# A DECREASE OF THE ANKLE-BRACHIAL PRESSURE INDEX (ABI) IS THE IMPORTANT RISK FACTORS FOR MORTALITY IN LONG-TERM DIALYSIS PATIENTS

Takayuki Abe1), Shigeru Otsubo2), 3), Naoki Kimata1), 4), Tomoki Okajima1), Yumi Otani1), Jun Murakami1), Iwakazu Kaneko1), Naoko Miwa4), Michio Mineshima1), Ken Tsuchiya3), 4), Takashi Akiba5) and Kosaku Nitta3)

- 1) Department of Clinical Engineering, Tokyo Women's Medical University, Tokyo, Japan
- 2) Department of Blood Purification, Sangenjaya Hospital, Tokyo, Japan
- 3) Department of Medicine, Kidney Center, Tokyo Women's Medical University, Tokyo, Japan
- 4) Department of Blood Purification, Kidney Center, Tokyo Women's Medical University, Tokyo, Japan
- 5) Sekikawa Hospital

# **OBJECTIVES**

Peripheral artery disease (PAD) is a regular complication of hemodialysis patients [1, 2] and is associated with poor outcomes[2-7]. Clinically, the ankle-brachial blood pressure index (ABI) is highly correlated with PAD of the lower extremities[8], and ABI is an independent predictor of all-cause mortality in hemodialysis patients [4-7].

A recent study also revealed that not only the value of the ABI itself, but also the rate of the reduction in the ABI was a risk factor for cardiovascular mortality[9].

In the present study, we investigated the effect of changes in the ABI on mortality and the factors that predict changes in the ABI in hemodialysis patients.

Cox proportional hazards analysis of the covariates for predicting mortality (univariate analysis)

Parameters (n=61)	Hazard Ratio (95% CI)	P value
ABI (per 0.1)*	0.43 (0.25 to 0.64)	<0.0001
ΔABI (per 0.1)*	0.62 (0.49 to 0.79)	< 0.001
Age (per 1 year)	1.06 (1.00 to 1.13)	0.048
Gender (M)	0.77 (0.34 to 1.64)	0.493
Duration of HD (per 1 year)	1.00 (0.91 to 1.09)	0.969
Albumin (per 1 g/dL)	0.12 (0.02 to 0.82)	0.032
Creatinine (per 1 mg/dL)	0.49 (0.32 to 0.71)	< 0.001
Uric acid (per 1 mg/dL)	0.42 (0.20 to 0.80)	0.008
Corrected Calcium (per 1 mg/dL)	1.14 (0.42 to 2.81)	0.789
Phosphate (per 1 mg/dL)	0.26 (0.09 to 0.65)	0.002
Ca x P (per 1 (mg/dL)2)	0.98 (0.92 to 1.03)	0.446
Total-Cholesterol (per 1 mg/dL)	1.00 (0.98 to 1.02)	0.919
Triglyceride (per 1 mg/dL)	1.00 (0.98 to 1.02)	0.916
c-reactive protein (per 1 mg/dL)	1.04 (0.29 to 1.75)	0.912
Hemoglobin (per 1 g/dL)	0.60 (0.25 to 1.44)	0.25
Hypertension (Y)	2.15 (0.89 to 9.36)	0.095
Diabetes mellitus(Y)	0.95 (0.22 to 2.29)	0.918

Cox proportional hazards analysis of the covariates for predicting composite endpoints including cardiovascular events and mortality (univariate analysis)

Parameters (n=61)	Hazard Ratio (95% CI)	P value
ABI (per 0.1)*	0.65 (0.50 to 0.86)	0.025
ΔABI (per 0.1)*	0.73 (0.61 to 0.90)	0.005
Age (per 1 year)	1.04 (1.00 to 1.09)	0.044
Gender (M)	1.25 (0.73 to 2.25)	0.419
Duration of HD (per 1 year)	1.03 (0.97 to 1.09)	0.357
Albumin (per 1 g/dL)	0.16 (0.05 to 0.64)	0.011
Creatinine (per 1 mg/dL)	0.67 (0.52 to 0.86)	0.001
Uric acid (per 1 mg/dL)	0.80 (0.51 to 1.21)	0.294
Corrected Calcium (per 1 mg/dL)	1.68 (0.74 to 3.64)	0.211
Phosphate (per 1 mg/dL)	0.60 (0.34 to 0.96)	0.033
Ca x P (per 1 (mg/dL)2)	0.96 (0.91 to 1.01)	0.084
Total-Cholesterol (per 1 mg/dL)	0.99 (0.97 to 1.00)	0.141
Triglyceride (per 1 mg/dL)	1.01 (1.00 to 1.02)	0.153
c-reactive protein (per 1 mg/dL)	1.44 (0.84 to 2.05)	0.154
Hemoglobin (per 1 g/dL)	0.52 (0.27 to 0.98)	0.042
Hypertension (Y)	1.40 (0.81 to 2.67)	0.239
Diabetes mellitus(Y)	0.62 ( 0.15 to 1.40)	0.299

Chronic Kidney Disease. Rehabilitation.

#### METHODS

A total of 61 consecutive patients receiving maintenance hemodialysis who successfully underwent ABI examinations in both 2005 and 2011 were retrospectively enrolled in this study. All patients were not treated by interventional or surgical repair for PAD during 2005 and 2011.

The blood pressure was measured after the patients had rested in a supine position for at least 5 min. Patients who failed to measure ABI of either side of legs were excluded from the study. The change in the ABI (2011 measurement versus 2005 measurement) was estimated. ⊿ABI was defined and estimated as the following formula.

#### **△** ABI=ABI in 2011 – ABI in 2005

We set the baseline at 2011 and investigated the patient outcomes. The clinical endpoints were defined as cardiovascular disease (CVD) events and death from any cause. The smaller values for either ABI or ⊿ABI among the two values of both legs were adopted and incorporated into the analysis. The data were expressed as the means  $\pm$  S.D or median (quartile) (interquartile range: IQR).

Cox proportional hazards model was used to examine predictors of overall outcomes. A simple regression analysis was used to examine the relationship between two continuous variables.

The relationship between the change of ABI (2005 to 2011) and other clinical parameters of 2005.

	r	P value
Age (year)	-0.1968	0.030
Duration of HD (year)	0.0762	0.404
Albumin (g/dL)	0.0490	0.592
Creatinine (mg/dL)	0.2079	0.022
Corrected Calcium (mg/dL)	0.0783	0.391
Phosphate (mg/dL)	-0.0371	0.685
Ca x P (mg/dL) <sup>2</sup>	0.0542	0.553
Total-Cholesterol (mg/dL)	-0.1385	0.128
Triglyceride (mg/dL)	-0.0002	0.998
c-reactive protein (mg/dL)	0.1233	0.176
Hemoglobin (g/dL)	-0.0134	0.884

HD: hemodialysis, Ca: calcium, P: phosphate, ABI: ankle-brachial blood pressure index, \*ABI and ⊿ABI: smaller values among both legs

△ABI=ABI in 2011 – ABI in 2005 ESKD: end-stage kidney disease Mean ± SD, Median (interquartile range: IQR)

Background characteristics of the study participants

	All patients (n= 61)
Age (year)	57.2 ± 13.2
Gender (M/F)	33 / 28
Duration of HD (year)	19.8 (11.1 – 28.1)
Albumin (g/dL)	$3.7 \pm 0.3$
Creatinine (mg/dL)	11.8 ± 2.3
Uric acid (mg/dL)	$7.7 \pm 1.4$
Corrected Calcium (mg/dL)	$9.4 \pm 0.8$
Phosphate (mg/dL)	$5.4 \pm 1.4$
Ca x P (mg/dL)2	50.6 ± 12.2
Total-Cholesterol (mg/dL)	159 ± 36
Triglyceride (mg/dL)	102 ± 41
c-reactive protein (mg/dL)	0.07 (0.05 - 0.16)
Hemoglobin (g/dL)	$10.7 \pm 0.8$
Primary Cause of ESKD, n (%)	
Chronic glomerulonephritis	43 (70.5)
Diabetic Nephropathy	5 (8.2)
Polycystic kidney disease	3 (4.9)
Chronic pyelonephritis	1 (1.7)
Post transplantation	8 (13.1)
Unknown and others	1 (1.7)
Complication, n (%)	
Hypertension	36 (59.0)
Diabetes mellitus	9 (14.8)

### RESULTS

The mean follow-up period was  $3.1 \pm 0.7$  years. During the follow-up period, 7 deaths were recorded. The causes of death were infection in 5 cases, cerebrovascular disease in one case, and unknown in one case. In the Cox proportional hazard analysis, predictive variables for mortality included ABI (0.43 [0.25–0.64]; [per 0.1 increase]), and the change of ABI (0.62 [0.49–0.79]; [per 0.1 increase]). Patient age was negatively correlated and the serum creatinine level was positively correlated with the change in the ABI (P = 0.030 and 0.022, respectively).

## CONCLUSIONS

We confirmed that not only the value of the ABI itself, but also the change in the ABI was a risk factor for mortality among hemodialysis patients. Careful attention should be given to changes in the ABI as well as the value of the ABI itself. The change in the ABI was negatively correlated with age and was positively correlated with the serum creatinine level. Careful observation of the ABI is needed for old patients and patients with a low serum creatinine level.

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