

## Insight into orthodontic appliance induced pain: Mechanism, duration and management

Yasemin Kartal, Omur Polat-Ozsoy

Yasemin Kartal, Omur Polat-Ozsoy, Department of Orthodontics, Baskent University, Faculty of Dentistry, 06500 Ankara, Turkey

**Author contributions:** Kartal Y wrote the manuscript; Polat-Ozsoy P helped writing the manuscript and edited the manuscript.

**Conflict-of-interest statement:** Authors deny any conflict of interest.

**Open-Access:** This article is an open-access article which was selected by an in-house editor and fully peer-reviewed by external reviewers. It is distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited and the use is non-commercial. See: <http://creativecommons.org/licenses/by-nc/4.0/>

**Correspondence to:** Omur Polat-Ozsoy, Professor, Department of Orthodontics, Baskent University, Faculty of Dentistry, 1<sup>st</sup> Street, Bahçelievler No.107, 06500 Ankara, Turkey. [omurorto@yahoo.com](mailto:omurorto@yahoo.com)  
Telephone: +90-31-22151336  
Fax: +90-31-22152962

Received: May 14, 2015

Peer-review started: May 19, 2015

First decision: September 2, 2015

Revised: November 4, 2015

Accepted: November 24, 2015

Article in press: November 25, 2015

Published online: March 27, 2016

### Abstract

Most of the orthodontic patients experience pain during treatment and this significantly influences their attitudes and the approach towards treatment. A number of factors that influence pain response include age, gender, personal pain threshold, mood and stress level of the person, cultural differences and types of orthodontic

treatment. Pain is a often overlooked subject by orthodontists, it is nevertheless important to understand the source and mechanism of the pain that occurs during treatment, as well as the methods for managing and controlling this pain. This review attempts to overview the mechanism, duration and current management strategies of orthodontic treatment.

**Key words:** Orthodontic appliance; Pain mechanism; Orthodontic treatment; Pain; Pain management

© **The Author(s) 2016.** Published by Baishideng Publishing Group Inc. All rights reserved.

**Core tip:** Pain during orthodontic treatment is an important concern for both clinicians and patients. Although it is not possible to completely eliminate pain during orthodontic treatment, it is still necessary to understand its causes and to minimize it to the greatest extent possible.

Kartal Y, Polat-Ozsoy O. Insight into orthodontic appliance induced pain: Mechanism, duration and management. *World J Anesthesiol* 2016; 5(1): 28-35 Available from: URL: <http://www.wjgnet.com/2218-6182/full/v5/i1/28.htm> DOI: <http://dx.doi.org/10.5313/wja.v5.i1.28>

### INTRODUCTION

Pain is a commonly encountered sensation in daily human life that is usually difficult to describe or diagnose, and which often represents an important problem that must be addressed through a multidisciplinary approach encompassing all branches of medicine.

Pain during orthodontic treatment is an important concern for both clinicians and patients<sup>[1,2]</sup>. Patient motivation and cooperation is an important factor in orthodontic treatment, while pain significantly influences

patient attitudes and the approach towards treatment. Studies indicate that 90% of orthodontic patients experience pain during treatment, and that 30% consider discontinuing or interrupting their treatment due to pain<sup>[3,4]</sup>.

The study of Abu Alhaijaa *et al.*<sup>[5]</sup> evaluated the relationship between personal characteristics, expectation of pain, and treatment compliance, reporting that individuals who experienced less pain during treatment generally displayed a more positive attitude, and that those sufficiently informed about treatment procedures had less expectations of pain.

In another study, 95% of the patients reported that they experienced pain in different stages of their treatment, and that this inevitably affected their diet<sup>[6]</sup>.

For these reasons, although pain is a subject that is often overlooked by orthodontists, it is nevertheless important to understand the source and mechanism of the pain that occurs during treatment, as well as the methods for managing and controlling this pain.

### **Mechanism of pain**

The International Association for the Study of Pain defines pain as "an unpleasant sensory and emotional experience associated with actual or potential tissue damage, or described in terms of such damage". Pain also has a strong motivational component, in that it not only triggers a withdrawal reflex, but also induces a highly organized avoidance and evasive behavior<sup>[7]</sup>. The motivational aspect of pain is an essential function, without which it would be difficult - if not impossible - for the human body to protect and sustain itself<sup>[7]</sup>.

In orthodontic treatments, the force transmitted by appliances allows the movement of the teeth within the alveolar bone<sup>[8]</sup>. However, this movement also has the effect of causing the compression and inflammation of the blood vessels and nerves within the periodontium. The perception of orthodontic pain is associated with changes in blood flow that occur due to inflammatory reactions following the application of force<sup>[9]</sup>. Studies indicate that periodontal pain consists of a combination of pressures, ischemia, inflammation, and oedema<sup>[10]</sup>. Davidovich and Shanfeld have reported that the application of force leads to acute inflammation, which, in turn, results in periodontal vasodilation and the sensation of pain<sup>[11]</sup>. It is known that the development of hyperalgesic resistance is associated with the release of various chemical mediators<sup>[12,13]</sup>. Studies have shown that the chemical mediators involved in the development of the hyperalgesic response include histamine, substance P, enkephalin, dopamine, serotonin, glycine, glutamate gamma-amino butyric acid, prostaglandins (PGs), leukotriene, and cytokines<sup>[9,12,13]</sup>. The studies in the literature concerning the increase in the level of these mediators have also demonstrated that the hyperalgesic response occurs following the application of force<sup>[9,14,15]</sup>. Recent studies have investigated the molecular basis of orthodontic pain by evaluating subjects such as the elevation in the level of various neuropeptides<sup>[9]</sup>.

Kato *et al.*<sup>[16]</sup> previously investigated in rats the distribution of the neurofibrils within the PDL [such as the neurofilament protein (NFP), calcitonin gene-related peptide (CGRP), vasoactive intestinal polypeptide and neuropeptides Y] following the application of force on the first molar. Three days after the application of force, they observed that the level of neurofibrils consisting of NFP and CGRP increased in both the compressed and strained sides, and that these levels returned to normal on the 14<sup>th</sup> day<sup>[9,16]</sup>.

Studies indicate that substance P - a sensory neuropeptide released from the peripheral nerve ends - and CGRP both regulate the secretion of proinflammatory cytokines released by monocytes, such as IL-1 $\beta$ , IL-6, and TNF- $\alpha$ <sup>[15,17,18]</sup>. Yamaguchi *et al.*<sup>[19]</sup> determined that the level of three major cytokines (IL-6, IL-8, and TNF- $\alpha$ ) released from the human dental pulp cells increased significantly in the 12 h following the application of mechanical force. They also reported that major neuropeptides, such as proinflammatory cytokines, might be involved in pulpal inflammation during orthodontic teeth movement.

### **Duration of pain**

The time of onset and duration of orthodontic pain was similar in most studies, with patients generally beginning to experience discomfort four hours after the application of orthodontic force<sup>[20]</sup>. In a study using the Visual analogue scale (VAS) to evaluate the level of pain that developed following the placement of separators, the highest intensity of pain was observed on the second day, while the pain fully subsided by the fifth day<sup>[21]</sup>. Nearly half of the patients evaluated during this study were compelled to change their diet habits and to use analgesics.

In a study using the VAS to evaluate pain in patients with arch wire and separators placed between their molars, Wilson *et al.*<sup>[20]</sup> reported that pain generally began four hours after the application of force, reaching its highest level 24 h later, and almost fully disappearing by the seventh day. On the other hand, Tuncer *et al.*<sup>[22]</sup> described that pain began two hours following the application of orthodontic elastics, reaching its highest level six hours later, and almost fully disappearing by the second day.

A previous study reported that although pain ended in most patients on the seventh day following the application of orthodontic force, 25% of the patients still continued to experience a certain level of pain<sup>[23]</sup>. The results of the said study indicated that orthodontic pain began two to six hours following the application of orthodontic pain, reaching its maximum level within the first two days, and then gradually decreasing until it completely disappeared by the seventh day.

## **FACTORS AFFECTING ORTHODONTIC PAIN**

Pain is a subjective finding, and different individuals may

display different pain responses to the same stimulus. There are a number of factors that are responsible for these differences in response. The main factors that influence pain response include age, gender, personal pain threshold, mood and stress level of the person, as well as cultural differences and the person's previous pain experiences<sup>[1,4,9,24,25]</sup>.

### Age

As orthodontic treatments generally involve different therapeutic procedures for different age groups, making comparisons regarding the effect of age is difficult, with studies on this subject generally providing somewhat contradictory results. However, Ngan *et al.*<sup>[26]</sup> previously reported that there were no statistically significant differences between adolescents and adults regards to pain. On the other hand, in their comprehensive and large-scale study on pre-adolescents, adolescents, and adults, Moerenhout and Brown<sup>[27]</sup> reported that adolescents exhibited higher levels of pain.

In recent years, there is a growing consensus that the relationship between pain and age should be evaluated by also taking into account the effect of age, since this relationship appears to be particularly affected by adolescence. Sandhu and Sandhu<sup>[28]</sup> determined in their study that girls between the ages of 14 and 17 experienced the highest levels of pain during orthodontic treatments. These authors emphasized that due to their synergistic interaction, the effects of age and gender on the level of pain during orthodontic treatment should be evaluated together rather than separately<sup>[28]</sup>.

These contradictory and conflicting results appear to stem not only from the fact that different orthodontic treatment methods are generally used for different age groups, but also from the fact pain is a multifactorial element that can be affected by gender differences, as well as the psychological and emotional state of the patients.

### Gender

Similar to age, gender is another factor that is unlikely to provide accurate assessments when used independently to evaluate pain. This is because even within the same gender, factors such as age group and cultural differences can significantly affect the level of pain that is experienced. Certain studies report that while there are no statistical differences between males and females within the 11-14 age group, a significant difference begins to be observed within the 14-17 age group. This change is reported to be associated with the hormonal changes experienced by females during adolescence<sup>[28,29]</sup>. Cultural differences similarly appear to cause significant variations in study results regarding the relationship between pain and gender. In a study evaluating the pain response of both male and female individuals, it was observed females generally found it easier to express and describe the pain they experienced compared to males<sup>[30]</sup>. These contradictory

results indicate that the perception of pain is affected not only by physiological differences, but also by cultural factors<sup>[31]</sup>.

Although certain studies evaluating the effect of gender on orthodontic pain describe that females exhibit higher levels of pain than males<sup>[6,32,33]</sup>, most studies from the orthodontics literature have not identified a gender-related difference in the perception of pain<sup>[34-37]</sup>.

### Emotional state

Dental anxiety ranks fifth among the objects and situations that are the most common sources of anxiety<sup>[38]</sup>. A study conducted by Hamurcu<sup>[39]</sup> compared the intensity of pain experienced with their level of anxiety, and determined that patients exhibiting higher anxiety scores also experienced more pain.

In a study comparing the level of pain experienced by patients at the beginning of orthodontic treatment with their personal characteristics, Bergius *et al.*<sup>[40]</sup> determined that individuals with dental anxiety experienced higher intensities of pain. A similar study observed that anxiety reduced the pain threshold, causing patients to perceive even the simplest procedures as painful<sup>[41]</sup>.

---

## THE EFFECT OF THE TYPE OF ORTHODONTIC TREATMENT ON PAIN

---

### Orthodontic separation

Orthodontic separation is a method applied prior to the placement of an orthodontic band, and is usually associated with significant pain for the patients<sup>[9,21,26,42]</sup>. In another study performed on 55 patients, 87% of the patients described pain following the placement of an orthodontic separator, while 72% required analgesics<sup>[37]</sup>. In a study evaluating motor and sensory changes following the placement of a separator by using an electromyograph (EMG), Michelotti *et al.*<sup>[43]</sup> observed a decrease in the pain threshold and motor output of the chewing muscles, and suggested that this was a protective mechanism to prevent further damage to the injured area.

### Dental archwire placement and activation

The pain that develops following the initial placement of an archwire has been the subject of numerous studies. These studies generally report that most patients begin to experience pain four hours after the application of the arch, with the level of pain reaching its peak within the first 24 h, and then gradually decreasing<sup>[6,26,32,34,42,44,45]</sup>.

No statistically significant differences have been identified between the perception of pain and the intensity, prevalence, and duration of the archwire usage<sup>[34,46,47]</sup>. In a study comparing the super-elastic nickel titanium wires with helical stainless steel wires, Sandhu *et al.*<sup>[48]</sup> reported no statistically significant differences in the level of pain experienced with these two wires. However, they suggested that the super-elastic wire caused more pain during the hours when the level of pain reached

its peak (between the 12<sup>th</sup> and 24<sup>th</sup> h), and that was probably due to the greater force applied by this type of wire. Although the current literature indicates that the application of either strong or weak forces by the wires does not lead to a significant difference in terms of the resulting level of pain, Sandhu's study nevertheless suggests that higher forces result in higher IL-1 beta concentrations, and that this engenders a difference in the level of pain observed during the peak period<sup>[48]</sup>. Ogura *et al*<sup>[49]</sup> similarly performed comparisons between weak and strong forces, and determined that during the period of maximum pain levels, biting while the teeth were exposed to stronger forces lead to higher levels of pain.

Previous studies evaluating the activity of chewing muscles following arch activation by using EMG identified a decrease in the masseter muscle activity, which is believed to be responsible for the reflex mechanism for avoiding harmful stimuli<sup>[50-52]</sup>. Murdock *et al*<sup>[44]</sup> and Erdinç *et al*<sup>[34]</sup> have reported that patients report greater pain in the posterior teeth than their anterior teeth during the leveling stage and chewing. In sum, most studies indicate that arch placement and activation can lead to pain, and adversely affect the daily activity and diet habits of patients<sup>[9]</sup>.

### Type of appliance

The level of pain caused by different types of appliances during orthodontic treatments has been evaluated in many studies. In studies comparing fixed and removable appliances, Oliver and Knapman<sup>[1]</sup> identified no significant differences between these two types of treatments, while Sergl *et al*<sup>[25]</sup> and Gianelly *et al*<sup>[53]</sup> reported that treatment with fixed appliances resulted in greater pain.

Various comparisons have been performed between fixed orthodontic treatments applied using different methods. Wu *et al*<sup>[54]</sup> and Caniklioglu *et al*<sup>[55]</sup> have performed comparisons between labial and lingual appliances, and reported no statistically significant difference with regards to the total level of perceived pain associated with these appliances. However, they also described greater pain on the tongue among patients who received lingual appliances, as well as greater pain on the lips and cheeks among patients who received labial appliances<sup>[54,55]</sup>. A recent study compared the application of a fixed labial appliances with the Invisalign® and determined that Invisalign® caused less pain<sup>[56]</sup>.

Shalish *et al*<sup>[57]</sup> have evaluated and compared the fixed lingual treatment, fixed labial treatment and Invisalign® treatment in adult patients, and determined that the most pain and general oral dysfunction occurred in the lingual apparatus group; that Invisalign® caused significant pain in the first day of treatment; and that Invisalign® was similar to conventional labial techniques in terms of general oral dysfunction<sup>[57]</sup>.

Bertl *et al*<sup>[58]</sup> have examined self-ligating brackets and

conventional brackets with respect to pain, and determined that self-ligating brackets caused significantly more pain. In contrast to Bertl *et al*<sup>[58]</sup>, Tecco *et al*<sup>[59]</sup> suggested that conventional brackets lead to stronger and more persistent pain, while self-ligating brackets tended to cause pain mainly during chewing and biting.

### Orthopedic forces

The main purpose of craniofacial orthopedics is to bring skeletal changes by applying significant forces to the craniofacial complex. Various publications report the occurrence of pain during rapid palatal expansion applied for the transversal skeletal development of the maxilla<sup>[60-62]</sup>. In such cases, patients generally describe a sensation of pain spreading across the craniofacial area<sup>[9]</sup>.

Headgear applications represent another treatment method used during the development stages of children to bring about skeletal and dental modification. Studies have demonstrated that patients with such appliances generally begin to experience pain approximately 24 h after initial application, with the sensation of pain and discomfort gradually decreasing after the 3<sup>rd</sup> day<sup>[63,64]</sup>.

Egolf *et al*<sup>[65]</sup> have reported that nearly 28% of patients using orthodontic elastics and headgear discontinue to wear them due to pain. Ngan *et al*<sup>[64]</sup> previously examined the chewing muscles of protraction headgear patients by using EMG, and determined that the pain associated with the orthopedic devices originated not from the muscle tissues, but instead from the acute inflammation caused by the accumulation of forces in the sutural areas.

### Skeletal anchorage systems

Recently, skeletal anchorage systems are being used for absolute anchorage. These devices can be grouped/classified as "mini-plate" and "mini-screws". Zawawi<sup>[66]</sup> reported that patients with mini-screw implants reported significantly less pain; that 32.5% of patients receiving mini-screw implants did not require any medication; that 59.1% of these patients only required a single-dose analgesic; and that patients generally preferred mini-screws instead of extraction.

Kuroda *et al*<sup>[67]</sup> previously compared the level of pain experienced with mini-plate and mini-screws. No significant differences were observed in terms of perceived pain levels between mini-plates and mini-screws inserted through incisions, while a significant difference was observed when mini-plates and mini-screws were implanted without using incisions, with the mini-plates resulting in noticeably more pain<sup>[67]</sup>. In agreement with Kuroda *et al*<sup>[67]</sup>'s findings, Kawaguchi *et al*<sup>[68]</sup> demonstrated that implanting mini-plates without using incisions resulted in three times greater pain than placing mini-screws without incisions. The abovementioned studies have generally suggested that the main causes of pain during the application of skeletal anchorage systems could mainly be associated



with sutures, periosteal separation, and incisions.

### Debonding

Many patients also describe pain when removing their fixed appliances. Various studies have shown that applying intrusive forces during the debonding of fixed appliances reduced the level of pain experienced. These studies have therefore recommended applying finger pressure, biting a cotton roll, or using an occlusal wax layer during the removal process in order to reduce pain<sup>[69,70]</sup>.

## PAIN MEASUREMENT

It is important that pain is measured by the use of standardized pain scales and by using common language due to its complex and subjective nature. Unfortunately, objective assessment methods are still developing and subjective assessment is still the commonly used method. As the pain perception varies among individuals, it is important to take the patients' own report into consideration. Ideally, a pain intensity scale must have a low rate of incorrect responses, should be easy to administer, and be sensitive with an adequate number of response categories and be statistically powerful to detect treatment effects.

VAS is considered to be superior to other pain scales in terms of reproducibility and ease of measurement. VAS is a numeric scale and consists of a horizontal or vertical 100 mm line that has "no pain" and "worst pain" labels on two endpoints; the patient is asked to mark on the line to show the degree of pain experienced. The distance between the low end of the scale and the patient's mark is used as the index of pain intensity<sup>[8]</sup>.

## MANAGEMENT AND CONTROL OF ORTHODONTIC PAIN

Although it is not possible to completely eliminate pain during orthodontic treatment, it is still necessary to understand its causes and to minimize it to the greatest extent possible. It is therefore important to take into consideration and avoid overlooking the patient's complaints during the treatment process, and to inform them beforehand about the pain the treatment may cause. A study performed by Krukemeyer *et al.*<sup>[4]</sup> determined that orthodontists tend to ignore or dismiss the pain caused by the treatment, and that they generally expect a lower level of pain and medication use than the level reported by patients. Krukemeyer *et al.*<sup>[4]</sup> also reported a general lower-than-necessary amount of medication use. Abu Alhaijaa *et al.*<sup>[5]</sup>, on the other hand, reported that patients sufficiently informed about the treatment process had a lower medication requirements.

Nonsteroidal anti-inflammatory drugs (NSAIDs) are usually the medication of choice in orthodontics to alleviate mild and moderate pain and inflammation,

although there is no standard protocol concerning the application of NSAIDs. Many drugs such as acetaminophen, ibuprofen, aspirin, and flurbiprofen have been used and determined to be effective in the management of orthodontic pain<sup>[71-76]</sup>. However, a number of previous studies have suggested that PGs, and especially prostaglandin E2 and prostaglandin E1, can affect bone remodeling and teeth movement<sup>[14,77-80]</sup>. Nevertheless, the general consensus in orthodontic pain management is that the application of a low-dose analgesic during the first days of treatment will not have a clinically significant effect on the movement of the teeth. Another point that needs to be taken into consideration during orthodontic treatment is the possibility that teeth movement might be affected in patients who have been regularly receiving NSAIDs for a long period of time due to a systemic condition. In such cases, acetaminophen should be preferred because it provides sufficient analgesia without affecting teeth movement<sup>[8]</sup>.

In recent times, there has been an increasing focus on preventing the development of a pain memory through preemptive drug administration. Steen Law *et al.*<sup>[76]</sup> previously assessed the effect of ibuprofen and placebo administered one hour prior to separator application, and determined that the ibuprofen administration significantly reduced the pain experienced by the patients. Polat and Karaman<sup>[72]</sup> similarly conducted a comprehensive study evaluating the administration of five different medication (placebo, ibuprofen, flurbiprofen, acetaminophen, naproxen sodium, and aspirin) one hour before and six hours after bracketing procedures. The lowest pain scores were observed in the naproxen sodium and aspirin groups, while the highest pain scores were observed in the acetaminophen group<sup>[72]</sup>. In another study of the same authors, a single preoperative dose of placebo, ibuprofen and naproxen sodium was applied, and - in agreement with the findings of their previous study - lower levels of pain were reported during the first day in the naproxen sodium group. However, the authors also described that a single-dose application was not sufficient, and that additional postoperative doses were also necessary<sup>[42]</sup>.

Non-pharmacological methods used for pain management include transcutaneous electrical nerve stimulation (TENS), laser applications, vibration, and chewing apparatuses. Profitt described that the use of chewing gum or biting blocks during application would help reduce pain<sup>[81]</sup>. This theory was investigated by Mohri *et al.*<sup>[82]</sup> by evaluating the relationship between chewing and the serotonergic (5-HT) neurons responsible for nociceptive transmissions. Mohri *et al.*<sup>[82]</sup> determined that the rhythmic behavior of chewing indeed suppressed the nociceptive response. Hwang *et al.*<sup>[83]</sup> similarly determined that biting blocks reduced pain in 56% of their patients; however, they also observed that in other patients, biting blocks had the effect of increasing the experienced pain. Murdock *et al.*<sup>[44]</sup>, on the other hand, compared the effect of analgesics and

biting blocks, and determined that these apparatuses were as effective as analgesics, and that they represent a good option for adolescents.

Laser - a highly popular technological application in recent times - is also being used in the management of orthodontic pain. Fujiyama *et al.*<sup>[84]</sup> reported that CO<sub>2</sub> laser applications are able to reduce orthodontic pain without affecting teeth movement. In another study, comparisons were performed between a low-energy gallium-arsenic-aluminum laser (LLLT) group, a placebo group, and a control group following the implantation of an arch wire. The study determined that the LLLT and placebo groups both experienced significantly less pain, although the difference between these two groups was not significant<sup>[85]</sup>. Although numerous alternative, non-pharmacological methods are being used for the management of orthodontic pain, it is known that pharmacological methods still represent the most effective approach.

## CONCLUSION

Although it is not possible to completely eliminate pain during orthodontic treatment, it is still necessary to understand its causes and to minimize it to the greatest extent possible. It is therefore important to take into consideration and avoid overlooking the patient's complaints during the treatment process, and to inform them beforehand about the pain the treatment may cause. During pain management, medication that ensures the maximum reduction of pain with the minimum side effects should be administered by employing the most effective methods. In particular, the decision regarding the choice of medication or approach for reducing pain should not be left to the patient's relatives. Although there are no controlled studies supporting low-energy laser and TENS applications, further studies and growing interest on these techniques might eventually bring a new dimension to orthodontic pain management.

## REFERENCES

- 1 Oliver RG, Knapman YM. Attitudes to orthodontic treatment. *Br J Orthod* 1985; **12**: 179-188 [PMID: 3863673 DOI: 10.1179/bjo.12.4.179]
- 2 Kluemper GT, Hiser DG, Rayens MK, Jay MJ. Efficacy of a wax containing benzocaine in the relief of oral mucosal pain caused by orthodontic appliances. *Am J Orthod Dentofacial Orthop* 2002; **122**: 359-365 [PMID: 12411880 DOI: 10.1067/mod.2002.126405]
- 3 Otasevic M, Naini FB, Gill DS, Lee RT. Prospective randomized clinical trial comparing the effects of a masticatory bite wafer and avoidance of hard food on pain associated with initial orthodontic tooth movement. *Am J Orthod Dentofacial Orthop* 2006; **130**: 6.e9-6.e15 [PMID: 16849064 DOI: 10.1016/j.jado.2005.11.033]
- 4 Krukemeyer AM, Arruda AO, Inglehart MR. Pain and orthodontic treatment. *Angle Orthod* 2009; **79**: 1175-1181 [PMID: 19852612 DOI: 10.2319/121308-632R.1]
- 5 Abu Alhaija ES, Aldaikki A, Al-Omairi MK, Al-Khateeb SN. The relationship between personality traits, pain perception and attitude toward orthodontic treatment. *Angle Orthod* 2010; **80**: 1141-1149 [PMID: 20677967 DOI: 10.2319/012710-59.1]
- 6 Scheurer PA, Firestone AR, Bürgin WB. Perception of pain as a result of orthodontic treatment with fixed appliances. *Eur J Orthod* 1996; **18**: 349-357 [PMID: 8921656 DOI: 10.1093/ejo/18.4.349]
- 7 McNeil C, Dubner R, Woda A. What is pain and how do we classify orofacial pain? In: Sessle BJ, Lavigne GJ, Lund JP, Dubner R. Orofacial Pain. From Basic Science to Clinical Management. Second Edition. Quintessence Publishing Co, 2008: 3-11
- 8 Polat O. Pain and discomfort after orthodontic appointments. *Semin Orthod* 2007; **13**: 292-300 [DOI: 10.1053/j.sodo.2007.08.010]
- 9 Krishnan V. Orthodontic pain: from causes to management--a review. *Eur J Orthod* 2007; **29**: 170-179 [PMID: 17488999 DOI: 10.1093/ejo/cjl081]
- 10 Furstman L, Bernick S. Clinical considerations of the periodontium. *Am J Orthod* 1972; **61**: 138-155 [PMID: 4500502 DOI: 10.1016/002-9416(72)90092-9]
- 11 Stanfeld J, Jones J, Laster L, Davidovitch Z. Biochemical aspects of orthodontic tooth movement. I. Cyclic nucleotide and prostaglandin concentrations in tissues surrounding orthodontically treated teeth in vivo. *Am J Orthod Dentofacial Orthop* 1986; **90**: 139-148 [PMID: 3017094 DOI: 10.1016/0889-5406(86)90046-6]
- 12 Erdine S. Ağrı mekanizmaları giriş. İçinde: Özcan İ (Ed). Ağrı: Baş-Boyun ve orofasiyal ağrılar. İstanbul: Nobel Tıp Kitabevleri, 2000; 17-32
- 13 Aldemir T. Akut Ağrı Fizyopatolojisi. In: Özcan İ. Ağrı Bas-Boyun ve Orofasiyal Ağrılar. 2nd ed İstanbul, Nobel Tıp Kitabevleri, 2006: 43-48
- 14 Yamasaki K, Shibata Y, Imai S, Tani Y, Shibasaki Y, Fukuhara T. Clinical application of prostaglandin E1 (PGE1) upon orthodontic tooth movement. *Am J Orthod* 1984; **85**: 508-518 [PMID: 6587784 DOI: 10.1016/0002-9416(84)90091-5]
- 15 Alhashimi N, Frithiof L, Brudvik P, Bakhiet M. Orthodontic tooth movement and de novo synthesis of proinflammatory cytokines. *Am J Orthod Dentofacial Orthop* 2001; **119**: 307-312 [PMID: 11244425 DOI: 10.1067/mod.2001.110809]
- 16 Kato J, Wakisaka S, Kurisu K. Immunohistochemical changes in the distribution of nerve fibers in the periodontal ligament during an experimental tooth movement of the rat molar. *Acta Anat (Basel)* 1996; **157**: 53-62 [PMID: 9096742 DOI: 10.1159/000147866]
- 17 Nicolay OF, Davidovitch Z, Shanfeld JL, Alley K. Substance P immunoreactivity in periodontal tissues during orthodontic tooth movement. *Bone Miner* 1990; **11**: 19-29 [PMID: 1702686 DOI: 10.1016/0169-6009(90)90012-5]
- 18 Norevall LI, Forsgren S, Matsson L. Expression of neuropeptides (CGRP, substance P) during and after orthodontic tooth movement in the rat. *Eur J Orthod* 1995; **17**: 311-325 [PMID: 8521925 DOI: 10.1093/ejo/17.4.311]
- 19 Yamaguchi M, Kojima T, Kanekawa M, Aihara N, Nogimura A, Kasai K. Neuropeptides stimulate production of interleukin-1 beta, interleukin-6, and tumor necrosis factor-alpha in human dental pulp cells. *Inflamm Res* 2004; **53**: 199-204 [PMID: 15105969 DOI: 10.1007/s00011-003-1243-z]
- 20 Wilson S, Ngan P, Kess B. Time course of the discomfort in young patients undergoing orthodontic treatment. *Pediatr Dent* 1989; **11**: 107-110 [PMID: 2762180]
- 21 Bondemark L, Fredriksson K, Ilros S. Separation effect and perception of pain and discomfort from two types of orthodontic separators. *World J Orthod* 2004; **5**: 172-176 [PMID: 15615136]
- 22 Tuncer Z, Ozsoy FS, Polat-Ozsoy O. Self-reported pain associated with the use of intermaxillary elastics compared to pain experienced after initial archwire placement. *Angle Orthod* 2011; **81**: 807-811 [PMID: 21446869 DOI: 10.2319/092110-550.1]
- 23 Tuncer Z. The evaluation of the effects of the preoperative and postoperative non-steroidal antiinflammatory analgesics on the pain induced by orthodontic tooth movement and inflammatory mediator levels. Phd's thesis. University of Baskent, 2011
- 24 O'Connor PJ. Patients' perceptions before, during, and after orthodontic treatment. *J Clin Orthod* 2000; **34**: 591-592 [PMID: 11314173]
- 25 Sergrl HG, Klages U, Zentner A. Pain and discomfort during orthodontic treatment: causative factors and effects on compliance. *Am J Orthod Dentofacial Orthop* 1998; **114**: 684-691 [PMID: 9844209 DOI: 10.1016/S0889-5406(98)70201-X]

- 26 Ngan P, Kess B, Wilson S. Perception of discomfort by patients undergoing orthodontic treatment. *Am J Orthod Dentofacial Orthop* 1989; **96**: 47-53 [PMID: 2750720 DOI: 10.1016/0889-5406(89)90228-X]
- 27 Brown DF, Moerenhout RG. The pain experience and psychological adjustment to orthodontic treatment of preadolescents, adolescents, and adults. *Am J Orthod Dentofacial Orthop* 1991; **100**: 349-356 [PMID: 1927986 DOI: 10.1016/0889-5406(91)70073-6]
- 28 Sandhu SS, Sandhu J. Orthodontic pain: an interaction between age and sex in early and middle adolescence. *Angle Orthod* 2013; **83**: 966-972 [PMID: 23705940 DOI: 10.2319/030113-174.1]
- 29 Greydanus DE, Bashe P. American Academy of Pediatrics. Caring for Your Teenager: The Complete and Authoritative Guide. New York, NY: Bantam Books, 2003
- 30 Hobara M. Beliefs about appropriate pain behavior: cross-cultural and sex differences between Japanese and Euro-Americans. *Eur J Pain* 2005; **9**: 389-393 [PMID: 15979019 DOI: 10.1016/j.ejpain.2004.09.006]
- 31 Bergius M, Kiliaridis S, Berggren U. Pain in orthodontics. A review and discussion of the literature. *J Orofac Orthop* 2000; **61**: 125-137 [PMID: 10783564 DOI: 10.1007/BF01300354]
- 32 Firestone AR, Scheurer PA, Bürgin WB. Patients' anticipation of pain and pain-related side effects, and their perception of pain as a result of orthodontic treatment with fixed appliances. *Eur J Orthod* 1999; **21**: 387-396 [PMID: 10502901 DOI: 10.1093/ejo/21.4.387]
- 33 Kvam E, Bondevik O, Gjerdet NR. Traumatic ulcers and pain in adults during orthodontic treatment. *Community Dent Oral Epidemiol* 1989; **17**: 154-157 [PMID: 2736897 DOI: 10.1111/j.1600-0528.1989.tb00012.x]
- 34 Erdinç AM, Dinçer B. Perception of pain during orthodontic treatment with fixed appliances. *Eur J Orthod* 2004; **26**: 79-85 [PMID: 14994886 DOI: 10.1093/ejo/26.1.79]
- 35 Riley JL, Robinson ME, Wise EA, Myers CD, Fillingim RB. Sex differences in the perception of noxious experimental stimuli: a meta-analysis. *Pain* 1998; **74**: 181-187 [PMID: 9520232 DOI: 10.1016/S0304-3959(97)00199-1]
- 36 Jones M, Chan C. The pain and discomfort experienced during orthodontic treatment: a randomized controlled clinical trial of two initial aligning arch wires. *Am J Orthod Dentofacial Orthop* 1992; **102**: 373-381 [PMID: 1456222 DOI: 10.1016/0889-5406(92)70054-E]
- 37 Bergius M, Berggren U, Kiliaridis S. Experience of pain during an orthodontic procedure. *Eur J Oral Sci* 2002; **110**: 92-98 [PMID: 12013568 DOI: 10.1034/j.1600-0722.2002.11193.x]
- 38 Akarslan Z, Erten H. Dis Hekimligi Korkusu ve Kaygısı. *Hacettepe Dis Hekimligi Fakültesi Dergisi* 2009; **1**: 62-68
- 39 Hamurcu KN. The Evaluation of the Relationship between Dental Anxiety and Coping Strategies of the Patients who Referred to Oral and Maxillofacial Surgery Clinic with Pain Complaints and their Pain Perception. Phd's thesis. University of Gazi, 2014. Available from: URL: <http://www.acikarsiv.gazi.edu.tr/index.php?menu=2&secim=10&YayinBIK=10715>
- 40 Bergius M, Broberg AG, Hakeberg M, Berggren U. Prediction of prolonged pain experiences during orthodontic treatment. *Am J Orthod Dentofacial Orthop* 2008; **133**: 339.e1-339.e8 [PMID: 18331926 DOI: 10.1016/j.ajodo.2007.09.013]
- 41 Litt MD. A model of pain and anxiety associated with acute stressors: distress in dental procedures. *Behav Res Ther* 1996; **34**: 459-476 [PMID: 8687368 DOI: 10.1016/0005-7967(96)00015-0]
- 42 Polat O, Karaman AI, Durmus E. Effects of preoperative ibuprofen and naproxen sodium on orthodontic pain. *Angle Orthod* 2005; **75**: 791-796 [PMID: 16279825]
- 43 Michelotti A, Farella M, Martina R. Sensory and motor changes of the human jaw muscles during induced orthodontic pain. *Eur J Orthod* 1999; **21**: 397-404 [PMID: 10502902 DOI: 10.1093/ejo/21.4.397]
- 44 Murdock S, Phillips C, Khondker Z, Hershey HG. Treatment of pain after initial archwire placement: a noninferiority randomized clinical trial comparing over-the-counter analgesics and bite-wafer use. *Am J Orthod Dentofacial Orthop* 2010; **137**: 316-323 [PMID: 20197167 DOI: 10.1016/j.ajodo.2008.12.021]
- 45 Shetty N, Patil AK, Ganeshkar SV, Hegde S. Comparison of the effects of ibuprofen and acetaminophen on PGE2 levels in the GCF during orthodontic tooth movement: a human study. *Prog Orthod* 2013; **14**: 6 [PMID: 24325834 DOI: 10.1186/2196-1042-14-6]
- 46 Jones ML, Richmond S. Initial tooth movement: force application and pain--a relationship? *Am J Orthod* 1985; **88**: 111-116 [PMID: 3861096 DOI: 10.1016/0002-9416(85)90234-9]
- 47 Fernandes LM, Ogaard B, Skoglund L. Pain and discomfort experienced after placement of a conventional or a superelastic NiTi aligning archwire. A randomized clinical trial. *J Orofac Orthop* 1998; **59**: 331-339 [PMID: 9857602 DOI: 10.1007/BF01299769]
- 48 Sandhu SS, Sandhu J. A randomized clinical trial investigating pain associated with superelastic nickel-titanium and multistranded stainless steel archwires during the initial leveling and aligning phase of orthodontic treatment. *J Orthod* 2013; **40**: 276-285 [PMID: 24297959 DOI: 10.1179/1465313313Y.0000000072]
- 49 Ogura M, Kamimura H, Al-Kalaly A, Nagayama K, Taira K, Nagata J, Miyawaki S. Pain intensity during the first 7 days following the application of light and heavy continuous forces. *Eur J Orthod* 2009; **31**: 314-319 [PMID: 19088059 DOI: 10.1093/ejo/cjn072]
- 50 Smith BR, Flanary CM, Hurst LL, Rugh JD. Effects of orthodontic archwire changes on masseter muscle activity. *J Dent Res* 1984; **63**: 258 (abstract)
- 51 Goldreich H, Gazit E, Lieberman MA, Rugh JD. The effect of pain from orthodontic arch wire adjustment on masseter muscle electromyographic activity. *Am J Orthod Dentofacial Orthop* 1994; **106**: 365-370 [PMID: 7942651 DOI: 10.1016/S0889-5406(94)70057-5]
- 52 Lund JP, Lamarre Y, Lavigne G, Duquet G. Human jaw reflexes. In Desmedt J E (ed.) Motor control mechanisms in health and disease. Raven Press, New York, 1983: 739-755
- 53 Gianelly AA, Goldman HM. Tooth movement. Biological basis of orthodontics. Lea and Febiger, Philadelphia, 1971: 116-204
- 54 Wu AK, McGrath C, Wong RW, Wiechmann D, Rabie AB. A comparison of pain experienced by patients treated with labial and lingual orthodontic appliances. *Eur J Orthod* 2010; **32**: 403-407 [PMID: 20018798 DOI: 10.1093/ejo/cjp117]
- 55 Caniklioglu C, Öztürk Y. Patient discomfort: a comparison between lingual and labial fixed appliances. *Angle Orthod* 2005; **75**: 86-91 [PMID: 15747820]
- 56 Miller KB, McGorray SP, Womack R, Quintero JC, Perelmutter M, Gibson J, Dolan TA, Wheeler TT. A comparison of treatment impacts between Invisalign aligner and fixed appliance therapy during the first week of treatment. *Am J Orthod Dentofacial Orthop* 2007; **131**: 302.e1-302.e9 [PMID: 17346581 DOI: 10.1016/j.ajodo.2006.05.031]
- 57 Shalish M, Cooper-Kazar R, Ivgi I, Canetti L, Tsur B, Bachar E, Chaushu S. Adult patients' adjustability to orthodontic appliances. Part I: a comparison between Labial, Lingual, and Invisalign™. *Eur J Orthod* 2012; **34**: 724-730 [PMID: 21750242 DOI: 10.1093/ejo/cjr0862]
- 58 Bertl MH, Onodera K, Čelar AG. A prospective randomized split-mouth study on pain experience during chairside archwire manipulation in self-ligating and conventional brackets. *Angle Orthod* 2013; **83**: 292-297 [PMID: 22827479 DOI: 10.2319/042312-338.1]
- 59 Tecco S, D'Attilio M, Tetè S, Festa F. Prevalence and type of pain during conventional and self-ligating orthodontic treatment. *Eur J Orthod* 2009; **31**: 380-384 [PMID: 19465738 DOI: 10.1093/ejo/cjp003]
- 60 Handelman CS. Nonsurgical rapid maxillary alveolar expansion in adults: a clinical evaluation. *Angle Orthod* 1997; **67**: 291-305; discussion 306-308 [PMID: 9267578]
- 61 Needleman HL, Hoang CD, Allred E, Hertzberg J, Berde C. Reports of pain by children undergoing rapid palatal expansion. *Pediatr Dent* 2000; **22**: 221-226 [PMID: 10846733]
- 62 Schuster G, Borel-Scherf I, Schopf PM. Frequency of and complications in the use of RPE appliances--results of a survey in the Federal State of Hesse, Germany. *J Orofac Orthop* 2005; **66**:

- 148-161 [PMID: 15827702 DOI: 10.1007/s00056-005-0431-6]
- 63 **Cureton SL**. Headgear and pain. *J Clin Orthod* 1994; **28**: 525-530 [PMID: 8617835]
  - 64 **Ngan PW**, Yiu C, Hagg U, Wei SH, Bowley J. Masticatory muscle pain before, during, and after treatment with orthopedic protraction headgear: a pilot study. *Angle Orthod* 1997; **67**: 433-437 [PMID: 9428961]
  - 65 **Egolf RJ**, BeGole EA, Upshaw HS. Factors associated with orthodontic patient compliance with intraoral elastic and headgear wear. *Am J Orthod Dentofacial Orthop* 1990; **97**: 336-348 [PMID: 2321602 DOI: 10.1016/0889-5406(90)70106-M]
  - 66 **Zawawi KH**. Acceptance of orthodontic miniscrews as temporary anchorage devices. *Patient Prefer Adherence* 2014; **8**: 933-937 [PMID: 25061281 DOI: 10.2147/PPA.S66133]
  - 67 **Kuroda S**, Sugawara Y, Deguchi T, Kyung HM, Takano-Yamamoto T. Clinical use of miniscrew implants as orthodontic anchorage: success rates and postoperative discomfort. *Am J Orthod Dentofacial Orthop* 2007; **131**: 9-15 [PMID: 17208101 DOI: 10.1016/j.ajodo.2005.02.032]
  - 68 **Kawaguchi M**, Miyazawa K, Tabuchi M, Fuyamada M, Goto S. Questionnaire survey on pain and discomfort after insertion of orthodontic buccal miniscrews, palatal miniscrews and, orthodontic miniplates. *Orthodontic Waves* 2014; **73**: 1-7 [DOI: 10.1016/j.odw.2013.09.001]
  - 69 **Williams OL**, Bishara SE. Patient discomfort levels at the time of debonding: a pilot study. *Am J Orthod Dentofacial Orthop* 1992; **101**: 313-317 [PMID: 1558060 DOI: 10.1016/S0889-5406(05)80324-5]
  - 70 **Rinchuse DJ**. Pain-free debonding with occlusal rim wax. *J Clin Orthod* 1994; **28**: 587-588 [PMID: 8617816]
  - 71 **Bernhardt MK**, Southard KA, Batterson KD, Logan HL, Baker KA, Jakobsen JR. The effect of preemptive and/or postoperative ibuprofen therapy for orthodontic pain. *Am J Orthod Dentofacial Orthop* 2001; **120**: 20-27 [PMID: 11455373 DOI: 10.1067/mod.2001.115616]
  - 72 **Polat O**, Karaman AI. Pain control during fixed orthodontic appliance therapy. *Angle Orthod* 2005; **75**: 214-219 [PMID: 15825785]
  - 73 **Chumbley AB**, Tuncay OC. The effect of indomethacin (an aspirin-like drug) on the rate of orthodontic tooth movement. *Am J Orthod* 1986; **89**: 312-314 [PMID: 3083687 DOI: 10.1016/0002-9416(86)90053-9]
  - 74 **Kehoe MJ**, Cohen SM, Zarrinnia K, Cowan A. The effect of acetaminophen, ibuprofen, and misoprostol on prostaglandin E2 synthesis and the degree and rate of orthodontic tooth movement. *Angle Orthod* 1996; **66**: 339-349 [PMID: 8893104]
  - 75 **Roche JJ**, Cisneros GJ, Acs G. The effect of acetaminophen on tooth movement in rabbits. *Angle Orthod* 1997; **67**: 231-236 [PMID: 9188968]
  - 76 **Steen Law SL**, Southard KA, Law AS, Logan HL, Jakobsen JR. An evaluation of preoperative ibuprofen for treatment of pain associated with orthodontic separator placement. *Am J Orthod Dentofacial Orthop* 2000; **118**: 629-635 [PMID: 11113797 DOI: 10.1067/mod.2000.110638]
  - 77 **Lee WC**. Experimental study of the effect of prostaglandin administration on tooth movement--with particular emphasis on the relationship to the method of PGE1 administration. *Am J Orthod Dentofacial Orthop* 1990; **98**: 231-241 [PMID: 2403075 DOI: 10.1016/S0889-5406(05)81600-2]
  - 78 **Klein DC**, Raisz LG. Prostaglandins: stimulation of bone resorption in tissue culture. *Endocrinology* 1970; **86**: 1436-1440 [PMID: 4315103]
  - 79 **Leiker BJ**, Nanda RS, Currier GF, Howes RI, Sinha PK. The effects of exogenous prostaglandins on orthodontic tooth movement in rats. *Am J Orthod Dentofacial Orthop* 1995; **108**: 380-388 [PMID: 7572849 DOI: 10.1016/S0889-5406(95)70035-8]
  - 80 **Yamasaki K**, Shibata Y, Fukuhara T. The effect of prostaglandins on experimental tooth movement in monkeys (*Macaca fuscata*). *J Dent Res* 1982; **61**: 1444-1446 [PMID: 6960050]
  - 81 **Proffit WR**. Contemporary Orthodontics, 3rd Edn. The C V Mosby Company, St Louis, 2000
  - 82 **Mohri Y**, Fumoto M, Sato-Suzuki I, Umino M, Arita H. Prolonged rhythmic gum chewing suppresses nociceptive response via serotonergic descending inhibitory pathway in humans. *Pain* 2005; **118**: 35-42 [PMID: 16202533 DOI: 10.1016/j.pain.2005.07.009]
  - 83 **Hwang JY**, Tee CH, Huang AT, Taft L. Effectiveness of therabite wafers in reducing pain. *J Clin Orthod* 1994; **28**: 291-292 [PMID: 8613507]
  - 84 **Fujiyama K**, Deguchi T, Murakami T, Fujii A, Kushima K, Takano-Yamamoto T. Clinical effect of CO(2) laser in reducing pain in orthodontics. *Angle Orthod* 2008; **78**: 299-303 [PMID: 18251609 DOI: 10.2319/033007-153.1]
  - 85 **Tortamano A**, Lenzi DC, Haddad AC, Bottino MC, Dominguez GC, Vigorito JW. Low-level laser therapy for pain caused by placement of the first orthodontic archwire: a randomized clinical trial. *Am J Orthod Dentofacial Orthop* 2009; **136**: 662-667 [PMID: 19892282 DOI: 10.1016/j.ajodo.2008.06.028]

**P- Reviewer:** Carrilho EP, Gokul S

**S- Editor:** Ji FF **L- Editor:** A **E- Editor:** Jiao XK







Published by **Baishideng Publishing Group Inc**

8226 Regency Drive, Pleasanton, CA 94588, USA

Telephone: +1-925-223-8242

Fax: +1-925-223-8243

E-mail: [bpgoffice@wjgnet.com](mailto:bpgoffice@wjgnet.com)

Help Desk: <http://www.wjgnet.com/esps/helpdesk.aspx>

<http://www.wjgnet.com>

