Correlation between Changes in Leg Blood Flow and Ankle-Brachial Pressure Index: A Study Using Laser Doppler Flowmeter —The 1st Report—

Kazuyoshi Suzuki, PT,1, 2 Miho Sekiguchi, MD, PhD, 2 Hirofumi Midorikawa, MD, PhD, 3 Koichi Sato, MD, PhD, 4 Kazuyoshi Akase,5 Renshi Sawada, D.Eng,5 and Shin-ichi Konno, MD, PhD2

Objective: The objective of this study was to use non-invasive laser Doppler flowmeter to measure changes in blood flow in peripheral vessels in the legs before and after stress induced by leg elevation stress test and investigate correlations with the ankle-brachial pressure index (ABI).

Methods: Subjects included 28 patients over 20 years of age (mean, 73 years) who reported chiefly of leg symptoms such as intermittent claudication, numbness, chills, or cramps had been examined at the study institution, and agreed to participate in the study. The ABI of both legs was measured, and patients were divided into two groups: low ABI (ABI ≤0.9) and normal ABI (ABI ≥0.9). Blood flow in the big toe was measured using a box-type laser Doppler flowmeter before, during, and after leg-elevation stress. Amplitude of the recorded waveform and changes in blood flow were compared.

Results: Average ABI was 1.09 ± 0.10 in the normal ABI group (33 legs) and 0.68 ± 0.17 in the low ABI group (21 legs). Amplitude before and during stress was significantly smaller in the low ABI group than in the normal ABI group (p <0.01), and there was a significant correlation with ABI before and during stresses (r = 0.4606, r = 0.5048, respectively; p <0.05). Change in blood flow during stress was significantly lower in the low ABI group than in the normal ABI group (p <0.05). There was a significant correlation between change in blood flow during stress and ABI in both groups (r = 0.5073; p <0.05). There was also a significant correlation between change in blood flow and change in amplitude in both groups (r = 0.5477; p <0.05).

Conclusion: Results of this study show, that comparing amplitude and change in blood flow before and after leg extension and elevation stress, there was a correlation between change in blood flow and amplitude, and ABI during stress. A box-type laser Doppler flowmeter may provide a means of screening for peripheral arterial disease.

Key words: peripheral arterial disease, ankle-brachial pressure index, laser Doppler flowmeter, intermittent claudication, lumbar spinal stenosis

1Division of Rehabilitation, Iwase General Hospital, Sukagawa, Fukushima, Japan
2Department of Orthopedic Surgery, Fukushima Medical University School of Medicine, Fukushima, Fukushima, Japan
3Division of Cardiac and Vascular Surgery, Southern Tohoku Research Institute for Neuroscience, Southern Tohoku General Hospital, Koriyama, Fukushima, Japan
4Department of Cardiovascular Surgery, Sukagawa Hospital, Sukagawa, Fukushima, Japan
5Department of Intelligent Machinery and Systems, Graduate School of Engineering, Graduate School of Systems Life Sciences, Microsystems and Bioengineering Lab, Kyushu University, Fukuoka, Fukuoka, Japan
INTRODUCTION

With the increasing aging of society, the incidence of arteriosclerosis-related disorders such as hypertension, and cardiovascular disease is rising. Peripheral arterial disease (PAD) and lumbar spinal stenosis (LSS) sometimes have similar symptoms and commonly occur around the same age; thus it is important to distinguish these underlying conditions.1, 2) The overall prevalence of PAD is 3%–10%, rising to 15%–20% over the age of 70 years.3–5) The presence of arterial lesions in the legs reflects arterial lesions throughout the body, and the survival rate is poor.6) It follows that PAD with intermittent vascular claudication requires early diagnosis and referral to a vascular specialist for early treatment. Measurement of the ankle-brachial pressure index (ABI) is an effective method of screening for PAD.7, 8) Pulse volume recording, toe brachial pressure index (TBI), Duplex ultrasonography, and magnetic resonance angiography are non-invasive methods for diagnosis of PAD.7) ABI does not show blood flow of the leg tissue directly. Instead, ABI is determined using laser Doppler flowmeter (LDF) along with blood pressure measurements.9, 10) However, conventional LDF can only be used at rest and with patients maintaining the same position. If a simple, non-invasive method to measure blood flow in the leg were available that allowed for continual leg movement, it would enable screening for vascular lesions. A micro wireless flowmeter integrated laser Doppler blood flowmeter (MILDF) has been developed.11–13) MILDF provides similar results to measurement of blood flow with conventional LDF. In addition, MILDF is wireless, portable, and allows monitoring of diurnal blood flow variation. The objective of this study was to use MILDF to measure changes in blood flow in the peripheral arteries of the leg before and after stress imposed by leg elevation stress test and to investigate their correlation with ABI.

SUBJECTS AND METHODS

Subjects were 28 patients over 20 years of age who had been examined at the study institution due to report chiefly related to leg symptoms such as intermittent claudication, numbness, chills, or cramps, and who provided informed consent to participate in the study. Subjects included 26 men and 2 women, with a mean age of 73 years (range, 57–91; most were in their 70s). This study was approved by the ethics committee of the facilities concerned.

ABI values were measured for both legs of the subjects. Subjects lay supine on a bed with blood pressure cuffs fitted to their upper arms and ankles on both sides, and their blood pressure was measured using a VaSera VS-1000 (Fukuda Denshi, Japan). Then, blood flow in both great toes was measured. The MILDF11–13) used in this study was a box type that measured 5.6 mm × 12.0 mm × 17.5 mm and weighed 3 g (Fig. 1). This is approximately 1/300 the size of conventional LDF. Blood flow signals detected by the sensor were transmitted to the main unit, where measurement values were displayed. The main unit was also a signal transmitter, capable of sending data directly to a computer by Bluetooth operating on an industrial, science, and medical (ISM) 2.4–2.485 GHz wireless band. The blood flow waveform was continuously displayed on the computer screen and recorded from the data sent to the computer. As conventional LDF uses optical fibers for the input-output of laser light, movements of the optical fiber can result in noise. In this study, the fact that MILDF was wireless meant noise could be minimized. In addition, the miniaturized main unit also has the advantages of portability as well as the capability for continuous measurement and measurement during movement. It provides similar results to those of conventional LDF11–13) Measurements were performed in the outcome patient room at room temperature of 18°C. Only the subject and two observers were in the room during measurements.

The blood flow sensor was directly attached to the skin on one side of the great toe by double-sided tape. Subjects lay in the supine position, and blood flow measurements were started after subjects had spent at least 5 min resting on the bed. The blood flow waveform was checked on the computer screen, and blood flow was continuously recorded before, during, and after stress, as follows. Blood flow was measured for 60 seconds (s) in a resting state (before stress). Next, leg extension and elevation stress was imposed for 60 s, and blood flow was measured while the stress was imposed (during stress). The leg extension and elevation stress comprised elevating the leg in an extended position at an angle of 60° from the bed.14) The leg was then returned to the bed, and blood flow was measured for 60 s after the stress had finished (after stress). Blood flow in the big toe of the opposite leg was also measured in the same way.

Blood flow waveform amplitude and change in blood flow were measured from the results obtained. Amplitude reflected each pulse beat. Amplitude was calculated from the waveform during each 60-s period before,
Correlation of ABI and Change in Blood Flow

Annals of Vascular Diseases Vol. 4, No. 2 (2011)

Results

Analysis was performed on 54 of 56 legs in which blood flow was measured, excluding 2 legs for which the recorded waveform could not be analyzed. The normal ABI group comprised 33 legs, with an average ABI of $1.09 \pm 0.10$. The low ABI group comprised 21 legs, with an average ABI of $0.68 \pm 0.17$.

Blood flow waveform characteristics and amplitude size

Blood flow waveforms were characterized by a large amplitude in the normal ABI group and by a small amplitude in the low ABI group (Fig. 2). Amplitude before stress was $5.7 \pm 2.8$ ml (min 100 g) in the normal ABI group and $3.6 \pm 2.5$ ml (min 100 g) in the low ABI group. Amplitude was significantly smaller in the low ABI group than the normal ABI group ($p<0.01$). There was a correlation between amplitude before stress and ABI ($r = 0.4606; p<0.05$) (Fig. 3a).

Change in amplitude during stress

The change in amplitude during stress was $73.9\% \pm 62.2\%$ in the low ABI group and $108.7\% \pm 42.6\%$ in the normal ABI group. The amplitude was significantly decreased in the low ABI group compared with the normal ABI group ($p<0.05$). The change in amplitude after stress was $100.9\% \pm 19.4\%$ in the low ABI group and $104.5\% \pm 23.8\%$ in the normal ABI group. There was no significant difference between the two groups. There was a correlation between change in amplitude during stress and ABI ($r = 0.5048; p<0.05$) (Fig. 3b).

Change in blood flow during stress

The change in blood flow during stress was $44.5\% \pm 23.2\%$ in the low ABI group and $66.0\% \pm 18.5\%$ in the normal ABI group. The blood flow was significantly decreased in the low ABI group compared with the normal ABI group ($p<0.05$). The change in blood flow after stress was $116.2\% \pm 30.2\%$ in the low ABI group and $105.3\% \pm 22.3\%$ in the normal ABI group. There was no significant difference in change in blood flow between the two groups. There was a correlation between change in blood flow during stress and ABI ($r = 0.5073; p<0.05$) (Fig. 4).

When the cut-off point of the blood flow change was defined as $60\%$, the flowmeter had a sensitivity of $71\%$ and specificity of $58\%$. When cut-off point was $65\%$, the flowmeter had a sensitivity of $81\%$ and a specificity of $55\%$.  

Fig. 1 Laser Doppler flowmeter.  
(a) Box-type blood flow sensor and (b) portable flowmeter main unit.
Fig. 2 Blood flow waveform characteristics.
The blood flow waveform in the normal ABI group was of large amplitude, whereas the amplitude was small in the low ABI group.

Change in amplitude and change in blood flow

There was a correlation between change in amplitude and change in blood flow ($r = 0.5477; p < 0.05$) (Fig. 5).

Discussion

Measurement of ABI is a useful testing method for PAD diagnosis. ABI values $\leq 0.9$ have 95% sensitivity for the arteriographic detection of PAD. Specificity for angiographically documented PAD is 90%. ABI measurement is an essential requirement for PAD screening. Although devices are available for the simple calculation of ABI, the laser Doppler flowmeter used in this study has the advantages of being non-invasive and enabling continuous blood flow measurement. In our results, we found differences between the normal ABI and low ABI groups in the blood flow waveforms recorded. The amplitude of the waveform was significantly smaller in the low ABI group compared with the normal ABI group. The change in amplitude as a result of the imposition of leg extension and elevation stress was also smaller in the low ABI group. In addition, there was a positive correlation between change in amplitude size before and during stress.

We found that the change in blood flow as a result of the imposition of leg extension and elevation stress was significantly lower in the low ABI group compared with the normal ABI group. This means that there is little variation in blood flow in patients with PAD despite the imposition of stress. When the cut-off point of blood flow change was 65%, MILDF had a sensitivity of 81% and specificity of 58% in this study. A decrease in blood flow of more than 65% during leg elevation stress compared with blood flow at the rest (before stress) might indicate PAD. MILDF might be useful in the screening of PAD; however, the results in the present study did not consider the severity of PAD. Furthermore, a study with a large sample size is needed to evaluate whether MILDF could show the severity of PAD. In addition, there was a correlation between amplitude and change in blood flow. These results suggest that blood vessels have weak reactivity in cases of suspected PAD. ABI reflects the stenosis or occlusion of large blood vessels in the legs, and there has been insufficient investigation to say whether or not it reflects blood flow in peripheral tissue. Nevertheless, the positive correlation between change in blood flow and ABI suggests the possibility that ABI can be predicted from change in blood flow before and after leg
Correlation of ABI and Change in Blood Flow

Fig. 3  Correlation between amplitude and ABI.

(a) Before stress
There was a correlation between amplitude and ABI before stress ($r = 0.4606; p < 0.05$).

(b) During stress
There was a correlation between change in amplitude (%) and ABI during stress ($r = 0.5048; p < 0.05$).

ABI, ankle-brachial pressure index
There was a correlation between change in amplitude and change in blood flow ($r = 0.5477; p < 0.05$).

**Fig. 4** Correlation between change in blood flow and ABI.

There was a correlation between change in blood flow and ABI ($r = 0.5073; p < 0.05$).

ABI, ankle-brachial pressure index

There was a correlation between change in amplitude and change in blood flow ($r = 0.5477; p < 0.05$).

**Fig. 5** Correlation between change in amplitude and change in blood flow.
extension and elevation stress test. This study included only a small number of patients, however, and was limited by the fact that it was not possible to calculate the odds ratio for amplitude and change in blood flow that would constitute screening for PAD. If a large-scale study were carried out, this would enable the prediction of ABI from amplitude and change in blood flow. There are cases, however, in which it is not possible to obtain an accurate value for ABI because of a rise in ankle blood pressure due to diabetes, kidney failure, and vascular calcification. Cases which show false-positive ABI results should be examined using the TBI, which results in fewer false-positive findings compared with ABI. If TBI obtained by measuring blood pressure at the great toe is less than 0.7, then PAD is suspected. In a comparison of TBI and ABI among patients who underwent examination in a specialist vascular outpatient clinic, both ABI and TBI were low in 34% of cases. In 17% of cases, ABI was normal, and only TBI was low. Because the MILDF used in this study measured peripheral blood flow in the great toe, it may also be applicable to screening for vascular lesions in patients with other vascular disorders causing peripheral stenotic lesions, diabetes, and kidney failure. The present study, however, did not investigate whether there was any correlation among TBI, amplitude, and change in blood flow. Further research is required to determine whether there is any correlation between TBI and values measured by LDF. Further research is also required to determine whether PAD screening is possible in patients with PAD and LSS, both of which have characteristic symptoms of intermittent claudication and similar ages of onset. LSS is complicated by PAD in 6.7% of cases, occurs mainly in elderly people, and has a high frequency of cardiovascular complications. Whether screening is possible in patients with LSS complicated by PAD requires further study.

The MILDF used in this study has also been reported as a useful testing device for evaluating therapeutic efficacy in skin disorders. It can also be anticipated to be of use as a tool for evaluating therapeutic efficacy in PAD patients. Further studies of blood flow waveform and changes in blood flow in PAD patients is required.

The MILDF used in this study can be used to screen for low ABI, by comparing amplitude and change in blood flow before and after leg extension and elevation stress. When the cut-off point of blood flow change using MILDF is 65%, sensitivity for PAD is 81%. Further study may enable LDF to be used as a simple PAD screening tool in outpatient clinics other than specialist vascular departments.

**References**

nese)