Optimization of Stent Deployment by Intravascular Ultrasound

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Intravascular ultrasound (IVUS) is a useful diagnostic method that provides valuable information in addition to angiography regarding the coronary vessel lumen, dimensions, plaque burden, and characteristics. The major use of IVUS in coronary intervention is to guide interventional strategies and assess optimal stent deployment. Since the introduction of the drug-eluting stent (DES), concerns about restenosis have decreased. However, high-risk lesion subsets are being routinely treated with DESs, and the incidence of suboptimal results after stent deployment, such as stent underexpansion, incomplete stent apposition, edge dissection, geographic miss, and the risk of stent thrombosis, have correspondingly increased. Thus, optimization of stent deployment under IVUS guidance may be clinically important. In this review, we focus on the potential role of IVUS in stent optimization during percutaneous coronary intervention and its clinical benefits.

Keywords: Angioplasty, balloon; Coronary stenosis; Mortality; Stents; Therapy; Ultrasonography

INTRODUCTION

The coronary angiogram (CAG) remains the gold-standard method for assessing coronary artery disease (CAD). However, the CAG has inherent pitfalls, such as only showing the vessel lumen as an X-ray shadow image, created by the injection of contrast medium, and of often visualizing a "side-view." Thus, the apparent degree of coronary stenosis can be affected by the projection angle due to lesion eccentricity. Additionally, diffuse coronary disease, lesion foreshortening, angulations, calcification, and vessel overlap can be challenges in the angiographic assessment of lesion severity. In some cases, an angiographically normal-looking coronary artery actually shows various degrees of atherosclerotic plaque by intravascular ultrasound (IVUS)

[1,2].

IVUS is an invasive imaging technique used to visualize coronary cross-sectional anatomy and is superior to CAG in assessing vessel size, calcium content, and lesion severity [3]. It also provides complementary procedural information in lesions requiring percutaneous coronary intervention (PCI) when determining adequate stent sizing, and confirming optimal stent deployment and apposition without edge tearing in real time. Thus, a growing number of interventional cardiologists attain optimal procedural results with reduced complications when using IVUS in PCI.

Although the routine use of IVUS in daily PCI remains controversial, stent optimization by IVUS during stenting procedures, especially in the era of drug-eluting stents

Received: January 31, 2012 Accepted: February 2, 2012

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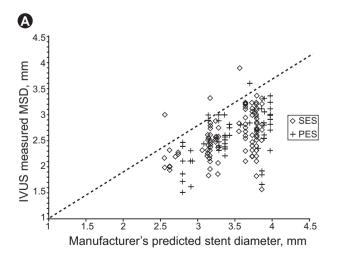
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(DESs), may have an important role in improving longterm clinical outcomes such as stent restenosis and stent thrombosis [4]. In this review, we focus on the potential roles of IVUS in stent optimization during PCI and its clinical benefits.

REALITY OF STENTING PROCEDURES IN DAILY PRACTICE

PCI has been the fastest growing method for the treatment of ischemic CAD over the past three decades. Coronary stents have emerged as the predominant form of PCI and are currently used in more than 90% of PCI procedures. Procedural success of PCI is usually determined by visual estimation by the operator, and usually, angiographic success after PCI is defined as the attainment of residual diameter stenosis of less than 30%, which is generally associated with at least a 20% improvement in diameter stenosis and relief of ischemia [5]. However, such subjective estimation of the severity of coronary artery stenosis is thought to be of limited reliability. Previous IVUS studies have demonstrated that visual estimation or quantitative angiographic analyses of vessel dimension for stent deployment appear inaccurate [6-9]. The post-dilatation clinical comparative (POSTIT) trial was designed to assess the achievement of optimal stent deployment by IVUS, according to normal-to-high pressure balloon dilation after bare metal stent implantation. Among 256 patients, only 14% of cases achieved optimal stent deployment with under 12-atmosphere pressure dilation and only 36% even with higher deployment pressures (> 14 atmospheres) [9]. Another IVUS study with an early-generation DES in 200 patients assessed stent expansion depending on the manufacturer's compliance chart as a guideline. In that study, the DES obtained only 75% of predicted minimal stent diameter and 66% of the predicted minimal stent area (MSA) [10] (Fig. 1). Based on these observations, angiographic success cannot always be linked with optimal stent expansion, despite higher pressure balloon inflation during the stenting procedure. In turn, stent optimization using a high-pressure balloon without IVUS guidance also has been associated with an increased risk of arterial perforation, probably secondary to vessel-balloon mismatch [11]. Thus, the operator should consider using IVUS guidance for high-pressure balloon inflation during stent deployment.

Apposition of stent struts to the vessel wall is also an important facet of stent optimization. Adequate stent expansion and adequate stent strut apposition have been reported to be important factors in reducing repeated revascularization due to stent restenosis or stent thrombosis [12,13]. In the DES era, incomplete stent apposition has been regarded as an important local factor in DES failure, probably due to reduced drug delivery to the vessel wall [14-17]. In a recent report, incomplete stent apposition was significantly associated with vessel/stent mismatch rather than stent underexpansion immediately after stent implantation [18]. Thus, adequate stent sizing by IVUS may be clinically important in preventing incomplete stent ap-



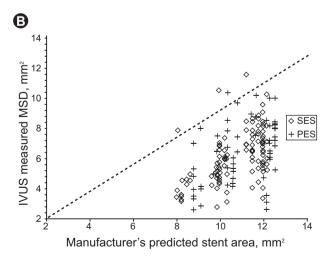


Figure 1. Intravascular ultrasound (IVUS)-measured minimum stent diameter (MSD, A) and minimum stent area (MSA, B) vs. predicted measurements from each manufacturer's compliance charts. SES, sirolimus-eluting stent; PES, paclitaxel-eluting stent. Reprint with permission from Elsevier Health Science Journals [10].

position and in optimizing initial stent deployment.

IMPORTANCE OF STENT OPTIMIZATION

IVUS has been used to detect suboptimal results after apparently angiographically successful stent deployment in both the DES and bare metal stent (BMS) eras (Fig. 2). IVUS predictors that are associated with increased adverse outcomes include smaller MSA, stent underexpansion, stent edge dissection, incomplete stent apposition, and incomplete lesion coverage [19-26]. In the BMS era, a major problem after stent implantation was stent restenosis, and the main mechanism of this phenomenon was a smaller MSA or stent underexpansion [21,26-30]. Several studies in the BMS era showed a beneficial effect of IVUS guidance on post-procedural angiographic results and stent restenosis during long-term follow-up, resulting from a larger MSA with a higher post-dilation balloon pressure [7,19,27,31]. Stent underexpansion, identified by IVUS, can be treated with appropriate post-balloon dilation. IVUS allows more aggressive intervention using a larger diameter balloon with confidence in terms of safety; thus, BMS implantation under IVUS guidance can provide a bigger MSA and more favorable clinical outcomes compared with angiography-guided PCI. DESs have led to a marked reduction in the rate of stent restenosis and the need for repeated revascularization compared with BMSs [32,33]. Because of their efficacy, high-risk lesions and clinical conditions, including bifurcation lesions, long lesions, calcified lesions, left main disease, diabetes, and multivessel

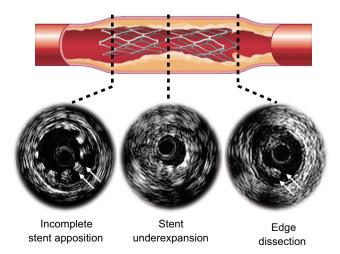


Figure 2. Stent-related complications after stent deployment.

disease, are now being treated routinely with DESs [34,35]. Thus, the risk of stent underexpansion, incomplete stent apposition, and incomplete lesion coverage increases and these suboptimal stent deployment conditions have been reported to be potent IVUS predictors of stent restenosis and stent thrombosis [13,17,24], suggesting that stent implantation under IVUS guidance still has a pivotal role even in the DES era. An important aspect of IVUS is determining appropriate reference segments that provide the landing zone for stent deployment. IVUS examination typically reveals a considerable amount of plaque, even in segments of the vessel that appear "normal" on the angiogram, known as "reference vessel" disease. Quantitative IVUS studies have demonstrated that the segment chosen as the "normal" reference site for the calculation of angiographic percent stenosis has an average of 30-50% of its cross-sectional area occupied by plaque [36]. By IVUS, the definition of reference segment is a cross-sectional image adjacent to the lesion that has < 40% plaque burden [37]. A previous study reported the association between clinical outcomes and longitudinal positioning of the stent in 162 consecutive patients with 180 lesions treated with sirolimus-eluting stent (SES) implantation [38]. In that study, stepwise IVUS criteria primarily targeting plaque burden < 50% were shown to be feasible and improved the rates of stent restenosis and target lesion revascularization (TLR) at 8 months follow-up.

HOW TO OPTIMIZE STENT DEPLOYMENT

IVUS is used frequently in PCI, but the use of IVUS cannot be directly related to stent optimization. Definitive guidelines for IVUS-guided stent optimization are not available and it is still performed at the operator's discretion. So, how do we perform stent optimization using IVUS, and what IVUS criteria are acceptable for current practice?

Although many IVUS criteria for stent optimization have been suggested, the basic concepts underlying them can be summarized briefly as minimizing the occurrence of IVUS-related predictors of adverse events after PCI, including stent underexpansion, incomplete stent apposition, edge dissection, and lesion undercoverage.

Stent underexpansion is defined as an area of inadequate stent expansion compared with the adjacent refer-

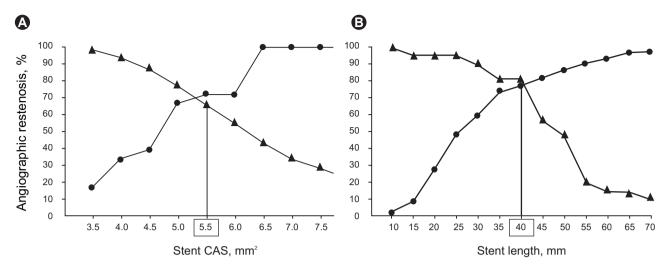


Figure 3. Sensitivity and specificity curves identified optimal cut-off values of final minimum stent cross-sectional area (CSA, A) and stent length (B) that predicted angiographic restenosis after sirolimus-eluting stent implantation: 5.5 mm² for final minimum stent CSA and 40 mm² for intravascular ultrasound-measured stent length. Triangle, sensitivity; circles, specificity. Reprinted with permission from Oxford University Press [22].

ence segments. However, a consensus definition of "adequate" expansion is still lacking.

In the BMS era, several randomized trials used various IVUS criteria for stent deployment optimization, yielding mixed results. The first large multicenter study, called MUSIC (multicenter ultrasound stenting in coronaries study), sought to define specific criteria for optimal stent deployment and demonstrate the feasibility and safety of IVUS-guided stent optimization [39]. In the MUSIC criteria, "adequate" expansion was defined as > 90% of the average reference cross-sectional area (CSA), or > 100% of a smaller reference CSA with complete apposition and symmetric expansion. Another large multicenter trial was AVID (angiographic versus IVUS direct stent placement); its defined IVUS optimization required complete stent apposition with stent CSA > 90% of the distal reference lumen area [40]. The results of the AVID trial showed a significant benefit of IVUS guidance in vessels 2.5-3.5 mm in size and in saphenous vein graft PCI. The TULIP (thrombotic activity evaluation and effects of ultrasound guidance in long intracoronary stent placement) Study also showed significant angiographic and clinical benefits of IVUS guidance using criteria such as stent CSA > distal reference lumen [41]. In contrast to the results mentioned above, the OPTICUS trial (optimization with IVUS to reduce stent restenosis) using the MUSIC criteria for IVUS guidance did not show a significant difference in 6- or 12-month clinical outcomes [42]. However, in this study,

only 56% of stents met all three MUSIC criteria. Many other trials in the BMS era used similar IVUS criteria, and two meta-analyses showed better outcomes of IVUSguided PCI than angiography-guided PCI, especially in terms of in-stent restenosis and target vessel revascularization (TVR), but not in mortality or myocardial infarction [43,44].

In contrast to the BMS era, for IVUS-guided PCI with DES, few randomized studies showing clinical efficacy, preventing TVR or restenosis, or stent optimization under IVUS guidance have been carried out. In the HOME DES trial, optimal stent deployment was defined as complete apposition of the stent struts, no edge dissection, and adequate stent expansion, defined as either MSA > 5.0 mm² or > 90% of the distal reference lumen area [45]. In that study, no significant benefit in terms of TVR or clinical events was reported. A similar finding was also noted in the AVIO (angiography versus IVUS optimization) study in which optimal stent expansion was > 70% of the CSA of the chosen balloon [46]. However, attention should be paid to avoid stent underexpansion. Increasing evidence indicates that IVUS-guided PCI may reduce the risk of stent thrombosis (ST) [47]. Stent underexpansion was demonstrated to be one of the major causes of this disastrous complication, although the cause of ST is multifactorial [13,24,48,49]. In a substudy of the SIRIUS trial, adequate patency was defined as a follow-up IVUS MSA > 4.0 mm². When the adequate post-interventional MSA of SESs was

defined as > 5.0 mm², the positive predictive value of patency was 90%, but the optimal cutoff value of BMS was defined as 6.5 mm [50]. Another study concerning SES failure also supported that MSA < 5.5 mm² and < 5.0 mm² were the most important predictors of SES failure (Fig. 3) [22,51]. Even if SES had a considerably lower optimal MSA threshold than BMS, these studies showed that underexpansion remained the main cause of stent failure in DESs; at least a MSA < 5.0 mm should be avoided in non-left main (LM) lesions. In LM lesions, optimal MSA was reported in the MAIN-COMPARE (revascularization for unprotected left main coronary artery stenosis: comparison of percutaneous coronary angioplasty versus surgical revascularization) study to be > 8.7 mm² to prevent TLR [34].

Edge dissection, which is complicated by lumen narrowing $< 4 \text{ mm}^2$ or dissection angle $\ge 60^\circ$, has been associated with an increased incidence of early ST [49]; thus, additional stents may be needed to prevent ST. However, a minor dissection, detected by IVUS, may not be associated with an increased incidence of ST [52,53]. Although no consensus exists on an optimal strategy, in minor dissection, careful observation without stenting can be helpful.

Overall, the results discussed above encourage ensuring good apposition of stent struts to the vessel wall, such that the stent struts are not surrounded by lumen, adequate stent expansion to obtain MSA at least > 6.5 mm² for BMSs and > 5.0 mm² for DESs or MSA > 90% of the distal reference lumen CSA, and lack of major dissections, intramural hematomas, and geographic misses.

OUTCOMES OF IVUS-GUIDED VERSUS ANGIOGRAPHY-GUIDED PCI

Numerous studies have evaluated the clinical benefits of IVUS-guided PCI compared with angiography-guided PCI in the BMS era [41-44]. The OPTICUS trial showed no significant difference between IVUS- and angiography-guided PCI groups in terms of 6- and 12-month rates of death, myocardial infarction (MI), and TLR in 550 patients meeting the MUSIC criteria [42]. In contrast, the TULIP study demonstrated favorable angiographic and clinical outcomes in patients with long coronary lesions (> 20 mm) treated with a BMS (> 3 mm) under IVUS guidance [41]. In a meta-analysis of 2,193 patients from seven randomized

trials, the rates of 6-month angiographic restenosis and target vessel revascularization were significantly lower in the IVUS-guided PCI group than the angiography-guided group (22% vs. 29%, p = 0.02 and 13% vs. 18%, p < 0.001, respectively), with no difference in the rates of death (2.4% vs. 1.6%, p = 0.18) or MI (3.6% vs. 4.4%, p = 0.51) [44], consistent with a previous meta-analysis [43].

To date, few studies have investigated the clinical benefits of DES optimization under IVUS guidance compared with that of BMSs. The HOME DES IVUS (long-term health outcome and mortality evaluation after invasive coronary treatment using drug-eluting stents with or without IVUS guidance) study was a randomized trial to investigate clinical outcomes of IVUS-guided PCI with DESs. Although the IVUS-guided strategy resulted in the frequent use of adjunctive balloons and a larger size balloon with higher pressure, no significant difference was observed in major adverse cardiac events or stent thrombosis in the study [45]. A similar retrospective study of IVUS-guided stent optimization also showed no significant difference in the incidence of in-stent restenosis or neointimal volume between IVUS- versus angiographyguided PCI [54]. Conversely, a study with a propensitymatched analysis in 884 patients treated with DESs showed a significant reduction in the stent thrombosis rate at both 30 days (0.5% vs. 1.4%, p = 0.046) and 12 months (0.7% vs. 2.0%, p = 0.014) in the IVUS-guided PCI group [47].

Recently, a large "real world" registry from two Korean centers reported long-term outcomes of both IVUS- and angiography-guided PCI using BMS or DES implantation [55]. In total, 8,371 patients who underwent coronary stenting under IVUS guidance (4,627 patients) or angiography guidance (3,744 patients) were consecutively enrolled, and 3-year adverse clinical outcomes were compared between the groups using a Cox regression model and propensity score matching. In the overall population, the 3-year adjusted incidence of mortality was significantly lower in the IVUS-guided PCI group compared with the angiography-guided PCI group (hazard ratio [HR], 0.70; 95% confidence interval [CI], 0.56 to 0.87; p = 0.001) (Fig. 4). In 2,715 matched pairs of the overall population, the IVUS-guided PCI group also had a lower mortality risk (HR, 0.71; 95% CI, 0.56 to 0.90; p = 0.005). However, IVUS-guided PCI did not influence the rates of myocardial infarction, target-vessel revascularization, or

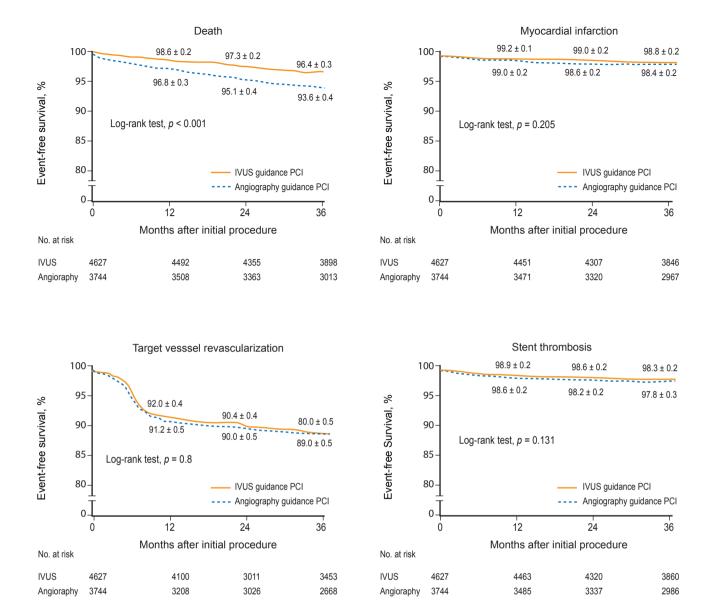


Figure 4. Kaplan-Meier event-free 3-year survival curves for death, myocardial infarction, target-vessel revascularization, and stent thrombosis in 8,371 patients following intravascular ultrasound (IVUS)- (n = 4,627) or angiography- (n = 3,744) guided percutaneous coronary intervention (PCI). Reprinted with permission from John Wiley and Sons [55].

stent thrombosis in the overall or in the 2,715 matchedpair populations. In the DES subpopulation, IVUS guidance significantly reduced the 3-year adjusted mortality rate (HR, 0.55; 95% CI, 0.36 to 0.78; p = 0.001), which was not the case in the BMS subpopulation (HR, 0.79; 95% CI, 0.59 to 1.05; p = 0.10). A propensity score matching analysis of 201 matched patients from the MAIN-COMPARE study also demonstrated the importance of IVUS-guided PCI in unprotected left main disease [34]. In this analysis, significantly lower incidence of 3-year mortality was

noted in the IVUS-guided PCI group compared with the angiography-guided PCI group (4.7% vs. 13.6%, p = 0.048), but no significant difference was detected in the rates of TVR or MI. Notably, this benefit was found only for DES, and the benefit in mortality appeared to be primarily associated with reduced sudden cardiac death related to late stent thrombosis. Similar findings were also observed in patients undergoing PCI of non-left main bifurcations with DESs [35]. Taken together, the benefit of IVUS guidance contributed primarily to decreased rates of stent

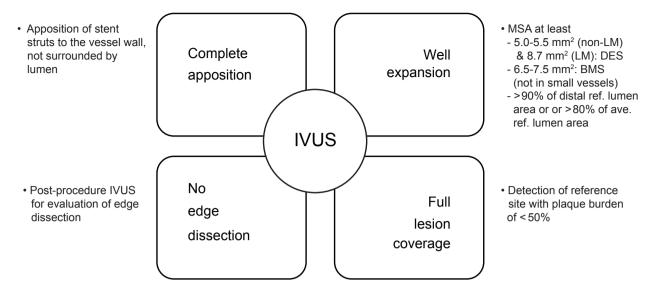


Figure 5. The possible intravascular ultrasound criteria for optimal stent deployment. LM, left main; DES, drug-eluting stent; BMS, bare metal stent.

restenosis and repeated revascularization in the BMS era, whereas reduction of the stent thrombosis rate with possible improvement in mortality have predominated in the DES era.

CONCLUSIONS

IVUS can provide direct cross-sectional images as well as longitudinal images of the coronary vessel wall. It has also contributed to our understanding of mechanisms in coronary atherosclerotic plaques and provided realtime information at stented segments after coronary interventions. Possible criteria for optimal stent deployment by IVUS are complete stent apposition to the vessel wall, adequate stent expansion, and full lesion coverage without edge dissection (Fig. 5). Recent data suggest that IVUS-guided PCI may reduce long-term mortality when compared with angiography-guided PCI, particularly after DES implantation; thus, the clinical importance of IVUSguided PCI raised in the BMS era persists in the DES era. Optimization of stent deployment by IVUS during PCI may be considered as a routine practice in daily PCI, especially for complex lesion intervention.

Conflict of interest

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

REFERENCES

- Nissen SE, Gurley JC, Grines CL, et al. Intravascular ultrasound assessment of lumen size and wall morphology in normal subjects and patients with coronary artery disease. Circulation 1991;84:1087-1099.
- Nissen SE, Gurley JC. Application of intravascular ultrasound for detection and quantitation of coronary atherosclerosis. Int J Card Imaging 1991;6:165-177.
- Waller BF, Pinkerton CA, Slack JD. Intravascular ultrasound: a histological study of vessels during life: the new 'gold standard' for vascular imaging. Circulation 1992;85:2305-2310.
- Schiele F, Meneveau N, Seronde MF, et al. Medical costs of intravascular ultrasound optimization of stent deployment: results of the multicenter randomized 'REStenosis after Intravascular ultrasound STenting' (RESIST) study. Int J Cardiovasc Intervent 2000;3:207-213.
- 5. Smith SC Jr, Feldman TE, Hirshfeld JW Jr, et al. ACC/AHA/ SCAI 2005 Guideline Update for Percutaneous Coronary Intervention-summary article: a report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines (ACC/AHA/SCAI Writing Committee to Update the 2001 Guidelines for Percutaneous Coronary Intervention). Circulation 2006;113:156-175.
- Nakamura S, Colombo A, Gaglione A, et al. Intracoronary ultrasound observations during stent implantation. Circulation 1994;89:2026-2034.
- Gorge G, Haude M, Ge J, et al. Intravascular ultrasound after low and high inflation pressure coronary artery stent implantation. J Am Coll Cardiol 1995;26:725-730.

- 8. Mudra H, Klauss V, Blasini R, et al. Ultrasound guidance of Palmaz-Schatz intracoronary stenting with a combined intravascular ultrasound balloon catheter. Circulation 1994;90:1252-1261.
- 9. Brodie BR, Cooper C, Jones M, Fitzgerald P, Cummins F; Postdilatation Clinical Compartative Study (POSTIT) Investigators. Is adjunctive balloon postdilatation necessary after coronary stent deployment? Final results from the POSTIT trial. Catheter Cardiovasc Interv 2003;59:184-192.
- 10. de Ribamar Costa J Jr, Mintz GS, Carlier SG, et al. Intravascular ultrasound assessment of drug-eluting stent expansion. Am Heart J 2007;153:297-303.
- 11. Alfonso F, Goicolea J, Hernandez R, et al. Arterial perforation during optimization of coronary stents using high-pressure balloon inflations. Am J Cardiol 1996;78:1169-1172.
- 12. Mintz GS, Weissman NJ. Intravascular ultrasound in the drugeluting stent era. J Am Coll Cardiol 2006;48:421-429.
- 13. Cook S, Wenaweser P, Togni M, et al. Incomplete stent apposition and very late stent thrombosis after drug-eluting stent implantation. Circulation 2007;115:2426-2434.
- 14. Hwang CW, Wu D, Edelman ER. Physiological transport forces govern drug distribution for stent-based delivery. Circulation 2001;104:600-605.
- 15. Papafaklis MI, Chatzizisis YS, Naka KK, Giannoglou GD, Michalis LK. Drug-eluting stent restenosis: Effect of drug type, release kinetics, hemodynamics and coating strategy. Pharmacol Ther 2011 Dec 22 [Epub]. http://dx.doi.org/10.1016/ j.pharmthera.2011.12.006.
- 16. Hwang CW, Edelman ER. Arterial ultrastructure influences transport of locally delivered drugs. Circ Res 2002;90:826-832.
- 17. Windecker S, Meier B. Late coronary stent thrombosis. Circulation 2007;116:1952-1965.
- 18. Kume T, Waseda K, Ako J, et al. Intravascular ultrasound assessment of postprocedural incomplete stent apposition. J Invasive Cardiol 2012;24:13-16.
- 19. Fitzgerald PJ, Oshima A, Hayase M, et al. Final results of the Can Routine Ultrasound Influence Stent Expansion (CRUISE) study. Circulation 2000;102:523-530.
- 20. Uren NG, Schwarzacher SP, Metz JA, et al. Predictors and outcomes of stent thrombosis: an intravascular ultrasound registry. Eur Heart J 2002;23:124-132.
- 21. Doi H, Maehara A, Mintz GS, et al. Impact of post-intervention minimal stent area on 9-month follow-up patency of paclitaxeleluting stents: an integrated intravascular ultrasound analysis from the TAXUS IV, V, and VI and TAXUS ATLAS Workhorse, Long Lesion, and Direct Stent Trials. JACC Cardiovasc Interv 2009;2:1269-1275.
- 22. Hong MK, Mintz GS, Lee CW, et al. Intravascular ultrasound predictors of angiographic restenosis after sirolimus-eluting stent implantation. Eur Heart J 2006;27:1305-1310.

- 23. Cheneau E, Leborgne L, Mintz GS, et al. Predictors of subacute stent thrombosis: results of a systematic intravascular ultrasound study. Circulation 2003;108:43-47.
- 24. Fujii K, Carlier SG, Mintz GS, et al. Stent underexpansion and residual reference segment stenosis are related to stent thrombosis after sirolimus-eluting stent implantation: an intravascular ultrasound study. J Am Coll Cardiol 2005;45:995-998.
- 25. Kimura M, Mintz GS, Carlier S, et al. Outcome after acute incomplete sirolimus-eluting stent apposition as assessed by serial intravascular ultrasound. Am J Cardiol 2006;98:436-442.
- 26. Hong MK, Lee CW, Kim JH, et al. Impact of various intravascular ultrasound criteria for stent optimization on the sixmonth angiographic restenosis. Catheter Cardiovasc Interv 2002;56:178-183.
- 27. Albiero R, Rau T, Schlüter M, et al. Comparison of immediate and intermediate-term results of intravascular ultrasound versus angiography-guided Palmaz-Schatz stent implantation in matched lesions. Circulation 1997;96:2997-3005.
- 28. Hoffmann R, Mintz GS, Mehran R, et al. Intravascular ultrasound predictors of angiographic restenosis in lesions treated with Palmaz-Schatz stents. J Am Coll Cardiol 1998;31:43-49.
- 29. Kasaoka S, Tobis JM, Akiyama T, et al. Angiographic and intravascular ultrasound predictors of in-stent restenosis. J Am Coll Cardiol 1998;32:1630-1635.
- 30. Moussa I, Moses J, Di Mario C, et al. Does the specific intravascular ultrasound criterion used to optimize stent expansion have an impact on the probability of stent restenosis? Am J Cardiol 1999;83:1012-1017.
- 31. Blasini R, Neumann FJ, Schmitt C, Bokenkamp J, Schomig A. Comparison of angiography and intravascular ultrasound for the assessment of lumen size after coronary stent placement: impact of dilation pressures. Cathet Cardiovasc Diagn 1997;42:113-119.
- 32. 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.
- 33. Colombo A, Drzewiecki J, Banning A, et al. Randomized study to assess the effectiveness of slow- and moderate-release polymer-based paclitaxel-eluting stents for coronary artery lesions. Circulation 2003;108:788-794.
- 34. 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.
- 35. Kim SH, Kim YH, Kang SJ, et al. Long-term outcomes of intravascular ultrasound-guided stenting in coronary bifurcation lesions. Am J Cardiol 2010;106:612-618.
- 36. Mintz GS, Painter JA, Pichard AD, et al. Atherosclerosis in angiographically "normal" coronary artery reference segments: an intravascular ultrasound study with clinical correlations. J Am

- Coll Cardiol 1995;25:1479-1485.
- Weissman NJ, Palacios IF, Nidorf SM, Dinsmore RE, Weyman AE. Three-dimensional intravascular ultrasound assessment of plaque volume after successful atherectomy. Am Heart J 1995;130:413-419.
- 38. Morino Y, Tamiya S, Masuda N, et al. Intravascular ultrasound criteria for determination of optimal longitudinal positioning of sirolimus-eluting stents. Circ J 2010;74:1609-1616.
- 39. de Jaegere P, Mudra H, Figulla H, et al. Intravascular ultrasound-guided optimized stent deployment: Immediate and 6 months clinical and angiographic results from the Multicenter Ultrasound Stenting In Coronaries Study (MUSIC Study). Eur Heart J 1998;19:1214-1223.
- 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.
- 41. Oemrawsingh PV, Mintz GS, Schalij MJ, Zwinderman AH, Jukema JW, van der Wall EE. Intravascular ultrasound guidance improves angiographic and clinical outcome of stent implantation for long coronary artery stenoses: final results of a randomized comparison with angiographic guidance (TULIP Study). Circulation 2003;107:62-67.
- Mudra H, di Mario C, de Jaegere P, et al. Randomized comparison of coronary stent implantation under ultrasound or angiographic guidance to reduce stent restenosis (OPTICUS Study).
 Circulation 2001;104:1343-1349.
- 43. Casella G, Klauss V, Ottani F, Siebert U, Sangiorgio P, Bracchetti D. Impact of intravascular ultrasound-guided stenting on long-term clinical outcome: a meta-analysis of available studies comparing intravascular ultrasound-guided and angiographically guided stenting. Catheter Cardiovasc Interv 2003;59:314-321.
- 44. Parise H, Maehara A, Stone GW, Leon MB, Mintz GS. Metaanalysis of randomized studies comparing intravascular ultrasound versus angiographic guidance of percutaneous coronary intervention in pre-drug-eluting stent era. Am J Cardiol 2011;107:374-382.
- 45. Jakabein J, Spacek R, Bystron M, et al. Long-term health outcome and mortality evaluation after invasive coronary treatment using drug eluting stents with or without the IVUS guidance. Randomized control trial: HOME DES IVUS. Catheter Cardiovasc Interv 2010;75:578-583.

- 46. Colombo A, Caussin C, Presbitero P, Chieffo A. AVIO: a prospective, randomized trial of intravascular-ultrasound guided compared to angiography guided stent implantation in complex coronary lesions [abstract]. J Am Coll Cardiol 2010;56:xvii.
- 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.
- 48. Okabe T, Mintz GS, Buch AN, et al. Intravascular ultrasound parameters associated with stent thrombosis after drug-eluting stent deployment. Am J Cardiol 2007;100:615-620.
- 49. Choi SY, Witzenbichler B, Maehara A, et al. Intravascular ultrasound findings of early stent thrombosis after primary percutaneous intervention in acute myocardial infarction: a Harmonizing Outcomes with Revascularization and Stents in Acute Myocardial Infarction (HORIZONS-AMI) substudy. Circ Cardiovasc Interv 2011;4:239-247.
- 50. 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.
- 51. Takebayashi H, Kobayashi Y, Mintz GS, et al. Intravascular ultrasound assessment of lesions with target vessel failure after sirolimus-eluting stent implantation. Am J Cardiol 2005;95:498-502.
- 52. Hong MK, Park SW, Lee NH, et al. Long-term outcomes of minor dissection at the edge of stents detected with intravascular ultrasound. Am J Cardiol 2000;86:791-795, A9.
- 53. Liu X, Tsujita K, Maehara A, et al. Intravascular ultrasound assessment of the incidence and predictors of edge dissections after drug-eluting stent implantation. JACC Cardiovasc Interv 2009;2:997-1004.
- 54. Park SM, Kim JS, Ko YG, et al. Angiographic and intravascular ultrasound follow up of paclitaxel- and sirolimus-eluting stent after poststent high-pressure balloon dilation: from the poststent optimal stent expansion trial. Catheter Cardiovasc Interv 2011;77:15-21.
- 55. 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 2011 Jul 29 [Epub]. http://dx.doi.org/10.1002/ccd.23279.