

Comparing the Efficacy of Intra-dermal Platelet Rich Plasma (PRP) Versus 50% Trichloroacetic Acid (TCA) using Cross Technique for Atrophic Acne Scars

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ABSTRACT

Objective: To compare the efficacy of intra-dermal platelet rich plasma (PRP) versus 50% trichloroacetic acid (TCA) using chemical reconstruction of skin scars (CROSS) technique in the treatment of atrophic acne scars.

Study Design: Non-randomised controlled trial.

Place and Duration of Study: Sheikh Zayed Hospital, Rahim Yar Khan, from October 2019 to April 2020.

Methodology: In this study, cases of either gender and age 20 to 40 years with atrophic acne scars were included. The severity of the scar was graded on the basis of global acne scarring grading system. The cases in group A were managed by monthly injections of 1 ml intra-dermal PRP every month; while those in Group B were given treatment with 50% TCA, which was applied by CROSS technique every month. Both treatments were offered for three months. They were assessed at every four weeks for initial three months. Then these cases were followed another three months and final outcome was seen at 6th month.

Results: In this study, there were 92 cases, 46 in each group. The mean age in group A and B was 27.72 ± 8.05 vs. 26.50 ± 8.20 years ($p = 0.474$). The mean global scar score at baseline was 36.07 ± 5.37 vs. 38.70 ± 4.80 ($p = 0.015$). The mean scar score at 4 weeks was 28.87 ± 5.27 vs. 29.00 ± 3.07 ($p = 0.885$), at 8 weeks 23.22 ± 4.10 vs. 23.11 ± 2.49 ($p = 0.878$), at 12 weeks 14.15 ± 3.05 vs. 17.57 ± 4.51 ($p < 0.001$), and at 24 weeks it was 7.09 ± 1.46 vs. 10.09 ± 3.58 ($p = < 0.001$).

Conclusion: PRP is significantly better than 50% TCA in reducing post-acne atrophic scars.

Key Words: Acne, Atrophic scar, Platelet rich plasma, 50% TCA.

How to cite this article: Mumtaz M, Hassan T, Shahzad MK, Hanif N, Anwar S, Anjum R. Comparing the Efficacy of Intra-dermal Platelet Rich Plasma (PRP) Versus 50% Trichloroacetic Acid (TCA) using Cross Technique for Atrophic Acne Scars. *J Coll Physicians Surg Pak* 2021; **31(01)**:55-59.

INTRODUCTION

Acne is one of the highly concerning entity encountered in adolescent population and post-acne scarring adds more to overall morbidity in terms of cosmetic disfigurement, and social and psychological stress. Its distribution is variable and it is more prevalent in young age, especially in females. The most common areas of involvement are those with rich distribution of pilosebaceous glands. The prevalence in general population is around 1%.^{1,2}

Acne scars are broadly subdivided into three major categories: atrophic, hypertrophic, and keloids: where atrophic scars are the highest in number. Atrophic scars are further classified into ice pick, rolling, and boxcars.

The underlying pathophysiology is multifactorial and largely relies upon chronic inflammation and scarring at the site of subcutaneous matrix. The major steps in scarring include inflammation, granulation formation and matrix remodelling. They are usually asymptomatic and might have some itching or other symptoms, if associated with super-added infection.^{3,4} The diagnosis is made clinically; and severity is further graded on the basis of various tools and scores. Tissue biopsy and histopathology is seldom needed, which guides towards particular subtype. The most commonly practised scales are Vancouver scar scale (VSS), Good MAN and Baron grading scale, patient and observer scar scale etc.^{5,6}

Early diagnosis and prompt treatment is the key to success. The treatment options are broadly divided into energy-based and non-energy-based, where former comprises lasers (both ablative and non-ablative), fractional, radiofrequency lasers; and latter contains derm abrasions, micro needling, fillers, peels and drugs. Each carries different degree of success and side effect profile.^{7,8}

Platelet rich plasma is a relatively newer entity; while, trichloroacetic acid was used conventionally over years in different strengths ranging from 35% to 100% in various application

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Received: September 14, 2020; Revised: December 19, 2020;

Accepted: December 30, 2020

DOI: <https://doi.org/10.29271/jcpsp.2021.01.55>

methods; and had shown promising results but side effect profile was the concern, which led to the search of newer therapies like PRP.^{9,10} The studies were also done on a particular group of scar and with smaller sample size of 25 to 30.

That is why this study was planned to compare newer PRP with conventional TCA, especially with a strength of 50% (for which data is scarce) in all types of atrophic acne scars to find the better agent.

METHODOLOGY

The approval of this study was taken from Ethical Review Committee via reference number 02/IRB/SZMC/SZH. This was a non-randomised controlled trial, that was carried out from October 2019 to April 2020 at Sheikh Zayed Hospital, Rahim Yar Khan. Cases of either gender with age 20 to 40 years suffering from atrophic type of acne scar lasting for less than one year and no active acne were included for which diagnosis was made clinically with history of acne vulgaris and scarring at the facial area. Exclusion criteria consisted of patients with scar duration more than one year, any bleeding disorder, presence of active acne on face, active infection (purulent discharge from the site), keloid tendency, pregnancy and lactating women. An informed written consent was taken from each subject to include in this study; and were also explained the procedures and side effect profiles of the drugs used in the study.

The sample size was calculated as 92 (46 in each group) by keeping the confidence level equal to 95%, power equal to 80%, and the very good response as 46.7% with PRP and 20% with 100% TCA as there was no one-to-one comparison with 50% TCA before.¹¹ Total 134 cases were analysed, 92 were finalised, which were divided into two groups. There was no lost-to-follow-up. Final data was analysed on these 46 cases of each group.

Then these cases were subdivided into two equal groups by simple lottery method. Thorough history was taken of each patient, including demographic history, disease history, age of onset, duration of disease and treatment history. Dermatological examination was performed and grading of acne scars was done according to Goodman and Baron global acne scarring grading system (Table I). Then mean score at baseline was calculated.

Patients were explained in detail about the procedure. Of cases in group A, complete blood count, including baseline platelet count, was performed. Area of interest in both the groups was wiped with spirit before procedure.

In group A, after cleaning the face with spirit, it was anaesthetised using a topical anaesthetic cream Anestin (10.56% lidocaine) for about 45 minutes. Autologous whole blood of 20mL was collected and pure platelet rich plasma was prepared, using a special sterile vacutainer tube containing an anticoagulant. PRP was obtained manually by two-step procedure, using a centrifuge machine. First spin was performed at 1500 revolu-

tions per minute (RPM) for 10 minutes. Second spin was performed at a rate of 4000 RPM for 10 minutes. Thus obtaining a two-part plasma. Upper two-thirds was platelet poor plasma (PPP) and was discarded. Lower 1/3rd was platelet rich plasma (PRP).

Just before injection, 0.1ml of calcium chloride was added for each 1 ml of PRP to activate the platelets. PRP was injected intradermally through a 30G needle (insulin syringe) deep to each scar on both cheeks. The amount injected was sufficient to elevate the scar and the end point was taken as the elevation of scar. Total amount injected was 3-4 mL depending on the number of scars. After injecting, the site was gently massaged and compressed for a few seconds to control the bleeding. Patients were given topical sunscreens. All the patients were assessed for side effects like pain, edema, erythema, hyperpigmentation and development of active acne lesions during the treatment.

In patients with group B, after cleaning the face with spirit, local anaesthetic was not used and patients were comfortable during the procedure. The skin was stretched to reach the bottom of scar and 50% TCA was then focally applied on entire depth of atrophic acne scar using a tooth pick; 50% TCA was selected as higher concentrations had more side effect profile. Care was taken to avoid spillage to the surrounding skin. The skin was monitored carefully until a frosted appearance was seen. Frosting was generally seen in 10-15 seconds and was a result of coagulation of epidermal and dermal proteins. Patients were instructed to wash their face with water immediately after 7-10 minutes. Sunscreen was applied after the procedure.

Both treatments were offered for only three months. They were assessed at every four weeks for initial three months. Then these cases were followed another three months and final outcome was seen at six month, which was assessed on the basis of global acne scarring grading system, and mean score was calculated.

SPSS version 24 was used for data analysis. Qualitative data, like gender, was presented as frequency and percentages; while quantitative data like age, BMI, duration of scar and global acne score were presented as mean \pm S.D. Chi-square and independent sample t-tests were used for categorical and numerical data stratification, respectively; and $p < 0.05$ was taken as significant.

RESULTS

In this study, there were 92 cases, 46 in each group. The mean age in group A and B was 27.72 ± 8.05 vs. 26.50 ± 8.20 years ($p = 0.474$). Table II further reveals BMI, duration of scar and gender distribution in both groups. The mean global scar score at baseline was 36.07 ± 5.37 vs. 38.70 ± 4.80 with $p = 0.015$. There was no significant difference in mean scar score at four weeks ($p = 0.885$), at eight weeks ($p = 0.878$), while there was significant difference at 12 weeks ($p < 0.001$) and at 24 weeks with p -value of < 0.001 , as displayed in Table III.

Table I: Goodman and Baron's quantitative scar scale.

Grade (type)	No. of lesions		
	1-10	11-20	>20
A. Milder scarring - macular erythematous, pigmented, mildly atrophic dish-like.			
B. Moderate scarring - moderately atrophic dish like, punched out small scars with, shallow bases but atrophic areas (<5 mm).	2 pts	4 pts	6 pts
C. Severe scarring - punched out with deep but normal bases, punched out with deep abnormal bases, linear or trpuged dermal scarring, deep and broad atrophic areas.	3 pts	6 pts	9 pts
D. Hyperplastic papular scars.	2 pts	4 pts	6 pts
E. Hyperplastic keloidal or hypertrophic scars.	Area <5 cm ² - 6 pts	Area 5-20 cm ² - 12 pts	Area >20 cm ² - 18 pts

Table II: Demographics of study subjects (n= 46 each).

Study variables	Group A	Group B	p-value
Age (years)	27.72±8.05	26.50±8.20	0.474
BMI (kg/m ²)	24.83±3.83	24.41±2.61	0.547
Duration of scar (months)	6.37±1.44	5.54±1.50	0.008
Gender			
Male	27 (58.7%)	22 (47.8)	0.296
Female	19 (41.3)	24 (52.2%)	

Table III: Comparison of mean acne score (n=46 in each group).

Global acne score	Group A	Group B	p-value
Pre-treatment	36.07±5.37	38.70±4.80	0.015
4 weeks	28.87±5.27	29.00±3.07	0.885
8 weeks	23.22±4.10	23.11±2.49	0.878
12 weeks	14.15±3.05	17.57±4.51	<0.001
24 weeks	7.09±1.46	10.09±3.58	<0.001

DISCUSSION

Post-acne scarring is a well known entity and is the most common subtype. Considering its asymptomatic nature, globally the most common reason to acquire therapies is cosmetic disfigurement. There are different treatment modalities, but are time-consuming as well as variable in efficacies. Longer duration of treatment and slow response to the therapy are the two major concerns for losing the patients on long term follow-up.^{12,13}

In the present study, intra-dermal PRP was significantly better at 12 and 24 weeks as compared to 50% TCA applied by CROSS technique, and it was seen that mean scar score at 12 weeks was 14.15 ± 3.05 vs. 17.57±4.51 (p <0.001); and at 24 weeks it was 7.09 ± 1.46 vs. 10.09 ± 3.58 with p value of <0.001. There was lack of data regarding one-to-one comparison of these two agents, and especially in this (50%) strength as TCA has been used in various strengths and combination, especially 70% and 100% with or without combination of other agents.^{9,11}

According to a study done by Nofal *et al.*, they compared PRP with 100% of TCA applied by same CROSS technique, and it was seen that PRP was better than TCA just like the findings of the present study. They further found that, in grade I of acne scar, there were no cases, in grade 2 efficacy of PRP vs. 100% TCA was 40% vs. 33% in grade 3, 33% vs. 40%, and in grade 4, it was 26.7% each.¹¹

Apart from one-to-one comparison, for which very little data was available, they also compared these modalities individually with other therapeutic agents as well. In another study done by Porwal *et al.*, they compared PRP combined with derma roller vs. derma roller alone and treatment assessment was done by Goodman and Baron's quantitative scores. It was found that a statistically significant difference was noted between the two groups after the treatment (p<0.05), favouring PRP and derma roller group. They further described that improvement was noted in 58.58% cases in PRP and derma roller combined and in 43.03% in derma roller alone; and on quantitative assessment, the mean acne score after third session was 18.58 ± 4.12 in PRP and derma roller combined as compared to 23.58±5.71 in derma roller alone group.¹⁴

Aamir *et al.*, on other hand, assessed the utility of 35% TCA on the basis of laid down criteria with CROSS technique in their nine cases, out of which seven were females. They found that five (55.5%) out of nine patients experienced good clinical response, while four patients had excellent, *i.e.* more than 70% improvement. Only one patient rated the results to be >25% (fair) despite the clinical improvement being more than 50%.¹⁵

In another study done by Kaur *et al.*, where they analysed multiple outcomes in cases managed with 50% TCA by using in CROSS technique. They revealed that in all the patients,

scar grading improved from grade 4 to grade 2 and results were graded excellent, good, and fair in 6, 3, and 1 patients respectively.¹⁶

According to another systemic review analysis done by Kravvas *et al.*, they described that all of their cases managed with micro-needling achieved some improvement in scar appearance in the range of 31–62%. Dermal fillers also led to favourable outcomes. In particular, treatment with PPMA led to improvement in 84% of patients. Utilising chemical peels, trichloroacetic acid (TCA) CROSS achieved >70% improvement in 73.3% of patients, whereas 20% glycolic acid did not lead to any improvement in 25%.¹⁷

Ramadan *et al.* performed a split-face study on 20 patients only with rolling acne scars, comparing the effects of subcision with those of 100% TCA CROSS chemical peel. They found a mean decrease in the size of scars following subcision and peeling by $0.3867 \pm 0.09 \text{ cm}^2$ and $0.08657 \pm 0.090 \text{ cm}^2$ ($p < 0.01$).¹⁸ In a study done by Narayanan *et al.*, the combination of subcision, 50% TCA with CROSS technique was used as in the present study was and the pre- and post-treatment outcome was assessed in the form of photographic record; and it was observed that out of 14 patients with grade 4 acne scars, 9 (64.3%) patients improved to grade 2, and 5 (35.7%) patients improved to Grade 3. Out of 10 patients with Grade 3 scars, 6 (60%) patients improved to grade 1, and 4 (40%) patients were improved to grade 2 at the end of the study. All five patients with Grade 2 scars showed significant improvement from baseline.¹⁹

Interestingly, in this study more males were seen with acne scarring as compared to females; though in adulthood it is more common in females. The reason for this could not be found. This study was limited to only three months. Treatment could have been extended beyond three months in both good or slow responders to attain further efficacy. No side effect profile data was collected in the scope of this study.

CONCLUSION

Platelet rich plasma (PRP) is recently proving its role in various modalities and is found significantly better than 50% TCA in reducing post-acne atrophic scars. Future studies can help in assessing its long term benefits and can also look for side effect profiles.

CONFLICT OF INTEREST:

The authors declared no conflict of interest.

ETHICAL APPROVAL: Ethical approval was taken from Institutional Review Board Sheikh Zayed Hospital, Rahim Yar Khan prior to the initiation of the study.

PATIENTS' CONSENT:

Informed consents were obtained from patients to publish the data concerning this case.

AUTHORS' CONTRIBUTION:

MM: Literature review, article writing, design, conception, data collection, analysis.

TH, MKS: Conception, design, analysis.

NH: Study design.

SA, RA: Conception.

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