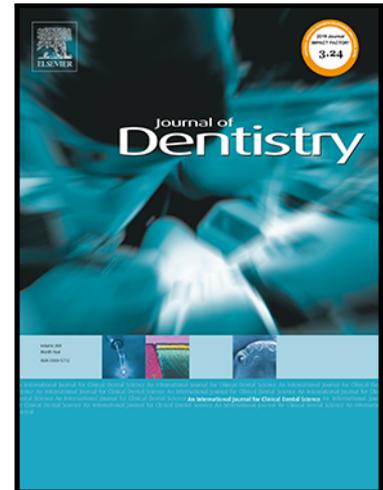


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Deep Learning for Caries Detection: A Systematic Review

Deep Learning for Caries Detection: A Systematic Review

Short Title: DL for Caries Detection

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Abstract

Objectives: Detecting caries lesions is challenging for dentists, and deep learning models may help practitioners to increase accuracy and reliability. We aimed to systematically review deep learning studies on caries detection.

Data: We selected diagnostic accuracy studies that used deep learning models on dental imagery (including radiographs, photographs, optical coherence tomography images, near-infrared light transillumination images). The latest version of the quality assessment tool for diagnostic accuracy studies (QUADAS-2) tool was used for risk of bias assessment. Meta-analysis was not performed due to heterogeneity in the studies methods and their performance measurements.

Sources: Databases (Medline via PubMed, Google Scholar, Scopus, Embase) and a repository (ArXiv) were screened for publications published after 2010, without any limitation on language.

Study selection: From 252 potentially eligible references, 48 studies were assessed full-text and 42 included, using classification (n=26), object detection (n=6), or segmentation models (n=10). A wide range of performance metrics was used; image, object or pixel accuracy ranged between 68%-99%. The minority of studies (n=11) showed a low risk of biases in all domains, and 13 studies (31.0%) low risk for concerns regarding applicability. The accuracy of caries classification models varied, i.e. 71% to 96% on intra-oral photographs, 82% to 99.2% on peri-apical radiographs, 87.6% to 95.4% on bitewing radiographs, 68.0% to 78.0% on near-infrared transillumination images, 88.7% to 95.2% on optical coherence tomography images, and 86.1% to 96.1% on panoramic radiographs. Pooled diagnostic odds ratios varied from 2.27 to 32767. For detection and segmentation models, heterogeneity in reporting did not allow useful pooling.

Conclusion: An increasing number of studies investigated caries detection using deep learning, with a diverse types of architectures being employed. Reported accuracy seems promising, while study and reporting quality are currently low.

Clinical significance: Deep learning models can be considered as an assistant for decisions regarding the presence or absence of carious lesions.

Keywords

Artificial intelligence, Machine learning, Neural Networks, Dental Caries, Dentistry, Systematic Review

1. Introduction

Artificial intelligence (AI) is defined as “the science and engineering of making intelligent machines” which can solve problems instead of humans [1]. Machine learning is one of the main subcategories of AI, enabling machines to learn and improve from experience without being explicitly programmed for a single task [2]. Typically, computers use example data that is extracted discriminant features, which are mostly handcrafted, from images in order to train machine learning systems [3].

As a subset of machine learning models, deep learning models are built based on neural networks (NNs), which are biologically inspired programming architectures that allow computers to learn by observing patterns in data [4]. These networks are composed of many layers, which transform input data (such as raw images) into outputs (such as diagnoses) while learning higher-level features automatically [3]. Convolutional NN (CNN), which is a modification of NN, has proven most successful for the analysis of images to date. They are designed to integrate spatial information and configuration of both 2D and 3D images [5].

CNNs are widely used in medical applications, including drug development, diagnostics, treatment, and associated processes [6, 7]. Deep learning has proven especially powerful for the analysis of complex data, like imagery [8], and has been used for image classification (i.e., labeling the image, e.g., signs of disease are present), detection (e.g. signs of disease are present in this area, usually indicated by a bounding box), and segmentation (i.e., signs of disease are present on these specific pixels). The accuracy of deep learning for medical image analysis has been found to match or, in some cases, surpass that of experts [9]. In dentistry, deep learning has been employed for image analysis in orthodontics [2], specifically landmark analysis on cephalometric radiographs [10], endodontics (detection of apical lesions), periodontology (periodontal bone loss) and cariology [9].

Dental caries is the most prevalent condition in the human population [11]. Caries lesions can be detected by visual and tactile means, while often this visual-tactile detection is supplemented by imaging strategies like radiography (the most common type of adjunct detection method), optical coherence tomography, quantitative light-induced fluorescence, intraoral scanner, or near-infrared light transillumination [12-14]. Evaluating any imagery for caries detection is a challenge for practitioners [15, 16]; dentists miss a substantial proportion of early caries lesions in radiographs [17] but also other image types [18-20], and show considerable variability in their diagnostic findings and treatment decisions.

The usage of AI, specifically deep learning, may support practitioners in caries lesion detection and diagnosis on imagery [15, 21-23]. Notably, the development in the field of deep learning of caries detection is highly dynamic; moreover, studies show substantial variability in methods and outcomes. We, therefore, aimed to systematically appraise studies using deep learning (specifically, NNs) for caries detection on dental imagery and compare their outcomes regarding

reported performance measurements including accuracy, specificity and sensitivity. We did not further specify the task (image classification, detection, segmentation) or the image modality. We further aimed to synthesize our findings and to evaluate the robustness of overall body of evidence.

2. Materials & Methods

This is a systematic review of diagnostic accuracy studies. Reporting of this study follows the PRISMA-DTA guideline [24]. The study protocol was registered at PROSPERO (CRD42019125491).

2.1. Eligibility Criteria

This systematic review answers the following PICO question: What are the applications and accuracy (outcome) of deep learning (intervention) for caries detection on dental imagery (population) in comparison with the reference test (additionally and, if available, also against the standard of care, i.e. human experts without deep learning)?

Studies reporting the following criteria were included:

P: Studies used deep learning models on dental imagery resulting from routine care, clinical studies or studies on extracted teeth;

I, C: Caries classification, detection, or segmentation models with a deep learning (NN) architecture, compared with a reference test

O: Any kind of accuracy estimate on image, tooth, surface or pixel level.

Studies with the following criteria were excluded: Studies without sufficient details on the data used for training and testing (e.g., dataset size, data modality, *etc.*); studies without a clear explanation of the deep learning model; studies that did not separate the accuracy of caries detection with that of detecting other pathologies; reviews.

2.2. Information sources and search

An electronic search was conducted in the following electronic databases up to 12th April 2021: Medline (via PubMed), Google Scholar, Scopus, Embase, and ArXiv. Search results were limited to publications after 2010, accounting for the fact that deep learning for image analysis has been popularized in 2012 by Krizhevsky et al. [25]. There was no limitation in the language. Each database was searched with adapted keywords. The search query for each database is described in Table 1. Journal articles and conference proceedings were screened to identify further studies. Moreover, manual cross-referencing of the bibliographies of included papers was conducted.

2.3. Study selection

For managing the citations, Endnote X9 (Clarivate, Philadelphia, USA) was used. Two independent reviewers performed title and abstract screening after removing duplicate papers (M.N and R.R). Any disagreement was resolved through consensus with a third reviewer (H.M.R). Then, two independent investigators evaluated full texts of eligible studies based on inclusion and exclusion criteria (M.N and R.R). Any disagreements or discrepancies between the two reviewers were resolved through consensus by a third investigator (H.M.R).

2.4. Data collection and extraction

Two reviewers (M.N. and E.M.) independently collected data from the included studies. A third reviewer (H.M.R.) revised the data collection for discrepancies and disagreements. The following data items were extracted: bibliographic details (name of authors and the year of publication), data modality, dataset size (train/valid/test, if given), inclusion and exclusion criteria on population and image level (if available), caries prevalence, labeling procedure (i.e. how the reference test was established), task (i.e. classification, detection, segmentation), pre-processing, augmentation, deep learning approach (NN architecture), loss function employed, hardware used, accuracy measures employed and resulting findings. If more than one NN architecture was used, we only reported on the most accurate one.

2.5. Risk of bias and applicability

Two reviewers (M.N. and E.M.) independently used the QUADAS-2 tool [26] for risk of bias assessment. The QUADAS-2 checklist contains four risk of bias domains, including data selection, index test, reference standard, and flow and timing. It also has three domains evaluating the applicability of a study on patient selection, index test, reference standard. Any disagreements or discrepancies between the two reviewers were resolved by a third investigator (H.M.R.).

In “patient selection”, limited information on the dataset being presented as well as unclear data-split strategies and resulting data leakage were considered to indicate high risk of bias. For “index test”, indicators were poor reporting on test reproducibility, insufficient information about the model construction and lack of robustness analyses of the model. For “reference standard”, indicators, were lack of information on reference standard definition and using only one examiner for establishing the reference test. Finally, in “flow and timing”, indicators were employing different reference standards across the same study and inappropriate intervals between the index test and reference standard. In the case of concerns regarding the applicability of the studies, we looked for the dataset used, the deep learning model and its performance, and the annotation procedure match review question in each domain, respectively. Leading questions used in the present study are reported in Table 2.

2.6. Synthesis

Due to a wide range of specific study designs and accuracy measures employed, we limited the quantitative synthesis to classification studies (as here, accuracy is always on image level; in detection and segmentation studies, accuracy can be determined on multiple levels, with accuracy measures only being limitedly comparable).

As very few studies reported the number of true positives (TPs), true negatives (TNs), false positives (FPs) and false negatives (FNs) samples, but reported sensitivity and specificity, we used the diagnostic odds ratios (DORs) as pooled outcome for classification studies, calculated as follows:

$$DOR = \frac{Sensitivity \times Specificity}{(1 - Sensitivity) \times (1 - Specificity)}$$

As a result, we limited the quantitative analysis to studies that reported both the specificity and the sensitivity. We further employed stratification according to detection threshold, i.e., enamel, enamel and dentin combined, only dentin, or unclear threshold.

3. Results

3.1. Study selection and study characteristics

Of 252 identified studies, a total of 48 studies were assessed full-text. Eleven of these studies were excluded after full-text assessment, the reasons for the exclusion are shown in Table S1. In total, following the manual search, 42 studies were included; the number of studies per year increased over the observation period, and the type of imagery became more diverse with time (Figure 1).

Individual studies are summarized in Tables 3, 4, and 5, with each table showing studies using different deep learning tasks (classification, detection, segmentation). Overall, seven different types of imagery were employed; intra-oral photographs (n=12), peri-apical radiographs (n=12), bitewing radiographs (n=8), optical coherence tomography images (n=4), panoramic radiographs (n=3), near-infrared light transillumination images (n=2), and cone-beam computed tomography (n=1) (Figure 2). Of the employed image datasets, three were publicly available; the international symposium on biomedical imaging (ISBI) 2015 challenge dataset of bitewing radiographs (used by n=4 studies) [27], a dataset of peri-apical radiographs (n=6 studies) [28], and a dataset of intra-oral photographs used in a Kaggle competition (n=1) [29]. One study reported that they employed peri-apical radiographs from a Kaggle competition, too, but the manuscript did not give further details towards the dataset.

Most studies used expert opinions to set the reference test (n=37). Specifically, one human expert (n=8), two (n=7), three or more (n=6) experts were involved in defining the reference test; 16 studies did not mention the number of experts involved. Two studies used other image modalities for establishing the reference test, namely fluorescent imaging [30] and micro-CT [31]. Four studies did not mention how the reference test was established.

Regarding the deep learning task, the most often chosen task was classification (n=26), followed by segmentation (n=10) and object detection (n=6). Various deep learning models were used (Figure 2). In classification studies, most of the studies used customized CNN structures (n=8), transfer learning models (n=8), or multi-layer perceptron (n=7). Regarding segmentation, auto-encoders were used (e.g., U-net) (n=8). Regarding detection, one-stage object detectors (e.g., YoLo) or two-stage object detectors (e.g., Faster R-CNN) were used in the majority of studies (n=4).

Classification studies by large reported on accuracy, sensitivity and specificity; other outcome measures were the F1-score, precision (positive predictive value) or the area-under-the receiver-operating-characteristics curve (AUC). In object detection and segmentation studies, outcome measures were more heterogeneous, and in addition to the mentioned ones, the mean Intersection over Union (mIoU) and the Dice coefficient index were employed (Tables 3-5).

3.2. Risk of bias and applicability

Detailed information about risk of bias and concerns regarding applicability are presented in Table S2 and Figure S2. Among the included studies, 11 (26.2%) were found to have low risk of biases in all four domains. Moreover, 13 studies (31.0%) were evaluated as low risk for concerns regarding applicability. The most problematic domain was “Reference Standard”, where only 20 studies (47.6%) and 22 studies (52.4%) were classified as low risk of bias and low risk of applicability concern, respectively.

3.3. Findings of the studies

The accuracy of deep learning for classifying images according to their caries status varied widely, i.e. 71% to 96% on intra-oral photographs, 82% to 99.2% on peri-apical radiographs, 87.6% to 95.4% on bitewing radiographs, 68.0% to 78.0% on near-infrared transillumination images, 88.7% to 95.2% on optical coherence tomography images, and 86.1% to 96.1% on panoramic radiographs.

As outlined, only classification studies were used for further synthesis. Of these, 17 studies could be pooled; the range of the reported sensitivity was 25% to 99.8% and specificity 17.2% to 100%, respectively. The majority of studies used both enamel and dentin lesions (i.e., a mixed threshold) or did not specify the threshold at all; those which defined the involvement as dentin as threshold found high sensitivity (above 90%) but mixed specificity (Figure 3). For enamel as threshold, the opposite was true; studies found high specificity (above 95%) but mixed sensitivity. For combined or other thresholds, no clear pattern evolved. Similarly, the training dataset size was not clearly associated with sensitivity or specificity. DORs varied from 2.27 to 32767; further analyses are displayed in Figure S2.

4. Discussion

The detection of early caries on imagery is challenging and marred by low sensitivity and inter-observer agreement [17]. Deep learning has been suggested to possibly assist in overcoming this problem, and in this systematic review, we systematically compiled and appraised studies involving deep learning for caries detection on images. We found a growing and increasingly promising body of evidence supporting deep learning for this task. Notably, though, studies were by large of limited quality and comparison across studies impeded by heterogeneity in conduct and reporting of the studies. A number of findings require more detailed discussion.

First, we found most studies to show relatively high accuracy, sensitivity, and specificity of deep learning for caries detection (usually exceeding 80%). Moreover, the few studies involving human experts as control group (and not only to establish the reference test) found deep learning similar or more accurate than humans. Generally, when considering the low accuracy of dentists especially for detecting early lesions [17], deep learning will likely yield useful accuracies for this task. Second, heterogeneity in study conduct and reporting was significant, impeding further comparisons across studies or quantitative synthesis. For example, studies employed a wide range of reference tests, with unclear impact on accuracy and bias. Particularly, a reference test defined by a single clinician only seems to come with serious bias, as any deep learning model will be as accurate (or worse) than this clinician in caries detection. Future studies are needed to (1) elaborate which strategies for establishing the reference test are most likely to yield the “truth” (e.g. majority vote, stepwise annotation, triangulation with second imagery), and (2) employ comprehensive and accurate data labeling to yield robust models [32]. Third, we had planned meta-analysis of accuracy outcomes if possible. For diagnostic accuracy studies, hierarchical models are recommended for such quantitative synthesis, but require the proportion TPs, TNs, FPs, and FNs as input data [33]. Since these values were reported by only a small minority of studies, this option was not available, though. Alternatively, separate pooling of sensitivity and specificity may be considered, but relies on the (unlikely) assumption that sensitivity and specificity are independent. Notably, this approach is also not recommended if studies used different diagnostic thresholds, as was the case here [33, 34], and requires estimates of variance of both measures (which, again, were unavailable in most studies). Hence, we eventually decided to compute summary DORs as a single parameter of accuracy [35]. However, these are hard to interpret by clinicians [33], and we only pooled classification studies assuming that DORs of detection or segmentation models are not meaningful from a clinical perspective (but also to account for the fact that for these studies, the level of the performance estimates was heterogeneous across studies). Fourth, one of the most critical shortcomings of the included studies was the lack of detail about the validation and calibration of the annotators and how the lack of inter-annotator agreement was handled. Generally, poor reporting should be addressed by future studies by adhering to minimum standards, e.g., STARD-AI; for diagnostic studies using AI models [36], CLAIM; a checklist for AI in medical imaging, [37], and recent reporting guidelines for dental AI [38]. Fifth, we found a wide range of imagery to be employed for deep learning tasks. Some, like near-infrared light transillumination or optical coherence tomography,

are clinically uncommon, and interpretation of such images may not be accurate in the hands of inexperienced examiners [18, 39-41]. The latter two types of imagery, however, are clinically promising, as they show high accuracy and do not require ionizing radiation, which is why deep learning seems useful to employ here overcoming the described “experience gap”. Last, it needs highlighting that nearly all included studies only determined the accuracy of deep learning, in a few cases comparing it against the standard of care (dentists without deep learning). Notably, the decisions derived from the diagnostic process are – eventually – more relevant to patients and payers alike, and it remains not understood if deep learning assistance will improve only accuracy, or also decision making and long-term treatment effectiveness and efficiency [42, 43]. Ideally, this impact and true usefulness of deep learning for caries detection should be explored in randomized, practice-based settings, and should also reflect aspects like lesion activity as an additional decision point (currently not at all considered by any of the included studies).

This study comes with a number of strengths and limitations. First, it comprehensively and systematically assessed studies on deep learning for caries detection, and hence allows to (at least narratively) synthesize and contrast them. Second, and as a limitation, our search was limited in its time horizon (which has been justified) and scope (only deep learning has been considered, while alternative image analysis methods have been used previously). Given that we yielded a large and diverse body of evidence, these restrictions seemed justified. Third, we had planned to conduct a meta-analysis, but given the heterogeneity and, more important, limited quality of reporting, we could not extract sufficient detail from most studies to perform this analysis (even if heterogeneity allowed).

5. Conclusions

An increasing number of studies investigated caries detection using deep learning, with a diverse type of architectures being employed. Reported accuracy seems promising, while study and reporting quality are currently low. Future should critically determine the reference test and rely on a comparable, broad and clinically meaningful outcome set.

Declaration of interests

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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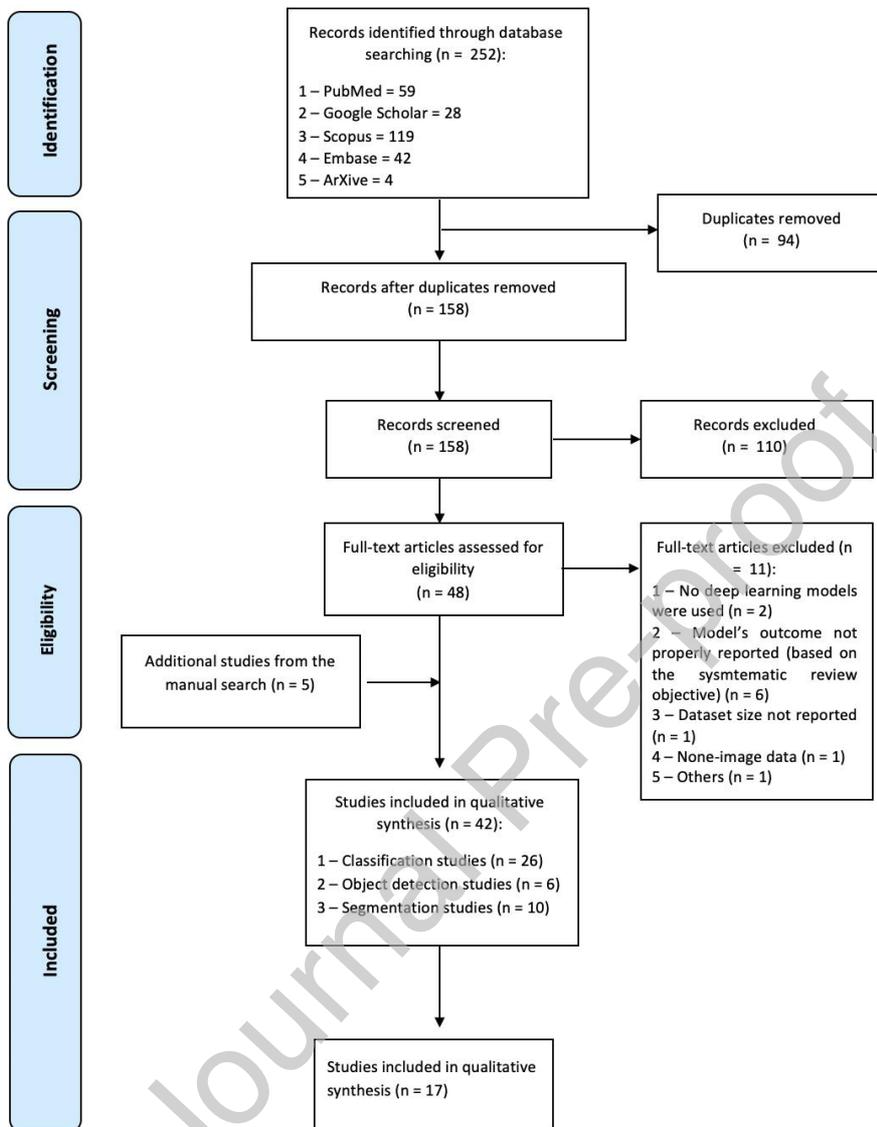


Figure 1 – Flowchart of the search

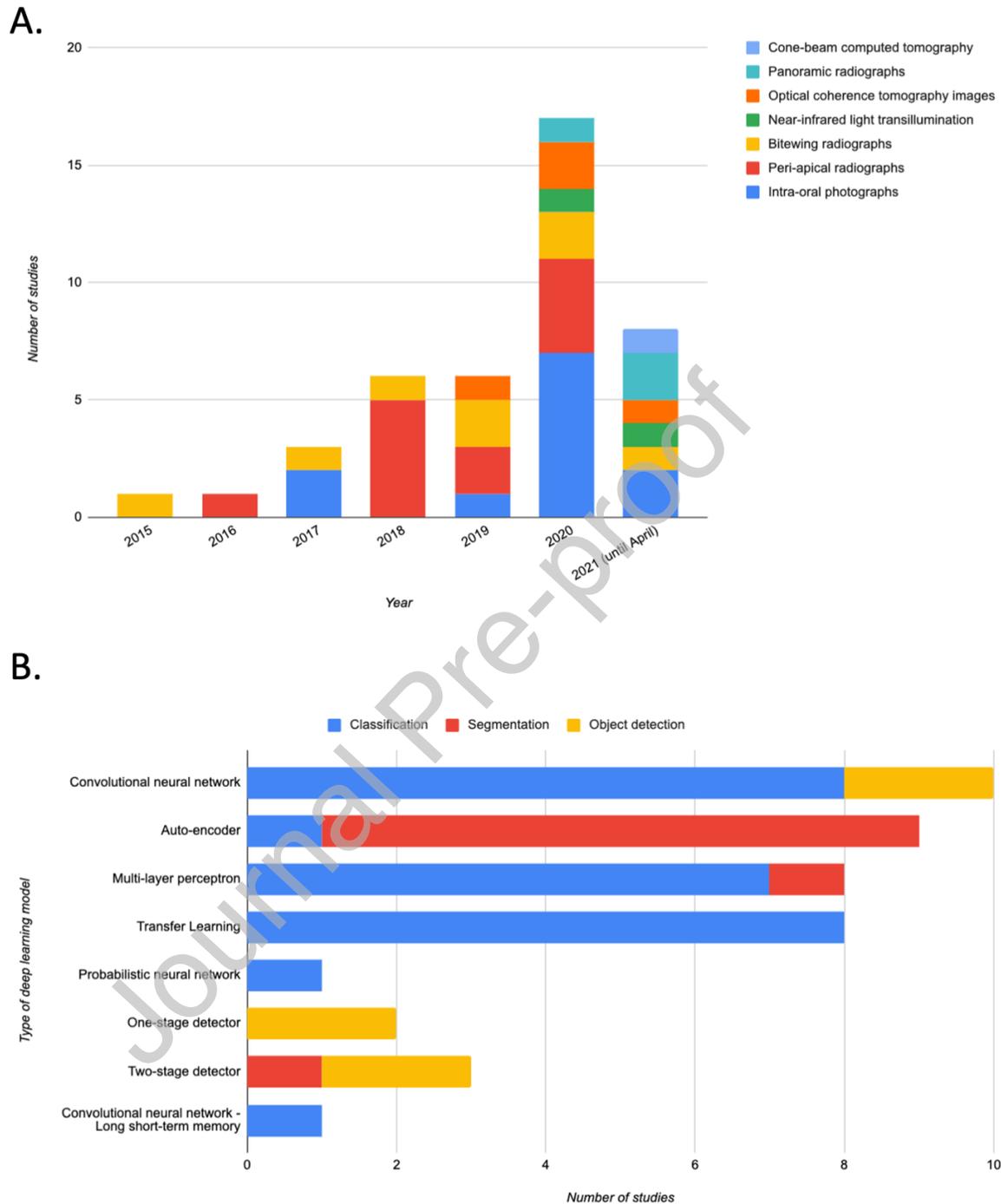


Figure 2 - Included studies; **A)** Number of deep learning studies for caries detection and image type employed. **B)** Tasks and modeling approaches involved (n=studies).

Table 1 – The results of the electronic search in the various databases

Database	Keywords	Results	Date
PubMed	("Artificial Intelligence" OR artificial intelligence [MeSH Terms] OR "deep learning" OR "deep learning"[Mesh Terms] OR "machine learning" OR "machine learning"[MeSH Terms] OR "neural network" OR "computer vision") AND caries	59	12 th April 2021
Google Scholar	allintitle:("artificial intelligence" OR "deep learning" OR "machine learning" OR "computer vision" OR "neural network") AND caries	28	12 th April 2021
Scopus	TITLE-ABS-KEY (("Artificial Intelligence" OR "deep learning" OR "machine learning" OR "neural network" OR "computer vision") AND ("caries")) AND (LIMIT-TO (PUBYEAR , 2021) OR LIMIT-TO (PUBYEAR , 2020) OR LIMIT-TO (PUBYEAR , 2019) OR LIMIT-TO (PUBYEAR , 2018) OR LIMIT-TO (PUBYEAR , 2017) OR LIMIT-TO (PUBYEAR , 2016) OR LIMIT-TO (PUBYEAR , 2015) OR LIMIT- TO (PUBYEAR , 2014) OR LIMIT-TO (PUBYEAR , 2012) OR LIMIT-TO (PUBYEAR , 2011) OR LIMIT-TO (PUBYEAR , 2010))	119	12 th April 2021
Embase	('artificial intelligence':ti,ab,kw OR 'deep learning':ti,ab,kw OR 'machine learning':ti,ab,kw OR 'neural network':ti,ab,kw OR 'computer vision':ti,ab,kw) AND 'caries':ti,ab,kw AND [2010-2021]/py	42	12 th April 2021
ArXiv	AND all="Artificial Intelligence" OR "deep learning" OR "machine learning" OR "neural network" OR "computer vision"; AND all=caries	4	12 th April 2021

Table 2 – Modified leading questions of QUADAS-2 for critical appraisal

Domain	Leading Question
PATIENT SELECTION	1. Were data imbalances (if there were any) addressed in the study?
	2. Did the study avoid inappropriate data exclusion? (e.g., difficult to diagnostic or considered as outliers without a specific outlier detection method)
	3. Was a consecutive or random sample of patients (or data) enrolled?
	4. Was the test dataset separate from the training and validation datasets?
	Applicability: Are there concerns that the included data and setting did not match the review question?
INDEX TEST	1. Were the deep learning method results interpreted without knowledge of the results of the reference standard?
	2. If a threshold was used, was it pre-specified?
	3. Was the method described in sufficient detail to reproduce the presented results?
	4. Did the study perform any robustness or sensitivity analysis of their model?
	Applicability: Are there concerns that the method, its conduct, or interpretation differed from the review question?
REFERENCE STANDARD	1. Were the reference standard results interpreted without knowledge of the results of the index test?
	2. Did the study use a gold standard?
	3. If not, was the annotation procedure described in the study and found to minimize bias?
	4. Did the study sufficiently report their limitations, biases, or issues around generalizability?
	Applicability: Are there concerns that the target condition as defined by the reference standard did not match the question?
FLOW AND TIMING	1. Were all data included in the analysis?
	2. Was there an appropriate interval between the index test and reference standard?
	3. Did all data have a reference standard?
	4. Did all data have the exact reference standard?

Table 3 - Summary of findings in the selected classification studies

Author, Year (Ref)	Data Modality	Dataset Size (Train/Valid /Test) *	Inclusion & Exclusion Criteria (if any)	Caries prevalence in the (test) dataset	Labeling Procedure	Pre-Processing	Augmentations	Model Structure	Loss Function	Hardware	Performance measure (for caries detection, mean or single)	Outcome
Leo 2021 [44]	Bitewing radiographs (from ISBI2015 [27])	80 bitewings, 480 tooth segments (300/180)	NA	NA	By 2 experienced dentists, 7 dental structures were annotated: caries, enamel, dentin, pulp, crown, restoration, root canal treatment. Based on the annotations, data were classified into 2 classes: healthy and carious teeth	Manual ROI extraction by cropping, Normalization, Selective median filter, Segmentation, Feature extraction	NA	Hybrid Neural Network (stacked sparse autoencoder + MLP)	NA	NA	Accuracy	0.96
Leo 2020 [45]	Bitewing radiographs (from ISBI2015 [27])	120 bitewings, 418 tooth segments (280/138)	NA	All: 56.70 % Test: 47.83 %	By 2 experienced dentists, 7 dental structures were annotated: caries, enamel, dentin, pulp, crown, restoration, root canal treatment. Based on the study's objective, extracted teeth were classified as healthy and carious teeth	ROI extraction (tooth crown), Grayscale, resize, Selective median filter	NA	Google Net inception v3	NA	NA	Accuracy	0.876
Tripathi 2019 [46]	Bitewing radiographs	800	NA	NA	NA	Extracting features using Local Binary Patterns	NA	MLP	NA	NA	Accuracy	0.954
											MSE	0.001
Sonavane 2021 [47]	Intra-oral photographs [29]	74 (48/12/14)	NA	All: 74.32 % Test: 71.43 %	Data were manually annotated into 2 classes: healthy and carious teeth	NA	Horizontal flip, Zoom	CNN	Binary cross-entropy loss	NA	Accuracy	0.714
Singh 2021 [48]	Intra-oral photographs	1500 (1200/300)	NA	NA	Data were manually annotated into 5 classes based on GV Black's classification + healthy teeth	Median filter for noise removal, Segmentation	NA	CNN-LSTM	Categorical cross-entropy	Intel Core (TM) i3 Lenovo platform with 3 GHz main processor or speed and 8	Accuracy	0.96
											Sensitivity	0.96
											Specificity	0.93
											Precision	0.95
											F1-score	0.95
											G-mean	0.94
AUC	0.96											

										GB memor y		
Wang 2020 [30]	Intra-oral photographs	7200 (6000/1200)	NA	NA	Classified by more than two dentists; Verification of the results was performed using the dual imaging system in both white and fluorescent modes. Data were annotated into 4 classes: Healthy, White Spot Lesion, Smashed, Plaque	NA	NA	T-Net CNN	NA	NA	Accuracy	0.944
											Specificity	0.972
											Sensitivity	0.962
Singh 2020 [49]	Intra-oral photographs	400	NA	100%	Data were manually annotated into 5 classes based on GV Black's classification + healthy teeth	Feature extraction using GIST descriptor, Dimension reduction, Feature selection	NA	Probabilistic neural network	NA	NA	Accuracy	0.89
											Sensitivity	0.92
											Specificity	0.90
Guijarro-Rodríguez 2020 [50]	Intra-oral photographs	2030	NA	NA	NA	Gaussian filter, Median filter, Segmentation with Sobel filter, Feature extraction, PCA	NA	MLP	NA	NA	Accuracy	0.80
Prajapati 2017 [51]	Intra-oral photographs	251 (180/45/26)	NA	All: 31.97% Test: 30.76%	By dentists and oral and maxillofacial radiologists, data were annotated into 3 classes: Caries, periapical infection, periodontitis	NA	NA	VGG16	NA	NA	Accuracy	0.8846-0.875
Holtkamp 2021 [40]	Near-infrared light transillumination images	226 ex vivo 1319 in vivo* 5-fold cross-validation * trained and tested on both datasets.	Inclusion : Posterior teeth	NA	Three expert dentists independently annotated images pixel-wise and the fourth expert dentist reviewed all annotated images. Based on the union of all annotated areas on the image, the reference	Feature extraction	Horizontal and vertical flip, Zoom and Shift	ResNet	Binary cross-entropy loss	NVIDIA Quadro RTX 6000 GPU	Accuracy	0.78
											F1-score	0.73
											sensitivity	0.76
											specificity	0.79
											PPV	0.70
											NPV	0.84
AUC	0.78											

Schwendicke 2020 [18]	Near-infrared light transillumination images	226* 10-fold cross-validation	Inclusion : Posterior teeth	All: 40.71 %	test was constructed. Two experienced dentists marked occlusal and/or proximal caries lesions pixel-wise, independently (following calibration procedure). Data were annotated into 2 classes: teeth with or without carious lesions.	NA	Random resize, Random rotations, Horizontal and vertical flip	Resnet18, Resnext50	NA	NA	AUC (Resnet18-Resnext50)	0.73-0.74
											Accuracy (Resnet18-Resnext50)	0.69-0.68
											Sensitivity (Resnet18-Resnext50)	0.46-0.59
											Specificity (Resnet18-Resnext50)	0.85-0.76
											PPV (Resnet18-Resnext50)	0.71-0.63
											NPV (Resnet18-Resnext50)	0.69-0.73
Salehi 2021 [19]	Optical coherence tomography images	81 (60/21)	Inclusion : human extracted premolar and molar teeth	NA	Data were manually annotated into 3 classes: Non-carious teeth, teeth with enamel caries and teeth with dentin caries	NA	Crop, Flip and Rotation	CNN	NA	NA	Accuracy	0.95
											Sensitivity	0.92
											Specificity	0.95
											PPV	0.92
											NPV	0.95
Salehi 2020 [20]	Optical coherence tomography images	81 (60/21)	Inclusion: Human extracted premolar and molar teeth	NA	Data were manually annotated into 3 classes: Non-carious teeth, teeth with enamel caries and teeth with dentin caries	Normalization	Crop, Flip, and Rotation	CNN	NA	NA	Accuracy	0.8873
											Sensitivity	0.7578
											Specificity	0.9621
											PPV	0.9203
											NPV	0.8731
Yu-Ping 2020 [31]	Optical coherence tomography images	748 cross-sectional images from 63 extracted teeth (599/149)	NA	All: 36.63 %	Micro-CT was used for the reference standard; Data were annotated into 3 classes: Non-carious teeth, teeth with enamel caries and teeth with dentin caries	NA	NA	ResNet-152	NA	NA	Accuracy	0.9521
											Sensitivity	0.9885
											Specificity	0.8983
											PPV	0.9348
											NPV	0.9815
Salehi 2019 [52]	Optical coherence tomography images	51* 5-fold cross-validation	Inclusion : Human permanent teeth	All: 66.67 %	Data were manually annotated into 3 classes: Non-carious teeth, teeth with enamel	NA	NA	CNN	NA	NA	Accuracy	0.91
											Sensitivity	0.9846
											Specificity	0.998
											PPV	0.999

						caries and teeth with dentin caries							
Riyadi 2020 [53]	PA radiographs	237 radiographs, 660 cropped segments* 10-fold cross-validation	Inclusion : Teeth treated by pulp capping approach	NA	NA	Manual ROI extraction (tertiary dentine), Edge cropping	NA	CNN	NA	NA	Accuracy	0.91	
Geetha 2020 [54]	PA radiographs	105* 10-fold cross-validation	NA	All: 46.68 %	By a clinician, data were manually annotated into 2 classes: healthy and carious teeth	Laplacian filter, Gaussian filter. Dilate and erode, Statistical feature extraction	NA	MLP	NA	NA	Precision	0.963	
											Recall	0.962	
											F-measure	0.962	
											MCC	0.924	
											AUC	0.983	
Patil 2019 [55]	PA radiographs [28]	120 (84/36)	NA	NA	By a dental expert	Contrast enhancement, Grey thresholding, Active contour	NA	MNP-ADA (MPCA + MLP)	NA	NA	Accuracy	0.95	
											Sensitivity	1	
											Specificity	0.5	
											Precision	0.95	
											F1-score	0.97	
MCC	0.69												
Sornam 2019 [56]	PA radiographs [28]	120	NA	NA	By a dental examiner, other than healthy .vs carious teeth classification, data were labeled into 5 classes based on caries severity	Segmentation, Gray-level co-occurrence matrix feature extraction	NA	MLP	NA	NA	Accuracy	0.99	
											MSE	0.003	
Patil 2018 [57]	PA radiographs [28]	120 (80/40)* 3-fold cross-validation	NA	NA	By a dental examiner	Contrast enhancement, Gray threshold, Active contour	NA	MPCA + Customized CNN	L1	NA	Accuracy	0.9	
											Sensitivity	0.93	
											Specificity	0.6	
											Precision	0.943	
Patil 2018 [58]	PA radiographs [28]	120 (80/40)* 3-fold cross-validation	NA	NA	By a dental examiner	Contrast enhancement, Gray threshold, Active contour	NA	MPCA + MLP	L1 loss	NA	Accuracy	0.95	
											Sensitivity	0.967	
											Specificity	0.75	
Lee 2018 [59]	PA radiographs	3000 (2400/600)	Exclusion: deciduous teeth	All & Test: 50%	By four calibrated board-certified dentists	Manual ROI extraction by cropping, Vertical flip	Rotation, Shear, Zoom, Width and Height shift, Horizontal flip	GoogleNet Inception V3	NA	NA	Accuracy	0.82	
											Sensitivity	0.81	
											Specificity	0.83	
											MSE	0.003	
Sornam 2018 [60]	PA radiographs [28]	120	NA	NA	By a dental examiner	Size Reduction, cropping, Smoothing, Sharpening and noise removal, GLCM feature extraction	NA	MLP	NA	NA	Accuracy	0.991	
											MSE	0.008	
Lakshmi 2021 [61]	Panoramic radiographs	1900 (1000/900)	NA	NA	NA	Histogram equalization, extracting the GLCM features	NA	AlexNet	NA	NA	Accuracy	0.960	
											Precision	0.785	
											Recall	0.842	
											F1 score	0.982	
Vinayahalingam 2021	Panoramic radiographs	500 segments	Inclusion : Third	All: 50%	By medical records,	Manual ROI extraction	Applied (augmentation)	MobileNet V2	NA	NA	Accuracy	0.87	

[62]		of the third molar region (320/80/100)	molars, minimum age of 16, presence of at least one third molar; Exclusion: Blurred and incomplete panoramics		then images were revalidated by two clinicians. If there was any disagreements, images were excluded.	by cropping, Histogram equalization	ons approaches were not mentioned				Precision	0.87
											F1-score	0.87
											Recall	0.87
											Specificity	0.86
Haghanifar 2020 [63]	Panoramic radiographs	1838 extracted tooth segments (80%/20%)	NA	All: 17.36%	Extracted teeth were classified as healthy and carious teeth by an oral and maxillofacial radiologist. Mild and severe dental caries are also categorized. Annotation was done by an oral and maxillofacial radiologist.	Re-sampling of the smaller class, Vertical edge filter, Gaussian and bilateral filter, Sauvola binarization, ROI extraction, CLAHE, Feature extraction	Horizontal flip, Vertical flip, Rotation, Brightness, Zoom, Width and Height shift	PaXNet (CNN + CapsuleNet)	NA	NA	Accuracy	0.860
											Precision	0.894
											Recall	0.506
											F0.5-score	0.78

* If only two numbers were written in this column usually it means Train/Test splits

Abbreviations: **ADA**, Adaptive Dragonfly Optimizer; **AUC**, Area Under Curve; **CLACHE**, Contrast Limited Adaptive Histogram Equalization; **CNN**, Convolutional Neural Network; **GLCM**, Gray Level Co-Occurrence Matrix; **GPU**, Graphics Processing Unit; **LSTM**, Long Short-Term Memory; **MCC**, Matthews Correlation Coefficient; **MLP**, Multi-Layer Perceptron; **MPCA**, Modified Principal Component Analysis; **MSE**, Mean Squared Error; **NA**, Not Available; **NPV**, Negative Predictive Values; **PA**, Peri-Apical; **PPV**, Positive Predictive Values; **ROI**, Region of Interest

Table 4 - Summary of findings in the selected object detection studies

Author, Year (Ref)	Data Modality	Dataset Size (Train/Valid /Test) *	Inclusion & Exclusion Criteria (if any)	Caries prevalence in the (test) dataset	Labeling Procedure	Pre-Processing	Augmentations	Model Structure	Loss Function	Hardware	Performance Measurements	Outcome
Srivastav 2017 [23]	Bitewing radiographs	3000 (2500/500)	NA	NA	By various dentists following clinical verification, annotations were as loose polygons around caries.	NA	NA	CNN	NA	NA	Recall	0.805
											Precision	0.615
											F1-Score	0.70
Zhang 2020 [64]	Intra-oral photographs	3932 (2507/300 / 1125)	None	All: 33.80 % Test: 35.20 %	By three board-certified dentists, each image annotated by a dentist independently with bounding boxes.	NA	Rotation, Hue, Saturation, and Exposure change	Single-shot detector	Smooth L1 (Localization) cross-entropy loss (Classification)	NA	AUC	0.856
											Image wise sensitivity	0.819
											Box-wise sensitivity	0.646
Javid 2020 [65]	Intra-oral photographs	90 (65/25)	NA	NA	By supervision of a dentist	Sharpening filter	NA	Mask RCNN (Based on ResNet50)	NA	NA	Precision	0.95
											Recall	0.96
Yu 2020 [66]	Intra-oral photographs	1368 (1052/316)	First permanent molar present	NA	By board-certified dentists	NA	Horizontal and vertical flip	ResNet-FPN, and two parallel task-specific subnetworks for region regression and region classification	Weighted Cross-entropy	Nvidia Titan XP GPU with 12 GB memory	Accuracy	0.952
											Sensitivity	0.898
											Specificity	0.961
Zhang 2020 [67]	Intra-oral photographs	1000 (800/200)	Inclusion: Anterior teeth	53.70 %	By dental practitioners, sample, annotation classes: normal teeth, 6 classes of caries based on ICDAS classification system, others (crown, bracket, etc)	NA	NA	Faster R-CNN (Based on ResNet50)	NA	Tesla K80 GPU	mAP	0.473
											AP50	0.665
											AP75	0.543
											AR	0.716
Choi 2016 [68]	PA radiographs	475 (380/95) * 5-fold cross-validation	NA	NA	Proximal caries was confirmed in periapical images by experts	Horizontal alignment, Cropping above the tooth top line and one-third of the image height from the bottom	Rotation, Left-right direction flip	CNN	NA	NA	F1-max	0.74
											FPs	0.88

* If only two numbers were written in this column usually it means Train/Test splits

Abbreviations: **AUC**, Area Under Curve; **CNN**, Convolutional Neural Network; **ICDAS**, International Caries Detection and Assessment System; **FP**, False Positive; **FPN**, Feature Pyramid Networks; **GPU**, Graphics

Processing Unit; **L1 loss**, Least Absolute Deviations; **NA**, Not Available; **PA**, Peri-Apical; **R-CNN**, Region Based Convolutional Neural Networks; **ROI**, Region of Interest

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Table 5 - Summary of findings in the selected segmentation studies

Author, Year (Ref)	Data Modality	Dataset Size (Train/Valid /Test) *	Inclusion & Exclusion Criteria (if any)	Caries prevalence in the (test) dataset	Labeling Procedure	Pre-Processing	Augmentations	Model Structure	Loss Function	Hardware	Performance Measurements	Outcome
Cantu 2020 [69]	Bitewing radiographs	3686 (3293/252/141)	Inclusion: Permanent teeth, crowns of min. one arch visible	NA	Three expert dentists independently annotated images pixel-wise and the fourth expert dentist reviewed all annotated images. Based on the union of all annotated areas on the image, the reference test was constructed.	NA	Flip, Center crop, Translation, Rotations, Gaussian-blur, Sharpening, Contrast, and Brightness	U-Net	Binary focal type	GeForce GTX 1080 Ti GPU using CUDA 10	Accuracy	0.80
											Sensitivity	0.75
											Specificity	0.80
											F1-score	0.73
											MCC	0.57
Kumar 2019 [70]	Bitewing radiographs	6000 (5000/1000)	NA	NA	By various dentists following clinical verification	NA	NA	U-Net	Cross-entropy loss	NA	Recall	0.73
											Precision	0.53
											F1-score	0.6142
Yun 2018 [71]	Bitewing radiographs [27])	120 (40/40/40)	NA	NA	By 2 experienced dentists, 7 dental structures were annotated: caries, enamel, dentin, pulp, crown, restoration, root canal treatment	NA	Rotation, Horizontal and vertical flip, Translation Transformation, Change of image gray value	Conditional generative adversarial network + U-Net	Arg min(G)max(D) [cGAN(GD) + Lambda*L1 (G)]	NA	Precision	0.546 – 0.418
											True positives	0.784 – 0.768
											True negatives	0.956 – 0.973
											Dice similarity	0.697 – 0.584
Ronneberger 2015 [72]	Bitewing radiographs [27])	80 (40/40)	NA	NA	By 2 experienced dentists, 7 dental structures were annotated: caries, enamel, dentin, pulp, crown, restoration, root canal treatment	NA	Translations Rotations, Deformation gray value variations, (Increased to 20,000 training images)	U-net	Softmax loss	Nvidia Titan GPU	Precision	0.453
											True positives	0.576
											True negatives	0.983
											Dice Similarity	0.564
Ezhov 2021 [73]	CBCT	4398	NA	NA	By dentists and oral and maxillofacial radiologists, then images	ROI localization, tooth localization and numeration	NA	U-net	NA	NA	Sensitivity	0.7285
											Specificity	0.9953

					were revalidated by the lead oral and maxillofacial radiologist, Labels were background, caries, metallic artifact and non-contrast filling.							
Moutselos 2019 [74]	Intra-oral photographs	88 (79/9)	Inclusion: Posterior permanent molar Exclusion: Teeth with hypoplastic, hypomineralised areas and with sealants	All: 100%	Occlusal surfaces were annotated pixel-wise into 7 classes: free of caries and 6 classes of caries based on the ICDAS II classification system	Superpixel segmentation	Horizontal and vertical flip, Rotations, Shear and piecewise affine	Mask R-CNN	NA	NA	micro F-measure	0.596, 0.625, 0.684
Casalegno 2019 [75]	Near-infrared light transillumination images	217 (185/32)	Exclusion: Images with dental restorations	NA	The overall agreement of 2 segmentation maps through manual labeling by experts; 5 dental structures were annotated: background, enamel, dentin, proximal caries, and occlusal caries	NA	Horizontal and vertical flip, Zoom, Rotation, Translation, Random contrast and brightness	U-net + VGG16	NA	NA	AUC	0.836 – 0.856
											mIOU	0.727
Khan 2021 [76]	PA radiographs	206 (132/44/30) * 4-fold cross-validation	Inclusion: Images used parallel technique, Exclusion: Images if at least 2 of 3 examiners did not agree	76.70% (In train + valid set)	By 3 experienced dentists. One examiner annotated data. The remaining 2 examiners evaluated the labels. Data were annotated into 3 classes: Caries, alveolar bone recession, and interradicular radiolucencies	NA	Magnification, Horizontal and vertical flip, Translation, Rotation	U-Net + Densenet 121	NA	NA	mIoU	0.383
											Dice coefficient (all)	0.434
											mIoU	0.194
											Dice coefficient (caries)	0.239
Jung 2020 [77]	PA radiographs (from Kaggle)	86 (70%/30%)	NA	NA	By oral and maxillofacial radiologist	NA	NA	DeepLabv3+ (An autoencoder based on	NA	GeForce RTX 2070 GPUs, 32.0	Accuracy	0.9847
											Loss	0.0378

					, Data were manually annotated into 6 classes: apical problem abrasion, caries, impaction, periodontal problem, sound tissue.			ResNet18)		GB RAM, AMD Ryzen 7 3800X 8-Core Processor or (3893 Mhz, 8 core, 16 logic processor)	Segmentation accuracy	0.20
											Specificity	0.9953
Rad 2018 [78]	PA radiographs [28]	NA * They extracted 155 teeth using segmentation	Inclusion: Age from 25 to 35	NA	By a dental examiner	NA	NA	MLP	NA	NA	Accuracy on image	0.9085
											Accuracy on segments	0.98

* If only two numbers were written in this column usually it means Train/Test splits

Abbreviations: **CBCT**, Cone-beam computed tomography; **ICDAS**, International Caries Detection and Assessment System; **GPU**, Graphics Processing Unit; **NA**, Not Available; **mIoU**, mean Intersection over Union; **MCC**, Matthews Correlation Coefficient; **MLP**, Multi-Layer Perceptron; **PA**, Peri-Apical; **R-CNN**, Region Based Convolutional Neural Networks; **ROI**, Region of Interest