#### **ORIGINAL ARTICLE**



# The effect of photobiomodulation therapy on nonsurgical periodontal treatment in patients with type 2 diabetes mellitus: a randomized controlled, single-blind, split-mouth clinical trial

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## Abstract

Photobiomodulation therapy (PBMT) is a method currently used in the treatment of hard and soft tissue injuries due to its accelerating and enhancing effects on healing. In this study, we aimed to evaluate the possible additional benefits of applying PBMT with nonsurgical periodontal treatment in type 2 diabetes mellitus (DM) patients with chronic periodontitis (CP). Twenty-two type 2 DM patients with CP were enrolled in this clinical split-mouth study. Probing pocket depth (PPD), gingival index (GI), plaque index (PI), and clinical attachment level (CAL) were measured by intracaliber clinician (H.G.) at baseline and at 1 m, 3 m, and 6 m after treatment. Gingival crevicular fluid (GCF) samples were collected at baseline and at 1 week and 1 m, 3 m, and 6 m after treatment. According to split-mouth design, one randomly selected quadrant was treated with PBMT + nonsurgical periodontal treatment (NSPT) and the other quadrant was treated only non-surgical periodontal treatment. PBMT was applied the test quadrant on NSPT day and first, third, and seventh day after treatment at an energy density of 7.64 J/cm<sup>2</sup>. Repeated measures analysis of variance test was used for the intragroup comparison and a "paired *t* test" in the intergroup comparison of the clinical and laboratory findings. Comparing the test and control quadrant after treatment, the test quadrant showed significant decrease in PPD at 1 month, 3 months, and 6 months; in GI at 3 months and 6 months; in CAL at month 6; in GCF at 1 week, 1 month, 3 months; and 6 months; in comparison to the control quadrant. In contrast, there was no statistically significant difference in PI data at all times. Within the limitation of this study, adjunct use of PBMT on NSPT in patient with DM may positively affect the clinical and biochemical parameters.

Keywords Photobiomodulation · Type 2 diabetes mellitus · Chronic periodontitis · Cytokine · Non-surgical periodontal treatment

# Introduction

Periodontal disease is an inflammatory disease of the toothsupporting tissue that is caused by specific microorganisms

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and causes periodontal ligament and alveolar bone degradation [1]. Although local systemic and environmental factors play a role in the development of periodontal disease, the primary etiological factors are microbial plaque (MP) and its products [2]. Diabetes mellitus (DM) is the leading systemic disease that increases the severity of periodontal disease [3]. Diabetes mellitus is a chronic metabolic disease that progresses with hyperglycemia and develops as a result of resistance against the effect of insulin, insulin deficiency, or sometimes a complete lack of insulin. It has four subgroups: type 1 DM (insulin-dependent diabetes), type 2 DM (non-insulin-dependent diabetes), gestational DM, and other specific types [4]. Among DM patients, 90–95% have type 2 diabetes. There is a bidirectional relationship between DM and periodontal disease and also DM may increase the prevalence, incidence, and severity of periodontal diseases. Similarly, periodontal diseases may complicate the metabolic control of DM [3, 5, 6].

The first stage in the treatment of periodontal disease is nonsurgical periodontal treatment; the aim of this treatment is to remove MP, which is the primary etiological factor. Therefore, nonsurgical periodontal treatment mainly aims to eliminate supra and subgingival accumulation, as well as provide effective personal plaque control, thus stopping the progression of the disease. However, nonsurgical periodontal treatment has some limitations and difficulties, depending on the technique employed, due to anatomical formation [7, 8]. Moreover, adjunct treatment modalities such as lasers on NSPT were introduced in order to change these difficulties and limitations [8, 9]. The use of laser applications has been raised as an alternative to supportive treatments to increase the efficacy of nonsurgical periodontal treatments and decrease the aforementioned limitations [10]. Researches related to PBMT are conducted at the level of in vitro or animal studies. According to in vitro studies examining the effects of PBMT administration on human gingival fibroblasts at different wavelengths and doses, PBMT administration has a positive effect on cell proliferation in gingival fibroblasts and results in increased FGF-b expression [11-13]. In another study Hakkı et al., it was reported that type-1 collagen mRNA expression increased significantly after laser photobiomodulation [14]. Cellular and animal studies have reported that PBMT increases fibroblast and osteoblast activation and contributes to tissue healing. The biostimulative effect of PBMT on tissue healing has drawn significant attention, not only in healthy individuals but also in patients with a systemic disease such as DM [15]. However, there are limited studies on the additional benefits of PBMT in patients with uncontrolled type 2 DM-in which periodontal disease progresses more severely [16]. In our study, we aimed to investigate and evaluate the possible effects of PBMT applied using a diode laser at the 980-nm wavelength on the wound healing process, in addition to the nonsurgical periodontal treatment of CP in uncontrolled type 2 DM patients in whom wound healing is harder and more impaired.

# Materials and methods

## **Patient recruitment**

For our study, we submitted an application to the Local Ethics Committee of the Dentistry Faculty at Gaziantep University and obtained approval dated September 06, 2014 per decision no 210. Overall, 22 individuals applied to the Gaziantep University Medical Faculty's Department of Endocrinology between July 2014 and July 2015 and were referred to the Periodontology Department of the Dentistry Faculty at Gaziantep University after being diagnosed with CP as a result of clinical and radiographic examinations, serving as the uncontrolled type 2 DM patients who were included in the study.

#### **Inclusion criteria**

- 1. Volunteering to participate in the study
- 2. Having been diagnosed with type 2 DM as per the ADA criteria at least 2 years before the study and having an HbA1c value of  $\geq 7$
- 3. Not having a systemic disease other than DM
- 4. Not having the systemic complications of diabetes
- 5. Not smoking or consuming alcohol
- 6. Having at least 20 teeth
- 7. Having at least two single-rooted teeth and a periodontal pocket depth of  $\geq$  5 mm on the left and right area of each (mandibula or maxilla) symmetrically
- 8. Not having used any antibiotics or long-term anti-inflammatory drugs within the previous 6 months
- 9. Not having received periodontal treatment within the previous year
- 10. Being able to fulfill the instructions for oral hygiene training
- 11. For female patients, not being pregnant or in the breastfeeding period
- 12. Not having predisposing factors such as a dental crown, bridge, filling, or cavity on teeth from which the GCF was to be collected

Third molar teeth were not included in the study.

All the patients who fulfilled the inclusion criteria signed the informed consent form after they were informed of the aim and contents of the study.

## **Clinical measurements**

We performed clinical and radiographic periodontal examinations on all individuals included in the study. To assess the periodontal condition of each individual in the clinical examination, probing pocket depth (PPD), clinical attachment level (CAL), plaque index (PI) [17], and gingival index (GI) [18] measurements were taken from six surfaces (mesio-buccal, mid-buccal, distobuccal, mesiopalatal, midpalatal, and distopalatal); all teeth by single intra-caliber clinician (H.G.). Kappa values of PPD, CAL, PI, and GI were 0.92, 0.89, 0.93, and 0.85, respectively. These measurements were performed by a different researcher (H.G.), who did not know the treatment protocol, at baseline, first, third, and sixth months after treatment using a Williams periodontal probe (Hu-Friedy, Chicago, IL, USA) that had millimetric calibration. We recorded the performed measurements in a chart format.

## Study design

This study is a randomized, controlled, single-blind, splitmouth clinical study. A single dentist performed the periodontal treatment and laser application on all the patients included in this study. Patients' quadrants were divided into two parts: "control" and "laser" side. Areas that received PBMT in addition to nonsurgical periodontal treatment constitute the laser side, and areas that only received nonsurgical periodontal treatment constitute the control side. The quadrant on which PBMT was to be applied was determined using the heads/tails randomization technique on treatment season by another clinician K.E. For control side, the device was held on the area for the same time period, purporting to apply PBMT as a placebo.

- Group I (control area): nonsurgical periodontal treatment
- Group II (laser area): nonsurgical periodontal treatment + photobiomodulation therapy

#### Treatment protocol

Before starting periodontal treatment, we informed all patients about the relationship between MP and periodontal disease, and we demonstrated tooth brushing using a modified bass technique, together with dental floss and/or an interdental brush, on an oral hygiene training model. We recommended that patients brush their teeth for 2 min at least twice a day.

A single dentist (S.S.Ö.) applied the nonsurgical periodontal treatment and PBMT on all patients included in the study. Scaling and root planing were performed under local anesthesia within 24 h on two halves of the quadrant for 45 min in the morning session and 45 min in the afternoon session. These procedures were performed using ultrasonic devices (Piezon, OEM Built-in Kit, EMS, Switzerland) and curettes (Gracey Curette, SG 1/2, 3/4, 5/6, 7/8, 11/12, 13/14; Gracey Curette, SAS 3/4, 11/12, 13/14; Hu-Friedy, USA). Then, the tooth surface was polished using paste (Sultan prophylaxy paste, Topex, Turkey) and a rubber brush (Kerr Manufacturing Co., Romulus, MI, USA).

#### Laser parameters

Photobiomodulation therapy was applied using a GaAlAs diode laser at the 980-nm wavelength (CHEESETM, GIGAA Laser, Wuhan Gigaa Optronics Technology Co., Ltd., China) and an PBMT application handpiece (Therapy handpiece, Wuhan Gigaa Optronics Technology Co., Ltd., China) on the buccal surface by adjusting distance as 10 mm.

PBMT was applied on the buccal side of each tooth on the selected side (on the maxilla and mandibula) for 15 s continuously at 0.4 W power when the handpiece adjusts 1 cm diameter. The laser spot size was approximately 0.785 cm<sup>2</sup>, which produces an energy density of approximately 0.5  $J/cm^2$ . PBMT application was performed for four sessions immediately after the nonsurgical periodontal treatment (day

0) and the first, third, and seventh days after treatment according to our previous study by Gündoğar et al. [9].

#### **Collection of GCF samples**

GCF samples were collected from one rooted tooth each side from previously determined teeth before starting the periodontal treatment and first, third, and sixth months after treatment by using paper strips (Periopaper, Proflow Inc., New York, USA) at morning times of the day. We collected samples from the mesiobuccal or distobuccal regions of the single-rooted tooth that had the deepest pocket in the control and laser areas. Sampled areas were isolated using sterile cotton rolls, and saliva contamination was prevented by removing the existing saliva using a saliva absorber and air-water spray. If there was subgingival plaque, we removed it using a sterile curette without touching the marginal gum. For sampling, periopaper strips (Periopaper®, OraFlow, Inc., PlainView, New York, USA) were inserted into the gingival pocket in the sulcus until a slight resistance was felt; they were held within the pocket for 30 s and GCF samples were collected. Afterwards, paper strips were placed in a previously calibrated and reset Periotron 8000 device to measure GCF volume (µl) immediately, and the measured volumes were noted. Samples contaminated with blood or saliva were not included in the study. Each sample was placed in 1.5 ml Eppendorf tubes and stored at -80 °C until the day of analysis.

We performed the IL-1 $\beta$  analysis of the gingival crevicular fluid in the laboratory of the Biochemistry Department of Gaziantep University's Medical Faculty, using R&D Systems (Human IL-1 $\beta$ /IL-1F2, DLB50, MN, USA) kits from the Biotek Instruments ELx800 device. We measured the IL-1 $\beta$  levels in the GCF samples (R&D Systems, ABD) using the ELISA method according to [19].

#### Sample size calculation

Sample size was calculated based on a two-sided hypothesis test with 0.05 type 1 error. The power analysis was the minimum sample size required to accept an average change of "0.2  $\pm$  0.2 mm" in the mean value of the clinical attachment level in two different treatment modalities (control vs. laser) according to our previous study [9] which was determined to be 22. A web page (www.p005.net) was used to calculate sample size.

## Statistical analysis

We used an SPSS for Windows version 22.0 software package to analyze the data obtained from our study, and we considered a value of p < 0.05 statistically significant. We tested the normal distribution of numerical data using the Shapiro–Wilk test, including a paired *t* test to compare two dependent measurements and a repeated measures analysis of variance to compare more than two dependent measurements. The mean  $\pm$  standard deviation values were given as the descriptive statistics.

## Results

All patients, 12 (54.5%) women and 10 (45.5%) men, completed the study and mean age was  $45.32 \pm 6.191$ . Post-treatment recovery was uneventful in all patients, and there were no side effects from the laser in any patient. The demographic data of the patients are provided in Table 1.

#### **Clinical data**

Although, there was no statistically significant difference according to baseline PPD, CAL, PI, and GI data between the control and laser areas (p < 0.05). We observed that posttreatment PPD data at months 1, 3, and 6 exhibited a statistically significant decrease in the laser area compared to the control area, whereas the change in CAL data was only statistically significant at month 6 of post-treatment in the laser area (p < 0.05). Regarding PI data, there were no statistically significant differences between groups. All groups showed statistically significant decrease in PPD and CAL data at the first, third, and sixth months after treatment compared to baseline (p< 0.05). The comparisons of the clinical periodontal parameters at the baseline and at the first, third, and sixth months after treatment are provided in Table 2.

## **Biochemical data**

There was no statistically significant difference, according to baseline GCFv data in the comparison of the control and laser sides. GCFv data at the first week and first, third, and sixth months after treatment data were statistically significantly lower in the laser area compared to the control area (p < 0.05). There was a statistically significant decrease in GCFv data at the first week, and at the first, third, and sixth months of post-treatment in comparison to the baseline; at the first, third, and sixth months of post-treatment in comparison to the first.

Table 1	Socio-demographic	data
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AGE	45.32 ± 6.19
DM diagnosis time	$6.14 \pm 1.83$
HbA1c	$8.39\pm0.94$
Body mass index	$25.42\pm0.96$
Number of teeth	$24.27\pm2.47$
Gender	
Woman	12 (%54,5)
Man	10 (%45,5)

week; and at the third and sixth months post-treatment in comparison to the first month (p < 0.05). There was no statistically significant difference according to baseline IL-1 $\beta$  data in the comparison of the control and laser sides (p < 0.05). We observed that there was a statistically significant decrease in the laser area compared to the control area in terms of IL-1 $\beta$  data at the third month after treatment (p < 0.05). The intergroup and intragroup variation and comparisons of IL-1 $\beta$  and GCF volume (GCFv) at baseline, at first week, and at first, third and sixth months after treatment are provided in Table 3.

## Discussion

Although MP is the primary etiological factor in the pathogenesis of periodontal disease, the host response against microorganisms is of vital importance in the onset and progression of the disease. The host response against MP was different in each individual. One of the factors that affect the host response is systemic disease. The leading systemic disease accepted as a risk factor for periodontal disease for a long time was DM [20, 21]. It is known that there is a bidirectional relationship between DM and periodontal disease and the negative effect of DM on periodontal tissue and the negative effect of periodontal disease on the metabolic control of DM [5]. DM patients exhibit an increase in some markers in GCF due to impaired neutrophil functions and advanced glycation end products (AGE) increase in periodontal tissue. Especially in patients with uncontrolled DM, the increase in these markers causes the disease to become even more severe [22]. In addition to DM being a risk factor for periodontitis, periodontitis also has a negative effect on DM patients' metabolic control. Periodontal disease negatively affects metabolic control by causing chronic inflammation in DM patients [23]. Taylor et al. demonstrated that severe periodontitis exacerbated hyperglycemia seen in DM patients; additionally, metabolic control could be ensured and hyperglycemia diminished with nonsurgical periodontal treatment [24].

Nonsurgical periodontal treatment is accepted as the gold standard in the treatment of periodontal diseases. Researchers have investigated additional protocols to increase the efficacy of existing treatment protocols. Today, the one that stands out the most among these protocols is laser application. It is accepted that laser applications could support or be an alternative to periodontal treatment [25]. PBMT has positive effects on wound healing. PBMT increases and accelerates the healing process in damaged or diseased tissue with bio-stimulation, normalizes the permeability of blood vessels, and boosts microcirculation by causing vasodilation [26]. Although there are studies showing that applying PBMT in addition to nonsurgical periodontal treatment provides good results, there are also studies that could not find a significant

#### Table 2 Comparison of clinical data

	Baseline	1 month	3 months	6 months		
PPD (mm)						
Control	$4{,}08\pm0.44$	$3.11 \pm 0.48*$	$2.99 \pm 0.42^{*},^{**}$	$2.90 \pm 0.4^{*},^{**},^{***}$		
Laser	$4.09\pm0.29$	$2.94 \pm 0.29^{a} \ast$	$2.72 \pm 0.32^{a_*},^{**}$	$2.60\pm0.3^{a}\!\!*,\!**,\!***$		
CAL (mm)						
Control	$4.50\pm0.65$	$3,25 \pm 0.47*$	$3.06 \pm 0.41^{*},^{**}$	$3.00 \pm 0.39^{*},^{**},^{**}$		
Laser	$4.52\pm0.75$	$3.22\pm0.43^*$	$2.93 \pm 0.36^{*},^{**}$	$2.79 \pm 0.32^{a} *, **, ***$		
PI						
Control	$2.89\pm0.60$	$0.83\pm0.24*$	$0.59 \pm 0.16^{*},^{**}$	$0.54 \pm 0.15^{*},^{**}$		
Laser	$2.88\pm0.68$	$0.81\pm0.24*$	$0.54 \pm 0.15^{*},^{**}$	$0.52 \pm 0.12^{*},^{**}$		
GI						
Control	$1.91\pm0.14$	$0.74\pm0.24*$	$0.52 \pm 0.17^*,^{**}$	$0.48 \pm 0.16^{*},^{**},^{***}$		
Laser	$1.92\pm0.17$	$0.73 \pm 0.27*$	$0.40 \pm 0.15^{a_*}, **$	$0.38 \pm 0.10^{a*}, **, ***$		

\*Significant difference from baseline (p < 0.05); \*\*significant difference from 1 month (p < 0.05); \*\*\*significant difference from 3 months (p < 0.05) a Significant difference between groups (control vs laser) (p < 0.05)

difference [27–31]. Therefore, additional randomized controlled studies are required to evaluate the effects of applying PBMT with nonsurgical periodontal treatment.

In our study, we aimed to investigate the effects of applying PBMT with a diode laser at the 980-nm wavelength along with nonsurgical periodontal treatment on clinical periodontal parameters (PI, GI, PPD, CAL), as well as GCFv and IL-1 $\beta$  levels in GCF in CP patients with type 2 DM. To the best of our knowledge, there are no studies that evaluate the effects of applying PBMT with lasers at the 980-nm wavelength along with nonsurgical periodontal treatment in CP patients with type 2 DM.

The host defense system is of vital importance in periodontal diseases. Therefore, we preferred the split-mouth study design to ensure the standardization of the study and eliminate possible environmental and local factors among individuals and host-dependent factors.

Al-Zahrani et al. divided CP patients with type 2 DM into three groups: group 1 only received nonsurgical periodontal treatment, group 2 received doxycycline, and group 3 received photodynamic laser therapy with PBMT (670 nm) in addition to nonsurgical periodontal treatment. These three groups yielded similar results in terms of periodontal parameters. As a result, the authors concluded that applying photodynamic laser therapy in addition to nonsurgical periodontal treatment in patients with type 2 DM did not have any additional benefits [32]. Macedo et al. divided DM patients into two groups, wherein group 1 received doxycycline and group 2 received photodynamic laser therapy at 660 nm in addition to nonsurgical periodontal treatment. These researchers obtained similar results in both groups in terms of periodontal parameters [33]. Obradovic et al. included 300 CP patients in their study, dividing the patients into three groups: patients with type 1 DM, patients with type 2 DM, and systemically healthy

patients. In this split-mouth study, patients in groups 1 and 2 received PBMT (670 nm, 5 mW, 14 min/day) for 5 days on the gingival tissue of the right premolar teeth immediately after nonsurgical periodontal treatment. In this study, we concluded that applying PBMT in addition to nonsurgical periodontal treatment in patients with DM reduced gingival inflammation and may make positive contributions to the healing process; further, PBMT could be used as a supportive therapy for nonsurgical periodontal treatment in CP patients with DM [16]. Qadri et al. investigated the short-term (8 weeks) effects of PBMT applied in addition to nonsurgical periodontal treatment in their split-mouth, double-blind controlled study. They applied a laser at 635 nm wavelength  $(4.4 \text{ J/cm}^2)$ , InGaAlP) on buccal papilla and at 830 nm wavelength (8.75 J/cm<sup>2</sup>, GaAlAs) on buccal and lingual section with slight contact to tissue, once a week for 6 weeks, 1 week after nonsurgical periodontal treatment. A significant decrease was observed in the laser application area in terms of PPD, PI, and GI values measured 1 week after the last laser application. The investigators thought this result might stem from the laser beam reducing  $PGE_2$  or from cellular ATP stimulation [34]. Aykol et al. used PBMT (808 nm, 4 J/cm<sup>2</sup>, GaAlAs) in addition to nonsurgical periodontal treatment in their parallel, randomized controlled study. They applied PBMT on days 1, 2, and 7 after nonsurgical periodontal treatment. They found that there was a statistically significant difference in the PPD, CAL, and GI values between the laser and control groups [35]. Sağlam et al. conducted a study on 30 CP patients who were systemically healthy and compared the group that only received nonsurgical periodontal treatment with the group that also received PBMT. As a result of the 6-month follow-up, they found that PI, GI, PPD, and CAL values were statistically significant in the group that received the PBMT application [19]. Despite the promising results, the use of laser in addition

Table 3

Comparison of biochemical data

	IL-1β pg/30s		GCF volume (µl)	
	Control	Laser	Control	Laser
Baseline	$60.99 \pm 49.28$	$72.80 \pm 45.54$	$0.84 \pm 0.21$	$0.87 \pm 0,20$
1 week	$25.29 \pm 13.93^{\dagger}$	$22.24 \pm 22.32^{\dagger}$	$0.36\pm0.16^\dagger$	$0.32 \pm 0.18^{*, \dagger}$
1 month	$29.17 \pm 20.64^{\dagger,\Omega}$	$26.52 \pm 25.06^{\dagger, \Omega}$	$0.30\pm0.11^{\dagger,\Omega}$	$0.26\pm0,18^{*\dagger,\Omega}$
3 months	$29.45\pm34.02^{\dagger,\Omega,}$	$15.28 \pm 12.06^{*\dagger, \ \Omega}$	$0.26\pm0.10^\dagger$	$0.22 \pm 0.06^{*,\dagger,\Omega}$
6 months	$26.02\pm23.16$	$18.11\pm17.58^\dagger$	$0.24\pm0.09^{\dagger}$	$0.20\pm0.08^{*,\dagger,\Omega}$

\*Significant difference between groups (p < 0.05); <sup>†</sup> significant difference from baseline (p < 0.05); <sup>Ω</sup> significant difference from 1 week (p < 0.05)

to periodontal treatment is still controversial. According to the results of some studies, the combined use of laser applications and nonsurgical periodontal treatment did not indicate superiority over nonsurgical periodontal treatment alone in terms of microbial and gingival parameters. Walter et al. conducted a split-mouth study on 35 systemically healthy CP patients and applied a 980-nm diode laser in addition to the classical treatment. They applied diode laser therapy on days 1, 3, and 7 after non-surgical periodontal therapy. They reevaluated PI, GI, CAL, and PPD values at weeks 6 and 18 and did not find a significant difference between the two groups in terms of PI, GI, CAL, and PPD 7-10 mm pockets. They only found a significant difference in PPD 4-6 mm pockets [36]. Makhlouf et al. conducted a split-mouth, double-blind, controlled short-term study using PBMT (830 nm, 3 J/cm<sup>2</sup>) in addition to nonsurgical periodontal treatment. They applied PBMT for a total of 10 sessions (i.e., three sessions at weeks 1 and 2, two sessions at week 3, and one session at weeks 4 and 5) after nonsurgical periodontal treatment. They did not find a statistically significant difference between the laser and control groups in terms of GI and PI values. The researchers reported that PPD exhibited a statistically significant difference in the short term (week 5 and month 3), whereas this difference was not seen in the long term (month 6) [28]. Lai et al. applied laser eight times in addition to nonsurgical periodontal treatment on 14 patients with moderate and severe CP. They did not find a statistically significant difference between the laser and control groups in terms of clinical parameters [27]. The fact that the results of our study were different from those of some studies was thought to stem from the use of different laser parameters and different patient groups.

Interleukin-1 beta is a proinflammatory cytokine that plays an important role in immuno-inflammatory response and causes increased bone degradation [37]. It was found that the level of IL-1 $\beta$  was higher in individuals with periodontitis compared to healthy individuals and that IL-1 $\beta$  levels exhibited a decline after the treatment of individuals with periodontitis. Correa et al. conducted a study of 23 CP patients with type 2 DM and 26 systemically healthy CP patients; they found a significant decline in IL-1 $\beta$  data obtained from the samples collected from deep and shallow areas at the 3-month follow-up in comparison to the baseline [38]. Navarro et al. included 10 CP patients with DM and 10 CP patients without DM in their study. They found a significant decrease in GCF total volumes and IL-1 $\beta$  data at months 3 and 6 after nonsurgical periodontal treatment [39]. Reviewing GCFv and IL-1 $\beta$  data from the control area in our study, we determined that the results were consistent with the results in the literature of nonsurgical periodontal treatment administered to CP patients with DM.

In our study, although there was no statistically significant difference in all clinical and biochemical data comparison of groups, laser groups showed statistically significant improvement of the PPD and CAL data in all times after treatment. However, regarding IL-1 $\beta$  values, there was no statistically significant difference at week 1 or months 1 and 6; laser side showed statistically significant decrease comparison with control side. These results were consistent with the results previously stated in studies. The effect of PBMT (980 nm) application in addition to nonsurgical periodontal treatment on clinical and biochemical parameters in patient with DM was investigated in our study, and we observed that the combined use of PBMT to nonsurgical periodontal treatment in DM patient resulted in a statistically significant benefit in terms of clinical and biochemical parameters between the groups.

## Conclusion

- Within the limitations of our study, we found that applying PBMT therapy with nonsurgical periodontal treatment to CP patients with type 2 DM is beneficial and can enhance tissue healing potential and reduce inflammation and these benefits have been shown clinically and biochemically.
- Although to the best of our knowledge this is the first biochemical and clinical study evaluating the effect of PBMT applied with a diode laser at the 980-nm wavelength in addition to nonsurgical periodontal treatment on IL-1β levels in CP patients with type 2 DM, there is

a need for further research on this subject to provide a better understanding of the clinical use of applying PBMT with nonsurgical periodontal treatments in CP patients with type 2 DM.

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#### **Compliance with ethical standards**

This clinical trial was approved by local ethics committee of the University of Gaziantep (date 09.06.2014 per decision no 210), and all the participants were given information about the research, and oral and written informed consent was obtained from all participants. All procedures performed in the present study were in accordance with the Helsinki declaration.

**Conflict of interest** The authors declare that they have no conflicts of interest in publishing this article.

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