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Original Article

The Effectiveness of Micro-osteoperforations during Canine Retraction: A Three-dimensional Randomized Clinical Trial

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Aim: A major challenge in orthodontics is decreasing treatment time without compromising treatment outcome. The purpose of this split-mouth trial was 10 to evaluate micro-osteoperforations (MOPs) in accelerating orthodontic tooth 11 12 movement. Materials and Methods: Eight patients of both genders were selected, age ranging between 15 and 40 years, with Class II Division 1 malocclusion. The 13 participants in this trial with MOPs were randomly allocated to either the right 14 15 or the left side, distal to the maxillary canine. First maxillary premolars were extracted as part of the treatment plan on both sides and then canine retraction 16 was applied. Miniscrews were used to support anchorage. MOP side received 17 (three small perforations) placed on the buccal bone, distal to the maxillary 18 canine, on randomly selected side using an automated mini-implant driver and 19 the other side was the control side. Blinding was used at the data collection and 20 analysis stages. The primary outcome was the rate of canine retraction measured 21 22 with a three-dimensional (3D) digital model from the baseline to the first 2 weeks 23 superimposed at the rugae area from the baseline to the first, second, and third months. The following secondary outcomes were examined: anchorage loss, 24 canine tipping, canine rotation, root resorption, plaque index, and gingival index. 25 Pain level, pain interference with the patients' daily life, patients' satisfaction 26 27 with the procedure and degree of ease, willingness to repeat the procedure, and recommendation to others were also evaluated. Results: No statistically significant 28 difference was observed in the rates of tooth movement between the MOP and 29 the control sides at all-time points (first month: P = 0.77; mean difference, 0.2 30 mm; 95% CI, -0.13, 0.18 mm; second month: P = 0.50; mean difference, -0.0831 mm; 95% CI, -0.33, 0.16 mm; third month: P = 0.76; mean difference, -0.0532 33 mm; 95% CI, -0.40, 0.29 mm). There were also no differences in anchorage loss, rotation, tipping, root resorption, plaque index, periodontal index, and pain 34 perception between the MOP and control sides at any time point (P > 0.05). 35 MOPs had no effect on the patients' daily life except for a feeling of swelling 36 on the first day (P = 0.05). Level of satisfaction and degree of easiness of the 37 procedure were high. Conclusion: According to our clinical trial, MOPs cannot 38 help in speeding up a canine retraction. 39

Keywords: Accelerating tooth movement, cementoenamel junction, marginal bone crest, micro-osteoperforation

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INTRODUCTION AND LITERATURE REVIEW

L onger treatment periods on orthodontic treatment are associated with root resorption and periodontal disease are common problems, which we need to decrease, so a shorter treatment period is important, especially for older patients, who typically require a longer treatment span because their metabolic rate is much slower than younger patients.

The new era in orthodontics aims to accelerate tooth movement without using any harmful surgical procedures,^[1-3] according to the familiar standard theory of activation of the inflammatory process and osteoclasts.^[4] Acceleration method is needed to decrease the side effects of treatment such as root resorption, caries, periodontal inflammation, and patient follow-up times.^[5,6]

Tooth movement usually arises from the arranged activity of alveolar bone remodeling system (osteoclasts and osteoblasts). The applied orthodontic force on the tooth causes tension in the surrounding Periodontal ligaments and the alveolar bone, this results in bone resorption and formation of area of tension and pressure on tooth surfaces. The speed of tooth movement in orthodontic treatment is mainly affected by the remodeling process. ^[4,7]

In general, three common methods^[8] are used before speeding up the tooth movement, the first method is smearing physical stimulating substance to initiate bone anabolic and catabolic process, for example, low-level laser therapy,^[9,10] vibration,^[11] and AcceleDent System.^[12,13] These procedures are not the paths that are stimulated throughout the usual orthodontic tooth movement. Rather, these stimulant-activated processes activated cells that are responsible for resorption (osteoclasts) or bone formation (osteoblasts).^[7,14] The second method is using chemicals or medication,^[15] for example, an injection of prostaglandins, vitamin D3, and osteocalcin locally in the place of the required movement. The third approach encourages natural bone remodeling in ways that are activated with surgical, minimal surgical methods, and stimulants during orthodontic treatments (such as cortectomies,^[7,16] piezocision,^[3,7,17-19] and micro-osteoperforations [MOPs]^[2,7,20-22]). Because of the need for the development of the previous approach mentioned and for more trials of these methods, we present our study, to evaluate MOPs as one of the simplest and safest methods to accelerate orthodontic 49 treatment and to improve patients' acceptance for the 50 treatment duration, especially adults. In 2016, systematic 51 review^[23] showed that minimally invasive method can 52 accelerate tooth movement with high difference from 53 the first month of application. 54

All research results of previous trials showed that orderly simple distress from applying MOPs will accelerate the tooth movements^[14] except one study,^[24] which showed that MOPs did not add any increase in tooth movement during canine retraction. Almost all studies showed that there is root tipping without any mention of loss of anchorage in the presence of temporary anchorage device (TAD) and no root resorption. 1

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We hypothesize that harnessing and amplifying the body's natural inflammatory response to orthodontic movement using MOPs in alveolar bone would produce a minimally invasive, safe, and easily performed protocol to accelerate tooth movement.

In our study, we wanted to inspect the effectiveness of MOPs in the acceleration of tooth movement during canine retraction as a minimally invasive method by measuring the anteroposterior displacement of canine in both sides by using three-dimensional (3D) digital models in the planed time frame as primary outcome and to measure the root resorption and pain intensity during canine retraction as secondary outcome.

MATERIALS AND METHODS

This study was a split-mouth randomized clinical trial with a 1:1 allocation. The research was accepted by the stomatology department of Xi'an Jiaotong University Hospital in Xi'an City, Shaanxi Province, China. Ethical approval was obtained from the ethics committee with approval number, NXjkqII [2017] No.006. The research sample involved eight patients of ages ranging between 15 and 40 years. The main inclusion criteria of the study were patients with Class II Division 1 malocclusion who required the extraction of both maxillary first premolars with full erupted canine. Table 1 summarizes the rest of the inclusion and exclusion criteria of the study. Patients who agreed to participate in the experiments were asked to fill out and sign the agreement and the ethics form after clarifying the purpose of the intervention and the associated risks and benefits.

42 All participant were referred to the periodontics 43 department for checking the periodontal and oral 44 hygiene condition before starting the orthodontic 45 treatment. The orthodontic treatment was carried 46 out by the same orthodontic resident (BA) using fixed 47 orthodontic appliance (McLaughlin, Bennt and Trevisi 48 prescription and self-ligating bracket) with maximum 49 anchorage support using miniscrews, also direct 50 anchorage was used by applying the force directly 51 from minisrew to canine to prevent mesial movement 52 of posterior teeth during canine retraction. Indirect 53 anchorage was applied by ligating upper second 54

	Table 1: Inclusion and exclusion criteria
Clinical diagnosis	Class II
Age / gender	15–40 years old, both men and women
Oral examination	Good oral hygiene; adequate attached gingiva thickness (1-2mm); no periodontal disease
Bracket type	Both self-ligating brackets and non-self-ligating brackets, self-ligating is better
The treatment plan	The first maxillary bilateral premolars need to be extracted(the experiment is only limited to the upper
	jaw, so whether the mandibular first premolar is removed from the teeth and does not affect the inclusion
	of the patient); And implant anchorage is needed in the posterior area
Anamnesis	No history of previous orthodontic treatment; No systemic disease (which effects bone cycle)No history
	of periodontal disease treatment
X-ray examination	No radiographic bone loss
	on-smoking

premolars to minisrew to prevent mesial movements of posterior teeth.

The treatment strategy was as follows: the extraction of the maxillary first premolar, which was carried out at the beginning of the treatment and then canine retraction was performed after finishing the alignment and leveling stage.

RANDOMIZATION

The intervention was randomly allocated to either right or left side with 1:1 allocation ratio.

To increase the unpredictability of the random allocation sequence, the sequences with either right or left were concealed in opaque envelope and shuffled. Each participant was asked to pick one sealed envelope to assign to the intervention either on the right or left side. The patient's model was divided into two groups (MOPs side and control side), then the trial group gets MOPs on the right or left side, the aim of the non-determined side was to avoid unequal occlusal forces. Power arms were used to advance the retraction on the maxillary canine brackets. After the extractions and complete alignment stage, impressions and cone beam computed tomography (CBCT) were taken. The canine retraction was attained with calibrated 150-g nickel-titanium closing coil springs attached from the TAD to supporting power arms on the bracket of the canine.

INTERVENTIONS

The trial group obtained three perforations; the flapless method was performed on the left or right side in the middle of the extraction space using automated mini-implant instrumentation, each perforation was around 1.5–2mm width and 5–7mm depth [Figure 1A and B]. No antibiotic prescription was given; the nonexperimental side of the same arch was considered as the control group.

PRIMARY OUTCOME

The experimental procedure has five stages: before the experiment, after two weeks, after one month, two months, and three months of canine retraction. At every stage, impressions and photographs were taken to monitor the rate of tooth movement.

Model assessment: The patient's name, date, and the number of visits were labeled and stored. Forty cast models were scanned in a three shape scanner. All cast measurements were made by Geomagic program (engineer software form 3D system). After reference points were determined (1 midline point, 2 and 3 on the molar cusp, 4 and 5 tip of the buccal cusp of a second premolar, 6 and 7 tip of the buccal cusp of canine), lines were drawn to measure the difference of tooth movement [Figure 2]. The distance between line 5 and 6 (which represented the difference in distance of tooth movement between the control and experimental group) was calculated.^[18,25]

SECONDARY OUTCOMES

Cone beam computed tomography assessment

Before CBCT scanning, the patient was placed in a seated position with head upright position so that the intersection lines were straight horizontal and vertical through the center of the region of interest. All CBCT images were taken with the following parameters: 47 mA, 120 kVp, 250 mm voxel resolution, and 16-cm field of view.

Root resorption

The measurements of root resorption before and after the procedure were carried out by determining a point on the crown tip and another point on the apex tip of the root to determine the total canine length [Figure 3A].^[26,27] The height of the bone was specified by measuring the distance between cementoenamel junction and marginal bone crest from the buccal and lingual sides by drawing two lines between them^[28] [Figure 3B]. All measurements were calculated by one examiner, the reliability coefficient was used to evaluate the measurement twice weekly, showing excellent agreement.

Pain intensity

The participating patients were asked to evaluate their level of discomfort and pain on the day of the procedure as well as on 2, 7, and 28 days after the procedure.


Figure 1: (A) MOP instruments hand-assisted mini-implant. (B) Micro-osteoperforations procedure





A numeric rating scale short form of questionnaire (https://www.physiopedia.com/Shortform_McGill_Pain_ Questionnaire) was applied. The patients were instructed to select a digit on a scale (from 0 to 10) to describe their pain with 0 for "no pain" and 10 for "severe pain."

STATISTICAL ANALYSIS

Data were handled and analyzed using the Statistical Package for the Social Sciences (SPSS) software, version 21 (IBM, Armonk, New York). Owing to the small sample size, all statistical analyses were performed using nonparametric tests. Wilcoxon signed-rank test was used to compare different measurement on both sides. To compare an individual measurement in different time points, repeated measures analysis of variance (ANOVA) was applied. A P value of < 0.05 was considered statistically significant. Reliability coefficient was used to evaluate the reliability of primary outcomes measurements by one examiner. All participants were randomly selected, six of 3D superimposed digital models and canine displacement were measured twice within two-week intervals, the reliability coefficient was found to be 0.9, which showed excellent superimposing and measurement agreement.

RESULTS

MODEL CAST ANALYSIS

All statistical tests in different time points revealed no significant differences between MOPs and control sides except for the test after three months where the displacement movement of the canine was higher in MOPs compared to the control side (P = 0.002) [Table 2]. The repeated measures ANOVA for the difference in distance between any two-time points revealed significant differences for MOPs (P = 0.007) and control sides (P = 0.002). Table 3 presents the differences between time points that were statistically significant.

CONE BEAM COMPUTED TOMOGRAPHY ANALYSIS FOR ROOT RESORPTION

Table 4 presents the measured canine lengths at baseline and after 3 months of the study. Neither MOPs (measurement difference = 0.03 mm) nor control (measurement difference = 0.05 mm) showed a significant difference between the two measurements [Figure 4]. Comparing these differences between both groups did not reveal a statistically significant difference (P = 0.886) [Figure 5].



Figure 3: (A and B) Total tooth length: the distance between root apex and tip of the crown. (C and D) Cementoenamel Junction to Marginal Alveolar Bone Crest Distance

CONE BEAM COMPUTED TOMOGRAPHY ANALYSIS FOR BONE HEIGHT

The results did not show any differences between both sides (MOPs and control) with regard to the bone height whether buccally or palatally. Neither the MOP side (measurement difference = 0.11 and 0.23 mm, for buccal and palatal, respectively) nor the control side (measurement difference = 0.1 and 0.05 mm, for buccal and palatal, respectively) showed a significant difference between the two measurements. Comparing these differences between both groups did not reveal a statistically significant difference (P = 1.00 and 0.393, respectively) [Figure 6].

PAIN ASSESSMENT

After 48 h of MOP application and starting canine retraction, no signs of trauma were detected in the relevant sides or in the control group. The distribution of the pain categories between the two groups was exactly the same; the patients reported moderate pain in MOP and control groups (35.5% each) [Table 5].

DISCUSSION

Our study evaluated a new method of speeding up the time required for orthodontic treatment and of decreasing the associated pain. In orthodontics, a common scenario is when two interventions are applied in the same patient in a splitmouth approach. In this case, we have paired observations, and the required sample size is smaller because of the reduced variability. The split-mouth technique design resembles the crossover design, more often encountered in trials in medicine, without the period effects.^[29] In our working example, we are interested in evaluating space closure differences in Class II Division 1 maxillary premolar extraction patients by using nickel-titanium on one side and a nickel-titanium with MOPs by miniscrew on the contralateral side. This is a paired observation case, with the participant serving as the control, as both intervention and comparison treatments are applied in each patient. This design is more efficient because the sites that receive the interventions are similar, thus reducing variance and sample size requirements.

The result of our study shows that the mean values of the canine displacements were nonsignificant. Our results were in disagreement with the findings from previous clinical and laboratory trial,^[14] which showed that application of MOPs could increase the rate of canine retraction during orthodontic treatment by more than one to two times and was better than the conventional orthodontic treatment method. However, our result was

Table 2: Mea between car points for n	n, standard deviations (SD) of the distance ine and second premolar in different time icro-osteoperforations and control groups				
Side group	MO	Ps	Con	trol	P-value
	Mean	SD	Mean	SD	
Before Exp.	10.83	1.25	10.96	0.76	0.808
After 2 w	10.32	1.18	10.85	1.06	0.545

1.26

0.89

1.21

9.72

9.10

8.70

9.79

9.09

9.28

0.67

1.08

1.23

0.818

0.945

0.002*

After 2	m
After 3	m

After 1 m

*P<0.005

micro-osteoperforations in different time points					
(I) MOPs	(J) MOPs	Mean Difference	Std. Error	Sig.	
After 1 m	After 2 m	(I-J) 6225	58230	201	
AITCI I III	After 2 m	.0223	.38230	.29.	
	After 2 m	3987-	.38230	.51	
	Arter 5 III Before Exp	1.0223	.38230	.00	
After 2 m	After 1 m	-1.1100-	58230	200	
	After 2 w	0223-	58230	.29	
	After 3 m	-1.2212-	58230	0. /0/	
	Refore Exp	_1 7325_*	58230	.49	
Δ fter 2 w	After 1 m	-1.7323-	58230	.00	
IIII 2 W	After 2 m	1 2212*	58230	.51	
	After 3 m	1.6212*	58230	.00	
	Before Exp	- 5113-	58230	38	
After 3 m	After 1 m	-1 0225-	58230	.08	
inter 5 m	After 2 m	- 4000-	58230	49	
	After 2 w	-1.6212-*	.58230	.00	
	Before Exp	-2 1325-*	58230	00	
Before Exp	After 1 m	1.1100	.58230	.06	
Serore Enpi	After 2 m	1.7325*	.58230	.00	
	After 2 w	.5113	.58230	.38	
	After 3 m	2.1325*	.58230	.00	

in agreement with a study published in 2018 that failed to show any significant difference for using MOPs during canine retraction over the conventional method.^[24]

It is well known that the timing of treatment and the age of patients have a significant role in orthodontic treatment. To minimize the influence of age on the rate of tooth movement and to get almost similar outcome during the study, we chose adults between 15 and 40 years. In this age range, the development is almost or completely finished.^[30] Gender of the patients is also another difficult factor that affects the treatment results due to the hormone implications, which is different among males and females. Similar to most of the studies, our study could not decrease this problem due to the limitation in patients' numbers.^[31,32] Occlusion factor was excluded from this

Table 4: Multiple comparisons using post hoc test of									
control group in different time points									
Dependent Variable: control side									
(I)	(J) Mean Std. Error S								
CONTROL	CONTROL	Difference (I-J)							
After 1 m	After 2 m	.6225	.58230	.292					
	After 2 w	5987-	.58230	.311					
	After 3 m	1.0225	.58230	.088					
	Before Exp.	-1.1100-	.58230	.065					
After 2 m	After 1 m	6225-	.58230	.292					
	After 2 w	-1.2212-*	.58230	.043					
	After 3 m	.4000	.58230	.497					
	Before Exp.	-1.7325-*	.58230	.005					
After 2 w	After 1 m	.5987	.58230	.311					
	After 2 m	1.2212*	.58230	.043					
	After 3 m	1.6212*	.58230	.009					
	Before Exp.	5113-	.58230	.386					
After 3 m	After 1 m	-1.0225-	.58230	.088					
	After 2 m	4000-	.58230	.497					
	After 2 w	-1.6212-*	.58230	.009					
	Before Exp.	-2.1325-*	.58230	.001					
Before Exp.	After 1 m	1.1100	.58230	.065					
	After 2 m	1.7325*	.58230	.005					
	After 2 w	.5113	.58230	.386					
	After 3 m	2.1325*	.58230	.001					

*P<0.005



Figure 4: (A) Position of canine before experiment, (B) Position of canine after one month

study for its effects on the speeding tooth movement.^[33] Any defect related to the bite or unequal occlusion was excluded. Many studies tested the occlusion interfering with canine destabilization in Class II Division 1 case. Because the canine is free from arch bite association, canine retraction is another possible reason that might affect the rate of the orthodontic movement.

All diseases related to the periodontal ligament or patients with bad oral health were excluded to remove any probability of inflammation from any cause other than the MOPS.^[29,34] So to make all these variables less and to reduce their effect in the treatment procedures, certain criteria have been selected [Table 1].

Many studies reported the effects of extraction timing on tooth movement. Some of them recommended 





Figure 6: illustrating the differences in Bone height of the buccal and palatal side in MOPs and control groups

Table 5: Before and after root resorption multiple comparison using post hoc test						
Side group	MOPs		Control		<i>P</i> -value	
	Mean	SD	Mean	SD		
Before	22.71	.71	22.76	1.06	0.842	
After	22.68	.71	22.71	1.14	0.886	
P-value	0.934	0.929				

delaying extraction until retraction or after six months.To make sure that the extraction did not affect the study results, we performed it at the beginning of the orthodontic treatment.

Pain and discomfort recorded in both sides appear to be exactly similar, meaning that pain is not a considerable concern with MOPs. Likewise, almost all patients were willing to redo the treatment to accelerate the orthodontic treatments when they were asked about that [Table 6].

There are many variables that lead to root resorption,
so if the study took a long time, it would have been
more challenging to control these variables. We took
the CBCT before and three months after the retraction.
We assessed the root resorption and bone height.

CONCLUSION

MOP modality is a less aggressive and safe procedure, but it does not significantly speed up the tooth movement during orthodontic treatment. Further well-designed studies with larger sample size for longer treatment time are highly recommended.

ETHICAL CONSIDERATIONS

All the procedures have been performed as per the ethical guidelines laid down by Declaration of Helsinki and approved by the review board of stomatology department ethics committee of College of Medicine, Xi'an Jiaotong University, Xi'an, Shaanxi Province, China (approval number: Xjkq11 2017, dated: January 7, 2017, with number of trial: 06).

DATA AVAILABILITY

Datasets related to this article will be available from the corresponding author (Xia HY, Email: 369 hyx@163. com) on request.

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DECLARATION OF PATIENT CONSENT

The authors certify that they have obtained all appropriate patient consent forms. In the form the patient(s) has/have given his/her/their consent for his/ her/their images and other clinical information to be reported in the journal. The patients understand that their names and initials will not be published and due efforts will be made to conceal their identity, but anonymity cannot be guaranteed.

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CONFLICTS OF INTEREST

There are no conflicts of interest.

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MOPs pain%	Mild		Moderate		Severe
Experiments side	62.50%		37.50%		0.00%
Control side	62.50% 37.50%			0.00%	
Table (5-A): Pain percentage calculated from the visual pain scalefrom 1–10, mild (1–3), moderate (4–6) and sever (7–10)					
MOPs pain Q	А	В	С	D	Patient No
Experimental effects on daily life?	5	2	1		8
Is the pain compared to extraction pain?	7	1			8
Is the pain compared with TAD pain?	8				8
Are you willing to do the MOPs again to accelerate your treatment?	7	1			8

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