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# Photon-counting detector computed tomography for the assessment of coronary stents and in-stent restenosis

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**BACKGROUND:** Photon-counting detector computed tomography (PCD-CT) offers enhanced spatial resolution and reduced blooming artefacts, potentially improving the evaluation of stented coronary vessels.

**AIMS:** This study aimed to assess the diagnostic performance of dual-source PCD-CT in detecting obstructive in-stent restenosis (ISR).

METHODS: We identified consecutive patients with prior coronary stent implantation who underwent clinically indicated coronary computed tomography angiography (CCTA) with PCD-CT and subsequent invasive coronary angiography within 90 days between 2023 and 2024. Obstructive ISR (≥50% diameter stenosis) was determined by visual assessment of CCTA and invasive quantitative coronary angiography (QCA) in a blinded fashion. The diagnostic performance of CCTA for ISR was compared with that of QCA.

**RESULTS:** A total of 283 stented lesions from 171 patients were included. Of these, only 3 lesions (1.1%) were deemed indeterminate by PCD-CT. Using invasive QCA as the reference standard, PCD-CT demonstrated a lesion-level sensitivity of 80.0%, specificity of 90.4%, positive predictive value (PPV) of 58.2%, negative predictive value (NPV) of 96.4%, and an overall diagnostic accuracy of 88.9% for detecting obstructive ISR. In a subgroup analysis according to the stent diameter (<3.00 mm [n=83] vs  $\geq$ 3.00 mm [n=108]), there were no significant differences in sensitivity (87.5% vs 86.7%; p=1.00), specificity (93.3% vs 92.5%; p=1.00), PPV (58.3% vs 65.0%; p=1.00), NPV (98.6% vs 97.7%; p=1.00), or overall diagnostic accuracy (92.8% vs 91.7%; p=1.00), respectively.

**CONCLUSIONS:** PCD-CT demonstrated good diagnostic performance for evaluating obstructive ISR using QCA as the reference standard, regardless of stent diameter.

**KEYWORDS:** computed tomography; coronary computed tomography angiography; diagnosis; in-stent restenosis; quantitative coronary angiography

oronary computed tomography angiography (CCTA) is increasingly used as a first-line diagnostic modality for patients with chest pain<sup>1,2</sup>. However, its effectiveness in assessing in-stent restenosis (ISR), particularly in smaller stents, remains limited due to artefacts from metallic struts and calcium<sup>3</sup>.

Photon-counting detector computed tomography (PCD-CT) is an advanced CT technology that may enhance the evaluation of stented lesions. Unlike conventional energy-integrating detector CT (EID-CT), PCD-CT utilises semiconductor detectors to directly convert photon energy into electronic signals, eliminating the need for septa required by scintillating detectors, thereby improving spatial resolution and reducing electronic noise and blooming artefacts<sup>4</sup>. These advantages have been demonstrated in clinical practice, with a recent large-scale study showing improved diagnostic performance of PCD-CT in evaluating patients with suspected coronary artery disease compared with EID-CT<sup>5</sup>.

Although preliminary studies have suggested the feasibility of PCD-CT in assessing coronary stents<sup>6,7</sup>, these were limited by small sample sizes and the inclusion of few patients with ISR. Therefore, this study sought to assess the diagnostic performance of PCD-CT in detecting obstructive ISR in routine clinical practice.

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#### **Methods**

#### STUDY DESIGN AND POPULATION

This was a retrospective, single-centre study that included all consecutive patients with prior coronary stent implantation who underwent clinically indicated CCTA with PCD-CT and subsequent invasive coronary angiography (ICA) within 90 days between January 2023 and December 2024 at St. Francis Hospital & Heart Center, Roslyn, NY, USA. Demographic and clinical characteristics were obtained through chart review. The study was conducted in accordance with the Declaration of Helsinki and approved by the Institutional Review Board of St. Francis Hospital & Heart Center. Informed consent was waived due to the minimal risk involved.

#### **CCTA IMAGE ACQUISITION AND ANALYSIS**

Details of PCD-CT image acquisition have been described previously<sup>5</sup>. PCD-CT was performed using the dual-source NAEOTOM Alpha (Siemens Healthineers). Collimation was 144×0.4 mm in standard mode and 120×0.2 mm in ultrahighresolution (UHR) mode, with reconstruction slice thicknesses of 0.4 mm and 0.2 mm, respectively. Scans were acquired at a tube voltage of 140 kVp per the manufacturer's recommendations<sup>5</sup>.

All patients received 0.4-0.8 mg of sublingual nitroglycerine as well as oral metoprolol (50-100 mg) and/or intravenous (IV) metoprolol (5-10 mg boluses every 5 minutes, up to

# Impact on daily practice

Photon-counting detector computed tomography (PCD-CT) demonstrated good diagnostic performance for evaluating in-stent restenosis in patients with prior coronary stents. PCD-CT can be used in patients with suspected obstructive coronary artery disease, whether in unstented or stented segments.

a maximum of 40 mg) as per institutional protocol to achieve a target heart rate of <65 bpm. A total volume of 60-100 mL of iso-osmolar, isotonic IV contrast was injected at a rate of 4-6 mL/s<sup>5</sup>. Standard reconstructions were performed using medium-smooth kernels and sharper kernels for all patients, which were available for the CT interpreting physician<sup>5</sup>.

CCTA images were analysed by cardiologists with Level 3 certification or equivalent expertise in cardiac CT, following the guidelines of the Society of Cardiovascular Computed Tomography<sup>8</sup>. Luminal stenosis was visually assessed and classified using the Coronary Artery Disease Reporting and Data System (CAD-RADS)<sup>9</sup>. Stented segments with ≥50% diameter stenosis (DS) were categorised as obstructive ISR. If the severity of stenosis could not be confidently determined because of suboptimal image quality or artefacts, the lesion was categorised as "indeterminate"<sup>5</sup>. To assess interobserver variability, a second imaging cardiologist independently analysed the first 60 stented segments in a blinded fashion.

#### **QUANTITATIVE CORONARY ANGIOGRAPHY**

All ICA was performed using standard techniques. In baseline ICA images, all stented lesions were analysed using quantitative coronary angiography (QCA) with dedicated software (Cath-QCA [TOMTEC]) by experienced analysts blinded to the CCTA findings. Overlapping stents were considered as a single lesion. Obstructive ISR was defined as a DS of  $\geq$ 50%, as assessed by QCA<sup>10</sup>.

#### STATISTICAL ANALYSIS

Continuous variables are presented as mean±standard deviation or median (interquartile range [IQR]), depending on the distribution tested by the Shapiro-Wilk test, and were compared using the t-test or the Mann-Whitney U test, respectively. Categorical variables are presented as number and proportion (%) and were compared using the chi-square test or Fisher's exact test, as appropriate.

The diagnostic performance of PCD-CT in detecting obstructive ISR was evaluated at both the stented-lesion and patient levels, excluding lesions with indeterminate readings, using QCA as the reference standard. Sensitivity, specificity, positive predictive value (PPV), negative predictive value (NPV), and overall diagnostic accuracy were derived from two-by-two contingency

#### **Abbreviations**

**CCTA** coronary computed tomography angiography

**CT** computed tomography

ICA invasive coronary angiography

ISR in-stent restenosis

**NPV** negative predictive value

**PCD-CT** photon-counting detector computed tomography

**PPV** positive predictive value

**QCA** quantitative coronary angiography

tables. The area under the receiver operating characteristic curve (AUC-ROC) was also calculated. To evaluate the impact of stent diameter on diagnostic performance, subgroup analyses were conducted among patients with available stent information, stratified by the mean stent diameter of the stented lesion (<3.00 mm vs ≥3.00 mm). Diagnostic performance metrics were compared between groups using Fisher's exact test or the chisquare test, and AUC-ROC comparisons were conducted using the DeLong test. Interobserver variability between imaging cardiologists in diagnosing ISR using PCD-CT was assessed using Cohen's kappa coefficient.

A two-sided p-value of less than 0.05 was considered statistically significant. All statistical analyses were performed using R, version 4.4.2 (R Foundation for Statistical Computing) and Stata 18.0 SE (StataCorp).

#### Results

#### STUDY POPULATION

A total of 398 consecutive patients with prior coronary stent implantation who underwent clinically indicated CCTA between 2023 and 2024 were identified. Among them, 184 patients subsequently underwent ICA within 90 days. After excluding 13 patients owing to missing or poor-quality coronary angiography images for QCA, the final study cohort consisted of 171 patients with 283 stented lesions (Supplementary Figure 1).

The mean age was  $67.5\pm9.8$  years, with 58 patients (33.9%) being female and 57 patients (33.3%) having diabetes mellitus **(Table 1)**. The majority of patients (81.3%) underwent PCD-CT in UHR mode. The median dose-length product

Table 1. Patient demographics and CT parameters.

	N=171
Demographics	
Age, years	67.5±9.8
Female	58 (33.9)
BMI, kg/m <sup>2</sup>	28.1 (25.2-31.2)
Hypertension	139 (81.3)
Hyperlipidaemia	139 (81.3)
Diabetes mellitus	57 (33.3)
Serum creatinine, mg/dL	0.97 (0.84-1.14)
Chronic kidney disease*	35 (20.5)
Atrial fibrillation	20 (11.7)
Prior MI	41 (24.0)
CABG	12 (7.0)
CT acquisition parameters	
UHR mode	139 (81.3)
Average heart rate, bpm	59.0 (55.0-64.0)
Total contrast, mL	95.0 (90.0-95.0)
Peak voltage, kVp	140 (140-140)
Dose-length product, mGy · cm	791.0 (677.5-905.0)
Total radiation, mSv	11.1 (9.5-12.7)

Data are presented as mean±standard deviation, median (interquartile range), or n (%). \*Defined as estimated glomerular filtration rate <60 mL/min/1.73 m². BMI: body mass index; bpm: beats per minute; CABG: coronary artery bypass graft surgery; CT: computed tomography; MI: myocardial infarction; UHR: ultrahigh-resolution

(DLP) was 791.0 mGy·cm (IQR 677.5-905.0 mGy·cm), and the total effective radiation dose from CCTA was 11.1 mSv (IQR 9.5-12.7 mSv). The indications for invasive coronary angiography are detailed in **Supplementary Table 1**.

#### STENTED SEGMENT CHARACTERISTICS

Stented lesions were most commonly located in the left anterior descending artery (44.5%), followed by the right coronary artery (32.5%) (**Table 2**). Among them, 3 stented lesions (1.1%) were classified as indeterminate by CCTA, and 55 (19.4%) were identified as having obstructive ISR. There was substantial interobserver agreement, with a Cohen's kappa of 0.74 (95% confidence interval [CI]: 0.47-1.00). By

Table 2. Characteristics of stented lesions.

Stented lesions	N=283
Stented vessel	
Left main	4 (1.4)
Left main trunk	1 (0.4)
Left main bifurcation	3 (1.1)
Left anterior descending artery	126 (44.5)
Left circumflex artery	61 (21.6)
Right coronary artery	92 (32.5)
Stented segment	
Proximal segment only	57 (20.1)
Including mid- to distal segments or branches	226 (79.9)
Stent information, per stented lesion* (n=19	3)
Number of stents	1 (1-2)
Mean diameter of stents, mm	3.00 (2.75-3.50)
<3.0 mm	83/193 (43.0)
≥3.0 mm	110/193 (57.0)
Total stent length, mm	23.0 (15.0-35.0)
Strut thickness <sup>†</sup>	
<100 μm	178 (94.2)
≥100 µm	11 (5.8)
CCTA findings	
Diameter stenosis	
0%	85 (30.0)
1-24%	72 (25.4)
25-49%	68 (24.0)
50-69%	28 (9.9)
70-99%	19 (6.7)
100%	8 (2.8)
Indeterminate	3 (1.1)
In-stent restenosis (≥50%)	55 (19.4)
QCA findings	
Diameter stenosis, %	26.7 (17.3-38.3)
In-stent restenosis (≥50%)	41 (14.5)

Data are presented as median (interquartile range), n/N (%), or n (%). \*Overlapped stents were considered to be one stented lesion. Among 283 stented lesions, no stent information was available for 90 lesions. \*Stent type and strut thickness data were available for 189 stented lesions. CCTA: coronary computed tomography angiography; QCA: quantitative coronary angiography

QCA, the median DS was 26.7% (IQR 17.3-38.3%), with 41 lesions (14.5%) classified as obstructive ISR (DS  $\geq$ 50%).

Among patients with available stent information (n=193), most stented lesions (70.0%) had a single stent (median: 1 [IQR 1-2]). The median stent diameter per stented lesion was 3.00 mm (IQR 2.75-3.50 mm), with a total stent length of 23.0 mm (IQR 15.0-35.0 mm) (Table 2). The majority (94.2%) of stents with available data had a strut thickness <100 µm. Stented lesions with a mean stent diameter of  $\geq$ 3.00 mm (n=110) were more frequently located in the right coronary artery than those with a mean stent diameter of <3.00 mm (n=83) (Table 3). The mean stent diameter of the smaller-stent group was 2.62 mm (IQR 2.50-2.75 mm), whereas that of larger-stent group was 3.33 mm (IQR 3.00-3.50 mm). The rate of indeterminate CCTA readings was similar between the two groups (<3.00 mm: 0% vs  $\geq$ 3.00 mm: 1.8%).

#### DIAGNOSTIC PERFORMANCE OF PCD-CT FOR ISR

Among stented lesions not classified as indeterminate (280 lesions from 168 patients), PCD-CT demonstrated good diagnostic performance for obstructive ISR at both the lesion and patient levels, using QCA as the reference standard (Figure 1). At the lesion level, PCD-CT achieved sensitivity of 80.0% (95% CI: 65.2-89.5), specificity of 90.4% (95% CI: 86.0-93.5), PPV of 58.2% (95% CI: 45.0-70.3), NPV of 96.4% (95% CI: 93.1-98.2) and overall diagnostic accuracy of 88.9% (95% CI: 84.7-92.1) with an AUC-ROC of 0.85 (95% CI: 0.79-0.92). At the patient level, PCD-CT demonstrated sensitivity of 77.1% (95% CI: 61.0-87.9), specificity of 88.0% (95% CI: 81.4-92.5), PPV of 62.8% (95% CI: 47.9-75.6), NPV of 93.6% (95% CI: 87.9-96.7),

and overall diagnostic accuracy of 85.7% (95% CI: 79.6-90.2) with an AUC-ROC of 0.83 (95% CI: 0.75-0.90).

The characteristics of false positive cases (23 stented lesions and 16 patients) are presented in **Supplementary Table 2** and **Supplementary Table 3**. The majority of these cases (78.3%) were graded as 50-69% stenosis by PCD-CT, suggesting a mild overestimation of lesion severity.

Subgroup analysis based on mean stent diameter (<3.00 mm [n=83] vs  $\geq$ 3.00 mm [n=108]) demonstrated comparable lesion-level diagnostic performance of PCD-CT for detecting obstructive ISR (AUC-ROC 0.90 [95% CI: 0.78-1.00] vs 0.89 [95% CI: 0.80-0.99], respectively; p=0.92), with no significant differences in any metrics (p=1.00) between the two stent size groups (**Figure 2**).

#### **Discussion**

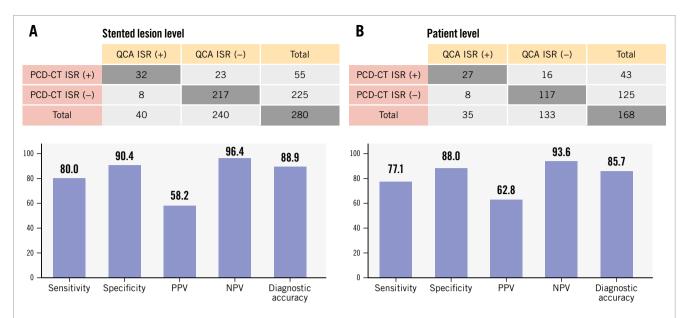
In this study, we assessed the diagnostic performance of PCD-CT in detecting obstructive ISR. Our key findings include the following: (1) among 171 consecutive patients with 283 stented lesions evaluated by PCD-CT followed by ICA, the vast majority of stented lesions were assessable by CCTA; (2) using QCA as the reference standard, PCD-CT demonstrated excellent diagnostic performance for obstructive ISR at both the stented-lesion and patient levels with high specificity and NPV; (3) the diagnostic accuracy of PCD-CT for obstructive ISR was similar for stented lesions with an average stent diameter of ≥3.00 mm and those <3.00 mm (Central illustration).

Advances in PCD-CT have enhanced spatial resolution while reducing blooming artefacts and image noise compared with EID-CT, offering significant advantages in evaluating

Table 3. Characteristics of stented lesions according to the mean stent diameter.

	Mean stent diameter		n value
	<3.0 mm (N=83)	≥3.0 mm (N=110)	<i>p</i> -value
Stented vessel			<0.001
Left main	0 (0)	3 (2.7)	
Left anterior descending artery	40 (48.2)	40 (36.4)	
Left circumflex artery	28 (33.7)	15 (13.6)	
Right coronary artery	15 (18.1)	52 (47.3)	
Stented segment			<0.001
Proximal segment only	5 (6.0)	37 (33.6)	
Including mid- to distal segments or branches	78 (94.0)	73 (66.4)	
Stent information			
Number of stents	1 (1-2)	1 (1-1)	0.02
Mean stent diameter, mm	2.62 (2.50-2.75)	3.33 (3.00-3.50)	< 0.001
Total stent length, mm	23.0 (15.0-39.5)	22.5 (15.0-33.0)	0.57
CCTA findings			
Indeterminate	0 (0)	2 (1.8)	NA
In-stent restenosis (≥50%)	12 (14.5)	20 (18.2)	0.58
QCA findings			
Diameter stenosis, %	27.6 (16.7-38.8)	24.7 (13.3-34.9)	0.33
In-stent restenosis (≥50%)	8 (9.6)	16 (14.5)	0.42

Data are presented as median (interquartile range) or n (%). CCTA: coronary computed tomography angiography; NA: not available; QCA: quantitative coronary angiography



**Figure 1.** Diagnostic performance of PCD-CT for obstructive ISR using QCA as the reference standard. Two-by-two contingency tables comparing the presence of obstructive ISR as assessed by PCD-CT and QCA are presented at (A) the stented-lesion level and (B) the patient level. Sensitivity, specificity, PPV, NPV, and overall diagnostic accuracy of PCD-CT are illustrated in bar graphs. ISR: in-stent restenosis; NPV: negative predictive value; PCD-CT: photon-counting detector computed tomography; PPV: positive predictive value; QCA: quantitative coronary angiography

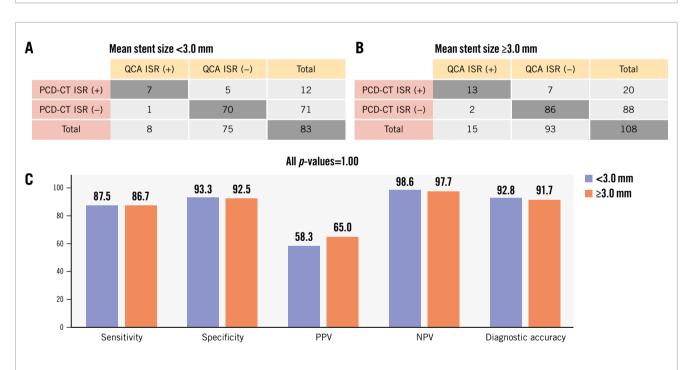


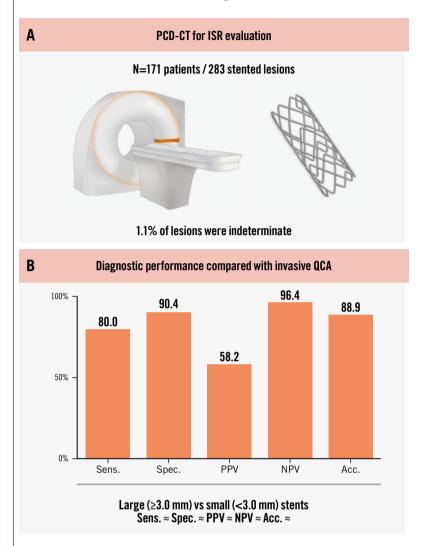
Figure 2. Stented lesion-level diagnostic performance of PCD-CT for ISR using QCA as the reference standard according to stent size. Two-by-two contingency tables comparing the presence of obstructive ISR as assessed by PCD-CT and QCA are presented based on mean stent diameter: (A) <3.00 mm and (B)  $\geq$ 3.00 mm. Sensitivity, specificity, PPV, NPV, and overall diagnostic accuracy of PCD-CT are depicted in bar graphs (C). ISR: in-stent restenosis; NPV: negative predictive value; PCD-CT: photon-counting detector computed tomography; PPV: positive predictive value; QCA: quantitative coronary angiography

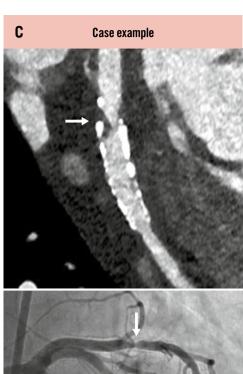
coronary artery disease<sup>4</sup>. In a recent study, the use of PCD-CT compared with EID-CT was associated with a lower referral rate for invasive coronary angiography and a higher rate of revascularisation among referred patients<sup>5</sup>. However,

the diagnostic performance of PCD-CT in evaluating ISR remains incompletely understood, with only a few small studies published to date<sup>6,7</sup>. Hagar et al analysed 44 stents from 18 patients undergoing CCTA with PCD-CT, reporting

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Assessment of obstructive ISR using PCD-CT.







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Among 283 stented lesions from 171 patients who underwent PCD-CT followed by invasive coronary angiography (A), PCD-CT showed good diagnostic performance for the detection of ISR using QCA as the reference standard (B). C) Case example. White arrows represent focal ISR in the proximal stent edge. acc: accuracy; ISR: in-stent restenosis; PCD-CT: photon-counting detector computed tomography; NPV: negative predictive value; PPV: positive predictive value; QCA: quantitative coronary angiography; sens: sensitivity; spec: specificity

a sensitivity of 100% and a specificity of 87.2-92.3%, compared with ICA<sup>7</sup>. However, this study primarily included patients undergoing CCTA prior to transcatheter aortic valve implantation assessment and included only five ISR cases. Qin et al reported similar findings among 25 stents in 12 patients, with a sensitivity of 75.0% and a specificity of 90.1%<sup>6</sup>. Beyond their small sample sizes, both studies relied on visual estimation of angiographic stenosis severity as the reference standard – a method known for its low reproducibility<sup>11</sup>.

The present study included 283 stented lesions from 171 consecutive patients who underwent clinically indicated

CCTA with PCD-CT in routine clinical practice. In this all-comers population, PCD-CT demonstrated excellent diagnostic performance for evaluating ISR, using QCA as the reference standard. Although the PPV remained modest at both the lesion and vessel levels, PCD-CT was particularly effective in ruling out obstructive ISR. This finding aligns with our recent study in patients without prior stents<sup>5</sup>, further supporting the potential role of PCD-CT in patients with suspected obstructive coronary artery disease, whether in unstented or stented segments.

Earlier evidence from EID-CT suggested that the diagnostic value of CCTA is limited in smaller stents<sup>3,12</sup>. The 2010

multisocietal Appropriate Use Criteria classified CCTA as "appropriate" only for asymptomatic patients with prior left main coronary stents ≥3.00 mm in diameter and "inappropriate" for stents <3.00 mm or of unknown diameter<sup>13</sup>. In contrast, the latest consensus from the Society of Cardiovascular Computed Tomography classified CCTA as "may be appropriate" for symptomatic patients with stents <3.00 mm<sup>14</sup>. In our analysis, we found that PCD-CT demonstrated consistent diagnostic performance for stents <3.00 mm compared with those ≥3.00 mm. These findings demonstrate that CCTA with PCD-CT is a viable option for evaluating patients with prior stent implantation, regardless of stent size (Figure 3, Figure 4) and suggest that further updates to these Appropriate Use Criteria may be warranted when PCD-CT is available.

#### Limitations

This study has several limitations. First, as a retrospective study, it was subject to inherent biases. In particular, not all

patients underwent ICA following CCTA, and the decision to proceed with ICA was based on clinical judgment rather than standardised criteria. This may have introduced verification bias and impacted the assessment of diagnostic performance, especially in underestimating false negatives among patients who were not referred for ICA. However, the NPV of CCTA is generally high, as demonstrated in the present study as well. Second, while this analysis represents the largest population studied to date, the number of ISR cases remains limited. Third, CCTA images were visually analysed, which may introduce reader subjectivity. However, these images were evaluated by the same group of experienced, boardcertified CT readers as part of routine clinical practice, and interobserver agreement among the readers was substantial. In addition, the primary objective of the present study was to evaluate the diagnostic performance of PCD-CT in real-world clinical practice, where visual assessment of CCTA images remains the standard approach. While quantitative analysis

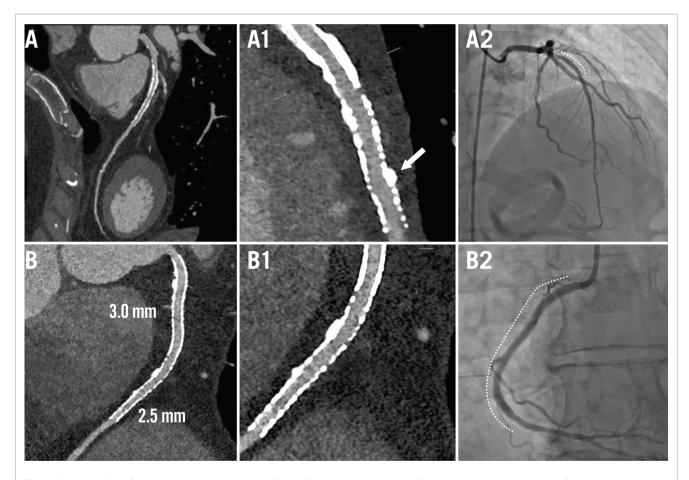


Figure 3. Examples of non-ISR cases. Case examples without ISR are presented using PCD-CT in UHR mode (UHR reconstruction: 0.2 mm slice, BV68 sharp kernel, quantum iterative reconstruction algorithm 4) and ICA. A) CCTA of the LAD in a patient with a prior DES implantation in the mid-LAD (Resolute [Medtronic]; 3.00×18 mm) and (B) CCTA of the RCA with two prior DES implants (Resolute Onyx [also Medtronic]; 3.0×38 mm proximally and 2.5×38 mm distally). A1) CCTA showed a patent stent in the mid-LAD with mild calcified plaque external to the stent (white arrow), confirmed by subsequent ICA (A2; dotted line). B1) CCTA demonstrated patent stents with clear visualisation of the stent lumen, confirmed by ICA (B2; dotted line). Dotted lines on the ICA indicate the stents. CCTA: coronary computed tomography angiography; DES: drugeluting stent; ICA: invasive coronary angiography; ISR: in-stent restenosis; LAD: left anterior descending artery; PCD-CT: photon-counting detector computed tomography; RCA: right coronary artery; UHR: ultrahigh-resolution

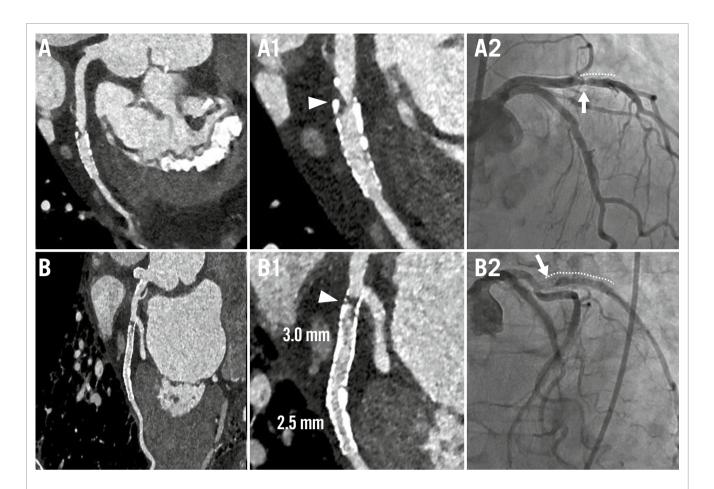


Figure 4. Examples of ISR cases. Case examples of ISR are presented using PCD-CT in UHR mode (UHR reconstruction: 0.2 mm slice, BV68 sharp kernel, quantum iterative reconstruction algorithm 4) and ICA. A) CCTA of a patient with a prior DES (type and size unknown) in the first obtuse marginal branch, who presented with chest pain. A1) CCTA revealed obstructive ISR at the proximal stent edge (white arrowhead), confirmed by ICA, which showed a 59.2% diameter stenosis by QCA (A2; white arrow). B) CCTA of a patient with two DES implants (XIENCE [Abbott]; 2.75×12 mm proximally and 2.50×12 mm distally) in the first obtuse marginal branch. B1) CCTA demonstrated obstructive ISR at the proximal stent edge (white arrowhead), confirmed by ICA, which revealed subtotal occlusion (B2; white arrow). Dotted lines on the ICA indicate the stents. CCTA: coronary computed tomography angiography; DES: drug-eluting stent; ICA: invasive coronary angiography; ISR: in-stent restenosis; PCD-CT: photon-counting detector computed tomography; QCA: quantitative coronary angiography; UHR: ultrahigh-resolution

of PCD-CT images could enhance objectivity, its application to in-stent assessment remains to be validated and may warrant future investigation. In contrast, ICA images, which served as the reference standard, were objectively evaluated using blinded QCA analysis. Fourth, angiographic DS was assessed using two-dimensional QCA, as three-dimensional QCA was not available. In addition, intravascular imaging was available in only a limited number of patients. Fifth, although institutional policy recommends UHR mode for patients with prior stents, PCD-CT was performed in both UHR (81.3%) and conventional modes (18.7%) due to high heart rates, arrhythmias, and the inability of patients to hold their breath for the scan time required for UHR, which represents a spectrum of real-life clinical patients. Sixth, stent information was unavailable in approximately one-third of cases, and among those with available data, the majority had a strut thickness <100 µm. Therefore, the present findings may not be applicable to stents with a strut thickness ≥100 µm. Finally, as a single-centre study conducted in a high-volume cardiac PCD-CT programme, the results may not be generalisable to institutions with different scanning protocols or practice patterns. Despite these limitations, this study provides the largest real-world evaluation of PCD-CT for ISR, offering valuable early insights into its diagnostic performance and clinical utility. A future study will evaluate the feasibility of using PCD-CT to assess ISR mechanisms and tissue characteristics, using angiography and intravascular imaging as reference standards.

# **Conclusions**

In this large-scale, single-centre study, PCD-CT demonstrated good diagnostic performance for evaluating obstructive ISR, particularly with high specificity and NPV, independent of stent diameter. These findings suggest that PCD-CT

technology may improve the management of patients with prior stents. Further multicentre studies with larger sample sizes are warranted to validate these findings.

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#### Conflict of interest statement

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# References

 Writing Committee Members; Gulati M, Levy PD, Mukherjee D, Amsterdam E, Bhatt DL, Birtcher KK, Blankstein R, Boyd J, Bullock-Palmer RP, Conejo T, Diercks DB, Gentile F, Greenwood JP, Hess EP, Hollenberg SM, Jaber WA, Jneid H, Joglar JA, Morrow DA, O'Connor RE,

- Ross MA, Shaw LJ. 2021 AHA/ACC/ASE/CHEST/SAEM/SCCT/SCMR Guideline for the Evaluation and Diagnosis of Chest Pain: A Report of the American College of Cardiology/American Heart Association Joint Committee on Clinical Practice Guidelines. *J Am Coll Cardiol*. 2021;78:e187-285.
- 2. Vrints C, Andreotti F, Koskinas KC, Rossello X, Adamo M, Ainslie J, Banning AP, Budaj A, Buechel RR, Chiariello GA, Chieffo A, Christodorescu RM, Deaton C, Doenst T, Jones HW, Kunadian V, Mehilli J, Milojevic M, Piek JJ, Pugliese F, Rubboli A, Semb AG, Senior R, Ten Berg JM, Van Belle E, Van Craenenbroeck EM, Vidal-Perez R, Winther S; ESC Scientific Document Group. 2024 ESC Guidelines for the management of chronic coronary syndromes. Eur Heart J. 2024;45:3415-537.
- 3. Narula J, Chandrashekhar Y, Ahmadi A, Abbara S, Berman DS, Blankstein R, Leipsic J, Newby D, Nicol ED, Nieman K, Shaw L, Villines TC, Williams M, Hecht HS. SCCT 2021 Expert Consensus Document on Coronary Computed Tomographic Angiography: A Report of the Society of Cardiovascular Computed Tomography. J Cardiovasc Comput Tomogr. 2021;15:192-217.
- Esquivel A, Ferrero A, Mileto A, Baffour F, Horst K, Rajiah PS, Inoue A, Leng S, McCollough C, Fletcher JG. Photon-Counting Detector CT: Key Points Radiologists Should Know. Korean J Radiol. 2022;23:854-65.
- 5. Sakai K, Shin D, Singh M, Malik S, Dakroub A, Sami Z, Weber J, Cao JJ, Parikh R, Chen L, Sosa F, Cohen DJ, Moses JW, Shlofmitz RA, Collet C, Shlofmitz E, Jeremias A, Khalique OK, Ali ZA. Diagnostic Performance and Clinical Impact of Photon-Counting Detector Computed Tomography in Coronary Artery Disease. J Am Coll Cardiol. 2025;85:339-48.
- Qin L, Zhou S, Dong H, Li J, Zhang R, Yang C, Liu P, Xu Z, Yan F, Yang W. Improvement of coronary stent visualization using ultra-high-resolution photon-counting detector CT. Eur Radiol. 2024;34:6568-77.
- 7. Hagar MT, Soschynski M, Saffar R, Molina-Fuentes MF, Weiss J, Rau A, Schuppert C, Ruile P, Faby S, Schibilsky D, von Zur Muehlen C, Schlett CL, Bamberg F, Krauss T. Ultra-high-resolution photon-counting detector CT in evaluating coronary stent patency: a comparison to invasive coronary angiography. Eur Radiol. 2024;34:4273-83.
- Leipsic J, Abbara S, Achenbach S, Cury R, Earls JP, Mancini GJ, Nieman K, Pontone G, Raff GL. SCCT guidelines for the interpretation and reporting of coronary CT angiography: a report of the Society of Cardiovascular Computed Tomography Guidelines Committee. J Cardiovasc Comput Tomogr. 2014;8:342-58.
- 9. Cury RC, Abbara S, Achenbach S, Agatston A, Berman DS, Budoff MJ, Dill KE, Jacobs JE, Maroules CD, Rubin GD, Rybicki FJ, Schoepf UJ, Shaw LJ, Stillman AE, White CS, Woodard PK, Leipsic JA. CAD-RADS™. Coronary Artery Disease Reporting and Data System: An Expert Consensus Document of the Society of Cardiovascular Computed Tomography (SCCT), the American College of Radiology (ACR) and the North American Society for Cardiovascular Imaging (NASCI). Endorsed by the American College of Cardiology. J Am Coll Radiol. 2016;13:1458-66.e9.
- Alfonso F, Coughlan JJ, Giacoppo D, Kastrati A, Byrne RA. Management of in-stent restenosis. EuroIntervention. 2022;18:e103-23.
- Zir LM, Miller SW, Dinsmore RE, Gilbert JP, Harthorne JW. Interobserver variability in coronary angiography. Circulation. 1976;53:627-32.
- 12. Pontone G, Rossi A, Guglielmo M, Dweck MR, Gaemperli O, Nieman K, Pugliese F, Maurovich-Horvat P, Gimelli A, Cosyns B, Achenbach S. Clinical applications of cardiac computed tomography: a consensus paper of the European Association of Cardiovascular Imaging-part I. Eur Heart J Cardiovasc Imaging. 2022;23:299-314.
- 13. Taylor AJ, Cerqueira M, Hodgson JM, Mark D, Min J, O'Gara P, Rubin GD; American College of Cardiology Foundation Appropriate Use Criteria Task Force; Society of Cardiovascular Computed Tomography; American College of Radiology; American Heart Association; American Society of Echocardiography; American Society of Nuclear Cardiology; North American Society for Cardiovascular Imaging; Society for Cardiovascular Angiography and Interventions; Society for Cardiovascular Magnetic Resonance. ACCF/SCCT/ACR/AHA/ASE/ASNC/NASCI/SCAI/SCMR 2010 Appropriate Use Criteria for Cardiology Foundation Appropriate Use Criteria Task Force, the Society of Cardiovascular Computed Tomography, the American College of Radiology, the American Heart

Association, the American Society of Echocardiography, the American Society of Nuclear Cardiology, the North American Society for Cardiovascular Imaging, the Society for Cardiovascular Angiography and Interventions, and the Society for Cardiovascular Magnetic Resonance. *J Cardiovasc Comput Tomogr.* 2010;4:407.e1-33.

14. Dai T, Wang JR, Hu PF. Diagnostic performance of computed tomography angiography in the detection of coronary artery in-stent restenosis: evidence from an updated meta-analysis. Eur Radiol. 2018;28:1373-82.

## Supplementary data

**Supplementary Table 1.** Indications for invasive coronary angiography.

**Supplementary Table 2.** Patient demographics and CT parameters in false positive cases.

**Supplementary Table 3.** Characteristics of stented lesions in false positive cases.

**Supplementary Figure 1.** Study flowchart. **Data sharing statement** 

The supplementary data are published online at: https://eurointervention.pcronline.com/doi/10.4244/EIJ-D-25-00257



# Supplementary data

Supplementary Table 1. Indications for invasive coronary angiography.

	N = 171
Obstructive lesions in stented segments	43 (25.1)
Indeterminate lesions in stented segments	3 (1.8)
Obstructive lesions in non-stented segments	107 (62.6)
Moderate 50-69%	58 (33.9)
Severe $\geq 70\%$	49 (28.2)
No obstructive lesions	18 (10.5)
Cardiomyopathy or heart failure	7 (4.1)
Persistent stable angina or anginal equivalent	6 (3.5)
Unstable angina	4 (2.3)
STEMI*	1 (0.6)

Data are presented in n (%).
\*Culprit lesion located outside of the stented segment.
STEMI, ST-segment elevation myocardial infarction

Supplementary Table 2. Patient demographics and CT parameters in false positive cases.

	N = 16
Demographics	
Age, year	$67.3 \pm 9.6$
Female	4 (25.0)
BMI, kg/m <sup>2</sup>	$28.5 \pm 4.3$
Hypertension	11 (68.8)
Hyperlipidemia	13 (81.3)
Diabetes mellitus	6 (37.5)
Serum creatinine, mg/dL	$1.1\pm0.2$
Chronic kidney disease*	4 (25.0)
Atrial fibrillation	1 (6.3)
Prior MI	5 (31.3)
CABG	1 (6.3)
CT acquisition parameters	
UHR mode	12 (75.0)
Average heart rate, bpm	$63.0 \pm 7.6$
Total contrast, mL	95.0 (90.0, 95.0)
Peak voltage, kVP	140 (140, 140)
Dose-length product, mGy·cm	706.0 (678.5, 825.0)
Total radiation, mSv	9.88 (9.5, 11.6)

Data are presented as mean  $\pm$  standard deviation, median (interquartile range), or number (%).

BMI, body mass index; BPM, beat per minute; CABG, coronary artery bypass graft surgery; CT, computed tomography; MI, myocardial infarction; UHR, ultra-high-resolution.

<sup>\*</sup> Defined as estimated glomerular filtration rate < 60 ml/min/1.73m<sup>2</sup>

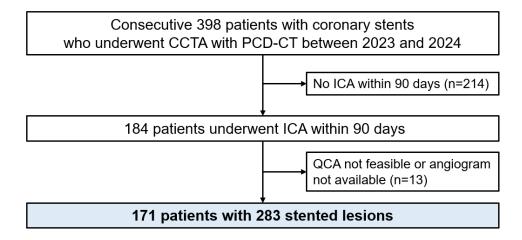
Supplementary Table 3. Characteristics of stented lesions in false positive cases.

Stented Lesions	N = 23
Stented vessel	
Left main	0 (0.0)
Left anterior descending artery	12 (52.2%)
Left circumflex artery	4 (17.4%)
Right coronary artery	7 (30.4%)
Stented segment	
Proximal segment only	6 (26.1%)
Including mid to distal segments or branches	17 (73.9%)
Stent information, per stented lesion* (n=12)	
Number of stents	1 (1, 1)
Mean diameter of stents, mm	$2.9 \pm 0.5$
< 3.0 mm	5 (41.7%)
≥ 3.0 mm	7 (58.3%)
Total stent length, mm	18.0 (15.0, 31.5)
Strut thickness	
< 100 μm	11 (91.7%)
≥ 100 µm	1 (8.3%)
CCTA findings	
Diameter stenosis	
50-69%	18 (78.26%)
70-99%	3 (13.04%)
100%	2 (8.70%)
QCA findings	
Diameter stenosis, %	$25.5 \pm 14.3$
In-stent restenosis ( $\geq 50\%$ )	23 (100.0%)

Data are presented as median (interquartile range), mean  $\pm$  standard deviation, or number (%).

CCTA, coronary computed tomography angiography; QCA, quantitative coronary angiography.

<sup>\*</sup> Overlapped stents were considered to be one stented lesion. Stent information was available for 12 stented lesions.



# Supplementary Figure 1. Study flowchart.

CCTA, coronary computed tomography angiography; ICA, invasive coronary angiography; PCD-CT, photon counting detector computed tomography; QCA quantitative coronary angiography

# **Data sharing statement**

The data that support the findings of this study are not openly available but are available from the corresponding author upon reasonable request.