

# Mineral trioxide aggregate in paediatric dentistry

VIDYA SRINIVASAN<sup>1</sup>, PAULA WATERHOUSE<sup>2</sup> & JOHN WHITWORTH<sup>3</sup>

<sup>1</sup>Department of Paediatric Dentistry, Edinburgh Dental Institute & Royal Hospital for Sick Children, Edinburgh, UK, and School of Dental Sciences, Newcastle University, Newcastle upon Tyne, <sup>2</sup>The Department of Child Dental Health and Paediatric Dentistry, and <sup>3</sup>The Department of Restorative Dentistry, School of Dental Sciences, Newcastle University, Newcastle upon Tyne, UK

*International Journal of Paediatric Dentistry* 2009; 19: 34–47

**Objective.** The aim of this study was to present a review of the reported literature on: (i) the physical and chemical properties; and (ii) clinical applications of mineral trioxide aggregate (MTA) in the practice of paediatric dentistry.

**Method.** Electronic literature search of scientific papers from January 1993 to June 2008 was carried out on the MEDLINE, Embase, Entrez Pubmed, and Scopus databases using specific key words. The search yielded 448 papers, out of which 100 were identified as conforming to the applied criteria.

These papers formed the basis of the review and the clinical scenarios presented which demonstrate the application of MTA in the practice of paediatric dentistry.

**Conclusion.** Paediatric dentists have successfully employed MTA in a variety of endodontic/restorative applications since the late 1990s. Clinical impressions have generally been favourable and support the findings of laboratory and animal-based investigations. Very few clinical studies have been reported so far in humans, and although these have been positive, the body of research is currently insufficient to enable a meaningful systematic review and meta-analysis.

## Introduction

Mineral trioxide aggregate (MTA) was developed at the Loma Linda University, California, USA, as a root-end filling material in surgical endodontic treatment<sup>1</sup>. Over the years, further research on the material has resulted in MTA being applied in various clinical situations in addition to its use as a suitable root-end filling material. The diverse application of MTA in the practice of paediatric dentistry is evident in its use as an apical barrier in immature non-vital teeth and in the coronal fragment of fractured roots, as a pulpotomy medicament in primary and permanent teeth, a pulp-capping agent in young permanent teeth, and as a repair material for perforation and resorptive defects.

## Method

An electronic literature search of scientific papers from January 1993 to June 2008 was carried out using the MEDLINE, Embase, Entrez Pubmed, and Scopus databases. These databases were used to search for the key words Mineral Trioxide Aggregate, MTA, Gray MTA, Grey MTA, White MTA, GMTA, WMTA, and mineral AND trioxide AND aggregate. All papers on MTA, which reported studies carried out *in vitro*, *in vivo*, and *ex vivo* on tissues, animals, and humans were included for reviewing. Papers not published in the English language and case reports were excluded. Use of the search key-words produced a total of 448 results. Manual checking of the reference list and application of the inclusion and exclusion criteria produced 100 citations used in this review.

## Review of MTA

The review is presented in two main sections. The first section reviews the literature which has contributed to our understanding of the

### Correspondence to:

Vidya Srinivasan, Department of Paediatric Dentistry, Edinburgh Dental Institute, Lauriston Building, Lauriston Place, Edinburgh EH3 9HA, UK. E-mail: vidya.srinivasan@nhslothian.scot.nhs.uk

physical and chemical properties of MTA. The second section reviews the clinical studies which have been reported in the search period with case reports depicting the applications of MTA in paediatric dentistry.

The various commercially available products employed in the studies which are reviewed in the first section of this paper include: (i) ProRoot MTA (Dentsply Tulsa Dental, Tulsa, OK, USA); (ii) White ProRoot MTA (Dentsply Tulsa Dental); (iii) MTA-Angelus (Solucos Odontologicas, Londrina, Brazil); (iv) MTA-Angelus Blanco (Solucos Odontologicas); and (v) MTA Bio (Solucos Odontologicas).

### Chemical and physical properties of MTA

*Chemical composition and structure.* Studies analysing the constituents of both grey and white ProRoot MTA have conclusively shown that both materials are similar to Portland cement, but with bismuth oxide added, presumably to make the materials radiopaque for dental use<sup>2-4</sup>. Portland cement itself is a mixture of dicalcium silicate, tricalcium silicate, tricalcium aluminate, gypsum, and tetracalcium aluminoferrite<sup>5,6</sup>.

It was reported that the amount of gypsum in ProRoot MTA is approximately half the amount found in Portland cements. Gypsum is an important determinant of setting time and is added by cement manufacturers to retard the setting time of the cement clinker in Portland cement. Modifying the gypsum content in the MTA mix can result in significant reduction of setting time, thereby reducing the number of treatment visits<sup>7</sup>. Dammaschke *et al.* described the marked differences between Portland cement and ProRoot MTA, by comparing the same chemical and physical surface, and bulk material characteristics<sup>6</sup>. They reported that ProRoot MTA contains lower levels of potentially toxic heavy metals (e.g. manganese and strontium), chromophores (iron oxide), aluminium, and potassium. In contrast to Portland cements, ProRoot MTA contains about 17–18wt% (= 2%) bismuth. Portland cements are composed of particles with a wide range of sizes, whereas ProRoot MTA showed smaller and more uniform particle size<sup>6</sup>.

Studies have compared the constituents of grey and white ProRoot MTA materials. There

are contradictory reports with respect to the iron and magnesium oxide phases in the grey and white forms. Asgary *et al.* reported that white MTA contained significantly less amounts of oxides of iron, aluminium, and magnesium than grey MTA<sup>8</sup>. There have been other studies, which have reported the complete absence of iron oxide in white MTA when compared to grey MTA<sup>9,10</sup>. When comparing the ProRoot MTA forms to MTA-Angelus, Song *et al.* also reported that MTA-Angelus had a lower content of bismuth oxide than the ProRoot MTAs<sup>10</sup>. There are no studies to date comparing the relative radiopacity of MTA-Angelus with the ProRoot MTAs.

*Setting/Hardening time.* There are few published reports of experimental data relating to the comparative setting times of the different forms of MTA. The setting time of grey ProRoot MTA was reported by Torabinejad *et al.* as 2 h and 45 min ( $\pm 5$  min)<sup>1</sup>. Islam *et al.* reported final setting times of 140 min (2 h and 20 min) for white MTA, and 175 min (2 h and 55 min) for grey MTA<sup>11</sup>. Although the manufacturers of MTA-Angelus claim that this material has a setting time of 10 min, there appears to be no independent evidence to confirm this.

Several studies have compared various modified forms of Portland cement and ProRoot MTA in an effort to identify a material with all the advantages of MTA and without its extended setting time. The presence of gypsum is reported to be the reason for the extended setting time of MTA. Therefore, studies<sup>7</sup> have reported modified forms of Portland cement without the gypsum and with added plasticizers, which reportedly do not affect the biocompatibility of MTA. In order to reduce the setting time, the effect of accelerators such as sodium phosphate dibasic ( $\text{Na}_2\text{HPO}_4$ ) and calcium chloride ( $\text{CaCl}_2$ ) are being investigated currently<sup>12-15</sup>. MTA Bio is one commercially available product which incorporates an accelerator of this sort, and is promoted as a rapid-setting material.

*Compressive strength.* Compressive strength is the capacity of a material to withstand axially directed pressure generating compressive stress as a result of compression force. Torabinejad

*et al.* reported comparable mean compressive strength after 21 days for ProRoot Grey MTA, IRM, and Super EBA<sup>1</sup>. The compressive strength of amalgam was higher than these materials after the same time period. Root-end fillings and apical barrier materials do not bear direct pressure during function, hence, their compressive strength is not thought as important as those materials used to repair or restore defects in load-bearing sites. The compressive strength of ProRoot grey MTA increased with time in the above study. The authors suggest that this increase over a period of time required the presence of moisture. Islam *et al.* reported greater compressive strength for the grey form of ProRoot MTA in comparison to the white form at 3 days and 28 days in a similar *in vitro* study<sup>11</sup>. Their study also showed that the compressive strength of the grey form of MTA was greater than that of Portland cement.

**Radiopacity.** An ideal restorative material should be more radiopaque than its surrounding structures when placed *in situ*, in order to allow the quality of the restoration or apical seal to be assessed. Several studies have confirmed that MTA is less radiopaque than Super EBA, IRM, amalgam, and conventional gutta-percha, but in the same range as zinc oxide–eugenol-based root canal sealers<sup>1,16,17</sup>.

**Setting conditions.** Studies have reported that an initial period of exposure to moisture or humidity is required for the MTA to achieve optimum flexural strength<sup>18,19</sup>. These authors recommend the placement of a moistened cotton pellet in the root canal for a period of time before placement of the permanent coronal seal when placing an apical barrier in immature teeth. Currently, there is insufficient literature regarding the implications of the setting conditions on the use of MTA as a pulp capping and pulpotomy medication. The role of moisture drawn in from the pulp or peri-radicular tissues is also unclear.

**Solubility.** The manufacturer recommends the use of 0.33 g of water with 1 g of ProRoot MTA to achieve an optimum mix of the material. Earlier studies showed no signs of solubility of ProRoot MTA in water when tested under

modified International Organization for Standardization (ISO) and American Dental Association specifications<sup>1</sup>. Fridland and Rosado demonstrated that both solubility and porosity of the material show a significantly increasing trend that follows the amount of water used when preparing the mix under ISO specifications<sup>20,21</sup>. The set or hardened/cured material, on exposure to water, was shown to release calcium as hydroxide, and the authors reported that their finding could explain the basis of the cementogenesis-inducing property of MTA. These studies also suggest that the water-to-powder ratio recommended by the manufacturer (0.33) would be the ideal proportion. Santos *et al.* reported that calcium and hydroxyl ions may be released from MTA Angelus during storage in moist conditions for periods up to 360 h<sup>22</sup>.

**Marginal adaptation and sealing ability.** An effective root-end filling material should ideally provide a hermetic apical seal, preventing the movement of tissue fluids into the root canal system and the egress of micro-organisms and their by-products from the root canal system<sup>23</sup>. Investigations have been carried out on extracted human teeth which were prepared and restored with the root-end filling materials whose marginal adaptation and sealing ability were investigated by various methods. Historically, investigators have evaluated quality of the apical seal by the degree of dye, radioisotope, or bacterial penetration; electrochemical means; scanning electron microscopy (SEM); or fluid filtration. Each technique has significant limitations that can result in errors<sup>24–27</sup>. As a result, the validity of such data is questionable.

The results of some studies demonstrate correlation with respect to marginal adaptation with SEM and apical seal dye penetration for different root-end fillings<sup>28,29</sup>. These studies compared amalgam, Super EBA, IRM, and MTA, and were the earliest investigations comparing MTA to other traditional root-end filling materials. There are, however, conflicting results in the literature relating to the correlation between marginal adaptation and sealing ability of different root-end filling materials<sup>30,31</sup>.

Several investigators report that the fluid filtration method has the advantage of

measuring the cumulative leakage of the entire tooth restoration interface and is therefore quantitative<sup>32–34</sup>. Bates *et al.* reported that MTA was superior to amalgam and comparable with Super EBA in preventing microleakage when used as a root-end filling by the fluid filtration method on extracted human teeth<sup>34</sup>.

Almost a decade later, Shipper *et al.* compared MTA with amalgam as a root-end filling material with a high and low vacuum SEM study on extracted human teeth<sup>35</sup>. Results showed that MTA demonstrated better marginal adaptation to the root end cavity wall than amalgam. This group of workers purport their findings to be linked to the inherent nature of MTA, and suggested that the expansion of the material during the hydration setting reaction contributed to the superior adaptation to dentine. The authors concluded that this expansion may play a role in the increased incidence of cracks at the interface compared to the amalgam specimens.

Investigators have also compared MTA with amalgam, Super EBA, and IRM with *in vitro* studies using specific bacterial leakage tests. The micro-organisms used in such tests have included *Staphylococcus epidermidis* and *Serratia marcescens*<sup>36,37</sup>. These studies have been consistent in reporting MTA as showing no or less leakage in comparison to the other three materials, respectively.

The peri-radicular environment may have varying pH from a neutral pH of 7.4 to an acidic pH as low as 5.0. An acidic pH has been shown to inhibit the setting reactions, affect adhesion, and increase solubility of materials placed to effect a root-end seal<sup>38–41</sup>. Roy *et al.* compared the sealing ability of materials tested by recording the linear dye leakage with Pelikan Ink under a surgical microscope, and reported that an acidic environment does not hinder the sealing ability of MTA, amalgam, Geristore, Super-EBA, CPS, and MTA with CPC matrix<sup>42</sup>.

The protein leakage and assay test is claimed to provide the advantage of eliminating the problems involved with radio-isotopes, dye, and bacteria during leakage identification. Valois and Costa studied the influence of the thickness of MTA on the sealing ability of root-end fillings *in vitro* by using a protein-dye

complex with Coomassie Blue G dye. It was shown that 4-mm-thick MTA was significantly more effective than others (1, 2, 3 mm) in preventing apical leakage<sup>43</sup>.

Calcium hydroxide intra-canal medication has been shown to affect the sealing ability of MTA<sup>44</sup>. The *in vitro* sealing efficiency of white and grey ProRoot MTA as apical barriers was investigated in simulated divergent apices using a dye tracer (basic fuchsin). A comparable apical seal with both forms of ProRoot MTA was reported. It was also shown that residual calcium hydroxide intra-canal medication could interfere with the adaptation of MTA to the root canal walls by being a mechanical obstacle, and also by chemically reacting with MTA, thus influencing its surface characteristics.

The use of the internal matrix concept to limit the flow of the MTA material (in the root-end and perforation situations) and improve its sealing ability has been investigated. Zou *et al.* reported an *in vitro* study to evaluate internal matrices as barriers to prevent the over-extension of MTA. They reported that calcium sulphate provided a successful barrier against over-extension of MTA, but significantly decreased its sealing ability, and Collaplug (collagen plug) did not prevent over-extension or improve its sealing ability<sup>45</sup>.

The validity of 'leakage studies' has recently been brought into sharp focus by a decision of the *Journal of Endodontics* to place a moratorium on the acceptance of such studies until more is known of their relevance<sup>46</sup>.

*Effect of compaction/condensation on MTA.* Condensation pressure or compaction is an uncontrolled variable when MTA is placed as an apical barrier in an immature tooth, surgical or non-surgical perforation repair, a pulp-capping material, or as a retrograde filling material in a root-end cavity. It is likely that the condensation pressure during the placement of MTA as an apical barrier will be much reduced to prevent the material from being forced into the periodontal ligament or pulp tissue in some of these situations. Nekoofar *et al.* reported no statistically significant effect of condensation pressure on the compressive strength of white ProRoot MTA, but there was a significant reduction in surface hardness<sup>47</sup>.

This group of workers also concluded that greater condensation pressures could limit the space for the ingress of water required to hydrate the material in order to achieve an adequate surface hardness.

*Effect of MTA on the strength and hardness of root dentine.* Studies using sheep and bovine teeth (*in vitro* and *in vivo*) have shown that teeth with an MTA apical barrier and MTA root filling showed higher fracture resistance in comparison to teeth that had calcium hydroxide placed as an intra-canal medicament<sup>48–51</sup>. It has been previously suggested that the conventional apexification technique involving the placement of calcium hydroxide in the root canal for a prolonged period of time may be responsible for an increased susceptibility of immature incisors to root fracture<sup>52</sup>. In 2002, Andreasen *et al.* reported that long-term calcium hydroxide dressings weaken the root structure possibly by neutralizing, denaturing, or dissolving the acidic components of dentine<sup>53</sup>. Because these components act as 'bonding agents' between the collagen network and the hydroxyapatite crystals, their destruction may render the tooth more prone to fracture. In light of this evidence, the placement of MTA as a one-step apical barrier after appropriate debridement of the root canal, may be a suitable alternative to traditional apexification or the induced apical barrier technique. As with any other material, inadequate debridement and canal preparation could lead to persistent symptoms even after the apical seal has been placed with MTA. Once the MTA placed as an apical seal has hardened, it becomes extremely difficult to remove, and persistent symptoms can result in the need for surgery or even tooth extraction.

*Antibacterial and antifungal activity.* As micro-organisms are the main aetiological factors in pulpitis and periodontitis, their elimination during treatment is essential. It is likely that even after caries removal and root canal debridement, micro-organisms with the potential to promote disease may persist, or new organisms may enter by coronal leakage<sup>54</sup>. The healing of tissues damaged by pulp or peri-apical pathology depends on the absence of irritating

agents originating from microbial metabolic products, or of chemical origin from the sealing materials. For such healing to occur, the materials placed in contact with healthy pulp (pulp capping, pulpotomy) and peri-apical tissues (apical barrier, root-end filling) should not damage the tissues and should ideally stimulate the deposition of hard tissue, therefore promoting biological sealing<sup>55,56</sup>. MTA materials fulfil this requirement adequately.

Several studies have been carried out to ascertain the antibacterial and antifungal properties of the MTA cements. Sipert *et al.* reported that MTA-Angelus did not inhibit the growth of *Escherichia coli* in an *in vitro* study<sup>57</sup>. In 2006, Al-Hazaimi *et al.* assessed the antibacterial effects of the grey and white MTA materials against *Enterococcus faecalis* and *Streptococcus sanguis in vitro*. They reported that lower concentrations of grey MTA were required than the white MTA to exert the same antibacterial effect against each of these micro-organisms<sup>57</sup>. Eldeniz *et al.* also obtained similar results<sup>58</sup>. *Enterococcus faecalis* is one of the organisms more likely to be found in cases of failed endodontic therapy than in primary infections, and it is likely that on this basis that all reported literature regarding antibacterial activity of root filling materials is pertaining to this organism.

*Reactions with other dental materials.* In an effort to offset the extended setting time of MTA, researchers have reported various alternatives to the placement of a moist cotton pellet over the setting MTA material. An *in vitro* study conducted by Nandini *et al.* reported that conventional glass ionomer cement can be layered over partially set MTA for a single-visit procedure and that the setting of MTA proceeds unhindered underneath the layer of glass ionomer<sup>59</sup>. In such a procedure, Ballal *et al.* reported their observations on a glass ionomer cement layered upon partially set MTA. The setting of both materials was unaffected, and the glass ionomer cement showed no signs of dehydration<sup>60</sup>. Tunc *et al.* evaluated the bond strength of a composite and a compomer to white MTA using different bonding systems<sup>61</sup>. They concluded that the total-etch one-bottle system mediated a stronger bond to white

MTA than the self-etch one-step system. The conclusions from these studies bear relevance to the use of MTA as a pulp-capping and pulpotomy wound-dressing material.

**Biocompatibility.** The biocompatibility of MTA has been reported widely over the past decade by researchers involved in *ex vivo* cell culture studies and *in vivo* studies in animals and humans.

- 1) Subcutaneous and intra-osseous evaluation: Studies in the late 1990s reported bacterial and cell culture assays, respectively, to conclude that MTA was not mutagenic or cytotoxic<sup>62,63</sup>. There have been several studies since then, which have tested samples of MTA as subcutaneous and intra-osseous implants in rats<sup>64–66</sup>, guinea pigs<sup>67</sup>, and rabbits<sup>68</sup>. These studies reported minimal inflammatory responses in the soft tissue and bone, and confirmed MTA to be capable of inducing osteogenesis.
- 2) Animal studies: The biocompatibility of MTA has also been studied *in vivo* as root-end fillings in dogs<sup>69–72</sup> and monkeys<sup>73,74</sup>. These studies reported satisfactory peri-apical tissue responses and healing with MTA. Animal studies have also reported MTA as a favourable pulp-capping material following traumatic exposures in monkeys and dogs<sup>75–77</sup>. MTA has been evaluated *in vivo* in rats as a pulpotomy medicament in comparison to formocresol and ferric sulphate, and reported to perform ideally as a pulpotomy agent, causing dentine bridge formation and simultaneously maintaining normal pulpal histology<sup>78</sup>.

**Cost implications and storage.** Important barriers to the widespread use of MTA in paediatric dentistry include its perceived cost, difficulties with storage, and the need for appropriate training.

The list prices of some commercial products are as follows:

- 1) ProRoot MTA (White) 5 Dose Pack (ProRoot MTA) – £210.00 (€269.93) (USD363.41)
- 2) ProRoot MTA (White) 2 Dose Pack (ProRoot MTA) – £91.00 (€116.97) (USD157.48)
- 3) MTA- Grey and White 1G Pack 7 Applications (MTA-Angelus) – £37.50 (€48.20) (USD64.89)

- 4) MTA-Grey 2G 14 Applications (MTA-Angelus) – £70.00 (€89.97) (USD 121.13) (Figures courtesy of QED, Peterborough, Cambs, UK).

ProRoot products are supplied in single-dose sachets, whereas Angelus products are supplied in double-sealed glass vials.

The composition of MTA is not unlike that of the cement used in the building industry to make concrete. Such a material should be kept dry during storage because moist air leads to the phenomenon of air setting, which reduces the strength of the mix. The presentation of ProRoot products as a 1 g sachet for single use, would result in considerable wastage of material, and the transfer of this material to a sealed container such as an Eppendorf tube (Eppendorf UK Ltd, Cambridge, UK) would extend the life of the material and allow more than one treatment to be completed from a single 'dose'.

The Angelus vials of material are marketed with guidance that 1 g may allow up to seven treatments, depending of course on the volume of material to be used.

### Clinical applications of MTA in paediatric dentistry

There is a paucity of *in vivo* human studies on the performance of MTA in its various clinical applications. This section will review these studies. Table 1 outlines the evidence available from human clinical studies for the use of MTA in its various clinical applications.

#### Pulp treatment in permanent teeth

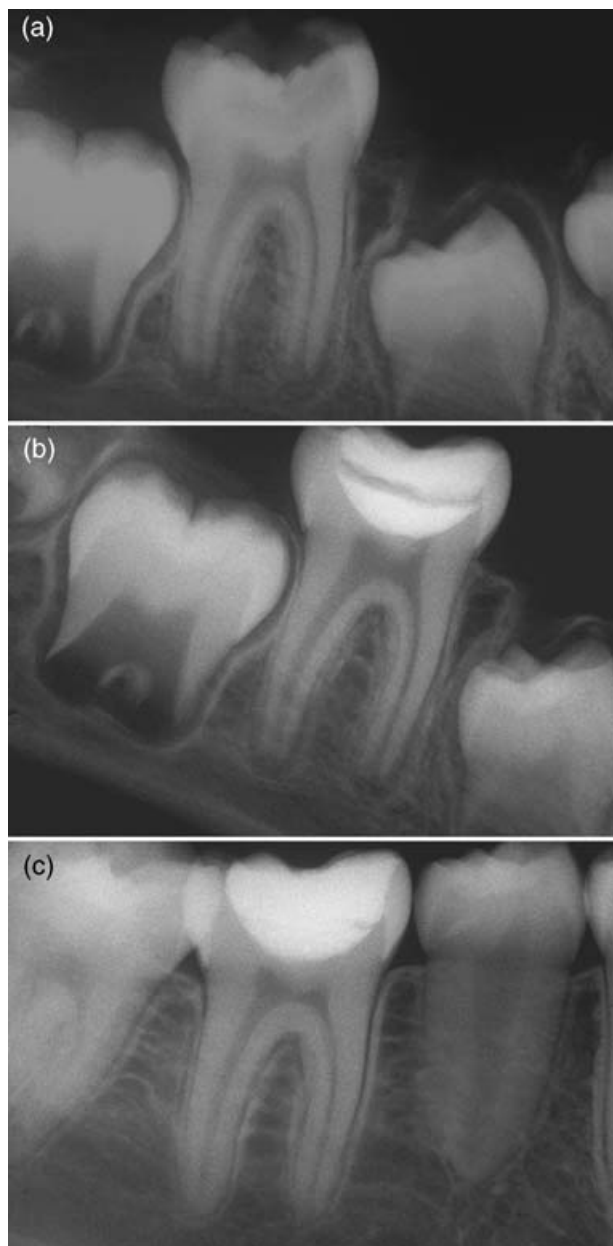
- (i) Pulp capping: There have been few studies to date on MTA as a pulp-capping agent in human permanent molars. Four prospective human clinical studies have compared MTA and calcium hydroxide as pulp-capping medicaments in third permanent molars following the *mechanical exposure* of healthy pulps<sup>79–82</sup>. Aeinehchi *et al.* reported a 0.28-mm-thick dentine bridge in teeth pulp capped with grey MTA at 2 months, and 0.43 thickness at 6 months in contrast to a 0.15-mm-thick dentine bridge noted with calcium hydroxide at 6 months<sup>79</sup>. This study also reported tissue inflammation and adjacent pulp tissue necrosis

**Table 1. Evidence from human clinical studies for the clinical applications of mineral trioxide aggregate (MTA).**

Clinical application	Studies	Evidence
Pulp capping – permanent teeth	Prospective studies comparing MTA to calcium hydroxide (Ca(OH) <sub>2</sub> )	<i>Studies in teeth capped after mechanical pulp exposures</i> (Validation by histopathology)
	– Aeinehchi <i>et al.</i> 2003 <sup>79</sup>	Thicker dentinal bridge, less pulpal inflammation with MTA
	– Iwamoto <i>et al.</i> 2006 <sup>80</sup>	No significant difference between MTA and Ca(OH) <sub>2</sub>
	– Nair <i>et al.</i> 2008 <sup>81</sup>	Thicker dentinal bridge, less inflammation with MTA
	– Accorinte Mde L <i>et al.</i> 2008 <sup>82</sup>	Initial healing is better with MTA; subsequent healing similar in MTA and Ca(OH) <sub>2</sub>
	MTA – observational study	<i>Study in teeth capped after carious pulpal exposures</i> Clinical success outcomes
Pulpotomy – permanent teeth	– Bogen <i>et al.</i> 2008 <sup>83</sup>	97.6% of the sample showed favourable outcomes; all immature teeth showed subsequent complete root formation
	MTA – observational study	<i>Studies in teeth with pulpal exposures due to caries/fractures</i> Clinical success outcomes
	– Barrieshi-Nusair and Qudeimat 2006 <sup>84</sup>	
	– Witherspoon <i>et al.</i> 2006 <sup>85</sup>	< 75% of the sample in both studies showed favourable outcomes
	Prospective studies comparing MTA to formocresol	
Pulpotomy – primary teeth	– El-Meligy and Avery 2006 <sup>86</sup>	Similar clinical and radiographic outcomes
	Prospective studies comparing MTA to formocresol	<i>Studies in teeth with carious pulpal exposures</i> Clinical success outcomes
	– Eidelman <i>et al.</i> 2001 <sup>87</sup>	Grey: > 90% over 6–30 months
	– Agamy <i>et al.</i> 2004 <sup>88</sup>	Grey: 100% over 1 year. White: 84.2% over 1 year
	– Holan <i>et al.</i> 2005 <sup>89</sup>	Grey: 97% over 4–72 months
	– Farsi <i>et al.</i> 2005 <sup>90</sup>	Grey: 100% over 2 years
	– Aeinehchi <i>et al.</i> 2007 <sup>91</sup>	Grey: 100% over 6 months
	– Noorollahian, 2008 <sup>92</sup>	White: 100% over 2 years
	Prospective longitudinal studies	
	– Maroto <i>et al.</i> 2005 <sup>93</sup>	Grey: 100% over 6 months
	– Maroto <i>et al.</i> 2007 <sup>94</sup>	White: 100% over 42 months
	Prospective study comparing MTA to Ca (OH) <sub>2</sub>	
	– Percinoto <i>et al.</i> 2006 <sup>95</sup>	Grey: 95 % over 1 year
	Prospective study comparing MTA to FC and Ca (OH) <sub>2</sub>	
	– Moretti <i>et al.</i> 2008 <sup>96</sup>	Grey: 100% over 2 years
Pulp Capping – Primary teeth	Prospective study comparing MTA to calcium hydroxide (Ca(OH) <sub>2</sub> )	Study in teeth with pulpal exposures due to caries
	– Tuna and Olmez 2007 <sup>97</sup>	Clinical and radiographic outcomes were comparable for MTA and Ca(OH) <sub>2</sub> . Histological validation recommended.
Apical barrier –permanent teeth		<i>Studies in non-vital immature teeth</i>
	– Simon <i>et al.</i> , 2008 <sup>98</sup>	Radiographic success outcome
	– Saris <i>et al.</i> 2008 <sup>99</sup>	81 % over 1 year
Root fractures – permanent teeth		76.5 % over 1 year
	Case reports only available at the time of this review	–

with calcium hydroxide at 6 months, and no pulp tissue inflammation adjacent to MTA with a near-regular odontoblastic layer for the same duration. These results were not significant due to the small sample size. A similar study by Iwamoto *et al.* compared white MTA with calcium hydroxide at 30 days and 136 days post-treatment<sup>80</sup>. At these evaluation periods, no significant difference was found

between the groups with regard to the clinical presentation and the histological status. Initial results indicate that both grey and white MTA may perform as well as calcium hydroxide in non-carious mechanical pulp exposures in permanent teeth with normal pulp tissue. In contrast, Nair *et al.* reported that MTA resulted in less pulpal inflammation and more predictable hard tissue barrier formation in permanent



**Fig. 1.** (a) Radiographic image prior to treatment in a mandibular molar with deep caries and immature apices in a 9-year-old patient (Bogen G, Kim JS, Bakland LK. Direct pulp capping with mineral trioxide aggregate. An observational study. *J Am Dent Assoc* 2008; **139**(3): 313. Copyright ©2008 American Dental Association. All rights reserved. Reprinted by permission.). (b) Radiographic image of the tooth with mineral trioxide aggregate as a pulp-capping agent. (Bogen G, Kim JS, Bakland LK. Direct pulp capping with mineral trioxide aggregate. An observational study. *J Am Dent Assoc* 2008; **139**(3): 313. Copyright ©2008 American Dental Association. All rights reserved. Reprinted by permission.). (c) Radiographic image of the tooth at 5.5-year recall showing a permanent restoration and evidence of complete root formation. The tooth exhibited a normal response to cold testing. (Bogen G, Kim JS, Bakland LK. Direct pulp

teeth in comparison to hard-setting calcium hydroxide<sup>81</sup>. Accorinte Mde *et al.* also reported their support of the safety of MTA for pulp capping in human teeth<sup>82</sup>. Their study reported that MTA seemed to heal the pulp tissue at a faster rate than calcium hydroxide cement, although after 60 days both materials reached similar and excellent results for pulp capping in human teeth.

Bogen *et al.* have reported an observational study where MTA was placed over *carious exposures* in permanent teeth with reversible pulpitis, over a 9-year period<sup>83</sup>. They followed 49 teeth over this period and reported favourable outcomes of 97.6% of the sample based on radiographic appearance, subjective symptoms, and cold testing. All teeth in younger patients (15/15) that initially had open immature apices showed complete root formation (apexogenesis) (Fig. 1).

(ii) Pulpotomy: At the time of this review, three studies had been reported on the outcome of MTA as a wound dressing following pulpotomy in permanent teeth<sup>84–86</sup>. Two of these were observational studies reporting a small sample of 28 and 23 teeth, respectively<sup>84,85</sup>. Barrieshi-Nusair and Qudeimat evaluated the success of grey MTA for partial pulpotomy in 28 cariously exposed young permanent first molars<sup>84</sup>. This was an observational study over a period of 12–26 months with an average of 17.5 months. The authors reported that 79% of the teeth tested positive to sensibility testing with no clinical or radiographic failures. Seven of the teeth which had immature apices at the beginning of treatment showed continued root maturation. A similar outcome was also reported by Witherspoon *et al.* in permanent molars with clinical signs of irreversible pulpal disease<sup>85</sup>. El-Meligy and Avery reported a study comparing MTA and calcium hydroxide as pulpotomy agents in young permanent teeth<sup>86</sup>. This was a split-mouth study, in which 15 pairs of young permanent molar teeth in the same number of children were followed-up for a period of 12 months.

capping with mineral trioxide aggregate. An observational study. *J Am Dent Assoc* 2008; **139**(3): 313. Copyright ©2008 American Dental Association. All rights reserved. Reprinted by permission.).



The authors reported similar clinical and radiographic success for MTA and calcium hydroxide as pulpotomy agents in immature permanent teeth, and concluded that MTA was a suitable alternative to calcium hydroxide.

*Pulp treatment in primary teeth.* (i) Pulpotomy: Several studies have evaluated MTA as a wound dressing following pulpotomy in primary teeth. Six of these studies have compared formocresol to the two forms of MTA (grey and white), and reported MTA to be an acceptable alternative to formocresol as a wound dressing in the pulpotomy of primary teeth<sup>87–92</sup>.

Maroto *et al.* have reported longitudinal and observational clinical studies on grey and white MTA<sup>93,94</sup>. Both studies reported favourably on the clinical outcomes with these materials as a pulpotomy medicament in primary teeth. Percinoto *et al.* have compared MTA to calcium hydroxide as a pulpotomy dressing, and reported both materials to be equally effective in primary teeth<sup>95</sup>. MTA has also been compared to formocresol and calcium hydroxide as a pulpotomy dressing in primary molars by Moretti *et al.* with satisfactory clinical and radiographic outcomes<sup>96</sup>.

The clinical outcome measures are similar in all the mentioned studies, but there is variation in the definition of radiographic 'success', making comparisons difficult. The radiographic outcome has been perceived to be successful in all of the mentioned studies if there have been no pathological signs of peri-apical and furcal radiolucencies, and there has been variable or no sign of reparative dentine bridge formation at the time of the reported recall. Sign of internal resorption has been recorded as a negative radiographic outcome in most of the studies. Root canal calcification of the primary molars has been observed and recorded, but not perceived as a negative radiographic outcome in the study reported by Maroto *et al.*<sup>93,94</sup>

(ii) Pulp capping: A single study was identified on the outcome of MTA pulp capping in primary teeth<sup>97</sup>. Tuna and Olmez reported that MTA was as successful as calcium hydroxide in direct pulp capping, and recommended further histological validation to support their findings.

*Root-end filling in immature permanent teeth.* Two prospective, observational studies have investigated MTA as an apical barrier in non-vital immature permanent incisors<sup>98,99</sup>. Simon *et al.* reported a range of follow-up periods from 6 to 36 months for 57 teeth, 14 of which were in patients under the age of 16, in which an apical plug of MTA was placed as a barrier. This group of workers reported a decrease in the size of the pre-existing peri-apical lesion in 81% of their cases<sup>98</sup>. A similar study by Saris *et al.* reported similar results in 17 non-vital permanent immature incisors<sup>99</sup>. The one-step placement of an MTA apical barrier was viewed as a promising alternative to traditional, multiple-visit apexification with calcium hydroxide. The advantages of a one-step MTA procedure were cited as reduced treatment time, reduced risk of calcium hydroxide-induced changes to dentine, and consequently reduced fracture risk, and the early placement of a sealing and possibly reinforcing coronal/intra-radicular restoration.

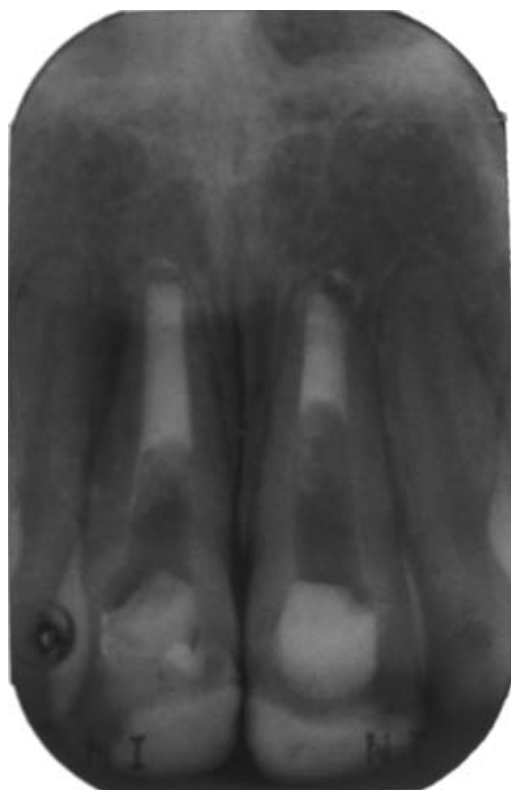
Figure 2 shows the radiographic appearance of an MTA apical barrier in the non-vital immature maxillary incisors of a 13 years old.

*Apical seal in the non-vital coronal portion of permanent teeth following root fracture.* Root canal treatment of the coronal fragment with calcium hydroxide followed by filling with gutta-percha is the traditional treatment of choice for non-vital root-fractured teeth<sup>100</sup>.

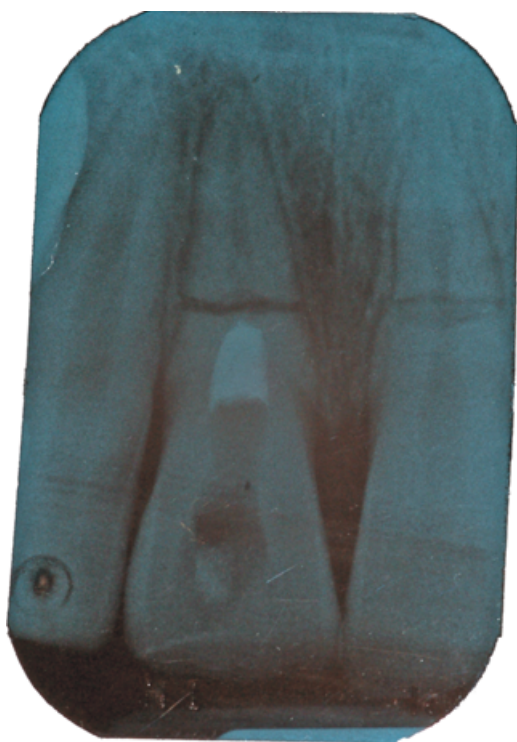
As in the case of open root apices, the use of calcium hydroxide has been promoted to induce hard-tissue barrier formation at the fracture site. The hard tissue barrier is then able to serve as a matrix for the condensation of gutta-percha and sealer. In this situation, MTA has the potential to offer all of the advantages noted for one-step root-end filling.

Currently, there have been no human or animal studies reported on the use of MTA as an apical barrier for the coronal fragment of the root-fractured tooth. There are, however, case reports explaining the technique with follow-up of up to 2 years.

Figure 3 illustrates the radiographic appearance of MTA placed as an apical seal in a coronal fragment of a fractured maxillary right central incisor in an 11 years old.



**Fig. 2.** Radiographic image of mineral trioxide aggregate placed as an apical barrier in a non-vital immature maxillary incisors in a 13 years old.



**Fig. 3.** Radiographic image of mineral trioxide aggregate placed as an apical seal in a coronal fragment of a maxillary right central incisor in an 11 years old.

## Conclusion

The paucity of human clinical studies on MTA is likely to change within the next decade. Considering that the material has been in routine clinical use for a little more than a decade, all aspects of its profile as a dental material are currently being investigated. Studies designed to conform with CONSORT guidelines in terms of power, blinding, control groups, and recall times have the potential to validate the clinical impressions of MTA as a useful and versatile material in the armamentarium of the paediatric dentist.

### What this paper adds

- This paper provides a synopsis of the key chemical and physical properties of MTA, many of which are of direct clinical relevance.
- This paper reviews the available evidence on the effectiveness of MTA in a range of clinical applications. It also highlights weaknesses in the literature and points to the need for further studies.

### Why this paper is important to paediatric dentists

- All paediatric dentists should be familiar with the fundamental properties of the materials they use on patients. This paper provides readers with helpful information on a relatively new restorative material which comes into direct contact with soft connective tissues (pulp and peri-apical tissues).
- Paediatric dentists may be hesitant to adopt a relatively new material into their clinical regimes, lacking confidence in the available evidence or having concerns about its cost and handling characteristics. This paper provides illustrated evidence on the success which can accompany the use of MTA in a range of settings including: (i) primary teeth, as a pulpotomy medicament; (ii) young permanent teeth, as a pulp-capping agent; (iii) immature traumatized permanent teeth, as an apical barrier; (iv) traumatized permanent teeth with root fractures, as an apical barrier of the coronal root fragment; and (v) permanent teeth, as a repair material for perforation and resorptive defects.

## References

- 1 Torabinejad M, Hong CU, McDonald F, Pitt Ford TR. Physical and chemical properties of a new root-end filling material. *J Endod* 1995; **21**: 349–353.
- 2 Torabinejad M, White TJ. Tooth filling material and use. US Patent Number 5,769,638.
- 3 Camilleri J, Montesin FE, Brady K, Sweeney R, Curtis RV, Pitt Ford TR. The constitution of mineral trioxide aggregate. *Dent Mater* 2005; **21**: 297–303.

- 4 Roberts HW, Toth JM, Berzins DW, Charlton DG. Mineral trioxide aggregate material use in endodontic treatment: a review of the literature. *Dent Mater* 2008; **24**: 149–164.
- 5 Sarkar NK, Caidedo R, Tirwik P, Moiseyeva R, Kawashima I. Physicochemical basis of the biologic properties of mineral trioxide aggregate. *J Endod* 2005; **21**: 731–738.
- 6 Dammaschke T, Gerth HUV, Zuchner H, Schafer E. Chemical and physical surface and bulk material characterization of white ProRoot MTA and two Portland cements. *Dent Mater* 2005; **21**: 731–738.
- 7 Camilleri J, Montesin FE, Di Silvio L, Pitt Ford TR. The chemical constitution and biocompatibility of accelerated Portland cement for endodontic use. *Int Endod J* 2005; **38**: 834–842.
- 8 Asgary S, Parirokh M, Egbbal MJ, Brink F. Chemical differences between white and gray mineral trioxide aggregate. *J Endod* 2005; **31**: 101–103.
- 9 Diamantti E, Kerezoudis NP, Gakis NB, Tsatsas V. Chemical composition and surface characteristics of grey and new white ProRoot MTA. *J Endod* 2003: Abstract R81.
- 10 Song J, Mante FK, Romanow WJ, Kim S. Chemical analysis of powder and set forms of Portland cement, gray ProRoot MTA, white ProRoot MTA, and gray MTA-Angelus. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod* 2006; **102**: 809–815.
- 11 Torabinejad M, Smith PW, Kettering JD, Pitt Ford T. Comparative investigation of marginal adaptation of mineral trioxide aggregate and other commonly used root-end filling materials. *J Endod* 1995; **21**: 295–299.
- 12 Islam I, Chang HK, Yap AUJ. X-ray diffraction analysis of mineral trioxide aggregate and Portland cement. *Int Endod J* 2006; **39**: 220–225.
- 13 Kogan P, He J, Glickman GN, Watanabe I. The effects of various additives on setting properties of MTA. *J Endod* 2006; **32**: 569–572.
- 14 Bortoluzzi EA, Broon NJ, Duarte MAH, Demarchi ACCO, Bramante CM. The use of a setting accelerator and its effect on pH and calcium ion release of mineral trioxide aggregate and white Portland cement. *J Endod* 2006; **32**: 1194–1197.
- 15 Bortoluzzi EA, Broon NJ, Bramante CM, Garcia RB, de Moraes IG, Bernardineli N. Sealing ability of MTA and radiopaque Portland cement with or without calcium chloride for root-end filling. *J Endod* 2006; **32**: 897–900.
- 16 Ding SJ, Kao CT, Shie MY, Hung CJ, Huang TH. The physical and cytological properties of white MTA mixed with  $\text{Na}_2\text{HPO}_4$  as an accelerant. *J Endod* 2008; **34**: 748–751.
- 17 Shah PMM, Chong BS, Sidhu SK, Pitt Ford T. Radiopacity of potential root end filling materials. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod* 1996; **81**: 476–479.
- 18 Danesh G, Dammaschke T, Gerth HUV, Zandbiglari T, Schafer E. A comparative study of selected properties of ProRoot mineral trioxide aggregate and two Portland cements. *Int Endod J* 2006; **39**: 213–219.
- 19 Walker MP, Diliberto A, Lee C. Effect of setting conditions on mineral trioxide aggregate flexural strength. *J Endod* 2006; **32**: 334–336.
- 20 Budig CG, Eleazer PD. *In vitro* comparison of the setting of dry ProRoot MTA by moisture absorbed through the root. *J Endod* 2008; **34**: 712–714.
- 21 Fridland M, Rosado R. Mineral trioxide aggregate (MTA) solubility and porosity with different water-to-powder ratios. *J Endod* 2003; **29**: 814–817.
- 22 Fridland M, Rosado R. MTA solubility: a long term study. *J Endod* 2005; **31**: 376–379.
- 23 Santos AD, Moraes JCS, Araujo EB, Yukimity K, Filho VV. Physico-chemical properties of MTA and a novel experimental cement. *Int Endod J* 2005; **38**: 443–447.
- 24 Harty FJ, Parkins BJ, Wengraf AM. The success rate of apicectomy. A retrospective study of 1016 cases. *Br Dent J* 1970; **129**: 407–413.
- 25 Matlof IR, Jensen JR, Singer L, Tabibi A. A comparison of methods used in root canal sealability studies. *Oral Surg Oral Med Oral Pathol* 1982; **53**: 203–208.
- 26 Kersten HW, Moorer WR. Particles and molecules in endodontic leakage. *Int Endod J* 1989; **22**: 118–124.
- 28 Haikel Y, Wittenmeyer W, Bateman G, Bentaleb A, Allenmann C. A new method for the quantitative analysis of endodontic microleakage. *J Endod* 1999; **25**: 172–177.
- 29 Torabinejad M, Higa RK, McKendry DJ, Pitt Ford T. Dye leakage of four root end filling materials: effects of blood contamination. *J Endod* 1994; **20**: 159–163.
- 27 Carpenter PL. *Microbiology*, 3rd edn. Philadelphia, PA: W.B. Saunders Company, 1972.
- 30 Abdal AK, Retief DH. The apical seal via the retro-surgical approach. I. A preliminary study. *Oral Surg* 1982; **53**: 614–621.
- 31 Yoshimura M, Marshall FJ, Tinkle JS. *In vitro* quantification of the apical sealing ability of retrograde amalgam. *J Endod* 1990; **16**: 5–12.
- 32 Inoue E, Yoshimura M, Tinkle JS, Marshall FJ. Apical dentin permeability and microleakage associated with root end resection and retrograde filling. *J Endod* 1991; **17**: 369–375.
- 33 Gilheany PA, Figdor D, Tyas MJ. Apical dentin permeability and microleakage associated with root end resection and retrograde filling. *J Endod* 1994; **20**: 22–26.
- 34 Bates C, Carnes DL, del Rio CE. Longitudinal sealing ability of mineral trioxide aggregate as a root end filling material. *J Endod* 1996; **22**: 575–578.
- 35 Shipper G, Grossman ES, Botha AJ, Cleaton-Jones PE. Marginal adaptation of mineral trioxide aggregate (MTA) compared with amalgam as a root-end filling material: a low-vacuum (LV) versus high-vacuum (HV) SEM study. *Int Endod J* 2004; **37**: 325–336.
- 36 Torabinejad M, Rastegar AF, Kettering J, Pitt Ford T. bacterial leakage of mineral trioxide aggregate as a root-end filling material. *J Endod* 1995; **21**: 109–112.
- 37 Fischer E, Arens DE, Miller C. Bacterial leakage of mineral trioxide aggregate as compared with zinc-free

- amalgam, intermediate restorative material, and Super-EBA as a root-end filling material. *J Endod* 1998; **24**: 176–179.
- 38 Soh G, Chew CL, Lee AS, Yeoh TS. Significance of hydrogen ion concentration on the dissolution of mercury from amalgam. *Quintessence Int* 1991; **22**: 225–228.
- 39 Arnold JW, Rueggeberg FA, Anderson RW, Weller RN, Borke JL, Pashley DH. The disintegration of Super EBA cement in solutions with adjusted pH and osmolarity. *J Endod* 1987; **23**: 663–668.
- 40 Gopferich A. Mechanisms of polymer degradation and erosion. *Biomaterials* 1996; **17**: 103–114.
- 41 Ferracane JL. Elution of leachable components from composites. *J Oral Rehabil* 1994; **21**: 441–452.
- 42 Roy CO, Jeanson BG, Gerrets TF. Effect of an acid environment on leakage of root-end filling materials. *J Endod* 2001; **27**: 7–8.
- 43 Valois CR, Costa ED Jr. Influence of the thickness of mineral trioxide aggregate on sealing ability of root-end fillings *in vitro*. *Oral Surg Oral Med Oral Pathol* 2004; **97**: 108–111.
- 44 Stefopoulos S, Tsatsas DV, Kerezoudis NP, Eliades G. Comparative *in vitro* study of the sealing efficiency of white vs gray ProRoot mineral trioxide aggregate formulas as apical barriers. *Dent Traumatol* 2008; **24**: 207–213.
- 45 Zou L, Liu H, Yin S, Li W, Xie J. *In vitro* evaluation of the sealing ability of MTA used for the repair of furcation perforations with and without the use of an internal matrix. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod* 2008; **105**: e61–e65.
- 46 The Editorial Board of Journal of Endodontics. Wanted: a base of evidence. *J Endod* 2007; **33**: 1401–1402.
- 47 Nekoofar MH, Adusei G, Sheykhrezae MS, Hayes SJ, Bryant ST, Dummer PMH. The effect of condensation pressure on selected physical properties of mineral trioxide aggregate. *Int Endod J* 2007; **40**: 453–461.
- 48 White DJ, Lacefield WR, Chavers LS, Eleazer PD. The effect of three commonly used endodontic materials on the strength and hardness of root dentin. *J Endod* 2002; **28**: 828–830.
- 49 Andreasen JO, Munksgaard EC, Bakland LK. Comparison of fracture resistance in root canals of immature sheep teeth after filling with calcium hydroxide or MTA. *Dent Traumatol* 2006; **22**: 154–156.
- 50 Bortoluzzi EA, Souza EM, Reis JMS, Esberard RM, Tanomaru-Filho M. Fracture strength of bovine incisors after intra-radicular treatment with MTA in an experimental immature tooth model. *Int Endod J* 2007; **40**: 684–691.
- 51 Hatibovic-Kofman S, Raimundo L, Zheng L, Chong L, Friedman M, Andreasen JO. Fracture resistance and histological findings of immature teeth treated with mineral trioxide aggregate. *Dent Traumatol* 2008; **24**: 272–276.
- 52 Cvek M. Prognosis of luxated non-vital maxillary incisors treated with calcium hydroxide and filled with gutta-percha. A retrospective clinical study. *Endod Dent Traumatol* 1992; **8**: 45–55.
- 53 Andreasen JO, Farik B, Munksgaard EC. Long-term calcium hydroxide as a root canal dressing may increase risk of root fracture. *Dent Traumatol* 2002; **18**: 134–137.
- 54 Ørstavik D. Antibacterial properties of root canal sealers, cements and pastes. *Int Endod J* 1981; **14**: 125–133.
- 55 Ørstavik D, Qvist V, Stoltze K. A multivariate analysis of the outcome of endodontic treatment. *Eur J Oral Sci* 2004; **112**: 224–230.
- 56 Sipert CR, Hussne RP, Nishiyama CK, Torres SA. *In vitro* antimicrobial activity of Fill Canal, Sealapex, Mineral Trioxide Aggregate, Portland cement and EndoRez. *Int Endod J* 2005; **38**: 539–553.
- 57 Al-Hazaimi K, Al-Shalan TA, Naghshbandi J, Oglesby S, Dimon JSH, Rotstein I. Antibacterial effect of two mineral trioxide aggregate (MTA) preparations against *Enterococcus faecalis* and *Streptococcus sanguis in vitro*. *J Endod* 2006; **32**: 1053–1056.
- 58 Eldeniz AU, Hadimli HH, Ataogly H, Ørstavik D. Antibacterial effect of selected root-end filling materials. *J Endod* 2006; **32**: 345–349.
- 59 Nandini S, Ballal S, Kandaswamy D. Influence of glass-ionomer cement on the interface and setting reaction of mineral trioxide aggregate when used as a furcal repair material using laser Raman spectroscopic analysis. *J Endod* 2006; **33**: 167–172.
- 60 Ballal S, Venkateshbabu N, Nandini S, Kandaswamy D. An *in vitro* study to assess the setting and surface crazing of conventional glass ionomer cement when layered over partially set mineral trioxide aggregate. *J Endod* 2008; **34**: 478–480.
- 61 Tunc ES, Sonmez IS, Bayrak S, Egilmez T. The evaluation of bond strength of a composite and a compomer to white mineral trioxide aggregate with two different bonding systems. *J Endod* 2008; **34**: 603–605.
- 62 Kettering JD, Torabinejad M. Investigation of mutagenicity and mineral trioxide aggregate and other commonly used root-end filling materials. *J Endod* 1995; **21**: 537–539.
- 63 Osario RM, Hefti A, Vertucci FJ, Shawley AL. Cytotoxicity of endodontic materials. *J Endod* 1998; **24**: 91–96.
- 64 Moretton TR, Brown CE Jr, Legan JL, Kafrawy AH. Tissue reactions after subcutaneous and intraosseous implantation of mineral trioxide aggregate and ethoxybenzoic acid. *Biomed Mater Res* 2000; **52**: 528–533.
- 65 Sumer M, Muglali M, Bodrumlu E, Guvenic T. Reactions of connective tissue to amalgam, intermediate restorative material, mineral trioxide aggregate mixed with chlorhexidine. *J Endod* 2006; **32**: 1094–1096.
- 66 Vosoughhosseini S, Lotfi M, Shahi S, *et al*. Influence of white versus gray mineral trioxide aggregate on inflammatory cells. *J Endod* 2008; **34**: 715–717.
- 67 Saidon J, He J, Zhu Q, Safavi K, Spangberg LSW. Cell and cell tissue reactions to mineral trioxide aggregate

- and Portland cement. *Oral Surg Oral Med Oral Pathol Oral Radiol Endod* 2003; **95**: 483–489.
- 68 Masuda YM, Wang X, Hossain M, *et al.* Evaluation of biocompatibility of mineral trioxide aggregate with an improved rabbit ear chamber. *J Oral Rehabil* 2005; **32**: 145–150.
  - 69 Torabinejad M, Hong C, Lee S, Monsef M, Pitt Ford T. Investigation of mineral trioxide aggregate for root-end filling in dogs. *J Endod* 1995; **21**: 603–608.
  - 70 Economides N, Pantelidou O, Kokkas A, Tziafas D. Short-term periradicular tissue response to mineral trioxide aggregate (MTA) as root-end filling material. *Int Endod J* 2003; **36**: 44–48.
  - 71 Baek S, Plenk H, Kin S. Periapical tissue responses and cementum regeneration with amalgam, Super EBA, and MTA as root-end filling materials. *J Endod* 2005; **31**: 444–449.
  - 72 Felipe WT, Felipe CS, Rocha MJC. The effect of mineral trioxide aggregate on the apexification and periapical healing of teeth with incomplete root formation. *Int Endod J* 2006; **39**: 2–9.
  - 73 Torabinejad M, Pitt Ford T, McKendry DJ, Abedi HR, Miller DA, Kariyawasam SP. Histologic assessment of mineral trioxide aggregate as a root-end filling in monkeys. *J Endod* 1997; **23**: 225–228.
  - 74 Panzarini SR, Holland R, de Souza V, Poi WR, Sonoda CK, Pedrini D. Mineral trioxide aggregate as a root canal filling material in reimplanted teeth. Microscopic analysis in monkeys. *Dent Traumatol* 2007; **23**: 265–272.
  - 75 Pitt Ford T, Torabinejad M, Abedi HR, Bakland LK, Kariyawasam SP. Using mineral trioxide aggregate as a pulp-capping material. *J Am Dent Assoc* 1996; **127**: 1491–1494.
  - 76 Faracoe IM, Holland R. Response of the pulp of dogs to capping with mineral trioxide aggregate or a calcium hydroxide cement. *Dent Traumatol* 2001; **17**: 163–166.
  - 77 Tziafas D, Pantelidou O, Alvanou A, Belibasakis G, Papadimitriou S. The dentinogenic effect of mineral trioxide aggregate (MTA) in short-term capping experiments. *Int Endod J* 2002; **35**: 245–254.
  - 78 Salako N, Joseph B, Ritwik P, Salonen J, John P, Junaid TA. Comparison of bioactive glass, mineral trioxide aggregate, ferric sulphate and formocresol as pulpotomy agents in rat molar. *Dent Traumatol* 2003; **19**: 314–320.
  - 79 Aeinehchi M, Eslami B, Ghanbariha M, Saffar AS. Mineral trioxide aggregate (MTA) and calcium hydroxide as pulp-capping agents in human teeth: a preliminary report. *Int Endod J* 2003; **36**: 225–231.
  - 80 Iwamoto CE, Adachi E, Pameijer CH, Barnes D, Romberg EE, Jeffries S. Clinical and histological evaluation of white ProRoot MTA in direct pulp capping. *Am J Dent* 2006; **19**: 85–90.
  - 81 Nair PNR, Duncan HF, Pitt Ford TR, Luder HU. Histological, ultrastructural and quantitative investigations on the response of healthy human pulps to experimental capping with mineral trioxide aggregate: a randomized controlled trial. *Int Endod J* 2008; **41**: 128–150.
  - 82 Accorinte Mde L, Holland R, Reis A, *et al.* Evaluation of mineral trioxide aggregate and calcium hydroxide cement as pulp-capping agents in human teeth. *J Endod* 2008; **34**: 1–6.
  - 83 Bogen G, Kim JS, Bakland LK. Direct pulp capping with mineral trioxide aggregate. An observational study. *J Am Dent Assoc* 2008; **139**: 305–315.
  - 84 Barrieshi-Nusair KM, Qudeimat MA. A prospective clinical study of mineral trioxide aggregate for partial pulpotomy in cariously exposed permanent teeth. *J Endod* 2006; **32**: 731–735.
  - 85 Witherspoon DE, Small JC, Harris GZ. Mineral trioxide aggregate pulpotomies: a case series outcome assessment. *J Am Dent Assoc* 2006; **137**: 610–618.
  - 86 El-Meligy OA, Avery DR. Comparison of mineral trioxide aggregate and calcium hydroxide as pulpotomy agents in young permanent teeth (apexogenesis). *Pediatr Dent* 2006; **28**: 399–404.
  - 87 Eidelman E, Holan G, Fuks AB. Mineral trioxide aggregate vs. formocresol in pulpotomized primary molars: a preliminary report. *Pediatr Dent* 2001; **23**: 15–18.
  - 88 Agamy HA, Bakry NS, Mounir MMF, Avery DR. Comparison of mineral trioxide aggregate and formocresol as pulp-capping agents in pulpotomized primary teeth. *Pediatr Dent* 2004; **26**: 302–309.
  - 89 Holan G, Eidelman E, Fuks AB. Long-term evaluation of pulpotomy in primary molars using mineral trioxide aggregate or formocresol. *Pediatr Dent* 2005; **27**: 129–136.
  - 90 Farsi N, Alamoudi N, Balto K, Mushayt A. Success of mineral trioxide aggregate in pulpotomized primary molars. *J Clin Pediatr Dent* 2005; **29**: 307–311.
  - 91 Aeinehchi M, Dadvand S, Fayazi S, Bayat-Movahed S. Randomized controlled trial of mineral trioxide aggregate and formocresol for pulpotomy in primary molar teeth. *Int Endod J* 2007; **40**: 261–267.
  - 92 Noorollahain H. Comparison of mineral trioxide aggregate and formocresol as pulp medicaments for pulpotomies in primary molars. *Br Dent J* 2008; **204**: E20. doi :10.1038/sj.bdj.2008.319.
  - 93 Maroto M, Barberia E, Planells P, Garcia-Godoy F. Dentin bridge formation after mineral trioxide aggregate (MTA) pulpotomies in primary teeth. *Am J Dent* 2005; **18**: 151–154.
  - 94 Maroto M, Barberia E, Vera V, Garcia-Godoy F. Mineral trioxide aggregate as pulp dressing agent in pulpotomy treatment of primary molars: 42-month clinical study. *Am J Dent* 2007; **20**: 283–286.
  - 95 Percinoto C, Castro AM, Pinto LM. Clinical and radiographic evaluation of pulpotomies employing calcium hydroxide and trioxide mineral aggregate. *Gen Dent* 2006; **54**: 258–261.
  - 96 Moretti ABS, Sakai VT, Oliveira TM, *et al.* The effectiveness of mineral trioxide aggregate, calcium hydroxide and formocresol for pulpotomies in primary teeth. *Int Endod J* 2008; **41**: 547–555.

- 97 Tuna D, Olmez A. Clinical long-term evaluation of MTA as a direct pulp capping material in primary teeth. *Int Endod J* 2008; **41**: 273–278.
- 98 Simon S, Rilliard F, Berdel A, Machtou P. The use of mineral trioxide aggregate in one-visit apexification treatment: a prospective study. *Int Endod J* 2007; **40**: 186–197.
- 99 Saris S, Tahmassebi JF, Duggal MS, Cross I. A clinical evaluation of mineral trioxide aggregate for root-end closure of non-vital immature permanent incisors in children – a pilot study. *Dent Traumatol* 2008; **24**: 79–85.
- 100 Cvek M, Mejare I, Andreasen JO. Conservative endodontic treatment of teeth fractured in the middle or apical part of the root. *Endod Traumatol* 2004; **20**: 261–269.

Copyright of International Journal of Paediatric Dentistry is the property of Blackwell Publishing Limited and its content may not be copied or emailed to multiple sites or posted to a listserv without the copyright holder's express written permission. However, users may print, download, or email articles for individual use.