scoring system used evaluates efficacy of treatment but not patient tolerance of treatment, which may affect options offered. **Figure 2.** Immunostaining of anti- β -catenin in (a) atrophic scar and (b) keloid. Sparse staining in (a) correlates with reduced growth factor activity compared to (b). Haemotoxylin and eosin staining.⁴⁷ Histopathology of atrophic scar (a) and keloid (b). Scale bar: 100 μ m



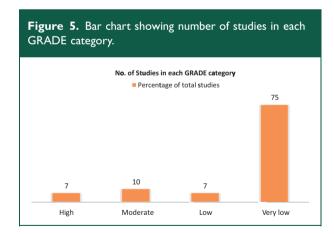




For example, most patients did not tolerate CO_2 ablative therapy well due to pain during and immediately after therapy as well as multiple side effects.

Conclusion

In summary, our review identifies an overall lack of published data regarding treatment of atrophic



scarring and poor study methodology. Further research at a molecular level may be able to better define atrophic scarring and allow targeting of therapy. Our review allows the possible identification of areas where further RCTs may be conducted. Whilst our review reveals various treatment options which may be utilised to treat atrophic scarring, the ultimate choice of a particular treatment modality will depend

Treatment	Type of study and highest score	Comment
CO ₂ ablative therapy	RCT (+5)	Significantly effective ($p < 0.0001$)
Laser 1450 nm diode laser/Nd YAG laser	RCTs (+5)	Both treatments significantly effective $(p = 0.008)$ but nil difference between two.
Long-pulse/combined 585/1064 nm	RCT (+4)	Both treatments significantly effective $(p < 0.001)$ but nil difference between two.
Glycolic acid/biweekly peels	RCT (+3)	Both treatments significantly effective $(p < 0.001)$ but nil difference between two.
Percutaneous collagen induction/trichloroacetic acid	RCT (+3)	Both treatments significantly effective $(p < 0.01)$ but nil difference between two.
Subcision	Observational (+2)	Significantly effective ($p < 0.041$)

Table 4. Hierarchy of therapy for atrophic scars based on GRADE scores.

 $Other \ treatments \ (scores \ less \ than \ +2): \ autologous \ fat \ transfer, \ dermabrasion, \ injectables, \ tretinoin \ iontophoresis, \ subcision \ and \ artefiller \ and \ triple \ therapy.$

RCT: randomised controlled trial.

Table 5. Pros and cons of each treatment modality.

Treatment	Pros	Cons
CO_2 ablative therapy	• Quick therapy	• Multiple therapies less well tolerated
		Side effects and longer downtime
Non-ablative laser	• Quick therapy	• May require increased treatment frequencies for end result
	• Minimal side effects	
	• Can also improve skin wrinkling	
Autologous fat transfer	• Use of patient's own fat	Dubious long-term maintenance of results
	• Good for forehead scars	
Dermabrasion	• Quick therapy	Dubious long-term maintenance of results
	• Minimal side effects	Multiple therapies required
Chemical peels	• Quick	Multiple therapies
	• Easy to administer	• Increase in acid concentration less well tolerated
		Side effects and longer downtime
Injectables	Quick	• Nil significant research into efficacy, new treat- ment modality
	• Easy to administer	
	• Initial results in literature are good	

Treatment	Pros	Cons
Subcision	• Well known technique	• Significant side effects and downtime
	• Easy to utilise	• Delay in seeing end results
		Discomfort during treatment
		Multiple therapies
Tretinoin iontophoresis	• Good initial results	• Technical procedure requiring specialist equip- ment and facilities
		• Dubious long-term treatment maintenance
		• Side effects

on an individual patient's circumstances, preference and aims of treatment outcome.

Declarations

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Guarantor: LP

Contributorship: LP – study design, data collection, analysis of results and writing of manuscript; DM – analysis of results and review of manuscript; KC – analysis of results and review of manuscript

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