

## Correlation between Proximal Abdominal Aortic Stiffness Measured by Ultrasound and Brachial-Ankle Pulse Wave Velocity

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**Background and Objectives:** The proximal portion of the abdominal aorta (AA) is characterized by minimal arteriosclerosis compared with other aortic segments. To assess the clinical usefulness of this characteristic, the correlation between ultrasonographically measured proximal AA stiffness and brachial-ankle pulse wave velocity (baPWV) was examined.

**Subjects and Methods:** 285 subjects were analyzed, half with hypertension and half with normal blood pressure. Proximal AA was examined using ultrasonography; strain, distensibility, elastic modulus, and the stiffness index were determined. After adjustment for age, gender, body mass index (BMI), systolic blood pressure (SBP), and heart rate, the relationships between baPWV and all these parameters were tested.

**Results:** The mean age of the study subjects was 58.1±12.8 years and the mean BMI was 24.3±3.8 kg/cm<sup>2</sup>. 58.9% of the subjects were female. 42.8% were hypertensive subjects. Among the hypertensive subjects, 56.0% were taking antihypertensive medication. Adjusted partial correlation coefficients for the relationship between baPWV with strain, distensibility, elastic modulus, and the stiffness index of the proximal AA were -0.203 (p=0.01), -0.121 (p=0.129), 0.304 (p=0.0001), and 0.299 (p=0.0001), respectively, in normotensive subjects. In the multivariate analyses, such correlations were observed mainly in the normotensive group, whereas there was no association among hypertensive subjects regardless of antihypertensive medication status.

**Conclusion:** baPWV is moderately correlated with the stiffness parameters for the proximal AA, mainly in normotensive subjects. (Korean Circ J 2013;43:391-399)

**KEY WORDS:** Abdominal aorta; Arteriosclerosis; Blood pressure; Elastic modulus; Pulse wave velocity.

### Introduction

Aortic stiffness is known to be associated with cardiovascular outcomes. Pulse wave velocity (PWV) is one of the best ways to mea-

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sure aortic stiffness. Aortic stiffness measured by PWV is regarded as one of the prognostic markers of cardiovascular diseases.<sup>1)</sup> In addition, aortic stiffness can also be assessed by analyzing strain in various parts of the aorta.<sup>2-5)</sup> This can be measured directly using ultrasound or magnetic resonance imaging,<sup>2)</sup> which are familiar tools in clinical practice. In general, the abdominal aorta (AA) is more susceptible to increased stiffness due to degenerative changes or aging than the more proximal or ascending parts, because the elastic fiber composition and the susceptibility to ischemic damage from atherosclerosis differ.<sup>6,7)</sup>

In a recent necropsy study of Korean subjects, the proximal portion of the AA was found to display minimal atherosclerosis compared to other segments such as the ascending or proximal parts or the distal part of the AA.<sup>8)</sup> This observation suggests that an examination of the proximal part of the AA may be more useful for the measurement of aortic stiffness, because any structural changes, such as

severe atherosclerosis or aneurysmal changes, can make it difficult to measure the strain.

Carotid-femoral PWV (cfPWV) is the standard measure of aortic stiffness. However, from a practical point of view, cfPWV has some limitations, such as requiring a long time for measurement, examiner variability, and inconvenience of exposing the femoral site.<sup>9)</sup> Practically, brachial-ankle PWV (baPWV), on the other hand, does not require exposure of the femoral site and is a more feasible measure of aortic stiffness than cfPWV. baPWV is a measure of the pulse wave propagation velocity from the point of the aorta where the pulse wave arrives during the time period of the same pulse wave travel to the brachial artery to the ankle. Although baPWV is also affected by the characteristics of the peripheral arteries and the inferior measure of the aortic stiffness compared to cfPWV, baPWV has been validated mainly by comparison with cfPWV, with which it shows a moderate correlation.<sup>10)</sup> Hence, at least in part, baPWV can reflect the characteristics of the aorta. Moreover, the validity of baPWV as a marker for cardiovascular diseases has been reported in population studies.<sup>11)</sup> To investigate the usefulness of measurement of proximal AA distensibility, we examined the relationship between the stiffness parameters for the proximal AA and baPWV as the partial measure of aortic stiffness.

## Subjects and Methods

Data were prospectively collected from 290 consecutive patients (selected from an initial cohort of 396) who underwent echocardiography after fasting, in hypertension clinics at the Hanyang University Medical Center, Seoul, Korea from February 1st to May 31st, 2009. During the study period, 114 cases were referred from a health promotion center and local clinic for an evaluation of atypical chest symptoms, palpitation, electrocardiographic changes without symptoms, and abnormal blood pressure or laboratory findings.

Clinical exclusion criteria were the refusal to provide informed consent, incomplete clinical data, patients with exposure to nicotine smoking or drinking coffee one hour before the study, and patients with significant skin diseases, joint deformity, renal insufficiency (estimated glomerular filtration rate <60 mL/min), atrial fibrillation, valvular heart disease, and unstable coronary artery disease.

Echocardiographic exclusion criteria were moderate or severe valvular dysfunction, systolic dysfunction characterized by an ejection fraction below 55% as indicated by M-mode measurement, regional wall motion abnormality, moderate or large pericardial effusion, cardiomyopathy, aortic aneurysm, right side heart failure, or pulmonary hypertension. Patients with an ankle-brachial index less than 0.95 were also excluded from the study.

Clinical information was collected via interviews with the partici-

pants and from medical records. Body mass index (BMI) was calculated as weight in kg/(height in meters)<sup>2</sup> and blood pressure was measured according to current guidelines. Blood chemistry data on fasting blood glucose, lipid profiles, creatinine, and hemoglobin were collected from patient medical records. Only current smokers were defined as smokers. Hypertensive patients were defined as those with systolic blood pressure (SBP) ≥140 mm Hg, diastolic blood pressure (DBP) ≥90 mm Hg, or those taking antihypertensive medications. Patients with diabetes mellitus were defined as those with fasting blood glucose >126 mg/dL, HbA1C >6.5%, or those taking anti-diabetic medications or insulin.

Dyslipidemia was defined as total cholesterol ≥240 mg/dL, low density lipoprotein-cholesterol (LDL-C) ≥160 mg/dL, high density lipoprotein-cholesterol <40 mg/dL in males and <50 mg/dL in females, or triglyceride ≥150 mg/dL. A history of ischemic heart disease and/or cerebrovascular accident was determined through interviews.

## Measurement of abdominal aortic stiffness

The proximal AA was examined at the epigastrium for changes in diameter during the cardiac cycle using an echocardiography machine (iE33, Philips®). Unlike the ascending aorta, the posterolateral wall motion of the proximal AA is limited by the spine, so that most of the wall excursion during a cardiac cycle is observed in the anterior wall. By targeting the spine and the diaphragm and tilting the projection slightly to the right side, proximal AA wall excursions were recorded just below the diaphragm, in the suprarenal portion of the AA (Fig. 1).

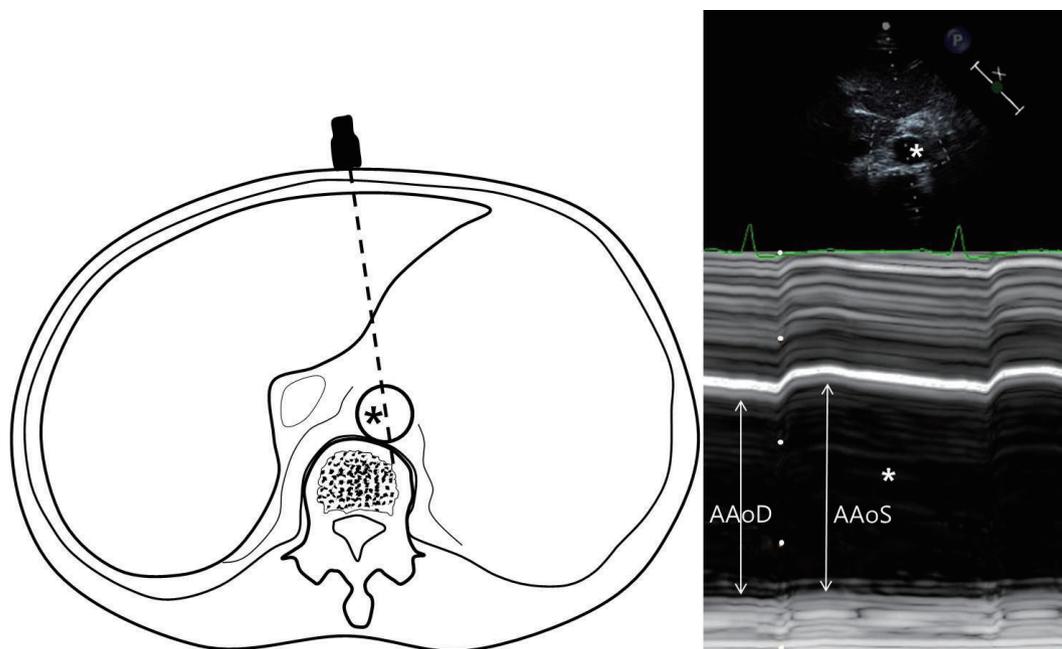
All participants were examined by a single examiner, and aortic dimensions were measured by a single individual in an off-line procedure using Infinity® DICOM Software (Infinity Inc., Korea). The minimal diastolic proximal AA diameter (AAoD) was measured at the time of QRS onset, and the maximal systolic proximal AA diameter (AAoS) was measured at the time of peak T wave in the electrocardiography recording from the trailing edge of the anterior wall to the leading edge of the posterior wall, over three cycles. Intra-observer Pearson correlation coefficients for repeated measurements of aortic dimensions in 30 subjects were 0.89 and 0.87, respectively, for diastolic and systolic diameters.

A preliminary study to calculate the Pearson correlation coefficient between proximal AA strain and aortic strain measured on the ascending aorta at a level 3 cm distal to the aortic annulus revealed a coefficient of 0.322 (p=0.018, n=50).

The stiffness parameters for the proximal AA were calculated as follows:

$$1) \text{ AA strain (\%)} = (\text{AAoS} - \text{AAoD}) / \text{AAoD} \times 100^{12)}$$

$$2) \text{ AA distensibility (mm Hg}^{-1} \times 10^{-3}) = \Delta \text{aortic luminal area} / \text{pulse pressure}$$



**Fig. 1.** Measurement of abdominal aortic strain. A sector probe was placed at the epigastrium targeting the spine, which typically shows a prominent anterior wall excursion and a relatively fixed posterior wall. \*Abdominal aorta abutting spine posteriorly. AAoD: minimal diastolic abdominal aortic dimension at the onset of the QRS wave, AAoS: maximal systolic abdominal aortic dimension.

$$= \pi \times [(AAoS/2)^2 - (AAoD/2)^2] / [(\pi \times AAoD/2)^2 \times \text{pulse pressure}]^{13)}$$

3) Elastic modulus (also known as Peterson's modulus) =  $(SBP - DBP) / [(AAoS - AAoD) / AAoD] \times 100^{14)}$

4) Aortic stiffness index =  $\ln(SBP/DBP) / [(AAoS - AAoD) / AAoD]^{15-18)}$

### Measurement of brachial-ankle pulse wave velocity

Brachial-ankle pulse wave velocity was measured upon the completion of echocardiography after five minutes of rest in the supine position using a VP-2000™ (Colin Co. Ltd., Komaki, Japan), as described in a previous study.<sup>19)</sup> The mean values of the right and left baPWV were used for statistical analysis. All measurements were performed by a single examiner. For 17 subjects in the repeated measurement, Cronbach alpha was 0.978 and the repeatability coefficient was 186.4 cm/sec.

Heart-carotid PWV (hcPWV), heart-femoral PWV (hfPWV), and cfPWV were also measured to allow for an assessment of the correlation between baPWV and those particular parameters. Measurements were performed on the left side using an applanation tonometry probe.<sup>20,21)</sup> The Pearson correlation coefficients for the relationship of baPWV with hcPWV, hfPWV, and cfPWV were 0.580 ( $p < 0.001$ ), 0.423 ( $p < 0.05$ ), and 0.577 ( $p < 0.001$ ), respectively.

The study protocol was reviewed and approved by the Institutional Review Board of the Hanyang University Medical Center in Seoul, and each participant provided informed consent.

### Statistical analysis

The sample size was calculated from the expected Pearson cor-

relation coefficients for the relationship of age with proximal AA strain and baPWV of 0.5 and 0.3. A two-tailed difference between the two correlation coefficients was expected to be detected at the level of  $\alpha = 0.05$ , with a value for  $1 - \beta$  of 0.9.<sup>22)</sup> The sample size was 279 when the correlation coefficient between proximal AA strain and baPWV was 0.3.

For the analysis of the correlation between baPWV and the stiffness parameters for the proximal AA partial correlation coefficients, adjusted for age, gender, and SBP, were calculated for all subjects, normotensive/hypertensive patients, and treated/untreated patients (with antihypertensive medication). Partial correlation coefficients according to gender were generated by adjusting for age and SBP.

Multiple regression analyses were performed to separately examine the factors influencing each proximal AA stiffness parameter. To test the main hypothesis, the parameters for the stiffness parameters for the proximal AA were set as the dependent variables in each multiple linear regression model. baPWV was set as an independent variable along with other independent variables, such as age, gender, BMI, SBP, heart rate, total cholesterol, and fasting blood glucose. baPWV values adjusted for age, gender, BMI, SBP, and heart rate were compared according to the quartiles of each proximal AA stiffness parameter using analysis of variance and general linear models. A  $p < 0.05$  was regarded as statistically significant.

All statistical analysis was performed using Statistical Package for the Social Sciences (SPSS) software package 15th edition (SPSS Inc., Chicago, IL, USA), and data are expressed as mean  $\pm$  standard deviation.

## Results

### General characteristics of the study subjects

The mean age of the study subjects was  $58.1 \pm 12.8$  years and the mean BMI was  $24.3 \pm 3.8$  kg/cm<sup>2</sup>. 58.9% of the subjects were female. Hypertension was noted in 122 subjects (42.8%). The proportion of current smokers was 13.6% (32 males and 7 females). Frequencies of essential hypertension, dyslipidemia, and diabetes were 49.5%, 58.9%, and 14.7%, respectively. Histories of coronary artery disease and stroke were observed in 10.5% and 2.4% of the subjects, respectively.

Table 1 shows the general characteristics of the subjects and compares these characteristics between males and females. As can be seen in the table, on average, females had higher pulse pressure, total cholesterol level, and LDL level than the male subjects.

Table 2 shows the comparison between hypertensive and normotensive subjects and between hypertensive subjects according to antihypertensive medication status. SBP in hypertensive subjects was the highest in untreated hypertensive at  $137.7 \pm 18.8$ , whereas the control rate of hypertension was 61/132 (46.2%). Among the hypertensive patients, 56.0% (74/122) were taking antihypertensive

medication. baPWV was also higher in hypertensive subjects. As shown in Table 2, the proportion of ischemic heart disease in treated hypertension subjects was higher than among normotensive or untreated subjects.

### Stiffness parameters for the proximal AA

As shown in Table 1, the parameters related to pulse pressure, i.e. distensibility, elastic modulus, and stiffness index, were significantly more affected among females than among males, even though baPWV was not significantly different between genders. Parameters for proximal AA stiffness were different among the normotensive, untreated hypertensive, and treated hypertensive groups. They were worst among untreated hypertensive subjects (Table 2).

### Correlation between brachial-ankle pulse wave velocity and stiffness parameters for the proximal AA

In a simple correlation analyses, the Pearson correlation coefficients for the relationship between baPWV and age, SBP, pulse pressure, heart rate, fasting blood glucose, total cholesterol, and creatinine were significantly positive (Table 3). The parameters of proximal AA stiffness were significantly correlated with baPWV in both nor-

**Table 1.** General characteristics of the study subjects and comparison between genders

	Overall (n=285)	Male (n=117)	Female (n=168)	p
Age (years)	58.1±12.8	56.6±13.9	59.1±12.0	0.102
Height (cm)	159.7±9.5	168.2±6.6	153.9±6.3	<0.0001
Body weight (kg)	62.2±11.2	68.8±10.2	57.5±9.3	<0.0001
Body mass index (kg/m <sup>2</sup> )	24.3±3.8	24.3±3.6	24.3±3.9	0.952
Systolic blood pressure (mm Hg)	127.7±18.5	126.7±15.2	128.4±20.5	0.467
Diastolic blood pressure (mm Hg)	77.4±10.9	78.8±11.1	76.4±10.6	0.067
Pulse pressure (mm Hg)	50.2±12.7	47.8±10.1	51.9±14.0	0.005
Heart rate (beats per minute)	67.2±11.6	65.8±11.0	68.1±12.0	0.111
Total cholesterol (mg/dL)	198.0±39.4	187.1±34.6	205.3±40.8	0.0003
Fasting blood glucose (mg/dL)	104.6±29.3	105.4±27.5	104.1±30.6	0.734
Creatinine (mg/dL)	0.92±0.22	1.07±0.22	0.81±0.13	<0.0001
Triglyceride (mg/dL)	154.1±94.0	159.7±103.1	149.7±86.3	0.434
HDL-C (mg/dL)	46.6±11.5	43.8±11.3	48.8±11.2	0.0013
LDL-C (mg/dL)	112.3±35.3	103.5±31.7	119.1±36.6	0.0012
Mean baPWV (cm/s)	1556.8±321.9	1526.1±300.4	1578.2±335.3	0.179
Aortic dimension (cm)	3.06±0.36	3.22±0.35	2.94±0.33	<0.0001
Systolic AA dimension (cm)	2.20±0.32	2.29±0.32	2.14±0.31	0.0002
Diastolic AA dimension (cm)	2.02±0.32	2.09±0.33	1.97±0.31	0.004
AA strain (%)	9.13±4.08	9.82±4.17	8.65±3.96	0.0182
AA distensibility (mm Hg <sup>-1</sup> ×10 <sup>-3</sup> )	4.15±2.29	4.59±2.37	3.84±2.1	0.0066
AA elastic modulus	7.40±6.33	6.48±6.16	8.03±6.39	0.0428
AA stiffness index	7.29±6.05	6.40±5.91	7.91±6.09	0.0395

p from Student t-tests to compare genders. HDL-C: high density lipoprotein-cholesterol, LDL-C: low density lipoprotein-cholesterol, AA: abdominal aortic, baPWV: brachial-ankle pulse wave velocity

**Table 2.** Comparison of the general characteristics of normotensive and hypertensive subjects

	Normotensive (n=163)	Untreated hypertensive (n=48)	Treated hypertensive (n=74)	p
Age (years)	56.4±13.6	57.2±13.4	62.4±9.4	0.003
Female (%)	58.2	66.6	55.4	0.45
Height (cm)	160.3±8.8	159.2±10.5	158.8±10.4	0.51
Body weight (kg)	60.8±10.7	64.0±12.5	63.9±11.1	0.06
BMI (kg/m <sup>2</sup> )	23.6±3.7	25.1±4.1	25.3±3.3	0.002
Systolic BP (mm Hg)	118.2±10.9	151.5±12.5	133.3±19.0	<0.001
Diastolic BP (mm Hg)	72.7±8.0	89.1±9.5	80.2±10.6	<0.001
Pulse pressure (mm Hg)	45.4±9.3	62.3±11.0	53.1±14.3	<0.001
Pulse rate (bpm)	66.3±11.4	68.6±9.0	68.1±13.4	0.347
Total cholesterol (mg/dL)	195.1±37.8	206.2±38.0	199.4±43.2	0.278
FBG (mg/dL)	105.2±32.1	101.7±20.4	105.1±28.0	0.814
Creatinine (mg/dL)	0.92±0.19	0.92±0.27	0.91±0.24	0.976
Triglyceride (mg/dL)	140.1±90.9	153.3±77.9	181.5±101.7	0.015
HDL-C (mg/dL)	47.9±11.5	49.9±13.2	42.6±9.5	0.0025
Mean baPWV (cm/s)	1445.8±280.7	1764.6±322.6	1666.4±305.3	<0.001
Aortic diameter (cm)	3.00±0.38	3.04±0.33	3.18±0.31	0.0023
Systolic AA (cm)	2.15±0.30	2.22±0.35	2.29±0.35	0.0068
Diastolic AA (cm)	1.96±0.30	2.06±0.33	2.12±0.34	0.0016
AA strain (%)	9.82±4.23	8.07±3.70	8.28±3.72	0.004
AA distensibility (mm Hg <sup>-1</sup> ×10 <sup>-3</sup> )	6.04±4.55	2.79±1.35	3.51±1.80	<0.001
AA elastic modulus	6.04±4.55	10.04±8.41	8.73±7.42	<0.001
AA stiffness index	6.44±4.74	8.68±7.92	8.30±7.01	0.0211
Ischemic heart disease (%)	5.5	8.3	25.6	0.006
History of stroke (%)	1.2	2.0	5.4	0.719
Diabetes mellitus (%)	12.7	10.4	21.6	0.554

p from Student t-tests or chi-square tests to compare groups. HDL-C: high density lipoprotein-cholesterol, AA: abdominal aortic, baPWV: brachial-ankle pulse wave velocity

normotensive and hypertensive subjects (Fig. 2).

In partial correlation analyses, after adjustments for age, gender, SBP, BMI, and heart rate, baPWV was consistently significantly correlated with strain, elastic modulus, and stiffness index in all subjects and in normotensive subjects (Table 4). When additionally adjusted for antihypertensive medication status, there was no significant correlation between baPWV and the stiffness parameters for the proximal AA.

### Multiple regression analyses for the stiffness parameters of the proximal AA

In the multiple regression analyses, baPWV was independently associated with the strain of the proximal AA or strain only among normotensive subjects, as shown in Table 5. Not antihypertensive medication status itself, but the level of blood pressure and heart rate, was associated with baPWV in the multiple linear regression model, including abdominal aortic strain.

### Comparison of brachial-ankle pulse wave velocity between the quartiles of the stiffness parameters of proximal AA

After adjustments for age, gender, BMI, SBP, and heart rate, baPWV was significantly higher among the lower quartiles of distensibility in the normotensive group (Q1, 1539±36 cm/sec; Q2, 1426±32 cm/sec; Q3, 1406±32 cm/sec; Q4, 1425±35 cm/sec, p for linear trend=0.0419). The baPWV among normotensive subjects was significantly higher with increasing quartiles of elastic modulus (Q1, 1428±35 cm/sec; Q2, 1404±31 cm/sec; Q3, 1426±32 cm/sec; Q4, 1539±36 cm/sec, p for linear trend=0.0489) (Fig. 3A). For the hypertensive group, with additional adjustments for antihypertensive medication status, baPWV was not significantly different among the quartile groups (Fig. 3B).

### Discussion

The main finding of this study is that baPWV has a moderate

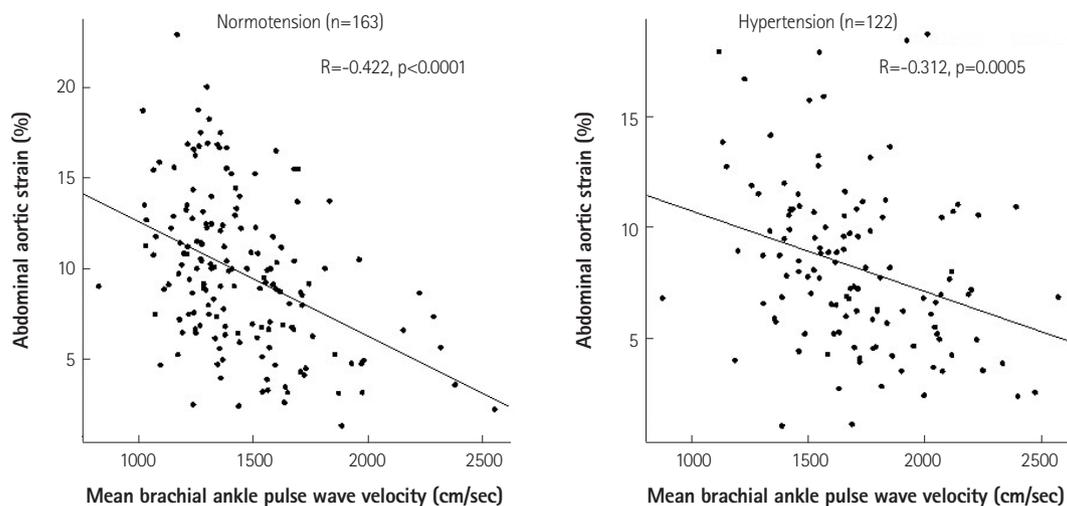
crude correlation with the stiffness parameters for proximal AA. However, when adjusted for age, sex, SBP, heart rate, and BMI, such correlations were observed only among normotensive subjects. The

antihypertensive medication status does not appear to be associated with baPWV in the multiple linear regression model, including abdominal aortic strain. Such a finding suggests that, in hypertensive

**Table 3.** Pearson correlation coefficients for the relationships between mean baPWV and proximal AA stiffness parameter

	All subjects (n=285)		Normotensive (n=163)		Hypertensive (n=122)	
	R	p	R	p	R	p
Age	0.573	<0.001	0.594	<0.001	0.537	<0.0001
BMI	0.119	0.043	0.132	0.092	-0.069	0.449
Systolic blood pressure	0.548	<0.001	0.457	<0.001	0.413	<0.0001
Pulse pressure	0.459	<0.001	0.307	<0.001	0.381	<0.0001
Heart rate	0.228	<0.001	0.126	0.108	0.311	0.0005
Fasting blood glucose	0.212	0.001	0.311	0.0003	0.141	0.154
Total cholesterol	0.213	0.0006	0.086	0.30	0.317	0.0007
Creatinine	0.130	0.051	0.287	0.0001	0.026	0.795
Strain	-0.413	<0.001	-0.422	<0.001	-0.311	0.0005
Distensibility	-0.5509	<0.001	-0.454	<0.001	-0.434	<0.0001
Elastic modulus	0.416	<0.001	0.487	<0.001	0.279	0.002
Stiffness index	0.351	<0.001	0.453	<0.001	0.217	0.016

baPWV: brachial-ankle pulse wave velocity, AA: abdominal aortic, BMI: body mass index



**Fig. 2.** Scatter plots for the correlation between proximal abdominal aortic strain and mean brachial-ankle pulse wave velocity according to hypertension status. The correlation was higher in normotensive subjects than in hypertensive subjects.

**Table 4.** Partial correlation coefficients for the relationships between mean baPWV and proximal AA stiffness parameters

	All subjects (n=285)		Normotensive (n=163)		Hypertensive (n=122)	
	R	p*	R	p*	R	p <sup>†</sup>
Pulse pressure	-0.107	0.072	-0.080	0.315	-0.180	0.056
Strain	-0.110	0.062	-0.203	0.010	0.019	0.837
Distensibility	-0.059	0.320	-0.121	0.129	0.053	0.576
Elastic modulus	0.097	0.109	0.304	0.0001	-0.079	0.405
Stiffness index	0.096	0.114	0.299	0.0001	-0.085	0.576

\*p adjusted for age, gender, systolic blood pressure, body mass index, and heart rate, <sup>†</sup>p adjusted additionally for antihypertensive medication status. baPWV: brachial-ankle pulse wave velocity, AA: abdominal aortic

subjects, not antihypertensive medication itself, but other factors, (such as age, SBP, and heart rate), may be more involved in the relationship between baPWV and proximal AA strain than for normotensive subjects. The relationship between baPWV and proximal AA stiffness parameters seemed to be linear only in the normotensive

group. The lack of any significant association between baPWV and proximal AA stiffness parameters in the hypertension group may be due to the effect of antihypertensive medication on blood pressure and heart rate. The role of the different classes of antihypertensive medication could not be demonstrated in the present study.

**Table 5.** Multiple linear regression analysis of factors associated with baPWV

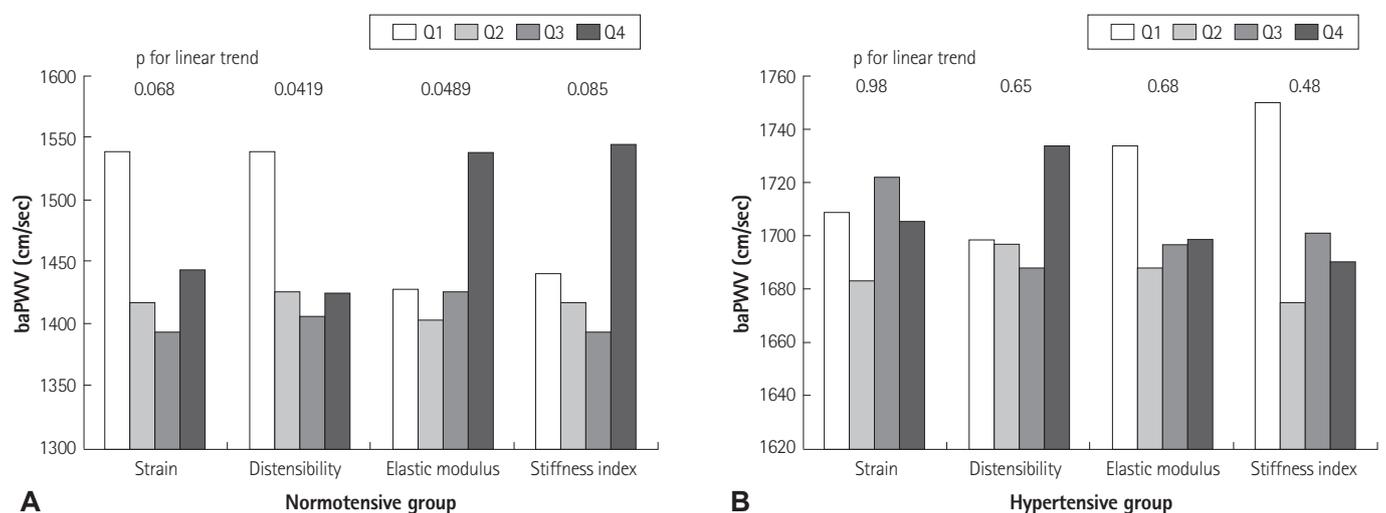
Group	Factor	baPWV	
		$\beta$	p
All subjects	Age	11.8	<0.0001
	SBP	7.5	<0.0001
	HR	4.4	0.0005
	FBS	0.9	0.0569
	CHOL	0.9	0.0177
	Proximal AA strain	-5.8	0.1610
Normotensive	Age	9.4	<0.0001
	Female	-63.8	0.1173
	SBP	7.7	0.0001
	HR	3.8	0.0237
	FBG	0.9	0.130
	Proximal AA strain	-11.0	0.048
Hypertensive	Age	15.1	<0.0001
	SBP	5.5	0.0002
	HR	5.9	0.0024
	CHOL	1.3	0.0265
	AHM	-76.2	0.168
	Proximal AA strain	-2.0	0.756-

baPWV: brachial-ankle pulse wave velocity, SBP: systolic blood pressure, HR: heart rate, FBS: fasting blood sugar, CHOL: total cholesterol, AA: abdominal aorta

Similar results were observed in a previous study, which showed that baPWV might have some limitations in predicting arterial stiffness in hypertensive patients with a history of cardiovascular disease.<sup>23)</sup> However, some authors suggest that baPWV can predict early arteriosclerotic changes.<sup>24)</sup> In addition, compared to the augmentation index, baPWV is reported to be useful among elderly patients, and some studies have shown that baPWV can predict aortic calcification.<sup>4)25)</sup>

The non-significant correlations between baPWV and proximal AA stiffness in the hypertensive group, regardless of antihypertensive medication status, can be explained by findings from previous studies. The stiffness parameters of the proximal AA do not have a linear relationship with age; a plateau can be observed in old age.<sup>26)27)</sup> It was reported that aortic distensibility and elastic modulus are more useful indicators in normal young subjects than among hypertensive and elderly subjects.<sup>28)</sup> baPWV was reported to underestimate aortic stiffness in hypertensive patients with coronary artery disease or target organ damage, as these subjects have lower femoral-ankle PWV.<sup>23)</sup>

In this study, we unexpectedly found that the stiffness parameters of the proximal AA were more affected in female subjects (Table 1 and 5). The higher pulse pressure in females is likely to be the reason for those differences. It is known that pulse pressure tends to be higher in females than males after the age of 50.<sup>29)</sup> In addition,



**Fig. 3.** baPWV according to the quartiles of the stiffness parameters of the proximal AA. A: least square means adjusted for age, gender, systolic blood pressure, heart rate, and body mass index according to the quartile groups of the stiffness parameters of the proximal AA in normotensive subjects. B: least square means additionally adjusted for antihypertensive medication status, according to the quartile groups of the stiffness parameters of the proximal AA in hypertensive subjects. There was no difference observed in baPWV between the quartiles of the stiffness parameters of the proximal AA in the hypertensive group. baPWV: brachial-ankle pulse wave velocity, AA: abdominal aorta.

we found that diastolic aortic dimensions were significantly lower among female subjects, which is consistent with a previous study suggesting that aortic arch diameter in the diastole was lower in females than in males ( $2.56 \pm 0.31$  mm vs.  $2.88 \pm 0.35$  mm).<sup>30)</sup> Although the smaller aortic size could not explain the pulse pressure difference completely, these differences between genders point to some gender-related differences in mechanical properties in response to aortic pressure changes, or a gender-related mismatch between ventricular function and aortic function.<sup>30)</sup> Furthermore, females had higher total cholesterol and LDL, consistent with a previous study performed in the general population and one of patients with familial hypercholesterolemia.<sup>16)31)</sup> Finally, there may be a difference in left ventricular function, which may affect proximal AA distensibility, as found in a previous study.<sup>32)</sup>

### Clinical Implications

Ultrasonography is a readily accessible clinical tool for examining proximal AA when combined with an examination of the heart or intra-abdominal organs. The worst quartiles of proximal AA stiffness in normotensive subjects had greatly increased baPWV values. The application of a conventional ultrasound examination of the proximal AA may be helpful in identifying normotensive subjects with arterial stiffness, but its potential application for hypertensive subjects is not supported by the findings of the present study. However, the proportion of subjects with ischemic heart disease history was about a quarter in the treated hypertension group, because, in our study, the subjects were recruited from a hypertension clinic affiliated with a cardiology division. Consequently, any generalized application of the negative findings among hypertensive subjects may be limited.

### Limitations of the study

There are some limitations of this study. Firstly, all subjects were examined after fasting. Thus, the usefulness of measuring proximal AA stiffness in fed subjects was not demonstrated. Additionally, cfPWV, which is more accurate than baPWV, was not used in this study because, during preliminary testing, baPWV was found to be superior in terms of both technical feasibility and general use in actual clinical practice. Using a VP-2000 device, hfPWV would be a far more accurate indicator of aortic stiffness than baPWV if a femoral artery probe can be applied. The correlation between PWV and abdominal aortic strain would have been improved had cfPWV been adopted. The result cannot be applied to cfPWV. The results of the study demonstrating the effects of antihypertensive medication on aortic strain in the multiple regression analysis should be interpreted with caution, because each of the classes of antihypertensive drugs were not considered. In the present study, the sample size was too

small to examine the effects of antihypertensive drug classes in the treated subjects. Further studies are warranted in this area of investigation.

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