

Critical Reviews in Oral Biology & Medicine

Application of Artificial Intelligence in Dentistry

Journal of Dental Research 2021, Vol. 100(3) 232-244 © International & American Associations for Dental Research 2020 Article reuse guidelines: sagepub.com/iournals-permissions DOI: 10.1177/0022034520969115 journals.sagepub.com/home/jdr

T. Shan¹, F.R. Tay², and L. Gu¹

Abstract

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Artificial intelligence (AI) is a technology that utilizes machines to mimic intelligent human behavior. To appreciate human-technology interaction in the clinical setting, augmented intelligence has been proposed as a cognitive extension of AI in health care, emphasizing its assistive and supplementary role to medical professionals. While truly autonomous medical robotic systems are still beyond reach, the virtual component of Al, known as software-type algorithms, is the main component used in dentistry. Because of their powerful capabilities in data analysis, these virtual algorithms are expected to improve the accuracy and efficacy of dental diagnosis, provide visualized anatomic guidance for treatment, simulate and evaluate prospective results, and project the occurrence and prognosis of oral diseases. Potential obstacles in contemporary algorithms that prevent routine implementation of Al include the lack of data curation, sharing, and readability; the inability to illustrate the inner decision-making process; the insufficient power of classical computing; and the neglect of ethical principles in the design of Al frameworks. It is necessary to maintain a proactive attitude toward Al to ensure its affirmative development and promote human-technology rapport to revolutionize dental practice. The present review outlines the progress and potential dental applications of Al in medical-aided diagnosis, treatment, and disease prediction and discusses their data limitations, interpretability, computing power, and ethical considerations, as well as their impact on dentists, with the objective of creating a backdrop for future research in this rapidly expanding arena.

Keywords: big data, clinical decision making, dental, informatics, machine learning, neural networks

Advancement in digitized technology for clinical examination has rendered health care data collection from dental patients less complex and cumbersome. Personalized dentistry requires taking massive data sets into consideration for each patient. Conventional statistical analytics relies on specific assumptions and handcrafted markers, making it impractical in dealing with such high-volume data (Ayala Solares et al. 2020). Artificial intelligence (AI) intends to reproduce the cognitive process of humans and can achieve the same outcome as medical professionals within a much shorter time frame. It excels in extracting information from historical data and benefits physicians by automating time-consuming tasks. Although the current development of AI is preliminary and medical tasks that contemporary AI can complete can almost be performed by humans, the emergence of AI in dentistry heralds an era of disruptive technology with the potential to reengineer the landscape in which dental clinical care is practiced. The present review outlines the progress and potential dental applications of AI and discusses their challenges as well as the potential impact on dentists. A glossary of AI terms used in the review is included in the Appendix.

Artificial Intelligence

AI is a new branch of applied computer science that endows machines with the ability to mimic intelligent human behavior. Two types of AI are available for general health care delivery: physical and virtual. Physical applications are represented by sophisticated robots or automated robotic arms (Wang et al. 2014). Virtual components are software-type algorithms that support clinical decision making.

Attempts at implementing AI were initially based on the assumption that human intelligence can be fully digitized and integrated into machines. As early as the 1970s, commentators predicted that AI would bring careers in medicine to an end (Maxmen 1976). However, the development of strong AI is limited by the finite storage and data-processing capacity of bit-based computers as well as the complexity of human cognition. Persuaded by these intractable barriers, weak (narrow) AI was developed to partially solve problems in distinct application areas by learning specific perception and thinking mechanisms instead of comprehensive intelligence (Park and Park 2018). Augmented intelligence, introduced by the American

¹Department of Operative Dentistry and Endodontics, Guanghua School of Stomatology, Hospital of Stomatology, Sun Yat-sen University, Guangdong Provincial Key Laboratory of Stomatology, Guangzhou, China ²The Dental College of Georgia, Augusta University, Augusta, GA, USA

A supplemental appendix to this article is available online.

Corresponding Authors:

F.R. Tay, The Dental College of Georgia, Augusta University, 1430 John Wesley Gilbert Drive, Augusta, GA 30912-0004, USA. Email: ftay@augusta.edu

L. Gu, Department of Operative Dentistry and Endodontics, Guanghua School of Stomatology, Hospital of Stomatology, Sun Yat-sen University, Guangdong Provincial Key Laboratory of Stomatology, Lingyuanxi Road, Guangzhou, Guangdong, 510275, China.

Email: gulisha@mail.sysu.edu.cn

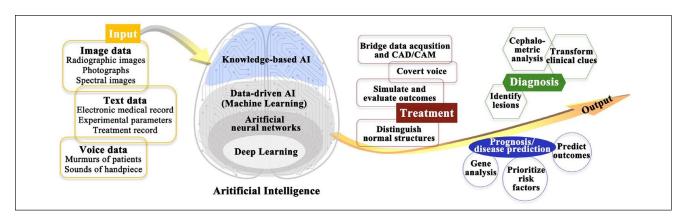


Figure 1. Overview of the hierarchy and major dental applications of artificial intelligence (Al). CAD/CAM, computer-aided design/computer-aided manufacturing.

Medical Association (2018), is an alternative conceptualization of narrow AI in health care, emphasizing its assistive and supplementary role to medical professionals (Crigger and Khoury 2019). Admittedly, human intelligence is still superior in complex cognitive tasks. Nonhominid intelligent systems guided by augmented intelligence are designed to cooperate with humans, augmenting human intelligence rather than replacing human expertise.

In dentistry, the applications of AI are mostly virtual, employing AI algorithms to distinguish between lesions and normal structures, prioritize risk factors, and simulate and evaluate prospective results. According to the Barcelona Declaration for the Proper Development and Usage of Artificial Intelligence in Europe (Steels and Lopez de Mantaras 2017), virtual AI methodologies are fundamentally divided into knowledge-based and data-driven AI (Fig. 1). Knowledgebased AI attempts to model human knowledge and is built in a top-down fashion from the self-reported concepts and knowledge that humans use to solve problems. However, knowledge acquisition and formalization are 2 major bottlenecks, which consume development time and require significant initial effort (Montani and Striani 2019). Conversely, data-driven AI, commonly known as machine learning (ML), commences in a bottom-up approach by training mathematical models with data derived from human activities. Because of the large amount of dental data available in electronic form, data-driven AI receives a lot of attention in dentistry.

Data-driven AI or ML may be divided as supervised, unsupervised, and semisupervised learning (Handelman et al. 2018). On the supervised platform, algorithms employ manually labeled training data sets to learn the correlations between data instances and labels, yielding the desired and known outcomes (Krittanawong et al. 2017). Support vector machines (SVMs), decision tree (DT), random forest (RF), and artificial neural networks (ANNs) are all ML algorithms that can be used in supervised learning. SVMs set up an imaginary high-dimensional space, place samples according to their features, and separate them by a hyperplane, resulting in data classification (Amasya et al. 2020). DT is a hierarchical data structure (Kok et al. 2019). Each node is a specific attribute, and branches are followed until a leaf node is reached. Classification

is completed once a leaf node is reached. RF is an extension of DT, in which each DT is independently trained and subsequently combined with others (Krittanawong et al. 2017). ANNs are highly interconnected models (Fig. 2) inspired by vertebrate nervous systems. Every neuron receives signals from neurons in the previous layer, performs mathematical transformation, and transmits signals to neurons in the next layer as output.

In unsupervised learning, algorithms are provided with unlabeled data, and they involve recognizing hidden data patterns that investigators may not have conceived, yielding unknown results. Principal component analysis and k-means clustering are common methods used in unsupervised learning. They employ discrete or continuous data as input to identify latent regularities (i.e., k-means clustering) or lower dimensional representations (i.e., principal component analysis). Deep neural networks, commonly known as deep learning (DL), is a subset of ML; they can also be operated in unsupervised scenarios. The term "deep" refers to multiple neural layers between the input and output layers (Park and Park 2018). Convolutional neural networks (CNNs; Fig. 2A) are the most widely used DL architecture in dentistry, employing a convolutional process to learn features contained within data (Hiraiwa et al. 2019). Although it is frequently mentioned that supervised learning is limited by the lack of annotated data, unsupervised learning has also been criticized for failing to identify the initial pattern. Semisupervised learning is an amalgamation of supervised and unsupervised learning that analyzes a collection of data while augmenting the pattern recognition abilities with a small amount of labeled data (Handelman et al. 2018). All these techniques offer the promise of more powerful endeavors to enhance human capability in dealing with massive and complex data.

Applications of AI in Dentistry

Medical-Aided Diagnosis

Diagnostic logic for a certain disease is based on a clinician's analysis of symptoms, diagnostic test results, and other factors, which are vulnerable to the clinician's imperfect memory and

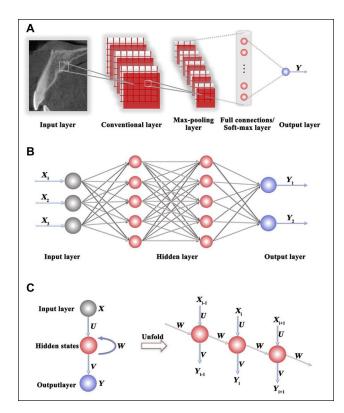


Figure 2. Topology of 3 artificial neural networks: (**A**) convolutional, (**B**) multilayer perceptron, and (**C**) recurrent. X_p , input data set; Y_p , output data set. U, V, and W are the weights.

cognitive bias. When "educated" with hundreds of thousands of cases, AI surpasses the clinical experience of even the most proficient specialist (Bouletreau et al. 2019). This infers that more accurate and efficient diagnosis will be achieved with the integration of AI into existing dental clinical workflow (Table 1).

Oral and Maxillofacial Surgery. ML algorithms, including SVM, ANN, RF, and k-nearest neighbors, have been experimentally investigated for their ability to identify cysts, benign tumors, oral cancer, and lymph node metastasis. Fed with handcrafted engineered features in cone beam computed tomography (CBCT), SVM was 94% accurate in differentiating periapical cysts from keratocystic odontogenic tumors (Yilmaz et al. 2017). A risk stratification model based on brush biopsy and cytology was also constructed with ML algorithms such as SVM and RF. Utilizing CNN to score the malignancy of cytology images derived from a telemedicine platform, this model showed high sensitivity in detecting oral malignant (93%) and high-grade potential malignant (73%) lesions (Sunny et al. 2019). Despite these excellent results, contemporary AI models for oral and maxillofacial surgery diagnosis focus on only 1 type of data, such as radiographic results or cytopathologic images. For highly accurate diagnosis, models that integrate more medical information about the patient are required.

Cariology and Endodontics. With the ability to perform automated lesion segmentation, DL with CNN has become the predominant AI component used in cariology and endodontic diagnostics. The segmentation process divides radiographs or images into multiple nonoverlapping regions using sets of rules, such as similar pixels or intrinsic features, to convert them into a meaningful form that can be conveniently analyzed (Anwar et al. 2018). Based on an encoder-decoder architecture (U-Net), DL segmented CBCT voxels into "lesion," "tooth structure," "bone," "restorative materials," and "background," achieving results comparable to those of clinicians in diagnosing periapical lesions (Setzer et al. 2020). Focusing on the binary presence or absence of lesions, DL identified proximal carious lesions from near-infrared transillumination images with an area under the receiver operating characteristic curve of 0.856 (Casalegno et al. 2019). Volumetric measurement in CBCT, following DL-based segmentation, was reported to be comparable to the results obtained from manual segmentation of periapical lesions (Orhan et al. 2020). However, this research failed to report outcomes such as the volume deviation of lesions and the Intersection over Union metric. This oversight challenges the reliability of the results.

Periodontics. Periodontal disease is a complex inflammatory disease contributed by multiple causal factors simultaneously and interactively. Focusing on periapical radiographs, CNN achieved 81.0% and 76.7% accuracy in diagnosing periodontally compromised premolars and molars (Lee et al. 2018). However, due to the hysteresis of imaging characteristics and the visual field of periapical radiographs, this technique cannot distinguish incipient lesions or make a final diagnosis of periodontal disease. DT and SVM performed well in classifying healthy periodontium, gingivitis, chronic periodontitis, and aggressive periodontitis by integrating a patient's medical history, clinical information, and radiographs. Although immunologic responses and microbial composition in relation to the etiology of periodontitis have not been fully elucidated, AIbased classifiers—such as a multilayer perceptron neural network (a class of feedforward ANN; Fig. 2B) focusing on leukocytes, interleukins, and IgG antibody titers, as well as an SVM focusing on the relative bacterial load—performed satisfactorily in distinguishing aggressive from chronic periodontitis (Feres et al. 2018). In addition to promoting our understanding of periodontitis, AI serves as a bridge to incorporate conventional indicators and immunologic and microbiological parameters into periodontal diagnosis.

Temporomandibular Joint Disorder. Clinical clues from a patient's complaint and history are important for diagnosing temporomandibular joint disorders (TMDs). Natural language processing is a technology that transforms natural human language into structured computer language. A natural language processing—based model was successful in differentiating TMD-mimicking conditions from genuine TMDs, according to the frequency of word usage in the patient's chief complaint and mouth-opening size (Nam et al. 2018). ANNs based on

 $\textbf{Table I.} \ \ \mathsf{Dental} \ \ \mathsf{Applications} \ \ \mathsf{of} \ \ \mathsf{Al} \ \ \mathsf{in} \ \ \mathsf{Diagnosis}.$

Diagnosed Disease	Clinical or Experimental Data	– Sample Size, Training Testing	g: Validation Method	Al Methods	Accuracy	Sensitivity	Specificity	AUC
	·	resting	validation Flethod	Ai Fiethods	Accuracy	Sensitivity	эреспісіту	700
Oral and n Oral cancer	naxillofacial surgery Fourier-transform infrared	34: —	k-fold cross-	PCA and LDA	95%	100%	89%	
oral cancer	spectra of salivary exosomes (Zlotogorski-Hurvitz et al.	J.,	validation	SVM	89%	100%	07/0	
	2019*) Hyperspectral images (Jeyaraj	100: —	7-fold cross-	CNN	91.4%	94%	91%	
	and Samuel Nadar 2019*)	100.	validation					
	X-ray images (Al-Ma'aitah and AlZubi 2018*)		Mean square error rate	GSOESNN	99.2%	>95%	>95%	
Oral squamous cell carcinoma	Laser endomicroscopy images (Aubreville et al. 2017*)	7,894 augmented 2 times: —	Leave-I-patient-out cross-validation	CNN LBP-based RF	88.3%	86.6% 84.7%	90%	0.955
carcinoma	(rtableville et al. 2017)	unics.	ci oss-vandacion	GLCM-based RF	81.4% 73.1%	77.5%	78.2% 69.5%	0.895 0.807
	Oral tissue histopathologic slides (Das et al. 2018*)	80: 20		CNN and RF	96.9% (keratin			
	Brush cytology specimens (McRae et al. 2020*)			k-nearest neighbors	96.9%			
	Cytology images (Sunny et al. 2019)	11,981: —		Pretrained ANN (inception V3)		73% (high potential malignant), 93% (malignant)		
ervical lymph node metastasis	CT images (Ariji et al. 2019*)	441 augmented to 21,362: —	5-fold cross- validation	Pretrained CNN (Alex Net)	78.20%	75.4%	81.0%	0.8
meloblastoma, keratocystic	Panoramic images (Poedjiastoeti and Suebnukarn 2018*)	400 augmented to 800: 100		Pretrained CNN (VGG-16)	83%	81.8%	83.3%	0.88
odontogenic tumor Dentigerous cysts,	Panoramic radiography (Lee	augmented to 200 648 augmented 100 times: 228		Pretrained CNN		96.1%	77.1%	0.914
keratocystic odontogenic tumor	et al. 2020*) CBCT images (Lee et al. 2020*)	592 augmented 100 times: 197		(GoogLeNet) Pretrained CNN		88.2%	77.0%	0.847
and periapical cysts eriapical cyst and	CBCT images (Yilmaz et al.	times: 197	10-fold cross-	(GoogLeNet) SVM	94%			
keratocystic	2017)		validation	ANN	92%			
odontogenic tumor				Naïve Baves	Inferior to ANN			
	y and endodontics					0.107		
Dental caries	Periapical radiographic images of premolar and molar (Lee et al. 2018*)	2,400: 600		Pretrained CNN (GoogLeNet)	82%	81%	83%	0.845
	Near-infrared transillumination images (Casalegno et al. 2019)	185: 32	Monte Carlo cross- validation	CNN				0.836 (occlusa 0.856 (proxim
	Near-infrared-light transillumination images	226 with augmentation: —	10-fold cross- validation	Pretrained CNN (Resnet18)	69%	85%	46%	0.73
	(Schwendicke et al. 2020*)	aug.noneacion.	vandacion	Pretrained CNN (Resnext50)	68%	76%	59%	0.74
oot fracture	Panoramic images (Fukuda et al. 2019*)	240: 60		Pretrained CNN (DetectNet)	93%			
	Periapical radiographs (Johari	180: 60		PNN	70%	93.3%	63.6%	
	et al. 2017*)	120: 120		PNN	65%	81.7%	61.3%	
		60: 180		PNN	63.9%	88.9%	59.3%	
	CBCT images (Johari et al.	180: 60		PNN	96.7%	93.3%	100%	
	2017*)	120: 120		PNN	95.0%	90%	100%	
eriapical pathosis	Panoramic radiographs (Ekert et al. 2019*)	60: 180 2,001 teeth from 85 panoramic	10-fold repetition	PNN CNN	93.3%	86.7% 65%	100% 87%	0.84
	CBCT images (Setzer et al. 2020)	images: — 20: —	5-fold cross- validation	Pretrained CNN (U-net)	93%	93%	88%	
	CBCT images (Orhan et al. 2020)	3,900: 153		CNN		0.89 estimated recall		
racked dental root, caries, hypodontia and bone resorption	X-ray images (Ngan et al. 2016*)			Fuzzy aggregation operators	93.0%			
	eriodontics							
ingivitis and periodontitis	Risk factors, clinical periodontal parameters (Ozden et al.	100: 50		SVM DT	98% precision 98% precision			
ania danaallo	2015*)	1.044		BPNN	46% precision			0.007
eriodontally compromised teeth	Periapical radiographs (Lee et al. 2018)	1,044 augmented to 104,400: 348		CNN	82.8% (premolar) 73.4% (molar)			0.826 (premo 0.734 (mo

(continued)

Table I. (continued)

	Field	6 I 6 T						
Diagnosed Disease	Clinical or Experimental Data	— Sample Size, Training Testing	Validation Method	Al Methods	Accuracy	Sensitivity	Specificity	AUC
Aggressive and chronic periodontitis	Clinical and immunologic data sets (Papantonopoulos et al. 2014*)		10-fold cross- validation	MLP neural network	90% to 98%			
	Microbial profiles (Feres et al. 2018)	2,740: 1,175		SVM		86%	79%	0.83
	TMDs							
TMD-mimicking conditions	Mouth opening size and the frequency of word usage in chief complaint (Nam et al. 2018)	29 (TMD-mimicking cases), 290 (TMD): —	10-fold cross- validation	Text-mining method	96.6%	69.0%	99.3%	
Bone changes and disc displacement	Magnetic resonance images (Iwasaki 2015*)	590: —	Resubstitution validation	BBN with path condition	99.8%			
			10-fold cross- validation	BBN with path condition	99.5%			
Deformation of condyles with osteoarthritis	CBCT images (Shoukri et al. 2019)	259: 34		ANN	97.1%			
Or	thodontics							
Cervical vertebral maturation	Measurement data from lateral	300: —	5-fold cross-	ANN	93.0% (CVS I)			
	cephalometric radiographs		validation	k-nearest neighbors	78.7% (CVS I)			
	(Kok et al. 2019)			Naïve Bayes	92.1% (CVS I)			
				DT	97.1% (CVS I)			
				SVM	84.8% (CVS I)			
				RF	91.8% (CVS I)			
	Measurement data from lateral	498: 149		ANN	0.926 (κ value)			
	cephalometric radiographs			SVM	0.874 (κ value)			
	(Amasya et al. 2020)			RF	0.908 (κ value)			
				DT	0.921 (κ value)			
Automated cephalometric analysis	Lateral cephalometric radiographs (Kunz et al. 2020*)	96.6% of 1,792 samples after augmentation: —		CNN	High correlation with humans $(r > 0.864)$			
	CBCT images (Gupta et al.	—: 30		A knowledge-based	2.01 ± 1.23 mm			
	2015)			algorithm	(mean error)			
	Others							
Oral lichen planus	Clinical data, normalized	81: —		SVM	0.76	0.77	0.75	0.87
	expression of interleukin-12			RF	0.86	0.90	0.80	0.92
	receptor β2 and tumor necrosis factor receptor			ANN	0.78	0.79	0.77	0.83
	superfamily member 8 (Jeon			LDA	0.82	0.87	0.75	0.88
	et al. 2015)			Naïve Bayes	0.80	0.77	0.85	0.87
Sjögren's syndrome	Ultrasonography images of parotid gland (Kise et al. 2020)	160 augmented to 8,000: 40	5-fold cross- validation	Pretrained CNN (VGG16)	89.5%	90.0%	89.0%	0.948
	Ultrasonography images of submandibular gland (Kise et al. 2020)	160 augmented to 8,000: 40	5-fold cross- validation	Pretrained CNN (VGG16)	84.0%	81.0%	87.0%	0.894

*References marked with an asterisk are included in the Appendix.

Al, artificial intelligence; ANN, artificial neural network; AÜC, area under the receiver operating characteristic curve; BBN, Bayesian belief network; BPNN, back propagation neural network; CBCT, cone beam computed tomography; CNN, convolutional neural network; CT, computed tomography; CVS, cervical vertebrae stage; DT, decision tree; GLCM, gray-level co-occurrence matrix; GSOESNN, gravitational search optimized echo state neural network; LBP, local binary pattern; LDA, linear discriminant analysis; MLP, multilayer perceptron; PCA, principal component analysis; PNN, probabilistic neural network; RF, random forest; SVM, support vector machine; TMD, temporomandibular disorder.

CBCT revealed a 91.2% degree of conformity as compared with clinician consensus in classifying condylar morphology (Shoukri et al. 2019). By integrating patients' chief complaints, clinical and biochemical indicators, and objective radiomic features into training data sets and collecting larger samples of these data sets, a computer-assisted diagnosis system is warranted to improve TMD diagnostic accuracy.

Orthodontics. Given the availability of clearly definable cephalometric landmarks, a knowledge-based algorithm has been developed to detect them. This algorithm revealed only

a 2.01-mm mean error for all 20 landmarks tested (Gupta et al. 2015). Diagnostic modeling based on lateral cephalometric radiographs was also constructed with ML algorithms, such as ANN, SVM, RF, and DT, to evaluate cervical vertebral maturation. Among the ML algorithms tested, ANN achieved the best results in classifying cervical vertebral maturation stage and DT for vertebral body shape classification (Amasya et al. 2020). In the future, hybrid approaches that combine knowledge-based algorithms and ML should be able to achieve more accurate and precise automated cephalometric analysis.

Others. Focusing on the expression of inflammatory cytokines genes, ANN, SVM, and RF were all capable of distinguishing oral lichen planus from other white lesions of the oral mucosa (Jeon et al. 2015). When used for recognizing steatosis (i.e., abnormal retention of lipids) of the salivary gland parenchyma in ultrasonography images, ANN was superior to inexperienced radiologists in differentiating patients with true Sjögren's syndrome from those with xerostomia (Kise et al. 2020).

Treatment

The versatility of AI in distinguishing anatomic structures and simulating prospective results renders it a valuable adjunct in providing anatomic guidance, planning treatment, and evaluating outcomes of dental treatment (Table 2). Although truly autonomous medical robotic systems are still beyond reach, these virtual AI applications have immense potential in simplifying clinical operations. They are the keystone in the development of precision dentistry.

Oral and Maxillofacial Surgery. To date, much of the clinical potential of AI relies on its ability to trace critical anatomic structures by analyzing patients' radiographic data or diffuse reflectance spectra produced by a laser scalpel. Examples of these anatomic structures include interdigitated tongue muscles, the mandibular canal, nerves, and the parotid gland. Although the difference between AI-based segmentation and expert measurement or true anatomic position is only in the millimeter range (Gerlach et al. 2014), it is still substantial for fine structures, such as neurovascular canals, and may lead to severe surgical complications. In terms of postoperative rehabilitation, an ML-based voice conversion technique has been used to transform nonaudible murmurs of source speakers into the normal speech of target speakers. By learning a set of bases that is representative of the entire set of voice and target dictionary, this technique adapted well to a limited amount of training data and acquired satisfactory short-time objective intelligibility scores (Fu et al. 2017). This is instrumental in improving the speech intelligibility of oral surgical patients.

Prosthodontics. Computer-aided design/computer-aided manufacturing (CAD/CAM) is gaining acceptance in prosthetic dentistry. Integrating AI with CAD/CAM improves its chairside application (Raith et al. 2017). ANNs based on panoramic radiographs, periapical radiographs, micro-computed tomography images, and 3-dimensional scanning of dental surfaces have been explored for tooth segmentation and classification. With >90% accuracy, such automatic classification is instrumental for bridging the gap between data acquisition and manufacturing in CAD/CAM technology. For abutment tooth preparation, an ablation system combining robotics and picosecond laser has been developed. In vitro results indicated that the average linear ablation errors of this system were only 0.06 mm for wax resin and 0.05 mm for dentin (Wang et al. 2014). However, the long ablation time for dentin (3.5 h)

precluded its clinical application. An automatic robotic system is still an underdeveloped area of research in fixed prosthodontics. In removable prosthodontics, a clinical decision support model that applies ontology and case-based reasoning was capable of recommending a design of individualized removable partial dentures (Chen et al. 2016). Unfortunately, this model makes recommendations based on the most similar case in the database. Because clinical scenarios are ever changing, a skeptical attitude must be maintained with respect to its output.

Orthodontics. Treatment need assessment and outcome scoring are the 2 potential applications of AI in orthodontics. The Bayesian network, based on orthodontics-related data, attained a high level of agreement with orthodontists in diagnosis of orthodontic treatment needs (Thanathornwong 2018). SVM enabled analysis of the fixation time of lay persons viewing patients and utilized observer attention as an indicator of orthodontic treatment need (Wang et al. 2016). CNNs achieved attractiveness scoring and apparent age estimation by characterizing specific facial traits (Patcas et al. 2019). Whereas professional appraisal of attractiveness relies on the perception of "ideal facial features," AI evaluation is a quantifiable representation of social attractiveness. Introducing AI into orthodontics is helpful in objectively and reproducibly interpreting facial appearance. Nevertheless, AI will never replace a patient's own perception and expectations.

Others. Fed with data extracted from experiments, a prediction model integrating ANN obtained 0.81 coefficient of determination in predicting the viability of dental pulp stem cells under different bacterial lipopolysaccharide concentrations and treatment times (Bindal et al. 2017). This may be a future tool for evaluating regenerative dentistry protocols in a simulated inflammatory microenvironment.

Disease Prediction and Prognosis

Recent advances in AI provide valuable frameworks for integrating all clinical symptoms—related attributes, including patient history, demographics, lifestyle, and clinical and gene factors. Classifier and predictive AI models help prioritize risk factors and predict long-term outcomes of dental diseases by exploring associations between diseases and patient data (Table 3).

Oral and Maxillofacial Surgery. In dental extraction scenarios, an ANN that established associations among personal, anatomic, and surgical factors was 98% accurate in predicting facial swelling after extraction of impacted mandibular third molars (Zhang et al. 2018). In oral cancer prognosis, ML-based predictive models that integrate all patient performance metrics have been constructed. Decision forest, SVM, and gradient boosting algorithms all reached ~0.70 area under the receiver operating characteristic curve in predicting occult nodal metastasis (Bur et al. 2019). Boosted DT, SVM, decision forest, and

Table 2. Dental Applications of AI in Treatment of Diseases.

Fie	lds		Consola Sina Tosiais a			
Aim	Function	— Data	Sample Size, Training: Testing	Assessment	Al Method	Results
Oral and maxi	llofacial surgery					
Provide anatomic guidance	Distinguish interdigitated tongue muscles (Ye et al. 2015*)	Limited diffusion MRI			Bayesian approach	
	Segmentate the mandibular canal (Gerlach et al. 2014)	CBCT images	13: —	Compared with histologic sections of the same region	ASM	The difference ranged from -3.45 to 3.27 mm
					AAM	The difference ranged from -4.44 to 4.44 mm
	Segmentate parotid gland (Yang et al. 2014*)	MRI		Compared with the physicians' manual contours	SVM combined with Atlas registration	The average volume differences were 7.98% (left) and 8.12% (right)
	Classifying different tissue types (Engelhardt et al. 2014*)	Diffuse reflected spectra of laser scalpel	1,000: 1,000	Calculated misclassification rate	ANN	0.26 misclassification rate 0.98 misclassification rate
					LDA QDA PDA	0.34 misclassification rate 0.27 misclassification rate 0.02 misclassification rate
					Random forests Classification and regression trees	0.34 misclassification rate 0.50 misclassification rate
Improve speech intelligibility of patients	Voice conversion (Fu et al. 2017)	Sentences uttered by oral surgical patients and target speakers	70: 40	Compared with conventional exemplar-based NMF method	Joint dictionary learning based non-NMF algorithm	Higher short-time objectiv intelligibility scores in th proposed algorithm
Prostho	odontics					
Classify tooth	Classify dental cusps (Raith et al. 2017)	3D surface scans of dental casts	119: 10		ANN	93.3% accuracy (cusp distance method), 93.5% accuracy (range image method)
	Classify and number tooth (Tuzoff et al. 2019*)	Panoramic radiographs	1,352: 222	Compared with experts	Pretrained CNN (VGG-16)	0.9941 sensitivity and 0.994 precision
	Detect and classify tooth (Zhang et al. 2018*)	Periapical radiographs	700: 200		Pretrained CNN (VGG-16)	95.8% accuracy and 96.1% recall in total
	Classify enamel, dentin, and pulp layer (Wang et al. 2017*)	Micro-computed tomography data sets			k-means++	0.83, 0.85, and 0.77 accura in classifying enamel, dentin, and pulp
Optimize conditions	Correlate resin composite hardness with light conditions (Deniz Arısu et al. 2018*)	Processing parameters and corresponding results			ANN	The mean square error value of the model was 0.0373
	Predict color outcomes of porcelain powder (Li et al. 2015*)	Parameters from spectrophotometer colorimetric instrument	75% of 119 sets of data: 25 of 119 sets of data	5%	Back propagation neural network with genetic algorithm	Smaller mean square error than back propagation neural network
Resin restoration removal and tooth preparation	Discriminate tooth and restorative materials (Zakeri et al. 2015*)	Cutting sounds of an air- turbine handpiece			SVM	89% accuracy for resin composite, 92% accuracy for amalgam
	Achieve tooth crown preparation (Wang et al. 2014)	Shape of the target tooth acquired by a laser scanner		Matched with expected tooth preparation	Automatic laser ablation system	Wax resin: 0.05-mm linear error, 4.33° angel error, and 0.1-mm depth error Dentin: 0.06-mm linear error, 0.5° angel error, and 0.1-mm depth error
	Achieve tooth preparation for porcelain laminate veneers (Otani et al. 2015*)	Image of tooth models scanned with 3D laser scanner	10: —	Compared with freehand tooth preparation	A robotic arm	The mean absolute deviation was 0.112 mm in the control group and 0.133 mm in the experimental group
Assist designing RPDs	Produce RPD designs according to the most similar cases (Chen et al. 2016)	Data about of oral conditions	104: —	Compared with professionals	An ontology-driven, case- based clinical decision support model	The mean average of precision was 0.61; AUC = 0.96; normalized discounted cumulative gain = 0.74
Measure masticatory efficiency	Identify patterns of masticated chewing gums (Vaccaro et al. 2018*)	Images of chewing gums obtained by flatbed scanner	400: —		Expert system	Mathews correlation coefficient score = 0.97

(continued)

Table 2. (continued)

I	Fields					
Aim	Function	— Data	Sample Size, Training: Testing	Assessment	Al Method	Results
	hodontics					
Assist therapeutic decisions	Evaluate the orthodontic treatment needs (Thanathornwong 2018)	Patients' oral examination data sets	800: 200	Compared with 2 orthodontists on the newly recruited patients	Bayesian network	A high degree of agreemen between the system and orthodontists (kappa value was 1.00 and 0.894
	Identify pretreatment patients from posttreatment and normal individuals (Wang et al. 2016)	The eye-tracking data of observers	440: —	Leave-I-out cross- validation	SVM	97.2% and 93.4% accuracy for classifying pretreatment patient from posttreatment patient and normal people, respectively
Evaluate aesthetic outcome	Assess facial attractiveness and apparent age (Patcas et al. 2019)	Pre- and posttreatment photographs	469: —		Pretrained CNN (VGG-16)	Facial attractiveness was scaled from 0 to 100; apparent age was compared with real age
	Assess facial attractiveness (Patcas et al. 2019*)	Frontal and left-side profile images		Compared with lay people, orthodontists, and oral surgeons	Pretrained CNN (VGG-16)	No significant differences with regard to the evaluation of attractiveness
End	dodontics					
Produce anatomic guidance	Identify root morphology (Hiraiwa et al. 2019)	Panoramic images	22,476: 64	Examined on CBCT images	Pretrained CNN (AlexNet) Pretrained CNN	87.4% accuracy, 77.3% sensitivity, 97.1% specificity 85.3% accuracy, 74.2%
					(GoogleNet)	sensitivity, 95.9% specificity
	Measure root canal curvature (Christodoulou et al. 2018*)	CBCT images	30: —		An algorithm for the 3D measurement	
Optimize conditions	Predict the viability of stem cells (Bindal et al. 2017)	Processing variables and cell viability			Fuzzy inference system with ANN	The root mean square error was 0.028855. The coefficient of determination was 0.811
Dental	implantology					
Classify implant types	Identify 4 types of implant fixtures (Kim et al.	Periapical radiographs	60% of 801 images: 20% of 801 images		Pretrained CNN (MobileNet-v2)	97% accuracy, 96% precisio
	2020*)				Pretrained CNN (ResNet-50)	98% accuracy, 98% precisio
					Pretrained CNN (ResNet-18)	98% accuracy, 98% precisio
					Pretrained CNN (GoogleNet)	93% accuracy, 92% precisio
					Pretrained CNN (SqueezeNet)	96% accuracy, 96% precisio

*References marked with an asterisk are included in the Appendix.

3D, 3-dimensional; AAM, active appearance model; AI, artificial intelligence; ANN, artificial neural network; ASM, active shape model; AUC, area under the receiver operating characteristic curve; CBCT, cone beam computed tomography; CNN, convolutional neural network; LDA, linear discriminant analysis; MRI, magnetic resonance imaging; NMF, negative matrix factorization; PDA, penalized discriminant analysis; QDA, quadratic discriminant analysis; RPD, removable partial denture; SVM, support vector machine.

naïve Bayes were \sim 70% accurate in predicting locoregional recurrence of squamous cell carcinoma (Alabi et al. 2020); these algorithms outperformed a contemporary clinical model that focused exclusively on tumor invasion depth. Because of the small sample size of training data sets and the large number of variables, the accuracy of these predictive models is inadvertently affected by irrelevant features (i.e., the overfitting problem) and is still below satisfaction.

Cariology and Endodontics. Predictive models based on ML algorithms, including SVM, RF, and k-nearest neighbors, were

used to identify individuals prone to tooth surface loss and root caries. SVM demonstrated the best performance, with 97% accuracy and 99.6% sensitivity, by analyzing information derived from the demographic, nutrition, lifestyle, and clinical data of patients (Hung et al. 2019). Utilizing 83 features from the American Association of Endodontists' "Endodontic Case Difficulty Assessment Form" as input, SVM and ANN obtained >90% accuracy in predicting the level of difficulty of cases requiring root canal treatment. This illustrated the potential of ML to improve the decision making of general dental practitioners in their referrals to endodontists (Mallishery et al. 2020).

 Table 3. Dental Applications of Al in Disease Prediction.

	Field	Sample Size, Training						
Prediction	Data	Testing	Validation Method	Al Methods	Accuracy	Sensitivity	Specificity	AUC
Oral and	maxillofacial surgery							
urvivability of patients with oral cavity squamous carcinoma	5 factors related to quantitative nuclear histomorphometric features of tissue sections (Lu et al. 2017*)	50: 65		Wilcoxon rank sum test and quadratic discriminant analysis	70.8%	61.5%	73.1%	0.72
lodal metastasis in early oral cavity squamous cell carcinoma	5 factors pertaining to clinical and pathologic features of patients (Bur et al. 2019)	1,570: 391	5-fold cross- validation	Decision forest Kernel SVM Gradient boosting		0.753 0.649 0.773	0.492 0.636 0.492	0.712 0.698 0.704
Stage, type, and survivability of oral	12 attributes regarding clinical details, personal history, and habits of	1,025	Leave-I-out method	PNN and general regression neural networks	70.0%	Benign and malignant 91.0% 55.7%		0.7491
cancer	patients (Sharma and Om 2015*)				67.0%	Surviva 91.0%	ability 55.7%	0.7491
	III for a superior to a secultariant	F09/ - C2F4	F 6-14	CV/M	409/	0.04	0.40	
-	I I features pertaining to clinical,	50% of 254	5-fold cross- validation	SVM	68%	0.84	0.60	
of oral tongue squamou cell carcinoma	s pathologic, and treatment features of patients (Alabi et al. 2020)	cases: 59	validation	Boosted DT	81%	0.79	0.83	
cell cal cilionia	or patients (Alabi et al. 2020)			Decision forest	78%	0.79	0.78	
				Naïve Bayes	70%	0.84	0.63	
y survival rate of	10 features selected from 17 features	31: —	5-fold cross-	Genetic programming	83.9%			0.834
patients with oral	pertaining to patients (Tan et al.		validation	SVM	64.8%			0.500
cancer	2016*)			Logistic regression	64.5%			0.500
ccurrence of	· · ·	70%: 30% of the 125		RF		100%	83.3%	0.973
bisphosphonate-related	tooth conditions (Kim et al.	patients		ANN		100%	76.7%	0.91
osteonecrosis after	2018*)			SVM		81.8%	86.7%	0.883
dental extraction				DT		90.9%	79.0%	0.82
				Logistic regression		90.9%	70.0%	0.84
icial swelling after impacted mandibular third molars extraction Cariolog	15 factors related to patients, the third molars, bone, and surgical conditions (Zhang et al. 2018) gy and endodontics	300: 100	5-fold cross- validation	ANN based on improved conjugate grads BP algorithm	98.00%			
oot caries	15 factors related to personal,	7,272: 1,818		SVM	97.1%	99.6%	94.3%	0.99
500 041105	nutrition, lifestyle, and clinical	7,272. 1,010		XGBoost	94.7%	100.0%	88.9%	0.98
	factors (Hung et al. 2019)			RF	94.1%	100.0%	87.5%	0.99
	,					97.1%	67.9%	0.88
				k-nearest neighbors	83.2%			
ooth surface loss index	II factors related to patients and	81: 15		Logistic regression ANN	74.2% 73.3%, ±5	77.1%	71.1%	0.818
	teeth (Al Haidan et al. 2014*)				scores			
ene expression of radicular cyst and periapical granuloma	Times of the gene associated to the diseases in the PubMed database (Poswar et al. 2015*)			MLP neural network				
Difficulty of endodontic	83 features from AAE case difficulty		10-fold cross-	SVM	94.8%	95.0%	94.6%	
cases	assessment form (Mallishery		validation	Deep neural network	93.4%	93.0%	93.8%	
	et al. 2020)							
	al implantology	7.17	10.611	DT	0.470	0.500	0.740	0.470
ilure of dental implants	20 attributes pertaining to patients	747: —	10-fold cross- validation	DT	0.679	0.590	0.768	0.670
	and surgeon (Liu et al. 2018*)		validation	SVM	0.628	0.581	0.675	0.628
				Logistic regressions	0.624	0.607	0.641	0.644
dividual implant mean bone levels	6 factors related to demographic and clinical features (Papantonopoulos et al. 2017)	237: —	10-fold resampling cross-validation	Ensemble selection		55%	91%	
	Age, tooth number, and implant surface (Papantonopoulos et al. 2017)	237	10-fold resampling cross-validation	SVM with Particle Swarm Optimization		62%	85%	
treatment	Dental data, patient self-behavior, health condition, and attitude (Alarifi and AlZubi 2018)			Memetic search optimization with genetic scale RNN	99.2%	97.6%	98.3%	
	Periodontics							
resence of oral malodor	16s ribosomal RNA sequence from microbiota in saliva (Nakano et al. 2018*)	90: —	Leave-I-out cross- validation	DL SVM	96.7% 78.9%	100% 77.8%	93.3% 80.0%	
Tooth mobility	9 factors related to patients (Yoon et al. 2018)		10-fold cross- validation	MLP and deep neural networks	88.4%			0.72
	Orthodontics							
erioperative blood loss in orthognathic surgery	Demographic, clinical and surgical data (Stehrer et al. 2019*)	760: 190		RF	7.4-mL mean difference			

(continued)

Table 3. (continued)

	Field	– Sample Size, Trainin	·			Sensitivity	Specificity	AUC
Prediction	Data	Testing	Validation Method	Al Methods	Accuracy			
Prosthodontics								
Longevity of dental restorations	Attributes related to patients and teeth (Aliaga et al. 2015)	4,336: 1,714	Leave-I-out validation	Bayesian network and MLP neural network	Error: 0.42 y (composites), 0.21 (amalgam)			
Debonding probability of CAD/CAM composite resin crowns	2D images captured from 3D stereolithography (Yamaguchi et al. 2019)	6,480: 2,160		CNN	98.5%			0.998
Facial deformation after complete denture prosthesis	3D pre- and postoperative models (Cheng et al. 2015*)	43: 5		BP neural network				

*References marked with an asterisk are included in the Appendix.

2D, 2-dimensional; 3D, 3-dimensional; AAE, American Association of Endodontists; Al, artifical intelligence; ANN, artificial neural network; AUC, area under the receiver operating characteristic curve; BP, back propagation; CAD/CAM, computer-aided design/computer-aided manufacturing; CNN, convolutional neural network; DL, deep learning; DT, decision tree; MLP, multilayer perceptron; PNN, probabilistic neural network; RF, random forest; RNN, recurrent neural network; SVM, support vector machine.

Dental Implantology. Predictive AI models are useful in 2 aspects of dental implantology. First, predictive models focusing on clinical outcome and individual bone levels (Papantonopoulos et al. 2017) have been constructed with ML algorithms. By simultaneously analyzing the patients' data, the implant system, and the surgeons' operations, recurrent ANN (Fig. 2C) with memetic search optimization attained 99.2% accuracy in success rate prediction (Alarifi and AlZubi 2018). Second, AI was introduced as a surrogate to contemporary technology in predicting the mechanical performance of a bioimplant system, relieving high computational cost in optimizing implant design variables. By automatically learning the explicit relationship among variables, support vector regression was almost equivalent to contemporary mathematical models in predicting stresses at the implant-bone interface (Li et al. 2019). Nevertheless, this study was conducted by using vertical loading and is not representative of a genuine chewing cycle. Application of AI in the uncertainty optimization of bioimplants still requires improvement.

Others. Based on patients' attributes, a multilayer perceptron neural network performed satisfactorily in predicting tooth mobility (Yoon et al. 2018) and longevity of resin composite restorations (Aliaga et al. 2015). CNNs achieved 98.5% accuracy in predicting the debonding probability of CAD/CAM resin composite crowns by analyzing images captured from abutment models (Yamaguchi et al. 2019). These systems are useful in forecasting prognosis and enable dentists to provide patients with the most suitable maintenance program.

Limitations and Challenges

Data Acquisition

Application of AI in dentistry is lagging behind medicine by several years (Schwendicke et al. 2019). Apart from limitations encountered in other medical fields, such as insufficient data curation and sharing (Hosny et al. 2018), the lack of

information on data processing, measuring, and validating is another blemish in dental AI research (Schwendicke et al. 2020). Sample size used for training and testing as well as the information for reference and comparative tests are sometimes unclear (Schwendicke et al. 2019), challenging the robustness, comparability, and generalizability of the results. It is imperative to improve data quantity, quality, and readability by standardizing methodology in data curation and reporting. Establishing an open-access standard data set, which contains comprehensive demographic, clinical, experimental, and treatment data, would be a crucial task in the next stage of AI development to facilitate evaluation and comparison of different algorithms.

Interpretability

Data-driven AI calculates output in a purely computational manner; however, it fails to illustrate the decision-making process in a medically acknowledgeable format. The lack of interpretability and transparency reflects the black-box nature of many ML approaches, which are not conducive to verification (Magrabi et al. 2019). Interpretability matters for 2 reasons. First, ensuring that the algorithm is a reasonable interpretation of medical incidents is important for the rapport between technology and humans. Failure to explain the inner working will inevitably disrupt practitioners' trust in the clinical value of AI. Second, the lack of transparency and interpretability makes it difficult to predict failures and generalize specific algorithms for similar contexts. Enhancing visualization has become a pivotal task. Rigorous scientific inquiry and humanistic treatment models that respond to patient and practitioner narrative exchanges are critical to understanding the clinical phenotypes of AI as well as its relationship to personalized care (Kulikowski 2019).

Computing Power

Extracting information from constantly updating medical and dental databases for the application of AI requires continuous upgrading of processing power. Because the computational power of classical computers has been largely saturated (Solenov et al. 2018), the insufficient computational resources in data processing have become one of the obstacles that constrain the efficacy of AI (Sunny et al. 2019). Quantum computing processes data by using quantum bits (qubits), in accordance with the laws and restrictions of quantum mechanics. Since quantum particles can exist in multiple states at the same time—a phenomenon known as superposition—quantum computation is exponentially faster than classical approaches (Ranjan and Hopper 2019). The ability to handle data simultaneously from very different data sets makes quantum computing a valuable platform to accelerate AI, especially ML (Sarma et al. 2019), from algorithms to data acquisition and modeling.

Ethical Considerations

Development of AI should ensure that such intelligent technologies do no harm to humans and the moral status of the machines themselves (Keskinbora 2019). Nevertheless, demands for health care privacy may no longer be attainable with a colossal scale of data sharing (Char et al. 2018). Incorporating AI into health care would inevitably replace some established services and potentially exacerbate current health inequalities (Fiske et al. 2019). These ethical paradoxes emphasize the need to establish clear guidelines for the manner in which AI is applied clinically. Judgment of legal responsibility is another ethical dilemma. At present, AI is not accountable, irrespective of whether it is employed in a supervised or unsupervised manner. The physician takes responsibility for each patient and for how information is used (Currie et al. 2020). Applying the same social and ethical norms acceptable to humans is inappropriate when the border of human responsibility is increasingly blurred by the advent of chatbot-based, unsupervised AI-based diagnosis (Hauser-Ulrich et al. 2020).

Impact of AI on Dentists

Dentistry is an excellent discipline for applying virtual AI algorithms because of its regular use of digitized imaging and electronic health records. While there is plenty of active discussion on how AI may revolutionize dental practice, concerns remain regarding whether AI will eventually replace dentists.

Contemporary AI excels in utilizing formalized knowledge and extracting information from massive data sets. However, it fails to make associations like a human brain and can only partially perform complex decision making in a clinical setting. Higher-level understanding that relies on the expertise of dentists is required, especially in ambiguous conditions, to conduct physical examinations, integrate medical histories, evaluate aesthetic outcomes, and facilitate discussion. Fleshless, machine-led dentistry does not embody the wreath of clinical care. Issues such as clinical intuition, ineffable perception, and empathy, which are keys to the delivery of personalized health care and professionalism, cannot be delivered by machines. Although there have been intense discussions

whether empathy should be incorporated into algorithms so that artificial emotions may be expressed by affective robotics, it is important to stress that effective communication between a dentist and a patient involves nonverbal evaluation of the person's hopes, fears, and expectations. Such channels of communication are intuitive and spontaneous. The most intriguing dimension of human-to-human interaction cannot simply be replicated in computer language. AI should be viewed as an augmentation tool to enhance and, at times, relieve dentists so that they can perform more valued tasks, such as integrating patient information and improving professional interactions (Recht and Bryan 2017). Pedagogy for dental students should walk hand-in-hand with AI development.

Conclusion

AI is progressing rapidly, with potential applications in diagnosis, treatment, and prognosis predictions. Although roadblocks that arise from data acquisition, interpretation, computing power, and moral issues exist and need to be overcome, AI is perceived to be an excellent adjunct for dentists. With thoughtful design and long-term clinical validation, AI can be user-friendly, transparent, reproducible, and unbiased. Future AI development should continue to consider human interest as its primary mission, with increasing capability in handling big data.

Author Contributions

T. Shan, contributed to data acquisition, analysis, and interpretation, drafted the manuscript; F.R. Tay, L. Gu, contributed to conception and design, critically revised the manuscript. All authors gave final approval and agree to be accountable for all aspects of the work.

Declaration of Conflicting Interests

The authors declared no potential conflicts of interest with respect to the research, authorship, and/or publication of this article.

Funding

The authors disclosed receipt of the following financial support for the research, authorship, and/or publication of this article: This work was supported by grant 81873712 from the National Nature Science Foundation of China, grant 2018A030313409 from the National Nature Science Foundation of Guangdong province, and grant 201904010057 from the Science and Technology Program of Guangzhou.

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