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Efficacy of Alb-PRF as Adjunctive to Non-surgical Periodontal Therapy in Management of Stage II Periodontitis

Abstract

Background:

This research evaluates the clinical effectiveness of local subgingival delivery of injectable Alb-PRF compared with mechanical debridement (MD) alone for the treatment of stage II periodontitis.

Methods:

Thirty patients diagnosed with stage II periodontitis through clinical and radiographic screening were randomly divided into two groups (n=15 each). Group I received mechanical debridement (MD) combined with Alb-PRF as a local drug delivery (LDD), while group II received MD alone. In group I, Alb-PRF was locally applied two weeks after the completion of mechanical debridement.

Periodontal measurement including plaque index (PI), gingival index (GI), bleeding on probing (BOP), probing depth (PD), and clinical attachment level (CAL) were recorded at baseline, three months and six months post treatment.

Results:

The Alb-PRF group demonstrated significantly greater clinical improvements than the MD alone group. At six months, the mean PD reduction was more substantial in the Alb-PRF group (from 3.77 ± 0.33 mm to 2.13 ± 0.21 mm) compared to the MD group (from 3.81 ± 0.28 mm to 2.65 ± 0.47 mm), with the difference being statistically significant ($P < 0.001$). Likewise, the Alb-PRF group achieved a significantly greater CAL gain demonstrating a reduction (from 4.13 ± 0.37 mm to 2.20 ± 0.15 mm) than the MD group (which improved from 4.22 ± 0.37 mm to 2.78 ± 0.40 mm) ($P < 0.001$).

Conclusion:

29 The adjunctive use of Alb-PRF with MD shows promising clinical benefits in the
30 treatment of stage II periodontitis, contributing to significant reductions in PD and gains in CAL
31 over a six-month period compared to MD alone.

32 **Keywords:**Alb-PRF, Stage II Periodontitis, Local drug delivery, Mechanical debridement,
33 Platelet concentrates.

34

35 **Introduction**

36 Periodontitis is a complex, multifactorial chronic inflammatory disease initiated by dental
37 plaque biofilm, leading to progressive destruction of the periodontal ligament and alveolar
38 bone.(1) This microbial challenge triggers an exaggerated host immune-inflammatory
39 response.(2) Clinically, periodontitis manifests through bleeding on probing (BOP), gingival
40 inflammation, increased probing depth (PD), clinical attachment loss (CAL), and radiographic
41 alveolar bone resorption.(3) While typically slow, periods of accelerated tissue breakdown can
42 occur.The primary objective of periodontal therapy is to arrest the progression by effectively
43 reducing the microbial burden.(4)

44 This is achieved through various therapeutic modalities, with scaling and root planing
45 (commonly referred to as mechanical debridement, MD) representing the most widely endorsed
46 non-surgical approach for subgingival biofilm removal..(4, 5) Despite their common use as an
47 adjunct to phase one therapy for targeting residual pathogens, systemic antimicrobial agents
48 exhibit considerable pharmacological obstacles. While systemic antimicrobials play a supportive
49 role in periodontal treatment, their clinical utility is compromised by site-specific inefficacy
50 linked to hepatic transformation and a higher susceptibility to systemic complications.(6, 7)

51 Moreover, conventional MD has inherent limitations, particularly in accessing deep and
52 intricate anatomy of tooth surfaces.(8, 9)Consequently, to overcome these limitations and
53 enhance treatment outcomes, the combined application of MD with localized subgingival drug
54 delivery has emerged as a synergistic approach. This strategy aims to improve treatment
55 responses in periodontitis and facilitate periodontal tissue health.(10) While a substantial
56 proportion of individuals diagnosed with periodontitis achieve favorable and enduring
57 therapeutic outcomes through MD alone, many cases exhibit an insufficient response to initial
58 treatment. Furthermore, if surgical interventions are not viable, the strategic integration of
59 adjunctive local therapies becomes crucial.(11)

60 Therefore, local drug delivery approaches have been proposed to fulfill two primary
61 objectives: to afford a mechanism for the protracted and precise delivery of pharmaceutical
62 agents or therapeutic biomolecules, and concurrently, to decrease the dosing regimen, thereby

63 fostering enhanced patient compliance and an improved quality of life.(12, 13)Effective personal
64 oral hygiene is fundamentally dependent upon the prior elimination of pre-existing dental plaque
65 and calculus. Consequently, professional oral prophylaxis and scheduled recall appointments for
66 professional mechanical plaque removal are essential components of comprehensive oral
67 care.(14)

68 Enhanced periodontal treatment outcomes can also be achieved through the adjunctive
69 use of local drug delivery systems. These modalities—including antimicrobial agents, host-
70 modulating compounds and autologous platelet concentration are administered directly into
71 periodontal pockets, providing targeted therapeutic effects that complement conventional MD
72 procedures.(15)Hence autologous platelet concentrates (APCs) have emerged as promising
73 biologically active agents in periodontal therapy, offering regenerative and anti-inflammatory
74 properties that enhance both surgical and non-surgical treatment outcomes.(16)

75 Among the various types of autologous platelet concentrates, platelet-rich fibrin (PRF)
76 stands out for its capacity to stimulate healing. This PRF contains platelets, stem-like cells,
77 cytokines, and growth factors within a structured fibrin network that supports targeted cell
78 migration and differentiation, leading to improved tissue repair.(17-19)

79 While platelet-rich fibrin (PRF) offers notable regenerative advantages, its clinical
80 efficacy in procedures requiring long-term scaffoldingsuch as guided bone regeneration (GBR)is
81 often limited by its rapid degradation, typically occurring within 2–3 weeks. To overcome this
82 limitation and expand its therapeutic effect, an advanced formulation known as albuminplatelet
83 rich fibrin (Alb-PRF), has been developed. This novel biomaterial involves the thermal
84 denaturation of the liquid portion of autologous plasma, primarily albumin, through a heated
85 process, resulting in the formation of a biocompatible, denser fibrin matrix. This structural
86 enhancement significantly prolongs the functional lifespan of the concentrate and enhances its
87 biological activity, making Alb-PRF a promising scaffold for sustained regenerative applications
88 in periodontal and bone therapies.(20, 21)

89 While various autologous platelet concentrates have been explored, the clinical potential
90 of Alb-PRF as a sustained-release scaffold in non-surgical periodontal therapy remains to be
91 fully elucidated. Therefore, the primary aim of this randomized controlled clinical trial was to
92 evaluate the clinical effectiveness of local subgingival delivery of Alb-PRF as an adjunct to MD
93 compared to MD alone in the treatment of patients with Stage II periodontitis.

94 **Materials and Methods**

95 Patients who attended the Department of Oral Medicine and Periodontology at Mansoura
96 University, were screened for eligibility to participate in the study. The study protocol was

97 approved by the institutional ethics committee (Ref: A0401024OM) and registered on
98 ClinicalTrials.gov (ID: NCT07080294).

99 Initially, the study included 30 patients diagnosed clinically and radiographically with stage II
100 periodontitis according to the 2017 World Workshop classification.(22) Patients were randomly
101 allocated into two equal groups (n=15 each): group I (test group): MD with Alb-PRF injection,
102 while group II (control group): MD only.

103

104 **Exclusion criteria**

- 105 (a) Periodontitis patients with stage I, III, IV.
- 106 (b) pregnant or lactating women.
- 107 (c) Systemic disease which could influence the outcome of therapy.
- 108 (d) Smoking.
- 109 (e) Patients with a history of periodontal treatment or oral infections for at least three months.

110 **Inclusion criteria**

- 111 (a) Patients with stage II (pockets less than 5 mm and CAL 3-4 mm).
- 112 (b) Systemically healthy individuals aged between 30 and 50 years.
- 113 (c) Good compliance of plaque control instructions following phase I therapy.
- 114 (d) Access to follow-up care and maintenance program.

115 **Sample Size Calculation:**

116 Sample size calculation was performed using G*Power software (version 3.1.9.7). The primary
117 outcome was the intergroup difference in PD reduction at 6 months. Based on a previous
118 randomized clinical trial with a similar design (23) an effect size of 0.25 was assumed. Using an
119 independent t-test, with a significance level of 0.05 and a power of 90%, the required total
120 sample size was calculated to be 30 patients (15 per group).

121

122 **Randomization, blinding, and allocation:**

123

124 Participants were randomly assigned to either the test or control group using a computer-
125 generated randomization sequence obtained from an external web-based tool
126 (www.randomizer.org). Allocation concealment was ensured by keeping the randomization
127 sequence inaccessible to both the operator and the participants until the time of intervention.

128 Due to the nature of the intervention, blinding of the operator and participants was not feasible.
129 However, all clinical periodontal measurements were performed by a single calibrated examiner
130 who was blinded to group allocation (single-blind design). Prior to the commencement of the
131 study, the examiner underwent calibration training to ensure intra-examiner reliability, achieving
132 an intraclass correlation coefficient (ICC) greater than 0.85, indicating excellent reproducibility.

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135 **Preoperative Phase:**Participants who fulfilled all inclusion criteria received comprehensive full-
136 mouth phase I periodontal therapy. Treatment was performed under local anesthesia, when it was
137 necessary, and was completed in two visits within a 15-day period. The therapy included both
138 ultrasonic and manual instrumentation, with supragingival mechanical debridement performed
139 using a piezoelectric ultrasonic scaler (Woodpecker Medical Instrument Co., Guilin, China).

140 Following initial therapy, participants were provided with standardized oral hygiene
141 instructions. These included training in the modified Bass tooth brushing technique using a soft-
142 bristled toothbrush and regular toothpaste twice daily. Interdental cleaning was also emphasized,
143 and patients were instructed to perform it daily using dental floss or interdental brushes.

144 **GelAdministration:** A total of 10 ml of whole blood was collected from each participant using
145 sterile plastic tubes (Vacutest Kima, Arzergrande, Italy) and centrifuged at 700 g for 8 minutes in
146 horizontal centrifuge (DLAPDM2424, China). Following centrifugation, the upper yellow
147 plasma layer was identified as the liquid plasma component. The uppermost portion of this layer,
148 corresponding to platelet-poor plasma (PPP), was aspirated into a sterile syringe. PPP was heated
149 at 75°C for 10 minutes to denature plasma proteins, cooled for approximately 2 minutes and then
150 mixed with the intermediate plasma fraction containing the buffy coat, rich in platelets and
151 leukocytes (liquid-PRF), using a sterile female–female luer lock connector to obtain a
152 homogenized injectable Alb-PRF, which was immediately applied at the treatment site in group
153 I.(20)Injection was administered using a 3 ml syringe with a blunt tip into the deepest
154 periodontal pocket after two weeks of the second MD visit, this timeframe was chosen to ensure
155 that initial post-instrumentation bleeding and acute inflammation had subsided, providing a more
156 stable subgingival environment for optimal gel retention. Patients were instructed to avoid
157 brushing or flossing at the treated sites for the first 48 hours to permit gel stabilization and
158 retention. Gentle brushing away from treated areas is acceptable.

159 **Clinical Periodontal Assessment**

160 At baseline, a comprehensive **fullmouth periodontal examination** was performed at six
161 sites per tooth for all present teeth (excluding third molars) using a standardized periodontal

162 probe (UNC-15). **Target sites** were subsequently selected based on specific inclusion criteria,
163 which required PD less than 5 mm and CAL 3 - 4 mm. While PI, GI, and BOP were monitored
164 **fullmouth** to assess the patient's overall compliance and response, the primary clinical outcomes
165 (PD and CAL) were recorded **sitespecifically** for the treated and control teeth at 3 and 6-month
166 follow-up intervals. This approach was adopted to evaluate the localized therapeutic efficacy of
167 the injected materials within the specific sites.(24-30)

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170 **Statistical analysis**

171 All statistical procedures were conducted using IBM SPSS Statistics, version 27.0 (IBM Corp.,
172 Armonk, NY, 2020).

173 Categorical variables are reported as frequencies (n) and corresponding percentages (%). Inter-
174 group comparisons for categorical data were performed using the Chi-square test. When the
175 assumption for the Chi-square test was not met (specifically, if more than 20% of the cells had an
176 expected count less than 5), the Fisher's exact test was applied.

177 Quantitative data were summarized using the mean and standard deviation (SD). The normality
178 of continuous variables was assessed using the Shapiro–Wilk test for sample sizes of 50 or fewer.
179 For normally distributed quantitative variables, the Student's t-test was employed to compare the
180 two study groups. Comparisons involving more than two time points or periods were analyzed
181 using Analysis of Variance (ANOVA) with repeated measures. Statistical significance for all tests
182 was determined at a two-sided p-value < 0.05.

183 **Results**

184 The study was included thirty participants 4 participants (26.7%) were male and 11
185 (73.3%) were female for group I, while group II consisted of 5 males (33.3%) and 10 females
186 (66.7%). The mean age was 39.20 ± 5.80 years for group I and 41.87 ± 8.04 years for group II.

187 Statistical analysis of this demographic data, presented in **Table 1**, indicated no
188 significant differences between the two groups in terms of sex distribution ($p = 1.000$) or mean
189 age ($p = 0.306$). These findings confirm that both groups were comparable in their baseline
190 characteristics, ensuring a well-balanced foundation for the clinical evaluation of the two
191 treatment modalities. The clinical parameters for both groups were evaluated at baseline, three
192 months, and six months, with a comparative analysis presented in **Table 2**.

193

194 Baseline analysis revealed no significant differences between group I and group II across all
195 parameters ($P > 0.05$). Post-intervention, both groups exhibited significant intra-group
196 improvements in all clinical indices at 3 and 6 months ($P < 0.001$). Inter-group comparisons
197 showed that group I achieved significantly lower full mouth PI at 3 months ($P = 0.007$) and
198 superior full mouth BOP reduction at 6 months ($P = 0.011$), with no significant differences in GI.
199 Notably, group I demonstrated markedly superior outcomes in site-specific PD reduction and
200 CAL gain at both follow-up intervals compared to Group II ($P < 0.001$).

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204 Discussion

205 The primary objective of the study was to assess the clinical efficacy of Alb-PRF as an adjunct
206 to MD in the management of stage II periodontitis. It is important to note that since Alb-PRF is a
207 relatively recent innovation in the field of regenerative dentistry, the clinical comparisons in this
208 discussion involve various generations of platelet concentrates. These include traditional PRF,
209 L-PRF and i-PRF, all of which share the foundational biological principle of using concentrated
210 autologous growth factors to enhance periodontal healing, similar to the Alb-PRF used in this
211 study.

212 Regarding the PI, Alb-PRF group showed a significant decrease from 2.13 ± 0.47 at baseline to
213 0.71 ± 0.15 at six months. Parallel to this, the GI demonstrated a superior reduction from $2.03 \pm$
214 0.36 to 0.69 ± 0.20 during the same period. These results were statistically superior to the
215 control group, which recorded higher mean scores for both PI (0.96 ± 0.49) and GI (0.80 ± 0.18)
216 at the end of the study. While both groups showed significant intragroup improvements,
217 intergroup analysis at six months revealed no statistically significant difference in PI ($p = 0.073$)
218 or GI ($p = 0.108$) between group I and group II. This synchronized full mouth suggests that the
219 overall patient compliance and the reduction in PD provided by Alb-PRF facilitated better
220 plaque control and reduced the niche for microbial colonization across the entire dentition. Our
221 findings align with the trends reported by Youssef et al. (2024)(31) regarding the benefits of
222 platelet concentrates in soft tissue healing. However, this study offers a distinct advantage over
223 Alghannam et al. (2024)(32), who observed a non-significant increase in PI and GI scores after
224 3 months followed by statistically significant decrease again after 6 months. The sustained
225 stability observed in the present study may be attributed to the thermal denaturation of albumin,
226 which results in the formation of a denser and more stable biological scaffold. This modified
227 structure has been shown to prolong the biological activity of the concentrate compared to
228 conventional PRF formulations(20), potentially allowing for a more sustained release of growth
229 factors and enhanced tissue healing. BOP resolution was further confirmed by the significant
230 reduction in full mouth bleeding measures. In group I, the mean BOP score dropped from $0.85 \pm$

231 0.26 at baseline to 0.23 ± 0.17 at three months, eventually reaching a low of 0.19 ± 0.19 at six
232 months. In contrast, group II showed significantly higher bleeding scores (0.37 ± 0.17) at the
233 end of the study ($p < 0.001$). These results are in harmony with Gürbüz and Yıldırım (2025)(33),
234 suggesting that the concentrated growth factors in the Alb-PRF matrix accelerate vascular
235 maturation and reduce tissue fragility more effectively than mechanical therapy alone, even in
236 challenging periodontal environments.

237 PD reduction at the target sites was a pivotal finding, with group I improving from 3.77 ± 0.33
238 mm to 2.13 ± 0.21 mm at six months, significantly outperforming the control group. These results
239 are consistent with Parwani et al (2024)(34), who used PRF derivatives to enhance MD. The
240 physical presence of Alb-PRF as a biological plug provides a mechanical advantage, preventing
241 the early collapse of the pocket wall and allowing for more stable healing of the periodontal
242 tissues.

243 Group I exhibited a highly significant gain in CAL at the treated sites, improving from $4.13 \pm$
244 0.37 mm at baseline to 2.20 ± 0.15 mm at six months. This outcome supports the regenerative
245 potential of Alb-PRF. The thermal processing of albumin creates a three-dimensional scaffold
246 that provides superior space-maintenance. This protects the blood clot and provides a protected
247 niche for the proliferation of periodontal ligament cells, explaining the absence of the "rebound
248 effect" often noted in studies using short-acting concentrates (i-PRF) like those investigated by
249 Alghannam et al. (2024).(32)

250 **Limitation**

251 Despite the significant clinical improvements observed, certain limitations of this study should
252 be acknowledged. First, the sample size was relatively small ($N=30$), which, although
253 statistically powered for clinical parameters, may limit the generalizability of the findings to a
254 broader population. Second, while a six-month follow-up period provided valuable short-term
255 data, longer-term studies are necessary to evaluate the long-term stability of Alb-PRF results.
256 Furthermore, the inclusion of microbiological analyses and radiographic assessments would
257 have provided a more comprehensive evaluation.

258 **Conclusion**

259 The application of Alb-PRF as an adjunct to MD shows promising clinical benefits compared to
260 MD alone in the treatment of stage II periodontitis.

261 **Ethical Approval and Informed Consent**

262 The study was approved by the Institutional Ethics Committee of Mansoura University (Ref:
263 A0401024OM). All participants provided written informed consent before participation.

264 **Funding**

265 No funding was received for this study.

266 **Conflict Of Interest**

267 The authors affirm that there are no conflicts of interest to disclose regarding the publication of
268 this paper.

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Table (1): Demographic characteristics of study populations

	Group I	Group II	p
Gender			
Male, N (%)	4 (26.7)	5 (33.3)	1.000
Female, N (%)	11 (73.3)	10 (66.7)	
Age (years)			
Range (min – max)	33.0 – 48.0	32.0– 64.0	0.306
Mean \pm SD.	39.20 \pm 5.80	41.87 \pm 8.04	

Data was expressed using Mean \pm SD. SD: Standard deviation
p: p value for comparing between the two studied groups

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Table (2): Comparison between the two studied groups according to clinical parameters in each period:

Clinical Parameters	Periods	MD+Alb-PRF (n = 15)	MD alone (n = 15)	P
Plaque Index (PI)	Baseline	2.13 ± 0.47	2.09 ± 0.35	0.823
	After 3M	0.76 ± 0.11	0.92 ± 0.18	0.007*
	After 6M	0.71 ± 0.15	0.96 ± 0.49	0.073
	P0	<0.001*	<0.001*	
		p1=0.001*,p2=<0.001*,p3=1.000	p1=0.001*,p2=0.001*,p3=1.000	
Gingival Index (GI)	Baseline	2.03 ± 0.36	1.98 ± 0.32	0.710
	After 3M	0.74 ± 0.14	0.85 ± 0.21	0.117
	After 6M	0.69 ± 0.20	0.80 ± 0.18	0.108
	P0	<0.001*	<0.001*	
		p1=0.001*,p2=<0.001*,p3=<1.000	p1=0.001*,p2=<0.001*,p3=<1.000	
Bleeding On Probing (BOP)	Baseline	0.85 ± 0.26	0.83 ± 0.15	0.825
	After 3M	0.23 ± 0.17	0.40 ± 0.21	0.825*
	After 6M	0.19 ± 0.19	0.37 ± 0.17	0.011*
	P0	<0.001*	<0.001*	
		p1=0.001*,p2=<0.001*,p3=<1.000	p1=0.001*,p2=<0.001*,p3=<1.00	

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Pocket Depth(PD)	Baseline	3.77 ± 0.33	3.81 ± 0.28	0.681
	After 3M	2.18 ± 0.19	2.82 ± 0.41	<0.001*
	After 6M	2.13 ± 0.21	2.65 ± 0.47	0.001*
	P0	<0.001*	<0.001*	
		<i>p1</i> <0.001*, <i>p2</i> <0.001*, <i>p3</i> =1.000	<i>p1</i> <0.001*, <i>p2</i> <0.001*, <i>p3</i> =0.891	
Clinical Attachment Level (CAL)	Baseline	4.13 ± 0.37	4.22 ± 0.37	0.498
	After 3M	2.27 ± 0.14	2.96 ± 0.34	<0.001*
	After 6M	2.20 ± 0.15	2.78 ± 0.40	<0.001*
	P0	<0.001*	<0.001*	
		<i>p1</i> <0.001*, <i>p2</i> <0.001*, <i>p3</i> =0.805	<i>p1</i> <0.001*, <i>p2</i> <0.001*, <i>p3</i> =0.834	

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428 Data was expressed using Mean ± SD. SD: Standard deviation
429 p: p value for Student t-test for comparing between the two studied groups
430 p0: p value for Post Hoc test (adjusted Bonferroni) for ANOVA with repeated measures for comparison between
431 three periods
432 p1: p value for comparing between the Baseline and After 3m
433 p2: p value for comparing between the Baseline and After 6m
434 p3: p value for comparing between the After 3m and After 6m
435 *: Statistically significant at $p \leq 0.05$

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437 Group I: MD with Alb-PRF injection group

438 Group II: MD only.

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