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# Pioneering Periapical Healing: The Novel Synergy of Mineral Trioxide Aggregate and Injectable Platelet-Rich Fibrin

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#### **Abstract**

This case report presents a novel non-surgical approach for managing a substantial periapical lesion associated with tooth 12 using a combination of injectable platelet-rich fibrin (i-PRF) and mineral trioxide aggregate (MTA). A 28-year-old male patient presented with pus discharge and intermittent swelling following a history of dental trauma. Clinical and radiographic assessments confirmed a large periapical cyst associated with tooth 12. The treatment involved root canal therapy with calcium hydroxide medication, leading to symptom relief. Subsequently, i-PRF combined with MTA was used as a regenerative material for periapical healing. Follow-up examinations at three, six, and nine months showed complete resolution of symptoms and radiographic evidence of lesion healing. This innovative technique demonstrates the potential of i-PRF and MTA synergy in the non-surgical management of periapical lesions, avoiding the risks associated with surgical interventions and promoting effective tissue healing.

Categories: Dentistry

**Keywords:** radicular cyst, non-surgical management, mineral trioxide aggregate, injectable platelet-rich fibrin, periapical lesion

#### Introduction

Traumatic injuries typically disrupt the blood supply to the dental pulp, leading to pulp tissue death and creating oxygen-deprived conditions that favor the growth of opportunistic microorganisms. Consequently, this can result in the formation of periapical lesions [1]. Periapical lesions generally arise as a response to the intrusion of microorganisms and their by-products into the root canal system, with their growth being facilitated through various processes such as the accumulation of fluid due to osmosis, the proliferation of epithelial cells, and molecular mechanisms [2]. Therefore, if the inflammatory exudates within the lesion are effectively removed to reduce hydrostatic pressure, and if the underlying microbiological cause is addressed through nonsurgical root canal treatment, these lesions may shrink and regress by apoptosis [3].

The ultimate objective of endodontic treatment is to restore the affected tooth to a healthy and functional state without undergoing surgical procedures [4]. Until recently, the conventional approach to managing endodontic periapical lesions, especially when they were large, involved surgical intervention. However, recent advances in our understanding of these lesions' origin, pathological characteristics, and clinical behavior, coupled with successful treatments in various clinical trials, have shifted the preference toward nonsurgical methods [5]. All inflammatory periapical lesions should be addressed using conservative nonsurgical methods, such as orthograde root canal therapy because it is less invasive, has more patient acceptability, and causes less psychological distress to the patients [6]. Surgical intervention is only recommended when non-surgical approaches prove ineffective [7]. Furthermore, surgical procedures come with several drawbacks such as fistula formation, healing by scar tissue formation, and sometimes swelling that limit their use in the treatment of periapical lesions [8]. It is worth noting that endodontic treatment for teeth with periapical lesions has been reported to achieve a success rate of 85% [9]. Additionally, there have been reports of a 94.4% incidence of complete and partial healing of periapical lesions following nonsurgical endodontic therapy [10]. This case report describes a novel technique for the non-surgical management of a large periapical lesion using a combination of injectable platelet-rich fibrin (i-PRF) and mineral trioxide aggregate (MTA).

#### **Case Presentation**

A 28-year-old male patient presented to the department with a chief complaint of pus discharge from the upper front region of the jaw for one week. The patient had started noticing this pus discharge and accompanying swelling from the upper front region of the jaw for one week, which appeared intermittently, sometimes causing mild discomfort. The patient gave a history of trauma (due to a self-fall) in the same region approximately 10 years ago resulting in mild pain in the upper front region, which was effectively relieved with medication. There was no history of heat or balm application. The past medical and dental history was not significant. On extra-oral examination, no gross asymmetry was detected. On intra-oral

examination, a sinus opening was seen associated with tooth 12 (Figure 1).



FIGURE 1: Pre-operative intraoral picture

Crowding was evident in the upper and lower jaws. Electric and thermal pulp vitality testing showed a negative response for tooth 12 with a normal response for teeth 11, 13 suggestive of non-vital tooth with 12. All teeth did not elicit tenderness during percussion testing. Subsequently, an intra-oral periapical (IOPA) radiograph was taken which revealed a large periapical radiolucent shadow associated with 12 (Figure 2A). Thereafter, a cone-beam computed tomography (CBCT) was performed with 12 regions. The coronal CBCT view showed a wide canal and a radiolucent shadow with well-corticated borders measuring 13.78 mm superior-inferiorly and a resorbed root apex with 12 (Figure 2B). The axial view showed perforation of the labial and lingual cortical plate and the lesion measuring 10.48 mm mesiodistally and 10.56 mm buccopalatally (Figures 2C, 2D). The sagittal view of 12 showed root resorption with an apical diameter of 0.82 mm and loss of buccal cortical plate (Figure 2E). The bony view showed a tunnel defect with 12 (Figures 2F, 2G).



# FIGURE 2: Pre-operative radiographic images

(A) Intra-oral periapical radiograph with 12 showing a large periapical radiolucent shadow, (B) coronal cone-beam computed tomographic view showing wide canal and a radiolucent shadow with well-corticated borders measuring 13.78 mm superior-inferiorly and resorbed root apex with 12, (C) axial view showing the lesion measuring 10.48 mm mesiodistally and 10.56 mm buccopalatally, (D) axial view showing perforation of the labial and lingual cortical plate, (E) sagittal view of 12 showing root resorption with an apical diameter of 0.82 mm and loss of buccal cortical plate, (F) bony view showing a tunnel defect with 12, and (G) bony view showing a tunnel defect with 12.

Following the radiographic criteria for the diagnosis of periapical cysts (Figure 3) [11], the findings were suggestive of a periapical cyst of size  $10.48 \text{ mm} \times 10.56 \text{ mm} \times 13.78 \text{ mm}$  with 12.

Criteria for radiographic diagnosis (CBCT) of cyst	
Location	Apex of the involved tooth
Periphery	Well-defined corticated border
Shape	Curved or circular
Internal structure	• Radiolucent
Effect on surrounding structures	Displacement and root resorption with curved outline
Effect on surrounding bone	Cortical plate perforation
	Cortical plate perforation

Considering the history, clinical examination, and clinical and radiographic investigation, a diagnosis of an

FIGURE 3: Criteria for radiographic diagnosis of radicular cyst

CBCT- Cone-beam computed tomography

Image Credits: Paridhi Agrawal

infected periapical cyst with 12 was made. A treatment plan was formulated and explained to the patient, and the procedure commenced following the patient's informed consent.

Under rubber dam isolation, access opening was performed with 12. Thereafter, patency filling was done and working length was determined using an apex locator (Root ZX2, J.W. Morita, Japan) which was then confirmed radiographically. The working length of 12 was found to be 19.5 mm (Figure 4).

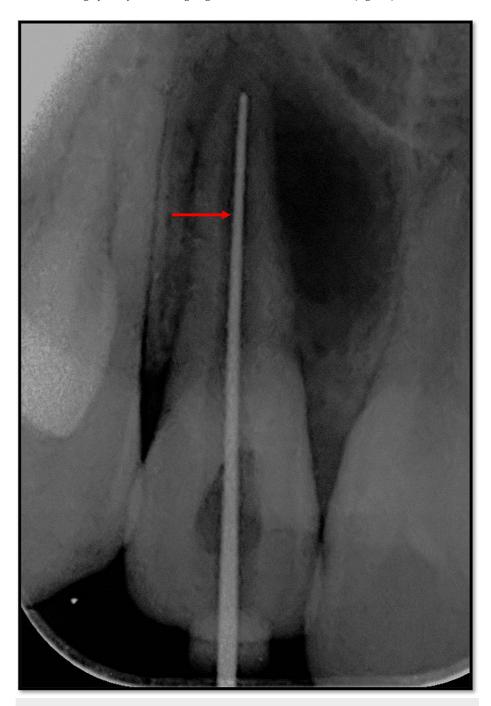


FIGURE 4: Radiographic confirmation of working length with 12

Considering the apical diameter of 0.82 mm, tooth 12 was lightly cleaned and shaped in circumferential strokes using a size 70 hand K-file (Mani, Japan). Over instrumentation two to three times 1mm beyond the apex was performed [12]. Ultrasonically activated irrigation using 0.9% normal saline, 2.5% sodium hypochlorite, and 2% chlorhexidine was performed followed by the placement of calcium hydroxide intracanal medicament dressing for 1 week with tooth 12. In the subsequent visit, complete healing of the sinus tract was seen. Following this, two additional applications of calcium hydroxide intracanal medication spaced at an interval of 1 week were done. In the fourth visit, the tooth was completely asymptomatic, there was no drainage from the canal, so it was decided to perform obturation with 12. A final irrigation with 17%

ethylenediamine tetraacetic acid (EDTA) and 0.9% normal saline was carried out with 12 and the canals were dried using absorbent paper points. Thereafter, 1.5 grams of MTA (ProRoot MTA, Dentsply Sirona, USA) was taken and placed in a 2.5mL syringe with 24 gauze needle. Miron's protocol [13] for the preparation of i-PRF was used. 9mL of the patient's blood was withdrawn and collected in plastic test tubes. It was then centrifuged at 700 rpm for three minutes. The obtained i-PRF in light yellow color was then extracted in the syringe containing MTA, lightly mixed, and placed in the periapical area of tooth 12. Subsequently, the canal was obturated using MTA mixed with i-PRF [14] in a wet sand consistency. The post-endodontic restoration was carried out using composite resin (3M Z250 XT, 3M, USA), and an IOPA radiograph was taken (Figure 5).

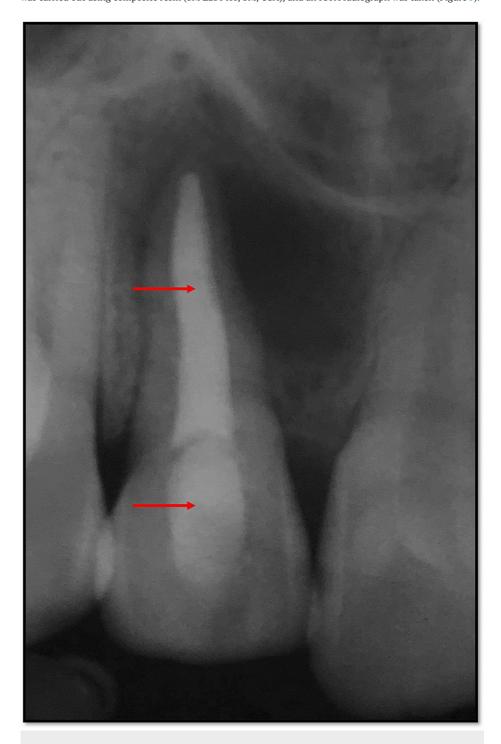


FIGURE 5: Radiograph showing mineral trioxide aggregate obturation and composite resin post-endodontic restoration with 12

As i-PRF is recognized for its sustained release of growth factors over a period of approximately 10 days [14], the patient was recalled after 10 days. During this appointment, a fresh mixture of MTA and i-PRF was

prepared using the same established protocol as previously mentioned. Subsequently, the mixture was slowly re-administered in the periapical area through the buccal mucosa, ensuring precise needle placement within the cystic cavity adjacent to tooth 12, as confirmed through both palpation and radiographic assessment (Figures *6A-6C*).



FIGURE 6: Extra-oral administration of combination of injectable platelet-rich fibrin and mineral trioxide aggregate

(A) Radiographic confirmation of needle placement in the cystic cavity, (B) re-administration of the mixture in the periapical area through the buccal mucosa, and (C) radiograph confirming the placement of the mixture in the periapical area.

The patient was then recalled for follow-ups after three, six, and nine months during which the clinical and radiographic examination revealed that the patient was completely asymptomatic, and the periapical lesion showed definite signs of healing (Figures 7A-7C).



FIGURE 7: Follow-up radiographs

(A) Three months follow-up, (B) six months follow-up, and (C) nine months follow-up

#### **Discussion**

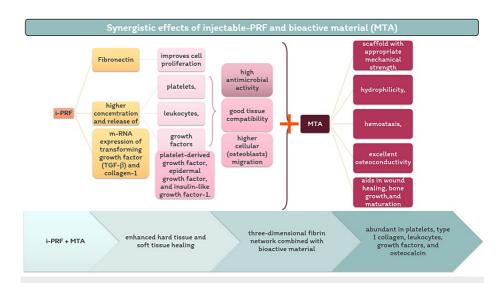
Periapical lesions are typically categorized as radicular cysts, dental granulomas, or abscesses [15]. Among these periapical lesions, the prevalence of cysts can range from 6% to 55% [16]. Similarly, the occurrence of granulomas varies between 9.3% and 87.1%, while abscesses are found in approximately 28.7% to 70.07% of cases [17]. It is important to note that based on clinical observations, larger-sized lesions are more likely to be radicular cysts. However, some of these sizable lesions may display characteristics resembling granulomas [18]. The conclusive and definite diagnosis of a cyst can be made only through histopathological findings. As shown in Figure 8, a primary clinical diagnosis of a radicular cyst can be approximately made based on certain facts [18,19].

# Criteria for preliminary diagnosis of radicular cyst periradicular lesion involving one or more teeth lesion is ≥200 mm<sup>2</sup>; with necrotic pulps; aspiration yielding a strawcolored fluid or drainage of presence of cholesterol such fluid through an crystals in the fluid; access; lesion is seen radiographically as a circumscribed, well-defined radiolucent area bound by a thin radiopaque line. FIGURE 8: Criteria for preliminary diagnosis of radicular cyst Image Credits: Paridhi Agrawal

Additionally, studies have indicated that the occurrence of cysts can be as high as 60%-67% in lesions with a diameter ranging from 10 to 20 mm [20,21]. When taking into account the volume of the lesion, there is an 80% likelihood of it being a cyst when the volume exceeds  $247 \text{ mm}^3$ . In cases where there is root displacement and the volume is less than  $247 \text{ mm}^3$ , there is still a 60% probability of it being a cyst [22].

Nonsurgical management of periapical lesions is the preferred approach when compared to surgical methods and should be contemplated in all cases [23]. This preference is rooted in the potential risks associated with surgical procedures, such as the risk of harming adjacent vital teeth and damage to anatomical structures in the proximity of the lesion. Additionally, the discomfort and pain often associated with surgical interventions can be avoided through nonsurgical means. Patient comfort and their reluctance towards surgical procedures, along with considerations like age and medical conditions that may limit surgical options, further support the choice of a nonsurgical approach. Surgical interventions should only be considered as a last resort when conventional methods have proven ineffective [23].

Calcium hydroxide is a widely utilized intra-canal medication due to its notable high alkalinity [24] and its ability to effectively combat bacteria [25]. It is generally recommended to apply calcium hydroxide within the root canals and surrounding periradicular tissues, especially in cases involving sizable and chronic periapical lesions producing 80.8% success rate. This substance is believed to exert a direct influence on inflamed tissues and epithelial cystic linings, ultimately promoting periapical healing and facilitating bone repair [26]. In our case, calcium hydroxide was introduced into the root canal for a duration of three weeks, resulting in the alleviation of symptoms. As shown in Figure 9, in this case, we have employed a novel technique utilizing the synergistic effect of i-PRF combined with MTA.



# FIGURE 9: Synergistic effects of injectable platelet-rich fibrin and mineral trioxide aggregate

PRF- platelet-rich fibrin, i-PRF- injectable platelet-rich fibrin, MTA- mineral trioxide aggregate

Image Credits: Paridhi Agrawal

An innovative formulation of Platelet-Rich Fibrin (PRF), referred to as "injectable" PRF or i-PRF, has been introduced with the aim of simplifying the application of concentrated platelets in a liquid form. This formulation can be used either independently or in combination with bioactive materials, offering promising prospects for the enhancement of both soft and hard tissue healing. It achieves this through the creation of a three-dimensional fibrin network that is abundant in platelets, type I collagen, leukocytes, growth factors, and osteocalcin [27].

i-PRF is prepared by employing a low-speed centrifugation concept, which significantly enhances the concentration of regenerative cells, leukocytes, platelets, and growth factors when compared to other PRF formulations that use higher centrifugation speeds [28]. Additionally, i-PRF plays a pivotal role in tissue healing by activating the host defense system due to its high platelet and leukocyte content. This high leukocyte count contributes to its antimicrobial potency against bacterial lipopolysaccharides when compared to other platelet concentrates. Furthermore, leukocytes are instrumental in wound healing by directing and recruiting various cell types [29]. Importantly, i-PRF contains a unique combination of proteins with diverse biochemical signals, which may work synergistically to exert a positive biological impact on tissue regeneration and mineralization. As a result, i-PRF is a promising option for tissue engineering, harnessing autologous cues to promote the regeneration of hard tissues [30].

The benefits of combining i-PRF with bioactive materials (such as MTA) include serving as a scaffold material with appropriate mechanical strength, hydrophilicity, and excellent osteoconductivity. It aids in wound healing, bone growth, maturation, material stabilization, and hemostasis, as well as improving the handling properties of bioactive materials [31]. Therefore, the combination of i-PRF and MTA for non-surgical healing of periapical lesion was used.

# **Conclusions**

This case report demonstrates the successful non-surgical management of a large periapical lesion through a novel combination of i-PRF and MTA. This innovative approach offers an effective alternative to traditional surgical interventions, emphasizing patient comfort and optimal tissue healing. By harnessing the regenerative potential of i-PRF and the osteoconductivity of MTA, this technique showcases promising prospects for periapical lesion treatment. Further research and clinical validation will be essential to establish its broader applicability in endodontic practice.

#### **Additional Information**

#### **Author Contributions**

All authors have reviewed the final version to be published and agreed to be accountable for all aspects of the work.

Concept and design: Paridhi Agrawal, Jay Bhopatkar, Pradnya Nikhade, Aditya Patel, Tejas Suryawanshi

**Acquisition, analysis, or interpretation of data:** Paridhi Agrawal, Jay Bhopatkar, Pradnya Nikhade, Aditya Patel, Tejas Suryawanshi

**Drafting of the manuscript:** Paridhi Agrawal, Jay Bhopatkar, Pradnya Nikhade, Aditya Patel, Tejas Suryawanshi

**Critical review of the manuscript for important intellectual content:** Paridhi Agrawal, Jay Bhopatkar, Pradnya Nikhade, Aditya Patel, Tejas Suryawanshi

Supervision: Tejas Suryawanshi

#### **Disclosures**

**Human subjects:** Consent was obtained or waived by all participants in this study. **Conflicts of interest:** In compliance with the ICMJE uniform disclosure form, all authors declare the following: **Payment/services info:** All authors have declared that no financial support was received from any organization for the submitted work. **Financial relationships:** All authors have declared that they have no financial relationships at present or within the previous three years with any organizations that might have an interest in the submitted work. **Other relationships:** All authors have declared that there are no other relationships or activities that could appear to have influenced the submitted work.

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