

Long-term outcomes of intravascular ultrasound-guided implantation of bare metal stents versus drug-eluting stents in primary percutaneous coronary intervention

Yun-Kyeong Cho¹, Seung-Ho Hur¹, Nam-Hee Park², Sang-Woong Choi¹, Ji-Hyun Sohn¹, Hyun-Ok Cho¹, Hyoung-Seob Park¹, Hyuck-Jun Yoon¹, Hyungseop Kim¹, Chang-Wook Nam¹, Yoon-Nyun Kim¹, and Kwon-Bae Kim¹

Departments of ¹Internal Medicine and ²Cardiothoracic Surgery, Keimyung University School of Medicine, Daegu, Korea

Background/Aims: While drug-eluting stents (DESs) have shown favorable outcomes in ST-segment elevation myocardial infarction (STEMI) compared to bare metal stents (BMSs), there are concerns about the risk of stent thrombosis (ST) with DESs. Because intravascular ultrasound (IVUS) guidance may help optimize stent placement and improve outcomes in percutaneous coronary intervention (PCI) patients, we evaluated the impact of IVUS-guided BMS versus DES implantation on long-term outcomes in primary PCI.

Methods: In all, 239 STEMI patients received DES (n = 172) or BMS (n = 67) under IVUS guidance in primary PCI. The 3-year incidence of major adverse cardiac events (MACEs) including death, myocardial infarction (MI), target vessel revascularization (TVR), and ST was evaluated.

Results: There was no difference in all cause mortality or MI. However, the incidence of TVR was 23.9% with BMS versus 9.3% with DES ($p = 0.005$). Thus, the number of MACEs was significantly lower with DES (11.0% vs. 29.9%; $p = 0.001$). The incidence of definite or probable ST was not different (1.5% vs. 2.3%; $p = 1.0$). IVUS-guided DES implantation (hazard ratio [HR], 0.25; 95% confidence interval [CI], 0.08 to 0.78; $p = 0.017$), stent length (HR, 1.03; 95% CI, 1.00 to 1.06; $p = 0.046$), and multivessel disease (HR, 3.01; 95% CI, 1.11 to 8.15; $p = 0.030$) were independent predictors of MACE.

Conclusions: In patients treated with primary PCI under IVUS guidance, the use of DES reduced the incidence of 3-year TVR versus BMS. However, all cause mortality and MI were similar between the groups. The incidence of ST was low in both groups.

Keywords: Ultrasonography, interventional; Myocardial infarction; Drug-eluting stents

Received: February 11, 2013
Revised: March 15, 2013
Accepted: June 28, 2013

Correspondence to
Seung-Ho Hur, M.D.

Department of Internal Medicine,
Keimyung University Dongsan
Medical Center, 56 Dalseong-ro,
Jung-gu, Daegu 700-712, Korea
Tel: +82-53-250-7949
Fax: +82-53-250-7034
E-mail: shur@dsmc.or.kr

INTRODUCTION

Numerous studies have shown that intravascular ultra-

sound (IVUS)-guided percutaneous coronary intervention (PCI) decreases the frequency of major adverse cardiac events (MACEs) and stent thrombosis (ST), mainly

in patients with stable coronary artery disease undergoing elective PCI [1-4]. However, in the setting of acute myocardial infarction (AMI), the benefits of IVUS guidance during PCI remain a matter of debate [5,6].

Drug-eluting stents (DESs) are a highly efficacious treatment for patients with coronary artery disease, markedly inhibiting neointimal hyperplasia [7-9], and they have demonstrated favorable clinical outcomes even in patients with high-risk clinical conditions such as AMI and diabetes [10]. However, there are also safety concerns about their use in AMI patients, because of an increased risk of ST [11].

Although bare metal stents (BMSs) are less effective than DESs for inhibiting neointimal proliferation [8,9], they are associated with similar clinical outcomes when adequate stent expansion can be achieved during an index procedure [12-14]. Furthermore, a recent meta-analysis in primary angioplasty reported that BMSs were not associated with an increased risk of very late ST [15]. Thus, we hypothesized that IVUS-guided PCI using BMSs would show similar efficacy and better safety at long-term follow-up than IVUS-guided DES implantation in AMI patients undergoing primary PCI.

METHODS

We analyzed data retrospectively from patients with ST-segment elevation myocardial infarction (STEMI) who underwent primary PCI for a *de novo* culprit lesion from January 2000 to July 2008.

During primary PCI, BMSs were used exclusively from January 2000 to May 2003, whereas DESs were implanted exclusively from June 2003 to July 2008. Regardless of stent type, all procedures were performed according to standard techniques via the femoral approach.

All patients were older than 18 years. To be eligible for primary PCI, patients had to meet the following criteria: symptoms present < 12 hours from onset to time of hospital arrival, and ST-segment elevation or a new left bundle branch block. All interventions were performed according to current standard guidelines. Procedural success in the infarct-related artery was defined as residual stenosis < 30% by visual estimation with thrombolysis in myocardial infarction (TIMI) grade 3 flow.

Patients were excluded if they had: intolerance or a contraindication to aspirin or thienopyridine, advanced heart failure or an ejection fraction < 30%, or another severe comorbidity. The patients were premedicated with aspirin 300 mg, which was continued indefinitely, and given a loading dose of ticlopidine (500 mg) or clopidogrel (300 to 600 mg) before PCI. The patients were advised to stay on dual antiplatelet therapy for a minimum of 3 months in cases of BMS and 12 months for DES.

IVUS (Atlantis, Boston Scientific Corp., Minneapolis, MN, USA) was performed and interpreted by the physician. IVUS images were obtained after administration of 200 mcg of nitroglycerin. After preinterventional or post-balloon IVUS was performed, stent size and diameter were determined according to IVUS parameters. When postdilation was required to optimize stent expansion or apposition, a balloon shorter than the stent length was used with careful positioning of the balloon inside the stent to avoid injury at the edge. Stent under-expansion was defined as minimal stent area (MSA) < 6.5 mm² for BMS and 5.0 mm² for DES [16]. Coronary angiography results were analyzed using a computer-assisted system for quantitative coronary angiography (QCA) analysis (Digital Cardiac Imaging System, Philips Medical Systems, Best, The Netherlands) using end diastolic frames and a contrast-filled guiding catheter for calibration. The percent diameter stenosis was defined as $[1 - (\text{minimal lumen diameter}/\text{reference vessel diameter})] \times 100$.

The primary endpoint was defined as the incidence of MACEs including all cause death, myocardial infarction (MI), target vessel revascularization (TVR), and ST at 3-year follow-up. MI was defined as an elevation in creatinine kinase-MB ≥ 3 times the upper normal value. TVR was defined as percutaneous or surgical revascularization of the stented vessel. ST was defined using the Academic Research Consortium definition [17].

Statistical analysis

Statistical analyses were performed using the SPSS software version 15.0 (SPSS Inc., Chicago, IL, USA). Continuous data are presented as means \pm standard deviation while categorical data are presented as frequencies. Continuous variables were compared using unpaired Student *t* tests. Categorical variables were compared

using chi-square and Fisher exact tests. The cumulative incidence of MACE was estimated according to the Kaplan-Meier method, and curves were compared using the log-rank test. Cox multivariate regression analyses were used to determine predictors of cardiac events. Variables with $p < 0.10$ on univariate analysis and classical risk factors such as age, gender, diabetes, hypertension, and hyperlipidemia, were entered into a multivariate regression analysis. These p values < 0.05 were considered to indicate statistical significance.

RESULTS

In total, 337 patients who underwent primary PCI with BMS (107 patients) or DES (230 patients) implantation were enrolled consecutively. In total, 239 STEMI patients received BMS (67 patients with 77 stents) or DES (172 patients with 221 stents) under IVUS guidance (Fig. 1). During enrollment, four types of DES were used: sirolimus-eluting stents (SEEs, 48.3%; Cypher, Cordis, Miami Lakes, FL, USA), paclitaxel-eluting stents (PESs, 29.0%; Taxus, Boston Scientific, Natick, MA, USA), zotarolimus-eluting stents (ZESs, 16.9%; Endeavor Sprint, Medtronic CardioVascular, Santa Rosa, CA, USA), and everolimus-eluting stents (EESs, 5.8%; Xience V, Abbott Vascular Devices, Santa Clara, CA, USA). Patients undergoing IVUS-guided BMS implantation had decreased left ventricular ejection fraction ($p < 0.001$).

The frequency of diabetes mellitus and hypertension did not differ between the groups (Table 1). Procedural characteristics are presented in Table 2. Infarct-related arteries and lesion type, by American College of Cardiology/American Heart Association classification, were similar between the DES and BMS groups. However, the presence of intracoronary thrombus by coronary angiography and performance of thrombus aspiration were significantly higher in the BMS group than the DES group (73.1% vs. 51.7%, $p = 0.010$; and 35.8% vs. 8.7%, $p < 0.001$, respectively). The reference vessel diameter and stent diameter were significantly larger in the BMS group than in the DES group (3.47 ± 0.43 mm vs. 3.23 ± 0.40 mm and 3.58 ± 0.42 mm vs. 3.23 ± 0.39 mm; all $p < 0.05$). Lesion length and stent length were longer in the DES group (28.5 ± 14.2 mm vs. 23.8 ± 11.7 mm and 32.0 ± 15.4 mm vs. 26.9 ± 12.5 mm; both $p < 0.01$). The minimal stent diameter by QCA and MSA by IVUS were significantly larger in the BMS group than the DES group (3.31 ± 0.44 mm vs. 2.89 ± 0.39 mm and 7.51 ± 2.15 mm² vs. 6.57 ± 2.16 mm²; both $p < 0.05$) (Table 3). However, the incidence of stent underexpansion was higher in the BMS group (37.3% vs. 22.1%; $p = 0.023$). Prescriptions of β blockers, angiotensin converting enzyme inhibitors or angiotensin receptor blockers, statins, and triple antiplatelet therapy were more frequent in the DES group (all $p < 0.05$).

Clinical outcomes

All patients were followed-up with face to face contact or by telephone and more than half of patients received angiographic follow-up examinations in each group (53.7% of BMS and 61.0% of DES). There was no difference in all cause mortality or MI between the DES and BMS groups at 1-, 2-, and 3-year follow-up (Table 4). However, the incidence of TVR was significantly lower in the DES group than the BMS group (6.4% vs. 17.9%, $p = 0.006$ at 1 year; 8.1% vs. 23.9%, $p = 0.002$ at 2 years; and 9.3% vs. 23.9%, $p = 0.005$ at 3 years, respectively). However, the incidence of TVR did not differ among the four DES types (8.6% of SEE, 10.0% of PES, 14.3% of ZES, and 0% of EES, $p = 0.736$ at 3 years). The cumulative incidence of MACE was significantly lower in the DES group (7.6% vs. 22.4%, $p = 0.003$ at 1 year; 9.3% vs. 29.9%, $p < 0.001$ at 2 years; and 11.0% vs. 29.9%, $p = 0.001$ at 3 years, respectively). The incidence of definite or probable ST did not differ between

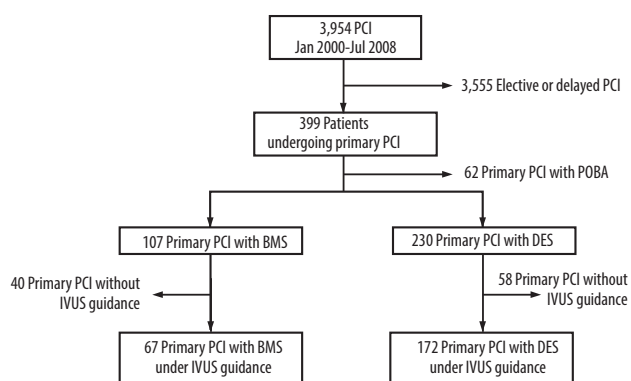


Figure 1. Patient recruitment and follow-up. PCI, percutaneous coronary intervention; POBA, plain old balloon angioplasty; BMS, bare metal stent; DES, drug-eluting stent; IVUS, intravascular ultrasound.

Table 1. Patient demographics

| | BMS (n = 67) | DES (n = 172) | p value |
|----------------------------------|--------------|---------------|---------|
| Age, yr | 61.4 ± 11.5 | 62.7 ± 10.8 | 0.394 |
| Male gender | 51 (76.1) | 135 (78.5) | 0.730 |
| Diabetes | 13 (19.4) | 34 (19.8) | 1.000 |
| Hypertension | 21 (31.3) | 62 (36.0) | 0.547 |
| Smoking | 45 (67.2) | 107 (62.2) | 0.550 |
| Hyperlipidemia | 9 (13.4) | 38 (22.1) | 0.150 |
| Prior MI | 0 | 9 (5.2) | 0.065 |
| Prior PCI | 1 (1.5) | 11 (6.4) | 0.187 |
| Prior CABG | 0 | 2 (1.2) | 1.000 |
| Renal Insufficiency ^a | 1 (1.5) | 3 (1.7) | 1.000 |
| LVEF, % | 44.0 ± 6.3 | 48.0 ± 8.8 | < 0.001 |
| Total cholesterol, mg/dL | 188.2 ± 35.7 | 197.9 ± 38.6 | 0.075 |
| High density lipoprotein, mg/dL | 43.4 ± 10.9 | 46.8 ± 13.8 | 0.072 |
| Triglyceride, mg/dL | 115.2 ± 67.2 | 90.5 ± 53.8 | 0.009 |
| Low density lipoprotein, mg/dL | 130.5 ± 36.8 | 131.3 ± 33.3 | 0.867 |
| Medication at discharge | | | |
| ACE inhibitor or ARB | 38 (56.7) | 122 (70.9) | 0.046 |
| β-Blocker | 36 (53.7) | 140 (81.4) | < 0.001 |
| Statin | 14 (20.9) | 109 (63.4) | < 0.001 |
| Triple antiplatelet therapy | 2 (3.0) | 47 (27.3) | < 0.001 |

Values are presented as mean ± SD or number (%).

BMS, bare metal stent; DES, drug-eluting stent; MI, myocardial infarction; PCI, percutaneous coronary intervention; CABG, coronary artery bypass surgery; LVEF, left ventricular ejection fraction; ACE, angiotensin converting enzyme; ARB, angiotensin receptor blocker.

^aRenal insufficiency defined as serum creatinine > 2 mg/dL.

the groups (1.7% vs. 1.5%, $p = 1.0$ at 1 and 2 years; 2.3% vs. 1.5%, $p = 1.0$ at 3 years, respectively). These clinical outcomes were seen at the time of the 1-year follow-up and were sustained during 3 years of follow-up. On univariate analysis, stent type, stent underexpansion, high density lipoprotein level, multivessel disease and the prescription of statin were associated with 3-year MACE. After adjustment for these parameters, IVUS-guided DES implantation was associated with a lower rate of 3-year MACE (hazard ratio [HR], 0.34; 95% confidence interval [CI], 0.13 to 0.90; $p = 0.030$) versus BMS. In addition, stent length (HR, 1.03; 95% CI, 1.00 to 1.06; $p = 0.023$) and multivessel disease (HR, 2.49; 95% CI, 1.00 to 6.18; $p = 0.049$) were independent predictors of 3-year MACE (Table 5).

DISCUSSION

The major finding of the present study was that despite the fact that IVUS-guided BMS implantation was associated with significantly larger MSA after stenting, IVUS-guided DES implantation showed better efficacy by diminishing the rate of TVR with similar safety and no increased risk of ST up to 3 years. In addition, IVUS-guided BMS versus DES implantation in patients with STEMI undergoing primary PCI showed similar long-term clinical outcomes to those seen in patients with stable coronary artery stenosis.

Because DESs have proven effective for inhibiting intimal hyperplasia in stable coronary lesions [9], STEMI has been treated with DESs [18-22]. Despite concerns regarding an increased risk of ST due to delayed healing [11], numerous studies have reported superior efficacy

Table 2. Procedural characteristics

| | BMS (n = 67) | DES (n = 172) | p value |
|--|-------------------------------------|--------------------------------------|---------|
| Door to balloon time, min | 106.8 ± 36.0 | 74.6 ± 29.0 | < 0.001 |
| Infarct related artery | | | 0.755 |
| Left anterior descending artery | 30 (44.7) | 85 (49.4) | |
| Left circumflex artery | 6 (9.0) | 16 (9.3) | |
| Right coronary artery | 31 (46.3) | 71 (41.3) | |
| Diseased vessel | | | 0.150 |
| One-vessel | 30 (44.7) | 101 (58.7) | |
| Two-vessel | 28 (41.8) | 54 (31.4) | |
| Three-vessel | 9 (13.5) | 17 (9.9) | |
| Lesion type by ACC /AHA classification | | | 0.179 |
| Type A/B1 | 16 (23.9) | 44 (25.6) | |
| Type B2/C | 51 (76.1) | 128 (74.4) | |
| Intracoronary thrombus | 49 (73.1) | 89 (51.7) | 0.010 |
| Thrombus aspiration | 24 (35.8) | 15 (8.7) | < 0.001 |
| Preprocedural TIMI flow | | | 0.712 |
| Grade 0/1/2/3 | 31 (46.4)/2 (2.9)/5 (7.2)/29 (43.5) | 94 (54.6)/4 (2.4)/10 (5.9)/64 (37.1) | |
| Minimal lumen diameter at preintervention, mm | 0.21 ± 0.30 | 0.20 ± 0.31 | 0.837 |
| Minimal stent diameter at postintervention, mm | 3.31 ± 0.44 | 2.89 ± 0.39 | < 0.001 |
| Reference vessel diameter, mm | 3.47 ± 0.43 | 3.23 ± 0.40 | < 0.001 |
| Lesion length, mm | 23.8 ± 11.7 | 28.5 ± 14.2 | 0.009 |
| Stent diameter, mm | 3.58 ± 0.42 | 3.23 ± 0.39 | 0.021 |
| Stent length, mm | 26.9 ± 12.5 | 32.0 ± 15.4 | 0.009 |
| Stent number per patient | 1.15 ± 0.36 | 1.28 ± 0.58 | 0.113 |
| Adjunctive balloon inflation | 8 (11.9) | 26 (15.1) | 0.681 |
| Maximum balloon diameter, mm | 3.80 ± 0.46 | 3.54 ± 0.44 | < 0.001 |
| Maximum balloon pressure, atm | 13.8 ± 2.2 | 15.2 ± 2.2 | < 0.001 |
| Final TIMI flow | | | 0.411 |
| Grade 0/1/2/3 | 0/0/2 (3.0)/65 (97.0) | 0/3 (1.7)/9 (5.2)/160 (93.1) | |
| Dissection | 2 (3.0) | 5 (2.9) | 1.000 |
| Abrupt closure | 8 (11.9) | 20 (11.6) | 1.000 |
| Glycoprotein IIb/IIIa inhibitor | 1 (1.5) | 11 (6.4) | 0.187 |

Values are presented as mean ± SD or number (%).

BMS, bare metal stent; DES, drug-eluting stent; ACC/AHA, American College of Cardiology/American Heart Association; TIMI, thrombolysis in myocardial infarction.

with DES versus BMS, driven mainly by reduced TVR or target lesion revascularization (TLR), without safety issues, in AMI patients during 3 to 5 years of follow-up [18-22]. A recent optical coherence tomography substudy of the Harmonizing Outcomes with Revascularization and Stents in Acute Myocardial Infarction (HORIZON-AMI) trial demonstrated that most struts (> 94%) were covered in PES at 13-month follow-up, suggesting

that the risk of ST may not be as high as previously anticipated, compared to BMS [23]. Our results are consistent with these findings. However, a recent meta-analysis of 13 randomized trials in primary angioplasty showed that although first-generation DES compared to BMS yielded a significantly lower incidence of TVR (12.7% vs. 20.1%; HR, 0.57; 95% CI, 0.50 to 0.66; *p* < 0.001) with no effect on mortality, reinfarction, or ST, the risk

Table 3. Intravascular ultrasound analysis

| | BMS (n = 67) | DES (n = 172) | p value |
|--------------------------------------|--------------|---------------|---------|
| Preintervention | | | |
| Proximal reference segment | | | |
| Vessel area, mm ² | 20.28 ± 5.79 | 17.31 ± 5.88 | 0.007 |
| Lumen area, mm ² | 11.28 ± 3.31 | 9.79 ± 3.50 | 0.020 |
| Plaque area, mm ² | 9.00 ± 3.26 | 7.65 ± 2.99 | 0.021 |
| Plaque burden, % ^a | 44.1 ± 8.1 | 44.4 ± 10.1 | 0.867 |
| Lesion site ^b | | | |
| Remodeling index ^c | 1.01 ± 0.15 | 1.04 ± 0.22 | 0.296 |
| Vessel area, mm ² | 17.07 ± 4.90 | 14.90 ± 4.81 | 0.004 |
| Lumen area, mm ² | 2.68 ± 0.70 | 2.62 ± 0.93 | 0.702 |
| Plaque area, mm ² | 14.83 ± 5.04 | 12.27 ± 4.47 | 0.003 |
| Plaque burden, % | 83.8 ± 5.5 | 81.3 ± 6.7 | 0.034 |
| Distal reference segment | | | |
| Vessel area, mm ² | 16.47 ± 8.60 | 11.75 ± 5.51 | 0.001 |
| Lumen area, mm ² | 9.34 ± 5.78 | 6.80 ± 3.09 | 0.003 |
| Plaque area, mm ² | 7.13 ± 3.62 | 4.95 ± 2.78 | 0.001 |
| Plaque burden, % | 43.0 ± 9.9 | 40.7 ± 9.2 | 0.228 |
| Postintervention | | | |
| Minimal stent area, mm ² | 7.51 ± 2.15 | 6.57 ± 2.16 | 0.005 |
| Stent underexpansion, % ^d | 37.3 | 22.1 | 0.023 |

Values are presented as mean ± SD.

BMS, bare metal stent; DES, drug-eluting stent.

^aPlaque burden was calculated as [(plaque area/vessel area) × 100].

^bLesions were assessed after balloon dilation in cases of preprocedural thrombolysis in myocardial infarction o flow.

^cThe remodeling index was calculated as vessel area at lesion site/mean reference vessel area.

^dStent underexpansion was defined as minimal stent area < 6.5 mm² for BMS and 5.0 mm² for DES.

of very late ST and late infarction were significantly higher in patients treated with DES (HR, 2.81; 95% CI, 1.28 to 6.19; *p* = 0.04 and HR, 2.06; 95% CI, 1.22 to 3.49; *p* = 0.03, respectively) [15]. Taken together, DES safety may still be inconclusive in AMI patients although long-term efficacy seems to be favorable for DES. Thus, larger populations with longer-term follow-up will be necessary to clarify this issue in the setting of AMI.

Although BMS showed less inhibition of neointimal hyperplasia than DES, adequate BMS expansion with or without IVUS guidance provided favorable clinical outcomes, similar to those for DES [12]. Furthermore, the benefit of DES use was limited to vessels ≤ 3 mm in size [13,24]. In the A Randomized Controlled Trial of Angiography versus Intravascular Ultrasound-Directed Bare Metal Coronary Stent Placement (AVID) trial, the final MSA was

7.55 ± 2.82 mm² in the IVUS-guided group and 12-month TLR was only 8.1% in 394 patients receiving elective BMS placement [12]. In the present study, the IVUS-guided BMS group had vessel sizes > 3 mm and a similar final MSA to the AVID trial, predicting that long-term clinical outcomes were comparable with those in the IVUS-guided DES group. However, IVUS-guided DES implantation showed a lower incidence of MACE at 3-year follow-up, driven primarily by a reduced TVR rate, suggesting that IVUS-guided BMS versus DES implantation in patients with STEMI undergoing primary PCI had similar long-term clinical outcomes to those seen in stable coronary artery stenosis [9,24].

STEMI has been considered an off-label DES use [25]. Moreover, several studies demonstrated that STEMI is a strong predictor for the development of early or late ST

Table 4. Clinical outcomes at 30 days, and 1, 2, and 3 years

| | BMS (n = 67) | DES (n = 172) | p value |
|---------------------------------------|--------------|---------------|---------|
| 30-Day outcomes | | | |
| MACE | 3 (4.5) | 4 (2.3) | 0.404 |
| All cause death | 2 (3.0) | 1 (0.6) | 1.000 |
| Nonfatal MI | 0 | 1 (0.6) | 1.000 |
| Target lesion revascularization | 1 (1.5) | 3 (1.7) | 1.000 |
| Target vessel revascularization | 1 (1.5) | 3 (1.7) | 1.000 |
| Definite or probable stent thrombosis | 1 (1.5) | 3 (1.7) | 1.000 |
| 1-Year outcomes | | | |
| MACE | 15 (22.4) | 13 (7.6) | 0.003 |
| All cause death | 2 (3.0) | 1 (0.6) | 0.191 |
| Nonfatal MI | 3 (4.5) | 2 (1.2) | 0.189 |
| Target lesion revascularization | 10 (14.9) | 11 (6.4) | 0.047 |
| Target vessel revascularization | 12 (17.9) | 11 (6.4) | 0.006 |
| Definite or probable stent thrombosis | 1 (1.5) | 3 (1.7) | 1.000 |
| 2-Year outcomes | | | |
| MACE | 20 (29.9) | 16 (9.3) | < 0.001 |
| All cause death | 3 (4.5) | 1 (0.6) | 0.068 |
| Nonfatal MI | 3 (4.5) | 4 (2.3) | 0.404 |
| Target lesion revascularization | 13 (19.4) | 13 (7.6) | 0.020 |
| Target vessel revascularization | 16 (23.9) | 14 (8.1) | 0.002 |
| Definite or probable stent thrombosis | 1 (1.5) | 3 (1.7) | 1.000 |
| 3-Year outcomes | | | |
| MACE | 20 (29.9) | 19 (11.0) | 0.001 |
| All cause death | 3 (4.5) | 1 (0.6) | 0.068 |
| Nonfatal MI | 3 (4.5) | 6 (3.5) | 0.713 |
| Target lesion revascularization | 13 (19.4) | 14 (8.1) | 0.025 |
| Target vessel revascularization | 16 (23.9) | 16 (9.3) | 0.005 |
| Definite or probable stent thrombosis | 1 (1.5) | 4 (2.3) | 1.000 |

Values are presented as number (%).

BMS, bare metal stent; DES, drug-eluting stent; MACE, major adverse cardiac event; MI, myocardial infarction.

[26,27]. The incidence of ST after DES implantation has been reported to be 3% to 5% of patients with STEMI undergoing primary PCI [19-22]. Because of large thrombotic burden and a higher chance of incompletely apposed struts in STEMI patients, the use of DES has a potential risk of ST and consequently adverse cardiac events. However, the present study showed a relatively low incidence (2.3%) of ST compared to previous studies [19-22]. A possible explanation is that with either BMS or DES, implantation under IVUS guidance might contribute to reducing the rate of ST.

In a previous study, we demonstrated that IVUS-guided PCI may reduce long-term mortality compared to angiography-guided PCI in real world practice [2], consistent with a study on unprotected left mains [3]. In these studies, 50% to 60% of the study population was diagnosed with acute coronary syndrome. Another study by Roy et al. [4] showed that IVUS-guided PCI significantly reduced the development of subacute ST after DES implantation.

The benefits of IVUS-guided PCI seem to be offset by AMI presentation. Because the number of treated le-

Table 5. Independent predictors of 3-year major adverse cardiac event

| | Univariate | | Multivariate | |
|----------------------|------------------|---------|------------------|---------|
| | HR (95% CI) | p value | HR (95% CI) | p value |
| Age, yr | 0.99 (0.96–1.02) | 0.600 | | |
| Male gender | 1.37 (0.57–3.30) | 0.489 | | |
| Diabetes | 1.52 (0.68–3.39) | 0.307 | | |
| Hypertension | 1.38 (0.68–2.79) | 0.368 | | |
| Hypercholesterolemia | 0.71 (0.28–1.80) | 0.464 | | |
| HDL, mg/dL | 0.95 (0.92–0.99) | 0.008 | | |
| Multivessel disease | 2.21 (1.09–4.47) | 0.027 | 2.49 (1.00–6.18) | 0.049 |
| Thrombus aspiration | 2.03 (0.90–4.61) | 0.090 | | |
| Stent type, DES | 0.29 (0.14–0.59) | 0.001 | 0.34 (0.13–0.90) | 0.030 |
| Stent length, mm | 1.02 (1.00–1.04) | 0.057 | 1.03 (1.00–1.06) | 0.023 |
| Stent underexpansion | 3.13 (1.43–6.88) | 0.004 | | |
| Use of statin | 0.36 (0.17–0.74) | 0.006 | | |

HR, hazard ratio; CI, confidence interval; HDL, high density lipoprotein; DES, drug-eluting stent.

sions and stent implantation were both higher and procedural time was prolonged in the IVUS-guided group, 1-year clinical outcomes did not differ between IVUS-guided PCI and angiography-guided PCI [5,6]. Whether the impact of IVUS-guided PCI is different depends on clinical presentation and a randomized clinical trial is needed.

This retrospective study has several limitations. First, the chronological difference between the use of BMS and DES and the fact that the decision for IVUS guidance during primary PCI was at the physician's discretion may have introduced selection bias. Second, although we evaluated predictors of MACE performing multivariate analysis, unmeasured confounders could affect the clinical results. Third, because few patients were treated with BMS or DES under IVUS guidance, the clinical events during 3 years of follow-up may be underestimated. The study was also underpowered to detect rarely occurring events, such as ST. Fourth, our finding that the use of DES under IVUS guidance showed better efficacy with no increased risk of ST was similar to that in other subsets of lesions or patients. Thus, the clinical implications of our results may be limited. Finally, a heterogeneous baseline, procedural characteristics, and medication patterns between BMS and DES patients might affect long-term outcomes. In fact, recent DES trials have included more complex lesions and/or high-risk

patients compared to previous BMS studies, accounting for the differences between baseline characteristics and medication patterns.

Although in the present study IVUS-guided BMS implantation was associated with a larger final MSA, IVUS-guided DES implantation appeared to be as safe as BMS and showed significant benefits for reducing the risk of TVR for up to 3 years in patients with STEMI undergoing primary PCI.

KEY MESSAGE

1. Even if intravascular ultrasound (IVUS) guidance, drug-eluting stent (DES) implantation in primary percutaneous coronary intervention had better efficacy compared with bare metal stent.
2. IVUS-guided DES implantation showed favorable long-term safety without increased risk of stent thrombosis in the ST-segment elevation myocardial infarction setting.

Conflict of interest

No potential conflict of interest relevant to this article is reported.

REFERENCES

1. Claessen BE, Mehran R, Mintz GS, et al. Impact of intravascular ultrasound imaging on early and late clinical outcomes following percutaneous coronary intervention with drug-eluting stents. *JACC Cardiovasc Interv* 2011;4:974-981.
2. Hur SH, Kang SJ, Kim YH, et al. Impact of intravascular ultrasound-guided percutaneous coronary intervention on long-term clinical outcomes in a real world population. *Catheter Cardiovasc Interv* 2013;81:407-416.
3. Park SJ, Kim YH, Park DW, et al. Impact of intravascular ultrasound guidance on long-term mortality in stenting for unprotected left main coronary artery stenosis. *Circ Cardiovasc Interv* 2009;2:167-177.
4. Roy P, Steinberg DH, Sushinsky SJ, et al. The potential clinical utility of intravascular ultrasound guidance in patients undergoing percutaneous coronary intervention with drug-eluting stents. *Eur Heart J* 2008;29:1851-1857.
5. Ahmed K, Jeong MH, Chakraborty R, et al. Role of intravascular ultrasound in patients with acute myocardial infarction undergoing percutaneous coronary intervention. *Am J Cardiol* 2011;108:8-14.
6. Maluenda G, Lemesle G, Ben-Dor I, et al. Impact of intravascular ultrasound guidance in patients with acute myocardial infarction undergoing percutaneous coronary intervention. *Catheter Cardiovasc Interv* 2010;75:86-92.
7. Grube E, Dawkins K, Guagliumi G, et al. TAXUS VI final 5-year results: a multicentre, randomised trial comparing polymer-based moderate-release paclitaxel-eluting stent with a bare metal stent for treatment of long, complex coronary artery lesions. *EuroIntervention* 2009;4:572-577.
8. Stettler C, Wandel S, Allemann S, et al. Outcomes associated with drug-eluting and bare-metal stents: a collaborative network meta-analysis. *Lancet* 2007;370:937-948.
9. Moses JW, Leon MB, Popma JJ, et al. Sirolimus-eluting stents versus standard stents in patients with stenosis in a native coronary artery. *N Engl J Med* 2003;349:1315-1323.
10. Stone GW, Lansky AJ, Pocock SJ, et al. Paclitaxel-eluting stents versus bare-metal stents in acute myocardial infarction. *N Engl J Med* 2009;360:1946-1959.
11. Nakazawa G, Finn AV, Joner M, et al. Delayed arterial healing and increased late stent thrombosis at culprit sites after drug-eluting stent placement for acute myocardial infarction patients: an autopsy study. *Circulation* 2008;118:1138-1145.
12. Russo RJ, Silva PD, Teirstein PS, et al. A randomized controlled trial of angiography versus intravascular ultrasound-directed bare-metal coronary stent placement (the AVID Trial). *Circ Cardiovasc Interv* 2009;2:113-123.
13. Kim HT, Nam CW, Hur SH, et al. Two-year clinical outcomes after large coronary stent (4.0 mm) placement: comparison of bare-metal stent versus drug-eluting stent. *Clin Cardiol* 2010;33:620-625.
14. Yan BP, Ajani AE, New G, et al. Are drug-eluting stents indicated in large coronary arteries? Insights from a multi-centre percutaneous coronary intervention registry. *Int J Cardiol* 2008;130:374-379.
15. De Luca G, Dirksen MT, Spaulding C, et al. Drug-eluting vs bare-metal stents in primary angioplasty: a pooled patient-level meta-analysis of randomized trials. *Arch Intern Med* 2012;172:611-621.
16. Sonoda S, Morino Y, Ako J, et al. Impact of final stent dimensions on long-term results following sirolimus-eluting stent implantation: serial intravascular ultrasound analysis from the sirius trial. *J Am Coll Cardiol* 2004;43:1959-1963.
17. Mauri L, Hsieh WH, Massaro JM, Ho KK, D'Agostino R, Cutlip DE. Stent thrombosis in randomized clinical trials of drug-eluting stents. *N Engl J Med* 2007;356:1020-1029.
18. Park KW, Kang SH, Chung WY, et al. 'Real world' comparison of drug-eluting stents vs bare metal stents in the treatment of unselected patients with acute ST-segment elevation myocardial infarction. *Circ J* 2010;74:1111-1120.
19. Stone GW, Witzenbichler B, Guagliumi G, et al. Heparin plus a glycoprotein IIb/IIIa inhibitor versus bivalirudin monotherapy and paclitaxel-eluting stents versus bare-metal stents in acute myocardial infarction (HORIZONS-AMI): final 3-year results from a multicentre, randomised controlled trial. *Lancet* 2011;377:2193-2204.
20. Violini R, Musto C, De Felice F, et al. Maintenance of long-term clinical benefit with sirolimus-eluting stents in patients with ST-segment elevation myocardial infarction 3-year results of the SESAMI (sirolimus-eluting stent versus bare-metal stent in acute myocardial infarction) trial. *J Am Coll Cardiol* 2010;55:810-814.
21. Vink MA, Dirksen MT, Suttorp MJ, et al. 5-year follow-up after primary percutaneous coronary intervention with a paclitaxel-eluting stent versus a bare-metal stent in acute ST-segment elevation myocardial infarction: a follow-up study of the PASSION (Paclitaxel-Eluting Versus Con-

- tional Stent in Myocardial Infarction with ST-Segment Elevation) trial. *JACC Cardiovasc Interv* 2011;4:24-29.
22. Spaulding C, Teiger E, Commeau P, et al. Four-year follow-up of TYPHOON (trial to assess the use of the CYPHer sirolimus-eluting coronary stent in acute myocardial infarction treated with Balloon angioplasty). *JACC Cardiovasc Interv* 2011;4:14-23.
 23. Guagliumi G, Costa MA, Sirbu V, et al. Strut coverage and late malapposition with paclitaxel-eluting stents compared with bare metal stents in acute myocardial infarction: optical coherence tomography substudy of the Harmonizing Outcomes with Revascularization and Stents in Acute Myocardial Infarction (HORIZONS-AMI) Trial. *Circulation* 2011;123:274-281.
 24. Stone GW, Ellis SG, Cox DA, et al. A polymer-based, paclitaxel-eluting stent in patients with coronary artery disease. *N Engl J Med* 2004;350:221-231.
 25. Brodie BR, Stuckey T, Downey W, et al. Outcomes and complications with off-label use of drug-eluting stents: results from the STENT (Strategic Transcatheter Evaluation of New Therapies) group. *JACC Cardiovasc Interv* 2008;1:405-414.
 26. de la Torre-Hernandez JM, Alfonso F, Hernandez F, et al. Drug-eluting stent thrombosis: results from the multicenter Spanish registry ESTROFA (Estudio ESpañol sobre TROMbosis de stents FARmacoactivos). *J Am Coll Cardiol* 2008;51:986-990.
 27. Beinart R, Abu Sham'a R, Segev A, et al. The incidence and clinical predictors of early stent thrombosis in patients with acute coronary syndrome. *Am Heart J* 2010;159:118-124.