Assessing Health-Related Quality of Life with Antimicrobial Photodynamic Therapy (APDT) and Low Level Laser Therapy (LLLT) after Third Molar Removal

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Abstract:

Introduction: The purpose of this study was to evaluate the antimicrobial photodynamic therapy (APDT) and low level laser therapy (LLLT) on wound healing, pain intensity, swelling problems, halitosis and the postoperative usage of analgesics after surgical removal of lower third molars.

Methods: One hundred and fifty patients, randomly divided into three groups were selected (50 per each group). The P1 group received the APDT after a third molar surgery, the P2 group received the LLLT and the C group (control group) was without any additional therapy after surgery. A photoactive substance was applied in the APDT study group before suturing. After 60 seconds the photosensitive substance was thoroughly washed with saline water and the laser light was applied in two intervals (30 seconds each). The irradiation power was 50 mW while the wavelength was 660 nm. The laser therapy in P2 group was performed before suturing and the laser light was applied also in two intervals (90 seconds each), the irradiation power was 90 mW while the wavelength was the same as in the first group – 660 nm. Postoperative follow-ups were scheduled on the third and the seventh day in patients who received laser therapy.

Results: The results of the postoperative evaluation showed that there was a statistically significant difference in the postoperative wound healing, pain intensity, swelling problems, halitosis and analgesics intake between patients in all three groups (p<0.001). The patients that were subjected to APDT (P1) had the least postoperative problems. After the laser therapy (P1 and P2) wound healing was without any complications, opposite from the patients from the C group (p<0.001). Postoperative application of a laser therapy significantly reduced patient’s use of analgesics over the observed period of time (p<0.001).

Conclusion: Both modalities of laser therapy significantly reduced postoperative problems after surgical removal of third lower molars with the best results in both laser groups.

Keywords: low level laser therapy; photodynamic therapy; postoperative complications; diode laser

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Introduction

Low power diode lasers, also called biostimulators, base their anti-inflammatory effects, accelerated wound healing and reduction of acute and chronic pain on photobiostimulation effects. Anti-inflammatory effect of the laser beam is based on the reduction of prostaglandin (PGE2) concentration, changing the path of arachidonic acid and reduced effect of tumor necrosis factors (TNFα) in acute inflammatory states\(^1\). Furthermore, the anti-inflammatory effect of laser irradiation is contributed by the change of permeability and the size of lymphatic and blood vessel lumen and collateral growth stimulation which enhances defensive mechanism for fighting infection\(^2,3\). Accelerated healing via laser effect is achieved by stimulation of natural biological processes\(^4\). Cells with low redox-state are acidic, but turn alkaline after laser therapy and its optimal regeneration is achieved. Healthy cells can not significantly increase their redox-states and they will not react to laser energy effects, while cells with low redox-states will be stimulated\(^5\). The most important effect the laser energy has on the cell is the increase of adenosine triphosphate (ATP)\(^6,7\). Analgesic effect mechanism of the laser beam is based on changing neurotransmitter activity, especially serotonin, beta-endorphin and acetylcolinesterase. It is also proven that transient varicosities along neurons are achieved. This way, a transmission signal interference\(^8\) and inhibition of complex reaction of action potential creation occur\(^9\). The enhanced effect cannot be achieved by using multiple therapies simultaneously. On the contrary, that would lead to wound healing inhibition (Arndt-Schulz law)\(^10\). Because of that, acute states (oedema and inflammation symptoms) are treated until symptoms are cured, while therapy in chronic states (wounds, paraesthesia, chronic pain) should not be applied more that once or twice a week\(^11,13\). The biostimulative effect of laser therapy is increasingly being used for reducing postoperative problems after many operative procedures in the area of the orofacial region and even after third molar removal, one of the most frequent dentoalveolar surgical procedures in oral and maxillofacial surgery\(^14\). The period after alveolectomy is often followed by problems such as pain, halitosis and swelling problems, problems with postoperative healing which greatly impair life quality and cause work incapacity. Such problems are the reason for increased postoperative application of analgesics and antibiotics\(^15,16\).

By comparing recent literature on the effects of different lasers therapy on postoperative problems after the removal of lower third molars, there are some differences between the results. Lopez-Ramirez et al. have proved on a sample of 20 patients after third molar removal on both sides that laser irradiation has positive effects on pain, swelling problems and trismus, although not statistically significant\(^17\). Statistically insignificant effects of laser irradiation on the aforementioned postoperative symptoms have also been proven by Braams et al\(^18\). Wahl et al. have proved that laser irradiation of soft tissue statistically significantly reduced postoperative problems\(^19\). Almost all of that researches used different methodology and different laser irradiation parameters which can be the cause of result discrepancy\(^20\).

The aim of this study was to evaluate the effects of anti-microbial photodynamic therapy (APDT) and low level laser therapy (LLLT) on wound healing, pain intensity, swelling problems and halitosis after the removal of lower third molars and postoperative analgesic usage.

Methods

Subjects

This study was approved by the Ethical Committee of the School of Dental Medicine, University of Zagreb, Croatia. The research consisted of 150 participants, 92 (61%) were females and 58 (39%) were men. The participants were separated into four age groups: under 18 (8%), 19 to 24 years (42%), 25 to 30 years (25%) and over 30 years (25%). The exclusion criteria in the present study were systemic diseases, with developing local infections, smokers, pregnant women, breastfeeding mothers and patients using contraception. All participants were patients of the Department of Oral Surgery School of Dental Medicine University of Zagreb with complete indications for removal of the lower third molar. Before the surgery, patients were informed in detail about the laser therapy, its application, effects, indications and side effects and signed a participation agreement in which they accepted laser therapy after surgery. All anamnestic data was collected and inserted into a Patient Questionnaire, in the section “Patient General Information”.

Procedures

All surgical procedures were performed under
local anesthesia (UbistesinTM, 3M ESPE, Espe Plazt, D-82229 Seefeld, Germany). Local anesthetic quantity in each procedure was two ampoules per patient. Molars were removed using surgical technique of mucoperiosteal flap with osteotomy. In order not to influence the trauma level by surgical experience of the operator, both surgeons performing the operations on all participants had more than 10 years experience. Surgical approach was buccal with the incision of mucoperiosteal flap. The incision was of sufficient size in order to allow good visibility, in order not to place the operative area under tension, to make it vascularized and to rest edges of the operative area on solid bone surface. After removing the bone with drills, tooth removal was performed using elevating instruments (pliers, elevators, MEDIN, a.s., Nové Město na Moravě, Czech Republic) in the appropriate direction.

Patients were divided into three groups using a random selection: P1 group consisted of 50 patients who received APDT, P2 group consisted of 50 patients who received LLLT, and the C group (control group) consisted of 50 patients who did not receive any kind of laser therapy. In all examined groups data had been obtained by using identical questionnaire. The laser HF (Hager and Werken GmbH and Co. Duisburg, Germany, 2009.) was used in the study. In the P1 group, before suturing, a photosensitive substance consisting of toluidine chloride powder (155µg/ml), water, sodium phosphate and hydroxymethyl cellulose (LaserHF Paro - PDT solution, Hager and Werken GmbH and Co., Duisburg) was applied in the postextraction socket. After 60 seconds, the Paro-PDT solution was thoroughly rinsed using saline solution and laser light was applied directly into the surgical site for 60 seconds (2x30 seconds) with constant laser beam irradiation over the surgical area. The laser probe was used in non-contact mode and was set at the distance of approximately 5mm of the bone surface. The irradiation intensity was 50 mW, with the wavelength of 660 nm. The laser therapy in the P2 group was performed in the same manner as in the P1 group before suturing, but without the application of the Paro-PDT solution and it lasted for 180 seconds (2x90 seconds). The irradiation intensity was 90 mW and the wavelength was 660 nm, the same as the P1 group. Laser settings for both therapy modalities were according to the manufacturer’s instructions and were not changed arbitrarily. All patients received identical postoperative instructions.

The postoperative follow-ups were scheduled to the third and to the seventh day in laser P1 and P2 groups. Healing and postoperative recovery was assessed by two questionnaires (one for patients and another for surgeons). Patients evaluated their postoperative problems with grades from 1 to 4 (1=no problems, 2=mild problems, 3=medium problems, 4=intensive problems). The following symptoms were assessed: pain, swelling problems and halitosis. Patients also recorded their postoperative analgesics usage using a dichotomous scale (1=yes, 2=no). Surgeons evaluated the type of post-extraction alveolus healing (1=normal, 2=acute inflammation, 3=acute inflammation followed by infected alveolus and 4=alveolar osteitis). Three different surgeons evaluated 20 same patients independently. There was no significant difference considering their assessment (p<0.01; one way ANOVA).

Patients were also contacted on the fourteenth day after the procedure via their electronic mail or a telephone for additional questions regarding possible problems and for evaluation of their satisfaction with the results of the surgical procedure.

**Statistical methods**

Statistical analyses were performed using the SPSS Statistical package (IBM, Chicago, Illinois, USA). A descriptive statistic was made (frequencies, arithmetic means, and standard deviations). The tests used in this research were χ² test (Fisher’s test in cases when contingency was smaller than 5 samples), independent Student t-test and one-way analysis of variance (ANOVA) with post-hoc Scheffe’s tests.

Significance of the differences between participating groups (P1, P2 and C) regarding patient’s evaluation of pain, swelling problems and halitosis were tested with one-way variance analysis (post-hoc Scheffe).

Significance of difference in usage of analgesics between participating groups (P1, P2 and C) was tested by χ² test (Fisher’s test in cases when contingency was smaller than 5 samples). Significance of differences of clinical states between participating groups (P1, P2 and C) for the type of post-extraction alveolus healing was made by χ² test (Fisher’s test in cases when contingency was smaller than 5 samples).

**Results**

Results of postoperative healing evaluation showed...
there was a statistically significant difference in postoperative wound healing between three groups on third and seventh day after removal of lower third molars (day three \( \chi^2 = 26.02, p < 0.001 \), day seven \( \chi^2 = 18.13, p < 0.001 \), Fisher’s exact test).

On the first postoperative day there was a significant difference between the groups regarding pain levels (\( F = 6.17, p = 0.003 \)), swelling problems (\( F = 8.21, p < 0.001 \)) and halitosis (\( F = 4.22, p = 0.016 \)). The post hoc Scheffe’s tests showed that significantly less pain and halitosis was experienced by patients in the group P1 compared to other two groups. Moreover, the patients from the group C had significantly larger swelling problems when compared to patients from both laser groups.

On the third postoperative day there was also a significant difference between the groups considering pain intensity (\( F = 10.71, p < 0.001 \)), swelling problems (\( F = 10.67, p < 0.001 \)) and halitosis (\( F = 7.23, p = 0.001 \)). The post hoc Scheffe’s tests showed that significantly more pain was experienced by patients in the group C than in the laser groups. The patients in the group P1 experienced significantly less swelling problems and halitosis compared to other two groups.

On the seventh postoperative day there was still a significant difference between the groups in pain intensity (\( F = 12.41, p < 0.001 \)), swelling problems (\( F = 11.62, p < 0.001 \)) and halitosis (\( F = 17.37, p < 0.001 \)). The post hoc Scheffe’s tests showed that significantly more pain and halitosis was experienced by patients in the group C compared to laser groups. The patients in the group P1 and P2 experienced significantly less swelling problems compared to the group C.

On the fourteenth postoperative day, there was a significant difference between the groups regarding pain intensity (\( F = 7.01, p = 0.001 \)), swelling problems (\( F = 6.13, p = 0.003 \)) and halitosis (\( F = 8.02, p < 0.001 \)). The post hoc Scheffe’s test showed that C group had significantly higher pain intensity, swelling problems and halitosis than laser groups. Average grade of postoperative pain intensity (Figure 1), swelling problems (Figure 2) and halitosis (Figure 3) evaluated on the first, third, seventh and fourteenth day was the highest in the control group.

**Discussion**

The results of the present study showed positive

![Figure 1. The distribution of pain intensity between groups: C= control group, P1= APDT group, P2= LLLT group](image-url)
effects of low power diode laser therapy for all postoperative problems after the removal of lower third molars. The absence of complications in postoperative healing of patients in both laser groups, opposite to the control group, indicated the anti-inflammatory effect and enhanced defensive cell reaction of that
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treatment. Postoperative healing complications were seen in 36% of cases in the control group, most of which was attributed to alveolar osteitis. Gbotolorun et al. also showed that alveolar osteitis was the most frequent complication seen in 14.2% of their patients. Average grades for pain, swelling problems and halitosis decreased exponentially through 14 days of postoperative monitoring in all three groups, which was previously expected. But the group with the highest average grade drop, and with the lowest postoperative pain, swelling problems and halitosis intensity, was the P1 group. Such results, apart from many positive effects of laser therapy, could be explained by the anti-microbial effect of the Paro-PDT solution in combination with laser light used in group P1 therapy. The ability to destroy microorganisms by implementing the Paro-PDT solution into the cell wall and subsequent creation of free radicals has led to a strong anti-bacterial effect even in hard reachable places. Mattiell et al. showed in their study the anti-microbial effect of Paro-PDT solution and efficiency of photodynamic therapy on Aggregatibacter actinomycetemcomitans (Aa) and Streptococcus sanguinis (Ss). Photodynamic therapy has displayed statistically significant reduction of Aa and Ss occurrences with in vitro samples when compared with the control group. P2 group patients had not exhibited significantly lower levels of pain, swelling problems and halitosis on the first and third postoperative days when compared with the control group. On the seventh and fourteenth postoperative day, the intensity of the problems was reduced and equalled the intensity of postoperative problems reported by P1 group patients, which are significantly better results when compared with the results obtained from group C patients on the same days. Such results may be explained by the cumulative effect of laser therapy by which every newly applied postoperative dosage stays in the tissue. That way each new dosage had stronger effect on the tissue. Considering this, future research should be performed using everyday postoperative LLLT laser therapy. Cumulative effect would be increased and postoperative problems would be reduced. While doing so, it is mandatory to follow manufacturer’s instructions which stated that laser therapy for treating acute problems can be used until problems disappear, while it should not be used for longer than fourteen days postoperatively in case of chronic problems. Significantly lower pain intensity and reduced postoperative analgesics usage in laser therapy proved its complex analgesic effect. These results cannot be compared with results presented by other authors since most authors in most recent literature evaluated the effect of analgesics as premedication or their influence on long-lasting local anesthetics on postoperative problems.

Expert literature is expanding every day and various new data on the effect of various laser therapies on postoperative problems after the removal of lower third molars is constantly added. The difference in reports can be caused by the type of the laser used, laser beam wave length, radiation dosage and time of exposure. It was shown that laser radiation applied extraorally better reduces postoperative problems when compared with intraorally applied therapy.

Marković et al. showed that better postoperative swelling problems reduction after surgical removal of lower third molars is accomplished by a combination of laser therapy and local intramuscular application of dexamethasone, while the combination of laser therapy and systematic dexamethasone application did not show statistically significant effects compared with laser therapy. In the future, a standard laser therapy should be determined, a therapy in which numerous proved positive effects of diode laser on soft tissue should be applied, while all other variations of the standard therapy that would include medicament usage or other variation regarding therapy application should be proven in future researches.

Conclusion

The present study has approved the diode laser usage for significant enhancement of patient’s postoperative problems after third molar surgery. Both laser therapy modalities significantly reduced postoperative pain, swelling problems, halitosis and analgesic usage. The least postoperative problems were present in the group which received antimicrobial photodynamic (APDT) and low level laser therapy (LLLT).

Conflicts of Interest

The authors deny any conflicts of interest.

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