

BÖLÜM 14

PERİODONTAL REJENERASYONDA TROMBOSİTTEN ZENGİN FİBRİN TÜREVLERİNİN KULLANIM ALANLARI VE ETKİNLİĞİ

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GİRİŞ

Trombosit türevleri lökosit ve fibrin içeriğine göre dört ana gruba ayrılırlar. Bunlar; saf trombositten zengin plazma, lökosit-trombositten zengin plazma, saf trombositten zengin fibrin ve lökosit-trombositten zengin fibrindir (L-TZF). Bu sınıflama, ilk nesil trombositten zengin plazma (TZP) ve ikinci nesil trombositten zengin fibrin (TZF) olarak iki gruba da ayrılabilir¹. İkinci nesil trombosit türevi olan TZF, ilk olarak Choukroun tarafından tanımlanmış otolog, trombinize olmayan, trombosit ve lökositten zengin yoğun ve stabil bir fibrin ağıdır. Sığır trombini gibi bir antikoagülan ilavesi gerektiren TZP'nın aksine, ilavesiz basit bir şekilde santrifüj edilerek oluşturulduğu için tamamen otologdur⁽²⁾. Uygun koşullarda hazırlanmış bir TZF membran, immün-enflamatuar cevap ve yara iyileşmesi ile ilgili çeşitli sitokin ve büyüme faktörlerini 10-14 gün boyunca salabilmektedir³. Pıhtılaşma sürecinin son aşamasında şekillenen üç boyutlu fibrin iskeleti aynı zamanda büyüme faktörleri için bir depo görevi görerek osteoblast, endotel hücreleri, kondrositler ve çeşitli fibroblast kaynaklarının çoğalmasını ve farklılaşmasını tetikler^(4,5). TZP'ya göre daha yavaş çözünerek trombosit ve sitokinlerin etkili bir şekilde bölgede kalmasına ve zamanla kademeli olarak salınmasına olanak sağlar⁽⁶⁾.

TZF'in Yara İyileşmesindeki Etkileri

TZF'in yara iyileşmesindeki rolünde lökosit ve trombosit sitokinlerinin önemli etkileri olmakla birlikte, iyileştirici potansiyelinden esas olarak sorumlu tutulan, bu elementleri destekleyen fibrin matriksdir⁽⁷⁾. Anjiojenik özellikleri, trombosit kaynaklı büyüme faktörü, dönüştürücü büyüme faktörü, insülin benzeri büyüme faktörü ve vasküler endotelial büyüme faktörü gibi matriks içerisine gömülü çeşitli büyüme faktörleri ve çeşitli sitokinleri barındıran 3 boyutlu fibrin matriks yapısı ile açıklanabilir. Bu moleküller düzenli ve yavaş bir şekilde salınarak yumu-

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SONUÇ

TZF'in periodontal rejenerasyonda kullanımı ile ilgili literatürde çok sayıda çalışma olmakla birlikte, sert-yumuşak doku iyileşmesi ve postoperatif ağrı üzerine olumlu etkileri sıklıkla rapor edilmiştir. Bununla birlikte, rejeneratif yöntemlerle TZF'in optimal etkisini elde etmek için protokolün standardizasyonu gereklidir. TZF'in doğru bir şekilde ve cerrahi alan başına yeterli sayıda kullanımı da bu teknikten fayda elde etmek için önemli olabilecek bir unsurdur. TZF ve türevleri, bildirilen iyi biyolojik etkileri, hazırlanma kolaylığı, düşük maliyeti ve tamamen otolog olması nedeniyle umut vadeden bir biyomateryal olma özelliğindedir.

KAYNAKÇA

1. 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 in biotechnology*. 2009;27(3):158-167.
2. 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 surgery, oral medicine, oral pathology, oral radiology, and endodontics*. 2006;101(3):e56-60.
3. Bielecki T, Dohan Ehrenfest DM, Everts PA, Wiczowski A. The role of leukocytes from L-PRP/L-PRF in wound healing and immune defense: new perspectives. *Current pharmaceutical biotechnology*. 2012;13(7):1153-1162.
4. Kang YH, Jeon SH, Park JY, et al. Platelet-rich fibrin is a Bioscaffold and reservoir of growth factors for tissue regeneration. *Tissue engineering. Part A*. 2011;17(3-4):349-359.
5. Dohan Ehrenfest DM, Diss A, Odin G, Doglioli P, Hippolyte MP, Charrier JB. In vitro effects of Choukroun's PRF (platelet-rich fibrin) on human gingival fibroblasts, dermal prekeratinocytes, preadipocytes, and maxillofacial osteoblasts in primary cultures. *Oral surgery, oral medicine, oral pathology, oral radiology, and endodontics*. 2009;108(3):341-352.
6. Kobayashi E, Fluckiger L, Fujioka-Kobayashi M, et al. Comparative release of growth factors from PRP, PRF, and advanced-PRF. *Clinical oral investigations*. 2016;20(9):2353-2360.
7. Michael Toffler NT, Dan Holtzclaw, Marco Del Corso, David Dohan Ehrenfest. Introducing Choukroun's platelet rich fibrin (PRF) to the reconstructive surgery milieu. *J Implant Adv Clin Dent*. 2009;1:22-31.
8. Dohan DM, Choukroun J, Diss A, et al. Platelet-rich fibrin (PRF): a second-generation platelet concentrate. Part II: platelet-related biologic features. *Oral surgery, oral medicine, oral pathology, oral radiology, and endodontics*. 2006;101(3):e45-50.
9. Clipet F, Tricot S, Alno N, et al. In vitro effects of Choukroun's platelet-rich fibrin conditioned medium on 3 different cell lines implicated in dental implantology. *Implant dentistry*. 2012;21(1):51-56.
10. Vahabi S, Vaziri S, Torshabi M, Rezaei Esfahrood Z. Effects of Plasma Rich in Growth Factors and Platelet-Rich Fibrin on Proliferation and Viability of Human Gingival Fibroblasts. *J Dent (Tehran)*. 2015;12(7):504-512.
11. Castro AB, Herrero ER, Slomka V, Pinto N, Teughels W, Quirynen M. Antimicrobial capacity of Leucocyte-and Platelet Rich Fibrin against periodontal pathogens. *Scientific reports*. 2019;9(1):8188.
12. 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 surgery, oral medicine, oral pathology, oral radiology, and endodontics*. 2006;101(3):e51-55.
13. Rowe SL, Lee S, Stegemann JP. Influence of thrombin concentration on the mechanical and morphological properties of cell-seeded fibrin hydrogels. *Acta biomaterialia*. 2007;3(1):59-67.

14. Richard Miron JC, Shahram Ghanaati. Controversies related to scientific report describing g-forces from studies on platelet-rich fibrin: Necessity for standardization of relative centrifugal force values. *International Journal of Growth Factors and Stem Cells in Dentistry*. 2018;1(3):80.
15. Tunali M, Ozdemir H, Kucukodaci Z, et al. A novel platelet concentrate: titanium-prepared platelet-rich fibrin. *Biomed Res Int*. 2014;2014:209548.
16. Tunali M, Ozdemir H, Kucukodaci Z, Akman S, Firatli E. In vivo evaluation of titanium-prepared platelet-rich fibrin (T-PRF): a new platelet concentrate. *The British journal of oral & maxillofacial surgery*. 2013;51(5):438-443.
17. Fujioka-Kobayashi M, Miron RJ, Hernandez M, Kandalam U, Zhang Y, Choukroun J. Optimized Platelet-Rich Fibrin With the Low-Speed Concept: Growth Factor Release, Biocompatibility, and Cellular Response. *Journal of periodontology*. 2017;88(1):112-121.
18. Masuki H, Okudera T, Watanebe T, et al. Growth factor and pro-inflammatory cytokine contents in platelet-rich plasma (PRP), plasma rich in growth factors (PRGF), advanced platelet-rich fibrin (A-PRF), and concentrated growth factors (CGF). *International journal of implant dentistry*. 2016;2(1):19.
19. Caruana A, Savina D, Macedo JP, Soares SC. From Platelet-Rich Plasma to Advanced Platelet-Rich Fibrin: Biological Achievements and Clinical Advances in Modern Surgery. *European journal of dentistry*. 2019;13(2):280-286.
20. Miron RJ, Fujioka-Kobayashi M, Hernandez M, et al. Injectable platelet rich fibrin (i-PRF): opportunities in regenerative dentistry? *Clinical oral investigations*. 2017;21(8):2619-2627.
21. Sarah Al-Maawi CH-V, Anna Orłowska, Ines Willershausen, Robert Sader, Richard J Miron, Joseph Choukroun, Shahram Ghanaati Biologization of Collagen-Based Biomaterials Using Liquid-Platelet-Rich Fibrin: New Insights into Clinically Applicable Tissue Engineering. *Materials*. 2019;12(23):3993.
22. Choukroun J, Ghanaati S. Reduction of relative centrifugation force within injectable platelet-rich-fibrin (PRF) concentrates advances patients' own inflammatory cells, platelets and growth factors: the first introduction to the low speed centrifugation concept. *Eur J Trauma Emerg Surg*. 2018;44(1):87-95.
23. Richard J. Miron JC. Platelet Rich Fibrin in Regenerative Dentistry: Biological Background and Clinical Indications. 2017;29(1):48.
24. Richard J. Miron JC. Platelet Rich Fibrin in Regenerative Dentistry: Biological Background and Clinical Indications, 2017;18(2):331.
25. Zhang J, Yin C, Zhao Q, et al. Anti-inflammation effects of injectable platelet-rich fibrin via macrophages and dendritic cells. *J Biomed Mater Res A*. 2020;108(1):61-68.
26. Polimeni G, Xiropaidis AV, Wikesjo UM. Biology and principles of periodontal wound healing/regeneration. *Periodontology 2000*. 2006;41:30-47.
27. Wang HL, Greenwell H, Fiorellini J, et al. Periodontal regeneration. *Journal of periodontology*. 2005;76(9):1601-1622.
28. Sculean A, Nikolidakis D, Nikou G, Ivanovic A, Chapple IL, Stavropoulos A. Biomaterials for promoting periodontal regeneration in human intrabony defects: a systematic review. *Periodontology 2000*. 2015;68(1):182-216.
29. Cortellini P, Tonetti MS. Clinical concepts for regenerative therapy in intrabony defects. *Periodontology 2000*. 2015;68(1):282-307.
30. Wang HL, Boyapati L. "PASS" principles for predictable bone regeneration. *Implant dentistry*. 2006;15(1):8-17.
31. Richard J Miron VM, Masako Fujioka-Kobayashi. Use of platelet-rich fibrin for the treatment of periodontal intrabony defects: a systematic review and meta-analysis. *Clinical oral investigations*. 2021.
32. Liang Chen YD, Guoping Cheng. Use of Platelet-Rich Fibrin in the Treatment of Periodontal Intrabony Defects: A Systematic Review and Meta-Analysis. *BioMed Research International*. 2021;13.
33. Chadwick JK, Mills MP, Mealey BL. Clinical and Radiographic Evaluation of Demineralized Freeze-Dried Bone Allograft Versus Platelet-Rich Fibrin for the Treatment of Periodontal Intrabony Defects in Humans. *Journal of periodontology*. 2016;87(11):1253-1260.

34. Mathur A, Bains VK, Gupta V, Jhingran R, Singh GP. Evaluation of intrabony defects treated with platelet-rich fibrin or autogenous bone graft: A comparative analysis. *European journal of dentistry*. 2015;9(1):100-108.
35. Shah M, Patel J, Dave D, Shah S. Comparative evaluation of platelet-rich fibrin with demineralized freeze-dried bone allograft in periodontal infrabony defects: A randomized controlled clinical study. *Journal of Indian Society of Periodontology*. 2015;19(1):56-60.
36. Agarwal A, Gupta ND, Jain A. Platelet rich fibrin combined with decalcified freeze-dried bone allograft for the treatment of human intrabony periodontal defects: a randomized split mouth clinical trail. *Acta odontologica Scandinavica*. 2016;74(1):36-43.
37. Bodhare GH, Kolte AP, Kolte RA, Shirke PY. Clinical and radiographic evaluation and comparison of bioactive bone alloplast morsels when used alone and in combination with platelet-rich fibrin in the treatment of periodontal intrabony defects-A randomized controlled trial. *Journal of periodontology*. 2019;90(6):584-594.
38. Sezgin Y, Uraz A, Taner IL, Culhaoglu R. Effects of platelet-rich fibrin on healing of intra-bony defects treated with anorganic bovine bone mineral. *Brazilian oral research*. 2017;31:e15.
39. Rosamma Joseph V, Raghunath A, Sharma N. Clinical effectiveness of autologous platelet rich fibrin in the management of infrabony periodontal defects. *Singapore dental journal*. 2012;33(1):5-12.
40. Patel GK, Gaekwad SS, Gujjari SK, S CV. Platelet-Rich Fibrin in Regeneration of Intrabony Defects: A Randomized Controlled Trial. *Journal of periodontology*. 2017;88(11):1192-1199.
41. Gummaluri SS, Bhattacharya HS, Astekar M, Cheruvu S. Evaluation of titanium-prepared platelet-rich fibrin and leucocyte platelet-rich fibrin in the treatment of intra-bony defects: A randomized clinical trial. *J Dent Res Dent Clin Dent Prospects*. 2020;14(2):83-91.
42. Ustaoglu G, Ugur Aydin Z, Ozelci F. Comparison of GTR, T-PRF and open-flap debridement in the treatment of intrabony defects with endo-perio lesions: a randomized controlled trial. *Medicina oral, patologia oral y cirugia bucal*. 2020;25(1):e117-e123.
43. Panda S, Sankari M, Satpathy A, et al. Adjunctive Effect of Autologous Platelet-Rich Fibrin to Barrier Membrane in the Treatment of Periodontal Intrabony Defects. *The Journal of craniofacial surgery*. 2016;27(3):691-696.
44. Lei L, Yu Y, Han J, et al. Quantification of growth factors in advanced platelet-rich fibrin and concentrated growth factors and their clinical efficacy as adjunctive to the GTR procedure in periodontal intrabony defects. *J Periodontol*. 2020;91(4):462-472.
45. Aydemir Turkal H, Demirem S, Dolgun A, Keceli HG. Evaluation of the adjunctive effect of platelet-rich fibrin to enamel matrix derivative in the treatment of intrabony defects. Six-month results of a randomized, split-mouth, controlled clinical study. *Journal of clinical periodontology*. 2016;43(11):955-964.
46. Gupta SJ, Jhingran R, Gupta V, Bains VK, Madan R, Rizvi I. Efficacy of platelet-rich fibrin vs. enamel matrix derivative in the treatment of periodontal intrabony defects: a clinical and cone beam computed tomography study. *Journal of the International Academy of Periodontology*. 2014;16(3):86-96.
47. Salvi GE, Mischler DC, Schmidlin K, et al. Risk factors associated with the longevity of multi-rooted teeth. Long-term outcomes after active and supportive periodontal therapy. *Journal of clinical periodontology*. 2014;41(7):701-707.
48. Bajaj P, Pradeep AR, Agarwal E, et al. Comparative evaluation of autologous platelet-rich fibrin and platelet-rich plasma in the treatment of mandibular degree II furcation defects: a randomized controlled clinical trial. *Journal of periodontal research*. 2013;48(5):573-581.
49. Kanoriya D, Pradeep AR, Garg V, Singhal S. Mandibular Degree II Furcation Defects Treatment With Platelet-Rich Fibrin and 1% Alendronate Gel Combination: A Randomized Controlled Clinical Trial. *Journal of periodontology*. 2017;88(3):250-258.
50. Sharma A, Pradeep AR. Autologous platelet-rich fibrin in the treatment of mandibular degree II furcation defects: a randomized clinical trial. *Journal of periodontology*. 2011;82(10):1396-1403.

51. Lohi HS, Nayak DG, Uppoor AS. Comparative Evaluation of the Efficacy of Bioactive Ceramic Composite Granules Alone and in Combination with Platelet Rich Fibrin in the Treatment of Mandibular Class II Furcation Defects: A Clinical and Radiographic Study. *Journal of clinical and diagnostic research : JCDR*. 2017;11(7):ZC76-ZC80.
52. Rani N, Kaushal S, Singh S, Nandlal, Khan MA, Pathak AK. Evaluation of the relative efficacy of autologous platelet-rich fibrin membrane in combination with beta-tricalcium phosphate (Septodont- resorbable tissue replacement) alloplast versus beta-TCP alloplast alone in the treatment of grade II furcation defects. *National journal of maxillofacial surgery*. 2018;9(2):196-204.
53. Basireddy A, Prathypaty SK, Yendluri DB, Potharaju SP. Demineralized freeze-dried bone allograft with or without platelet-rich fibrin in the treatment of mandibular Degree II furcation defects: A clinical and cone beam computed tomography study. *Journal of Indian Society of Periodontology*. 2019;23(3):242-248.
54. Tarallo F, Mancini L, Pitzurra L, Bizzarro S, Tepedino M, Marchetti E. Use of Platelet-Rich Fibrin in the Treatment of Grade 2 Furcation Defects: Systematic Review and Meta-Analysis. *Journal of clinical medicine*. 2020;9(7).
55. Borie E, Olivi DG, Orsi IA, et al. Platelet-rich fibrin application in dentistry: a literature review. *International journal of clinical and experimental medicine*. 2015;8(5):7922-7929.
56. Eren G, Kantarci A, Sculean A, Atilla G. Vascularization after treatment of gingival recession defects with platelet-rich fibrin or connective tissue graft. *Clinical oral investigations*. 2016;20(8):2045-2053.
57. Sameera S, Nagasri M, Aravind Kumar P, Indeevar P, Raviraj K, Musalaiah S. Comparison of two surgical techniques in the treatment of multiple gingival recessions sandwiched with a combination of A-PRF and L-PRF. *Saudi Dent J*. 2018;30(3):183-189.
58. Dixit N, Lamba AK, Faraz F, Tandon S, Aggarwal K, Ahad A. Root coverage by modified coronally advanced flap with and without platelet-rich fibrin: A clinical study. *Indian journal of dental research : official publication of Indian Society for Dental Research*. 2018;29(5):600-604.
59. Gupta S, Banthia R, Singh P, Banthia P, Raje S, Aggarwal N. Clinical evaluation and comparison of the efficacy of coronally advanced flap alone and in combination with platelet rich fibrin membrane in the treatment of Miller Class I and II gingival recessions. *Contemporary clinical dentistry*. 2015;6(2):153-160.
60. Kuka S, Ipci SD, Cakar G, Yilmaz S. Clinical evaluation of coronally advanced flap with or without platelet-rich fibrin for the treatment of multiple gingival recessions. *Clinical oral investigations*. 2018;22(3):1551-1558.
61. Padma R, Shilpa A, Kumar PA, Nagasri M, Kumar C, Sreedhar A. A split mouth randomized controlled study to evaluate the adjunctive effect of platelet-rich fibrin to coronally advanced flap in Miller's class-I and II recession defects. *Journal of Indian Society of Periodontology*. 2013;17(5):631-636.
62. Thamaraiselvan M, Elavarasu S, Thangakumaran S, Gadagi JS, Arthie T. Comparative clinical evaluation of coronally advanced flap with or without platelet rich fibrin membrane in the treatment of isolated gingival recession. *Journal of Indian Society of Periodontology*. 2015;19(1):66-71.
63. Miron RJ, Moraschini V, Del Fabbro M, et al. Use of platelet-rich fibrin for the treatment of gingival recessions: a systematic review and meta-analysis. *Clinical oral investigations*. 2020;24(8):2543-2557.
64. Oncu E. The Use of Platelet-Rich Fibrin Versus Subepithelial Connective Tissue Graft in Treatment of Multiple Gingival Recessions: A Randomized Clinical Trial. *The International journal of periodontics & restorative dentistry*. 2017;37(2):265-271.
65. Culhaoglu R, Taner L, Guler B. Evaluation of the effect of dose-dependent platelet-rich fibrin membrane on treatment of gingival recession: a randomized, controlled clinical trial. *Journal of applied oral science : revista FOB*. 2018;26:e20170278.
66. Jankovic S, Aleksic Z, Klokkevold P, et al. Use of platelet-rich fibrin membrane following treatment of gingival recession: a randomized clinical trial. *The International journal of periodontics & restorative dentistry*. 2012;32(2):e41-50.

67. Uzun BC, Ercan E, Tunali M. Effectiveness and predictability of titanium-prepared platelet-rich fibrin for the management of multiple gingival recessions. *Clin Oral Investig*. 2018;22(3):1345-1354.
68. Mancini L, Tarallo F, Quinzi V, Fratini A, Mummolo S, Marchetti E. Platelet-Rich Fibrin in Single and Multiple Coronally Advanced Flap for Type 1 Recession: An Updated Systematic Review and Meta-Analysis. *Medicina (Kaunas)*. 2021;57(2).
69. Ucak Turer O, Ozcan M, Alkaya B, Surmeli S, Seydaoglu G, Haytac MC. Clinical evaluation of injectable platelet-rich fibrin with connective tissue graft for the treatment of deep gingival recession defects: A controlled randomized clinical trial. *J Clin Periodontol*. 2020;47(1):72-80.
70. Abirami T, Subramanian S, Prakash PSG, Victor DJ, Devapriya AM. Comparison of Connective Tissue Graft and Platelet Rich Fibrin as Matrices in A Novel Papillary Augmentation Access: A Randomized Controlled Clinical Trial. *European journal of dentistry*. 2019;13(4):607-612.
71. Jankovic S, Aleksic Z, Milinkovic I, Dimitrijevic B. The coronally advanced flap in combination with platelet-rich fibrin (PRF) and enamel matrix derivative in the treatment of gingival recession: a comparative study. *The European journal of esthetic dentistry : official journal of the European Academy of Esthetic Dentistry*. 2010;5(3):260-273.
72. Panda S, Satpathy A, Chandra Das A, et al. Additive Effect of Platelet Rich Fibrin with Coronally Advanced Flap Procedure in Root Coverage of Miller's Class I and II Recession Defects-A PRISMA Compliant Systematic Review and Meta-Analysis. *Materials (Basel)*. 2020;13(19).
73. Hofmanner P, Alessandri R, Laugisch O, et al. Predictability of surgical techniques used for coverage of multiple adjacent gingival recessions--A systematic review. *Quintessence Int*. 2012;43(7):545-554.
74. Izol BS, Uner DD. A New Approach for Root Surface Biomodification Using Injectable Platelet-Rich Fibrin (I-PRF). *Med Sci Monit*. 2019;25:4744-4750.
75. Nurden AT. Platelets, inflammation and tissue regeneration. *Thrombosis and haemostasis*. 2011;105 Suppl 1:S13-33.
76. Bahammam MA. Effect of platelet-rich fibrin palatal bandage on pain scores and wound healing after free gingival graft: a randomized controlled clinical trial. *Clinical oral investigations*. 2018;22(9):3179-3188.
77. Ustaoglu G, Ercan E, Tunali M. The role of titanium-prepared platelet-rich fibrin in palatal mucosal wound healing and histoconduction. *Acta odontologica Scandinavica*. 2016;74(7):558-564.
78. Sila Çağrı İşler AU, Janset Sengul. Evaluation Of the Patients Oral Health Related Quality of Life After Harvesting Free Gingival Graft. *Cumhuriyet Dental Journal*. 2019;22(1):11-22.
79. Sousa F, Machado V, Botelho J, Proenca L, Mendes JJ, Alves R. Effect of A-PRF Application on Palatal Wound Healing after Free Gingival Graft Harvesting: A Prospective Randomized Study. *Eur J Dent*. 2020;14(1):63-69.
80. Ozcan M, Ucak O, Alkaya B, Keceli S, Seydaoglu G, Haytac MC. Effects of Platelet-Rich Fibrin on Palatal Wound Healing After Free Gingival Graft Harvesting: A Comparative Randomized Controlled Clinical Trial. *The International journal of periodontics & restorative dentistry*. 2017;37(5):e270-e278.
81. Femminella B, Iaconi MC, Di Tullio M, et al. Clinical Comparison of Platelet-Rich Fibrin and a Gelatin Sponge in the Management of Palatal Wounds After Epithelialized Free Gingival Graft Harvest: A Randomized Clinical Trial. *Journal of periodontology*. 2016;87(2):103-113.
82. Sharma V, Kumar A, Puri K, Bansal M, Khatri M. Application of platelet-rich fibrin membrane and collagen dressing as palatal bandage for wound healing: A randomized clinical control trial. *Indian journal of dental research : official publication of Indian Society for Dental Research*. 2019;30(6):881-888.
83. Lektetur Alpan A, Torumtay Cin G. PRF improves wound healing and postoperative discomfort after harvesting subepithelial connective tissue graft from palate: a randomized controlled trial. *Clinical oral investigations*. 2020;24(1):425-436.