

Is the Atherogenic Index of Plasma (AIP) a Cardiovascular Disease Marker?

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SOUHRN

V diagnostice a stanovování prognózy kardiovaskulárních onemocnění (KVO) již byla použita řada ukazatelů. Aterogenní index plazmy (AIP) se vypočítává jako logaritmus molárních koncentrací triglyceridu (TG) a lipoproteinů o vysoké hustotě (high-density lipoprotein, HDL). Vynikající predikční hodnota AIP je výsledkem těsného vztahu mezi AIP a velikostí lipoproteinových částic. AIP lze jednoduše vypočítat pomocí typického lipidového profilu. Jedná se o přesnější prediktor velikosti lipoproteinových částic než velikosti částic jednotlivých lipidů nebo poměru TG a HDL.

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ABSTRACT

Numerous indicators have been used to diagnose and prognosticate cardiovascular disease (CVD). The atherogenic index of plasma (AIP) is a logarithmic conversion of triglyceride (TG) to high-density lipoprotein (HDL) molar concentrations. The close association between AIP and the size of lipoprotein particles may account for its excellent predictive value. AIP may be computed simply using a typical lipid profile. It is a better predictor of lipoprotein particle size than that of individual lipids or the ratio of TG to HDL.

Introduction

Cardiovascular disease (CVD) causes one third of fatalities globally, and the risk factors for CVD are growing.¹ The most fundamental effort in combating the CVD epidemic is preventing risk factors and understanding their interactions. Obesity, inactivity, poor nutrition, and smoking are major CVD risk factors.² Among them, the plasma lipid profile is the key CVD predictor.³

The LDL/HDL ratio is widely used to assess CV risk since there is a clear link between high LDL-C and low HDL-C.^{4,5} Conversely, elevated TG levels have been linked to increased LDL-C particles and CV risk.⁶ Thus, atherogenic

dyslipidemia (high LDL/HDL ratio and elevated TG) raises CV risk.⁷

Better atherogenic dyslipidemia indicators are replacing old rates.⁵ The atherogenic index of plasma (AIP) is

$$\text{Atherogenic index of plasma} = \log \frac{[\text{Triglyceride}]}{[\text{HDL cholesterol}]}$$

Fig. 1 – The AIP calculation.

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a powerful predictor of atherosclerosis.⁸⁻¹⁰ AIP stands for protective and atherogenic lipoprotein interaction.¹¹ AIP is calculated as $\log(\text{TG}/\text{HDL-C})$ (Fig. 1).¹² AIP readings below 0.11 are related to a low risk of CVD, whereas those between 0.11 and 0.21 are associated with an intermediate and increasing risk.¹³ We conducted a review to assess the relationship between AIP and CVD.

A novel CV predictive biomarker

Regular exercise has been shown to reduce the risk of heart disease, stroke, hypertension, and diabetes.¹⁴ According to Tariq M. Ali Rajab et al., diabetes dyslipidemia is characterized by a rise in TG and a reduction in HDL-C.¹⁵ Another study indicated that hyperglycemia, aberrant lipid profiles, and HTN all contributed to atherosclerosis progression.¹⁶ Various studies have shown a high association between AIP and lipoprotein particle size, making AIP an indication of atherogenic lipoprotein status.¹²

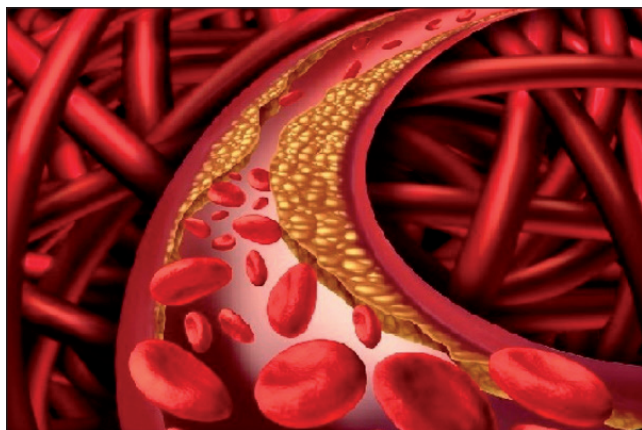


Fig. 2 – AIP may predict acute coronary events.

AIP may be used to evaluate CV risk factors and predict acute coronary events (Fig. 2).¹⁷ AIP may also be used as a screening technique when all atherogenic parameters are normal.⁵ The AIP index rises with changes in other CV risk factors. According to Niroumand et al., regular exercise and a balanced diet are suggested. AIP should be used regularly as a CVD indicator, especially in patients with high CV risk factors. It's a simple statistic to calculate, especially when other lipid levels are normal.¹⁸

For example, a study of 340 healthy women found a significant connection between AIP and the Framingham risk score (FRS), indicating an AIP involvement in early coronary artery disease (CAD) identification.¹⁹ A recent study of young individuals identified an association between AIP and CV risk factors.²⁰ A higher AIP value enhanced the incidence of CAD in HIV patients and RA/SLE women.^{21,22} The AIP value may better reflect metabolic dysfunction than a single metric like LDL-C. This subfraction of LDL-C is known as sdLDL.²³

Unsurprisingly, a higher AIP score may raise the risk of CAD in adults. It has to be replicated in a larger adult sample to determine the AIP's potential as a CAD risk predictor.²⁴ Fernández-Macias et al. show

that AIP may be used as a CVD biomarker in Mexico, because strong relationships were found between asymmetric dimethylarginine (ADMA) and adipocyte-fatty acid binding protein (FABP4) concentrations and AIP in people.²⁵

Cardiovascular diseases

The severity of OSA illness corresponds with AIP but not sleep quality or quantity. Although this was an experimental goal, AIP did not outperform other lipid measures in detecting dyslipidemia in OSA. The AIP value can vary drastically in OSA.²⁶ In OSA, the apnea-hypopnea index (AHI) is linked to AIP and the apoB/apoAI ratio. According to Cao et al., the AIP and apoB/apoAI ratio rose with OSA severity, perhaps contributing to the significant CVD risk in OSA.²⁷

AIP may be a reliable indicator of acute ischemic stroke, particularly the stroke subtype of large-artery atherosclerosis.²⁸ According to Kim et al., high AIP predicts IHD in non-diabetic Koreans. As an alternative to isolated TG levels or other lipid markers, AIP is increasingly used to stratify distinct cardiometabolic risks.²⁹ Research by Abacoglu et al. is significant in revealing that AIP is superior to TG/HDL-C and equal to stent length in predicting ST. It also proposes reducing LDL-C as well as serum TG and HDL levels to avoid ST. If AIP is found in regular lipid measurements, treatment choices may need to be changed or expanded.³⁰ AIP was related to angiographic progression of CAD irrespective of recognized risk factors.³¹

T2DM was linked to AIP transitions. Preventive measures are required to address T2DM at an early dyslipidemic stage.³² When it comes to AIP and coronary heart disease risk factors, Korean men are no exception. Obesity, diabetes, and lipid metabolic indices are all beneficial. Consumption of milk and dairy products was greater among those with low AIP. The AIP quartiles differed in total fat intake but not in saturated fat consumption. AIP recommends further research on the causal link between coronary heart disease risk factors and dietary intake. It is also vital to include both male and female genders in research for comparative analysis.³³

Based on a large-scale clinical experiment with 10,251 randomized T2DM patients, AIP is a strong predictor of CV events. Major adverse cardiovascular events (MACEs) were more common in diabetics with high AIP. The outcomes of this research helped design a unique MACE bio-indicator for use in high-risk populations.³⁴ AIP is better than normal lipid profiles in assessing CAD risk, and this finding may offer a theoretical foundation for successful CV prophylaxis in T2DM patients.³⁵

Due to its capacity to detect metabolic dysfunction, AIP may be used to predict poor outcomes in non-diabetic individuals following percutaneous coronary intervention (PCI). The HR-AIP index association seems J-shaped. The AIP index cutoff of 0.18 might be utilized to calculate hazard ratios (HRs) for non-diabetic patients following PCI. We need further multi-center research on non-diabetic individuals with metabolic dysfunction to confirm the AIP index's utility in this setting.³⁶

Adolescents from Spain had the highest AIP. The ROC AUC findings imply that all atherogenic indicators can predict MetS. More research is required to fully understand this forecasting ability.³⁷ The best diagnostic cutoff values for CAD and SS >23 were 2.035 and 2.23, respectively. Using AIP as a biomarker may help avoid CAD in Chinese patients. In clinical diagnosis and therapy, clinicians should consider patients' AIP values.³⁸

Conclusion

This review is noteworthy since it indicates that AIP is more predictive of CVD than TG or HDL-C. This also means that, in addition to LDL-C, serum TG and HDL levels must be examined while attempting to prevent CVD. If AIP is explored in conjunction with conventional lipid measurements, treatment techniques may need to be altered or increased. This review is important because it sheds light on multicenter, prospective studies with a lot of patients.

Conflict of interest

The authors declare that there is no conflict of interest regarding the publication of this paper.

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References

- Askin L, Askin HS, Tanriverdi O. The caveolin levels in cardiovascular disease. *Cor Vasa* 2021;63:579–582.
- Mensah GA, Brown DW. An overview of cardiovascular disease in the United States. *Health Aff (Millwood)* 2007;26:38–48.
- Kanth PS, Patil BS, Bagali S, et al. Atherogenic Index as a Predictor of Cardiovascular Risk among Women with Different Grades of Obesity. *IJCRIMPH* 2012;4:1767–1774.
- Igweh JC, Nwagha IU, Okaro JM. The Effects of Menopause on the Serum lipid profile of Normal Females of South East Nigeria. *Nig J Physiol Sci* 2005;20:48–53.
- Mudhaffar SK. Atherogenic Index of Plasma (AIP) As a Parameter in Predicting Cardiovascular Risk in Males Compared To the Conventional Dyslipidemic Indices (Cholesterol Ratios). *Karbala J Med* 2013;6:1506–1513.
- Guerin M, Legoff W, Lassel TS, et al. Proatherogenic role of elevated CE transfer from HDL to VLDL and dense LDL in type 2 diabetics. *Arterioscler Thromb Vasc Biol* 2001;21:282–287.
- Manninen V, Tenkanen L, Koshinen P, et al. Joint effects of serum triglyceride and LDL cholesterol and HDL cholesterol concentrations on coronary heart disease risk in the Helsinki Heart Study: implications for treatment. *Circulation* 2002;85:37–45.
- Njajou O, Kanaya AM, Holvoet P, et al. Association between oxidized LDL-C, obesity and type 2 diabetes. *Diabetes Metab Res Rev* 2009;25:733–739.
- Daniels LB, Laughlin G, Sarno MJ. Lp-PLA2 is an independent predictor of incident coronary heart disease in apparently healthy older population. *J Am Coll Cardiol* 2008;51:913–919.
- Ali Rajab TM. Comparative study for Atherogenic Index of Plasma (AIP) in patient with type 1 Diabetes Mellitus, type 2 Diabetes Mellitus, Betathalassemia and Hypothyroidism. *Int J Chem Res* 2012;2.
- Dobiášová M, Frohlich J, Šedová M, et al. Cholesterol esterification and atherogenic index of plasma correlate with lipoprotein size and findings on coronary angiography. *J Lipid Res* 2011;52:566–571.
- Dobiášová M, Frohlich J. The plasma parameter log (TG/HDL) as an atherogenic index: correlation with lipoprotein particle size and esterification rate in apoB-lipoprotein-depleted plasma (FER HDL). *Clin Biochem* 2001;34:583–588.
- Dobiasova M. AIP – atherogenic index of plasma as a significant predictor of cardiovascular risk: from research to practice. *Vnitr Lek* 2006;52:64–71.
- Flier JS. Biology of Obesity. In: Kasper DL, Fauci AS, Longo DL, Braunwald E, Hauser SL, Jameson JL. eds. *Harrison's Principles of Internal Medicine*. 17th ed. New York, NY: McGraw-Hill; 2008:462–464.
- Kavitha G, Ramani G, Dhass PK, Aruna RM. Oxidative stress, interleukin (il-6) and atherogenic index of plasma in diabetic nephropathy. *I JABPT* 2011;2:211–217.
- Wassink AM, Graaf VG, Sabita Y, et al. Metabolic syndrome and incidence of type 2 diabetes in patients with manifest vascular disease. *Diabetes Vasc Dis Res* 2008;5:114–122.
- Dobiasova M, Urbanova Z, Samanek M. Relation between particle size of HDL and LDL Lipoproteins and cholesterol Esterification rate. *Physiol Res* 2005;54:159–165.
- Niroumand S, Khajedaluae M, Khadem-Rezaian M, et al. Atherogenic Index of Plasma (AIP): A marker of cardiovascular disease. *Med J Islam Repub Iran* 2015;29:240.
- Fernandez-Macias JC, Ochoa-Martinez AC, Varela-Silva JA, PerezMaldonado IN. Atherogenic index of plasma: novel predictive biomarker for cardiovascular illnesses. *Arch Med Res* 2019;50:285–294.
- Kammar-Garcia A, Lopez-Moreno P, Hernandez-Hernandez ME, et al. Atherogenic index of plasma as a marker of cardiovascular risk factors in Mexicans aged 18 to 22 years. *Proc (Bayl Univ Med Cent)* 2020;34:22–27.
- Noumegni SR, Nansseu JR, Bigna JJ, et al. Atherogenic index of plasma and 10-year risk of cardiovascular disease in adult Africans living with HIV infection: a cross-sectional study from Yaounde, Cameroon. *JRSM Cardiovasc Dis* 2017;6:2048004017740478.
- Hammam N, Abdel-Wahab N, Gheita TA. Atherogenic index of plasma in women with rheumatoid arthritis and systemic lupus erythematosus: a 10-year potential predictor of cardiovascular disease. *Curr Rheumatol Rev* 2021;17:122–130.
- Ivanova EA, Myasoedova VA, Melnichenko AA, et al. Small dense low-density lipoprotein as biomarker for atherosclerotic diseases. *Oxid Med Cell Longev* 2017;2017:1273042.
- Wu J, Zhou Q, Wei Z, et al. Atherogenic Index of Plasma and Coronary Artery Disease in the Adult Population: A Meta-Analysis. *Front Cardiovasc Med* 2021;8:817441.
- Fernández-Macías JC, Ochoa-Martínez AC, Varela-Silva JA, Pérez-Maldonado IN. Atherogenic Index of Plasma: Novel Predictive Biomarker for Cardiovascular Illnesses. *Arch Med Res* 2019;50:285–294.
- Bikov A, Meszaros M, Kunos L, et al. Atherogenic Index of Plasma in Obstructive Sleep Apnoea. *J Clin Med* 2021;10:417.
- Cao B, Fan Z, Zhang Y, Li T. Independent association of severity of obstructive sleep apnea with lipid metabolism of atherogenic index of plasma (AIP) and apoB/apoA1 ratio. *Sleep Breath* 2020;24:1507–1513.
- Liu H, Liu K, Pei L, et al. Atherogenic Index of Plasma Predicts Outcomes in Acute Ischemic Stroke. *Front Neurol* 2021;12:741754.
- Kim JJ, Yoon J, Lee YJ, et al. Predictive Value of the Atherogenic Index of Plasma (AIP) for the Risk of Incident Ischemic Heart Disease among Non-Diabetic Koreans. *Nutrients* 2021;13:3231.
- Özcan AÖ, Yıldırım A, Koyunsever NY, et al. Relationship between atherogenic index of plasma and stent thrombosis in patients with acute coronary syndrome. *Anatol J Cardiol* 2022;2:112–117.
- Shui X, Chen Z, Wen Z, et al. Association of Atherogenic Index of Plasma With Angiographic Progression in Patients With Suspected Coronary Artery Disease. *Angiology* 2022;73:927–935.

32. Yi Q, Ren Z, Bai G, et al. The longitudinal effect of the atherogenic index of plasma on type 2 diabetes in middle-aged and older Chinese. *Acta Diabetol* 2022;59:269–279.
33. Shin HR, Song S, Cho JA, Ly SY. Atherogenic Index of Plasma and Its Association with Risk Factors of Coronary Artery Disease and Nutrient Intake in Korean Adult Men: The 2013-2014 KNHANES. *Nutrients* 2022;14:1071.
34. Fu L, Zhou Y, Sun J, et al. Atherogenic index of plasma is associated with major adverse cardiovascular events in patients with type 2 diabetes mellitus. *Cardiovasc Diabetol* 2021;20:201.
35. Zhou K, Qin Z, Tian J, et al. The Atherogenic Index of Plasma: A Powerful and Reliable Predictor for Coronary Artery Disease in Patients With Type 2 Diabetes. *Angiology* 2021;72:934–941.
36. Zheng Y, Li C, Yang J, et al. Atherogenic index of plasma for non-diabetic, coronary artery disease patients after percutaneous coronary intervention: a prospective study of the long-term outcomes in China. *Cardiovasc Diabetol* 2022;21:29.
37. Fernández-Aparicio Á, Perona JS, Schmidt-RioValle J, Padez C, González-Jiménez E. Assessment of Different Atherogenic Indices as Predictors of Metabolic Syndrome in Spanish Adolescents. *Biol Res Nurs* 2022;24:163–171.
38. Wang L, Chen F, Xiaoqi C, et al. Atherogenic Index of Plasma Is an Independent Risk Factor for Coronary Artery Disease and a Higher SYNTAX Score. *Angiology* 2021;72:181–186.