http://bjas.bu.edu.eg

Role of Platelet Rich Plasma in Healing of Chronic Leg Ulcers

E.M.Kilany, H.M.Sobaih, M.T.Younes and A.K.Meselhy

General Surgery Dept., Faculty of Medicine, Benha Univ., Benha, Egypt

E-Mail:Ahmed Kameel@gmail.com

Abstract

Platelet-rich plasma (PRP) is characterized as a segment of the plasma part of autologous blood having a platelet fixation above standard. PRP likewise has been alluded to as platelet-improved plasma, platelet-rich concentrate, autologous platelet gel, and platelet releasate. To research the advantages of infusing platelet rich plasma in mending of constant leg ulcers. This is a solitary – focus, planned, randomized, controlled examination that was directed in General Surgery Department, Faculty of Medicine, Benha University incorporating 100 patients giving persistent leg ulcers and introduced to us between January 2018 to June 2018. This investigation was completed in 100 patients with ongoing leg ulcers were remembered for this examination, patients were arbitrarily partitioned in two equivalent gatherings An and B. The magnificent outcomes appeared in ongoing investigations have just served to affirm the great results depicted in our examination. In ten patients that we treated, the outcomes got following consummation of the uses of PRP were acceptable or phenomenal. The utilization of PRP in remedial cycles identified with Plastic, Reconstructive and Esthetic Surgery is presently a genuine alternative in generally open and private emergency clinics in numerous nations; circumstance that shows the advances that have happened over ongoing years in tissue recovery treatment.

Keywords: Platelet rich plasma, Healing of chronic leg ulcers.

1. Introduction

The significant elements of platelets are forestalling intense blood misfortune and fixing vascular dividers and nearby tissues after injury.

During wound recuperating, platelets are actuated by contact with collagen, presented to the circulatory system after endothelial injury. Platelets discharge put away intercellular go betweens and cytokines from the cytoplasmic pool and delivery their α -granule content after accumulation. This discharge is extraordinary in the primary hour and platelets keep orchestrating more cytokines and development factors from their mRNA holds for at any rate an additional 7 days.

In excess of 800 distinct proteins are emitted into the encompassing media, having a paracrine impact on various cell types: myocytes, ligament cells, mesenchymal foundational microorganisms from various starting points, chondrocytes, osteoblasts, fibroblasts and endothelial cells. Cell multiplication, angiogenesis and cell relocation are invigorated, bringing about tissue recovery. There are additionally reports affirming that platelets emit antimicrobial peptides, proposing an antimicrobial impact. Different properties were at that point demonstrated for platelets identified with their mitigating and pain relieving impacts.

Platelet-determined items incorporate platelet-rich plasma (PRP), which can be utilized with or without past platelet enactment. Such arrangements have been utilized since the 1970s and they have been progressively famous since the 1990s. From that point forward, various methods of planning PRP have arisen: from traditional blood centrifugation to business frameworks; initiated by adding collagen, calcium or potentially thrombin, by glass contact or by freezing cycles; applied as platelet suspension or as a gel; and the technique keeps on expanding. PRP fills in as a development factor agonist and has both mitogenic and chemotactic properties. It contains a significant level of platelets and a full supplement of thickening and development factors.

Notwithstanding use in the therapy of persistent skin and delicate tissue ulcerations, distributions with respect to the utilization of PRP incorporate periodontal and oral medical procedure, maxillofacial medical procedure, muscular and injury medical procedure, restorative and plastic medical procedure, spinal medical procedure, heart sidestep a medical procedure, and consumes.

PRP is anything but difficult to create with insignificant exertion and can be set up varying at the purpose of care. In a two-venture measure, entire blood from the patient is first centrifuged to isolate the plasma from stuffed red platelets and afterward further centrifuged to isolate PRP from platelet-helpless plasma. This concentrate is then enacted with the expansion of thrombin or calcium, bringing about a coagulated platelet gel. Clinically important PRP contains in any event 1,000,000 platelets for each microliter. Lesser focuses can't be depended on to improve wound mending, and more prominent fixations have not been appeared to build wound recuperating.

2. Aim of the work

Aim of this work is

- To investigate the benefits of injecting platelet rich plasma in healing of chronic leg ulcers.
- To emphasize the use of platelet rich plasma in chronic leg ulcers based on the clinical outcomes.
- To determine the effect of platelet rich plasma and how to use in chronic leg ulcers.

3. Patients and methods

This is a single –center, prospective, randomized, controlled study that was conducted in General Surgery Department, Faculty of Medicine, Benha University including 100 patients presenting with chronic leg ulcers and presented to us between JAN 2018 to JUNE 2018.

This study was carried out in 100 patients with chronic leg ulcers were included in this study, patients were randomly divided in two equal groups A & B.

- **Group A**: includes 50 patients which PRP was used upon their wounds.
- **Group B**: includes 50 patients with traditional methods of dressing.

The study was ethically approved from the ethical committee of general surgery department prior to its conduct.

Weekly follow up was performed up to 12 weeks by measuring the maximum vertical and horizontal diameters of the ulcer and comparing the results in an excel sheet for every patient with a registered photography at each follow up session for each patient. Minor and major complications were assessed during the follow up and recorded.

3.1 Inclusion criteria

- Adults from 18 to 65 years old
- Sex: Both genders.
- Any chronic leg ulcer for example (Diabetic ulcer, venous ulcer, ischaemic ulcer, chronic non healing ulcers etc.

3.2 Exclusion criteria

- Patients with uncontrolled DM, cardiac diseases, Liver cell failure patients, immunocompromised patients ...etc.
- Patients with low platelet count below normal level (ITP, liver cell failure, blood dyscrasias).
- Collagen disease.

•

- HB level below 10 gm %.
 - Operative technique:
 - 1. Pre-operative
 - History taking.
 - Clinical examination
 - Investigations: ECG, routine laboratory investigations: (CBC, Coagulation profile, Liver functions, kidney functions, RBS, HCVAb, HBVsAg

2. Operative techniques

Blood samples were collected from the patients in the outpatient clinic then PRP was prepared and PRP paste was put over the wound.

3. Post-operative care and follow-up

Dressings were carried out every week by injection and paste. All patients were followed up and evaluated every week till 12 weeks then 6 months for pain, wound infection, decrease in ulcer size and increase in discharge.

So, this study was performed according to the following steps:

1 st step	Reviewing literature dealt with research topic
2 nd step	Selection of patients according to inclusion and exclusion criteria
3 rd step	Statistical analysis for collected data
4 th step	Writing down the founded data fulfilling the thesis

3.3 Administrative design

- This study was endorsed by moral advisory group of Benha University.
- Collected information was introduced in tables and reasonable diagrams and investigated by the PC programming utilizing proper techniques.
- Discussion was done on outcomes contrasted with related significant writing and logical examination to clarify the explanations behind giving such outcomes.

Data collection

3.4 General measures to be followed in both groups:

- Full vascular appraisal was done to evaluate the requirement for a revascularization strategy before the treatment.
- The segment information and the related co morbidities were explored.
- Laboratory examinations and x-beam foot were done just as blood vessel and venous duplex for all patients.
- Meticulous ulcer appraisal was accomplished for all patients as respects starting size of the ulcer, surface zone figuring (length X width), site of the ulcer i.e.: (plantar forefoot, plantar mid foot, plantar rear foot,

dorsum mid foot, dorsum forefoot), granulation tissue, shape, edge, base.

- Initial photographs are taken for all ulcers just as progressive photographs were taken for each case at each booked meeting each week all through the development for viable examination of the result
- All cases had a negligible debridement preceding commencement of the focused on treatment.
- Cases in the two gatherings were scrubbed with ordinary saline and saline bandage dressing was utilized as an essential dressing.
- All cases with plantar foot ulcers were exposed to offloading utilizing in-shoe orthoses to ease plantar pressing factor at the locales of ulceration.
- All patients were agreed for one of the focused on treatment.
- All cases are told to visit our unit at booked meeting each week for a maximal subsequent time of 12 weeks.

3.5 PRP preparation & application

• Approximately 50 ml of venous blood is gathered from every patient at the hour of the treatment, in an at first arranged sterile vacuum vials, i.e.: (2 cylinders for every patient) with anticoagulant, for

example citrate dextrose A to forestall the early actuation of the platelets.

- A specific tabletop gadget in an office in the outpatient center is utilized that is committed for computerized division of the PRP (platelet rich plasma) from the PPP (platelet helpless plasma) and the RBC's.
- Centrifugation is done as an initial turn for division of blood into plasma, white platelets and red platelets (1,000 rpm for 10 min) and a second turn for convergence of platelets and leukocytes (1, 500 rpm for 10 min) in PRP concentrate and PPP supernatant.
- A 50 ml of the patient's blood can yield 7-10 ml PRP.
- The upper bit of the plasma, compares to the platelet-helpless plasma (PPP), is gathered in a petri-dish to shape a gel subsequent to adding CaCl2 at 10% for 5-10 minutes that was utilized as an essential dressing with saline bandage later in the wake of infusing the forenamed ulcers.
- The lower bit of the plasma compares to PRP contains the concentrated platelets and a "buffy coat" layer containing a raised degree of leukocytes in a dainty layer of plasma.
- Both the concentrated platelets and the "buffy coat" are stacked in 7-10 insulin needles and are prepared for infusion of the ulcers.
- Prior to the infusion cycle, CaCl2 at 10% is added to help enactment of the platelet which is fundamental to start the ordinary physiological recuperating course at the site of tissue injury.
- Then, infusion is done at the mending edge and the floor of the ulcer with ensuing mechanical actuation for the platelets at the hour of infusion.
- The arranged PRP ought to be deliberately dealt with to dodge early actuation of the concentrated platelets that may prompt an arrival of development factors from the a-granules (degranulation) of the platelets.
- After-at that point, PPP gel is applied to the ulcer straightforwardly and the injury is covered with saline bandage.

• The measure is rehashed at regular intervals up to multiple times for every patient till accomplishing total recuperating of the tended to ulcers in the PRP arm of the investigation.

3.6 Primary endpoints

These include improvement in the total surface area of the ulcer and its rate of complete healing.

Secondary endpoints

The occurrence of infection and major limb amputations as well as assessment of the cost effectiveness of both modalities of treatment.

3.7 Statistical analysis

Information were coded and entered utilizing the measurable bundle SPSS (Statistical Package for the Social Sciences) rendition 25. Information was summed up utilizing mean, standard deviation, middle, least and greatest in quantitative information and utilizing recurrence (tally) and relative recurrence (rate) for absolute information. Correlations between quantitative factors were finished utilizing the non-parametric Mann-Whitney test (1). For looking at absolute information, Chi square (2) test was performed. Definite test was utilized rather when the normal recurrence is under 5 (2). Endurance bends were plotted by the Kaplan-Meier technique and analyzed utilizing the log-rank test.

4. Results

4.1Demographic data

- 1. Age
 - Group A: 50 60 with mean $54.90 \pm (SD) 2.37$.
 - Group B: 50- 60 with mean $54.75 \pm (SD) 3.75$.

2. Gender

- **Group A:** 35 patients (70%) were males, and 15 patients (30%) were females.
- **Group B:** 40 patients (80%) were males, and 10 patients (20%)were females.

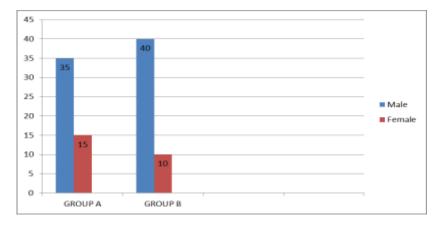
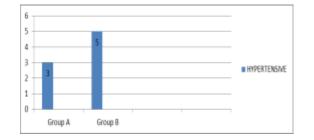


Fig (1) Showing gender in both groups.

1. Comorbidities

- a. All patients in both groups were diabetic and medically controlled.
- b. All patients in both groups had no history of stroke or ischemic heart disease.
- c. Hypertension.
 Group A: 3 patients (6%) were hypertensive.
 Group B: 5 patients (10%) were hypertensive.





2. Clinical examination prior to treatment

- a. All patients in both groups had pedal pulses.
- b. Patients who had previous bypass surgery for revascularization.

Group A: 8 patients (16%) had bypass surgery. **Group B:** 5 patients (10%) had bypass surgery.

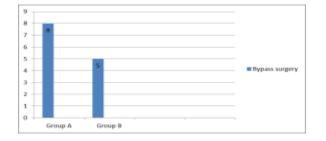
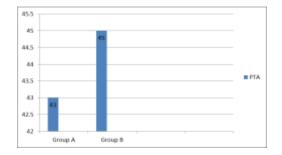


Fig (3) Showing number of patients who had bypass surgery in both groups.

3. Patients who had PTA ± stenting

Group A: 43patients had PTA (86%).

Group B: 45 patients had PTA (90%).



4. Site of ulcer

Fig (4) Showing number of patients who had PTA in both groups.

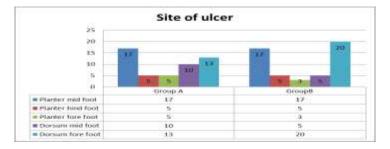


Fig (5) Showing different sites of ulcer in both groups.

4.2 Primary outcome Size of the ulcer

Table (1) Showing size of the ulcer represented by surface area in cm².

	Group										
	Group A (activated PRP)						Group B (standard dressings)				
	Mean	SD	Median	Minimum	Maximum	Mean	SD	Median	Minimum	Maximum	
Surface area	15.15	5.64	20.00	9.00	21.00	14.50	5.57	14.50	9.00	20.00	0.385
Follow up 1 w Surface area	11.65	5.04	14.00	6.00	20.00	14.50	5.57	14.50	9.00	20.00	0.002
Follow up 2 w Surface area	9.05	3.81	12.00	5.00	16.00	11.35	5.23	12.50	6.00	20.00	0.001
Follow up 3w Surface area	4.90	3.42	6.00	2.00	16.00	10.89	4.95	9.00	6.00	16.00	< 0.001
Follow up 4w Surface area	2.25	2.44	2.00	1.00	12.00	9.50	4.59	9.00	4.00	16.00	< 0.001
Follow up 5w Surface area	1.64	1.47	1.00	1.00	6.00	5.78	3.79	5.50	1.00	12.00	< 0.001
Follow up 6w Surface area	2.00	.00	2.00	2.00	2.00	2.78	1.96	2.00	1.00	6.00	1.000
Follow up 7w Surface area	1.00	.00	1.00	1.00	1.00	1.11	.52	1.00	.50	2.00	1.000
Follow up 8w Surface area	.50	.00	.50	.50	.50	.65	.24	.50	.50	1.00	0.529
Follow up 9w Surface area	•					.50	.00	.50	.50	.50	

It was observed that P value was statistically significant at the first five weeks while there was no difference detected between the groups starting from the sixth to the ninth week.

1.Time of complete healing.

We observed ≥ 50 % reduction in total surface area of the size of the ulcer in group A occurred at 2.5 weeks with a mean 7.57 ±SD 2.93 cm² compared to 4.5 weeks with a mean 7.25±SD 4.19 cm² in group B, with statistically significant P value < 0.001.

Table (2) Means and Medians for healing time.

While \geq 90 % reduction in total surface area of the size of the ulcer in group A occurred at 5 weeks with a mean 1.64 ±SD 1.47 cm² compared to 7 weeks with a mean 1.11±SD 0.52 cm² in group B, with statistically significant P value < 0.001.

From the previous data we can conclude that the complete healing rate was 96% (n=48) achieved in the sixth week for group A being a cutoff point while it was by the ninth week for 78% (n=39) cases with a significant P value < 0.01.

Group		Mean time	of healing	Median				
-	Estimate	Std. Error	95% Confidence Interval		Estimate	Std. Error	95% Confidence Interval	
			Lower Bound	Upper Bound	-		Lower Bound	Upper Bound
Group A (activated PRP)	5.700	.144	5.418	5.982	6.000	.069	5.865	6.135
Group B (standard dressings)	8.944	.119	8.711	9.178	9.000	.139	8.728	9.272
Overall	7.237	.209	6.827	7.647	6.000	.436	5.146	6.854

Table (3) Overall comparison between healing of ulcer in the 2 groups.

	Chi-Square	Df	P value
Log Rank (Mantel-Cox)	59.673	1	< 0.001
 Secondary outcome 1. Infection Group A: Five patients (10%) had superficial wound infection during the whole course of follow up. 	infection during from superfici	the whole co al infection u	tients (46%) had wound purse of follow up ranging up to deep infection and statistically significant <

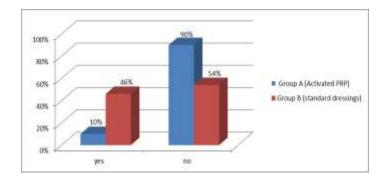


Fig.6 Infection in both groups.

2. Amputation

Group A: No major amputations were recorded in this arm.

 Table (4) Cost effectiveness in both groups.

Group B: Only five cases (10%) that had deep infection merged to extensive surgical debridement.

3. Cost effectiveness

	Group											
	Group A (activated PRP)							Group B (standard dressings)				
	Mean	SD	Median	Minimum	Maximum	Mean	SD	Median	Minimum	Maximum		
cost (\$)	247.50	29.85	250.00	200.00	300.00	437.50	70.48	400.00	400.00	600.00	< 0.001	

Concerning cost effectiveness, the mean cost for group A was 4455 Egyptian pound (247.50 \$) while the mean cost for group B was 7875 Egyptian pound (437.50 \$).

5.Discussion

Skin ulceration is a typical clinical issue. Because of populace maturing and the expansion of danger factors for atherosclerosis, for example, tobacco, weight and diabetes, there is an unmistakable pattern to the expansion danger of persistent ulcers. The social and monetary impacts are inescapable. The European Union allots 2 % of its wellbeing yearly financial plan for treating wounds, while a British report exhibited that 4 months of wandering consideration with a few treatments cost somewhere in the range of 200 and 2,000 pounds and that 40 million pounds a year go to just treating these injuries. It is assessed that, in the United States, costs identified with care of patients with pressure ulcer are above 1.3 billion dollars per year [3].

As of now, the U.S. Food and Drug Administration (FDA) affirm the utilization of PRP gel, under the management of a medical services Professional. The PRP gel is reasonable for oozing injuries, for example, leg ulcers, pressure ulcers, and diabetic ulcers and for the administration of precisely or carefully debrided twisted, in the U.S [4].

Platelet-rich plasma (PRP) is characterized as a part of the plasma portion of autologous blood having a platelet fixation above gauge. PRP likewise has been alluded to as platelet-advanced plasma, platelet-rich concentrate, and autologous platelet gel. PRP have been utilized to treat wounds since 1985 [5].

For over 20 years, PRP gel has been utilized for animating injury recuperating. Autologous PRP is created by cytokines, development factors, chemokines and fibrin framework got from a patient's blood. The system of activity for PRP gel is believed to be the subatomic and cell enlistment of ordinary injury mending reaction like that seen with platelet actuation [6].

This examination was directed to assess the viability of PRP in advancement of recuperating of constant ulcers, other than its preventive activity, which lessens removal rates and careful impedance. Based on the most recent 10 years of examination, the consequences of the precise audit with meta-investigation distributed via Carter et al. recommend that PRP treatment can emphatically affect wound mending and related factors, for example, agony and disease in both constant and intense cutaneous injuries [6].

A ulcer to be delegated persistent or intense, there ought to be a clear lucidity concerning a firm portrayal of the time span to outline chronicity. As no unmistakable agreement in the writing, most concurred that an intense ulcer ought to mend in a span of under multi month, while in a large portion of headstrong ulcers, a half year's term or considerably more is a satisfactory depiction [7].

In our examination, all cases were diabetic. Other comorbidities present in our investigation as presence of mechanical factor in a large portion of the patients (80%), Hypertension (HTN) in 13% of the patients.

Shockingly we found that DM and mechanical factor are inconsequential in connection of pace of recuperating. Generally in light of the fact that all diabetic patients remembered for our investigation should be controlled with clinical treatment prior to being remembered for the examination, and we keep patients with pressure ulcers from utilizing their appendages so no pressing factor showed at ulcers during treatment. These are starter clarifications with no logical proof.

No confusions occurred in our investigation, and few patients felt some torment during infusion.

Various components assume a part in injury mending, so it's anything but a solitary one-player show. It is probably going to be an intricate transaction multidiscipline amicability measure. These components incorporate diabetes, fringe blood vessel sickness, vasculitis disorder, actual pressing factor, and maturing [8]. Neighborhood wound variables including bacterial colonization, low thickness of helpful cytokines, significant levels of incendiary cytokines, and irregularity in framework metalloproteinases are constantly noted to draw out injury recuperating, making an endless loop for the injury to mend [8].

Subsequently, during treatment of such convincing issue, every one of these components ought to be drawn closer for victories. However, a few injuries may stay unmoved to the standard ordinary treatment, for example dressings, careful debridement and even skin joining as these modalities can't uphold these injuries with development factors (GF) expected to tweak the mending cycle [9].

Explores contributed with novel injury treatments like development factor items, negative pressing factor wound treatment (NPWT), parasite treatment, bioengineered tissue or skin substitutes, and hyperbaric oxygen treatment to help the mending cycle in this stubborn armamentarium [10].

Among these confident devices, autologous platelet rich plasma (PRP) is viewed as a reasonable technique in regarding non-mending ulcers as they are considered as a hotspot for development factors that can improve wound recuperating [11].

Platelets are wealthy in cytokines, which help fix tissue harm by invigorating multiplication, relocation, and separation of dermal fibroblasts and through neoangiogenesis [12]. After tissue injury, α - granules inside the platelets tie with the platelet plasma layer and delivery their substance into the environmental factors. Significant cytokines for wound mending, including platelet-determined development factor, (PDGF) changing development factor-b, (TGF) vascular endothelial development factor, (VEGF), fundamental fibroblast development factor, (BFGF) and epidermal development factor, (EGF) are delivered during this cycle [13]. These development factors invigorate the focused on cells, and furthermore interface with one another and enlarge their capacities [14].

Platelets play in injury recuperating an additional part as they fill in as a cross section of fibrin that frames a natural framework. That controls the mesenchymal cells to move from the base and the edges of the injury, at long last developing into a granulation tissue [15]. They additionally contain antibacterial proteins, which lessen bacterial colonization [16]. This is a solitary – focus, planned, randomized, controlled investigation that was directed in General Surgery Department, Faculty of Medicine, Benha University incorporating 100 patients giving constant leg ulcers and introduced to us between January 2018 to June 2018. This examination was completed in 100 patients with ongoing leg ulcers was remembered for this investigation, patients were arbitrarily separated in two equivalent gatherings A &B.

- Group A: incorporates 50 patients, which PRP was utilized upon their injuries.
- Group B: incorporates 50 patients with conventional strategies for dressing.

All cases had an insignificant debridement before treatment. Here in, our examination the segment information for example age, sex, DM, HTN were homogenously appropriated between the two gatherings.

All cases in the two gatherings were non-ischemic after effective revascularization either by or ER. The vast majority of the cases had the ulcer found principally on the grower mid foot (n=17 for bunch A just as gathering B), dorsum forefoot (n=13 for bunch A, n= 20 for bunch B).

In our examination, we determined the surface territory for the ulcer in the two gatherings utilizing a notable numerical recipe and was communicated in cm2 unit i.e.: the underlying mean surface zone in gathering A was $15.15 \pm SD5.64 \text{ cm } 2$ while the underlying mean surface region in gathering B $14.5 \pm SD 5.57 \text{ cm2}$.

We began the treatment for the two gatherings after full vascular appraisal. Patients were encouraged to go to at our unit at planned meetings consistently as long as 12 weeks, where in every meeting the complete surface region of the ulcer was estimated precisely and contrasted with sequential photography with the past outcomes for every patient. The photos were taken by an all around prepared assessor who was blinded for the technique for treatment in the two gatherings.

The focused on ulcers in gathering A were infused intermittently at regular intervals in the mending edge and the floor of the ulcer up to multiple times throughout the given treatment. While patients in gathering B were told to utilize soggy dressings with or without collagenase salve if necessary.

We noticed ≥ 50 % decrease in all out surface territory of the size of the ulcer in gathering A happened at 2.5 weeks with a mean 7.57 ±SD 2.93 cm2 contrasted with 4.5 weeks with a mean 7.25±SD 4.19 cm2 in gathering B, with genuinely huge P esteem < 0.001. Shockingly ≥ 90 % decrease in complete surface region of the size of the ulcer in gathering A happened at 5 weeks with a mean 1.64 ±SD 1.47 cm2 contrasted with 7 weeks with a mean 1.11±SD 0.52 cm2 in gathering B, with genuinely huge P esteem < 0.001.

In an investigation performed by Kontopodis et al. (17), they expressed a decrease >50% in ulcer zone in their treated gathering with PRP (bunch A) in 86% cases (n= 36/42) versus 73% cases (n=22/30) in the gathering who got a standard consideration (bunch B), this distinction was not measurable critical (P = 0.23).

Likewise Rate of ulcer region decrease >90% was essentially higher in gathering A. It merits referencing that the pace of complete recuperating for ulcers in gathering A was accomplished in 23 patients (46%) at the fifth week, while 25 patients (half) were mended totally by the 6th week and just two patients (4%) recuperated in the 10th week. 28 %(n=14) cases in gathering B indicated total mending rate by eighth week and half (n=25) cases were recuperated by 10th week while 22% (n=11) cases were recuperated in the 10th week.

P esteem was measurably critical < 0.001 between the gatherings.

From the past information we can presume that the total recuperating rate was 96% (n=48) accomplished in the 6th week for bunch A being a cutoff point while it was by the 10th week for 78% (n=39) cases with a critical P esteem < 0.01. Just five cases (10%) that had profound contamination converged to broad careful debridement.

P esteem was genuinely huge and rises to 0.041.

Along these lines, the appendage rescue rate was 100% in gathering A rather than 90% in gathering B with huge P esteem equivalent 0.041. Kakagia et al. [18] demonstrated Mean mending rate fundamentally higher with PRP than without PRP in a randomized forthcoming examination directed on 51 diabetic patients.

6. Conclusion

- The utilization of PRP in restorative cycles identified with Plastic, Reconstructive and Esthetic Surgery is right now a genuine alternative in generally open and private medical clinics in numerous nations; circumstance that shows the advances that have happened over ongoing years in tissue recovery treatment.
- We are focussing here on the territory of regenerative tissue therapy to be specific in therapy of ongoing ulcers, however we ought not fail to remember the gigantic potential that PRP could have as a haemostatic, mitigating and pain relieving substance.
- The fantastic outcomes appeared in late investigations have just served to affirm the great results depicted in our examination. In ten patients that we treated, the outcomes acquired after fulfillment of the uses of PRP were acceptable or phenomenal.
- PRP is an incredible weapon for treating persistent ulcers, giving mending, diminishing disease rates, other than its preventive activity, which decreases removal rates.
- It is imperative to feature that PRP treatment needs new multi driven, randomized, twofold visually impaired examinations to improve the degree of logical proof of the genuine advantage of this promising treatment, indicating the immediate and circuitous impacts (paracrine impact) of PRP in mending of persistent injuries. Other than that, wellbeing of this treatment in a long haul ought to likewise be illustrated. These outcomes can

contribute for the meaning of sort of advantage with this treatment for the few kinds of ulcers.

References

- Y.H.Chan. Biostatistics102: Quantitative Data Parametric & Non-parametric Tests. Singapore Med J, Vol.44(8),pp.391-396,2003.
- [2] Y.H.Chan. Biostatistics 103: qualitative data-tests of independence. Singapore Med J, Vol.44(10), PP.498-503,2003.
- [3] E.Anitua, M.Sánchez, G.Orive. The importance of understanding what is platelet-rich growth factor (PRGF) and what is not. J Shoulder Elbow Surg, Vol.20, PP.e23-e24,2011.
- [4] S.S.Gale, F.Lurie, T.Treadwell. Dominate Wounds. Wounds: a compendium of clinical research and practice, Vol.26(1),pp.1,2014.
- [5] R.E.Marx. Platelet-rich plasma (PRP): what is PRP and what is not PRP? Implant Dent, Vol.10(4), PP.225–228,2001.
- [6] M.J.Carter, C.P.Fylling, W.W.Li. Analysis of run-in and treatment data in a wound outcomes registry: clinical impact of topical platelet-rich plasma gel on healing trajectory. International wound journal, Vol.8(6), PP.638-650,2011.
- [7] C.P.Minniti, J.Eckman, P.Sebastiani. Leg ulcers in sickle cell disease. American journal of hematology, Vol.85(10), PP.831-833,2010.
- [8] L.M.Rappl. Effect of platelet rich plasma gel in a physiologically relevant platelet concentration on wounds in persons with spinal cord injury. International wound journal, Vol.8(2), PP.187-195,2011.
- [9] S.Sarvajnamurthy, S.Suryanarayan, L.Budamakuntala. Autologous platelet rich plasma in chronic venous ulcers: study of 17 cases. Journal of cutaneous and aesthetic surgery, Vol.6(2),pp.97,2013.
- [10] D.Baltzis, I.Eleftheriadou, A.Veves. Pathogenesis and treatment of impaired wound healing in diabetes mellitus: new insights. Advances in therapy, Vol.31(8), PP.817-836,2014.
- [11] D.H.Suresh, S.Suryanarayan, S.Sarvainamurthy. Treatment of a Non-healing diabetic foot ulcer with platelet rich plasma. J Cutan Aesthet Surg, Vol.7(4), PP.229–31,2014.
- [12] S.Mehta, J.T.Watson. Platelet rich concentrate: basic science and current clinical applications. J Orthop Trauma, Vol.22(6), PP.432–438. ,2008.
- [13] C.Y.Su, Y.P.Kuo, H.L.Nieh. Quantitative assessment of the kinetics of growth factors released from platelet gel. Transfusion, Vol.48(11),pp.2414– 20,2008.
- [14] S.P.Bennett, G.D.Griffiths, A.M.Schor. Growth factors in the treatment of diabetic foot ulcers. British Journal of Surgery, Vol.90(2), PP.133-146,2003.
- [15] D.H.Kim, J.Y.Kim, S.H.Seo. Recalcitrant cutaneous ulcer of comorbid patient treated with platelet rich plasma: A case report. Journal of Korean medical science, Vol.27(12), PP.1604-1606,2012.

- [16] T.M.Bielecki, T.S.Gazdzik, J.Arendt. Antibacterial effect of autologous platelet gel enriched with growth factors and other active substances: an in vitro study. J Bone Joint Surg Br, Vol.89(3), PP.417–420,2007.
- [17] N.Kontopodis, E.Tavlas, G.Papadopoulos Effectiveness of Platelet-Rich Plasma to Enhance Healing of Diabetic Foot Ulcers in Patients With Concomitant Peripheral Arterial Disease and Critical Limb Ischemia: The International Journal of Lower Extremity Wounds, Vol. 15(1) 45–51,2016.
- [18] D.D.Kakagia, K.J.Kazakos, K.C.Xarchas. Synergistic action of protease-modulating matrix and autologous growth factors in healing of diabetic foot ulcers. A prospective randomized trial. Journal of Diabetes and its Complications, Vol.21(6),pp.387-391, 2007.
- [19] V.R.Driver, J.Hanft, C.P Fylling. Autologel Diabetic Foot Ulcer Study Group. A prospective, randomized, controlled trial of autologous plateletrich plasma gel for the treatment of diabetic foot ulcers. Ostomy Wound Manage, Vol.52(6), PP.68-74,2006.
- [20] G. Saldalamacchia, E. Lapice, V. Cuomo. A controlled study of the use of autologous platelet gel for the treatment of diabetic foot ulcers. Nutrition, Metabolism and Cardiovascular Diseases, Vol.14(6), PP.395-396,2004.

- [21] G.Q. Shan, Y.N. Zhang, J. Ma. Evaluation of the effects of homologous platelet gel on healing lower extremity wounds in patients with diabetes. The international journal of lower extremity wounds, Vol.12(1), PP.22-9 ,2013.
- [22] N. Kontopodis, E. Tavlas, G. Papadopoulos. Effectiveness of Platelet-Rich Plasma to Enhance Healing of Diabetic Foot Ulcers in Patients With Concomitant Peripheral Arterial Disease and Critical Limb Ischemia: The International Journal of Lower Extremity Wounds, Vol.15(1), PP.45–51,2016.
- [23] R.Serra, G. Buffone, A. Dominijanni. Application of platelet-rich gel to enhance healing of transmetatarsal amputations in diabetic dysvascular patients. International wound journal, Vol.10(5), PP.612-615,2013.
- [24] D.L. Villela, V.L.C. Santos. Evidence on the use of platelet-rich plasma for diabetic ulcer: a systematic review. Growth factors,vol.28(2), PP.111-116,2010.
- [25] F.DeLalla, G. Pellizzer, M. Strazzabosco. Randomized prospective controlled trial of recombinant granulocyte colony-stimulating factor as adjunctive therapy for limb-threatening diabetic foot infection. Antimicrobial agents and chemotherapy, Vol.45(4), PP.1094-1098. ,2001.