## The relationship of the underlying lipidic plaque at the implanted newer-generation drug-eluting stents with future stent-related events: insights from the REASSURE-NIRS registry

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**Background:** Lipid-rich plaque is an important substrate causing acute coronary events. Near-infrared spectroscopy (NIRS) imaging has been shown to visualize lipidic coronary plaque at non-culprit site associated with future coronary events. Given that histopathological studies reported that the unstable plaque underlying the implanted drug-eluting stent (DES) could cause neoatherosclerosis formation, we hypothesized that NIRS-based evaluation of lipidic plaque burden behind the implanted DES may clinically predict the occurrence of stent failure in patients with CAD receiving PCI.

**Purpose:** We aimed to investigate the relationship of stent-related events' risk with lipidic plaque materials behind the implanted DES imaged by NIRS/intravascular ultrasound (NIRS/IVUS) imaging.

**Methods:** The REASSURE-NIRS registry is an on-going multi-center registry to enroll CAD subjects receiving NIRS/IVUS-guided PCI. In this registry data, 406 lesions in 379 CAD subjects (ACS/non-ACS=150/229) receiving new-generation DES were analyzed. Minimum stent area (MSA) after PCI and maximum lipid-core-burden index in any 4mm-segment within the implanted stents (in-stent maxLCBI4mm) were measured. A 3-year lesion-oriented composite outcome [LOCO: culprit lesion-related MI + ischemia-driven target lesion revascularization (ID-TLR)] was compared in subjects stratified according to the tertile of in-stent maxLCBI4mm.

Results: The mean value of in-stent maxLCBI4mm was 221, and 17% of lesions exhibited in-stent maxLCBI4mm >400. Patients with a greater in-stent maxLCBI4mm were more likely to exhibit a higher LDL-C level (p=0.026) with a longer stent length (p<0.001) and a smaller MSA (p=0.033) (Picture 1). Over 95% of entire study subjects received a statin. During the observational period (median=726 days), the frequency of LOCO up to 3 years was 3.4% in entire study subjects (culprit lesion-related MI=1.0%, ID-TLR=2.8%). Kaplan-Meier curve analysis demonstrated that the occurrence of LOCO did not increase in association with instent maxLCBI4mm (log-rank p-value=0.25, Picture 2). In addition, in-stent maxLCBI4mm did not associate with each component of LOCO (culprit lesion-related MI: p=0.502, ID-TLR: p=0.872). Receiver Operating Characteristic analysis revealed that the predictive ability of in-stent maxLCBI4mm for the occurrence of LOCO was unsatisfactorily (c-statistics=0.486).

**Conclusion:** The amount of underlying lipidic materials at culprit lesions receiving new-generation DES implantation did not necessarily predict future stent-related events. Clinical significance of maxLCBI4mm behind the implanted DES may be different from that at naïve non-culprit plaques.

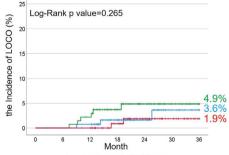
|                                     | in-stent maxLCBI4mm |                |               |         |
|-------------------------------------|---------------------|----------------|---------------|---------|
|                                     | Low tertile         | Middle tertile | High tertile  | p value |
|                                     | (0-124)             | (126-272)      | (276-799)     |         |
| Background characteristics          |                     |                |               |         |
| Age, years                          | 70 ± 10             | 68 ±11         | 69 ± 11       | 0.415   |
| Male                                | 79%                 | 81%            | 80%           | 0.940   |
| ACS                                 | 36%                 | 37%            | 45%           | 0.309   |
| Statin use at discharge             | 95%                 | 95%            | 97%           | 0.579   |
| Laboratory data at administra       | tion                |                |               |         |
| LDL-C, mg/dL                        | 94 ± 35             | 95 ± 37        | 105 ± 35      | 0.026   |
| HbA1c, %                            | $6.2 \pm 0.7$       | $6.3 \pm 0.9$  | $6.3 \pm 0.9$ | 0.622   |
| Angiographical findings after       | PCI                 |                |               |         |
| LAD                                 | 52%                 | 64%            | 62%           | 0.100   |
| Total stent length, mm              | 25.6 ± 11.1         | 30.9 ± 14.4    | 32.7 ± 15.8   | <0.001  |
| Percent diameter stenosis, %        | 7 (2-12)            | 7 (3-14)       | 7 (1-14)      | 0.815   |
| Intravascular ultrasound findi      | ings after PCI      |                |               |         |
| Minimum stent area, mm <sup>2</sup> | 6.34 ± 2.26         | 5.80 ± 2.18    | 5.69 ± 2.07   | 0.033   |

LAD; left anterior-descending artery, LCBI; lipid-core burden index, PCI; percutaneous coronary intervention

Background and lesion characteristics

## LOCO; Lesion-oriented clinical outcomes

=culprit lesion-related MI + ischemia-driven target lesion revascularization



in-stent maxLCBI4mm Low (0-124) Middle (126-272) High (276-799)

LCBI; lipid-core burden index

Kaplan-Meier analysis for LOCO