

PRF-From self to self.

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Short Communication

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PRP**ABSTRACT**

PRF, which belongs to a new second generation of platelet concentrates, with simplified processing, and not requiring biochemical blood handling, has several advantages over traditionally prepared PRP, which has been widely used for accelerating soft tissue and hard tissue healing. However, the preparation being strictly autologous, the amount of PRF obtained is limited. Choukroun's platelet-Rich Fibrin (PRF) incorporates leucocytes, platelets and a wide range of healing proteins within a dense fibrin matrix. It has a strong fibrin architecture and slow releasing growth factors and glycoproteins over several days. It is a natural bioactive membrane, which can enhance soft/hard tissue healing, at the same time, can also protect surgical sites, grafted materials from external aggressions. This article describes the evolution of this second generation platelet concentrate and its multiple uses in various surgical procedures. We have attempted to give an overview of PRF including its uses, advantages, its preparation & its microscopic structures.

INTRODUCTION

Periodontal diseases are among the most prevalent diseases worldwide. They are the major cause of tooth loss in adults [1]. The goal of periodontal therapy includes not only the arrest of periodontal disease progression, but also the regeneration of structures lost due to disease. One of the most important and, at present, unsolved problems in clinical periodontics is the predictable successful treatment of periodontitis affected furcations of multi-rooted teeth [2]. Since several therapeutic approaches that involve conservation, resection or regeneration are proposed, a proper diagnosis of these lesions is demanding. To overcome the prevailing healing limitations in furcation defects, the principles of tissue engineering were applied using a purified growth factor together with an osteoconductive scaffold to stimulate the patient's own cells toward a regenerative response. More recently, the use of growth factors and bone morphogenic proteins (BMPs) have shown promising results in the treatment of intra-bony defects. The use of fibrin glue [1] or platelet concentrate [2,3] during periodontal surgical procedures is one of the current treatment concepts used to accelerate wound healing and tissue maturation [4]. Choukroun's platelet-rich fibrin (PRF) a 2nd generation platelet concentrate [5], was defined as an autologous leukocyte and platelet-rich fibrin biomaterial [6,7,8]. PRF was developed in France by Choukroun et al [9]. In 2001. It is biocompatible, bioresorbable and plays an essential role in wound repair, not only for hemostasis but also provides a matrix for migration of tissue-forming cells like fibroblasts and endothelial cells, which are involved in angiogenesis and that are responsible for remodeling of the new tissue. In the normal wound-healing process, platelets are trapped within the fibrin matrix and are subsequently activated so that growth factors like platelet-derived growth factor (PDGF), transforming growth factor (TGF- β) and insulin-like growth factor I (IGF-I) are set free [10], which could stimulate the mitogenic response of the periosteum during bone [11]. The essence of platelet-rich fibrin (PRF) modified by Choukroun and colleagues [5] is a fibrin matrix in which the platelet cytokines and cells are trapped and may be delivered after a certain time [12].

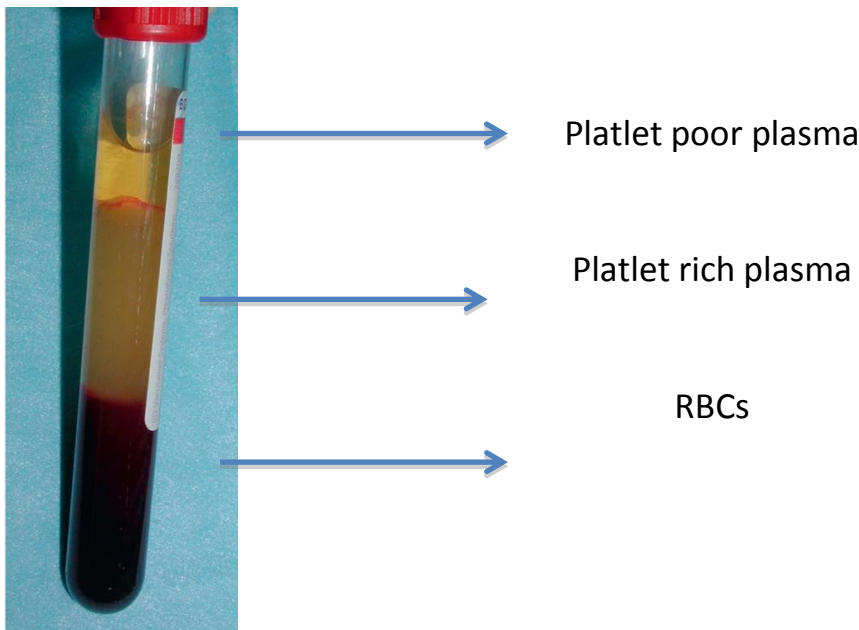
Definition of PRF

PRF can be considered as an autologous healing biomaterial, incorporating in a matrix of autologous fibrin most leukocytes, platelets & growth factors harvested from a simple blood sample [5].

PRF preparation

A blood sample of 10 mL in test tubes without an anticoagulant is centrifuged using a tabletop centrifuge machine for 12 min at 2500 rpm or 10 min at 3000 rpm. The resultant product which is obtained after centrifugation could be seen in 3 distinct layers, a red blood cell (RBC) base at the bottom, a cellular plasma (platelet-poor plasma [PPP]) as a supernatant, and a PRF clot in the middle. It can be used directly as a clot or after compression as a membrane (Fig. 1).

Figure 1



PRF: as seen microscopically (Figure 2)

The PRF clot can be described as composed of two main parts observable with the naked eye: a fibrin yellow portion, constituting the main body, and a red portion located at the end of the clot (full of RBCs). Between these two areas, a whitish layer called the “buffy coat” (similar to the whitish layer in PRP technologies) can be observed with the naked eye and concentrates cell corpuscles requiring identification. The PRF clot at a low magnification showed that the clot presented a concavity in its middle part. This is caused by matrix shrinkage due to fixation. In the red part of the PRF clot, RBCs are enmeshed in the fibrin network. RBC shapes are normal, but the fibrin-strand network appears immature. At the junction between the red and yellow parts of the PRF clot (the buffy coat area), the SEM examination showed leukocytes that clearly appeared as spherical structures with irregular surface (Figure 3) [13]. Platelet aggregates appeared very clearly along the fibrin strands. Beyond the buffy coat base, two distinguished different areas: the first area is composed of thick fibrin strands and a few scattered RBCs (probably from contamination during clot handling). The fibrin network appeared to be mature. The second area corresponded to the platelet veins. This area contained platelets and fibrin that formed large and dense clusters due to extensive aggregation and clotting. This aggregate formed a solid and thick mesh. Therefore, platelets seemed to be highly activated during the PRF-preparation protocol. At a low magnification, the PRF membrane surface showed the print of the gauze threads. Fibrin is a physiologic glue; therefore, the compression of the fibrin clot into a membrane provided a very compact matrix. In the fibrin, one end of the membrane is clearly organized in parallel strands that appeared very thick and dense. It is impossible to distinguish cellular elements trapped within this condensed network.

Figure 2

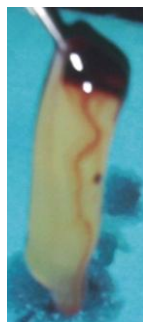
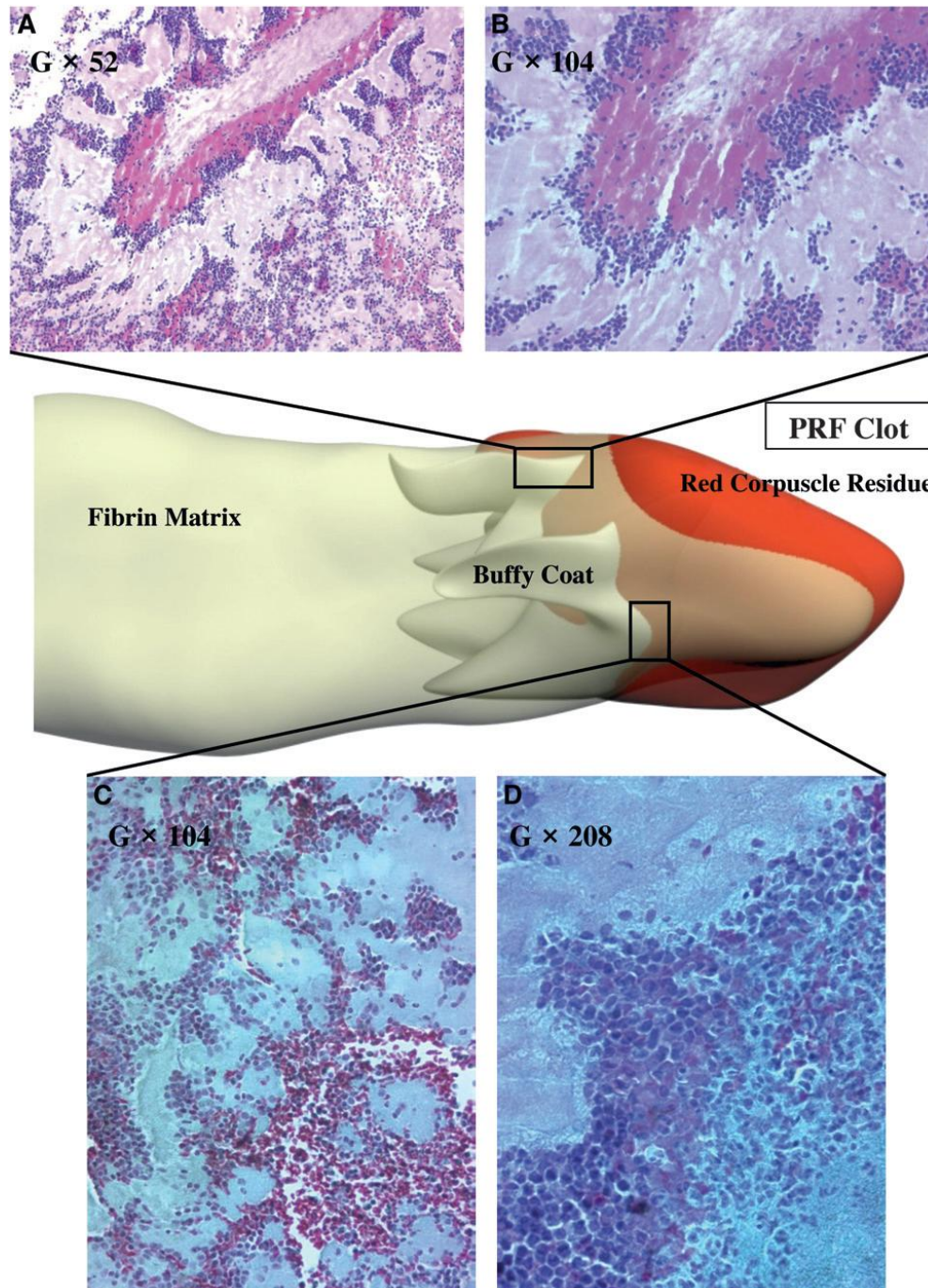


Figure 3



Saluja H, Dehane V, Mahindra U. Platelet-Rich fibrin: A second generation platelet concentrate and a new friend of oral and maxillofacial surgeons. *Ann Maxillofac Surg* 2011;1:53-7.

Distribution of cells in PRF

The highest platelet/leukocyte density is found in the first millimeter of the yellow clot, just after the red clot. The platelet/leukocyte distribution becomes increasingly scarce as they move away from the red clot, platelets or leukocytes beyond the first half of the yellow clot is not seen. In the first 2 mm located beyond the yellow/red border, the platelet/leukocyte distribution is homogeneous throughout the clot width.

Uses of PRF

- For the treatment of 2 and 3 wall infrabony defects.
- Grade II and grade-III furcation involvements.

- In the treatment of miller’s class-I and class-II gingival recession.
- For the improvement of soft tissue healing [14,15,16]
- For bone graft protection and remodeling
- Often mixed with graft materials.
- Socket preservation [17,18,19]
- It is also useful for Schneiderian membrane protection
- As a sole osteoconductive filling material during a sinus-lift
- In transfusion of PRP.
- In prosthodontics PRF can serve as a resorbable membrane that can be used in pre-prosthetic surgery as well as in implantology to cover bone augmentation sites [20].

Table 1: The advantages of Platelet-rich fibrin over Platelet-rich plasma and disadvantages of Platelet-rich fibrin

Advantages of PRF over PRP	Disadvantages of PRF
Ease of preparation/application	Amount available is low, because of autologous blood
No biochemical handling of blood required	Quick handling of blood is needed, immediately after collection
Simplified and cost effective process	
Use of bovine thrombin and anticoagulants not required. Thus, the use of vital fibrin as an autologous scaffold for periosteal cell or stem cell transplantation and consequently for bone tissue engineering is an obvious option	
Favorable healing due to slow polymerization	
More efficient cell migration and proliferation	
PRF has supportive effect on immune system	
PRF helps in haemostasis	
Contains large quantity of platelet and leukocyte cytokines	
Powerful healing potential on both soft & hard tissues.	

CONCLUSION

Thus, with this article we can conclude that the new and recent generation of platelet concentrate-PRF, would be a good friend to Periodontists in the near future. It has a list of its benefits, & intraoral applications. This material is already being used widely in France, and considering its advantages, its popularity should increase here too. More clinical, histological and statistical studies are now required to understand the benefits of this new platelet concentrate better. However, it cannot be ignored that since it is obtained from an autologous blood sample, the quantity of PRF produced is low and only a limited volume can be used. This fact limits the systematic utilization of PRF, as in general surgery. Also though the potential applications of PRF are broad, however, an accurate working knowledge of the biomaterial, its biology, efficiency & limits are necessary to optimize its use in daily practice. Hence additional randomized clinical trials evaluating the use & performance of PRF are warranted.

REFERENCES

1. Gibble JW, Ness PM. Fibrin glue: The perfect operative sealant? *Transfusion*. 1990;30:741-747.
2. Marx RE, Carlson ER, Eichstaedt RM, Schimmele SR, Strauss JE, Georgeff KR. Platelet-rich plasma: Growth factor enhancement for bone grafts. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod*. 1998; 85:638-646.
3. Dohan DM, Choukroun J. PRP, cPRP, PRF, PRG, PRGF, FC. . . How to find your way in the jungle of platelet concentrates? *Oral Surg Oral Med Oral Pathol Oral Radiol Endod*. 2007;103:305-306.
4. Man D, Plosker H, Winland-Brown JE. The use of autologous platelet-rich plasma (platelet gel) and autologous platelet-poor plasma (fibrin glue) in cosmetic surgery. *Plast Reconstr Surg*. 2001;107:229- 237.
5. Dohan DM, Choukroun J, Diss A, et al. Platelet-rich fibrin (PRF): A second-generation platelet concentrate. Part I Technological concepts and evolution. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod*. 2006;101:e37-e44.
6. Dohan Ehrenfest DM, Rasmusson L, Albrektsson T. Classification of platelet concentrates: From pure platelet-rich plasma (P-PRP) to leucocyte- and platelet- rich fibrin (L-PRF). *Trends Biotechnol*. 2009;27: 158-167.
7. Dohan DM, Choukroun J, Diss A, et al. Platelet-rich fibrin (PRF): A second-generation platelet concentrate. Part II: Platelet-related biologic features. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod*. 2006; 101:e45-e50.
8. Dohan DM, Choukroun J, Diss A, et al. Platelet-rich fibrin (PRF): A second-generation platelet concentrate. Part III: Leucocyte activation: A new feature for platelet concentrates? *Oral Surg Oral Med Oral Pathol Oral Radiol Endo*. 2006;101:e51-e55.

9. Choukroun J, Adda F, Schoeffler C, Vervelle A. Anopportunity in perio-implantology: The PRF (in French). *Implantodontie*. 2001;42:55-62.
10. Schliephake H. Bone growth factors in maxillofacial skeletal reconstruction. *The Int J Oral Maxillofac Surg*. 2002;31: 469- 484.
11. Gruber R, Karreth F, Frommlet F, Fischer MB, Watzek G. Platelets are mitogenic for periosteum-derived cells. *J Orthopaedic Res*. 2003;21: 941-948.
12. Mosesson MW. Fibrinogen and fibrin structure and functions. *J Thromb Haemostasis*. 2005;3: 1894-1904.
13. Saluja H, Dehane V, Mahindra U. Platelet-Rich fibrin: A second generation platelet concentrate and a new friend of oral and maxillofacial surgeons. *Ann Maxillofac Surg*. 2011;1:53-7.
14. Choukroun J, Diss A, Simonpieri A, et al. Platelet-rich fibrin (PRF): A second-generation platelet concentrate. Part IV: Clinical effects on tissue healing. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod*. 2006;101:e56-e60.
15. Saadoun AP, Touati B. Soft tissue recession around implants: Is it still unavoidable? - Part II. *Pract Proced Aesthet Dent*. 2007;19:81-87.
16. Del Corso M, Sammartino G, Dohan Ehrenfest DM. Choukroun's platelet-rich fibrin membranes in periodontal surgery: Understanding the biomaterial or believing in the magic of growth factors? (letter to the editor). *J Periodontol*. 2009;80:1694-1697; author reply 1697-1699.
17. Choukroun J, Diss A, Simonpieri A, et al. Platelet-rich fibrin (PRF): A second-generation platelet concentrate. Part V: Histologic evaluations of PRF effects on bone allograft maturation in sinus lift. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod*. 2006;101:299-303.
18. Simonpieri A, Del Corso M, Sammartino G, Dohan Ehrenfest DM. The relevance of Choukroun's platelet rich fibrin and metronidazole during complex maxillary rehabilitations using bone allograft. Part I: A new grafting protocol. *Implant Dent*. 2009;18:102-111.
19. Simonpieri A, Del Corso M, Sammartino G, Dohan Ehrenfest DM. The relevance of Choukroun's platelet rich fibrin and metronidazole during complex maxillary rehabilitations using bone allograft. Part II: Implant surgery, prosthodontics and survival. *Implant Dent*. 2009;18:220-229.
20. Choukroun J, Diss A, Simonpieri A, Girard MO, Schoeffler C, Dohan SL, et al. Platelet-rich fibrin (PRF): a second-generation platelet concentrate. Part V: histologic evaluations of PRF effects on bone allograft maturation in sinus lift. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod*. 2006b;101: 299-303.