

Coronary Imaging Modalities for Forecasting the “Eruption of the Volcano”

Nobuaki Suzuki, MD, PhD; Ken Kozuma, MD, PhD

Coronary plaque rupture can be likened to a volcanic eruption, given its abrupt occurrence and the “natural disaster” that follows the episode. Numerous cardiovascular researchers have aimed to identify vulnerable atherosclerotic plaques, which lead to plaque rupture and coronary thrombosis, because they are the major cause of acute myocardial infarction (AMI) and sudden cardiac death.¹ Morphological studies from autopsy have suggested the importance of necrotic core size, inflammation, and fibrous cap thickness.² These clinical features are similar to the amount of magma, volcanic tremor, and the distance from magma to the ground surface forecasting the explosion of the volcano. Thin-cap fibro-atheroma (TCFA), which is characterized by a large necrotic core with an overlying thin-fibrous cap measuring $<65\ \mu\text{m}$, is known to lead to AMI.³

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Coronary imaging modalities are playing an important role in interventional cardiology.⁴ According to the contemporary all-comer trials, intravascular ultrasound (IVUS) is the most frequently used and is overriding all other imaging modalities in the daily clinical settings of Japan.⁵ Recent pivotal data have shown that IVUS guidance reduces the clinical events in percutaneous coronary intervention (PCI).⁶ The major strength of IVUS is its ability to quantitatively and qualitatively assess atherosclerotic plaque, which enables adequate stent expansion in PCI procedures. In particular, radiofrequency signal-based IVUS (RF-IVUS) has facilitated automated plaque assessment and contributed to the building of evidence regarding the efficacy of PCI strategies.⁷

On the other hand, IVUS has reduced spatial resolution compared with the emerging newer modality of optical coherence tomography (OCT).⁸ One of the major limitations of IVUS in regard to poor spatial resolution is that IVUS is incapable of clearly detecting the fibrous cap thickness of plaques. The advantages and disadvantages of IVUS vs. OCT in the evaluation of vulnerable plaques are summarized in Table. In particular, OCT can show subtle changes such as neovascularization.⁹

In this issue of the Journal, Koga et al¹⁰ present crisp data showing that coronary lesions with greater absolute necrotic areas categorized by iMap-IVUS are closely associated with OCT-derived TCFA (OCT-TCFA). The authors evaluated 86 coronary lesions from 73 patients with stable angina pectoris

Table. Advantages and Disadvantages of IVUS vs. OCT in the Evaluation of Vulnerable Plaques

	IVUS	OCT
Evaluation of “micro” findings		
Fibrous cap thickness	×	◎
Neovascularization	×	○
Macrophages	×	○
Evaluation of “macro” findings		
Thrombus	△	◎
Spotty calcification	◎	◎
Positive remodeling	◎	×
Necrotic area	◎	△
Clinical utility		
Market penetration	◎	△
For patients with chronic kidney disease	◎	△
For ostial lesions	◎	×

◎, excellent; ○, good; △, fair; ×, poor; IVUS, intravascular ultrasound; OCT, optical coherence tomography.

using iMap-IVUS and OCT. They defined OCT-TCFA as lipid-rich plaques with $<65\ \text{mm}$ -thick fibrous caps, which were subsequently identified in 22 (26%) lesions. Significantly larger percentages of necrotic area, absolute lipidic and necrotic areas, and a smaller percentage of fibrotic area were found in OCT-TCFA than in non-TCFA. Multivariate analysis showed that absolute necrotic area was an independent predictor of OCT-TCFA. The area under the receiver-operator characteristics curve of the absolute necrotic area required to identify OCT-TCFA was 0.86. The sensitivity, specificity, positive, and negative predictive values of absolute necrotic area $\geq 7.3\ \text{mm}^2$ for identifying OCT-TCFA were 77%, 88%, 68%, and 92%, respectively. The data from the present study are comparable with those of previous similar investigations using virtual histology-IVUS, which is a variation of the RF-IVUS system.¹¹

The results of the present study are indeed promising, particularly for interventionists who perform IVUS-guided PCI. On the basis of this study, it is possible to assess the risk of future coronary events in daily clinical settings by reviewing the absolute necrotic area measured using iMap-IVUS. It may also be possible to clarify the pharmaceutical stabilization of vulnerable plaques by analyzing the serial changes of absolute necrotic area as a surrogate marker. By considering the several

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Department of Medicine, Teikyo University School of Medicine, Tokyo, Japan

Mailing address: Ken Kozuma, MD, PhD, Department of Medicine, Teikyo University School of Medicine, 2-11-1 Kaga, Itabashi-ku, Tokyo 173-8605, Japan. E-mail: PXE00364@nifty.com

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limitations of OCT, the evidence of the present study will likely lead to much future clinical evidence in daily clinical settings. For example, OCT-guided PCI requires higher levels of contrast medium, whereas a surrogate marker for the detection of TCFA using iMap-IVUS assessment enables clinical studies of patients with chronic kidney disease, which is known as a powerful risk of ischemic events. Furthermore, new tissue characterization imaging techniques, such as near-infrared spectroscopy, will improve IVUS imaging in the detection of vulnerable plaque.¹² Therefore, IVUS should continue to be a valuable imaging modality in the daily clinical setting.

There are several limitations associated with the study. First, the present data only show that the abundant necrotic area detected using iMap-IVUS frequently exists where OCT-TCFA appears,¹⁰ and therefore must be validated with respect to clinical relevance. Notably, a previous report suggested that most of the vulnerable plaques identified using intracoronary imaging modalities may be clinically silent, and/or the prevalence of vulnerable plaques identified using imaging modalities is overestimated.⁸ Regarding the possibility of overestimation of TCFA, it is known that a low or lack of signal in plaques located in regions oblique to an eccentric wire can be explained by the intrinsic properties of time-domain OCT rather than by the presence of lipid components.¹³ Second, the accuracy of IVUS measurement is a concern. A previous study showed that the measurement value determined using IVUS may be greater than reality.¹⁴ Third, the risk prediction for primary prevention of plaque rupture using invasive imaging modality is unrealistic. We should keep in mind that modern imaging technologies visualizing vulnerable plaques have yet to demonstrate improved risk prediction compared with conventional methods.¹ To foresee the future “explosion of volcano” more effectively, the innovation of non-invasive modalities also needs to be promoted.¹⁵

References

1. Arbab-Zadeh A, Fuster V. The myth of the “vulnerable plaque”: Transitioning from a focus on individual lesions to atherosclerotic disease burden for coronary artery disease risk assessment. *J Am Coll Cardiol* 2015; **65**: 846–855.
2. Finn AV, Nakano M, Narula J, Kolodgie FD, Virmani R. Concept of vulnerable/unstable plaque. *Arterioscler Thromb Vasc Biol* 2010; **30**: 1282–1292.
3. Coletta J, Suzuki N, Nascimento BR, Bezerra HG, Rosenthal N, Guagliumi G, et al. Use of optical coherence tomography for accurate characterization of atherosclerosis. *Arq Bras Cardiol* 2010; **94**: 250–254, 268–272, 254–259 (in English, Portuguese, Spanish).
4. Nagoshi R, Shinke T, Otake H, Shite J, Matsumoto D, Kawamori H, et al. Qualitative and quantitative assessment of stent restenosis by optical coherence tomography: Comparison between drug-eluting and bare-metal stents. *Circ J* 2013; **77**: 652–660.
5. Natsuaki M, Kozuma K, Morimoto T, Kadota K, Muramatsu T, Nakagawa Y, et al. Biodegradable polymer biolimus-eluting stent versus durable polymer everolimus-eluting stent: A randomized, controlled, noninferiority trial. *J Am Coll Cardiol* 2013; **62**: 181–190.
6. Witzencbichler B, Maehara A, Weisz G, Neumann FJ, Rinaldi MJ, Metzger DC, et al. Relationship between intravascular ultrasound guidance and clinical outcomes after drug-eluting stents: The assessment of dual antiplatelet therapy with drug-eluting stents (ADAPT-DES) study. *Circulation* 2014; **129**: 463–470.
7. Kawaguchi R, Oshima S, Jingu M, Tsurugaya H, Toyama T, Hoshizaki H, et al. Usefulness of virtual histology intravascular ultrasound to predict distal embolization for ST-segment elevation myocardial infarction. *J Am Coll Cardiol* 2007; **50**: 1641–1646.
8. Fujii K, Hao H, Ohyanagi M, Masuyama T. Intracoronary imaging for detecting vulnerable plaque. *Circ J* 2013; **77**: 588–595.
9. Suzuki N, Kozuma K, Kyono H, Otsuki S, Fu Q, Hosogoe N, et al. Predominant microvessel proliferation in coronary stent restenotic tissue in patients with diabetes: Insights from optical coherence tomography image analysis. *Int J Cardiol* 2013; **168**: 843–847.
10. Koga S, Ikeda S, Miura M, Yoshida T, Nakata T, Koide Y, et al. iMap-intravascular ultrasound radiofrequency signal analysis reflects plaque components of optical coherence tomography-derived thin-cap fibroatheroma. *Circ J* 2015; **79**: 2231–2237.
11. Kubo T, Nakamura N, Matsuo Y, Okumoto Y, Wu X, Choi SY, et al. Virtual histology intravascular ultrasound compared with optical coherence tomography for identification of thin-cap fibroatheroma. *Int Heart J* 2011; **52**: 175–179.
12. Brugaletta S, Sabaté M. Assessment of plaque composition by intravascular ultrasound and near-infrared spectroscopy: From PROSPECT I to PROSPECT II. *Circ J* 2014; **78**: 1531–1539.
13. Suzuki N, Guagliumi G, Bezerra HG, Sirbu V, Rosenthal N, Musumeci G, et al. The impact of an eccentric intravascular ImageWire during coronary optical coherence tomography imaging. *EuroIntervention* 2011; **6**: 963–969.
14. Kubo T, Akasaka T, Shite J, Suzuki T, Uemura S, Yu B, et al. OCT compared with IVUS in a coronary lesion assessment: The OPUS-CLASS study. *JACC Cardiovasc Imaging* 2013; **6**: 1095–1104.
15. Noguchi T, Kawasaki T, Tanaka A, Yasuda S, Goto Y, Ishihara M, et al. High-intensity signals in coronary plaques on noncontrast T1-weighted magnetic resonance imaging as a novel determinant of coronary events. *J Am Coll Cardiol* 2014; **63**: 989–999.