## Research Article

# The Relationship between Inter-Arm Blood Pressure Difference and Coronary Artery Disease Severity Calculated by the SYNTAX Score

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Received 4 February 2018; Revised 28 July 2018; Accepted 28 August 2018; Published 13 September 2018

Academic Editor: Franco Veglio

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*Objectives.* The inter-arm systolic blood pressure difference (IASBPD) is closely related to cardiovascular mortality and morbidity. The SYNTAX score indicates the extent and complexity of coronary artery disease, which are determined by coronary angiography. The aim of our study is to examine the relationship between the IASBPD (which is easily calculated in routine practice) and the SYNTAX score. *Methods.* 104 patients were included in this cross-sectional study. The IASBPD was calculated by blood pressure measurements obtained simultaneously from both arms. The SYNTAX score was calculated by coronary angiography. *Results.* Patients were divided into two groups: those with a high SYNTAX score ( $\geq 20$ ) and those with a low SYNTAX score (< 20). The mean IASBPD values were significantly higher in the group with a high SYNTAX score ( $\geq 20$ ) (p<0.001). The patients with IASBPD $\geq 10$  were more likely to have a high SYNTAX score compared to the patients with IASBPD<10 (p<0.001). Multiple logistic regression analysis revealed that only the IASBPD values were found to be independently associated with high SYNTAX score (OR: 1.717 (CI: 1.307-2.257), p<0.001). *Conclusion.* The IASBPD values obtained by only blood pressure measurements are closely related to the extent of coronary artery disease.

#### 1. Introduction

The Synergy between PCI with TAXUS and Cardiac Surgery (SYNTAX) score is a commonly used tool for both shortand long-term risk classifications in patients undergoing percutaneous coronary intervention (PCI) [1, 2]. Although its prognostic value has been proven, it only provides an assessment on the image. It does not include clinical variables that are closely related to long- and short-term outcomes [3, 4]. However, the scores including clinical data require a lot of data and increase the risk of occurrence of incorrect results due to their complex structures.

The inter-arm systolic blood pressure difference (IAS-BPD) is associated with increased subclinical atherosclerosis in different areas of the arterial tree. According to a meta-analysis, inter-arm systolic blood pressure difference (IASBPD)≥10 mmHg was found to be related to cerebrovascular disease, vascular damage, and increased cardiovascular mortality [5]. It is a noninvasive and inexpensive indicator of arterial stiffness. It was found that the anklebrachial index (ABI) showed both the extent of coronary artery disease and the post-PCI prognosis regardless of the SYNTAX score [6]. Studies have shown that blood pressure is not only related to narrowing of the coronary arteries but also affects the structure and content of atheromatous plagues [7]. However, IASBPD measurement can be simply applied to all patients without the need for Doppler. Moreover, hypertension (HT) guidelines recommend measuring blood pressure in both arms. In this study, we examined the relationship between the IASBPD and the SYNTAX score and evaluated the hypothesis that information can be received about the extent of coronary artery disease with a simple clinical instrument and thus about the prognosis.

#### 2. Method

Our study had a cross-sectional study design. The IASBPD was calculated by obtaining blood pressure measurements in patients before coronary angiography (CAG). From among patients undergoing planned elective CAG because of the symptoms of ischemic heart disease and ischemic findings on the treadmill test or myocardial perfusion scintigraphy, 104 patients who met inclusion/exclusion criteria were included consecutively in the study. We excluded patients with a history of myocardial infarction, PCI, or coronary arterial bypass surgery. Patients with a history of congenital heart disease, severe valvular disease, cardiomyopathy, atrial fibrillation, impulse conduction disturbances or aortic coarctation, hemiplegia, and pulseless disease and those with history of transradial coronary intervention were not included. Informed consent was obtained from all patients. The study was approved by Institutional Ethics Committee and the investigation conformed to the principles of the Declaration of Helsinki.

2.1. Angiographic Analysis and Calculation of the SYNTAX Score. All patients underwent CAG through transfemoral approach. Coronary anatomy was examined by two experienced invasive cardiologists. As a result of these evaluations, the same opinion was reached. The SYNTAX score for each patient was calculated by scoring all coronary lesions with a stenosis diameter as 50% in vessels 1.5 mm, using the SYN-TAX score algorithm, which is described in full elsewhere [8]. SYNTAX scores  $\geq$ 20 were categorized as "high", SYNTAX scores <20 were categorized as "low". To calculate SYNTAX score II, the anatomical SS was combined with the following variables: age, creatinine clearance, LVEF, presence of ULM disease, peripheral vascular disease, female sex, and chronic obstructive pulmonary disease [4]

Height and body weight values were measured to calculate the Body Mass Index (BMI). HT was defined as systolic blood pressure  $\geq$  140 mm Hg and/or diastolic blood pressure  $\geq$  90 mm Hg or use of antihypertensive medication. Diabetes Mellitus was defined as fasting blood glucose level  $\geq$ 126 mg/dL or use of insulin or an oral hypoglycemic medication. Coronary artery disease was assessed from patients' medical reports. Hyperlipidemia was defined by a total cholesterol of greater than 240 mg/dL and triglyceride of greater than 200 mg/dL. Current smoking was defined as smoking at least one cigarette per day in the year preceding the examination.

2.2. Measurement of Blood Pressure. The patients were made to sit for 5 minutes, with feet flat on the ground and back supported before BP measurements were taken. Two separate oscillometric blood pressure (BP) monitors with their own cuffs were used randomly, placed at the appropriate location on the arm and measured by the same trained nurse [9]. BP devices were validated by European Society of Hypertension (Omron HEM-7001-E; Omron Corp., Tokyo, Japan). An appropriate cuff was selected for each individual depending on their arm circumference. Simultaneous BP measurements were made three times by changing cuff and therefore BP monitor between arms each with 5 minutes interval. The IABPD was defined as the absolute difference between the blood pressure measurements in each arm [9]. Exaggerated IASBPD was defined as > 10 mmHg difference in systolic blood pressure (SBP) between two arms at all three measurements [10, 11]. Mean BP was calculated for both right arm and left arm using all 3 measurements. The difference was noted as the mean absolute IASBPD [9]. In order to evaluate reproducibility, BP measurements for 50 patients were performed by the same nurse. The intraclass correlation coefficient was calculated as 0.88 for BP difference between arms.

2.3. Laboratory Analyses. Blood collection was performed after a 12-hour fasting. All biochemical analyses were performed within the first two hours after blood collection. Glucose, creatinine, total cholesterol, high-density lipoprotein cholesterol, low-density lipoprotein cholesterol (LDL), and triglyceride levels were measured. Blood count analysis was performed using an automated cell counter (Sysmex XE-2100, Kobe, Japan).

2.4. Statistical Analysis. SPSS 17.0 for Windows (SPSS 17.0, Chicago, Illinois) software package was used in all analyses. The continuous variables were expressed as mean ± standard deviation (SD) (for parameters with normal distribution) and median (interquartile range, IQR) (for parameters without normal distribution), and categorical variables were expressed as percentages. The Chi-Square test was used to compare the categorical variables between the groups. Analysis of normality was performed with the Kolmogorov-Smirnov test. The independent samples t-test was used to compare continuous variables with normal distribution, and the Mann–Whitney U test was used to compare the continuous variables without a normal distribution. Correlations were sought with the Spearman's and Pearson correlation test. Binary logistic regression analysis (backward stepwise method) was performed to identify independently associated factors with high SYNTAX score ( $\geq 20$ ). Variables with a **p** value <0.1 in univariate analysis were incorporated in the binary logistic regression analysis. Receiver-operating characteristic (ROC) curve analysis was performed to determine the optimum IASBPD cut-off value for predicting high SYNTAX score. A two-sided **p** value <0.05 was considered significant within a 95% confidence interval (CI).

#### 3. Results

A total of 104 patients were included in our study. The mean age of the patients was  $58.9 \pm 9.5$  years. 67 (64.4%) were male. While 99 (95.2%) patients had dominant right hand, the remaining 5 (4.8%) were left-handed. The mean SYNTAX value for all patients was  $9.5 \pm 8.2$ . In addition, 15 of the patients (14.4%) had a SYNTAX score  $\geq 20$ .

When evaluated for the inter-arm blood pressure difference, the mean right arm systolic blood pressure (SBP) (p=0.012) and mean left arm systolic blood pressure (SBP)

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TABLE 1: Comparison of groups with high and low SYNTAX scores in terms of demographic and biochemical data.

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Variables	SYNTAX≥20 N=15	SYNTAX<20 N=89	р
Gender (male) (n, %)	13 (%86.7)	54 (%60.7)	0.052
Right-hand dominance	12 (%85.7)	81 (%96.4)	0.092
(n, %)			
Diabetes mellitus	4 (%26.7)	38 (%42.7)	0.242
(n, %)			
Hypertension	6 (%40.0)	50 (%56.2)	0.245
(n, %)			
Hyperlipidemia ( <b>n, %</b> )	3 (%20.0)	23 (%25.8)	0.629
Smoking (n, %)	5 (%35.7)	28 (%31.8)	0.776
BMI (kg/m <sup>2</sup> )	28.9 (3.6)	29.1 (6.8)	0.085
Mean of right arm SBP (mmHg)	134.4±12.6	124.4±14.4	0.012
Mean value of left arm SBP (mmHg)	133.4±9.7	124.6±13.7	0.019
Mean value of right arm	74.6±5.0	72.9±10.4	0.562
DBP (mmHg)			
Mean value of left arm	73.6±15.7	72.0±13.1	0.682
DBP (mmHg)			
Mean IASBPD (mmHg)	11.6±2.3	5.5±2.6	<0.001
Mean IADBPD (mmHg)	5.3 ±1.7	4.2±2.9	0.188
Exaggerate IASBPD	12 (%80)	4 (%4.5)	<0.001
(>10 mmHg)	12 (7000)	1 (701.3)	<0.001
Glucose (mg/dL)	120.6 (23.9)	122.9 (34.9)	0.816
TC (mg/dL)	207.4 (70.6)	195.0 (50.0)	0.410
LDL-C (mg/dL)	152.3 (45.4)	125.4 (38.2)	0.028
HDL-C (mg/dL)	43.3±9.2	44.2 ±11.7	0.993
Triglyceride (mg/dL)	174.0±17.0	162.3±21.2	0.773
Creatinine (mg/dL)	$0.8 \pm 0.1$	0.7±0.2	0.819
SYNTAX score	27.3±3.1	6.5±3.9	<0.001

Variables that fit a normal distribution were expressed as mean  $\pm$  SD. Variables that did not fit a normal distribution were expressed as median (interquartile range).

BMI: body mass index, DBP: diastolic blood pressure, HDL-C: high-density lipoprotein cholesterol, IASBPD: inter-arm systolic blood pressure difference, IADBPD: inter-arm diastolic blood pressure difference, LDL-C: low-density lipoprotein cholesterol, SBP: systolic blood pressure, TC: total cholesterol.

(p=0.019) and mean IASBPD values were significantly higher in the group with a high SYNTAX score ( $\geq$ 20) compared to the group with a low SYNTAX score ( $\geq$ 20) (Table 1). The number of those with IASBPD $\geq$ 10 was significantly greater in the group with a high SYNTAX score ( $\geq$ 20) compared to the group with a low SYNTAX score ( $\leq$ 20) (12 (80.0%) versus 4 (4.5%), p<0.001). When biochemical data were examined, LDL levels were found to be higher in the group with a high SYNTAX score ( $\geq$ 20) compared to the group with a low SYNTAX score ( $\leq$ 20) (152.3 (45.4) versus 125.4 (38.2), p=0.028).

Considering the relation between inter-arms blood pressure and SNYTAX II score, among the people whose SYN-TAX II score are more than 20, mean of right arm SBP (135.1 $\pm$ 19.3 versus 126.4 $\pm$ 14.8 mmHg, p=0.020), mean value of left arm SBP (134.5 $\pm$ 19.2 versus 122.7 $\pm$ 17.0 mmHg, p=0.002), and mean IASBPD (10.6 $\pm$ 6.6 versus 6.8 $\pm$ 5.1 mmHg, p=0.003) values are measured higher.

Correlation was found between SYNTAX score and mean right arm SBP (r=0.333, p=0.001), IASBPD (r=0.736, p<0.001), and LDL levels (r=0.228, p=0.025).

Univariate regression analysis showed that LDL, IASBPD, gender, BMI, and exaggerated IASBPD (>10 mmHg) were significant at the p<0.1 level. However, since IASBPD and exaggerated IASBPD (>10 mmHg) are relative variables, the IASBPD values from these two variables were included in multiple logistic regression analysis. Multiple logistic regression analysis revealed that only the IASBPD values were

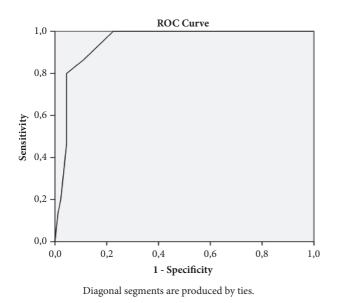


FIGURE 1: ROC (receiver-operating characteristic) curve in predicting the optimum inter-arm systolic blood pressure difference cut-off value for predicting high SYNTAX score.

found to be independently associated with high SYNTAX score (OR: 1.717 (CI: 1.307-2.257), p<0.001).

The receiver-operating characteristic analysis showed that the optimal IASBPD cut-off value for predicting high SYNTAX score was 8.5 mmHg with a sensitivity of 86.0% and specificity of 89.1% (AUC = 0.94; 95% CI: 0.90-0.99) (Figure 1).

#### 4. Discussion

In this study, we found a close association between the interarm blood pressure difference and the SYNTAX score (which indicates the severity and extent of coronary artery disease). We can obtain important data about the atherosclerosis severity of coronary artery disease and thus about the prognosis by blood pressure measurement in both arms, which is a simple and inexpensive method routinely recommended by HT guidelines.

Blood pressure measurement in both arms is recommended for each patient. It is important for determining the true value of blood pressure and thus the need for antihypertensive treatment especially in those with cardiovascular diseases. As in our study, simultaneous blood pressure measurement in both arms has a vital significance in determining the IASBPD [12]. Sequential measurements have a tendency to overestimate difference compared to simultaneous measurements. Recent studies have confirmed that the IASBPD is closely associated with cardiovascular mortality and peripheral arterial disease [9, 10]. The relationship between poor outcome and arterial stiffness in cardiovascular diseases has been demonstrated in different patient populations [13, 14]. The increase in arterial stiffness is systemic but often affects the arterial tree at different levels [14, 15]. Possibly, the changes in the amount of damage in

elastic fibers are due to atherosclerosis at different levels in the arterial system, and the different aortic reflection wave responses cause significant IASBPD.

The SYNTAX score provides information on the choice of optimal revascularization type through coronary anatomy and on the short- and long-term prognosis in those undergoing coronary interventions. However, the SYNTAX score gives a prognosis according to only coronary anatomy. On the other hand, systemic atherosclerosis has proven to be closely associated with prognosis in those undergoing PCI [16]. It was seen that patients with peripheral arterial disease had a poor post-PCI prognosis and that combining the anklebrachial index (ABI) and the SYNTAX score improved prognostic prediction [6, 17]. This demonstrates that evaluation of systemic atherosclerosis with the SYNTAX score improves prognostic prediction. Although the ABI is a noninvasive method, it is a more complex method than blood pressure measurement in both arms, which requires the use of Doppler ultrasound devices. The fact was that a close relationship between the IASBPD and the SYNTAX score was about the extent and complexity of coronary artery in our daily practice and about the prediction of PCI results with additional studies. Although the clinical SYNTAX score including clinical variables gives more clear prognostic information than the classic SYNTAX score, the clinical SYNTAX score includes complex calculation methods. The IASBPD is easy and simple to measure and is a convenient method for daily practice.

#### 5. Limitations

Our study has some limitations. The number of the patients was relatively low. It was conducted in a single center. Since the SYNTAX score is relatively low, cut-off values may be different for groups with complex and diffuse coronary artery disease. Routine radiological evaluation was not performed for the aorta and its branches in those with IASBPD>10 mmHg. However, performing radiological examinations in all these patients is not appropriate in terms of both time and cost. In addition, most of these patients do not have pathologies such as anomaly, arthritis, and aneurysm in their arterial system [18]. Power analysis was not performed to calculate the sample size.

#### 6. Conclusion

We found that the IASBPD that would be obtained by a simple and easy blood pressure measurement was associated with the extent of coronary artery disease obtained by coronary angiography. This situation, which is a reflection of systemic involvement of atherosclerosis, gives information about the extent of coronary artery disease in patients and also may provide us with information on prognosis as a result of prospective studies to be performed.

#### **Data Availability**

The data used to support the findings of this study are available from the corresponding author upon request.

#### **Conflicts of Interest**

There are no conflicts of interest regarding the publication of this article.

#### References

- P. W. Serruvs, M.-C. Morice, A. P. Kappetein et al., "Percutaneous coronary intervention versus coronary-artery bypass grafting for severe coronary artery disease," *The New England Journal of Medicine*, vol. 360, no. 10, pp. 961–972, 2009.
- [2] M. Valgimigli, P. W. Serruys, K. Tsuchida et al., "Cyphering the Complexity of Coronary Artery Disease Using the Syntax Score to Predict Clinical Outcome in Patients With Three-Vessel Lumen Obstruction Undergoing Percutaneous Coronary Intervention," *American Journal of Cardiology*, vol. 99, no. 8, pp. 1072–1081, 2007.
- [3] S. Garg, G. Sarno, H. M. Garcia-Garcia et al., "A new tool for the risk stratification of patients with complex coronary artery disease the clinical SYNTAX score," *Circulation: Cardiovascular Interventions*, vol. 3, no. 4, pp. 317–326, 2010.
- [4] V. Farooq, D. van Klaveren, E. W. Steyerberg et al., "Anatomical and clinical characteristics to guide decision making between coronary artery bypass surgery and percutaneous coronary intervention for individual patients: Development and validation of SYNTAX score II," *The Lancet*, vol. 381, no. 9867, pp. 639–650, 2013.
- [5] C. E. Clark, R. S. Taylor, A. C. Shore, O. C. Ukoumunne, and J. L. Campbell, "Association of a difference in systolic blood pressure between arms with vascular disease and mortality: A systematic review and meta-analysis," *The Lancet*, vol. 379, no. 9819, pp. 905–914, 2012.
- [6] Y. Ueki, T. Miura, Y. Miyashita et al., "Predictive Value of Combining the Ankle-Brachial Index and SYNTAX Score for the Prediction of Outcome After Percutaneous Coronary Intervention (from the SHINANO Registry)," *American Journal* of Cardiology, vol. 117, no. 2, pp. 179–185, 2016.
- [7] M. Iannaccone, G. Quadri, S. Taha et al., "Prevalence and predictors of culprit plaque rupture at OCT in patients with coronary artery disease: A meta-Analysis," *European Heart Journal—Cardiovascular Imaging*, vol. 17, no. 10, pp. 1128–1137, 2016.
- [8] G. Sianos, M-A. Morel, and A. P. Kappetein, "The SYNTAX Score: an angiographic tool grading the complexity of coronary artery disease," *EuroIntervention: Journal of EuroPCR in Collab*oration with the Working Group on Interventional Cardiology of the European Society of Cardiology, vol. 1, pp. 219–227, 2005.
- [9] E. Belen, E. Ozal, A. Bayyigit, S. Gunaydin, and A. Helvaci, "Interarm systolic blood pressure difference is associated with myocardial injury after noncardiac surgery," *Kardiologia Polska*, vol. 74, no. 7, pp. 674–680, 2016.
- [10] C. E. Clark, R. S. Taylor, A. C. Shore, and J. L. Campbell, "The difference in blood pressure readings between arms and survival: Primary care cohort study," *BMJ*, vol. 344, no. 7851, 2012.
- [11] R. Agarwal, Z. Bunaye, and D. M. Bekele, "Prognostic significance of between-arm blood pressure differences," *Hypertension*, vol. 51, no. 3, pp. 657–662, 2008.
- [12] W. J. Verberk, A. G. H. Kessels, and T. Thien, "Blood pressure measurement method and inter-arm differences: A metaanalysis," *American Journal of Hypertension*, vol. 24, no. 11, pp. 1201–1208, 2011.

- [13] T. Shoji, M. Emoto, K. Shinohara et al., "Diabetes mellitus, aortic stiffness, and cardiovascular mortality in end-stage renal disease," *The Journal of the American Society of Nephrology*, vol. 12, pp. 2117–2124, 2001.
- [14] K. Cruickshank, L. Riste, S. G. Anderson, J. S. Wright, G. Dunn, and R. G. Gosling, "Aortic pulse-wave velocity and its relationship to mortality in diabetes and glucose intolerance: an integrated index of vascular function?" *Circulation*, vol. 106, no. 16, pp. 2085–2090, 2002.
- [15] S. Hatsuda, T. Shoji, K. Shinohara et al., "Regional arterial stiffness associated with ischemic heart disease in type 2 diabetes mellitus," *Journal of Atherosclerosis and Thrombosis*, vol. 13, no. 2, pp. 114–121, 2006.
- [16] A. E. Bortnick, K. C. Epps, F. Selzer et al., "Five-year followup of patients treated for coronary artery disease in the face of an increasing burden of co-morbidity and disease complexity (from the nhlbi dynamic registry)," *American Journal of Cardiology*, vol. 113, no. 4, pp. 573–579, 2014.
- [17] K. A. Burek, K. Sutton-Tyrrell, M. M. Brooks et al., "Prognostic importance of lower extremity arterial disease in patients undergoing coronary revascularization in the Bypass Angioplasty Revascularization Investigation (BARI)," *Journal of the American College of Cardiology*, vol. 34, no. 3, pp. 716–721, 1999.
- [18] A. Kimura, J. Hashimoto, D. Watabe et al., "Patient characteristics and factors associated with inter-arm difference of blood pressure measurements in a general population in Ohasama, Japan," *Journal of Hypertension*, vol. 22, no. 12, pp. 2277–2283, 2004.