

Research Article

Effect and Safety Analysis of PRP and Yifu Combined with Ultrapulsed CO₂ Lattice Laser in Patients with Sunken Acne Scar

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Objective. To investigate the effect and safety of PRP and Yifu combined with ultrapulsed CO₂ lattice laser in patients with sunken acne scar. **Methods.** 700 subjects were selected from the group of patients with sunken acne scar treated in our hospital from November 2010 to December 2020. They were divided into control group ($n = 350$) and study group ($n = 350$). The grouping was mainly based on the random number table method. Patients in the control group were treated with ultrapulse CO₂ lattice laser, while those in the study group were treated with ultrapulse CO₂ lattice laser combined with PRP and Yifu. The clinical effect, scar improvement and quality of life before and after treatment, and adverse events during treatment were compared between the two groups. The clinical effect was categorized into cure after treatment, significant effect, effective, and ineffective. The total effective rate = 1 – ineffective rate. **Results.** After treatment, the total effective rate of the study group (81.43%) was higher than that of the control group (70.00%). After treatment, ECCA, VSS scores, daily activities, symptoms and feelings, work and study, leisure and entertainment, interpersonal relationship, treatment status, and total scores were all lower in both groups than before treatment, and the study group was lower than the control group. During the treatment, the incidence of adverse events in the study group (17.33%) was lower than that in the control group (28.57%), $P < 0.05$. **Conclusion.** PRP and Yifu combined with ultrapulse CO₂ lattice laser in the treatment of sunken acne scar can effectively improve the scar, reduce the incidence of adverse events, and the treatment effect is obvious, which can improve the quality of life of the patients.

1. Introduction

Acne is a chronic inflammatory skin disease that occurs on the sebaceous glands of hair follicles caused by a variety of causes. The main causes are related to abnormal hormone secretion, excessive sebaceous secretion, bacterial infection of propionibacterium acne, keratosis and blockage of hair follicle sebaceous ducts, and other factors. Women with endocrine system disorders and irregular menstruation and adolescent youths are at high risk [1, 2]. Acne is easy to relapse and aggravate repeatedly, resulting in complications such as pigmentation, persistent erythema, and sunken scar, among which sunken acne scar is more serious [3], mostly like orange peel, ice cone, meteorite crater, and so on. It greatly affects the normal life and mood of the patients [4, 5].

The treatment cycle of sunken acne scar is long. At present, in addition to drug therapy, physiotherapy is also increasingly used in the treatment of acne. Ultrapulsed CO₂ lattice laser can accelerate the formation of keratinocytes, promote the proliferation and rearrangement of facial collagen and elastic fibers, and improve the symptoms of sunken scar, but the effect is slow when used alone [6]. Platelet-rich plasma (PRP) is a synergistic effect of a variety of growth factors released after platelet activation, which promotes the proliferation and differentiation of local repair cells and the synthesis of extracellular matrix, thus enhancing the ability of tissue regeneration and repair [7, 8]. At present, it has been widely used in clinical repair of chronic wound, bone and soft tissue injury [9–11]. Yifu is used to treat chronic skin ulcer wounds and burn wounds [12], in which the

effective ingredient is recombinant human surface growth factor [13], which can accelerate the speed of cell wound repair [14]. The purpose of this study is to investigate the effect and safety of PRP and Yifu combined with ultrapulsed CO₂ lattice laser on scar improvement in patients with sunken acne scar.

2. Materials and Methods

2.1. General Information. The basic data of the following two groups of patients was calculated and compared, suggesting that the difference in the data is not significant in the statistical study ($P > 0.05$), and then there was comparability between the two groups. 700 subjects were selected from the group of patients with sunken acne scar treated in our hospital from November 2010 to December 2020. They were divided into control group ($n = 350$) and study group ($n = 350$). The grouping was mainly based on the random number table method. The control group included the following: age 15–31 years, average 20.14 ± 2.07 years, 66 males and 84 females, and the course of disease ranged from 0.5 to 1.5 years, with an average of 0.98 ± 0.11 years. The study group included the following: age 16–32 years, mean 20.20 ± 2.11 years, 64 males and 86 females, and the course of disease ranged from 1 to 2 years, with an average of 1.09 ± 0.08 years. Diagnostic criteria: the diagnostic criteria of sunken acne scar in the recommended guidelines for clinical treatment of keloid are used as a reference. Inclusion criteria: those who meet the diagnostic criteria; those diagnosed by microscopic examination; those diagnosed by sex hormone test; those with informed consent of patients, etc. Exclusion criteria: patients with hypertrophic scar, patients with multiple infectious diseases, mental disorders, etc. The Medical Ethics Committee of our hospital has approved the implementation of this study.

2.2. Method. The patients in the control group were smeared with the appropriate amount of compound lidocaine cream (Tongfang Pharmaceutical Group Co., Ltd., Chinese medicine standard H20063466, specification: lidocaine 25 mg ropivacaine 25 mg/g) and then received ultrapulsed CO₂ lattice laser, using pulsed carbon dioxide laser therapy apparatus (Beijing Hongqiang Furei Technology Co., Ltd., national equipment injection 20163241822, model: 10600AH). The full coverage electroacupuncture mode was selected to adjust the parameters around the scar with the frequency of 4 Hz for lattice, 20 min/times, once every 15 days. In the study group, on the basis of the control group, 40 ml of autologous elbow venous blood was taken 2 hours before treatment, and PRP was extracted by density gradient centrifugation. After the completion of ultrapulse CO₂ lattice laser treatment, the wound was evenly smeared with PRP for 3 times with an interval of 1 month. At the same time, apply Yifu on the whole face after the laser lattice (national medicine standard S20020112, specification: 100,000 IU, Guilin Huanowi Genome Pharmaceutical Co., Ltd.).

2.3. Observation Index. ① Clinical effect (including the cure after treatment): sunken scar basically disappeared, flat; significant effect: most of sunken scar disappeared, flat; effective: sunken scar improved compared with before treatment, but still not improved; ineffective: sunken scar did not improve or even aggravated. According to the consensus on early treatment of scar (version 2020), the total effective rate = 1–ineffective. ② Improvement of scar (including the acne scar weight score (ECCA)) before and after treatment reflects the severity of the scar, and the total score is 100 points; the higher the score is, the more serious the scar is; the Vancouver scar scale (VSS) score reflects the scar recovery, and the total score is 10; the higher the score is the worse the recovery is. ③ Quality of life (Including dermatology quality of life index (DLQI)) before and after treatment is divided into daily activities, symptoms and feelings, work and study, leisure and entertainment, interpersonal relationships, and treatment; each score is 3; the total score is 18; the higher the score is, the lower the quality of life is. ④ Adverse events include the occurrence of erythema, edema, and pigmentation during treatment, and the incidence of adverse events = (erythema + edema + pigmentation) cases/total cases $\times 100\%$.

2.4. Statistical Method. Statistical analysis was carried out with SPSS26.0 software, the measurement data were expressed as mean \pm standard deviation ($\bar{x} \pm S$), the frequency was expressed as percentage (%), and the counting data were tested by χ^2 test. Continuous variables were compared using Student's *t*-test. $P < 0.05$ was regarded as statistically significant.

3. Result

3.1. Clinical Effect. After treatment, it was found that the total clinical effective rate of patients in the control group (70.00%) was significantly lower than that in the study group (81.43%). After calculation, $P < 0.05$, as detailed in Table 1.

3.2. Scar Improvement. Compared with those before treatment, the scores of ECCA and VSS in the two groups decreased after treatment, and those in the study group were lower than those in the control group ($P < 0.05$), as detailed in Table 2.

3.3. Quality of Life. After treatment, the daily activities, symptoms and feelings, work and study, leisure and entertainment, interpersonal relationship, treatment, and total score in the study group were lower than those in the control group ($P < 0.05$), as detailed in Table 3.

3.4. Adverse Reaction. During the treatment, the incidence of adverse reactions in the study group was 17.33%, which was lower than that in the control group (28.67%). It was calculated that $P < 0.05$, as detailed in Table 4.

TABLE 1: Comparison of the clinical efficacy of the two groups (n (%)).

Groups	n	Cure	Significant effect	Effective	Invalid	Total effective
Control group	350	47 (13.43)	82 (23.43)	116 (33.14)	105 (30.00)	245 (70.00)
Study group	350	56 (16.00)	91 (26.00)	138 (39.43)	65 (18.57)	285 (81.43)
χ^2						5.232
P						<0.05

TABLE 2: Comparison of the improvement of scar between the two groups ($\bar{x} \pm s$, score).

Groups	n	ECCA		VSS	
		Before treatment	After treatment	Before treatment	After treatment
Control group	350	89.12 \pm 7.08	69.21 \pm 3.15*	8.41 \pm 1.14	5.65 \pm 2.86*
Study group	350	89.83 \pm 6.98	49.38 \pm 2.69*	8.39 \pm 1.21	3.55 \pm 1.72*
T		0.875	8.631	0.147	7.707
P		>0.05	<0.05	>0.05	<0.05

Note. Compared with before treatment, * $P < 0.05$.

4. Discussion

It is clinically believed that the pathogenesis of acne is complex and diverse [15], but it is mainly due to the high level of androgen in the patient's body, and the sebaceous gland is affected by too much androgen secretion, which increases the amount of sebum produced. When combined with exfoliated secretions from the epidermis, it will block pores, resulting in acne [16, 17]. Like greasy, high calorie eating habits, high ambient temperature, staying up late, and high pressure are all inducing factors of acne [18], the typical clinical symptoms of patients are blackhead, pimples, nodular pustules, and cysts [19]. Sunken acne scar is caused by severe acne, repeated attacks, and improper nursing during the inflammatory period, which not only affects the facial beauty of the patients. At the same time, it brings inconvenience to the normal social communication of the patients [20, 21]. Treatments are divided into general treatment, drug therapy, physiotherapy, and other treatments [22]. In physiotherapy, photodynamic therapy, red and blue light irradiation, and intense pulsed light have a certain effect on the elimination of acne red marks [23]. Ultrapulsed CO₂ lattice laser uses a gas laser to emit array-like light beams through an electroacupuncture laser, which can form some tiny thermal injury areas on the skin, where keratinocytes can regenerate quickly and accelerate wound healing, but the curative effect is slow when used alone [24].

The essence of PRP is to separate platelet and plasma by centrifugation and add thrombin to form gel to promote wound healing [25]. The therapeutic mechanism is that platelets can release brain-derived neurotrophic factor, vascular endothelial growth factor, and other growth factors after activation [26, 27]. A large number of studies have proved that PRP can promote the regeneration of cartilage and bone tissue, as well as anti-inflammatory and antibacterial effects [28]. In recent years, it has been widely used in bone and soft tissue reconstruction surgery, plastic surgery, diabetic foot refractory ulcers, and chronic skin ulcers [29]. Yifu is a kind of recombinant human epidermal growth factor for external use. It is mainly derived from

polypeptides in mammals and is mainly used in the treatment of wounds, residual wounds, and ulcerative wounds after skin burns. The effect and safety of recombinant human epidermal growth factor extracted from natural substances are basically the same as those secreted by human body, and there are no side effects. At the same time of improving the symptoms of sunken scar, it also has a certain impact on the quality of life of patients [30].

In the results of this study, the total effective rate of the study group after treatment was higher than that of the control group, and the DLQI score was lower than that of the control group, and during the treatment, the incidence of adverse reactions in the study group was also lower than that in the control group, indicating that PRP and Yifu combined with ultrapulse CO₂ lattice laser can effectively improve the therapeutic effect of patients with sunken acne scar, reduce the occurrence of adverse reactions, and then improve the quality of life of patients. ECCA and VSS are often used to evaluate the degree of scar caused by various skin problems. Yifu can promote the synthesis of nucleic acid and hydroxyproline in the cells around the skin wound, and at the same time, it can participate in inducing epidermal cells to mature and differentiate into stem cells and carry out scar epithelial cell proliferation and granulation tissue differentiation and development, so as to improve the scar condition and restore wound leveling [31, 32]. In the results of this study, the scores of ECCA and VSS in the study group after treatment were lower than those in the control group, indicating that PRP and Yifu combined with ultrapulse CO₂ lattice laser can improve the scar condition of patients with sunken acne scar and facilitate the recovery of the disease.

In summary, PRP and Yifu combined with ultrapulsed CO₂ lattice laser can effectively improve the scar of patients with sunken acne scar, reduce the incidence of adverse reactions, and the treatment effect is obvious, which can improve the quality of life of the patients. However, this study did not analyze the impact of inflammatory response in patients, and a multisample size and multicenter in-depth study is needed.

TABLE 3: Comparison of the scores of quality of life between the two groups ($\bar{x} \pm s$, score).

Groups	n	Daily activities		Symptoms and feelings		Work and study		Leisure and entertainment		Interpersonal relationship		Treatment condition		Total score	
		Before treatment	After treatment	Before treatment	After treatment	Before treatment	After treatment	Before treatment	After treatment	Before treatment	After treatment	Before treatment	After treatment	Before treatment	After treatment
Control group	350	2.23 ± 0.46	1.16 ± 0.24*	2.48 ± 0.22	1.73 ± 0.19*	2.49 ± 0.23	1.71 ± 0.18*	1.64 ± 1.08	1.48 ± 0.55*	2.44 ± 0.33	1.58 ± 0.38*	2.17 ± 0.51	1.69 ± 0.46*	13.27 ± 1.35	10.06 ± 0.51*
Study group	350	2.28 ± 0.39	0.51 ± 0.13*	2.51 ± 0.26	1.30 ± 0.09*	2.43 ± 0.19	1.08 ± 0.20*	1.66 ± 1.10	0.79 ± 0.24*	2.45 ± 0.40	1.03 ± 0.16*	2.12 ± 0.49	0.59 ± 0.30*	13.22 ± 1.33	8.63 ± 0.34*
T		1.015	29.166	1.079	25.050	2.463	28.676	0.159	14.083	0.236	16.337	0.866	24.531	0.323	28.573
P		>0.05	<0.05	>0.05	<0.05	>0.05	<0.05	>0.05	<0.05	>0.05	<0.05	>0.05	<0.05	>0.05	<0.05

Note. Compared with before treatment, * $P < 0.05$.

TABLE 4: Comparison of the occurrence of adverse reactions between the two groups (n (%)).

Groups	N	Erythema	Edema	Pigmentation	Total occurrence
Control group	350	30 (8.57)	35 (10.00)	35 (10.00)	100 (28.57)
Study group	350	21 (6.00)	25 (7.33)	14 (4.00)	60 (17.33)
χ^2					5.439
P					<0.05

Data Availability

The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

Disclosure

Wei Yu is the co-first author.

Conflicts of Interest

The authors declare that there are no conflicts of interest regarding the publication of this article.

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References

- [1] A. K. Gupta, S. G. Versteeg, J. Rapaport, A. K. Hausauer, N. H. Shear, and V. Piguet, "The efficacy of platelet-rich plasma in the field of hair restoration and facial aesthetics—a systematic review and meta-analysis," *Journal of Cutaneous Medicine and Surgery*, vol. 23, no. 2, pp. 185–203, 2019.
- [2] F. Chellini, A. Tani, S. Zecchi-Orlandini, and C. Sassoli, "Influence of platelet-rich and platelet-poor plasma on endogenous mechanisms of skeletal muscle repair/regeneration," *International Journal of Molecular Sciences*, vol. 20, no. 3, 2019.
- [3] J. Emer, "Platelet-rich plasma (PRP): current applications in dermatology," *Skin Therapy Letter*, vol. 24, no. 5, pp. 1–6, 2019.
- [4] F. Veronesi, S. Pagani, P. Torricelli et al., "PRP and MSCs on tenocytes artificial wound healing: an in vitro study comparing fresh and frozen PRP," *Histology and Histopathology*, vol. 33, no. 12, pp. 1323–1334, 2018.
- [5] Y. Zhang, Y. Liu, B. Cai et al., "Improvement of surgical scars by early intervention with carbon dioxide fractional laser," *Lasers in Surgery and Medicine*, vol. 52, no. 2, pp. 137–148, 2020.
- [6] T. Long, A. Gupta, S. Ma, and S. Hsu, "Platelet-rich plasma in noninvasive procedures for atrophic acne scars: a systematic review and meta-analysis," *Journal of Cosmetic Dermatology*, vol. 19, no. 4, pp. 836–844, 2020.
- [7] A. F. Tetila, M. R. S. Breda, R. M. B. Nogueira, G. A. Nai, and C. B. Laposy, "The use of platelet-rich plasma and rosuvastatin in wound healing in rabbits: a longitudinal study," *Advances in Skin and Wound Care*, vol. 32, no. 9, pp. 1–5, 2019.
- [8] E. Schoenberg, J. V. Wang, C. B. Zachary, and N. Saedi, "Treatment of acne scars with PRP and laser therapy: an up-to-date appraisal," *Archives of Dermatological Research*, vol. 311, no. 8, pp. 643–646, 2019.
- [9] A. K. Hausauer and S. Humphrey, "The physician's guide to platelet-rich plasma in dermatologic surgery part I: definitions, mechanisms of action, and technical specifications," *Dermatologic Surgery*, vol. 46, no. 3, pp. 348–357, 2020.
- [10] F. Parra, D. E. Morales-Rome, R. Campos-Rodríguez, T. R. Cruz-Hernández, and M. E. Drago-Serrano, "Effect of platelet-rich plasma on patients after blepharoplasty surgery," *Orbit*, vol. 37, no. 2, pp. 81–86, 2018.
- [11] M. Asif, S. Kanodia, and K. Singh, "Combined autologous platelet-rich plasma with microneedling verses microneedling with distilled water in the treatment of atrophic acne scars: a concurrent split-face study," *Journal of Cosmetic Dermatology*, vol. 15, no. 4, pp. 434–443, 2016.
- [12] R. Alves and R. Grimalt, "A review of platelet-rich plasma: history, biology, mechanism of action, and classification," *Skin Appendage Disorders*, vol. 4, no. 1, pp. 18–24, 2018.
- [13] M. El-Domyati, H. Abdel-Wahab, and A. Hossam, "Microneedling combined with platelet-rich plasma or trichloroacetic acid peeling for management of acne scarring: a split-face clinical and histologic comparison," *Journal of Cosmetic Dermatology*, vol. 17, no. 1, pp. 73–83, 2018.
- [14] N. Wu, H. Sun, Q. Sun et al., "A meta-analysis of fractional CO₂ laser combined with PRP in the treatment of acne scar," *Lasers in Medical Science*, vol. 36, no. 1, pp. 1–12, 2021.
- [15] S. M. Refahee, M. A. Aboulhassan, O. Abdel Aziz et al., "Is PRP effective in reducing the scar width of primary cleft lip repair? a randomized controlled clinical study," *The Cleft Palate-Craniofacial Journal*, vol. 57, no. 5, pp. 581–588, 2020.
- [16] A. Kaushik and M. S. Kumaran, "Platelet-rich plasma: the journey so far," *Indian Dermatol Online Journal*, vol. 11, no. 5, pp. 685–692, 2020.
- [17] G. H. Sasaki, "Micro-needling depth penetration, presence of pigment particles, and fluorescein-stained platelets: clinical usage for aesthetic concerns," *Aesthetic Surgery Journal*, vol. 37, no. 1, pp. 71–83, 2017.
- [18] J. U. Shin, S.-W. Kang, J. J. Jeong, K.-H. Nam, W. Y. Chung, and J. H. Lee, "Effect of recombinant human epidermal growth factor on cutaneous scar quality in thyroidectomy patients," *Journal of Dermatological Treatment*, vol. 26, no. 2, pp. 159–164, 2015.
- [19] E. Schoenberg, M. O'Connor, J. V. Wang, S. Yang, and N. Saedi, "Microneedling and PRP for acne scars: a new tool in our arsenal," *Journal of Cosmetic Dermatology*, vol. 19, no. 1, pp. 112–114, 2020.
- [20] S. Min, J. Y. Yoon, S. Y. Park, J. Moon, H. H. Kwon, and D. H. Suh, "Combination of platelet rich plasma in fractional carbon dioxide laser treatment increased clinical efficacy of for acne scar by enhancement of collagen production and modulation of laser-induced inflammation," *Lasers in Surgery and Medicine*, vol. 50, no. 4, pp. 302–310, 2018.
- [21] I.-H. Huang and Y.-C. Huang, "Comment on "platelet-rich plasma and its utility in the treatment of acne scars: a systematic review,"" *Journal of the American Academy of Dermatology*, vol. 82, no. 1, pp. e29–e32, 2020.

- [22] Z. Chen, Y. Wu, N. Turxun, Y. Shen, and X. Zhang, "Efficacy and safety of platelet-rich plasma in the treatment of severe burns," *Medicine*, vol. 99, no. 45, Article ID e23001, 2020.
- [23] R. Alves and R. Grimalt, "Platelet-rich plasma and its use for cicatricial and non-cicatricial alopecias: a narrative review," *Dermatologic Therapy*, vol. 10, no. 4, pp. 623–633, 2020.
- [24] P. Woo and T. Murry, "Short-term voice improvement after repeated office-based platelet-rich plasma PRP injection in patients with vocal fold scar, Sulcus, and Atrophy," *Journal of Voice*, 2021.
- [25] M. E. Jones, J. Mclane, R. Adenegan, J. Lee, and C. A. Ganzer, "Advancing keloid treatment: a novel multimodal approach to ear keloids," *Dermatologic Surgery*, vol. 43, no. 9, pp. 1164–1169, 2017.
- [26] S. Li, Y. Liu, Z. Huang, Y. Kou, and A. Hu, "Efficacy and safety of nano-silver dressings combined with recombinant human epidermal growth factor for deep second-degree burns: a meta-analysis," *Burns*, vol. 47, 2020.
- [27] A. J. Singer, J. Toussaint, W. T. Chung, S. McClain, V. Raut, and L. Rosenberg, "The effects of platelet rich plasma on healing of full thickness burns in swine," *Burns*, vol. 44, no. 6, pp. 1543–1550, 2018.
- [28] P. Karakol and M. Bozkurt, "Recent strategic approach in postburn extremity scars and contractures," *Journal of Plastic Surgery and Hand Surgery*, vol. 55, pp. 1–16, 2020.
- [29] V. Belebecha, R. Casagrande, M. R. Urbano et al., "Effect of the platelet-rich plasma covering of polypropylene mesh on oxidative stress, inflammation, and adhesions," *International Urogynecology Journal*, vol. 31, no. 1, pp. 139–147, 2020.
- [30] N. F. Agamia, O. Sorrow, M. Alrashidy, A. A. Tawfik, and A. Badawi, "Clinical and histopathological comparison of microneedling combined with platelets rich plasma versus fractional erbium-doped yttrium aluminum garnet (Er: YAG) laser 2940 nm in treatment of atrophic post traumatic scar: a randomized controlled study," *Journal of Dermatological Treatment*, vol. 32, pp. 1–8, 2020.
- [31] Z. F. Yin and S. M. Wang, "Effect of rhEGF on the levels of serum EGF and inflammatory cytokines in patients with oral and maxillofacial trauma," *Shang Hai Kou Qiang Yi Xue*, vol. 26, no. 6, pp. 654–657, 2017.
- [32] Z. Gong, C. Wang, L. Ni et al., "An injectable recombinant human milk fat globule-epidermal growth factor 8-loaded copolymer system for spinal cord injury reduces inflammation through NF- κ B and neuronal cell death," *Cytotherapy*, vol. 22, no. 4, pp. 193–203, 2020.