Micro-computed tomography in preventive and restorative dental research: A review

Mehrsima Ghavami-Lahiji 12.*, Reza Tayefeh Davalloo 2, Gelareh Tajziehchi 2, Paria Shams 2

¹Dental Sciences Research Center, School of Dentistry, Guilan University of Medical Sciences, Rasht, Iran ²Department of Restorative Dentistry, School of Dentistry, Guilan University of Medical Sciences, Rasht, Iran

ABSTRACT

Purpose: The use of micro-computed tomography (micro-CT) scans in biomedical and dental research is growing rapidly. This study aimed to explore the scientific literature on approaches and applications of micro-CT in restorative dentistry.

Materials and Methods: An electronic search of publications from January 2009 to March 2021 was conducted using ScienceDirect, PubMed, and Google Scholar. The search included only English-language articles. Therefore, only studies that addressed recent advances and the potential uses of micro-CT in restorative and preventive dentistry were selected.

Results: Micro-CT is a tool that enables 3-dimensional imaging on a small scale with very high resolution. In this method, there is no need for sample preparation or slicing. Therefore, it is possible to examine the internal structure of tissue and the internal adaptation of materials to surfaces without destroying them. Due to these advantages, micro-CT has been recommended as a standard imaging tool in dental research for many applications such as tissue engineering, endodontics, restorative dentistry, and research on the mineral density of hard tissues and bone growth. However, the high costs of micro-CT, the time necessary for scanning and reconstruction, computer expertise requirements, and the enormous volume of information are drawbacks.

Conclusion: The potential of micro-CT as an emerging, accurate, non-destructive approach is clear, and the valuable research findings reported in the literature provide an impetus for researchers to perform future studies focusing on employing this method in dental research. (*Imaging Sci Dent 2021; 51: 341-50*)

KEY WORDS: X-Ray Microtomography; Dentistry, Operative; Review Literature

Introduction

After the invention of X-rays in 1895 by Wilhelm Roentgen, it was hypothesized that X-rays could be used to diagnose some common conditions, such as fractures and foreign objects, and visualize the body's internal structures in a non-invasive way. Decades later, after a wider recognition of the importance of this technology, X-ray images became an integral part of routine patient care.^{1,2}

Allan Cormack and Godfrey Hounsfield developed computer-assisted tomography in 1970, for which they recei-

Published online October 15, 2021

*Correspondence to : Prof. Mehrsima Ghavami-Lahiji

Department of Restorative Dentistry, School of Dentistry, Academic Complex of Guilan University of Medical Sciences, Fuman-Saravan Road, Rasht, Iran Tel) 98-33486406, E-mail) dr.m.ghavami@gmail.com ved the Nobel Prize in Physiology or Medicine. In the late 1970s, computed tomography (CT) entered into widespread use in clinical settings.³ CT is a 3-dimensional (3D) imaging method that produces 3D images composed of 2-dimensional (2D) projections. It involves obtaining X-ray projection images from many angles of view around an axis through an object. Then, it applies a tomographic reconstruction algorithm to produce a group of thin tomographic images of continual transaxial slices through the object. The transaxial images are made up of volume elements termed voxels. An object is usually scanned by rotating it around a vertical axis within a system comprising a stationary X-ray source and an X-ray imaging array.^{3,4} Three levels of microscopic CT are defined based on their spatial resolution: mini-CT, micro-CT, and nano-CT. However, "micro-CT" is used as a generic term for all of them.⁴

Received April 9, 2021; Revised July 4, 2021; Accepted July 9, 2021

Copyright © 2021 by Korean Academy of Oral and Maxillofacial Radiology

This is an Open Access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (http://creativecommons.org/licenses/by-nc/3.0) which permits unrestricted non-commercial use, distribution, and reproduction in any medium, provided the original work is properly cited. Imaging Science in Dentistry · pISSN 2233-7822 eISSN 2233-7830

Micro-CT is a high-resolution type of CT. Lee Feldkamp developed the first micro-CT system in the early 1980s to evaluate structural defects in ceramic materials. Early micro-CT scanners were not widely available.⁵ Compact commercial systems rapidly became essential components of academic and industrial research laboratories for various investigations and applications, with the potential of becoming a convincing substitute for histological sectioning.⁶

Micro-CT equipment consists of various essential components, including a 90-150 kVA X-ray tube, a filter and collimator, a computer-controlled electric motor, a CCD camera for translating X-ray image data, an image intensifier apparatus, a specimen stand, and a computer.^{1,3} The 2 main types of micro-CT scans are *in vivo* and *ex vivo* scans. An object is placed on a stationary platform with an X-ray tube and an optical X-ray detector spins around it for *in vivo* scans. Instead, for *ex vivo* scanning, an object is placed on a rotating platform with the X-ray tube and the optical X-ray detector is in a fixed position.⁷ Although rotating object scanners produce ultra-high-resolution images, they are not technically suitable for small-animal imaging (Fig. 1).⁸

An electric rotation stage is employed to specifically monitor the amount of sample rotation relative to the optical X-ray detector and X-ray tube to scan the sample for *ex vivo* micro-CT. The overall motion of the micro-CT scanner is regulated by a computer. Three-dimensional images are reconstructed using lead shielding, which protects the

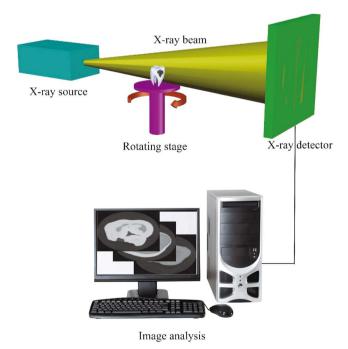


Fig. 1. Schematic representation of micro-computed tomography imaging.

operators during the scanning procedure.⁷ Clinical CT scanners typically produce images composed of 1 mm³ voxels, while micro-CT systems have a much better spatial resolution. The voxel range of micro-CT (5-50 μ m) is nearly 1 million times smaller in volume than the CT voxels. Thus, micro-CT is expected to have better diagnostic accuracy.^{5,9}

Micro-CT provides many advantages in comparison with other methods, but it also has some restrictions. The advantage of these systems is convenient in-depth exploration of the X-ray imaging modality due to its narrow, monochromatic X-ray with high flux.⁴ In contrast to microscopic methods, micro-CT is a nondestructive approach, meaning that the internal properties of the same sample can be tested several times, and samples remain intact for further biological and mechanical testing.⁵ Another benefit of micro-CT is the possibility of multiple scanning and image processing using specific software. Sample preparation for micro-CT is not an issue, unlike when other destructive methods are used. The overall advantages of micro-CT facilitate more precise measurements.¹

Some limitations of micro-CT relate to the scanning and reconstruction processing time, the high costs of this technique, and the necessity of computer expertise.^{1,10} The image file sizes are large (around 3 GB). As a result, data acquisition can be a problem when it is necessary to store and retrieve large volumes of files.¹¹ The radiation dose from micro-CT scans is also sufficient to impact experimental outcomes by provoking changes in the immune response and other biological pathways. Although these changes usually are not lethal,¹² the high radiation dose prevents the use of micro-CT in clinical settings.^{1,6}

In recent years, the utility of micro-CT has increased in experimental and preclinical bone and dental studies due to the broad range of technological advances in X-ray sources and X-ray imaging arrays.⁴ Micro-CT systems provide an opportunity to analyze microstructures, defects, and differences in density and morphology. Several types of samples encompassing mineralized hard tissues, soft tissues, and solid or liquid materials - can be examined using micro-CT.^{5,10}

For more than 2 decades, cone-beam computed tomography (CBCT) has been routinely used in clinical settings in dentistry. CBCT was developed as a method of gathering images more quickly, at a lower radiation dose and a lower cost.¹³ Compared to CBCT, micro-CT has smaller voxel sizes and a longer exposure, enabling the detection of more subtle and smaller changes.¹⁴ Quantitative gray value measurements in CBCT, in contrast to micro-CT, are unreliable and therefore should generally be avoided. Because of the geometry of the beam and flat-panel detectors, artifacts, and variation in scanning circumstances, CBCT-based density measurements are inevitably erroneous.¹⁵ Therefore, micro-CT images are anticipated to have higher accuracy for diagnostic measurements to detect carious lesions, periodontal disease, root resorption, and intra-operative imaging than CBCT images.¹⁰

The areas of implementation of micro-CT include biomedical and bioengineering applications and tissue engineering. Micro-CT has also facilitated dental research in various fields, such as endodontic studies of changes in bone structure and inflammatory root resorption due to bacterial infection of the dental apex, the evaluation of implants and surrounding bone, analyses of the growth and development of craniofacial bones, and studies investigating tooth-restoration interfaces.^{1,10,16}

The purpose of this review article was not to cover all areas of micro-CT in dentistry, as this would not be possible within a single review, but to focus on the potential applications of micro-CT in preventive, diagnostic, and restorative dental research. This review addressed micro-CT approaches for the quantitative evaluation of tooth or dental material structures in recent research.

Remineralizing potential of tooth lesions under different treatment modalities

Several indirect methods have been introduced to estimate the mineral content of dental structures, such as optical coherence tomography (as a modality to measure optical properties),¹⁷ microhardness,¹⁸ and nanoindentation.¹⁹ Direct approaches to measuring mineral content include transverse microradiography and micro-CT. CT is a nondestructive method of 3D tissue reconstruction for biological systems, whereas the traditional gold-standard method of transverse microradiography is destructive and time-consuming. Micro-CT has recently been used for quantitative research on dental hard tissue structures, and good agreement has been found with transverse microradiography in evaluations of mineral loss and the depth of lesions.²⁰

The mineral density profiles obtained by micro-CT analyses furnish spatial insights into mineral distribution and can be utilized to monitor alterations in the mineral content of dental hard tissues, including enamel and dentin lesions.¹⁹

Bagheri et al.¹⁹ used micro-CT to analyze the influence of surface impregnation of bovine enamel by leucine-rich amelogenin peptide (LRAP) on the remineralization capability of eroded lesions. They found that LRAP exerted a regulatory effect on the pattern of remineralization *in* *vitro*, improving lesion depth and mineral loss of the enamel lesions in a dose-dependent manner. In a similar study, Bächli et al.²¹ examined the ability of dentin matrix proteins and enamel matrix derivatives to remineralize experimentally produced dentin lesions.

Neve et al.²² promoted artificial dentin lesions in a multispecies biofilm model. After establishing caries lesions, they subjected washed samples to several bioactive types of cements, such as BiodentineTM, conventional glass ionomer cement, mineral trioxide aggregate, and Portland cement. Mineral density changes were then calculated using micro-CT and reported in comparison with zinc-oxide eugenol (ZOE) and control groups.

Zan et al.²³ investigated remineralization in 3 types of restorations by means of micro-CT at the root dentin around the restorative materials after a period of cariogenic challenge.²³

In the study of Liu et al.,²⁴ tooth blocks were randomly allocated to 4 groups (silver or potassium fluoride solutions, silver nitrate solution, and deionized water), and then were immersed in a buffered demineralization solution at pH 4.4. Mean lesion depth was measured by micro-CT. Another study recommended the adjunctive application of silver diamine fluoride with fluoride toothpaste for the remineralization of incipient caries lesions on smooth tooth surfaces based on the results of a micro-CT study on lesion depth and mineral density.²⁵

Occlusal and proximal caries detection

Dentists usually diagnose caries primarily by a visual examination and bitewing radiography. Other clinical caries detection techniques have been proposed to provide objective measurements, such as the DIAGNOdent Pen (LF pen, Kavo, Biberach, Germany) and CarieScan PRO (CarieScan Ltd, Scotland, UK). The first relies on laser fluorescence, and the latter is based on electrical impedance spectroscopy.²⁶

Histological slicing has been traditionally considered the gold-standard method for determining the depth of caries in *in vitro* studies to validate clinical methods. However, some parts of the specimen may be lost due to the thickness of the blade during the histological slicing procedure. In addition, the limited capability of a microscope to visualize damaged tissue should not be ignored.²⁷

In recent years, a substantial emphasis has been placed on new methods for caries detection to investigate the depth of caries. The micro-CT technology does not destroy specimens and can evaluate nearly 1000 dental sections simultaneously. Thus, micro-CT was considered the gold standard in recent studies.²⁷⁻³⁰ However, this method is not feasible in routine clinical practice because a long time is needed to scan, reconstruct and measure images, and micro-CT also requires a high dose of radiation.²⁷

Lederer et al.³¹ investigated the validity of bitewing radiography and near-infrared reflection to detect caries and compared them with micro-CT as a reference standard. In their study, near-infrared reflection was not found to be suitable for the reliable detection of proximal caries lesions. Micro-CT is a reliable and alternative nondestructive method compared with histology for the validation of caries detection methods.⁶

Evaluation of tooth structure mineral density

The density of minerals plays an essential role in providing the mechanical properties of enamel and dentin. Several studies have described the local mechanical properties of some areas of the tooth; however, it is necessary to examine the tooth's entire structure, from the crown to the apex, as a macrostructure from a mechanical point of view.³²

Hayashi-Sakai et al.³² examined mineral density distribution patterns in permanent and deciduous tooth structures to obtain a standard mineral density value for each type of tooth using micro-CT. They found that the mineral density of permanent enamel for each tooth gradually decreased towards the cementoenamel junction, and the mineral density of dentin in permanent teeth was lower than the density distribution of enamel in all areas.

Djomehri et al.³³ investigated various dental tissues, including normal, hypomineralized, and hypermineralized carious dentin, normal cementum, and cementum affected by periodontitis. They measured mineral density variations obtained from micro-CT in conjunction with elemental mapping obtained from X-ray fluorescence microscopy in the related tissues to obtain insights into normal and pathologic biomineralization processes.

Evaluation of thickness

Enamel thickness has long been important in anthropological studies and studies of human evolution due to its phylogenetic value.^{5,34} Enamel thickness was also thought to be important in interpreting occlusal loading regimens, as well as in the presentation of chronic diseases such as metabolic disorders or developmental defects (e.g., enamel hypoplasia) in the evolutionary history of humans.³⁴

Several methods exist for measuring the thickness of tooth

enamel, including preparation of physical sections of the tooth, which has been widely used in a number of studies. However, the destructive nature of this approach prompted an ongoing debate regarding whether or not to cut teeth into thin sections, especially in scarce or extinct fossil samples. Other problems with this method, such as sample orientation, may cause some of the data to be less than ideal. Recent research has also used CT to quantify enamel thickness.¹ The use of micro-CT systems has become an effective and non-destructive method for measuring the thickness of tooth enamel.

Micro-CT provides optimal sensitivity in measuring the thickness of tooth enamel, while also not causing irreparable damage to specimens. Using this method, in addition to quantifying the thickness of the enamel, the volume of tooth enamel, dentin, and pulp can also be determined.^{1,35} Some studies have employed this method to measure the crown thickness and cement space at different points.³⁶

Biofilm formation

Recently, a method to identify low X-ray absorption materials was developed and used successfully to identify "artificial" dental biofilms created around the tooth surface.³⁷ Dual-energy micro-CT can detect delicate changes in the attenuation coefficient of an object utilizing 2 image stacks taken after data acquisition at 2 different input energies.

Pires et al.³⁷ employed saliva specimens from a patient with a carious lesion as a natural source of bacteria to induce artificial caries in sound dentin. They then quantified bio-film formation using a novel dual-energy micro-CT method and characterized the demineralization potential of the formed biofilm in the sound dentin *in vitro*.

Evaluation of polymerization shrinkage and stress

Current dental resin composites have become popular due to increasing esthetic demands and are preferred over amalgam for direct restorations. These materials undergo volume loss when they are polymerized. Polymerization shrinkage generates stresses that have a major impact on the marginal integrity of resin-based restorations, causing debonding from the surrounding tooth structure, and hence, gap formation. Several techniques have been introduced to evaluate polymerization shrinkage by measuring linear and volume changes. High-resolution microtomography is a non-contact method that can be employed to obtain actual 3D information from the cavity during polymerization.³⁸

Some researchers added radiopaque zirconia fillers to experimental resin composites as markers to monitor polymerization shrinkage in bonded and unbonded class I cavities. Filler movement was investigated using micro-CT imaging.³⁹⁻⁴¹ The idea of tracing radiopaque fillers in a resin composite using micro-CT was suggested by Cho et al.,⁴¹ who showed that boundary conditions and cavity depth affect the amount and direction of polymerization shrinkage. In their study, unbonded cavities presented free and higher deformation than bonded cavities, in contrast to the findings reported by Chiang et al.⁴⁰ An examination of the displacement vectors revealed that the filler's movement was towards the light curing source, except for the upper region of the bonded cavities, where the displacement vectors were oriented inward to the bottom of the cavities. The shrinkage effects of polymerization were more noticeable in the deepest section of the cavity.41

Polymerization shrinkage was measured differently by micro-CT in other studies.^{42,43} Kamalak et al.⁴³ measured polymerization shrinkage by means of micro-CT as the ratio of the total volume of the composite resin before curing to the total volume of the composite resin after curing. Another study used a superimposition tool to evaluate volume changes of composite resin before and after curing with different irradiance modes of a light-curing unit. The results of that study did not confirm the effect of varying curing modes of the same light-curing unit on volumetric shrinkage.⁴²

Evaluation of marginal and internal adaptation of restorative materials

Defects such as voids, gaps, and bubbles can reduce the mechanical properties of resin-based materials. Although the structure of the restorative material may initially have appropriate mechanical properties, defects can reduce the performance of the restorations under fatigue load and decrease the durability of the restorative material, leading to fractures and clinical failure.

Void formation can occur as a result of the viscosity of the material, the diameter of the applicator tip, and trapping of air inside the bulk composite during the manufacturing process or between the layers of a resin composite in the incremental method.⁴⁴ Oglakci et al.⁴⁵ evaluated gap formation volume using micro-CT in premolars restored with different bulk-fill composites, with and without a resin-modified glass-ionomer cement as liner. In the study by Demirel et al.⁴⁴ on void formation in class II restorations, a paste-like bulk-fill resin composite, Filtek One BulkFill, produced

fewer internal voids than conventional flowable and pastelike resin composites. The most void formation was found in the group that received a layer of 1-mm conventional flowable restorative followed by conventional paste, like a restorative. The group that received 4 mm of Sonicfill SDR flowable bulk-fill resin composite followed by a pastelike resin composite also showed high void formation. The porous structure of the material, lower filler content, and chemical composition of the resin matrix may have a substantial impact on the material's microporosity and contribute to higher void formation. Some researchers have also investigated internal and marginal adaptation after curing of different adhesive restorative materials.^{16,46-48}

Since the success of ceramic restorations is influenced by factors such as the margin seal and the thickness of the cement adhesive, Bayrak et al.⁴⁹ measured the internal fit and marginal adaptation of inlays designed by various computer-aided design software programs. They also emphasized that non-standardized sections in the micro-CT system due to the non-standardization of specimens could significantly impact the statistical results.

Evaluation of microleakage

The marginal adaptation of dental restorations discussed in the last section is of considerable importance in the longevity of the treatment. It has been demonstrated that structural defects at the bonding interface, as well as the action of polymerization shrinkage, can play a role in disturbing marginal adaptation. This leads to marginal staining and increases the risk of postoperative sensitivity and recurrent caries.⁵⁰⁻⁵²

Microleakage is defined as the clinically undetectable percolation of substances, such as fluids (most commonly saliva), bacteria, bacterial byproducts, and different molecules and ions, between a restorative material and cavity wall of the tooth.⁵² A variety of *in vitro* methods have been suggested to image and measure microleakage. These methods include compressed air, electrochemical approaches, neutron activation analysis, reversible radioactive adsorption, radioisotopes, bacterial infiltration, scanning electron microscopy, and the dye-penetration method. Of these methods, the dye-penetration method is most widely used, but micro-CT has been introduced as the most recent method of microleakage assessment.⁵²⁻⁵⁴

The dye-penetration method entails immersion of a restored tooth in a dye solution. Then, it is necessary to cut through the restoration and assess the leakage that is visually detectable on the section under a microscope. The extent of leakage may be estimated quantitatively using a length scale. Nevertheless, leakage is often evaluated more subjectively using a system with a range of values for leakage as graded by an operator. Moreover, the major drawback of this method is that the results provide 2D images and do not take the whole tooth-restoration interface into account. In addition, dye leakage in different sections taken from other areas of the restoration varies considerably.⁵²

Like the compressed air method, some other methods can produce results that reveal the entire tooth-restoration interface, although it is not possible to discern the path and location of the leakage, which is needed to determine the mechanism and cause of microleakage.⁵²

To overcome all these shortcomings, micro-CT has recently been suggested to evaluate microleakage and restorations' internal and external adaptation. Micro-CT allows an accurate analysis without sectioning the samples due to the capacity of X-rays to penetrate the sample. In addition, regardless of the shape or dimensions of the sample, micro-CT makes it possible to examine the internal aspects predictably and reliably. Using micro-CT, 3D microleakage analysis is possible, as well as repeated evaluations over time.^{46,55} Therefore, micro-CT is now used in various studies to evaluate microleakage.

Zhao et al.⁵² evaluated the efficacy of micro-CT in detecting marginal leakage in class V restorations and reported that the accuracy of microleakage detection by micro-CT was comparable to that of the conventional microscope method in the dentin region. However, due to the lower radiopacity contrast of the silver nitrate in the enamel, it had less accuracy in the enamel area.

Tuwirqi et al.⁵⁶ used micro-CT to assess the penetration depth and microleakage of ICON resin infiltrant and conventional sealant in pits and fissures. They found that ICON revealed less microleakage and more penetration than conventional sealants. Two different techniques for bulk-fill resin composite in class II restorations were compared by Tosco et al.⁵⁴ using micro-CT, and the authors concluded that the observed microleakage was closely related to the positioning technique of the resin composite.

Bonding to cementum/dentin is considered a challenge in restorative dentistry due to the nature of the substrate. The study performed by Zavattini et al.⁴⁶ on class II cavities with micro-hybrid, flowable, and preheated composite resins showed that the majority of the microleakage detected by CT scans and an AgNO₃ tracer occurred at the cementum/ dentin margin rather than at the enamel. They reported that the use of a flowable material should be avoided in the dentin/cementum margin and suggested lining the cavity with

preheated composite-based material to reduce the incidence of microleakage.

Vohra et al.⁵⁷ compared the marginal integrity of ceramic crowns luted with bioactive and resin cements using micro-CT. The study's findings indicated that bioactive cements could provide an adhesive interface with less volumetric microleakage, better adhesion, and comparable bond strength compared with resin luting agents.

Endodontic studies

Micro-CT is also becoming increasingly important in the evaluation of root canal systems. Thirteen years after the development of micro-CT technology, the use of micro-CT was proposed for endodontic research in the field of dentistry.⁵⁸ Since then, researchers have explored the use of micro-CT in root canal systems. However, no comprehensive, detailed review of micro-CT applications in the area of restorative dentistry has yet been published, although this article attempted to supplement that gap in the present section. The following is a brief overview of the important applications of micro-CT in endodontic studies. Micro-CT allows the development of color-coded 3D models of root canals utilizing co-registered datasets from before and after preparation and high-accuracy image registration software. Endodontic research can now be conducted precisely and noninvasively thanks to the availability of micro-CT systems. A micro-CT scan can provide extensive information; slices can be reconstructed in any plane, and data can be depicted as 2D or rendered 3D images. Internal and external anatomy may be seen at the same time or separately. Images can be graded on both a qualitative and quantitative basis. Analyses of root canal morphology,^{59,60} cracks,^{61,62} and the shaping,⁶³ cleaning,⁶⁴ and filling abilities⁶⁵⁻⁶⁷ of different systems have been conducted using this technique. Table 1 presents a summary of the information discussed in this article.

Discussion

This review article presented examples of micro-CT applications in dental research, focusing on restorative, diagnostic, and preventive dentistry. Micro-CT is now being used extensively in experimental restorative dentistry, as it can create detailed instructive images of teeth and dental materials.

Although micro-CT is not suitable for clinical use, it has become a powerful tool for experimental and preclinical research in the last decade because it provides a non-invasive, reproducible, and accurate technique for the qualitative and

Table 1. Application of micro-computed tomography in r	recent studies
--	----------------

Field of measurement	Procedure/substrate			
Remineralizing potential following treatment modalities	 Effect of leucine-rich amelogenin peptide on eroded lesions in bovine enamel¹⁹ Effect of dentin matrix proteins and enamel matrix derivatives on dentin lesions²¹ Effect of several bioactive cements on carious dentin²² Impact of adhesive/composite or resin-modified glass ionomer cement restorative materials around bovine root dentin following cariogenic challenge²³ Impact of silver or potassium fluoride solutions and silver nitrate solution on enamel following demineralization solution²⁴ Effect of silver diamine fluoride and fluoride toothpaste on incipient caries lesions²⁵ 			
Occlusal and proximal caries detection	 Validity determination of bitewing radiography and visual examinations with micro-CT for occlusal caries²⁷ Validity determination of digital radiography and visual examinations with micro-CT for occlusal caries²⁹ Validity determination of bitewing radiography and near-infrared reflection with micro-CT for proximal caries³¹ 			
Tooth structure mineral density	 Mineral density changes of permanent and deciduous teeth³² Mineral density changes in normal and diseased tissue affected by periodontitis or caries³³ 			
Evaluation of thickness	 Determination of enamel thickness³⁴ Validity determination of optical coherence tomography with micro-CT for remaining dentin thickness³⁵ Measurements of crown thickness and cement space under monolithic zirconia crowns³⁶ 			
Biofilm formation	• Quantification of biofilm formation on dentin and its demineralization potential after a cariogenic approach ³⁷			
Polymerization shrinkage and stress	 Addition of traceable glass beads to monitor polymerization shrinkage of composite in cylindric cavity with or without an adhesive^{39,40} Addition of radiopaque fillers to monitor polymerization shrinkage of composite in class I cavity with or without an adhesive⁴¹ Ratio of the total volume of the composite before curing compared to after curing⁴³ Use of a superimposition tool to evaluate volume changes of composite before and after curing⁴² 			
Marginal and internal adaptation	 Adaptation of a nanocomposite resin with and without using Biodentine or resin-modified glass ionomer for pulp protection¹⁶ Gap formation under different bulk-fill composites compared to conventional composite, with or without capping⁴⁴ Gap formation under different bulk-fill/conventional composites, with and without resin-modified glass cement liner^{45,48} Adaptation of composite resins bonded using different adhesive strategies after thermo-mechanical load cycling⁴⁷ Adaptation of inlays designed by various computer-aided design software programs⁴⁹ Adaptation of lithium disilicate and zirconia crowns made by different manufacturing systems⁵¹ 			
Evaluation of microleakage	 Comparison of microleakage among micro-hybrid, flowable, and preheated composite resins in class II cavities⁴⁶ Marginal leakage detection of class V restorations⁵² Comparison of the effect of direct and indirect bonding techniques on microleakage under orthodontic brackets⁵³ Comparison of microleakage in different positioning techniques for bulk-fill in class II restorations⁵⁴ Validation of a 3-dimensional protocol for assessment of microleakage class II cavities using regular and flowable bulk-fill composites⁵⁵ Evaluation of penetration depth and microleakage of ICON resin infiltrant versus conventional sealant in pits and fissures⁵⁶ Marginal integrity of ceramic crowns luted with bioactive and resin cements⁵⁷ 			
Analysis of root canal morphology	 Analysis of different teeth with complex anatomy⁶⁰ Comparison of the accuracy of micro-CT and cone-beam computed tomography in detecting accessory canals in primary molars⁵⁹ 			
Evaluation of cracks	 Development of a method to produce a comprehensive finite element tooth model using data retrieved from micro-CT⁶¹ Measurements of crack depth using cone-beam computed tomography and panoramic radiography and validation of the measurements by micro-CT⁶² 			

Table	1.	Continued

Field of measurement	Procedure/substrate		
Evaluation of root canal preparation	• Shaping abilities of 3 nickel-titanium rotary files in 3-dimensionally-printed mandibular molars ⁶³		
Evaluation of cleaning of root canal	• Introduction of a validated approach for differentiating between debris and dentin tissue following instrumentation ⁶⁴		
Evaluation of filling of root canal	 Filling ability and sealer apical extrusion of a sealer injection system compared to a conventional obturation system⁶⁵ Impact of various test model sizes on the flow and volumetric filling of endodontic materials⁶⁶ Evaluation of the void of root canal filling over time in the single gutta-percha cone technique using calcium silicate sealer⁶⁷ 		

Micro-CT: micro-computed tomography

quantitative assessment of teeth and dental materials. Dedicated software to extract data on the mineral density of tooth structure and tooth-restoration interfaces can collect a substantial quantity of information from scans that would be impossible to obtain using traditional methods. Data can be shown in 2D or 3D renderings, and slices can be produced in any plane. Micro-CT has also been considered as the gold standard in recent studies to determine the validity of other caries detection methods. Moreover, dual-energy micro-CT has been used to identify low X-ray absorption materials such as dental biofilms.

Furthermore, several studies have used micro-CT to measure polymerization shrinkage in different ways. A few examples include tracing radiopaque fillers before and after polymerization, volumetric measurement of composite bulk before and after polymerization, and the use of superimposition tools.

The prospect of using micro-CT as an emerging, accurate, and non-destructive method is obvious, and the promising results obtained in extant studies encourage researchers to conduct more studies on this method in dental research. It is expected that additional milestones will be reached in dental research using this method. However, the high cost of micro-CT, the time required for scanning and reconstruction, the need for computer expertise, and the large volume of files are disadvantages that limit its use in some cases. Moreover, the role of standardization of samples and system parameters cannot be ignored in achieving reliable results.

Acknowledgements

The authors would like to acknowledge the Preclinical Lab, Core Facility, Tehran University of Medical Sciences, Tehran, Iran, for providing *in vitro* and *in vivo* imaging and image processing services for research projects and for holding various workshops and symposiums to increase researchers' awareness.

Conflicts of interest: None

References

- Erpaçal B, Adıgüzel Ö, Cangül S. The use of micro-computed tomography in dental applications. Int J Dent Res 2019; 9: 78-91.
- Howell JD. Early clinical use of the x-ray. Trans Am Clin Climatol Assoc 2016; 127: 341-9.
- Boerckel JD, Mason DE, McDermott AM, Alsberg E. Microcomputed tomography: approaches and applications in bioengineering. Stem Cell Res Ther 2014; 5: 144.
- 4. Ritman EL. Current status of developments and applications of micro-CT. Annu Rev Biomed Eng 2011; 13: 531-52.
- Swain MV, Xue J. State of the art of Micro-CT applications in dental research. Int J Oral Sci 2009; 1: 177-88.
- Boca C, Truyen B, Henin L, Schulte AG, Stachniss V, De Clerck N, et al. Comparison of micro-CT imaging and histology for approximal caries detection. Sci Rep 2017; 7: 6680.
- Liao CW, Fuh LJ, Shen YW, Huang HL, Kuo CW, Tsai MT, et al. Self-assembled micro-computed tomography for dental education. PLoS One 2018; 13: e0209698.
- Fouladi M, Gholami K, Ghadiri H. LOTUS-invivo micro computed tomography system for imaging of small animals and exvivo biological samples. Frontiers Biomed Technol 2020; 7: 134-7.
- Cengiz IF, Oliveira JM, Reis RL. Micro-CT a digital 3D microstructural voyage into scaffolds: a systematic review of the reported methods and results. Biomater Res 2018; 22: 26.
- Campioni I, Pecci R, Bedini R. Ten years of micro-CT in dentistry and maxillofacial surgery: a literature overview. Appl Sci 2020; 10: 4328.
- Paterson GL, Sykes D, Faulwetter S, Merk R, Ahmed F, Hawkins LE, et al. The pros and cons of using micro-computed tomography in gross and micro-anatomical assessments of polychaetous annelids. Mem Mus Vic 2014; 71: 237-46.
- 12. Willekens I, Buls N, Lahoutte T, Baeyens L, Vanhove C, Cave-

liers V, et al. Evaluation of the radiation dose in micro-CT with optimization of the scan protocol. Contrast Media Mol Imaging 2010; 5: 201-7.

- Anderson PJ, Yong R, Surman TL, Rajion ZA, Ranjitkar S. Application of three-dimensional computed tomography in craniofacial clinical practice and research. Aust Dent J 2014; 59 Suppl 1: 174-85.
- 14. Gulibire A, Cao Y, Gao A, Wang C, Wang T, Xie X, et al. Assessment of true vertical root fracture line in endodontically treated teeth using a new subtraction software - a micro-CT and CBCT study. Aust Endod J 2021; 47: 290-7.
- 15. Kivovics M, Szabó BT, Németh O, Iványi D, Trimmel B, Szmirnova I, et al. Comparison between micro-computed tomography and cone-beam computed tomography in the assessment of bone quality and a long-term volumetric study of the augmented sinus grafted with an albumin impregnated allograft. J Clin Med 2020; 9: 303.
- 16. El Ashiry EA, Alamoudi NM, Farsi NM, Al Tuwirqi AA, Attar MH, Alag HK, et al. The use of micro-computed tomography for evaluation of internal adaptation of dental restorative materials in primary molars: an in-vitro study. Int J Pharm Res Allied Sci 2019; 8: 129-37.
- 17. Hariri I, Sadr A, Nakashima S, Shimada Y, Tagami J, Sumi Y. Estimation of the enamel and dentin mineral content from the refractive index. Caries Res 2013; 47: 18-26.
- 18. Behroozibakhsh M, Shahabi S, Ghavami-Lahiji M, Sadeghian S, Nazari NS. Evaluation of crystalline changes and resistance to demineralization of the surface of human dental enamel treated with Er: YAG laser and fluoride using x-ray diffraction analysis and Vickers microhardness. Laser Phys 2018; 28: 065602.
- 19. Bagheri GH, Sadr A, Espigares J, Hariri I, Nakashima S, Hamba H, et al. Study on the influence of leucine-rich amelogenin peptide (LRAP) on the remineralization of enamel defects via micro-focus x-ray computed tomography and nanoindentation. Biomed Mater 2015; 10: 035007.
- Hamba H, Nikaido T, Sadr A, Nakashima S, Tagami J. Enamel lesion parameter correlations between polychromatic micro-CT and TMR. J Dent Res 2012; 91: 586-91.
- Bächli K, Schmidlin PR, Wegehaupt F, Paqué F, Ramenzoni L, Botter S. Remineralization of artificial dentin caries using dentin and enamel matrix proteins. Materials (Basel) 2019; 12: 2116.
- 22. Neves AB, Bergstrom TG, Fonseca-Gonçalves A, Dos Santos TM, Lopes RT, de Almeida Neves A. Mineral density changes in bovine carious dentin after treatment with bioactive dental cements: a comparative micro-CT study. Clin Oral Investig 2019; 23: 1865-70.
- 23. Zan KW, Nakamura K, Hamba H, Sadr A, Nikaido T, Tagami J. Micro-computed tomography assessment of root dentin around fluoride-releasing restorations after demineralization/remineralization. Eur J Oral Sci 2018; 126: 390-9.
- 24. Liu B, Lo E, Li C. Effect of silver and fluoride ions on enamel demineralization: a quantitative study using micro-computed tomography. Aust Dent J 2012; 57: 65-70.
- 25. Punyanirun K, Yospiboonwong T, Kunapinun T, Thanyasrisung P, Trairatvorakul C. Silver diamine fluoride remineralized artificial incipient caries in permanent teeth after bacterial pH-cycling in-vitro. J Dent 2018; 69: 55-9.

- 26. Mortensen D, Dannemand K, Twetman S, Keller MK. Detection of non-cavitated occlusal caries with impedance spectroscopy and laser fluorescence: an *in vitro* study. Open Dent J 2014; 8: 28-32.
- Özkan G, Kanli A, Başeren NM, Arslan U, Tatar I. Validation of micro-computed tomography for occlusal caries detection: an *in vitro* study. Braz Oral Res 2015; 29: S1806-832420150 00100309.
- Park YS, Ahn JS, Kwon HB, Lee SP. Current status of dental caries diagnosis using cone beam computed tomography. Imaging Sci Dent 2011; 41: 43-51.
- 29. Simões TC, Marques LC, de Sá AT, Maciel SM, Poleti ML, Prado FS, et al. Performance of methods for detecting occlusal caries lesions: ICDAS X radiological image. Res Soc Dev 2020; 9: e1859108490.
- Abogazalah N, Ando M. Alternative methods to visual and radiographic examinations for approximal caries detection. J Oral Sci 2017; 59: 315-22.
- Lederer A, Kunzelmann KH, Heck K, Hickel R, Litzenburger F. In-vitro validation of near-infrared reflection for proximal caries detection. Eur J Oral Sci 2019; 127: 515-22.
- 32. Hayashi-Sakai S, Sakamoto M, Hayashi T, Kondo T, Sugita K, Sakai J, et al. Evaluation of permanent and primary enamel and dentin mineral density using micro-computed tomography. Oral Radiol 2019; 35: 29-34.
- 33. Djomehri SI, Candell S, Case T, Browning A, Marshall GW, Yun W, et al. Mineral density volume gradients in normal and diseased human tissues. PLoS One 2015; 10: e0121611.
- Olejniczaka AJ, Grineb FE. High-resolution measurement of Neandertal tooth enamel thickness by micro-focal computed tomography. S Afr J Sci 2005; 101: 219-20.
- 35. Majkut P, Sadr A, Shimada Y, Sumi Y, Tagami J. Validation of optical coherence tomography against micro–computed tomography for evaluation of remaining coronal dentin thickness. J Endod 2015; 41: 1349-52.
- 36. Nakamura K, Mouhat M, Nergård JM, Lægreid SJ, Kanno T, Milleding P, et al. Effect of cements on fracture resistance of monolithic zirconia crowns. Acta Biomater Odontol Scand 2016; 2: 12-9.
- 37. Pires PM, Santos TP, Fonseca-Gonçalves A, Pithon MM, Lopes RT, Neves AA. A dual energy micro-CT methodology for visualization and quantification of biofilm formation and dentin demineralization. Arch Oral Biol 2018; 85: 10-5.
- Ghavami-Lahiji M, Hooshmand T. Analytical methods for the measurement of polymerization kinetics and stresses of dental resin-based composites: a review. Dent Res J (Isfahan) 2017; 14: 225-40.
- Chiang YC, Hickel R, Lin CP, Kunzelmann KH. Shrinkage vector determination of dental composite by μCT images. Compos Sci Technol 2010; 70: 989-94.
- Chiang YC, Rösch P, Dabanoglu A, Lin CP, Hickel R, Kunzelmann KH. Polymerization composite shrinkage evaluation with 3D deformation analysis from μCT images. Dent Mater 2010; 26: 223-31.
- 41. Cho E, Sadr A, Inai N, Tagami J. Evaluation of resin composite polymerization by three dimensional micro-CT imaging and nanoindentation. Dent Mater 2011; 27: 1070-8.
- 42. Atria PJ, Sampaio CS, Cáceres E, Fernández J, Reis AF, Gian-

nini M, et al. Micro-computed tomography evaluation of volumetric polymerization shrinkage and degree of conversion of composites cured by various light power outputs. Dent Mater J 2018; 37: 33-9.

- 43. Kamalak H, Kamalak A. Evaluation of polymerization shrinkage of dental composites by microcomputed tomography. Biomed Res 2018; 29: 844-52.
- 44. Demirel G, Baltacıoğlu İH, Kolsuz ME, Ocak MOrhan K. Micro-computed tomography evaluation of internal void formation of bulk-fill resin composites in Class II restorations. Polym Compos 2019; 40: 2984-92.
- 45. Oglakci B, Kazak M, Donmez N, Dalkilic EE, Koymen SS. The use of a liner under different bulk-fill resin composites: 3D GAP formation analysis by x-ray microcomputed tomography. J Appl Oral Sci 2020; 28: e20190042.
- 46. Zavattini A, Mancini M, Higginson J, Foschi F, Pasquantonio G, Mangani F. Micro-computed tomography evaluation of micro leakage of Class II composite restorations: an *in vitro* study. Eur J Dent 2018; 12: 369-74.
- 47. Han SH, Park SH. Micro-CT evaluation of internal adaptation in resin fillings with different dentin adhesives. Restor Dent Endod 2014; 39: 24-31.
- Kwon OH, Park SH. Evaluation of internal adaptation of dental adhesive restorations using micro-CT. Restor Dent Endod 2012; 37: 41-9.
- 49. Bayrak A, Akat B, Ocak M, Kuluçarslan MA, Özcan M. Microcomputed tomography analysis of fit of ceramic inlays produced with different CAD software programs. Eur J Prosthodont Restor Dent (in press).
- 50. Pushpa R, Suresh BS. Marginal permeability of one step selfetch adhesives: effects of double application or the application of hydrophobic layer. J Conserv Dent 2010; 13: 141-4.
- 51. Riccitiello F, Amato M, Leone R, Spagnuolo G, Sorrentino R. In vitro evaluation of the marginal fit and internal adaptation of zirconia and lithium disilicate single crowns: micro-CT comparison between different manufacturing procedures. Open Dent J 2018; 12: 160-72.
- 52. Zhao XY, Li SB, Gu LJ, Li Y. Detection of marginal leakage of Class V restorations *in vitro* by micro-computed tomography. Oper Dent 2014; 39: 174-80.
- 53. Öztürk F, Ersöz M, Öztürk SA, Hatunoğlu E, Malkoç S. Micro-CT evaluation of microleakage under orthodontic ceramic brackets bonded with different bonding techniques and adhesives. Eur J Orthod 2016; 38: 163-9.
- 54. Tosco V, Vitiello F, Furlani M, Gatto ML, Monterubbianesi R, Giuliani A, et al. Microleakage analysis of different bulk-filling techniques for class II restorations: μ-CT, SEM and EDS evaluations. Materials (Basel) 2020; 14: 31.
- 55. Rizzante FA, Sedky RA, Furuse AY, Teich S, Ishikiriama SK, Mendonça G. Validation of a method of quantifying 3D leakage

in dental restorations. J Prosthet Dent 2020; 123: 839-44.

- 56. Al Tuwirqi AA, Alshammari AM, Felemban OM, Ali Farsi N. Comparison of penetration depth and microleakage of resin infiltrant and conventional sealant in pits and fissures of permanent teeth *in vitro*. J Contemp Dent Pract 2019; 20: 1339-44.
- 57. Vohra F, Altwaim M, Alshuwaier AS, Alomayri A, Al Deeb M, AlFawaz YF, et al. Bond integrity and microleakage of dentinbonded crowns cemented with bioactive cement in comparison to resin cements: *in vitro* study. J Appl Biomater Funct Mater 2020; 18: 2280800020905768.
- 58. Nielsen RB, Alyassin AM, Peters DD, Carnes DL, Lancaster J. Microcomputed tomography: an advanced system for detailed endodontic research. J Endod 1995; 21: 561-8.
- 59. Acar B, Kamburoğlu K, Tatar İ, Arıkan V, Çelik HH, Yüksel S, et al. Comparison of micro-computerized tomography and conebeam computerized tomography in the detection of accessory canals in primary molars. Imaging Sci Dent 2015; 45: 205-11.
- 60. Grande NM, Plotino G, Gambarini G, Testarelli L, D'Ambrosio F, Pecci R, et al. Present and future in the use of micro-CT scanner 3D analysis for the study of dental and root canal morphology. Ann Ist Super Sanita 2012; 48: 26-34.
- Chen G, Fan W, Mishra S, El-Atem A, Schuetz MA, Xiao Y. Tooth fracture risk analysis based on a new finite element dental structure models using micro-CT data. Comput Biol Med 2012; 42: 957-63.
- 62. Wang S, Xu Y, Shen Z, Wang L, Qiao F, Zhang X, et al. The extent of the crack on artificial simulation models with CBCT and periapical radiography. PLoS One 2017; 12: e0169150.
- 63. Tufenkci P, Orhan K, Celikten B, Bilecenoglu B, Gur G, Sevimay S. Micro-computed tomographic assessment of the shaping ability of the One Curve, One Shape, and ProTaper Next nickeltitanium rotary systems. Restor Dent Endod 2020; 45: e30.
- 64. Robinson JP, Lumley PJ, Claridge E, Cooper PR, Grover LM, Williams RL, et al. An analytical Micro CT methodology for quantifying inorganic dentine debris following internal tooth preparation. J Dent 2012; 40: 999-1005.
- 65. Tanomaru-Filho M, Torres FF, Pinto JC, Santos-Junior AO, Tavares KI, Guerreiro-Tanomaru JM. Micro-computed tomographic evaluation of a new system for root canal filling using calcium silicate-based root canal sealers. Restor Dent Endod 2020; 45: e34.
- 66. Torres FF, Guerreiro-Tanomaru JM, Chavez-Andrade GM, Pinto JC, Berbert FL, Tanomaru-Filho M. Micro-computed tomographic evaluation of the flow and filling ability of endodontic materials using different test models. Restor Dent Endod 2020; 45: e11.
- 67. Kim JC, Moe MM, Kim SK. A micro-computed tomographic evaluation of root canal filling with a single gutta-percha cone and calcium silicate sealer. Restor Dent Endod 2020; 45: e18.