

### Automated detection and classification of coronary atherosclerotic plaques on coronary CT angiography using deep learning algorithm

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**Background:** Coronary artery disease (CAD) is the leading cause of mortality worldwide. Recent advances in deep learning technology promise better diagnosis of CAD and improve assessment of CAD plaque buildup. The purpose of this study is to assess the performance of a deep learning algorithm in detecting and classifying coronary atherosclerotic plaques in coronary computed tomographic angiography (CCTA) images.

**Methods:** Between January 2019 and September 2020, CCTA images of 669 consecutive patients with suspected CAD from Nanjing Drum Tower Hospital Clinical College of Nanjing University of Chinese Medicine were included in this study. There were 106 patients included in the retrospective plaque detection analysis, which was evaluated by a deep learning algorithm and four independent physicians with varying clinical experience. Additionally, 563 patients were included in the analysis for plaque classification using the deep learning algorithm, and their results were compared with those of expert radiologists. Plaques were categorized as absent, calcified, non-calcified, or mixed.

**Results:** The deep learning algorithm exhibited higher sensitivity, specificity, positive predictive value (PPV), negative predictive value (NPV), and accuracy {92% [95% confidence interval (CI): 89.5–94.1%], 87% (95% CI: 84.2–88.5%), 79% (95% CI: 76.1–82.4%), 95% (95% CI: 93.4–96.3%), and 89% (95% CI: 86.9–90.0%)} compared to physicians with ≤5 years of clinical experience in CAD diagnosis for the detection of coronary plaques. The algorithm's overall sensitivity, specificity, PPV, NPV, accuracy, and Cohen's kappa

for plaque classification were 94% (95% CI: 92.3–94.7%), 90% (95% CI: 88.8–90.3%), 70% (95% CI: 68.3–72.1%), 98% (95% CI: 97.8–98.5%), 90% (95% CI: 89.8–91.1%) and 0.74 (95% CI: 0.70–0.78), indicating strong performance.

**Conclusions:** The deep learning algorithm has demonstrated reliable and accurate detection and classification of coronary atherosclerotic plaques in CCTA images. It holds the potential to enhance the diagnostic capabilities of junior radiologists and junior intervention cardiologists in the CAD diagnosis, as well as to streamline the triage of patients with acute coronary symptoms.

**Keywords:** Coronary computed tomographic angiography (CCTA); deep learning; atherosclerotic plaques; coronary artery disease (CAD)

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### Introduction

Coronary artery disease (CAD) is the leading cause of mortality worldwide (1,2). Accurate diagnosis of CAD presence and severity is pivotal in determining appropriate clinical management for symptomatic patients (3). Traditionally, digital subtraction angiography has served as the gold standard for diagnosing CAD (4). Other coronary plaque imaging methods, such as intravascular ultrasound and optical coherence tomography, have also been employed to identify culprit coronary atherosclerosis (5,6). However, these invasive imaging techniques may entail complications such as arrhythmia, myocardial infarction, or strokes (7,8).

In contrast, coronary computed tomographic angiography (CCTA) is a non-invasive technique capable of accurately detecting obstructive CAD and evaluating atherosclerotic plaque burden in the coronary artery (7,9-13). Despite its reliability as a diagnostic tool, utilizing CCTA for coronary artery assessment remains timeconsuming and requires extensive expertise (14,15).

Recent advancements in deep learning technology have shown promising results in enhancing the efficiency of CAD plaque assessment (14,16-24). For the optical coherence tomography images, Liu *et al.* proposed a framework which automatically segmented lumen and calcified plaque, by leveraging deep learning techniques (25). For CCTA images, a significant contribution to this field is the development of a deep learning-based technology that leverages an enhanced three-dimensional (3D) U-Net architecture. This advanced version of the 3D U-Net incorporates a bottle-neck design for fully automated segmentation, alongside a connected growth prediction model (CGPM) for the repair of segmentation fractures (3,26). Crucial to its design, this architecture is augmented with attention mechanisms and residual modules, which are not explicitly mentioned but integral to its operation. The attention mechanisms enable focused analysis of salient image features critical for the accurate identification of obstructive CAD, while the residual modules facilitate the training of deeper networks by alleviating the vanishing gradient problem, thus enhancing the model's ability to learn complex patterns in the data. This nuanced architecture has demonstrated the capability to accurately and effectively identify obstructive CAD (21), showcasing the potential of deep learning in revolutionizing CAD plaque assessment. However, the widespread application of this technology necessitates rigorous validation of its diagnostic efficiency. Moreover, it is imperative to benchmark the accuracy and reliability of this deep learning approach against the diagnostic performance of physicians at various levels of expertise.

This study has a dual purpose: firstly, to determine the capability of deep learning technology in detecting coronary plaques by comparing it with physicians at varying diagnostic levels; secondly, to assess its ability to distinguish between different types of plaques. We present this article in accordance with the STARD reporting checklist (available at https://qims.amegroups.com/article/view/10.21037/ qims-23-1513/rc).

### **Methods**

The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by the Institutional Review Board of Nanjing Drum Tower Hospital Clinical College of Nanjing

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Figure 1 A flow diagram of study population. CCTA, coronary computed tomographic angiography.

University of Chinese Medicine (No. 2022-547-01), and individual consent for this retrospective analysis was waived. Between January 2019 and September 2020, CCTA images of 773 consecutive patients with suspected CAD from Nanjing Drum Tower Hospital Clinical College of Nanjing University of Chinese Medicine were collected. First, of 773 patients, 38 patients with poor quality CCTA images were excluded. Second, 59 patients with a history of coronary stenting or coronary artery bypass grafting were excluded. Thirdly, 7 patients with severe cardiac or coronary anatomical variants were excluded. Finally, CCTA images from a total of 669 patients were included in this study (Figure 1). These patients were divided into two groups to evaluate the performance of the deep learning algorithm: Group 1, consisting of 106 patients, for plaque detection, and Group 2, comprising 563 patients, for plaque type characterization.

In Group 1, the algorithm's results were compared with assessments made by physicians with varying levels of expertise: a cardiovascular research radiologist (K.Y., <2 years of clinical experience), a junior cardiovascular radiologist (W.C., 3 years of clinical experience), an intervention cardiologist (R.G., 5 years of clinical experience), and a senior cardiovascular radiologist (H.Y., 5 years of clinical experience). Diagnostic outcomes from two expert cardiovascular radiologists (H.L. and D.M., >10 years of clinical experience) were considered the gold standard for comparison. Any disparities were resolved through a consensus process among them. Moreover, these two expert radiologists interpreted CCTA images from the remaining 563 patients (Group 2), categorizing plaques as absent, calcified, non-calcified, or mixed calcified per segment, following Society of Cardiovascular Computed Tomography (SCCT) guidelines (27,28) (see *Figure 2*). The algorithm's outputs were then compared to these expert interpretations. All physicians were blind to the patients' clinical information.

### Imaging acquisition and analysis

All CCTA scans were conducted using a 256-slice computed tomography (CT) scanner (Brilliance iCT, Philips Healthcare, Cleveland, OH, USA) using electrocardiography (ECG)-gated prospective-triggered technology. The following acquisition parameters were applied: a tube voltage of 100 kVp, detector collimation of 128×0.625 mm, a rotation time of 0.27 seconds and automatic tube current modulation enabled (DoseRight with index setting of 16, Philips Healthcare). A total of 65 mL of iopromide with an iodine concentration of 370 mg/mL (Ultravist, Bayer, Guangzhou, China) was intravenously administered at 4-5 mL/s, followed by 50 mL of normal saline at same injection speed. The bolus tracking technique was employed, with a delay of 7 seconds after reaching a trigger threshold of 130 Hounsfield unit (HU) in the proximal descending aorta.

As per SCCT guidelines (27,28), atherosclerotic plaque analysis in the coronary artery segments was performed



**Figure 2** Classification of coronary atherosclerotic plaques using CTA by two experienced cardiovascular radiologists on consensus as internal reference for deep learning. (A) Absence of plaques in the left anterior descending artery; (B) calcified plaques (arrow); (C) non-calcified plaque (arrow); and (D) mixed plaque (calcified and non-calcified, open arrow). CTA, computed tomographic angiography.



Figure 3 Schematic diagram of the 18 coronary artery segments. 1, proximal right coronary artery (pRCA); 2, middle right coronary artery (mRCA); 3, distal right coronary artery (dRCA); 4, right posterior descending artery (R-PDA); 5, left main (LM); 6, proximal left anterior descending (pLAD); 7, middle left anterior descending (mLAD); 8, distal left anterior descending (dLAD); 9, diagonal 1 (D1); 10, diagonal 2 (D2); 11, proximal circumflex (pCx); 12, obtuse marginal 1 (OM1); 13, mid and distal left circumflex (LCx); 14, obtuse marginal 2 (OM2); 15, posterior descending artery from LCx (L-PDA); 16, right posterior lateral branch (R-PLB); 17, ramus intermedius (RI); 18, posterior lateral branch from LCx (L-PLB).

using a dedicated workstation (IntelliSpace Portal, Philips Healthcare). The evaluation covered 18 coronary artery segments: proximal right coronary artery (pRCA, 1), middle right coronary artery (mRCA, 2), distal right coronary artery (dRCA, 3), right posterior descending artery (R-PDA,

4), left main (LM, 5), proximal left anterior descending (pLAD, 6), middle left anterior descending (mLAD, 7), distal left anterior descending (dLAD, 8), diagonal 1 (D1, 9), diagonal 2 (D2, 10), proximal circumflex (pCx, 11), obtuse marginal 1 (OM1, 12), mid and distal left circumflex (LCx, 13), obtuse marginal 2 (OM2, 14), posterior descending artery from LCx (L-PDA, 15), right posterior lateral branch (R-PLB, 16), ramus intermedius (RI, 17), and posterior lateral branch from LCx (L-PLB, 18) (*Figure 3*).

The interpretation of CCTA images involved the following steps:

- (I) Image loading and review: loading the CCTA image series onto the workstation for reviewing the original axial images to assess quality and identify anatomical landmarks.
- (II) Heart segmentation stage: manual correction or resegmentation, if required.
- (III) Coronary extraction: verification, correction, or manual definition of coronary artery segmentation, vessel centerlines, and their labels. A straightened multi-planar reconstruction (MPR) image of the selected artery was provided.
- (IV) Coronary analysis: including plaque detection, diameter and area measurements of stenosis, and selection of reference locations. Volume-rendered images, curved MPR images, straightened MPR images, and cross-section images of vessels were provided for analysis.
- (V) Plaque analysis: calculation and display of plaque content as the percent (%) of calcified plaque, mixed plaque, and non-calcified plaque based on

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**Figure 4** Schematic diagram of the deep learning-based plaque detection algorithm workflow. (A) Stack of transverse images; (B) segmentations of aorta and coronary arteries (red area); (C) volume rending image of aorta and coronary arteries; (D) straightened rendering image of the left anterior descending artery; (E) architect of three-dimensional U-Net2 model; and (F) two probability curves corresponding to calcified and noncalcified plaques, respectively.

pre-defined HU thresholds. The interpretation time was recorded from image loading to plaque detection or plaque analysis.

### Deep learning algorithm

A deep learning-based system (Coronary Doc, ShuKun Technology, Beijing, China) was employed for coronary artery segmentation and plaque detection. This system primarily utilized 3D U-Net1 architecture with a bottleneck design for fully automated segmentation and a CGPM for segmentation fracture repair (26,29). After transferring the original axial images to the system, various image reconstructions, such as multiple planar reformat, straightened rendering, curve planar reformat, maximum intensity projection, and volume rendering images, were created based on the segmentation.

Further enhancing the 3D U-Net1 architecture, a dedicated 3D U-Net model (U-Net2) was trained specifically for recognizing plaques within the coronary arteries. Data cubes sized 60×60 (within the cross-sectional plane) ×128 (along the centerline) were extracted from the straightened arteries and input into the model. This model consisted of a five-layer encoding structure, a fivelayer decoding structure, and a max-pooling output layer. It underwent 100 epochs of training using stochastic gradient descent as the optimizer, starting with a learning rate of 0.01 and halving it every 20 epochs. The output layer of U-Net2 generated two probability curves, each 128 pixels in length, corresponding to calcified and non-calcified (soft) plaques (16,18,21,22,26,30,31) (Figure 4). In this context, a calcified plaque referred to a plaque with a CT value of  $\geq$ 130 HU, commonly used to quantify calcification burden in clinical trials and identify calcified plaques in CCTA

Table 1 Demogra	phics of patients	with suspected and	known CAD
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Demographics	All	Values <sup>†</sup>	All	Values <sup>†</sup>
No. of patients	106		563	
Female		39 (36.8)		225 (40.0)
Age, years		63±10		61±12
Indication				
Chest pain		19 (17.9)		139 (24.7)
Chest tightness		31 (29.2)		163 (29.0)
CAD		28 (26.4)		155 (27.5)
Lower extremity edema		6 (5.7)		18 (3.2)
Shortness of breath		10 (9.4)		14 (2.5)
Others		12 (11.3)		74 (13.1)
History of CAD	70	52 (74.3)	400	330 (82.5)
BMI, kg/m <sup>2</sup>	72		374	
<23 (normal)		25 (34.7)		85 (22.7)
23.00–27.49 (overweight)		21 (29.2)		122 (32.6)
≥27.5 (obesity)		26 (36.1)		167 (44.7)
Ever smoke (former and current)	53	14 (26.4)	271	94 (34.7)
Diabetes	93	33 (35.5)	292	106 (36.3)
Hypertension	96	67 (69.8)	364	271 (74.5)
Hyperlipidemia	83	23 (27.7)	270	90 (33.3)
Cardiovascular events	72	56 (77.8)	249	190 (76.3)

<sup>†</sup>, data are presented as mean  $\pm$  SD or n (%). The indication "CAD" represents the suspicious diagnosis made by the doctor during the outpatient visit, based on the patient's symptoms, laboratory tests, and other relevant factors. A history of CAD refers to a patient who has previously been diagnosed with CAD. Group 1: for plaque detection; Group 2: for plaque type characterization. CAD, coronary artery disease; SD, standard deviation; BMI, body mass index.

images. Conversely, a non-calcified plaque was defined by a CT value of <130 HU, while a mixed plaque denoted the overlap between calcified and non-calcified plaque positions. Plaque locations were determined by areas in the probability curve exceeding the cut-off value.

This deep learning-based system was integrated into the picture archiving and communication system (PACS), automatically analyzing coronary axial computed tomographic angiography (CTA) images transferred to PACS. It provided preliminary results, including comprehensive 3D reconstructions for curved plane reconstruction (CPR), lumen, and multi-cross sections, along with location identification, qualitative and quantitative plaque analyses. The interpretation time using this system involved opening the initial report, reviewing reconstructed images, and confirming the final report.

### Statistical analysis

Statistical analysis was conducted to evaluate the deep learning model's performance using R (version 4.1.1, R Foundation for Statistical Computing, Vienna). Sensitivity, specificity, positive predictive value (PPV), negative predictive values (NPV), accuracy, and Cohen's kappa were calculated. Cohen's kappa>0.61 was considered to have strong consistency. Categorical variables were presented as counts or percentages, while continuous variables were described as means with standard deviations (SD). McNemar's test was utilized for continuous, two-sided, paired variables, with a two-tailed P value <0.05 considered significant.

### **Results**

The 669 patients were divided into two groups to evaluate the performance of the deep learning algorithm: Group 1, consisting of 106 patients, for plaque detection, and Group 2, comprising 563 patients, for plaque type characterization. Patient demographics are summarized in *Table 1*.

## Coronary plaque detection efficiency of deep learning compared to physicians

The average sensitivity, specificity, PPV, NPV, and accuracy of the deep learning system were 92% [95% confidence interval (CI): 89.5-94.1%], 87% (95% CI: 84.2-88.5%), 79% (95% CI: 76.1-82.4%), 95% (95% CI: 93.4-96.3%), and 89% (95% CI: 86.9-90.0%), respectively. Comparative data for cardiovascular radiologists with varying experience levels were as follows: those with 5 years of clinical experience achieved an average sensitivity of 82% (95% CI: 78.3-84.7%), specificity of 86% (95% CI: 84.0-88.3%), PPV of 77% (95% CI: 73.6-80.4%), NPV of 89% (95% CI: 87.2-89.3%), and accuracy of 85% (95% CI: 82.8-86.4%); those with 3 years of clinical experience had an average sensitivity of 80% (95% CI: 76.4-83.2%), specificity of 82% (95% CI: 79.7-84.6%), PPV of 72% (68.1-75.2%), NPV of 88% (95% CI: 85.6-89.9%), and accuracy of 81%

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Table 2 Efficiency of plaques in the coronary artery segments on CCTA identified by deep learning in comparison with physicians with different levels of proficiency

		Ref. st	andard											Cli	nical e	exper	ience	•					
Segments	Total	(2 radiolo more than clinical ex	ogists with 10 years of xperience)	D	eep l	earnii	ng	Ra	diolo 5 ye	gist v ears	vith	Ra	diolo 3 ye	gist v ears	vith	Ra	diolo ≤2 y	gist v /ears	vith	lr care	nterve diolog 5 ye	entior gist w ars	า /ith
		Positive	Negative	TP	ΤN	FP	FN	TP	ΤN	FP	FN	TP	ΤN	FP	FN	TP	ΤN	FP	FN	TP	ΤN	FP	FN
pRCA	106	56	50	54	47	3	2	49	46	4	7	49	43	7	7	50	42	8	6	44	44	6	12
mRCA	106	45	61	40	46	15	5	35	53	8	10	35	48	13	10	42	54	7	3	37	51	10	8
dRCA	97	19	78	16	59	19	3	13	61	17	6	15	62	16	4	16	74	4	3	16	77	1	3
R-PDA	95	32	63	29	62	1	3	27	61	2	5	24	56	7	8	22	50	13	10	25	59	4	7
LM	106	41	65	40	49	16	1	28	56	9	13	31	55	10	10	20	43	22	21	32	59	6	9
pLAD	106	67	39	62	38	1	5	64	35	4	3	58	34	5	9	63	37	2	4	58	28	11	9
mLAD	106	55	51	52	36	15	3	47	38	13	8	46	44	7	9	51	44	7	4	49	47	4	6
dLAD	106	12	94	10	81	13	2	7	80	14	5	7	59	35	5	2	82	12	10	5	84	10	7
D1	106	35	71	32	58	13	3	23	60	11	12	27	62	9	8	14	42	29	21	27	55	16	8
D2	93	32	61	29	55	6	3	23	47	14	9	22	47	14	10	15	41	20	17	21	50	11	11
рСх	106	56	50	53	48	2	3	56	48	2	0	50	42	8	6	39	35	15	17	48	39	11	8
OM1	101	27	74	26	72	2	1	25	64	10	2	23	60	14	4	15	54	20	12	21	67	7	6
LCx	106	27	79	25	59	20	2	11	68	11	16	17	66	13	10	20	66	13	7	19	73	6	8
OM2	78	12	66	12	63	3	0	10	60	6	2	7	59	7	5	6	56	10	6	5	56	10	7
L-PDA	11	5	6	4	4	2	1	4	3	3	1	3	3	3	2	2	4	2	3	1	3	3	4
R-PLB	81	26	55	23	55	0	3	23	50	5	3	22	51	4	4	19	44	11	7	18	45	10	8
RI	45	11	34	9	32	2	2	10	29	5	1	10	28	6	1	6	24	10	5	5	24	10	6
L-PLB	25	12	13	9	10	3	3	11	13	0	1	10	12	1	2	10	11	2	2	7	9	4	5
Sensitivity, %				92	2 (89.	5–94	.1)	82	2 (78.	3–84	.7)	80	) (76.	4–83	.2)	72	2 (68.	4–75	.9)	78	78 (73.1–80		2)
Specificity, %				87	7 (84.	2–88	.5)	86	6 (84.	0–88	.3)	82	2 (79.	7–84	.6)	79	9 (76.	9–81	.9)	86	(83.8	8–88.	2)
PPV, %				79	9 (76.	1–82	.4)	77	7 (73.	6–80	.4)	72 (68.1–75.2)			.2)	67 (62.7–70.2)			76	(72.0	)–79.	2)	
NPV, %				98	5 (93.	4–96	.3)	89	9 (89.	3–87	.2)	88	8 (85.	6–89	.9)	84	4 (81.	0–85	.8)	87	(84.5	5–88.	8)
Accuracy, %				89	9 (86.	9–90	.0)	85	5 (82.	8–86	.4)	81	(79.	5 –83	.3)	77	7 (74.	8–78	.9)	83	(80.8	8–84.	6)

Data in parentheses are 95% confidence interval. CCTA, coronary computed tomographic angiography; TP, true positive; TN, true negative; FP, false positive; FN, false negative; PPV, positive predictive value; NPV, negative predictive value; pRCA, proximal right coronary artery; mRCA, middle right coronary artery; dRCA, distal right coronary artery; R-PDA, right posterior descending artery; LM, left main; pLAD, proximal left anterior descending; mLAD, middle left anterior descending; dLAD, distal left anterior descending; D1, diagonal 1; D2, diagonal 2; pCx, proximal circumflex; OM1, obtuse marginal 1; LCx, mid and distal left circumflex; OM2, obtuse marginal 2; L-PDA, posterior descending artery from LCx; R-PLB, right posterior lateral branch; RI, ramus intermedius; L-PLB, posterior lateral branch from LCx.

(95% CI: 79.5-83.3%); while the research cardiovascular radiologist with less than 2 years of clinical experience achieved an average sensitivity of 72% (95% CI: 68.4-75.9%), specificity of 79% (95% CI: 76.9%-81.9%), PPV of 67% (95% CI: 62.7-70.2%), NPV of 84% (95% CI: 81.0-85.8%), and accuracy of 77% (95% CI: 74.8-78.9%).

The intervention cardiologist with five years of clinical experience reached an average sensitivity of 78% (95% CI: 73.1–80.2%), specificity of 86% (95% CI: 83.8–88.2%), PPV of 76% (95% CI: 72.0–79.2%), NPV of 87% (95% CI: 84.5–88.8%), and accuracy of 83% (95% CI: 80.8–84.6%). For detailed results, refer to *Table 2*.



**Figure 5** Boxplots and scatterplots of interpretation time by radiologists and deep learning. (A) There is significant reduction in interpretation time by deep learning for the classification of coronary artery atherosclerosis (mean of 298 s) as opposed to 658 s by radiologists; (B) there is only one plaque that shows longer interpretation time by deep learning than by radiologist. AI, artificial intelligence.

### Deep learning time efficiency on plaque classification

When radiologists classified plaques per patient, interpretation time ranged from 280 to 995 s, with a mean of  $658\pm150$  s. In contrast, using the deep learning model, interpretation time ranged from 125 to 689 s, with a mean of 298±60 s. Significantly shorter interpretation time was observed with deep learning compared to radiologists (McNemar's test, P<0.001) (*Figure 5*).

### Deep learning performance on plaque classification

Among 563 patients, there were 1,233 calcified plaques, 416 non-calcified plaques, and 130 mixed plaques. The overall sensitivity, specificity, PPV, NPV, and accuracy were 94% (95% CI: 92.3–94.7%), 90% (95% CI: 88.8–90.3%), 70% (95% CI: 68.3–72.1%), 98% (95% CI: 97.8–98.5%), and 90% (95% CI: 89.8–91.1%), respectively, demonstrating substantial agreement ( $\kappa$ =0.74, 95% CI: 0.70–0.78) between the deep learning model and two expert cardiovascular radiologists in coronary atherosclerosis classification (*Table 3*).

For calcified plaques, sensitivity, specificity, PPV, NPV, and accuracy were 93% (95% CI: 91.9–94.7%), 96% (95% CI: 95.6–96.5%), 81% (95% CI: 78.6–82.8%), 99% (95% CI: 98.5–99.0%) and 96% (95% CI: 95.2–96.1%), respectively (*Table 4*). Lowest PPV, particularly in the dLAD and R-PLB areas, was the primary factor contributing to the overall lower PPV in characterizing calcified plaques with deep learning. In characterizing non-calcified plaques, mean sensitivity, specificity, PPV, NPV, and accuracy were 84% (95% CI: 79.7–87.0%), 95% (95% CI: 94.7–95.7%), 48% (95% CI: 44.6–52.1%), 99% (95% CI: 98.8–99.3%) and 95% (95% CI: 94.1–95.1%), respectively (*Table 4*). Similarly, the lowest PPV in the dLAD and R-PDA areas contributed to the lower average PPV in characterizing non-calcified plaques. Concerning mixed plaques, mean sensitivity, specificity, PPV, NPV, and accuracy were 72% (95% CI: 62.8–78.9%), 99% (95% CI: 99.4–99.7%), 79% (95% CI: 70.1–85.6%), 99% (95% CI: 99.2–99.6%), 81% (95% CI: 79.7–81.4%), respectively (*Table 4*).

### Discussion

In our study, the employed deep learning-based system (Coronary Doc, ShuKun Technology, Beijing, China) offers multiple advantages over conventional algorithms. It utilizes a 3D U-Net1 architecture, providing a more comprehensive spatial context for accurate plaque detection compared to existing 2D models (21,29). Additionally, it significantly reduces subjectivity and time consumption associated with manual or semi-automatic plaque detection methods, thus enhancing efficiency and reproducibility. It has been tested on an NVIDIA GeForce GTX 1080 Ti graphics card with which it demonstrates the ability to complete the diagnosis process for CAD within 120 seconds. This timeframe includes the full range of computations, from initial image processing to the final output. Using more advanced graphics cards can further reduce this computation time, indicating that the model's efficiency improves with better hardware. This scalability suggests the approach is not only effective with current technology but will also benefit from future advancements in graphics processing unit (GPU) capabilities. Furthermore, the integration of a CGPM for segmentation fracture repair notably improved plaque Table 3 Overall statistics of all plaques in the coronary artery segments on CCTA identified by deep learning in comparison with experienced radiologists

		Ref. sta than 1(	ndard (2 rac 0 years of c	liologists √ linical exp	with more erience)	Δ	eep le	arning		Concitation of	Coccelline Cocce				
Segments	Total	Non- plaque	Calcified	Non- calcified plaques	Mixed plaques	Ц	N	FР	N	Зепыцицу (95% CI), %	(95% CI), %	(95% Cl), %	(95% CI),%	Acculacy (95% Cl), %	к (95% СІ)
pRCA	563	364	113	66	20	179	299	75	10	95 (90.2–97.3)	80 (75.5–83.8)	70 (64.4–75.9)	97 (93.9–98.3)	85 (81.7- 87.6)	0.74 (0.69–0.79)
mRCA	563	377	129	49	8	168	291	97	7	96 (91.6–98.2)	75 (70.3–79.2)	63 (57.2–69.1)	97 (95.0–98.9)	82 (78.1- 84.5)	0.68 (0.62–0.73)
dRCA	526	412	75	35	4	108	356	60	N	98 (92.9–99.7)	86 (81.7–88.7)	64 (56.5–71.4)	99 (97.8–99.9)	88 (85.2–90.7)	0.73 (0.66–0.79)
R-PDA	485	445	32	9	2	40	407	38	0	100 (89.1–100)	91 (88.4–93.8)	51 (39.8–62.7)	100 (98.9–100)	92 (89.4–94.3)	0.65 (0.55–0.75)
LM	563	430	108	19	9	122	412	22	7	94 (88.7–97.6)	95 (92.3–96.7)	85 (77.6–89.9)	98 (96.4–99.3)	95 (92.7–96.4)	0.87 (0.82–0.91)
pLAD	563	291	157	73	42	220	258	54	31	88 (82.8–91.3)	83 (77.9–86.6)	80 (75.0–84.7)	89 (85.0–92.5)	85 (81.7–87.6)	0.76 (0.71–0.81)
mLAD	563	311	176	62	14	222	250	72	19	92 (87.8–95.1)	78 (72.6–81.9)	75 (70.1–80.2)	93 (89.0–95.5)	84 (80.6–86.7)	0.74 (0.69–0.79)
dLAD	563	540	18	5	0	21	507	33	N	91 (70.5–98.5)	94 (91.4–95.7)	39 (26.2–53.1)	99 (98.4–99.9)	94 (91.5–95.5)	0.52 (0.39–0.66)
5	554	475	65	13	-	76	442	33	с	96 (88.5–99)	93 (90.3–95.1)	70 (60.1–77.9)	(9.66–6.76) 66	93 (91.1–95.3)	0.79 (0.71–0.85)
D2	399	354	35	10	0	44	342	12	-	98 (86.8–99.9)	97 (94.0–98.2)	79 (65.2–88.0)	99 (98.1–99.9)	97 (94.5–98.1)	0.86 (0.78–0.93)
pCx	563	425	113	17	80	124	384	44	7	92 (85.6–95.7)	90 (86.4–92.3)	74 (66.4–80.1)	97 (94.9–98.5)	90 (87.5–92.4)	0.77 (0.71–0.83)
OM1	493	449	37	7	0	44	413	36	0	100 (90.0–100)	92 (90.0–94.2)	55 (43.5–66.0)	100 (98.9–100)	93 (90.0–94.7)	0.69 (0.58–0.78)
LCX	563	438	100	21	4	118	397	44	4	97 (91.3–98.9)	90 (86.7–92.6)	73 (65.2–79.3)	99 (97.3–99.7)	91 (88.9–93.5)	0.79 (0.73–0.85)
OM2	303	280	19	4	0	23	264	16	0	100 (82.2–100)	94 (90.7–96.6)	59 (42.2–74.2)	100 (98.2–100)	95 (91.5–96.8)	0.72 (0.59–0.85)
L-PDA	78	64	9	ი	5	12	59	5	N	86 (56.2–97.5)	92 (82.0–97.1)	70 (44.0–88.6)	97 (87.6–99.4)	91 (82.3–95.9)	0.74 (0.56–0.92)
R-PLB	462	422	18	15	7	30	417	10	5	86 (69.0–94.6)	98 (95.6–98.8)	75 (58.5–86.8)	96 (97–99·6)	97 (94.7–98.1)	0.79 (0.61–0.97)
Ы	258	229	25	4	0	28	221	8	-	96 (80.4–99.8)	96 (93.0–98.4)	78 (60.4–89.3)	99 (97.1–99.9)	97 (93.4–98.3)	0.85 (0.75–0.94)
L-PLB	101	78	7	7	6	14	72	12	ო	82 (55.8–95.3)	86 (76–92.1)	54 (33.7–72.9)	96 (87.9–99.0)	85 (76.8–90.9)	0.63 (0.48–0.79)
The mean 90% (95% 95% CI: 0.	sensitiv CI: 89. 70–0.78	ity, spec 8–91.19 \$). CCTA	cificity, PPV, %). Overall, v, coronary c	, NPV, and there is si computed	d accuracy ubstantial tomograp	/ are 9 agreer hic an	4% (9. ment b giogra	5% CI etwee ohy; TI	: 92.3 n dee P, true	-94.7%) 90% (( ep learning and positive; TN, tn	95% CI: 88.8-90 the radiologists i ue negative; FP, f	.3%), 70% (95% n the identificatio alse positive; FN	Cl: 68.3–72.1%) on of the coronal , false negative; F	), 98% (95% CI: ry atherosclerotic PPV, positive prec	97.8–98.5%), and plaques (ĸ=0.74, dictive value; NPV,
descending	arterv.	e value; : LM. lef	Cl, contide t main: pLA	nce interv D. proxim	al; pHCA, ial left ante	proxim erior de	al righ scend	t coro ina: m	LAD.	artery; mHCA, m middle left ante	niddle right coron rior descendina:	ary artery; dHCA dLAD. distal left	, distal right corc anterior descend	onary artery; K-PI lina: D1. diadonal	JA, right posterior 1:D2. diaαonal 2:
pCx, proxii	nal circ	umflex;	OM1, obtu:	se margin	al 1; LCx,	mid ar	nd dist	al left	circur	mflex; OM2, obt	use marginal 2; L	-PDA, posterior	descending arter	y from LCx; R-PI	-B, right posterior
lateral brar	ich; RI,	ramus ir	ntermedius;	L-PLB, p(	osterior lat	eral br	anch f	rom L(	č.						

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Table 4 Identification of coronary calcified plaques, coronary non-calcified plaques and coronary mixed plaques on CCTA using deep learning in comparison with expert radiologists

O a marca and a	Tatal	Cord	onary calc	ified plac	ques	Corona	ary non-ca	alcified pl	laques	Core	onary mix	ed plaque	es	
Segments	Iotal -	TP	TN	FP	FN	TP	TN	FP	FN	TP	TN	FP	FN	
pRCA	563	105	435	15	8	57	438	59	9	17	542	1	3	
mRCA	563	122	414	20	7	40	442	72	9	6	550	5	2	
dRCA	526	74	421	30	1	32	462	29	3	2	521	1	2	
R-PDA	485	32	436	17	0	6	459	20	0	2	482	1	0	
LM	563	101	440	15	7	17	537	7	2	4	557	0	2	
pLAD	563	141	383	23	16	53	463	27	20	26	517	4	16	
mLAD	563	162	365	22	14	49	453	48	13	11	547	2	3	
dLAD	563	17	523	22	1	4	547	11	1	-	-	-	-	
D1	554	62	471	18	3	13	527	14	0	1	552	1	0	
D2	399	34	356	8	1	10	385	4	0	-	-	-	-	
рСх	563	104	435	15	9	14	519	27	3	6	553	2	2	
OM1	493	37	435	21	0	7	472	14	0	0	492	1	0	
LCx	563	95	441	22	5	19	522	20	2	4	557	2	0	
OM2	303	19	273	11	0	4	294	5	0	-	-	-	-	
L-PDA	78	5	70	2	1	3	73	2	0	4	72	1	1	
R-PLB	462	14	436	8	4	10	445	2	5	6	455	0	1	
RI	258	24	231	2	1	4	248	6	0	-	-	-	-	
L-PLB	101	4	91	3	3	6	89	5	1	4	88	4	5	
Sensitivity (95% CI), %			93 (91.9	94.7)			84 (79.7	7–87.0)		72 (62.8–78.9)				
Specificity (95% Cl), %		96 (95.6–96.5)					95 (94.7	/-95.7)		99 (99.4–99.7)				
PPV (95% CI), %			81 (78.6	6–82.8)			48 (44.6	6–52.1)		79 (70.1–85.6)				
NPV (95% CI), %			99 (98.5	5–99.0)			99 (98.8	8–99.3)			99 (99.2-	-99.6)		
Accuracy (95% CI), %			96 (95.2	2–96.1)			95 (94.1	-95.1)			81 (79.7-	-81.4)		

CCTA, coronary computed tomographic angiography; TP, true positive; TN, true negative; FP, false positive; FN, false negative; CI, confidence interval; pRCA, proximal right coronary artery; mRCA, middle right coronary artery; dRCA, distal right coronary artery; R-PDA, right posterior descending artery; LM, left main; pLAD, proximal left anterior descending; mLAD, middle left anterior descending; dLAD, distal left anterior descending; D1, diagonal 1;D2, diagonal 2; pCx, proximal circumflex; OM1, obtuse marginal 1; LCx, mid and distal left circumflex; OM2, obtuse marginal 2; L-PDA, posterior descending artery from LCx; R-PLB, right posterior lateral branch; RI, ramus intermedius; L-PLB, posterior lateral branch from LCx; PPV, positive predictive value; NPV, negative predictive value.

detection and segmentation (21,22). The dedicated 3D U-Net model (U-Net2) specifically trained for coronary arteries offers enhanced precision in plaque detection, providing a more detailed view of plaque morphology, composition, and distribution.

For deep learning-based coronary plaque detection models, the most basic function is to identify plaques as well as stenosis in the vessel lumen. Only when this basic function achieves a high degree of stability, the accurate classification of plaques may be further realized. In the first part of this study, the deep learning-based model demonstrated the significantly enhanced efficiency in coronary artery plaque detection when compared to radiologists or an intervention cardiologist with up to 5 years of clinical experience. In the second part of this study, the model exhibited high consistency in classifying coronary atherosclerotic plaques compared to expert radiologists while achieving this in a notably shorter interpretation time. Furthermore, our analysis revealed not only good sensitivity and accuracy but also excellent NPV in diagnosing coronary atherosclerotic plaques. This is particularly valuable in excluding culprit atherosclerotic plaques.

The implications of our study are significant. Firstly, in daily practice, prompt diagnosis and appropriate treatment of coronary atherosclerosis are crucial to prevent myocardial infarction and sudden death. In densely populated countries like China, efficiently identifying individuals at risk of CAD and providing effective treatment to those with the disease are imperative. CCTA, more accurately detecting CAD (32), holds clinical significance in identifying high-risk CAD patients using efficient deep learning techniques. Moreover, for CAD patients, quickly determining the location and hemodynamic status of culprit blood vessels may allow cardiovascular physicians sufficient time for precise treatment. Secondly, variations in CAD therapy among hospitals or physicians may lead to different outcomes for CAD patients. Therefore, accurate and precise CAD identification is crucial. The deep learning technology in our study holds the potential to minimize disparities and enhance the precision of CAD treatment among different medical institutions. Thirdly, leveraging the capabilities of deep learning to distinguish plaque types and its excellent negative prediction ability may help avoid unnecessary standardized treatments for some CAD patients. Additionally, the extensive use of deep learning could become an indispensable tool for training physicians in CAD diagnosis and treatment. The efficacy of our study's deep learning for diagnostics suggests that it can provide relatively reliable results and potentially help physicians of different expertise levels shorten the time required for CAD diagnosis training.

There are limitations in this study. Firstly, as a retrospective study, there might be data insufficiencies, leading to the exclusion of some clinically significant indicators from the analysis. Secondly, we did not evaluate whether deep learning can effectively assess the degree of coronary artery stenosis or analyze plaque burden and stability, both crucial CAD predictors. Additionally, the single-hospital dataset suggests a need for a larger, multi-center study to validate and expand our findings. Moreover, while our model has shown promising results, we acknowledge using a dataset with a high plaque prevalence might influence the model's performance, potentially creating bias towards identifying plaques. Validation of our algorithm on datasets with varied plaque prevalence will ensure its robustness and generalizability for broader clinical use. Finally, the study also highlighted a lower PPV in the identification of coronary artery plaques, primarily attributed to the lower PPV in identifying calcified plaques in the dLAD and R-PLB, and non-calcified plaques in the dLAD and R-PDA. To improve the detection of lipid-rich and calcified plaques in these smaller branches, further training of this automated system with a larger number of plaques is warranted. Such plaque characterization may assist in identifying vulnerable plaques (33) and categorizing associated risks (34,35).

### Conclusions

In conclusion, the deep learning algorithm demonstrates high efficacy in detecting coronary plaques, comparable to expert radiologists and superior to junior radiologists or interventional cardiologists. Moreover, the automated classification of coronary plaques using deep learning is time-saving and reliable, potentially aiding in triaging patients with acute coronary symptoms. This study lays the foundation for further development and utilization of deep learning in cardiac diagnostics and treatment, offering promising avenues for improving patient care.

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### Footnote

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*Conflicts of Interest:* All authors have completed the ICMJE uniform disclosure form (available at https://qims. amegroups.com/article/view/10.21037/qims-23-1513/coif). Z.S. and X.C. are employees of Philips Healthcare. C.Z. is an employee of Shukun (Beijing) Network Technology. The other authors have no conflicts of interest to declare.

*Ethical Statement:* The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. The study was conducted in accordance with the Declaration of Helsinki (as revised in 2013). The study was approved by the Institutional Review Board of Nanjing Drum Tower Hospital Clinical College of Nanjing University of Chinese Medicine (No. 2022-547-01), and individual consent for this retrospective analysis was waived.

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