



“ADMINISTRATION OF PLATELET RICH PLASMA- A SHIFT IN ACCELERATED ORTHODONTICS– A REVIEW “

Orthodontology

Dr. Saraa Angel .L* BDS.,MDS Consultant Orthodontist, Delhi. *Corresponding Author

Dr. Abhijeet Jaiswal

BDS.,MDS, Senior Resident,AIIMS, Raipur.

ABSTRACT

Increasing awareness towards self-appearance and aesthetics has led to increase in overall patients willing to undergo orthodontic treatment. Various surgical and non surgical methods have been tried to reduce the overall orthodontic treatment time. Painless, non invasive procedures are preferred over the traumatic methods. Platelet Rich Plasma (PRP) injection has recently gained popularity due to its safe, relatively painless and economical alternative in accelerating the tooth movement. This article briefs on the researches done in animal and human population so far using PRP and their cumulative effects on tooth movement.

KEYWORDS

PRP. Accelerated Orthodontics, Rate of tooth movement

INTRODUCTION:

Prolonged treatment time has always been an area of concern for both patient and orthodontists. In spite of the fact of losing interest and motivation to treatment, it also has various adverse effects like enamel demineralization¹, dental caries², periodontitis³, and orthodontically induced external root resorption⁴ due to prolonged inflammatory reaction and poor oral hygiene.

The average resorption rate of human osteoclasts is $\sim 0.6\text{mm}/\text{mo}$ ⁵ theoretically and clinically it is only $0.34\text{ mm}/\text{mo}$ ⁶. Hence the osteoclasts plays an important role in orthodontic tooth movement⁷. Various procedures have been attempted in inducing inflammation by increased insult to bone by surgical techniques like corticotomy⁸, periodontal ligament distraction⁹, periodontically accelerated osteogenic orthodontics(PAOO)¹⁰, surgery first approach¹¹, micro osteo-perforation¹² etc. Other non-surgical methods like local injection of cytokine or growth factors like prostaglandin E¹³, misoprostol¹⁴, Vitamin D¹⁵, PTH¹⁶, VEGF¹⁷, PDGF¹⁸ etc. were tried to mimic the body's inflammatory response. The limitations of these biomodulators are that these may need multiple administration and their efficacy in human population is yet to be proved.

Platelet rich plasma (PRP):

Platelet rich plasma (PRP) is the concentration of autologous platelets suspended in the plasma. The baseline platelet count in any individual will be around $200,000 \pm 75000/\mu\text{L}$ in whole blood and a prepared PRP must contain at least 1 million/ μL PRP in a 6 ml aliquot to have a "therapeutic value"¹⁹. Hence the working definition of PRP according to Marx^{19,20} is that it must contain 5 folds increase in platelet counts than the whole blood i.e the baseline counts.

Classification :

PRP can be classified as Pure PRP (without leukocytes) or Leukocyte rich PRP (With leukocytes and small amount RBC's)²¹. Often Platelet concentrate is misunderstood as PRP but it is not PRP as it is devoid of plasma²².

Processing of PRP:

Generally, PRP is prepared by "Double centrifugation method"¹⁹. The desired amount of blood is collected in a tube with an anticoagulant and subjected to the "Soft Spin", that sediments the RBC's at the bottom with the platelets and WBC's in the middle buffy coat layer and the Platelet poor plasma (PPP) at the top. The PPP along with the buffy coat is pipetted out and transferred to a new tube and the RBC's are discarded. The tube is resubjected to the "Hard Spin" to obtain platelets pelleted down in the PPP with small amount of RBC's and reasonable amounts of WBC's. The excess PPP is removed and platelet pellets are suspended in the small amount of plasma to get the PRP. The dilution can be altered to get the desired concentration of PRP.

According to the protocol documented by Liou²³, 60ml of blood was used to obtain PRP with 9.5 to 12.5 times the baseline count. Other studies have documented PRP preparation from only 5ml of blood. For

orthodontic purposes freshly prepared PRP is used and it is found to be sterile for almost 8 hours with viable growth factors^{19,21,22}. It is used either in activated or non-activated forms. Once activated using thrombin or calcium the PRP increases into a highly viscous gel platelet gel and releases a bolus of growth factors immediately. Studies suggests that almost 70 % of growth factors are released in 10 minutes and 95% of growth factors(GF) are released in one hour²⁴. Amount of calcium used, the source of thrombin and the method of activation determine their properties to an extent. Lacoste²⁵ revealed that the higher concentration of calcium inhibits the GF release. The source of thrombin if bovine may lead to spongiform encephalopathy, bleeding complications and even anaphylactic shock²⁶. While non-activated platelets tend to release growth factors in a more consistent manner during their life span of 5-7 days as they come in contact with soluble type 1 collagen²⁷.

Route of Injection:

PRP can be injected through various routes including intra-ligamentary²⁸, submucosally in the buccal and lingual mucosa and gingiva²³, and into the buccal vestibule²⁹. Smaller area, technique sensitivity, lack of proper landmarks and high resistance offered during injection are few drawbacks of intra-ligamentary approach but when administered correctly it is the most effective route³⁰.

Advantages :

PRP is advantageous as it is simple, safe and cost effective. They are autologous in origin and hence has no fear of immune reaction or disease transmission and hypersensitivity sequelae³¹. It has an antibacterial effect due to the presence of leukocytes hence limits post surgical inflammation³². The antibacterial activity may be due to high pH of 6.5 -6.7²⁰. In dentistry PRP is used more locally in minimal quantities hence systemic uptake related complications are minimal.

Limitations:

Highly variable platelet counts in every individual with a lack of single protocol that yields similar levels is not yet formulated. The other variables for uncertain yield of platelets are centrifugation speed, time, initial platelet counts, centrifugation kit (Manual /Automatic), FDA approved machinery availability, efficiency in pipetting, preservation and viability of platelets²⁰.

PRP in Dentistry:

The use of PRP as an Autologous Fibrin Adhesive was reported as early as 1994 by Tayapongsak¹⁹ in mandibular reconstruction and by Whitman in 1997³³. The positive response propelled the research of PRP due to its effects on cellular proliferation. PRP consists of various growth factors like, VEGF, TGF β_1 , TGF β_2 , IGF-1, FGF, PDGF AA, PDGF AB, PDGF BB, PDEGF²⁰, PF- 4, RANTES³² in their alpha granules along with other proteases, anti-proteases, and inflammatory mediators. Platelets have pro-inflammatory and anti-inflammatory action which is dose-dependent^{19,20,22}.

PRP in Orthodontics :Animal Studies:

Author	Sample	Sample Size	PRP concentration	Outcome
Gulec et al (AJODO)	Sprague Dawley rats	81	High PRP- 5 times the baseline	High PRP- 1.7 times increased
Rashid et al (Orthodontics and Craniofacial Research)	Mongrel Dogs	6	Moderate PRP – 2 the baseline Levels not mentioned	Moderate PRP – 1.4 times increased Tooth movement increased by two times PRP group
Erliera et al (Advances in health sciences in research)	Guinea pigs	24	Twice the baseline counts	No significant tooth movement in PRP group
Nakornnoi et al (KJO)	White Rabbits	23	7.5 times the baseline counts approximately	The rate of tooth movement increased the first 14 days but not significant after 14 days
Akbulut et al (AJODO)	Wistar male albino rats	48	5 times the baseline count	No significant tooth movement in PRP group

Human Studies:

Author	Sample size	Follow up	PRP concentration	Outcome
ElTimamy (Angle Orthod)	16 female patients	3 months	Not specified	15% increased rate in First month and 5% increased in Second month Rate in PRP side decreased by 40% in third month
Ahmad S,³⁸ Hassan H (IJAR)	28 patients	3 months	Not Specified	PRP side was 1 times faster in first month and 0.5 times faster in second month

CONCLUSION:

Literature suggests that PRP may have a transient acceleratory effect on the rate of tooth movement but only for a short period of duration and may need repeated administration to sustain the accelerating potential. Hence studies with uniform protocol and increased sample size are needed to support the judicious use of PRP in the future.

REFERENCES

- Miles, P. (2017), "Accelerated orthodontic treatment - what's the evidence?," *Aust Dent J*, Mar;62 Suppl 1:63-70.
- Jung, W.S., Kim, H., Park, S.Y., Cho, E.J., and Ahn, S.J. (2014), "Quantitative analysis of changes in salivary mutans streptococci after orthodontic treatment," *Am J Orthod Dentofac Orthop*, May;145(5):603-9.
- Corbacho de Melo, M.M., Cardoso, M.G., Faber, J., and Sobral, A. (2012), "Risk factors for periodontal changes in adult patients with banded second molars during orthodontic treatment," *Angle Orthod*, Mar;82(2):224-8.
- Brezniak, N., and Wasserstein, A. (2002), "Orthodontically induced inflammatory root resorption. Part II: The clinical aspects," *Angle Orthod*, Apr;72(2):180-4.
- Roberts, W.E. (1988), "Bone tissue interface," *J Dent Educ*, Dec;52(12):804-9.
- Roberts, W.E., Goodwin, W.C. Jr., and Heiner, S.R. (1998), "Cellular response to orthodontic force," *Dent Clin North Am*, Jan;25(1):3-17.
- Huang, H., Williams, R.C., and Kyrkanides, S. (2014), "Accelerated orthodontic tooth movement: molecular mechanisms," *Am J Orthod Dentofacial Orthop*, Nov;146(5):620-32.
- Chung, K.R., Oh, M.Y., and Ko, S.J. (2001), "Corticotomy-assisted orthodontics," *J Clin Orthod*, May;35(5):331-9.
- Kumar, P.S., Saxena, R., Patil, S., Keluskar, K.M., Nagaraj, K., and Kotrashetti, S.M. (2009), "Clinical investigation of periodontal ligament distraction osteogenesis for rapid orthodontic canine retraction," *Aust Orthod J*, Nov;25(2):147-52.
- Wilcko, W.M., Wilcko, T., Bouquet, J.E., and Ferguson, D.J. (2001), "Rapid orthodontics with alveolar reshaping: two case reports of decrowding," *Int J Periodontics Restorative Dent*, Feb;21(1):9-19.
- Liou, E.J., Chen, P.H., Wang, Y.C., Yu, C.C., Huang, C.S., and Chen, Y.R. (2011), "Surgery-first accelerated orthognathic surgery: postoperative rapid orthodontic tooth movement," *J Oral Maxillofac Surg*, Mar;69(3):781-5.
- Alikhani, M., Raptis, M., Zoldan, B., Sangsuwon, C., Lee, Y.B., Alyami, B., Corpodian, C., Barrera, L.M., Alansari, S., Khoo, E., and Teixeira, C. (2013), "Effect of micro-osteoperforations on the rate of tooth movement," *Am J Orthod Dentofacial Orthop*, Nov;144(5):639-48.
- Spielmann, T., Wieslander, L., and Hefti, A.F. (1989), "Acceleration of orthodontically induced tooth movement through the local application of prostaglandin (PGE1)," *Schweiz Monatsschr Zahnmed*;99(2):162-5.
- Sekhvat, A.R., Mousavizadeh, K., Pakshir, H.R., and Aslani, F.S. (2002), "Effect of misoprostol, a prostaglandin E1 analog, on orthodontic tooth movement in rats," *Am J Orthod Dentofacial Orthop*, Nov;122(5):542-7.
- Collins, M.K., and Sinclair, P.M. (1988), "The local use of vitamin D to increase the rate of orthodontic tooth movement," *Am J Orthod Dentofacial Orthop*, Oct;94(4):278-84.
- Soma, S., Matsumoto, S., Higuchi, Y., Takano-Yamamoto, T., Yamashita, K., and Kurisu, K. (2000), "Local and chronic application of PTH accelerates tooth movement in rats," *J Dent Res*, Sep;79(9):1717-24.
- Kohno, S., Kaku, M., Tsutsui, K., Motokawa, M., Ohtani, J., Tenjo, K., Tohma, Y., Tokimasa, C., Fujita, T., Kawata, T., and Tanne, K. (2003), "Expression of vascular endothelial growth factor and the effects on bone remodeling during experimental tooth movement," *J Dent Res*, Mar;82(3):177-82.
- Marden, L.J., Fan, R.S., Pierce, G.F., Reddi, A.H., and Hollinger, J.O. (1993), "Platelet-

- derived growth factor inhibits bone regeneration induced by osteogenin, a bone morphogenetic protein, in rat craniotomy defects," *J Clin Invest*, Dec;92(6):2897-905.
- Marx, R.E., Carlson, E.R., Eichstaedt, R.M., Schimmele, S.R., Strauss, J.E., and Georgeff, K.R. (1998), "Platelet-rich plasma: Growth factor enhancement for bone grafts," *Oral Surg Oral Med Oral Pathol Oral Radiol Endod*, Jun;85(6):638-46.
- Marx, R.E. (2004), "Platelet-rich plasma: evidence to support its use," *J Oral Maxillofac Surg*, Apr;62(4):489-96.
- Dohan Ehrenfest, D.M., Rasmusson, L., and Albrektsson, T. (2009), "Classification of platelet concentrates: from pure platelet-rich plasma (P-PRP) to leucocyte- and platelet-rich fibrin (L-PRF)," *Trends Biotechnol*, Mar;27(3):158-67.
- Marx, R.E. (2001), "Platelet-rich plasma (PRP): what is PRP and what is not PRP?" *Implant Dent*, 10(4):225-8.
- Liou, E.J. (2016), "The development of submucosal injection of platelet rich plasma for accelerating orthodontic tooth movement and preserving pressure side alveolar bone," *APOS Trends Orthod*, Jan 1;6:5-11.
- Jeon, Y.R., Jung, B.K., Roh, T.S., Kang, E.H., Lee, W.J., Rah, D.K., Lew, D.H., and Yun, I.S. (2016), "Comparing the Effect of Nonactivated Platelet-Rich Plasma, Activated Platelet-Rich Plasma, and Bone Morphogenetic Protein-2 on Calvarial Bone Regeneration," *J Craniofac Surg*, Mar;27(2):317-21.
- Lacoste, E., Martineau, I., and Gagnon, G. (2003), "Platelet concentrates: effects of calcium and thrombin on endothelial cell proliferation and growth factor release," *J Periodontol*, Oct;74(10):1498-507.
- Han, B., Woodell-May, J., Ponticelli, M., Yang, Z., and Nimmi, M. (2009), "The effect of thrombin activation of platelet-rich plasma on demineralized bone matrix osteoinductivity," *J Bone Joint Surg Am*, Jun;91(6):1459-70.
- Fufa, D., Shealy, B., Jacobson, M., Kevy, S., and Murray, M.M. (2008), "Activation of platelet-rich plasma using soluble type I collagen," *J Oral Maxillofac Surg*, Apr;66(4):684-90.
- Rashid, A., ElSharaby, F.A., Nassef, E.M., Mehanni, S., and Mostafa, Y.A. (2017), "Effect of platelet-rich plasma on orthodontic tooth movement in dogs," *Orthod Craniofac Res*, May;20(2):102-110.
- Güleç, A., Bakkalbaşı, B.Ç., Cumbul, A., Uslu, Ü., Alev, B., and Yarat, A. (2017), "Effects of local platelet-rich plasma injection on the rate of orthodontic tooth movement in a rat model: A histomorphometric study," *Am J Orthod Dentofacial Orthop*, Jan;151(1):92-104.
- Çağlaroğlu, M., and Erdem, A. (2012), "Histopathologic investigation of the effects of prostaglandin E2 administered by different methods on tooth movement and bone metabolism," *Korean J Orthod*, Jun;42(3):118-28.
- Graziani, F., Ivanovski, S., Cei, S., Ducci, F., Tonetti, M., and Gabriele, M. (2006), "The in vitro effect of different PRP concentrations on osteoblasts and fibroblasts," *Clin Oral Implants Res*, Apr;17(2):12-9.
- El-Sharkawy, H., Kantarci, A., Deady, J., Hasturk, H., Liu, H., Alshahat, M., and Van Dyke, T.E. (2007), "Platelet-rich plasma: growth factors and pro- and anti-inflammatory properties," *J Periodontol*, Apr;78(4):661-9.
- Freymiller, E.G., and Aghaloo, T.L. (2004), "Platelet-rich plasma: ready or not?" *J Oral Maxillofac Surg*, Apr;62(4):484-8.
- Sufamap, E., Sofyanti, E., and Ilyas, S. (2018), "The Effect of Platelet-Rich Plasma to Orthodontic Tooth Movement. In International Dental Conference of Sumatera Utara 2017 (IDCSU 2017) Feb 7. Atlantis Press.
- Nakornnoi, T., Leethanakul, C., and Samruajbenjakun, B. (2019), "The influence of leukocyte-platelet-rich plasma on accelerated orthodontic tooth movement in rabbits," *Korean Journal of Orthod*, 49, 372-380.
- Akbulut, S., Yagci, A., Yay, A.H., and Yalcin, B. (2019), "Experimental investigation of effects of platelet-rich plasma on early phases of orthodontic tooth movement," *Am J Orthod Dentofacial Orthop*, Jan;155(1):71-79.
- El-Timamy, A., El Sharaby, F., Eid, F., El Dakrouy, A., Mostafa, Y., and Shaker, O. (2020), "Effect of platelet-rich plasma on the rate of orthodontic tooth movement," *Angle Orthod*, 90, 354-361.
- Ahmad, S. and Hassan, H. (2019), "Does PRP injection accelerate orthodontic tooth movement when retracting upper canines? A split-mouth-design randomized controlled trial," *International journal of Advanced research*, (in press).