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# Correlation between LCX-QFR and clinical outcomes following a single-stent strategy for left main bifurcation lesions

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**Objective:** The aim of this study was to investigate the quantitative flow ratio (QFR) outcomes in the left circumflex artery (LCX) following the placement of a crossover stent from the left main coronary artery (LM) to the left anterior descending artery (LAD) in LM bifurcation lesions. In addition, we sought to assess the relationship between these QFR results and clinical prognoses.

**Background:** The treatment approach for LM bifurcation lesions remains a topic of debate, with the LM-LAD single-stent technique being one possible option. QFR, a fractional flow reserve calculation method derived from angiography that does not require pressure guide wires, could serve as an alternative functional assessment of the LCX. This study aims to evaluate the clinical outcomes of postoperative LCX by utilizing QFR measurements, addressing a current gap in the relevant literature on this topic.

**Methods:** This study was a retrospective, single-center analysis of patients with LM bifurcation lesions who underwent percutaneous coronary intervention (PCI) guided by intravascular ultrasound. QFR values were derived from angiographies. The primary endpoint was the 1-year rate of major adverse cardiac events, defined as a composite of cardiovascular death, target bifurcation-related myocardial infarction (MI), or target bifurcation revascularization. The secondary clinical endpoint was defined as the persistence or recurrence of angina pectoris after PCI.

**Results:** We analyzed 91 patients from a total of 180 who were screened for LM bifurcation lesions. All patients completed the 1-year follow-up. The pre- and post-PCI QFR values were  $0.89 \pm 0.09$  and  $0.86 \pm 0.11$ , respectively. Subgroup analysis showed that 74 patients were in the postoperative QFR  $\geq 0.80$  group, whereas 17 patients were in the QFR <0.80 group. In addition, 32 patients had a  $\Delta$ QFR  $\geq 0$ , and 58 patients had a  $\Delta$ QFR <0. Nine patients (9.9%) achieved the primary endpoint, including one patient with non-ST elevation myocardial infarction who received revascularization in both the LM-LAD and LCX arteries. In addition, nine patients (9.9%) reported no substantial improvement in their chest pain symptoms. Post-LCX-QFR <0.8 was associated with a higher 1-year incidence of cardiovascular death or MI (P = 0.036).  $\Delta$ QFR proved to be a robust predictor of the 1-year incidence of the primary endpoint, with an incidence of 15.3% in the  $\Delta$ QFR  $\geq 0$  group compared to 0% in the  $\Delta$ QFR <0 group (area under the curve: 0.822; 95% CI: 0.728–0.895, P < 0.001), especially when  $\Delta$ QFR  $\leq -0.03$ .

**Conclusions:** After the LM-LAD single-stent strategy for LM bifurcation lesions, a  $\Delta$ QFR of LCX  $\leq$ -0.03 was associated with a higher risk of 1-year main adverse cardiac events, indicating the superior prognostic value of the post-PCI physiological assessment.

KEYWORDS

left main coronary artery disease, quantitative flow ratio fraction, percutaneous coronary intervention, main adverse cardiac events, left main bifurcation lesions

# 1 Introduction

Coronary artery bifurcation lesions (CBLs) are common and carry a higher risk of main cardiac events and restenosis after percutaneous coronary intervention (PCI) compared to lesions alone (1). The optimal treatment strategy for coronary bifurcation anatomy remains a subject of debate. CBLs with extensive atherosclerosis involving a large and significantly diseased side branch (SB) may benefit from an elective two-stent bifurcation technique (2). However, trials involving bifurcated lesions have shown that a systematic dual drug-eluting stent (DES) strategy offers no advantage, and in fact, a more sophisticated approach may lead to more severe long-term mortality (3). A provisional single-stent strategy may be superior to the two-stent method and is considered a feasible treatment option for left main coronary artery (LM) bifurcation lesions, although there are significant differences in the predictors, functional significance, and luminal changes for compromised SB after main vessel dilation (4, 5).

According to the "provisional" strategy, SB intervention should be considered only when the SB result is suboptimal. However, a suboptimal result for the left circumflex artery (LCX) ostium has not been standardized. The utilization of fractional flow reserve (FFR) to guide interventions in the LCX has been proposed as a means to enhance clinical outcomes by reducing unnecessary side branch interventions. However, it should be noted that FFR-guided LCX intervention may pose technical challenges, such as difficulty accessing side branches through stent struts, and its superiority over angiography-guided LCX intervention remains unclear (6, 7).

Therefore, we conducted a comprehensive analysis of the alterations in LCX-quantitative flow ratio (QFR) values following intravascular ultrasound (IVUS)-guided LM to left anterior descending artery (LAD) crossover stenting, and explored their correlation with clinical outcomes.

# 2 Methods

#### 2.1 Population

This was a single-center, retrospective, observational study of patients with significant *de novo* LM stem coronary artery disease, defined as angiographic stenosis >50%, who presented with distal bifurcation lesions classified as Medina (1,1,0/1,1,1) and had mild-to-moderate LCX disease with angiographic stenosis <70%. All the DESs were implanted from the LM to the LAD. The study was

conducted at Peking University Third Hospital between January 2019 and December 2023. Adults (≥18 years old) who were planning to undergo PCI treatment based on coronary angiography (CAG) findings were eligible for inclusion.

The main exclusion criteria encompassed: (1) use of a systematic dual-stent strategy or provisional intervention in the LCX ostium, (2) presence of acute myocardial infarction (MI) or severe left ventricular dysfunction, (3) history of bypassed vessels, (4) diffuse LCX disease extending >7 mm from the LCX ostium, (5) an LCX-angiographic diameter <2.5 mm, and (6) other conditions that prevented post-PCI QFR analysis.

## 2.2 PCI procedure

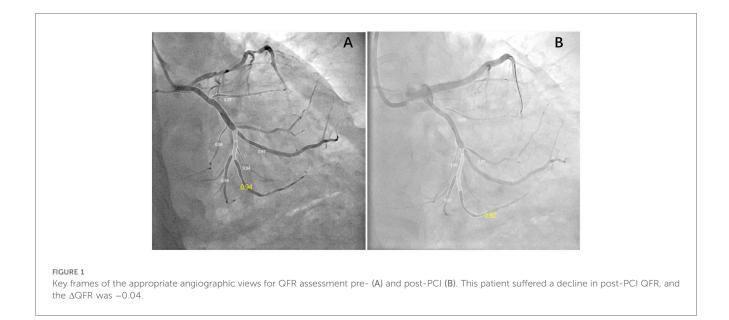
All patients received a loading dose of antiplatelet treatment before the procedure. All patients underwent DES implantation from the LM to the LAD via a simple crossover intervention technique, with stent diameter and length selected by expert operators based on the reference vessel diameter, culprit lesion length, plaque burden (PB), and partly IVUS-based measurements. During PCI, the type of DES employed, stenting techniques, utilization of intravascular ultrasound, and application of poststent adjunctive balloon inflation were left to the discretion of the operating physicians. In cases where the LCX flow restriction occurred during LM-to-LAD stent implantation, the surgeon used balloon dilation to improve it.

All patients received standard dual antiplatelet therapy and other secondary prevention drugs for coronary heart disease after discharge. During follow-up, drugs and dosages were adjusted according to each patient's blood pressure, heart rate, clinical symptoms, comorbidities, and the operator's discretion.

### 2.3 QFR and rQFR analysis

A certified operator, qualified to conduct QFR analysis, performed all measurements using QFR software [AngioPlus 2.0.1.0 and PStation (Pulse Medical Technology, Shanghai, China)], while a second, more experienced operator reviewed and made corrections to all analyses as needed. The results provided by the second operator were utilized for the analyses.

Eligible patients undergoing LM-LAD stenting were enrolled. Angiographic images with optimal contrast opacification and clear visualization of all coronary artery branches, including the ostium of the LCX, were selected. QFR analysis was performed



in a blinded fashion. Independent operators, blinded to each other's analyses, selected appropriate angiographic views and keyframes for QFR assessment to minimize interobserver variability. This approach ensured an objective and unbiased assessment of LCX ostial stenosis (Figure 1). QFR measurements, including hemodynamic parameters and FFR-derived indices, were acquired. These parameters included blood flow velocity, microvascular resistance, QFR, resting QFR (rQFR), and the difference between pre- and post-PCI QFR ( $\Delta$ QFR). These parameters were employed to evaluate the impact of LM-LAD stenting on LCX hemodynamics.

# 2.4 IVUS imaging

IVUS was performed either pre- and post-PCI or only postoperatively, according to the surgeon's experience with LM-LAD stent implantation. After the appropriate selection of a guiding catheter based on the target vessel's lesion characteristics, the IVUS catheter was advanced distal to the lesion, and a pullback was performed to acquire intravascular images. In conjunction with CAG findings, quantitative IVUS analysis was performed on the proximal and distal segments of the LM, LAD, and partly LCX arteries. The measured parameters included lumen diameter, lumen area, plaque burden, and lesion length. Poststenting IVUS was performed within the LM-LAD segment to evaluate stent apposition, stent expansion (minimum stent area), and lumen dimensions (diameter and area). The decision to perform an IVUS assessment of the LCX ostium was made based on intraoperative findings and clinical judgment.

#### 2.5 Clinical outcomes

The primary endpoint was the 1-year rate of major adverse cardiac events (MACE), which was defined as a composite of cardiovascular death, target bifurcation-related MI, or target bifurcation revascularization. All deaths were assumed to be of cardiac origin unless a clear non-cardiac cause was reported. Target bifurcation-related MI was defined as any MI with angiographic evidence confirming that the culprit lesion corresponded to the previously treated target bifurcation. Target bifurcation revascularization was defined as repeat percutaneous intervention or bypass surgery for culprit target lesions exhibiting restenosis or occlusion at the LCX ostium or within 5 mm of the distal or proximal edge of the implanted stent. The secondary clinical endpoint was performed at 2, 4, 6, 8, 10, and 12 months via outpatient clinical visits or telephone interviews.

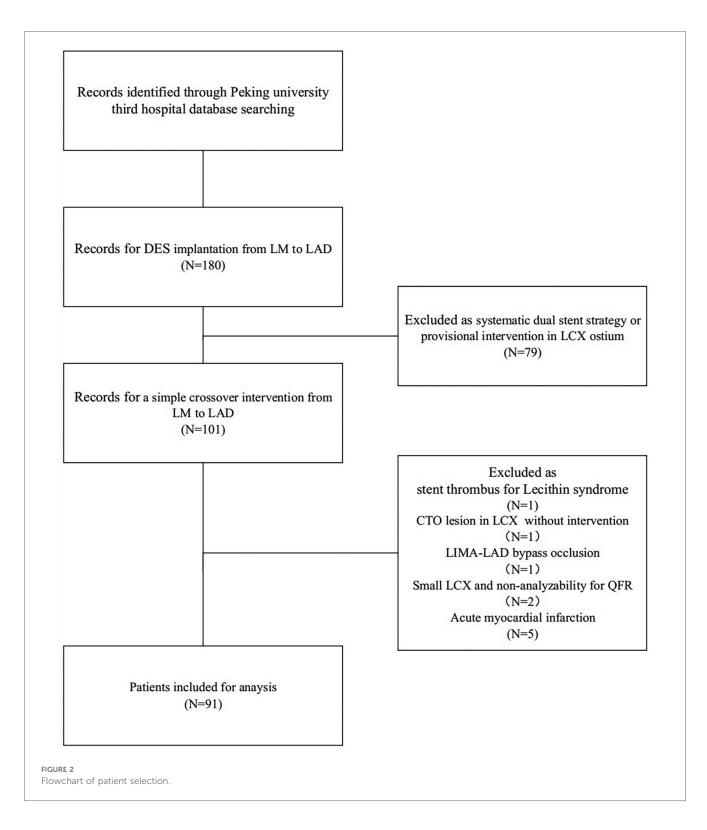
### 2.6 Statistical analysis

Data were processed using SPSS 21.0 (SPSS, Inc., an IBM Company), GraphPad Prism 8.0.0 (GraphPad Software, San Diego, CA, USA), MedCalc 22.014 (MedCalc Software Ltd., Belgium), and STATA 18.0 (StataCorp, USA). Intergroup comparisons for normally distributed data were made using *t*-tests or one-way ANOVA. The counting data rate (%) or constituent ratio was expressed, and Pearson's chi-square test was used for comparison between groups. A receiver operating characteristic (ROC) curve analysis was performed to determine the optimal cutoff values for parameters that independently forecast  $\Delta$ QFR following stenting in the LCX. The Kaplan–Meier method was employed to determine the time to clinical endpoints and survival rates. A forest plot was generated to display independent predictors of postoperative QFR. All *P*-values were two-tailed, and *P* < 0.05 was considered statistically significant.

# **3** Results

### 3.1 Study population

A comprehensive review of patients who underwent LM-LAD DES implantation at Peking University Third Hospital between



2019 and 2024 identified a total of 180 patients. Among them, 79 patients were treated with a dual-stent strategy or underwent provisional intervention at the LCX ostium and were therefore excluded. This left 101 patients who received a straightforward crossover intervention from the LM to LAD. Ultimately, 91 patients met the inclusion criteria (refer to Figure 2). Of these 91 patients, IVUS was utilized both preoperatively and postoperatively in 53 patients; specifically, 5 patients underwent

pre-PCI IVUS evaluation to guide treatment strategy, while 33 patients received IVUS assessment conducted exclusively during postdilatation. Notably, only 14 patients had preoperative IVUS measurements taken from the LCX to the LM.

The pre-PCI LCX-QFR was  $0.89 \pm 0.09$  among the 91 patients. Following PCI, a subgroup analysis was performed to evaluate whether the LCX-QFR exceeded 0.80 and to examine changes in the LCX-QFR values. This analysis revealed that 74 patients fell

Variables	Total (N = 91 patients)	Post- QFR ≥ 0.80 (N = 74 patients)	Post- QFR<0.80 (N = 17 patients)	<i>P</i> -value	$\Delta QFR \ge 0$ (N = 32 patients)	$\Delta QFR < 0$ (N = 59 patients)	<i>P</i> -value
Age (years)	$68 \pm 9$	$67 \pm 9$	$70 \pm 9$	0.389	$67 \pm 9$	$69 \pm 9$	0.340
Male patients, n (%)	67 (73.6)	54 (73.0)	13 (76.5)	0.516	22 (68.8)	45 (76.3)	0.296
Cardiovascular risk factors							
Hypertension, n (%)	68 (74.7)	53 (71.6)	15 (88.2)	0.170	22 (68.8)	46 (78.0)	0.265
Diabetes mellitus, n (%)	33 (36.3)	26 (35.1)	7 (41.2)	0.450	7 (21.9)	26 (44.1)	0.045*
Hypercholesterolemia, n (%)	39 (42.9)	33 (44.6)	6 (35.3)	0.384	14 (43.8)	25 (42.4)	0.500
Smoker, n (%)	44 (48.4)	38 (51.4)	6 (35.3)	0.407	17 (53.1)	27 (45.8)	0.283
Clinical presentation							
SCAD	7 (7.7)	5 (6.8)	2 (11.8)	0.389	0 (0)	7 (11.9)	0.042*
UAP	84 (92.3)	69 (93.2)	15 (88.2)	0.389	32 (100)	52 (88.1)	
Creatinine, µmol/L	$83 \pm 21$	$82 \pm 21$	86 ± 20	0.435	$78 \pm 17$	85 ± 23	0.136
HbA1c (%)	$6.5 \pm 1.1$	$6.5 \pm 1.1$	$6.8 \pm 1.2$	0.439	$6.3 \pm 0.8$	$6.7 \pm 1.2$	0.117
LDL-C (mmol/L)	$2.12\pm0.81$	$2.15\pm0.81$	$2.02\pm0.85$	0.574	$2.06\pm0.89$	$2.20 \pm 0.77$	0.583
Left ventricular ejection fraction (%)	$65 \pm 10$	$66 \pm 10$	$63 \pm 12$	0.234	$66 \pm 11$	65 ± 9	0.444
Body mass index (kg m <sup>2</sup> )	$22.9\pm6.7$	$23.2\pm6.33$	$21.8\pm8.3$	0.441	$23.0\pm6.5$	22.3 ± 6.9	0.924

TABLE 1 Clinical characteristics of patients with post-QFR  $\geq$ 0.80 or <0.80 and  $\triangle$ QFR  $\geq$ 0 or <0.

UAP, unstable angina pectoris; SCAD, stable coronary artery disease; LDL-C, low-density lipoprotein cholesterol. \**P* < 0.05.

into the postoperative QFR  $\geq 0.80$  category, while 17 patients were classified in the QFR <0.80 group. In addition, 32 patients had a  $\Delta$ QFR  $\geq 0$ , and 58 patients had a  $\Delta$ QFR <0. The baseline clinical, angiographic, procedural, and quantitative coronary angiography (QCA) characteristics of the patients in these subgroups are detailed in Tables 1, 2. A comparison indicated that patients with a  $\Delta$ QFR <0 had a higher prevalence of diabetes mellitus (P = 0.042) and a greater incidence of stable coronary artery disease (SCAD) (P = 0.045). Furthermore, patients in the postoperative QFR  $\geq 0.80$  subgroup exhibited elevated pre-PCI QFR levels (P < 0.01) and a larger LM area following stenting (P = 0.016).

### 3.2 Clinical follow-up

Complete 1-year follow-up data were available for all 91 patients, and the results are presented in Table 3 and Figures 3, 4. Nine patients (9.9%) achieved the primary endpoint, which included one individual who experienced a non-ST elevation myocardial infarction (NSTEMI) outside of the hospital at 11 months. This patient subsequently underwent revascularization involving the LM-LAD and LCX arteries. In addition, nine patients (9.9%) who did not meet the primary endpoint during the 1-year follow-up reported no substantial improvement in their chest pain symptoms and were prescribed antianginal medication in an outpatient setting.

A post-LCX-QFR <0.8 was associated with a higher 1-year incidence of cardiovascular death or MI (P = 0.036). However, no statistically significant difference was observed in the primary endpoint compared with the QFR  $\geq 0.8$  group (P = 0.235). All nine patients with persistent angina pectoris were in the  $\Delta$ QFR <0 group (P = 0.020).

 $\Delta$ QFR proved to be a robust predictor of the 1-year incidence of the primary endpoint, with an incidence of 15.3% in the  $\Delta$ QFR  $\geq 0$  group compared to 0% in the  $\Delta$ QFR <0 group [area under the curve (AUC): 0.822; 95% CI: 0.728-0.895, P < 0.001]. The optimal cutoff value of  $\Delta$ QFR for predicting adverse events was  $\leq -0.03$ . Using this cutoff, the Kaplan-Meier survival analysis showed that the  $\Delta$ QFR  $\leq -0.03$  group had a significantly increased risk of experiencing the 1-year primary endpoint compared to the  $\Delta$ QFR >-0.03 group (P = 0.003). However, no statistically significant differences were observed in 1-year cardiovascular death (P = 0.248), target bifurcation-related MI (P = 0.248), revascularization in LM-LAD (P = 0.831), revascularization in LCX (P = 0.082), or revascularization in both arteries (P = 0.100) (Figure 3).

## 3.3 Predictors for low QFR

The findings from the multivariate regression analysis were illustrated in a forest plot, indicating that the IVUS evaluation results of pre/post-LM-LAD are not significant independent predictors of a postoperative QFR <0.8 or  $\Delta$ QFR  $\leq$ -0.03 (Figure 5). However, a larger post-LAD minus lumen area (MLA) appears to be likely related to a decline in QFR (*P* = 0.064).

# 4 Discussion

LM-CBLs are rarely focal. IVUS data demonstrate that, in over 90% of cases, plaques extend from the LM into the proximal LAD. Furthermore, in 66% of cases, these plaques reach the proximal LCX, and in 62% of cases, they involve both the LAD and LCX branches. The provisional single-stent strategy has emerged as a TABLE 2 Lesion and procedural characteristics of the study population.

Variables	Total (N = 91 patients)	Post- QFR $\geq$ 0.80 ( $N = 74$ patients)	Post- QFR < 0.80 (N = 17 patients)	<i>P</i> -value	$\Delta QFR \ge 0$ ( $N = 32$ patients)	∆QFR < 0 ( <i>N</i> = 59 patients)	<i>P</i> -value
Baseline							
LM							
MLA (mm <sup>2</sup> )	$6.46 \pm 3.74$	$6.63 \pm 4.01$	$5.53 \pm 1.58$	0.424	5.43 ± 3.38	$6.93 \pm 3.84$	0.162
MLD (mm)	$2.38\pm0.74$	$2.41 \pm 0.79$	$2.20 \pm 0.33$	0.430	$2.17 \pm 0.68$	$2.48 \pm 0.74$	0.136
PB (%)	$67 \pm 12$	66 ± 13	71 ± 6	0.278	$71 \pm 11$	66 ± 13	0.147
LAD							
MLA (mm <sup>2</sup> )	$3.86 \pm 1.51$	3.79 ± 1.37	$4.24 \pm 2.16$	0.414	$4.33 \pm 1.67$	3.66 ± 1.41	0.129
MLD (mm)	$1.93 \pm 0.37$	1.91 ± 0.34	$2.09 \pm 0.51$	0.188	$2.05 \pm 0.46$	$1.89 \pm 0.32$	0.148
PB (%)	72 ± 8	72 ± 8	71 ± 10	0.701	$70 \pm 6$	72 ± 9	0.424
After LM-LAD stent imp	lantation					1	
LM .							
MLA (mm <sup>2</sup> )	$11.4 \pm 2.74$	$11.8 \pm 2.84$	9.97 ± 1.55	0.016*	11.7 ± 2.96	11.3 ± 2.62	0.459
MLD (mm)	$3.54 \pm 0.49$	3.60 ± 0.52	3.30 ± 0.26	0.031*	$3.57 \pm 0.54$	$3.53 \pm 0.47$	0.730
LAD							
MLA, (mm <sup>2</sup> )	8.11 ± 2.04	8.21 ± 1.94	$7.64 \pm 2.49$	0.315	7.98 ± 1.95	8.17 ± 2.11	0.689
MLD (mm)	$3.02 \pm 0.40$	$3.05 \pm 0.37$	$2.87 \pm 0.53$	0.101	$3.02 \pm 0.39$	$3.01 \pm 0.42$	0.906
PCI procedural data							
LM-LAD stent diameter (mm)	$3.61\pm0.40$	3.64 ± 0.39	$3.53\pm0.44$	0.331	$3.58\pm0.42$	3.63 ± 0.39	0.518
Postdilatation performed (%)	70 (76.9)	58 (78.4)	12 (70.6)	0.492	25 (78.1)	45 (76.3)	0.315
Postdilataion diameter (mm)	$3.90 \pm 0.41$	3.89 ± 0.67	3.75 ± 0.38	0.442	$3.93\pm0.41$	3.82 ± 0.72	0.493
LCX physiology assessm	nent						
Pre-PCI LCX-QFR	$0.89 \pm 0.09$	$0.92 \pm 0.05$	$0.77 \pm 0.10$	<0.01*	$0.89 \pm 0.06$	$0.89 \pm 0.10$	0.473
Post-PCI LCX-QFR	$0.86 \pm 0.11$	$0.90 \pm 0.05$	$0.69 \pm 0.10$	< 0.01*	$0.90 \pm 0.06$	$0.83 \pm 0.12$	< 0.01*

\*P < 0.05.

TABLE 3 Clinical outcomes for the study population at 1 year.

Variables	Total (N = 91 patients)	Post- QFR $\geq$ 0.80 (N = 74 patients)	Post- QFR<0.80 (N = 17 patients)	<i>P</i> -value	$\Delta QFR \ge 0$ ( $N = 32$ patients)	$\Delta QFR < 0$ ( $N = 59$ patients)	<i>P</i> -value
Primary endpoint	9 (9.9%)	6 (8.1%)	3 (17.6%)	0.235	0 (0.0%)	9 (15.3%)	0.020*
Cardiovascular death	1 (1.1%)	0 (0.0%)	1 (5.9%)	0.036*	0 (0.0%)	1 (1.7%)	0.459
Target bifurcation-related MI	1 (1.1%)	0 (0.0%)	1 (5.9%)	0.036*	0 (0.0%)	1 (1.7%)	0.459
Target bifurcation revascularization in LM-LAD	2 (2.2%)	2 (2.7%)	0 (0.0%)	0.493	0 (0.0%)	2 (3.4%)	0.292
Target bifurcation revascularization in LCX	4 (4.4%)	3 (4.1%)	1 (5.9%)	0.740	0 (0.0%)	4 (6.8%)	0.132
Target bifurcation revascularization both in LM-LAD and LCX	2 (2.2%)	1 (1.4%)	1 (5.9%)	0.251	0 (0.0%)	2 (3.4%)	0.292
Secondary endpoint							
Angina pectoris	9 (9.9%)	7 (9.5%)	2 (11.8%)	0.774	0 (0.0%)	9 (15.3%)	0.020*

\*P < 0.05.

potentially effective approach for the interventional management of left main bifurcation lesions. Postoperative physiological assessments can elucidate the functional significance and clinical implications of left main bifurcation lesions, offering the potential to enhance clinical outcomes in such cases (8). Previous research has shown that the use of rewire guidewires, which are advanced through stent struts into lateral branches, can complicate the procedure and is associated with a failure rate nearing 10%. QFR may serve as a reliable alternative for postoperative hemodynamic evaluation in patients with left main bifurcation lesions. A QFR threshold of <0.80 has been associated with a higher risk of cardiovascular death following

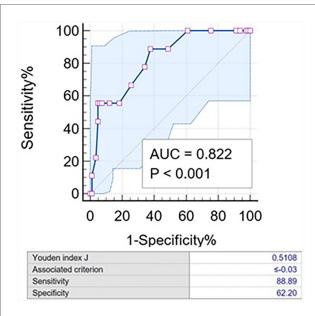


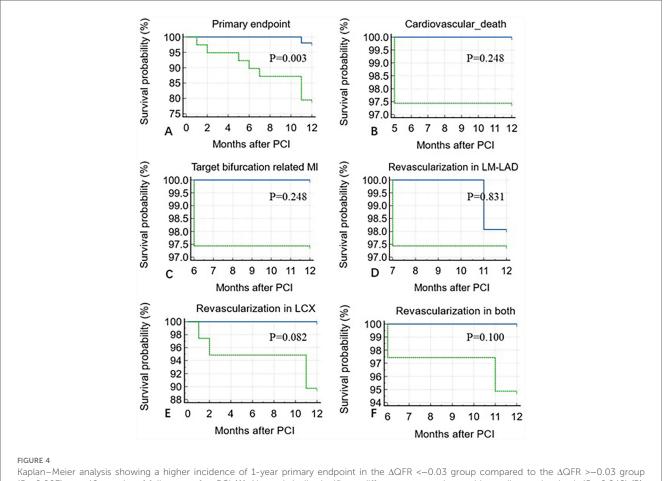
FIGURE 3

Characteristic curve of  $\Delta$ QFR for the 1-year incidence of the primary endpoint. The AUC was 0.822 (95% CI: 0.694–0.951). The diagnostic sensitivity and specificity of the  $\Delta$ QFR  $\leq$ -0.03 were 88.9% and 62.2%, respectively. AUC, area under the curve.

LM bifurcation stenting (9, 10). Therefore, we examined the connection between LCX-QFR and clinical results following single-stent placement for LM bifurcation lesions using IVUS guidance at our clinical center.

The major findings from the present study are as follows: (1) functional ischemia, as assessed by QFR ( $\Delta$ QFR < 0), was observed in approximately 65% of patients; (2) the IVUS-based lumen evaluation data from pre/post-LM-LAD did not effectively predict LCX-QFR outcomes; (3) patients in the  $\Delta$ QFR <0 group had a significantly increased risk of achieving the 1-year primary endpoint and persistent or recurrent angina pectoris, especially when  $\Delta$ QFR  $\leq$ -0.03; and (4) a post-LCX-QFR <0.8 was associated with a higher 1-year incidence of cardiovascular death or MI.

In a limited, single-center study involving 43 patients, LM-LAD crossover stenting led to 42% of cases showing angiographic diameter stenosis greater than 50%, an average reduction of 26% in the minimal lumen area, and a 7% occurrence of positive FFR at the LCX ostium after stenting (9). Post-IVUS analysis revealed that the MLA of the LCX ostium decreased to  $\leq 4.0 \text{ mm}^2$  following (provisional) crossover stenting, indicating the potential benefit of a twostent approach for distal LM bifurcation lesions. The reduced minimal lumen area, along with increased vessel eccentricity



#### (P = 0.003) over 12 months of follow-up after PCI (A). No statistically significant differences were observed in cardiovascular death (P = 0.248) (B),

target bifurcation-related MI (P = 0.248) (C), revascularization in LM-LAD (P = 0.831) (D), LCX (P = 0.082) (E), or both (P = 0.100) (F).

Characteristic	А	OR (95% CI)	P value
LM MLA	H <b>P</b> 1	1.40 (0.43 , 5.11)	0.577
LM MLD	•	0.16 (0.00 , 26.20)	0.491
LM PB	÷	0.97 (0.77, 1.18)	0.779
LAD MLA		→ 7.16 (0.55 , 207.00)	0.185
LAD MLD		0.00 (0.00, 5.81)	0.192
LAD PB	÷	0.93 (0.76 , 1.11)	0.462
Post-LM MLA	H=	1.67 (0.43, 8.92)	0.488
Post-LM MLD	H	3.10 (0.00 , 5.74)	0.754
Post-LAD MLA	<b>H</b> 4	0.42 (0.07, 1.59)	0.254
Post-LAD MLD		+ 17.00 (0.04 , 35.47)	0.398
<b>Post-Dilatation</b>			
Yes	÷	1.00 (Reference)	
No	•	0.19 (0.02, 1.51)	0.127
		٦ 36	
Characteristic	В	OR (95% CI)	P value
LM MLA	<b>⊢−=</b> ¦ (	0.65 (0.27, 1.42)	0.289
LM MLD	► <b></b>	11.70 (0.45 , 554.00)	0.166
LM PB	<b>i</b>	1.02 (0.88, 1.19)	0.831
	1	1.02(0.00, 1.19)	0.051
LAD MLA	F	1.25 (0.34 , 4.66)	0.732
	⊢ = →		
LAD MLA		1.25 (0.34 , 4.66)	0.732
LAD MLA LAD MLD		1.25 (0.34 , 4.66) 1.37 (0.02 , 112.00)	0.732 0.881
LAD MLA LAD MLD LAD PB		1.25 (0.34 , 4.66) 1.37 (0.02 , 112.00) 0.97 (0.85 , 1.09)	0.732 0.881 0.575
LAD MLA LAD MLD LAD PB Post-LM MLA		1.25 (0.34 , 4.66) 1.37 (0.02 , 112.00) 0.97 (0.85 , 1.09) 1.59 (0.81 , 3.44)	0.732 0.881 0.575 0.202
LAD MLA LAD MLD LAD PB Post-LM MLA Post-LM MLD		1.25 (0.34 , 4.66) 1.37 (0.02 , 112.00) 0.97 (0.85 , 1.09) 1.59 (0.81 , 3.44) 0.43 (0.01 , 16.60)	0.732 0.881 0.575 0.202 0.634
LAD MLA LAD MLD LAD PB Post-LM MLA Post-LM MLD Post-LAD MLA		1.25 (0.34 , 4.66) 1.37 (0.02 , 112.00) 0.97 (0.85 , 1.09) 1.59 (0.81 , 3.44) 0.43 (0.01 , 16.60) 0.29 (0.06 , 0.92)	0.732 0.881 0.575 0.202 0.634 0.064
LAD MLA LAD MLD LAD PB Post-LM MLA Post-LM MLD Post-LAD MLA Post-LAD MLD		1.25 (0.34 , 4.66) 1.37 (0.02 , 112.00) 0.97 (0.85 , 1.09) 1.59 (0.81 , 3.44) 0.43 (0.01 , 16.60) 0.29 (0.06 , 0.92)	0.732 0.881 0.575 0.202 0.634 0.064

FIGURE 5

Forest plot suggesting that IVUS evaluation results of pre/post-LAD are not significant independent predictors of postoperative QFR <0.8 (A) or  $\Delta$ QFR  $\leq$ -0.03 (B). LM, left main coronary artery; LAD, left anterior artery; MLA, minus lumen area; MLD, minus lumen diameter; PB, plaque burden.

and stable plaque mass, suggests that carina shift is the primary cause of acute lumen loss in the LCX. Nevertheless, carina shift is a localized occurrence and is infrequently linked to functional flow issues in the LCX (5). In our study, only 14 patients in the single-stent crossover group underwent a pre-PCI IVUS evaluation of the LCX. Following their surgeries, all of these patients declined additional assessments and treatments, relying solely on angiographic signs of thrombolysis in myocardial infarction (TIMI) III flow in the LCX. However, our retrospective analysis revealed a notable decrease in QFR. This decrease in QFR may indicate issues with microvascular function, ostial LCX remodeling, or issues related to carina shift, prompting the surgeon to evaluate the necessity for additional vascularization. A study involving 83 patients highlighted a significant visual-functional mismatch following LM-LAD stenting. The presence of a positive FFR after stenting was associated with a notably increased rate of target lesion failure over a 5-year follow-up period, with 33.4% in the positive FFR group compared to 10.7% in the negative FFR group. Furthermore, some researchers reported a comparable MACE rate of 18.1% at 1 year in their randomized study guided by FFR and angiography. They noted a somewhat increased revascularization rate in both the main and side branch vessels for angiography-guided PCI (11). Consequently, in conjunction with our research findings, we propose that patients with compromised LCX function following a single-stent strategy for LM-LAD should be considered for balloon dilatation at the LCX ostium to enhance both blood flow and overall prognosis. These patients should also receive closer clinical follow-up, repeat functional assessment, and alternative revascularization strategies when necessary.

Furthermore, no statistically significant differences were observed in the rates of 1-year cardiovascular mortality, target vessel revascularization, or related myocardial infarction, which may be attributable to the limited sample size of our investigation.

# **5** Limitations

Our study has several limitations that must be considered. First, the retrospective and single-center nature of this analysis, which was conducted using images not originally intended for QFR analysis, presents inherent limitations. Second, the sample size of our study was relatively small, and the cumulative incidence of 1-year endpoints was low. Therefore, our findings should be confirmed in larger prospective studies in the future. In addition, to date, only a 1-year follow-up has been completed. Follow-up is planned to continue for up to 3 years to further assess long-term outcomes. In addition, we performed IVUS assessment of LCX only in a small proportion of patients before PCI, which resulted in an inability to provide anatomical parameters that influence functional status after LCX. Finally, this study adopted a cutoff value of 0.80 for QFRguided ischemia, although the precise cutoff for QFR remains debated, which could have impacted the final outcomes.

# 6 Conclusion

Our research indicates that the LM-LAD single-stent strategy is associated with an increased incidence of decline in LCX-QFR. Furthermore, a  $\Delta$ QFR value of  $\leq$ -0.03 is correlated with the occurrence of major adverse cardiac events within 1 year, while a post-QFR value of <0.8 is linked to a heightened risk of cardiogenic death and myocardial infarction over the same period. Future investigations may enhance our understanding of optimal surgical approaches for LM bifurcation lesions by comparing anatomical outcomes guided by postprocedural physiology and intraluminal imaging, ultimately aiming to improve clinical prognoses.

# Data availability statement

The original contributions presented in this study are included in the article/Supplementary Material, further inquiries can be directed to the corresponding authors.

# **Ethics statement**

Ethical approval was not required for this study involving humans because the study used retrospectively collected data.

# References

The present study was conducted in accordance with the local legislation and institutional requirements. Written informed consent for participation was not required from the participants or the participants' legal guardians/next of kin in accordance with the national legislation and institutional requirements.

# Author contributions

C-DZ: Writing – original draft, Writing – review & editing. J-SZ: Writing – original draft. L-YH: Data curation, Writing – review & editing. X-YX: Data curation, Writing – review & editing. Y-PW: Funding acquisition, Resources, Writing – review & editing. MC: Funding acquisition, Resources, Writing – review & editing.

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# Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

# Generative AI statement

The author(s) declare that no Generative AI was used in the creation of this manuscript.

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