

Caries Research

Manuscript:	CRE-0-0-0
Title:	Diagnosis of interproximal caries lesions in bitewing radiographs using a deep convolutional neural network-based software.
Authors(s):	Ángel García-Cañas (Co-author), Mónica Bonfanti-Gris (Co-author), Sergio Paraíso-Medina (Co-author), Francisco Martínez-Rus (Corresponding Author), Guillermo Pradíes (Co-author)
Keywords:	Artificial Intelligence, Caries detection, Computer-Aided Diagnosis, Convolutional Neural Networks
Type:	Research Article

Research Article

Diagnosis of interproximal caries lesions in bitewing radiographs using a deep convolutional neural network-based software.

Ángel García-Cañas^a, Mónica Bonfanti-Gris^a, Sergio Paraíso-Medina^b, Francisco Martínez-Rus^a,
Guillermo Pradíes^a

^a Department of Conservative and Prosthetic Dentistry, Faculty of Dentistry, Complutense University of Madrid, Madrid, Spain.

^b Department of Informatic Systems and Languages. Faculty of Software Engineering, Polytechnic University of Madrid, Madrid. Spain.

Short Title: Diagnosis of caries using artificial intelligence.

Corresponding Author:

Francisco Martínez-Rus

Department of Conservative and Prosthetic Dentistry

Faculty of Dentistry, Complutense University of Madrid

Plaza Ramón y Cajal, s/n.

Madrid, 28040, Spain

Tel. +34 91 3941923

Email: framarti@ucm.es

Number of Tables: 2

Number of Figures: 3

Word Count: 3959

Keywords: Artificial Intelligence; Caries Detection; Computer-Aided Diagnosis; Convolutional Neural Networks.

Abstract

The aim of this study was to evaluate the diagnostic reliability of a web-based artificial intelligence program for the detection of interproximal caries in bitewing radiographs. Three hundred bitewing radiographs of patients were subjected to the evaluation of a convolutional neural network-based software. Four different models were established to analyze its evaluations according to the confidence threshold (0-100%) offered by the program: model 1 (values > 0% were considered positive and values of 0% were considered negative), model 2 (values $\geq$ 25% were considered positive and values < 25% were considered negative), model 3 (values $\geq$ 50% were considered positive and values < 50% were considered negative), and model 4 (values $\geq$ 75% were considered positive and values < 75% were considered negative). For enamel caries, clinical validation included a combination of clinical-visual and radiography assessments. For dentin caries, clinical validation was performed by instrumentally accessing to the cavity. The accuracy rate (A), sensitivity (S), specificity (E), positive predictive value (PPV), negative predictive value (NPV), positive likelihood ratio (PLR), negative likelihood ratio (NLR), and areas under receiver operating characteristic curves (AUC) were calculated for the four models of agreement with the software. Models showed the following results respectively: A= 70.8%, 82%, 85.6%, 86.1%; S= 87%, 69.8%, 57%, 41.6%; E= 66.3%, 85.4%, 93.7%, 98.5%; PPV=42%, 57.2%, 71.6%, 88.6%; NPV=94.8%, 91%, 88.6%, 85.8%; PLR= 2.58, 4.78, 9.05, 27.73; NLR= 0.2, 0.35, 0.46, 0.59; AUC=0.767, 0.777, 0.753, 0.701. The deep convolutional neural network-based software tested may highlight suspicious interproximal areas, increasing clinical efficiency by focusing human effort towards specific sites that need further evaluation, showing the model 2 the best results to differentiate between healthy teeth and decayed teeth.

Introduction

Dental caries is defined as a non-communicable multifactorial dynamic disease, biofilm-mediated and diet-modulated, which results in net mineral loss of dental hard tissues. It is determined by biological, behavioral, psychosocial and environmental factors and is considered the most prevalent condition worldwide, since it affects practically the entire population at some point in their life [Selwitz et al., 2007; Lee et al., 2018].

The visual-tactile method is the most frequently used technique for the diagnosis of dental caries. However, it must be complemented with the performance of radiographic tests. The radiological test of choice for the diagnosis of this condition is the bite-wing radiograph, which has a higher sensitivity than panoramic radiographs, especially for the detection of interproximal lesions in posterior teeth (molars and premolars), where clinical detection either visually or through a dental explorer is specially challenging [Selwitz et al., 2007; Schwendicke et al., 2015; Prados-Privado et al., 2020]. However, although it is considered to be a highly reliable method for the detection of dental caries, final diagnosis can be influenced by the operators' subjectivity, as well as by various factors including both monitor and viewing conditions [Lee et al., 2018].

During the last decades, great technological advances have been developed in all areas of our lives. This technological revolution has also affected the healthcare field, where computer-aided diagnosis has significantly evolved due to the progress of big data and artificial intelligence [Tuzoff et al., 2019]. Artificial intelligence (AI) is considered a field of engineering that implements concepts and solutions to solve complex situations [Hamet and Trembay, 2017]. Machine learning and deep learning are two subfields of artificial intelligence. These procedures participate in the study and construction of algorithms that are capable of learning from information and making predictions [Mupparapu et al., 2018]. Machine learning allows algorithms to identify complex patterns among a large amount of data establishing their own rules for detecting similar patterns in new data sets. Deep learning, instead, is a specialized type of machine learning that is based on a layered learning model that processes information. The basis of deep learning are the neural networks, a system that mimics the functioning of the human brain and its neurons [Mupparapu et al., 2018; Prados-Privado et al., 2020]. Big data refers to the storage and processing of massive amounts of structured, semi-structured and unstructured data with great potential to be extracted and organized in a way that provides valuable information by using artificial intelligence.

The development of this technology has shown certain difficulties in dentistry due to the learning curve involved in the process, as well as the limited availability of large data sets of medical images. Despite this, in recent years there have been wide-ranging advances in the field that have led researchers and

health professionals to recognize the important role of AI. This reinforces the idea that it could definitely be used for the diagnosis of dental caries, among other oral pathologies [Mazurowski et al., 2019]. Recently, an automatic web-based artificial intelligence software that generates an automatic diagnosis of radiological images (orthopantomography and intraoral radiographs) by means of machine learning technologies (Denti.Ai; Denti.Ai Technology Inc, Toronto, ON, Canada) was released and made available on the internet. As it was previously stated, the early diagnosis of dental caries continues to be a challenge for dentists. Artificial intelligence can perform these tasks in a simple, fast and precise way, thus helping the dentist in their daily practice, reducing work time and improving clinical decision making [Prajapati et al., 2017]. Although at present artificial intelligence in dentistry has been used mainly for the detection and identification of anatomical structures and teeth, the scientific literature about its use in dental caries diagnosis is scarce [Lee et al., 2018]. Bearing this in mind, the aim of this research was to compare the efficacy of a deep convolutional neural network-based software against the gold standard on detecting proximal caries lesions in enamel and dentin. The null hypothesis was that no significant differences would be found in the diagnostic performance of a machine learning software versus the clinical validation.

Material and methods

Study design and patient selection

This study was designed as an analytical, observational and cross-sectional study. The Standards for Reporting Diagnostic Accuracy (STARD) statement and the Checklist for Artificial Intelligence in Medical Imaging (CLAIM) were followed in planning, conducting and reporting this investigation [Bossuyt et al., 2015; Mongan et al., 2020; Schwendicke et al., 2021]. The sample size was calculated using a standard formula for prevalence studies [Naing et al., 2006] as follows:

$$n = \frac{Z^2 P(1 - P)}{d^2}$$

where n is sample size, Z is the value of standard normal deviation corresponding to the level of confidence (for a 95% confidence level, the value of Z is 1.96), P is the expected prevalence, and d is the absolute precision. Assuming that the mean prevalence of un-treated caries in permanent teeth in Spain is 19.46% [Bravo-Pérez et al., 2020], the calculated sample size was 241 to achieve a confidence level of 95% and an absolute precision of 5%. Based on this data, a final sample size of 300 bitewing radiographs was included.

Between July 2019 and July 2020, 150 patients aged between 18 and 85 years (81 women, 69 men, mean age: 43 years) who came for a check-up of their oral health status in a private dental clinic were recruited in the study. The inclusion and exclusion criteria are listed in Table 1. All patients were

evaluated and treated by the same trained clinician (A.G.-C.). The intra-examiner Kappa coefficient value prior to the start of the inspections was considered acceptable since it was found within a range of 0.81 and 0.85 ($p = 0.04$).

Clinical-visual examination

The dentist carefully conducted clinical-visual examination of each tooth using standard conditions. The lesion severity was scored according to the International Caries Detection and Assessment System (ICDAS) [Ekstrand et al., 2018]. The criteria were (0) sound tooth, (1) first visual change in enamel, (2) distinct visual change in enamel, (3) localized enamel breakdown, (4) underlying dark shadow originating from dentin, (5) distinct cavity with visible dentin, and (6) extensive distinct cavity with visible dentin. Codes of 1, 2, and 3 were used for the diagnosis of enamel caries, and codes of 4, 5 and 6 were used for the diagnosis of dentin caries.

Radiography examination

A total of 300 digital bitewing radiographs of premolars and permanent molars -both upper and lower- were performed by the same clinician during the appointment. Bitewing radiographs were acquired and processed following identical conditions. The X-ray unit (CS 2100; Carestream, Rochester, NY, USA) was adjusted to the irradiation parameters of 60 kV, 7 mA, and 0.125 s, with a focus-sensor distance of 20 cm. The images were obtained using a radiovisiography sensor (RVG 6100; Carestream) positioned with a bitewing holder (XCP DS Fit; Dentsply Sirona, Charlotte, NC, USA) and the manufacturer's software (CS Imaging Version 8; Carestream). The presence of proximal enamel or dentin caries was determined using the International Caries Classification and Management System (ICCMS) [Ismail et al., 2015]: (0) no radiolucency, (1) radiolucency in the outer 1/2 of the enamel, (2) radiolucency in the inner 1/2 of the enamel $\pm$ enamel-dentin junction, (3) radiolucency limited to the outer 1/3 of dentin, (4) radiolucency reaching the middle 1/3 of dentin, (5) radiolucency reaching the inner 1/3 of dentin, clinically cavitated, and (6) radiolucency into the pulp, clinically cavitated. For enamel caries, codes of 1 or 2 were used; for dentin caries, codes of 3, 4, and 5 were used in the classification. All radiographs were anonymized and stored in an encrypted digital folder.

Radiography assessment using a deep convolutional neural network-based software

The pre-trained convolutional neural network-based software Denti.AI was used to analyze all radiographs. Denti.AI is an automated online software based on artificial intelligence that is capable of generating an automatic diagnosis of radiological images using deep learning technologies. This program is compatible with orthopantomographs and intraoral radiographs, both periapical and bitewing. To carry out the task of detecting and classifying dental structures found in the radiographs, this program uses different detection modules: first, the image is processed to establish the limits of

each tooth present and then marks the structures found with bounding boxes. The tooth numbering module then classifies each delimited tooth region according to the *Federation Dentaire Internationale* (FDI) dental nomenclature. Regarding the diagnosis of pathologies (carious and periapical lesions) and the detection and classification of dental treatments (composite or amalgam fillings, root canal treatments, crowns and implants, etc), the program is executed by a coding system based on a gray scale and offers a confidence threshold for each find (from 0 to 100%) [Tuzoff et al, 2019]. Figure 1 shows an overview of caries detection process used.

Clinical validation

For enamel caries lesions, the gold standard was established by combining the clinical-visual and digital bitewing radiography assessments. With a consensus about enamel caries, the cavity was not opened for ethical reasons [Schwendicke et al., 2019]. For dentin caries, the gold standard was established by opening the cavity and the application of caries detector dye (Kurakay Caries Detector; Kuraray Noritake Dental, Tokyo, Japan). Detected dentin caries lesions were opened with a conical diamond bur and hand excavator until the soft carious tissue was completely removed. After the cavity was opened, lesions were restored with a filling material appropriate for the cavity.

Data analysis

The results obtained from the machine learning software were recorded in a spreadsheet (Microsoft Excel; Microsoft Corp, Redmond, WA, USA), where the following information was included: side, tooth, location, presence or absence of caries and confidence threshold (0-100%) offered by the software. Since the program offered the evaluation in a % bar representation, to obtain a reasonable and practice interpretation of the outcomes, it was decided to establish four models to analyze the results offered by the machine learning software:

- Model number 1, where values greater than 0% were considered positive, and values equal to 0% were considered negative.
- Model number 2, where values greater than or equal to 25% were considered positive, and values less than 25% were considered negative.
- Model number 3, where values greater than or equal to 50% were considered positive, and values less than 50% were considered negative.
- Model number 4, where values greater than or equal to 75% were considered positive, and values less than 75% were considered negative.

The diagnostic performance of the deep convolutional neural network-based software in determining enamel and dentin caries according to the gold standard was expressed in terms of accuracy rate (A), sensitivity (S), specificity (E), positive predictive value (PPV), negative predictive value (NPV), positive

likelihood ratio (PLR), negative likelihood ratio (NLR), receiver operating characteristic (ROC) curves and areas under the curves (AUC). Accordingly, the following values were calculated: true positives (TP), false positives (FP), true negatives (TN), and false negatives (FN). Data was analyzed using the software SPSS 26.0 for Mac (IBM Corporation, Armonk, NY, USA).

Results

Figure 2 shows the distribution of the analyzed teeth in this study. Out of a total of 1323 teeth, the first molar was the most studied tooth in the sample (42%), followed by the second premolar (32%), second molar (20%) and finally the first premolar (6%). The reference test showed 579 positive evaluations (caries) and 2067 negative evaluations (healthy), resulting in a sample prevalence of 21.9%. Table 2 shows the values of overall accuracy, specificity, positive predictive value, positive likelihood ratio and negative likelihood ratio, which increase as the model raises the threshold confidence; and sensitivity and negative predictive value, which decrease as the model raises the threshold confidence offered by the four models established for the AI web-based software. A comparison of the ROC curves offered by the four models established for the software can be seen in Figure 3. The area under the curve for model 1 was 0.767 (95% CI 0.746-0.787), for model 2 was 0.777 (95% CI 0.753-0.800), for model 3 was 0.753 (95% CI 0.727-0.779), and for model 4 was 0.701 (95% CI 0.673-0.728). The Pearson's chi-squared test indicated significant differences among the four models of the machine learning software versus the gold standard ($p < 0.001$).

Discussion/Conclusion

This study evaluated the diagnostic performance of a deep convolutional neural network-based software against the gold standard for detecting proximal caries lesions in enamel and dentin. The null hypothesis was rejected because statistical differences were obtained between the results obtained from the machine learning software versus the clinical validation, being the model 2 the best alternative when compared with the gold standard.

The caries diagnosis through bitewing radiographs present some limitations due to the lack of agreement among professionals -especially in those inexperienced- which result in a considerable amount of false positives that can cause iatrogenesis and false negatives that can cause the progression of the lesion and finally more serious pathologies [Srivastava et al., 2017; Cantu et al., 2020]. The introduction of artificial intelligence programs to assist in caries detection and diagnosis could reduce the aforementioned shortcomings, providing the professional with a more reliable and accurate diagnosis [Lee et al., 2018; Geetha et al., 2020; Prados-Privado et al., 2020].

Currently, several scientific studies related to the use of convolutional neural networks in the diagnosis of caries in intraoral radiographs have been published [Yu et al., 2006; Li et al., 2007; Albahbah et al.,

2016; Srivastava et al., 2017; Lee et al., 2018; Cantu et al., 2020; Geetha et al., 2020]. All of them showed positive results in favor of artificial intelligence, but the web-based program tested in this study (Denti.Ai) was not included in any of them. The results of our study show similar results compared to related works, which share a similar methodology [Yu et al., 2006; Li et al., 2007; Albahbah et al., 2016; Srivastava et al., 2017; Lee et al., 2018; Cantu et al., 2020; Geetha et al., 2020]. The highest overall accuracy is presented by model 4 with 86.1%, followed by model 3 (85.6%), model 2 (82%) and model 1 (70.8%). The same happened with the specificity values, model 4 had the highest capacity to detect healthy teeth with 98.5%, followed by model 3 (93.7%), model 2 (85.4%) and model 1 (66.3%). As the confidence threshold was increased, the program improved its ability to detect true negatives and therefore increased the percentage of total correct answers. However, the opposite occurred for sensitivity values. The program worsened its ability to detect decayed teeth: model 1 showed the best results (87%), followed by model 2 (69.8%), model 3 (57%) and model 4 (41.6%). Both testing parameters - sensitivity and specificity -, are inherent properties of the test regardless of the disease prevalence in the sample studied and offer us an approximation of the program capacity to identify caries. However, both values are given by inverse probability weighting, which limits the possibility of making intuitive clinical interpolations for each patient [Gansky 2003].

Regarding the probability of presenting dental caries with a positive result, as the value of the threshold confidence that determined the presence of cavities increased, the PPV increased. This may be due to the fact that the higher the percentage offered by the web-based software, the greater the probability of success, and, as a result, model 4 offered a percentage of 88.6% and model 1 of 42%. The difference between models is less for the NPV, being 94.8% for model 1 and 85.8% for model 4. One aspect to be highlighted is that these values, despite they are extremely useful when making clinical decisions and transmitting information to patients, are highly influenced by the condition prevalence in the sample studied. When the prevalence is low, NPV increases and PPV decreases, being the opposite when the prevalence is high. For this reason, PPV and NPV should not be used as rates to compare two different diagnostic methods, nor to extrapolate results obtained in other studies.

For this reason, it is necessary to determine alternative statistical tests that do not depend on the prevalence of the disease and offer a clinical utility that allows to reflect the diagnostic strength of the test, such as the likelihood ratio. The likelihood ratio is the likelihood that a given test result would be expected in a patient with the target characteristic compared to the likelihood that that same result would be expected in a patient without the target characteristic. According to the ranges of likelihood ratio values [Hayden and Brown, 1999], model 4 presented a highly relevant PLR and a bad NLR, model 3 a good PLR and a regular NLR, model 2 a regular PLR and NLR, and model 1 a regular PLR and a good NLR.

Another parameter independent of prevalence and with high clinical utility is the ROC curve, obtained from the sensitivity and specificity values of the test. The area under this curve is the best indicator of the predictive ability of the test, allowing to establish comparisons between different diagnostic tests. In this research, model 2 offered the highest value of the area under the curve (0.777), followed by model 1 (0.767), model 3 (0.753), and finally model 4 (0.701).

Although the null hypothesis was rejected, we consider that the web-based software analyzed can be used as a reliable tool in the radiological diagnosis of interproximal caries. Model 2 deserves special consideration, since it offers the best results to differentiate between healthy teeth and decayed teeth. However, it should not be ignored the importance of analyzing the data set from a global point of view, in order to take advantage of the benefits offered by each model. Similarly, it must be beared in mind that technology does not establish direct causal relationships, so the information provided must be interpreted with caution by users [Chen et al., 2020]. It is important to highlight that these programs must be developed and implemented in collaboration with experts from the different areas of health and computer engineering to guarantee the least possible number of errors [Mupparapu et al, 2018]. Also, it should be noted that the final responsibility for diagnosis and therapeutic intervention lies with the professional and never with technologies [Chen et al., 2020].

Concerning the methodology of dental caries scientific studies, it is important to establish an initial reference test to compare the results offered by the artificial intelligence program. The method considered as gold standard for the diagnosis of caries lesions is the histopathological examination [Downer, 1975]. Another reference test can be the one established in this work, the diagnosis of an experienced dentist based on the clinical examination according to the ICDAS criteria [Ekstrand et al., 2018], radiographic tests according to the ICCMS criteria [Ismail et al., 2015] and the subsequent cavity opening and obturation. At the same time, the level of intra- and inter-observer agreement of the expert dentists in charge of evaluating radiographic images should be taken into consideration. The scientific literature reports that Kappa values between 0.81 and 1.00 indicate excellent concordance [Rigby, 2000].

As to the data set that is used as training of the artificial intelligence program, they are based on evaluations from human professionals, so the quality of the input information is critical for the correct development of technology [Prados-Privado et al., 2020]. Nowadays, there is research where artificial intelligence applications, used in areas such as breast cancer diagnosis [McKinney et al., 2020] or skin cancer diagnosis [Haenssle et al., 2018], have showed better results than human professionals.

There were some limitations in the present study. First the sources of random error such as the variability of the teeth, the trained dentist and the radiology tools [Yu et al., 2006]. In turn, non-

Diagnosis of caries using artificial intelligence.

probability sampling leads to a worse distribution of the sample variability. Another limitation to be pointed out is the exclusion of temporary teeth, which prevents the data from being applied to the child and adolescent population who have at least one temporary tooth. Finally, carious lesions have not been differentiated according to their depth, which could mean obtaining relevant data about the reliability of the artificial intelligence program and the stage of the disease. Hence, a further study is necessary by training the system with a much larger dataset since the higher the number of subjects used to train the neural network, the higher the accuracy it will have for testing unknown objects. Deep learning techniques can be applied to numerous other tasks than carious detection. Several exciting prospects are currently being studied, for example, using machine learning technologies to detect pathologies, such as periodontitis and periapical cysts. Admittedly, to achieve expert-level results for such a task, these new systems would require larger datasets for training than used in this project.

The purpose of artificial intelligence is not to exclude the input of the clinician from the process of radiographic dental caries diagnosis but to highlight suspicious areas, increasing clinical efficiency by focusing human effort towards specific sites that need further evaluation. Thus, the deep convolutional neural network-based software tested is capable of being used as a reliable tool in the radiological diagnosis of interproximal caries, showing the model 2 the best results to differentiate between healthy teeth and decayed teeth.

Diagnosis of caries using artificial intelligence.

271 **Statement of Ethics**

272 This study protocol was reviewed and approved by the Ethics Committee of the Complutense
273 University of Madrid, approval number 21/303-E. All patients provided written informed consent to
274 participate in the study according to the Declaration of Helsinki.

275 **Conflict of Interest Statement**

276 The authors have no conflicts of interest to declare.

277 **Funding Sources**

278 This research did not receive any specific grant from funding agencies in the public, commercial or
279 non-for-profit sectors.

280 **Author Contributions**

281 G.P. and S.P.-M conceived and designed the study. A.G.-C. and M.B.-G. performed the study. G.P., S.P.-
282 M and F.M.-R. analyzed and interpreted the data. A.G.-C., M.B.-G, and F.M.-R wrote the paper. All of
283 the authors approved the final version of this paper and are accountable for all aspects of this work.

284 **Data Availability Statement**

285 The data that support the findings of this study are not publicly available due to their containing
286 information that could compromise the privacy of research participants but are available from the
287 corresponding author [F.M.-R.] on reasonable request.

References

- Albahbah AA, El-Bakry HM, Abd-Elgahany S. Detection of caries in panoramic dental X-ray images using back-propagation neural network. *Int J Electron Commun Comput Eng*. 2016;7(5):250-6.
- Bossuyt PM, Reitsma JB, Bruns DE, Gatsonis CA, Glasziou PP, Irwig L, Lijmer JG, Moher D, Rennie D, de Vet HC, Kressel HY, Rifai N, Golub RM, Altman DG, Hooft L, Korevaar DA, Cohen JF; STARD Group. STARD 2015: an updated list of essential items for reporting diagnostic accuracy studies. *BMJ*. 2015 Oct 28;351:h5527.
- Bravo-Pérez M, Almerich-Silla JM, Canorea-Díaz E, Casal-Peidró E, Cortés-Martinicorena FJ, Gómez-Santos G et al. Encuesta de Salud Oral en España 2020. *RCOE*. 2020;25:12-69.
- Cantu AG, Gehrung S, Krois J, Chaurasia A, Rossi JG, Gaudin R, et al. Detecting caries lesions of different radiographic extension on bitewings using deep learning. *J Dent*. 2020 Sep;100:103425.
- Chen YW, Stanley K, Att W. Artificial intelligence in dentistry: current applications and future perspectives. *Quintessence Int*. 2020;51(3):248-57.
- Downer MC. Concurrent validity of an epidemiological diagnostic system for caries with the histological appearance of extracted teeth as validating criterion. *Caries Res*. 1975;9(3):231-46.
- Ekstrand KR, Gimenez T, Ferreira FR, Mendes FM, Braga MM. The International Caries Detection and Assessment System - ICDAS: A Systematic Review. *Caries Res*. 2018;52(5):406-419.
- Gansky SA. Dental data mining: potential pitfalls and practical issues. *Adv Dent Res*. 2003 Dec;17:109-14.
- Geetha V, Aprameya KS, Hinduja DM. Dental caries diagnosis in digital radiographs using back-propagation neural network. *Health Inf Sci Syst*. 2020 Jan 3;8(1):8.
- Haenssle HA, Fink C, Schneiderbauer R, Toberer F, Buhl T, Blum A, et al. Man against machine: diagnostic performance of a deep learning convolutional neural network for dermoscopic melanoma recognition in comparison to 58 dermatologists. *Ann Oncol*. 2018 Aug 1;29(8):1836-42.
- Hamet P, Tremblay J. Artificial intelligence in medicine. *Metabolism*. 2017 Apr;69S:S36-S40.
- Hayden SR, Brown MD. Likelihood ratio: A powerful tool for incorporating the results of a diagnostic test into clinical decisionmaking. *Ann Emerg Med*. 1999 May;33(5):575-80.

Diagnosis of caries using artificial intelligence.

Ismail AI, Pitts NB, Tellez M, Authors of International Caries Classification and Management System (ICCMS). The International Caries Classification and Management System (ICCMS™) An Example of a Caries Management Pathway. BMC Oral Health. 2015;15 Suppl 1(Suppl 1):S9.

Lee JH, Kim DH, Jeong SN, Choi SH. Detection and diagnosis of dental caries using a deep learning-based convolutional neural network algorithm. J Dent. 2018 Oct;77:106-111.

Li W, Kuang W, Li Y, Li Y, Ye W. Clinical X-Ray Image Based Tooth Decay Diagnosis using SVM. In: 6th International Conference on Machine Learning and Cybernetics. Hong Kong; 2007; p. 1616-19.

Mazurowski MA, Buda M, Saha A, Bashir MR. Deep learning in radiology: An overview of the concepts and a survey of the state of the art with focus on MRI. J Magn Reson Imaging. 2019 Apr;49(4):939-954.

McKinney SM, Sieniek M, Godbole V, Godwin J, Antropova N, Ashrafian H, et al. International evaluation of an AI system for breast cancer screening. Nature. 2020 Jan;577(7788):89-94.

Mongan J, Moy L, Kahn CE Jr. Checklist for Artificial Intelligence in Medical Imaging (CLAIM): A Guide for Authors and Reviewers. Radiol Artif Intell. 2020 Mar 25;2(2):e200029.

Mupparapu M, Wu CW, Chen YC. Artificial intelligence, machine learning, neural networks, and deep learning: Futuristic concepts for new dental diagnosis. Quintessence Int. 2018;49(9):687-8.

Naing L, Winn T, Rusli BN. Practical issues in calculating the sample size for prevalence studies. Arch Orofac Sci. 2006;1:9-14.

Prados-Privado M, García Villalón J, Martínez-Martínez CH, Ivorra C, Prados-Frutos JC. Dental Caries Diagnosis and Detection Using Neural Networks: A Systematic Review. J Clin Med. 2020 Nov 6;9(11):3579.

Prajapati SA, Nagaraj R, Mitra S. Classification of dental diseases using CNN and transfer learning. In: 5th Int. Symp. Comput. Bus. Intell. ISCBI 2017. Institute of Electrical and Electronics Engineers Inc. New York; 2017; p. 70-4.

Rigby AS. Statistical methods in epidemiology. v. Towards an understanding of the kappa coefficient. Disabil Rehabil. 2000 May 20;22(8):339-44.

Schwendicke F, Tzschope M, Paris S. Radiographic caries detection: A systematic review and meta-analysis. J Dent. 2015 Aug;43(8):924-33.

Diagnosis of caries using artificial intelligence.

Schwendicke F, Splieth C, Breschi L, Banerjee A, Fontana M, Paris S, Burrow MF, et al. When to intervene in the caries process? An expert Delphi consensus statement. *Clin Oral Investig*. 2019 Oct;23(10):3691-3703.

Schwendicke F, Singh T, Lee JH, Gaudin R, Chaurasia A, Wiegand T, Uribe S, Krois J; IADR e-oral health network and the ITU WHO focus group AI for Health. Artificial intelligence in dental research: Checklist for authors, reviewers, readers. *J Dent*. 2021 Apr;107:103610.

Selwitz RH, Ismail AI, Pitts NB. Dental caries. *Lancet*. 2007 Jan;369(9555):51-9.

Srivastava MM, Kumar P, Pradhan L, Varadarajan S. Detection of tooth caries in bitewing radiographs using deep learning, *ArXiv*. DOI: 10.48550/arXiv.1711.07312.

Tuzoff DV, Tuzova LN, Bornstein MM, Krasnov AS, Kharchenko MA, Nikolenko SI, Sveshnikov MM, Bednenko GB. Tooth detection and numbering in panoramic radiographs using convolutional neural networks. *Dentomaxillofac Radiol*. 2019 May;48(4):20180051.

Yu Y, Li Y, Li Y, Wang J, Lin F, Ye W, Tooth Decay Diagnosis using Back Propagation Neural Network. 5th International Conference on Machine Learning and Cybernetics. Dalian; 2006, p. 3956-59.

Figure Legends

Fig. 1. Flowchart of the detection of dental caries in the deep convolutional neural network-based software: First, the radiological images were uploaded to the platform, which were immediately anonymized. The identification number of the patient, the gender, the date of the image and the date of birth of the patient were selected. Once all the fields were completed, the save option was clicked. Although the program offers the possibility of rotating the image, modifying brightness and contrast, and applying a negative or sharpening filter, none of these parameters were modified for the study. Once the definitive image was obtained, the analyze option was clicked and in a total of four seconds the program analyzed the radiological image, identifying the teeth present, calculating the distance from the alveolar bone to the to the amelocemental junction, detecting previous treatments and pathologies by offering a percentage that indicated the confidence threshold, and marking the findings in the image through green and red dashed lines. Finally, the program produced a written report that contained the data resulting from the process.

Fig. 2. Analyzed teeth distribution.

Fig. 3. Comparison of the ROC curves of models 1, 2, 3 and 4 of the deep convolutional neural network-based software.

Table Legends

Table 1. Inclusion and exclusion criteria employed in this study.

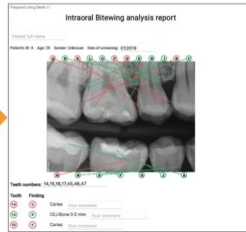
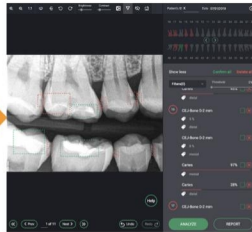
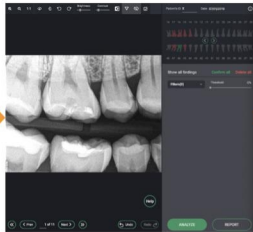
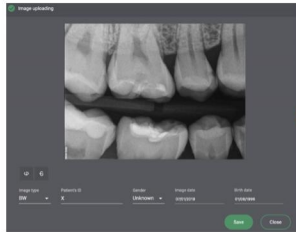
Table 2. Values of accuracy (A), sensitivity (S), specificity (E), positive predictive value (PPV), negative predictive value (NPV), positive likelihood ratio (PLR), negative likelihood ratio (NLR) showed by the four models of machine learning software.

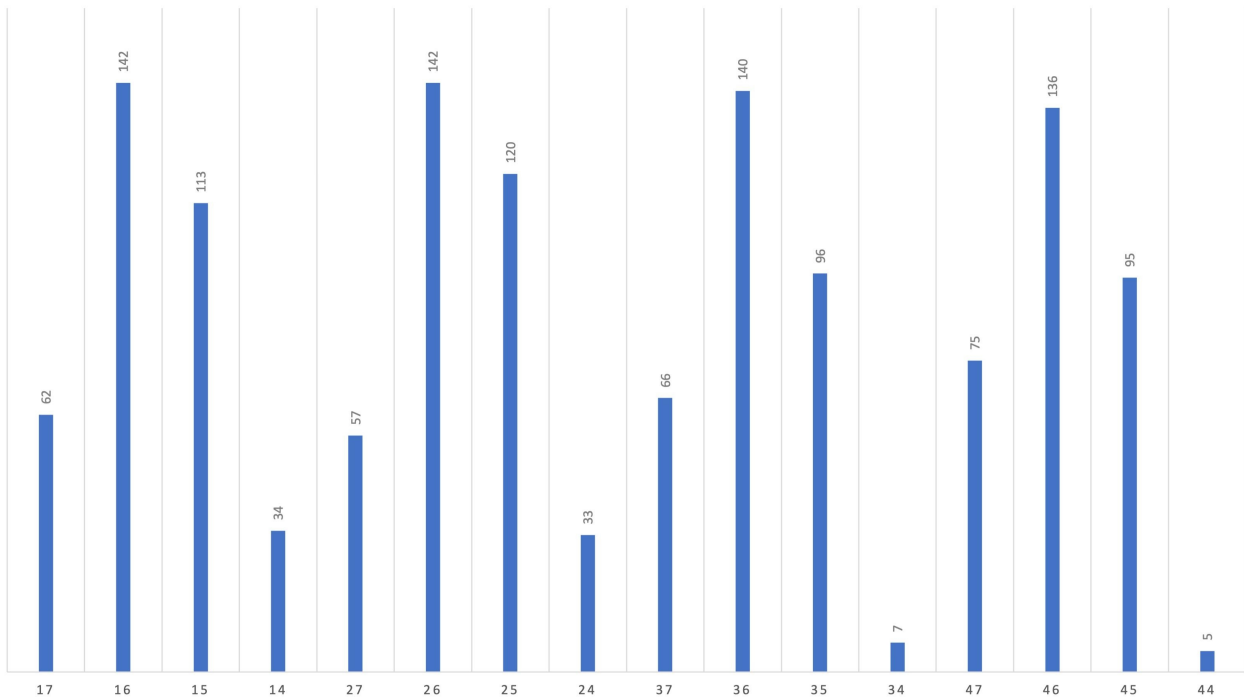
Table 1. Inclusion and exclusion criteria employed in this study.

Inclusion criteria
Patients who came for a check-up of their oral health status.
Age > 18 years.
Completely erupted permanent dentition.
No fixed orthodontic apparatus or prosthetic restoration.
Taking posterior bitewing radiographs.
Exclusion criteria
Patients who refused to participate in the research.
Posterior edentulous.
Dentition mixed o temporary.
Pregnant.
Bitewing radiographs where the interproximal limits were not clearly differentiated.

Table 2. Values of accuracy (A), sensitivity (S), specificity (E), positive predictive value (PPV), negative predictive value (NPV), positive likelihood ratio (PLR), negative likelihood ratio (NLR) showed by the four models of machine learning software.

Model	Overall Accuracy	Sensitivity	Specificity	Positive Predictive Value	Negative Predictive value	Positive Likelihood Ratio	Negative Likelihood Ratio
1	70.8%	87%	66.3%	42%	94.8%	2.58	0.20
2	82%	69.8%	85.4%	57.2%	91%	4.78	0.35
3	85.6%	57%	93.7%	71.6%	88.6%	9.05	0.46
4	86.1%	41.6%	98.5%	88.6%	85.8%	27.73	0.59





Receiver Operating Characteristic (ROC) Curves

