

The Estimated Absolute Risk of Coronary Artery Disease and Subclinical Atherosclerosis

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See article vol. 28: 1266-1274

Over several decades, epidemiological and experimental studies have revealed that atherosclerotic diseases are multifactorial diseases. Traditionally, their risk has been expressed using relative risks¹⁾. However, because atherosclerotic risk factors frequently co-exist in a single individual, global assessment of atherosclerotic risk is considered to be clinically preferable, leading to the evaluation of absolute risk that takes multiple risk factors into account. Thus, several organizations including the European Society of Cardiology and European Atherosclerosis Society (ESC/EAS), and the American College of Cardiology and the American Heart Association (ACC/AHA) have proposed guidelines using risk charts to estimate the absolute risk^{2,3)}. These changes have resulted in an increased awareness about the importance of absolute risk assessment and comprehensive management of risk factors. Japan Atherosclerosis Society (JAS) also proposed the comprehensive risk management guidelines for prevention of atherosclerotic cardiovascular diseases^{4,5)}. In the JAS Guidelines 2012, the 10-year absolute risk of death due to coronary artery disease (CAD), based on the NIPPON DATA80 Risk Assessment Chart, was applied to stratify individuals into different levels of risk categories, which were used to determine interventions for primary prevention of CAD. In 2017, JAS Guidelines 2017 applied the 10-year absolute risk of CAD incidence based on the Suita score for the stratification^{6,7)}.

Numerous equations have been invented for the estimation of absolute risk and published globally. In addition to the predictivity of the outcome, there are several points that need to be considered when we apply the equation to clinical guidelines. First, the equation should be developed based on the

representative population samples to which the equation is to be applied, because the absolute risk would be estimated from the model that incorporates the average levels of risk factors in the population. The outcome criteria, such as the subtypes of atherosclerotic cardiovascular diseases to be included and whether the incidence and/or mortality are to be assessed, are key factors that need to be considered when we apply an equation. Furthermore, the differences between the population for which the risk assessment score had been created and the present population to which the score would be applied should be noted, as there may be differences in the level of atherosclerosis or the incidence and mortality of CAD. For example, NIPPON DATA80 Risk Assessment Chart was based on the study (NIPPON DATA80) for which baseline survey was conducted in 1980 when statin was not available, and CAD mortality was assessed as the outcome when the coronary intervention procedures were limited. Thus, newly applying the established estimation equation to guidelines requires a set of processes that clarifies the existence or non-existence of discrepancies between the current and previous populations.

Subclinical atherosclerosis could be a possible parameter to be used to assess the existence of such discrepancies between populations, because it shares risk factors with hard cardiovascular events such as myocardial infarction and stroke, and it strongly predicts the risks of occurrence of such events. Thus, subclinical atherosclerosis parameters such as coronary artery calcification and carotid intima-media thickness have been examined to be concordant with the historically established estimation equations that were applied in the guidelines^{8,9)}. In this issue of the Journal of Atherosclerosis and Thrombosis, Sata *et al.* have provided additional evidence regarding the categories in the comprehensive lipid and risk management guidelines proposed by the JAS (JAS

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Received: April 30, 2021 Accepted for publication: May 6, 2021

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Guidelines 2017)¹⁰, using a measure of subclinical atherosclerosis. They investigated the association of the cardio-ankle vascular index (CAVI), an arterial stiffness parameter, to the estimated 10-year absolute risk of CAD and the risk categories of JAS Guidelines 2017, in 1,973 general Japanese. Their cross-sectional data obtained in 2014 showed that the mean CAVI (proportion of CAVI \geq 9.0 and multivariable adjusted Odds ratio of CAVI \geq 9.0) was concordant with the estimated 10-year absolute risk of CAD when JAS Guidelines 2017 were employed. These findings suggest that the 10-year absolute risk of CAD estimation equation in JAS Guidelines 2017 was almost applicable to the contemporary populations, even though the baseline survey of the Suita study (in which Suita score was developed) had been conducted around 1990. They also studied about the appropriateness of category III of the guidelines which includes individuals with diabetes mellitus, chronic kidney disease, non-cardiogenic cerebral infarction, and peripheral artery disease all together irrespective of the estimated absolute risk value. For men, they observed that the mean CAVI values, proportion of CAVI \geq 9.0 and multivariable adjusted Odds ratio of CAVI \geq 9.0 were similar between the groups with/without diabetes mellitus, chronic kidney disease, and peripheral artery disease. Their findings supported the classification in JAS Guidelines 2017. However, for women, those in category III with diabetes mellitus had higher mean CAVI than those without diabetes mellitus, although the mean CAVI were similar between the groups with/without chronic kidney disease. These diseases were very heterogeneous in severity. Because their study population largely consisted of relatively healthy individuals, further studies are required to clarify whether patients with severe conditions are to be classified into category III or into secondary prevention groups which require more rigorous risk factor control.

In summary, Sata *et al.* presented that a subclinical measure of atherosclerosis, or arterial stiffness assessed by CAVI was concordant with the estimated 10-year absolute risk of CAD as described in JAS Guidelines 2017, suggesting that the estimation equation could be applicable for the present population. Further studies, using other subclinical atherosclerosis parameters in the present population, are also recommended in order to support the estimated absolute risk equation and classification of JAS Guidelines 2017. However, lipid and comprehensive risk management would be beneficial, and the individuals classified in the higher risk category should have their risk factors adequately controlled in order to prevent cardiovascular disease.

Conflict of Interest

None.

References

- 1) Ueshima H, Sekikawa A, Miura K, Turin TC, Takashima N, Kita Y, Watanabe M, Kadota A, Okuda N, Kadowaki T, Nakamura Y and Okamura T. Cardiovascular disease and risk factors in Asia: a selected review. *Circulation*, 2008; 118: 2702-2709
- 2) Catapano AL, Graham I, De Backer G, Wiklund O, Chapman MJ, Drexel H, Hoes AW, Jennings CS, Landmesser U, Pedersen TR, Reiner Ž, Riccardi G, Taskinen MR, Tokgozoglu L, Verschuren WM, Vlachopoulos C, Wood DA and Zamorano JL. 2016 ESC/EAS Guidelines for the Management of Dyslipidaemias: The Task Force for the Management of Dyslipidaemias of the European Society of Cardiology (ESC) and European Atherosclerosis Society (EAS) Developed with the special contribution of the European Association for Cardiovascular Prevention & Rehabilitation (EACPR). *Atherosclerosis*, 2016; 253: 281-344
- 3) Goff DC, Jr., Lloyd-Jones DM, Bennett G, Coady S, D'Agostino RB, Gibbons R, Greenland P, Lackland DT, Levy D, O'Donnell CJ, Robinson JG, Schwartz JS, Shero ST, Smith SC, Jr., Sorlie P, Stone NJ, Wilson PW, Jordan HS, Nevo L, Wnek J, Anderson JL, Halperin JL, Albert NM, Bozkurt B, Brindis RG, Curtis LH, DeMets D, Hochman JS, Kovacs RJ, Ohman EM, Pressler SJ, Sellke FW, Shen WK, Smith SC, Jr. and Tomaselli GF. 2013 ACC/AHA guideline on the assessment of cardiovascular risk: a report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines. *Circulation*, 2014; 129: S49-S73
- 4) Kinoshita M, Yokote K, Arai H, Iida M, Ishigaki Y, Ishibashi S, Umemoto S, Egusa G, Ohmura H, Okamura T, Kihara S, Koba S, Saito I, Shoji T, Daida H, Tsukamoto K, Deguchi J, Dohi S, Dobashi K, Hamaguchi H, Hara M, Hiro T, Biro S, Fujioka Y, Maruyama C, Miyamoto Y, Murakami Y, Yokode M, Yoshida H, Rakugi H, Wakatsuki A, Yamashita S, Committee for E and Clinical Management of A. Japan Atherosclerosis Society (JAS) Guidelines for Prevention of Atherosclerotic Cardiovascular Diseases 2017. *J Atheroscler Thromb*, 2018; 25: 846-984
- 5) Teramoto T, Sasaki J, Ishibashi S, Birou S, Daida H, Dohi S, Egusa G, Hiro T, Hirobe K, Iida M, Kihara S, Kinoshita M, Maruyama C, Ohta T, Okamura T, Yamashita S, Yokode M, Yokote K and Japan Atherosclerosis S. Executive summary of the Japan Atherosclerosis Society (JAS) guidelines for the diagnosis and prevention of atherosclerotic cardiovascular diseases in Japan -2012 version. *J Atheroscler Thromb*, 2013; 20: 517-523
- 6) NIPPON DATA80 Research Group. Risk Assessment Chart for Death From Cardiovascular Disease Based on a 19-Year Follow-up Study of a Japanese Representative Population NIPPON DATA80. *Circ J*, 2006; 70: 1249-

- 1255
- 7) Nishimura K, Okamura T, Watanabe M, Nakai M, Takegami M, Higashiyama A, Kokubo Y, Okayama A and Miyamoto Y. Predicting coronary heart disease using risk factor categories for a Japanese urban population, and comparison with the framingham risk score: the suita study. *J Atheroscler Thromb*, 2014; 21: 784-798
 - 8) Kathiresan S, Larson MG, Keyes MJ, Polak JF, Wolf PA, D'Agostino RB, Jaffer FA, Clouse ME, Levy D, Manning WJ and O'Donnell CJ. Assessment by cardiovascular magnetic resonance, electron beam computed tomography, and carotid ultrasonography of the distribution of subclinical atherosclerosis across Framingham risk strata. *Am J Cardiol*, 2007; 99: 310-314
 - 9) Kadota A, Miura K, Okamura T, Fujiyoshi A, Ohkubo T, Kadowaki T, Takashima N, Hisamatsu T, Nakamura Y, Kasagi F, Maegawa H, Kashiwagi A and Ueshima H. Carotid intima-media thickness and plaque in apparently healthy Japanese individuals with an estimated 10-year absolute risk of CAD death according to the Japan Atherosclerosis Society (JAS) guidelines 2012: the Shiga Epidemiological Study of Subclinical Atherosclerosis (SESSA). *J Atheroscler Thromb*, 2013; 20: 755-766
 - 10) Sata M, Okamura T, Harada S, Sugiyama D, Kuwabara K, Hirata A, Takeuchi A, Iida M, Kato S, Matsumoto M, Kurihara A and Takebayashi T. Association of the estimated coronary artery incidence risk according to the Japan Atherosclerosis Society Guidelines 2017 with cardio ankle vascular index. *J Atheroscler Thromb*, 2021; 28: 1266-1274