Pain management in endodontics

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Abstract
Pain is a stimuli that causes the greatest fear in any patient who requires an endodontic treatment. This fear if tackled efficiently not only improves the success of the treatment but also creates a strong dentist patient relationship. Pain management can be done effectively by using various pharmacological regimes such as anxiolytics, local anaesthetics, NSAIDs and opioids. These drugs have proven to reduce pain and help the patient overcome the fear associated with endodontic treatment. This article briefly discusses the drugs, its dosage and properties that are commonly used to manage endodontic pain.

Keywords: Anxiolytics, Local anaesthesia, NSAIDs, Opioids, Pain pathway.

Introduction
The fear of pain is the greatest fear that makes most of the patients hesitant to approach a dentist for the treatment. Therefore managing pain is an art that every dentist should persist to gain the confidence of patients and complete the treatment in the most satisfactory manner.

The management of pain represents both a challenge and an opportunity for the endodontist. It is a challenge due to pharmacological, behaviourial and practice management issues.

Endodontic pain management encompasses all aspects of treatment: preoperative pain control includes accurate diagnosis and anxiety reduction; intraoperative pain control revolves around effective local anaesthetic and operative techniques; and postoperative pain management can involve a variety of pharmacologic agents. Therefore this review throws a light on the common analgesics used in endodontic practise which includes anti anxiety drugs, local anaesthetics and non steroidal anti inflammatory drugs (NSAIDs).

Pain and its pathways
Inflammatory mediators such as bradykinin, serotonin, prostaglandins, cytokines and H+ are released from damaged tissue which stimulate the nociceptors that carry the noxious stimuli to the central nervous system. This stimuli is carried to the brain by the free nerve endings of primary afferent Aδ and C fibres.

Aδ fibres are lightly myelinated and smaller diameter fibres that respond to mechanical and thermal stimuli. They are responsible for the rapid, sharp pain and initial reflex response to acute pain. C fibres are unmyelinated, smallest diameter fibres which demonstrate the slowest conduction. They respond to chemical, mechanical and thermal stimuli and their activation leads to slow burning pain. These fibres carry the nociceptor signals to higher centres in the brain by the the spinothalamic tract and the spinoreticular tract. While the spinothalamic tract is associated with pain localisation, the emotional aspect of pain is due to the spinoreticular tract.1–3

Anxiolytics
Anxiolytics or psycholeptics are drugs that cause soothing effects and decrease the anxiety level. In endodontics it can be used in cases of trauma, swelling like abscess or cellulitis or in patients in grave fear and apprehension. The commonly used anxiolytics are benzodiazepine (BZD) such as Diazepam, Lorazepam and Triazolam.

Benzodiazepine
BZD in endodontics have proved to be advantageous due to their inherent properties such as skeletal muscle relaxation, patients consciousness is not lost, no effect on respiration, cardiovascular functions or any other body sytem, less sleep distortion and low abuse liability.4

Mechanism of action
BZD has the affinity for the GABAα Subunit of Gamma amino butyric acid (GABA), which is an inhibitory neurotransmitter. These BZD bind at the β subunit of GABA and cause increase in the frequency of chloride channel opening and influx of Chloride ions. This influx further result in a hyperpolarized state and decreases the firing rate of the neurons thus causing the anxiolytic effect.4

Commonly used BZD
The BZD that is most commonly preferred in dental practise is triazolam as it has a fast onset of action, short duration of effect, short elimination half-life and early recovery.5

Diazepam is amongst the oldest used anti anxiety drugs. Though its efficacy as an anxiolytic and sedative
is commendable it causes sedation hours later after the initial effect which restricts its use in endodontics. It is rapidly absorbed and has a fast onset of action. Lorazepam is a commonly used sedative. Due to its poor lipid solubility its onset of action ranges within an uncertain limit of 1 to 6 hours. Since it has a longer duration of action it can be given in cases of endodontic surgery or where multiple root canal therapies have to be performed.

Though BZD are relatively safe drugs side effects may include dizziness, vertigo, ataxia, disorientation, amnesia, impairment of psychomotor skills therefore patient should not drive after its use. All the BZD are excreted in urine, crosses the placenta and are secreted in milk therefore are contraindicated in pregnancy and lactating mothers.

Dosage

Diazepam 5-10mg, 6-12 hrs
Lorazepam: 0.5-1.0 mg, 12-24 hrs
Triazolam: 0.125-0.25 mg, 24 hrs

Local Anaesthesia (LA)

LA causes loss of sensation in a circumscribed area of the body by depression of excitation in nerve endings or an inhibition of the conduction process in the peripheral nerves. They are classified according to their duration of action as intermediate or long duration, based on type of intermediate chain as amide or ester, based on surface anaesthesia as soluble and insoluble. The commonly used LA agents are amides such as lignocaine, benzocaine, prilocaine and articaine.

Mechanism of action

According to the specific receptor theory LA binds to specific receptors present in the sodium channel and blocks the influx of sodium ions into their channels. This prevents the increase in permeability of the nerve membrane to sodium that is responsible for nerve conduction. Displacement of calcium from the ion channels is said to be the primary factor responsible for competitively binding of LA at the channel and decreasing the sodium conductance.

Commonly used LA agents

Lignocaine

Lignocaine is the most versatile LA that serves both for surface application as well as injection and is available in a variety of forms. Its anaesthetic effect is more intense, long lasting and causes its action in 3 minutes. Vasoconstrictors are added to prolong its duration of action and therefore 2 percent with 1:100,000 epinephrine vasoconstrictor, and

2 percent with 1:50,000 epinephrine are commonly used which have a have a pulpal duration of 1-1.5 hours and a soft-tissue range of three to five hours.

Benzocaine

Topical anaesthetics may be indicated to minimize the sensation of needle insertion or for very brief relief from painful mucosal lesions. 20% benzocaine is most commonly used topical anaesthesia. They are available in gel as well as solution form. They produce long-lasting anaesthesia without systemic toxicity.

Prilocaine

Prilocaine is an alternative to patients in whom vasoconstrictor is contraindicated as it has mild vasodilatory effect compared to other amides. It is less toxic, less potent than lidocaine and its duration of action is slightly longer. It is available as 4% prilocaine with 1:200,000 epinephrine and 4% prilocaine (plain). Contraindicated in patients with methemoglobinemia and true amide type allergy.

Articaine

Following lignocaine, 4% solution of articaine with 1:100,000 epinephrine is the most common anaesthetic agent used. It has a faster duration of action of 1.4 to 3.6 minutes, longer profound level of anaesthesia and increased tissue diffusability. It provides pulp anaesthesia for 1 hour with soft-tissue anaesthesia for 2-4 hours. They should be used with caution in inferior alveolar and lingual nerve block injections as they show an increased risk of nerve paresthesia.

All the amides are degraded in liver microsomes and excreted by the kidneys. Hypersensitivity reactions like rashes, angioedema, dermatitis, asthma may occur. Vasoconstrictor containing LA should be avoided for patients with ischemic heart disease, cardiac arrhythmia, thyrotoxicosis, uncontrolled hypertension, and those receiving β blockers or tricyclic antidepressants.
Non steroidal anti inflammatory drugs
The major analgesic drug class for treating endodontic pain are the nonsteroidal anti-inflammatory analgesics and acetaminophen. They produce their analgesic and anti-inflammatory effect by inhibiting the enzyme cyclooxygenase and are mainly useful in the initial pain management that has an inflammatory component.

Mechanism of action
Prostaglandins and thromboxane A2 (TXA2) collectively termed as prostanoids are formed when arachidonic acid is metabolized by cyclooxygenase enzyme. They exist in different isoenzyme forms such as COX-1 and COX-2. These 2 cyclooxygenase enzymes are the target site of NSAIDs. Depending on their site of action they may be classified as non selective cox inhibitors such as ibuprofen, diclofenac and ketorolac, selective cox-2 inhibitors such as celecoxib and analgesic- antipyretics with poor anti-inflammatory action like acetaminophen.4

Ibuprofen
First member of class of Propionic acid derivative and is commonly used NSAID in endodontics. They are well absorbed orally and inhibit platelet function therefore should not be taken with anticoagulants. Contraindicated in pregnancy and peptic ulcer patients.11
Dosage12: 200-400 mg, 4-6hourly

Diclofenac
Diclofenac exists in two forms known as Diclofenac Sodium and Diclofenac Potassium. Diclofenac potassium has an immediate release as they are rapidly soluble in water while Diclofenac sodium has delayed release. Hence for acute and severe pain, it is better to take Diclofenac potassium than Diclofenac sodium.13
Dosage12: 50mg, 8 hourly

Ketorolac
It’s a novel pyrrolo pyyrole derivative with potent analgesic and modest anti-inflammatory activity. It has an efficacy equivalent to that of morphine without the side effects usually associated with opioids. It is advised to use Ketorolac in chronic pain conditions and its use for more than 5 days should be avoided as it may cause gastrointestinal bleeding, peptic ulceration, renal failure, or hematological dysfunction due to inhibited platelet aggregation with thromboxane inhibition.14
Dosage12: 10 mg 4-6 hourly for short term management of moderate pain.

Piroxicam
Piroxicam is a reversible inhibitor of COX which is long-acting with good anti-inflammatory and analgesic-antipyretic action. It also decreases the production of IgM rheumatoid factor and leucocyte chemotaxis and therefore can inhibit inflammation in diverse ways.11
Dosage12: 40mg once a day followed by 20 mg OD

Celecoxib
The COX-2 selectivity of celecoxib is modest with equal analgesic and anti inflammatory efficacy as the conventional NSAIDs. Its added advantage is it has low ulcerogenic potential which gives minimal GIT irritation and no increase in bleeding time.15 It is slowly absorbed with elimination half life of 11 hours.16
Dosage12: 100-200 mg bid

Acetaminophen
Also known as paracetamol (PCT) is the safest analgesic that blocks the prostaglandin pathway within the central nervous system. It has a good analgesic and antipyretic action but no anti inflammatory effect as it does not act directly on the peripheral tissue.17 It has good oral absorption, effect lasts for 3-5 hrs and is excreted rapidly in urine. PCT does not cause any significant gastric irritation, mucosal erosion, bleeding and also does not affect platelet function or clotting factors.4 The major adverse effect of acetaminophen is hepatotoxicity therefore in patients with chronic alcoholism its maximum dose should be limited to 2 g per day compared to 4g per day.17
Dosage12: 325-650 mg for every 4-6 hourly.

Adverse effects and contraindications of NSAIDs
They may cause gastrointestinal disturbances such as gastric irritation, erosions, peptic ulcers, renal impairment, headache, mental confusion, bleeding, asthma exacerbation and anaphylactic reactions. Therefore they are contraindicated in patients with gastrointestinal diseases, bleeding disorders, asthma, hypersensitivity, third trimester of pregnancy and chronic renal disease as they are majorly excreted via the kidneys and rest by bile. It should not be given to patients consuming drugs like diuretics or beta-blockers, ACE inhibitors, anticoagulants like warfarrin, methotrexate, digoxin and alcohol.7,18

Opioids
Opioids are indicated in cases of moderate to severe pain if PCT and NSAIDs are ineffective. Pertaining to the side effects associated with opioids it is advised to use it in combination regime with PCT or NSAIDs with doses of the latter two to being higher. Commonly used opioids are codeine, oxycodone, hydrocodone and tramadol.

Mechanism of Action
Opioids are G protein coupled receptors that act on \( \mu, K \) and \( \delta \) receptors present on the neurons in the CNS and in peripheral tissues. They cause their analgesic effect by inhibiting the neurotransmitter release either by inhibiting the calcium entry, by enhancing outward
movement of potassium ions or by inhibiting adenylate cyclase.⁴

**Commonly used opioids**

**Codeine**

Codeine is methyl-morphine that partly gets converted to morphine in the body that is responsible for its analgesic effect. Codeine has good activity orally and its effect lasts for 4-6 hours. Abuse liability is low and causes constipation when used in analgesic doses. Used with aspirin or PCT in mild to moderate pain.¹⁹

Dosage²⁰: 180-200mg every 4-6 hourly.

**Oxycodone and hydrocodone**

They are semi synthetic opioids indicated for moderate to severe pain. Onset of action is 10-30 minutes with instant action of 3-6 hours that starts reducing pain in 10-15 minutes on empty stomach. Hydrocodone is available orally and has onset of action of 10 to 20 minutes with effect lasting for 4-8hrs.⁴

Dosage of oxycodone and hydrocode²⁰: 20-30 mg, 4-8 hourly.

**Tramadol**

Tramadol is a synthetic analogue of codeine which has an action on the µ-opioid receptor that helps to reduce pain. It’s effective for moderately severe pain, especially if patient is already on NSAIDs for acute pain management. Its effect lasts for 4-6 hrs, is available in combination with PCT and decreases the risk of constipation.²¹

Dosage²¹: 50-100mg. 4-6 hourly

**Adverse effects and contraindications of opioids**

Opioids cause sedation, respiratory depression, dependence, nausea, miosis, and constipation.¹⁷ They are contraindicated in patients with severe chronic respiratory disease, severe inflammatory bowel disease and concurrent use of alcohol.⁷

**Conclusion**

Careful assessment of the type and intensity of pain helps to select the pharmacological regime required that will aid in the success of an endodontic treatment. Judicious and intelligent use of anti anxiety drugs to relieve stress, local anaesthesia to give a pain free endodontic experience and NSAIDs to prevent the after effects increases the standard of dental treatment. Combination therapy of opioids with analgesics should be ascertained in cases of moderate to severe pain when NSAIDs alone have proved to be unhelpful.

**Funding:** No funding sources.

**Conflict of interest:** None declared.

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**How to cite the article:** Pain management in endodontics, Khatri M., Ghivari S, Pujar M., Vagaralli H., Uppin V. IP Indian J Conserv Endod. 2018;3(3):65-68.