

Analgesics Prescribed In Patients Undergoing Impacted Lower Third Molar Surgery: An Institutional Study

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ABSTRACT

Pain and swelling are two of the main problems experienced by patients who have undergone surgical removal of impacted third molars. The primary obligation of oral health care providers is not only to restore function, but also to relieve pain.

Aim: The aim of the study is to assess the most prescribed analgesics in patients undergoing impacted lower third molar surgery.

Materials and Methods: A total of 1654 records of patients who underwent extraction procedure in 38 and 48 of age groups 10-70 years were collected from the DIAS along with the prescribed analgesics and formulated in the Excel spreadsheet according to the gender and age group of the patients. Descriptive statistics and chi square tests were performed. The p value < 0.05 is considered statistically significant. The analysis is done using SPSS software and the results were obtained.

Results: Out of 1654, about 50.28% of patients were prescribed with a combination of Paracetamol and aceclofenac after extraction procedure either in 38 or 48. 50.38% of the cases were extraction procedures in 38. Majority of the cases were seen in males (30.83%).

Conclusion: It can be comprehended from the study that the most commonly prescribed analgesics to the patients after the extraction of the impacted third molars include a combination drug consisting of Paracetamol and aceclofenac. These are noticed to be effective and balance the patient's analgesic requirements without the potential for adverse effects. These drugs tend to produce significant pain relief and improve the quality of patient's life in the immediate postoperative period. Proper prescribing

practices can help manage tolerance issues, adverse events, as well as common and uncommon side effects.

Keywords: NSAIDs, paracetamol, third molar extractions, innovative technique

INTRODUCTION

The pain and swelling are two of the most common problems experienced by patients who have undergone surgical removal of impacted third molars [1]. These problems result from inflammation due to any surgical trauma[2]. The ultimate primary obligation and ultimate responsibility of oral health care providers is not just to restore function [3], but also to relieve pain[4]. Pain is the most common complaint out of all others often occurring with inflammatory processes after a tooth extraction [5]. The removal of the impacted third molar along with the resultant tissues and cellular destruction brings about the release and production of several biochemical mediators involved in the pain process , in particular , histamine , bradykinin and the prostaglandins [6]. There are numerous analgesics available , and the recent introduction of new agents provides even more options from which to choose than the others [7]. Certain complications include stomach irritation , indigestion [8] , tachycardia , nausea, insomnia , metallic taste in the mouth are present, but it all depends on the dose and mode of administration of the drug[9]. The most prevalent methods involve the administration of analgesics like Non-steroidal anti-inflammatory drugs(NSAIDs)[10] .

NSAID has a very short onset and provides a long duration of analgesia, and numerous other studies have promoted its use in minor oral surgery [11]. The use of NSAID is however associated with several serious treatment side effects, with considerable associated morbidity and mortality. Most of these side effects may be prevented by careful consideration of the patient's risk factors and by subsequent implementation of preventive strategies [12]. Their therapeutic uses include anti-inflammatory , antipyretic , anti dysmenorrhea , antiplatelet action, pain relief, reduce inflammation and reduce fever ,chronic Inflammation ,osteoarthritis , rheumatoid arthritis, ankylosing spondylitis [13].

Our team has extensive knowledge and research experience that has translated into high quality publications [1,14–32]. The aim of this paper is to highlight the most widely analgesics and their protocols applied to the dental field, especially after surgical removal of impacted lower molar teeth in the age group of 10-70 years of age.

MATERIALS AND METHODS

This is a single centered retrospective study done in a private dental institution, Chennai. The data was collected from the dental hospital management system (DIAS). 1654 records of patients from the age group of 10-70 years who particularly underwent extraction procedure in 38 and 48 procedures were

collected. The list of the analgesics which were prescribed were also added. Ethical clearance for this study was obtained from the Institutional review board.

The data included a varied population predominantly the males and females of the south Indian population. All of the case sheets were reviewed and were cross verified by another examiner. The internal validity of the diagnosed cases were as per criteria, medical history, chief complaints and clinical findings. The data collected was tabulated under following parameters : Age, gender, analgesics consumed, extraction procedure performed on 38/48. The independent variable includes age, gender and dependent variables were analgesics, extraction procedure in 38 and 48. The data analysis was done using the SPSS software of version 19. Descriptive statistics and chi square tests were performed. The p value < 0.05 is considered statistically significant.

RESULTS

From (figure 1), it can be comprehended that 26.92% of males were prescribed a combination of Paracetamol and aceclofenac when compared to females which was about 22.19%. The combination of Paracetamol and aceclofenac were the most commonly prescribed analgesics after third molar surgery which accounted for about 50.28%. In (figure 2), it was observed that 27.66% of the patients consumed a combination of paracetamol and aceclofenac after the extraction of the third molars particularly in the age category of 20-30 years. In 20-30 years of age group it can be noticed that 30.23% of patients underwent extraction in 38 rather than 48 (26.06%) (figure 3). 30.83% of the male patients underwent extraction in 38 when compared to 48 (25.70%) (figure 4).

DISCUSSION

Nonsteroidal anti-inflammatory drugs (NSAIDs) play a vital role in reducing pain and edema by suppressing the formation of prostaglandins, by inhibiting the activity of the enzyme Cyclooxygenase (COX-1 and COX-2) [33]. NSAIDs are effective in alleviation of pain, fever and inflammation .They produce analgesic and anti-inflammatory actions by the inhibition of cyclo-oxygenase, thereby reducing the synthesis of arachidonic acid metabolites such as prostaglandins and thromboxanes [34].These are readily absorbed from stomach or small intestine ,widely distributed in all tissues and are bound to plasma proteins therefore, analgesics can displace other drugs from these proteins thereby increase their toxicity [35].

However in various research articles it was noted that the NSAIDs which are extremely effective for the management of acute dental pain possess several adverse effects. These adverse effects include gastrointestinal complaints and somnolence [36]. Cumulative consumption of NSAIDs over a lifetime increases the risk of end-stage renal disease, Dyspepsia, Gastric mucosal damage, Increased bleeding, Possible renal impairment, Anaphylactoid reactions [37].

A recent research study has succeeded in demonstrating the analgesic efficacy of single oral doses of tramadol, for impacted third molar extraction, with an acceptable incidence and severity of side effects, over the first 6 hours following extraction. Tramadol drug was found to be more effective postoperatively than preoperatively [38]. Paracetamol containing ibuprofen has been evaluated in several different formulations. One recent modification include the use of gel caps that provide faster absorption and

therefore a quicker onset for meaningful analgesia that occurs about 25-30 minutes after ingestion[39]. The selective cox-2 inhibitor has higher effect than the conventional NSAIDs and has low gastro intestinal and high cardiovascular side effects than to the conventional NSAIDs [40]. In recent studies it is described the analgesic effect of combination of dexamethasone and diclofenac K that the potency and dosage of dexamethasone within the first 24 h was adequate to enhance the efficacy of diclofenac K[41]. It appears that steroids are preferably administered preoperatively, as they extend the coverage up to 24 - 48 hours after surgery .

According to studies, it was confirmed that NSAIDs are comparatively more efficient in controlling pain when administered before the onset of the inflammatory process. The NSAIDs used preoperatively were efficient in controlling pain in the surgery of impacted mandibular third molars. However, when single 150-mg aceclofenac dose was administered after surgery, it was found that there was no statistically significant difference from placebo administration in the control of postoperative pain. The criteria which was adopted to determine the study time points were based on literature reports that indicate the greatest pain experienced in the 3–6-h period after surgery to extract the impacted third molars. In the following research, patient collaboration was necessary, which further limited the assessment period [42].

Further clinical studies are required to state specific guidelines, and oral surgeons involved in third molar surgery should evaluate the local and general health conditions of the patients before prescribing any drugs for patients.



Error Bars: 95% Cl

Fig. 1. Bar graph representing the association between the gender of the patients and the analgesic consumed post lower third molar surgery. X axis represents the gender of the patients and Y axis represents the total number of patients who took analgesics after the treatment. Majority of the male patients were prescribed with a combination of Paracetamol and aceclofenac rather than the single analgesics such as Diclofenac, ketorolac, tramadol, paracetamol, or aceclofenac after the extraction procedures. Chi square test was done and the association was found to be statistically significant. Pearson

chi square value:55.337; p value: 0.00(<0.05). Hence it is statistically significant, proving that there was an association with gender and the prescribed medications after the extraction procedure.



Error Bars: 95% CI

Fig. 2. Bar graph representing the association between the age of the patients and the analgesic consumed post lower third molar surgery. X axis represents the age of the patients ranging from 10-70years and Y axis represents the total number of patients who consumed the analgesics after the procedure. Higher numbers of consumption of a combination of Paracetamol and aceclofenac were noticed for the age group of 20-30 years. Chi square test was done and the association was found to be statistically significant. Pearson chi square value:54.609; p value:0.00(<0.05). Hence it is statistically significant, proving that there was an association of age with respect to the analgesics consumed after the procedure.





Fig. 3. Bar graph representing the association between age of the patients and the impacted mandibular third molars. The X axis represents the patients who underwent the procedure which ranged from 10-70 years and the Y axis represents the total number of patients who underwent extraction of third molars. It was noted that the majority of the patients underwent the procedure in the age group of 20-30 years. A higher number of extractions were done for 38 when compared to 48 in the same age group. Chi square test was done and the association was found to be statistically not significant. Pearson chi square value:1.471; p value:0.106 (<0.05). Hence it is statistically not significant, proving that there was no association between age of the patients with respect to the extraction done in 38 or 48.



Error Bars: 95% CI

Fig. 4. Bar graph representing the association between gender of the patients and the impacted mandibular third molars. X axis represents the gender of the patients and the Y axis represents the total number of patients who underwent extraction of third molars. Blue represents '38', Green represents '48'. It was observed that a large number of male patients underwent extraction procedures on 38 rather than 48 when compared to females. Chi square test was done and the association was found to be statistically not significant. Pearson chi square value:0.230; p value:0.973(<0.05). Hence it is statistically not significant, proving that there was no association between gender of the patients with respect to the extraction procedure done on 38 or 48.

CONCLUSION

It can be comprehended from the study that the most commonly prescribed analgesics to the patients after the extraction of the impacted third molars include a combination drug consisting of Paracetamol and aceclofenac. These are noticed to be effective and balance the patient's analgesic requirements without the potential for adverse effects. These drugs tend to produce significant pain relief and improve the quality of patient's life in the immediate postoperative period. Proper prescribing practices can help manage tolerance issues, adverse events, as well as common and uncommon side effects.

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No potential conflict of interest relevant to this article was reported.

REFERENCES

- [1] J PC, Pradeep CJ, Marimuthu T, Krithika C, Devadoss P, Kumar SM. Prevalence and measurement of anterior loop of the mandibular canal using CBCT: A cross sectional study. Clinical Implant Dentistry and Related Research 2018;20:531–4. https://doi.org/10.1111/cid.12609.
- [2] Owobu T, Bamgbose BO, Kaura AM, Amole IO, Asaumi J. Assessment of pain and swelling following surgical extraction of impacted mandibular third molar using complete and partial wound closure techniques in a tertiary institution. Journal of Dentomaxillofacial Science 2019;4:15. https://doi.org/10.15562/jdmfs.v4i1.855.
- [3] Wahab PUA, Abdul Wahab PU, Madhulaxmi M, Senthilnathan P, Muthusekhar MR, Vohra Y, et al. Scalpel Versus Diathermy in Wound Healing After Mucosal Incisions: A Split-Mouth Study. Journal of Oral and Maxillofacial Surgery 2018;76:1160–4. https://doi.org/10.1016/j.joms.2017.12.020.
- [4] López-Ramírez M, Vílchez-Pérez MÁ, Gargallo-Albiol J, Arnabat-Domínguez J, Gay-Escoda C. Efficacy of low-level laser therapy in the management of pain, facial swelling, and postoperative trismus after a lower third molar extraction. A preliminary study. Lasers in Medical Science 2012;27:559–66. https://doi.org/10.1007/s10103-011-0936-8.
- [5] Minwah B. Pre-Emptive Analgesic Effect of Tramadol and Ibuprofen After Impacted Mandibular Third Molar Extraction. Al-Rafidain Dental Journal 2016;17:76–85. https://doi.org/10.33899/rden.2016.164141.
- [6] Kikuchi M. Intranasal Ketorolac Spray: an Effective Alternative to Opiod Analgesics following Third Molar Surgery. Journal of Oral and Maxillofacial Surgery 2014;72:e110. https://doi.org/10.1016/j.joms.2014.06.194.
- [7] Haas DA. An update on analgesics for the management of acute postoperative dental pain. J Can Dent Assoc 2002;68:476–82.

- [8] Aggarwal BB, Kunnumakkara AB. Molecular Targets and Therapeutic Uses of Spices: Modern Uses for Ancient Medicine. World Scientific; 2009.
- [9] Lim D, Ngeow WC. A Comparative Study on the Efficacy of Submucosal Injection of Dexamethasone Versus Methylprednisolone in Reducing Postoperative Sequelae After Third Molar Surgery. Journal of Oral and Maxillofacial Surgery 2017;75:2278–86. https://doi.org/10.1016/j.joms.2017.05.033.
- [10] Ong KS, Seymour RA, Chen FG, Ho VCL. Preoperative ketorolac has a preemptive effect for postoperative third molar surgical pain. International Journal of Oral and Maxillofacial Surgery 2004;33:771–6. https://doi.org/10.1016/j.ijom.2004.01.020.
- [11] Giglio JA, Campbell RL. Comparison of etodolac, zomepirac, and placebo for relief of pain after oral surgery. J Oral Maxillofac Surg 1986;44:765–70. https://doi.org/10.1016/0278-2391(86)90150-3.
- [12] Meek IL, Van de Laar MAFJ, E Vonkeman H. Non-Steroidal Anti-Inflammatory Drugs: An Overview of Cardiovascular Risks. Pharmaceuticals 2010;3:2146–62. https://doi.org/10.3390/ph3072146.
- [13] Mishra H, Khan F. A double-blind, placebo-controlled randomized comparison of pre and postoperative administration of ketorolac and tramadol for dental extraction pain. Journal of Anaesthesiology Clinical Pharmacology 2012;28:221. https://doi.org/10.4103/0970-9185.94892.
- [14] Prostran M. Clinical Trials in Vulnerable Populations. BoD Books on Demand; 2018.
- [15] Mudigonda SK, Murugan S, Velavan K, Thulasiraman S, Krishna Kumar Raja VB. Non-suturing microvascular anastomosis in maxillofacial reconstruction- a comparative study. J Craniomaxillofac Surg 2020;48:599–606. https://doi.org/10.1016/j.jcms.2020.04.005.
- [16] Narayanasamy RK, Muthusekar RM, Nagalingam SP, Thyagarajan S, Ramakrishnan B, Perumal K. Lower pretreatment hemoglobin status and treatment breaks in locally advanced head and neck squamous cell carcinoma during concurrent chemoradiation. Indian J Cancer 2021;58:62–8. https://doi.org/10.4103/ijc.IJC_656_18.
- [17] Wang H, Chinnathambi A, Alahmadi TA, Alharbi SA, Veeraraghavan VP, Krishna Mohan S, et al. Phyllanthin inhibits MOLT-4 leukemic cancer cell growth and induces apoptosis through the inhibition of AKT and JNK signaling pathway. J Biochem Mol Toxicol 2021;35:1–10. https://doi.org/10.1002/jbt.22758.
- [18] Li S, Zhang Y, Veeraraghavan VP, Mohan SK, Ma Y. Restorative Effect of Fucoxanthin in an Ovalbumin-Induced Allergic Rhinitis Animal Model through NF-κB p65 and STAT3 Signaling. Journal of Environmental Pathology, Toxicology and Oncology 2019;38:365–75. https://doi.org/10.1615/jenvironpatholtoxicoloncol.2019030997.
- [19] Ma Y, Karunakaran T, Veeraraghavan VP, Mohan SK, Li S. Sesame Inhibits Cell Proliferation and Induces Apoptosis through Inhibition of STAT-3 Translocation in Thyroid Cancer Cell Lines (FTC-133). Biotechnology and Bioprocess Engineering 2019;24:646–52. https://doi.org/10.1007/s12257-019-0151-1.
- [20] Bishir M, Bhat A, Essa MM, Ekpo O, Ihunwo AO, Veeraraghavan VP, et al. Sleep Deprivation and Neurological Disorders. BioMed Research International 2020;2020:1–19. https://doi.org/10.1155/2020/5764017.
- [21] Fan Y, Maghimaa M, Chinnathambi A, Alharbi SA, Veeraraghavan VP, Mohan SK, et al. Tomentosin Reduces Behavior Deficits and Neuroinflammatory Response in MPTP-Induced Parkinson's Disease in Mice. J Environ Pathol Toxicol Oncol 2021;40:75–84. https://doi.org/10.1615/JEnvironPatholToxicolOncol.v40.i1.70.

- [22] Zhang C, Chen Y, Zhang M, Xu C, Gong G, Veeraraghavan VP, et al. Vicenin-2 Treatment Attenuated the Diethylnitrosamine-Induced Liver Carcinoma and Oxidative Stress through Increased Apoptotic Protein Expression in Experimental Rats. Journal of Environmental Pathology, Toxicology and Oncology 2020;39:113–23. https://doi.org/10.1615/jenvironpatholtoxicoloncol.2020031892.
- [23] Gan H, Zhang Y, Zhou Q, Zheng L, Xie X, Veeraraghavan VP, et al. Zingerone induced caspasedependent apoptosis in MCF-7 cells and prevents 7,12-dimethylbenz(a)anthracene-induced mammary carcinogenesis in experimental rats. J Biochem Mol Toxicol 2019;33:e22387. https://doi.org/10.1002/jbt.22387.
- [24] Saravanakumar K, Park S, Mariadoss AVA, Sathiyaseelan A, Veeraraghavan VP, Kim S, et al. Chemical composition, antioxidant, and anti-diabetic activities of ethyl acetate fraction of Stachys riederi var. japonica (Miq.) in streptozotocin-induced type 2 diabetic mice. Food Chem Toxicol 2021;155:112374. https://doi.org/10.1016/j.fct.2021.112374.
- [25] Veeraraghavan VP, Hussain S, Balakrishna JP, Dhawale L, Kullappan M, Ambrose JM, et al. A Comprehensive and Critical Review on Ethnopharmacological Importance of Desert Truffles: Terfezia claveryi, Terfezia boudieri, and Tirmania nivea. Food Reviews International 2021:1–20. https://doi.org/10.1080/87559129.2021.1889581.
- [26] Wei, Wei W, Li R, Liu Q, Seshadri VD, Veeraraghavan VP, et al. Amelioration of oxidative stress, inflammation and tumor promotion by Tin oxide-Sodium alginate-Polyethylene glycol-Allyl isothiocyanate nanocomposites on the 1,2-Dimethylhydrazine induced colon carcinogenesis in rats. Arabian Journal of Chemistry 2021;14:103238. https://doi.org/10.1016/j.arabjc.2021.103238.
- [27] Sathya S, Ragul V, Veeraraghavan VP, Singh L, Niyas Ahamed MI. An in vitro study on hexavalent chromium [Cr(VI)] remediation using iron oxide nanoparticles based beads. Environmental Nanotechnology, Monitoring & Management 2020;14:100333. https://doi.org/10.1016/j.enmm.2020.100333.
- [28] Chandrasekar R, Chandrasekhar S, Sundari KKS, Ravi P. Development and validation of a formula for objective assessment of cervical vertebral bone age. Prog Orthod 2020;21:38. https://doi.org/10.1186/s40510-020-00338-0.
- [29] Ramakrishnan M, Dhanalakshmi R, Subramanian EMG. Survival rate of different fixed posterior space maintainers used in Paediatric Dentistry - A systematic review. Saudi Dent J 2019;31:165–72. https://doi.org/10.1016/j.sdentj.2019.02.037.
- [30] Felicita AS, Sumathi Felicita A. Orthodontic extrusion of Ellis Class VIII fracture of maxillary lateral incisor – The sling shot method. The Saudi Dental Journal 2018;30:265–9. https://doi.org/10.1016/j.sdentj.2018.05.001.
- [31] Su P, Veeraraghavan VP, Mohan SK, Lu W. A ginger derivative, zingerone—a phenolic compound induces ROS-mediated apoptosis in colon cancer cells (HCT-116). Journal of Biochemical and Molecular Toxicology 2019;33. https://doi.org/10.1002/jbt.22403.
- [32] Wan J, Feng Y, Du L, Veeraraghavan VP, Mohan SK, Guo S. Antiatherosclerotic Activity of Eriocitrin in High-Fat-Diet-Induced Atherosclerosis Model Rats. J Environ Pathol Toxicol Oncol 2020;39:61–75. https://doi.org/10.1615/JEnvironPatholToxicolOncol.2020031478.
- [33] Vane JR, Botting RM. The mechanism of action of aspirin. Thrombosis Research 2003;110:255–8. https://doi.org/10.1016/s0049-3848(03)00379-7.
- [34] Vane JR, Botting RM. The history of anti-inflammatory drugs and their mechanism of action. New

Targets in Inflammation 1996:1–12. https://doi.org/10.1007/978-94-011-5386-7_1.

- [35] Hersh EV, Dionne RA. Nonopioid Analgesics. Pharmacology and Therapeutics for Dentistry 2017:257–75. https://doi.org/10.1016/b978-0-323-39307-2.00017-5.
- [36] Edwards JE, McQuay HJ, Andrew Moore R, Collins SL. Reporting of Adverse Effects in Clinical Trials Should Be Improved. Journal of Pain and Symptom Management 1999;18:427–37. https://doi.org/10.1016/s0885-3924(99)00093-7.
- [37] Perneger TV, Whelton PK, Klag MJ. Risk of Kidney Failure Associated with the Use of Acetaminophen, Aspirin, and Nonsteroidal Antiinflammatory Drugs. New England Journal of Medicine 1994;331:1675–9. https://doi.org/10.1056/nejm199412223312502.
- [38] Hargreaves K, Abbott PV. Drugs for pain management in dentistry. Australian Dental Journal 2005;50:S14–22. https://doi.org/10.1111/j.1834-7819.2005.tb00378.x.
- [39] Bamgbose BO, Akinwande JA, Adeyemo WL, Ladeinde AL, Arotiba GT, Ogunlewe MO. Effects of coadministered dexamethasone and diclofenac potassium on pain, swelling and trismus following third molar surgery. Head & Face Medicine 2005;1. https://doi.org/10.1186/1746-160x-1-11.
- [40] Mouedden ME, El Mouedden M, Meert TF. Pharmacological evaluation of opioid and non-opioid analgesics in a murine bone cancer model of pain. Pharmacology Biochemistry and Behavior 2007;86:458–67. https://doi.org/10.1016/j.pbb.2007.01.003.
- [41] Brogan S, Mandyam S, Drennan DA. Non-Opioid Analgesics. Pharmacology and Physiology for Anesthesia 2013:272–90. https://doi.org/10.1016/b978-1-4377-1679-5.00016-8.
- [42] Rang HP, Dale MM, Ritter JM, Flower RJ. Antihelminthic drugs. Rang & Dale's Pharmacology 2007:712–7. https://doi.org/10.1016/b978-0-443-06911-6.50055-4.