The Effects of Controlled Physical Training on Peripheral Circulation Following Intervenional Treatment of Coronary Artery Disease*

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Abstract

Background. Controlled physical training induces specific changes in the peripheral circulatory system and can lead to positive changes in the vascular perfusion of the lower extremities.

Objectives. The aim of the study was to evaluate changes in peripheral circulation in the calf in patients with acute coronary disease (ACD) undergoing controlled physical training. Impedance plethysmography was used to monitor peripheral circulation during the training.

Material and Methods. A total of 90 patients were divided into three study groups. Group 1 (n = 30) participated in a two-week cardiac rehabilitation program consisting of interval training on a cycle ergometer and exercise to improve the participants' general physical condition. Group 2 (n = 30) went through the same cardiac rehabilitation program for four weeks. The control group (n = 30) was assigned breathing exercises, active free exercises of the peripheral joints and different muscle groups, and relaxation exercises. All the patients underwent impedance plethysmography tests before and after the training sessions.

Results. In Group 1, the systolic slope (PSlope) increased by 2%, pulse wave amplitude (PAmpl) increased by 4.2%, crest time (CT) increased by 1.5% and propagation time (PT) decreased by 1.2% (p > 0.05). In Group 2, the PSlope and PAmpl increased by 19% and 17% respectively, while the CT and PT decreased by 8% and 6.5% respectively (p < 0.05). In the control group, only the CT decreased, by 5% (p < 0.05).


Key words: controlled physical training, peripheral circulation, impedance plethysmography.
system, which in turn have hemodynamic, anatomic and metabolic consequences in the central and peripheral circulation. This obliges the coronary rehabilitation staff to monitor and evaluate not only changes in the coronary arteries, but also in the peripheral circulation. This enables the rehabilitation program to be fully effective and to achieve the aims of decreasing rehospitalization, improving patients’ psychophysical condition and allowing patients more fully independent functioning [2].

Every physical workout is associated with changes in the cardiovascular system. Functional adaptive changes include decreases in the heart rate and increases in cardiac output, left ventricular stroke volume, the arteriovenous oxidation difference and arterial tension. This occurs simultaneously with pump muscle activation responsible for maintaining peripheral venous return and diastolic filling of the heart [3]. The central adaptive mechanisms are closely connected with the peripheral vascular and skeletal system. This cooperation is required to provide proper blood flow at rest as well as during exercise. The most important function of both the central and peripheral systems is to provide proper distribution of oxygenated blood to all body organs, especially to working muscles. The aim of this mechanism is to redistribute the blood flow through different vascular areas, especially to supply the right amount of oxygen to working muscles during exercise. The activation of the sympathetic nervous system leads to increases in the rate of metabolism and the local accumulation of metabolic products (lactate, adenosine, hydrogen ions, potassium ions, CO2), an increase in muscle temperature, hypoxia and osmotic pressure. During physical exercise coronary perfusion is almost five times faster. However, the most spectacular changes can be observed in skeletal muscles, which, during physical exercise, are provided with 80–85% of the left ventricular output [4]. In recent years, a number of hypotheses have arisen involving the pathogenesis of cardiovascular disease with peripheral pathomechanisms, such as neurohormonal activation, impaired anabolic-catabolic balance, impaired autonomic balance, muscle dysfunction or the activation of pro-inflammatory cytokines. These hypotheses are associated with patients’ reduced exercise tolerance and the progression of vascular disease [5]. Chronic functional disorders of the heart and its structural changes translate into a cardiovascular disorder adaptation in the form of lower peripheral blood redistribution. The blood flow through the skeletal muscles and the amount of oxygen provided both during exercise and at rest becomes limited. This is connected with a decrease in capillary density in the muscles, peripheral arterial vasoconstriction, increased activity ergoreceptors, impaired endothelial vasodilation function and reduced relaxation of peripheral vessels. An important role is played by changes in the quality, type and architecture of muscle fibers [6].

It is understood that regular exercise training can result in the development of collateral circulation, and post-exercise changes within it can be caused by the development of new collateral vessels or the expansion of existing ones. Improved endothelial vasodilator function is one of the reasons for the increase in blood flow. Some authors suggest that chronic increased shear forces can stimulate the release of growth factors such as vascular endothelial growth factor (VEGF) and basic fibroblast growth factor (bFGF). The result is stimulation of the vascular adaptation processes, and a resulting increase in collateral blood vessels [7]. As an effect of physical activity, which plays a role in flow-dependent vasodilatation, secretion of nitric oxide (NO) is observed to increase. This is due to increases in the activity and expression of endothelial NO synthase and the increased bioavailability of L-arginine; as well as to increases in the activity and expression of superoxide dismutase, catalase and peroxidase, and antioxidants (vitamins E and C), which inhibit the degradation of NO. As blood flow increases, shear force intensifies. Nitric oxide mediates the expansion of arteries, reducing shear force to its initial level; decreasing the amount of NO in the vessel walls leads to an increase in leukocyte adhesion and aggregation of platelets, initiating the process of atherosclerosis [8]. It is believed that long-term physical training has the effect of increasing the synthesis of NO and prostacyclin. Improvement of vascular endothelial vasodilator function, which has been observed in response to exercise training in patients with cardiovascular disease (CVD) and confirmed by experiments in animals, leads to increased blood flow as well as increased release of NO. This leads to a reduction in vascular resistance and improves patients’ peripheral blood flow [9, 10].

Monitoring cardiovascular activity is a necessary element of supervising the patient during cardiac rehabilitation. An important element of this supervision is to evaluate the efficiency of the peripheral circulation, especially in patients who have undergone coronary artery bypass grafting (CABG). Despite the availability of tests designed to evaluate vascular perfusion (Doppler ultrasound, thermodilution) their use is a significant nuisance for both the subject and the examiner, and the equipment and required training are quite costly.

The human body as a biological material is characterized by electrical resistance and electrical...
reactance. The result of these two is impedance. Each tissue is different in terms of impedance during the flow of alternating current, which differentiates its functional state [11]. Thanks to these relationships it is possible to use bioimpedance, or, to be more precise, a method of measuring the volume and pace of local blood flow, i.e. impedance plethysmography, in the diagnostics of the peripheral circulatory system. Changes in the volume and the pace of blood flow can be detected by measuring changes in the electrical resistance in particular body parts. The blood demonstrates better conduction properties than fixed body parts, which are mostly composed of membranous structures and therefore have insulating properties. The arterial pulse wave is very small but regular, so it is easily detected and measured [12]. Registration of the plethysmographic curve is a useful method for the patient. The authors therefore believe that it could be used to evaluate peripheral circulation in patients who are undergoing cardiac rehabilitation.

The aim of the study was to evaluate changes in peripheral circulation in the calf induced by controlled physical exercise performed by patients with acute coronary disease (ACD). Impedance plethysmography was used to monitor the peripheral circulation during different types of physical training.

Material and Methods

Ninety patients, aged 45–78 (mean: 61 ± 7.1) were included in the study. Among them were 67 men and 27 women who suffered from ACD. All the patients were referred for cardiac rehabilitation at the Clinic of Internal Diseases and Cardiac Rehabilitation of the Medical University of Lodz no more than one month after the occurrence of acute coronary disease.

The selection criteria included a diagnosis of ACD (myocardial infarction, coronary artery bypass grafting surgery, percutaneous intervention), a lack of absolute and relative contraindications to physical exercise, and a lack of comorbidities (symptomatic atherosclerosis of lower extremities, diagnosed advanced diabetes, neurological incidents). The patients were qualified for the training program on the basis of the results of an exercise test performed on a Cardiovit CS-200 treadmill (Schiller AG, Baar, Switzerland), following the Bruce protocol; during the test the patients’ blood pressure was monitored and they underwent electrocardiogram examination. Achieving an exercise workload of ≥ 5 MET was the criterion for inclusion in the study. This result qualified the patients for at least phase II cardiac rehabilitation according to the guidelines of the Polish Cardiac Society [14]. The occurrence of at least one of the following cardiac rehabilitation risk stratification factors was grounds for exclusion from the study groups: ejection fraction (EF) < 40%, horizontal depression of the ST segment on exertion, complex ventricular arrhythmia at rest and on exertion, physical efficiency < 5 MET, a pathological reaction to physical training in terms of systolic arterial pressure or heart rate. All the patients included in the study were undergoing pharmacological treatment in compliance with the guidelines of the Polish Cardiac Society, and had not changed their treatment in the previous three weeks.

Training Sessions

After the initial tests, the participants were randomly assigned to three groups of 30 patients. Group 1 consisted of 18 men and 12 women with an average age of 61.18 ± 8.53 years. Group 2 consisted of 22 men and 8 women; average age 59.69 ± 9.24 years. Group 3 (the control group) consisted of 20 men and 10 women with a mean age of 75.56 ± 7.50 years.

Group 1 participated in a two-week cardiac rehabilitation program, and Group 2 participated in a four-week program. Both rehabilitation programs consisted of sessions of controlled interval training on Ergoline cycle ergometers (Ergoline GmbH, Bitz, Germany). The training sessions were held five times a week before noon. Each interval training session lasted 32 min and was conducted according to a pre-established schedule. It started with a warm-up with about a 10-Watt load, followed by five 4-min intervals during which the load was gradually increased. The load in each interval increased up to the middle of the session; next, it decreased to the initial level; and there were five 2-min periods of rest with a 10-Watt load. The peak load (and maximum heart rate) during the initial training sessions was equal to 50% of the load (and heart rate) during the electrographic exercise test. In addition to interval exercises, breathing exercises, resistance training exercises for the small muscle groups of the upper and lower limbs and relaxation exercises were also introduced. These exercises were initiated after the interval training and lasted 20 min. During the entire training session the exertion did not exceed category 11–13 on the 6–20 Borg scale.

Group 3 (the control group) consisted of patients who were not admitted to the program of interval training because of the presence of one of the cardiac rehabilitation risk stratification character-
istics. The control group performed only breathing exercises, general fitness exercises and relaxation exercises. Each exercise session lasted 30 min and did not exceed 8–9 points on the Borg scale. The average stay in the group was 18 ± 32 days.

Before and after each exercise session, the heart rate and blood pressure of all the patients in all three groups were checked.

**Evaluation of Blood Flow in the Lower Extremities**

Before and after the two- and four-week training programs, all the patients underwent impedance plethysmography tests to evaluate peripheral circulation in the lower extremities.

Six months after the completion of the training programs the impedance plethysmography tests were repeated to evaluate the long-term effects.

Measurements of the blood flow were made in accordance with the instruction manual of the Niccomo cardiac output monitor (Medis Medizinische Messtechnik GmbH, Ilmenau, Germany). During the test the authors evaluated the following plethysmographic parameters (Fig. 1):

![Diagram of a plethysmographic curve](image1)

**Fig. 1.** Diagram of a plethysmographic curve [15]

- **PT [ms]** – propagation time;
- **PAmpl [‰]** – pulse wave amplitude;
- **CT [ms]** – crest time;
- **PSlope [‰/s]** – systolic slope.

Four electrodes were placed on the patient’s lower limb. The yellow electrodes in the calf area are the so-called measuring electrodes; the black electrodes on the dorsum of the foot and the distal anterior area of the thigh are the so-called voltage electrodes. The measurements were taken with the patient at rest in a lying position, after a 10-min rest. During the examination the patient was lying flat while electrodes placed on his lower limb measured the blood flow parameters within the examined area (Fig. 2). The final values were gathered after 8 min.

The data analysis was performed using PQStat software (v. 1.4.2.324, PQStat Software, Poznań, Poland). In the first stage of the analysis descriptive statistics characterizing the study group were generated. For the interval variables, average values with 25–75% confidence intervals were calculated, as well as the standard deviation, median and upper and lower limits of quartiles. The results of the analyzed parameters were compared using the Kruskal-Wallis test, a post-hoc Dunn test and Wilcoxon’s test. The Friedman test for dependent variables was used to analyze the data obtained at the three measurements (before, immediately after and six months after the completion of the training program). Differences were considered statistically significant when the p-value was < 0.05, and were assumed to be highly significant at p < 0.01.

The study was approved by the Ethical Review Board of the Medical University of Lodz (number RNN/226/11/KB).

**Results**

The baseline clinical data of the study participants are presented in Table 1. Other than the ages of patients, there was no statistically significant difference between the three groups in terms of baseline clinical variables and treatment.

In all three groups the authors observed changes in the four plethysmographic parameters studied (PSlope, PAmpl, CT and PT) in comparison to their baseline values. In Group 1 PSlope increased by 2%, PAmpl by 4.2% and CT by 1.5%, while PT decreased by 1.2%. The change in the mean values of the flow parameters appeared to be highly statistically significant (Table 2).

In Group 2, after four weeks of rehabilitation, PSlope and PAmpl parameters increased by 19% and 17% respectively, while CT and PT decreased by 8% and 6.5% respectively. The mean values of the flow parameters appeared to be highly statistically significant (Table 3).

At the end of cardiac rehabilitation (after an average of 18 days), in the control group there was
### Table 1. Baseline clinical data of the studied patients

<table>
<thead>
<tr>
<th>Demographic and medical variables</th>
<th>group 1 (n = 30)</th>
<th>group 2 (n = 30)</th>
<th>group 3 (n = 30)</th>
<th>p-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sex male : female</td>
<td>18 : 12</td>
<td>22 : 8</td>
<td>20 : 10</td>
<td></td>
</tr>
<tr>
<td>Age (year)</td>
<td>61.18 ± 8.53</td>
<td>59.69 ± 9.24</td>
<td>75.56 ± 7.50</td>
<td>&lt; 0.0001</td>
</tr>
<tr>
<td>BMI (kg/m²)</td>
<td>26.83 ± 5.12</td>
<td>27.02 ± 3.72</td>
<td>26.54 ± 4.18</td>
<td>ns.</td>
</tr>
<tr>
<td>Duration of illness (years)</td>
<td>14.04 ± 3.6</td>
<td>13.17 ± 2.1</td>
<td>12.92 ± 1.9</td>
<td>ns.</td>
</tr>
<tr>
<td>Number of patients treated with CABG</td>
<td>23</td>
<td>19</td>
<td>14</td>
<td>ns.</td>
</tr>
<tr>
<td>Number of patients treated with PCI</td>
<td>7</td>
<td>11</td>
<td>16</td>
<td>ns.</td>
</tr>
<tr>
<td>Medications:</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>beta blockers</td>
<td>30</td>
<td>30</td>
<td>30</td>
<td>ns.</td>
</tr>
<tr>
<td>statins</td>
<td>30</td>
<td>30</td>
<td>30</td>
<td>ns.</td>
</tr>
<tr>
<td>ACE inhibitors</td>
<td>30</td>
<td>30</td>
<td>30</td>
<td>ns.</td>
</tr>
<tr>
<td>aspirin</td>
<td>30</td>
<td>30</td>
<td>30</td>
<td>ns.</td>
</tr>
</tbody>
</table>

Values are expressed as mean ± SD; ns. = not significant.

### Table 2. Values of the studied parameters in Group 1

<table>
<thead>
<tr>
<th>Plethysmographic parameters</th>
<th>Period</th>
<th>Descriptive statistics</th>
<th>Wilcoxon test</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td>average</td>
<td>sd</td>
</tr>
<tr>
<td>PAmpl [%]</td>
<td>before</td>
<td>0.56</td>
<td>0.12</td>
</tr>
<tr>
<td></td>
<td></td>
<td>0.58</td>
<td>0.16</td>
</tr>
<tr>
<td></td>
<td>after</td>
<td>5.79</td>
<td>1.36</td>
</tr>
<tr>
<td></td>
<td></td>
<td>5.82</td>
<td>1.38</td>
</tr>
<tr>
<td>PSlope [%/s]</td>
<td>before</td>
<td>170.32</td>
<td>16.46</td>
</tr>
<tr>
<td></td>
<td></td>
<td>172.36</td>
<td>21.65</td>
</tr>
<tr>
<td>CT [ms]</td>
<td>before</td>
<td>269.82</td>
<td>39.01</td>
</tr>
<tr>
<td></td>
<td></td>
<td>265.55</td>
<td>36.95</td>
</tr>
</tbody>
</table>

### Table 3. Values of the studied parameters in Group 2

<table>
<thead>
<tr>
<th>Plethysmographic parameters</th>
<th>Period</th>
<th>Descriptive statistics</th>
<th>Wilcoxon test</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td></td>
<td>average</td>
<td>sd</td>
</tr>
<tr>
<td>PAmpl [%]</td>
<td>before</td>
<td>0.54</td>
<td>0.17</td>
</tr>
<tr>
<td></td>
<td></td>
<td>0.63</td>
<td>0.14</td>
</tr>
<tr>
<td></td>
<td>after</td>
<td>5.59</td>
<td>1.65</td>
</tr>
<tr>
<td></td>
<td></td>
<td>6.56</td>
<td>1.56</td>
</tr>
<tr>
<td>PSlope [%/s]</td>
<td>before</td>
<td>168.65</td>
<td>15.27</td>
</tr>
<tr>
<td></td>
<td></td>
<td>159.12</td>
<td>14.07</td>
</tr>
<tr>
<td>CT [ms]</td>
<td>before</td>
<td>297.08</td>
<td>36.27</td>
</tr>
<tr>
<td></td>
<td></td>
<td>278.77</td>
<td>37.21</td>
</tr>
</tbody>
</table>

Values are expressed as mean ± SD; ns. = not significant.
a 3% increase in PSlope, a 2.5% increase in PAmpl, and a 1% decrease in PT, but these values were not statistically significant. The only statistically significant change in the control group parameters was the CT, which decreased by 5% (Table 4).

Highly significant differences were found between Groups 1 and 2 at the end of the training program in terms of PSlope and PAmpl (p = 0.0003), as well as CT (p = 0.0233). The difference in the values of the propagation time (PT) were not statistically significant (p = 0.2481).

**Discussion**

Among the participants in the present study, men constituted a distinct majority (70%). This reflects epidemiological data, which shows that twice as many men as women suffer and die from coronary artery disease (CAD) [16]. This is confirmed by the data obtained in the Reversing Atherosclerosis with Aggressive Lipid Lowering (REVERSAL), the Comparsion of Amlodipine Versus Enarapril to Limit Occurences of Thrombosis (CAMELOT) and ACAT Intravascular Atherosclerosis Treatment Evaluation (ACTIVATE) studies, conducted among 978 people (251 female and 727 male).
Effects of Exercise on Peripheral Circulation

The present study investigated the effects of exercise on peripheral circulation in patients undergoing myocardial revascularization or coronary artery bypass graft (CABG) surgery. CABG had been performed on 23 patients in Group 1, 19 patients in Groups 2 and 14 patients in Group 3. Percutaneous coronary intervention (PCI) had been performed on 7 patients in Group 1, 11 in Group 2 and 16 in Group 3. Among the many treatments for heart disease in recent years, invasive methods are being used more and more frequently. This is indicated by the increasing number of centers performing open-heart procedures, as well as by the increase in the number of such procedures performed at cardiac centers in Poland [18]. In this situation, comprehensive cardiac rehabilitation plays an important role: Rehabilitation can contribute to sustaining the positive effects of expensive surgery.

Physical training, especially endurance training, is associated with increased capillary density, improved muscle fiber composition and improvement of the metabolism. The amount of type IIA muscle fiber increases, while type IIB fiber decreases. This process leads to a substantial increase in the cross-sectional area of all types of muscle fibers and an increase in the ratio of fibers to capillary vessels [19]. This is of vital importance, due to the fact that the muscle pump, integrally connected with the function of foot, lower leg and thigh muscles, is the main driving force causing blood movement in the blood vessels of lower limbs. The function of the joint pump, which is dependent on flexion and extension movements (mainly in the ankle joint) causing compression on the venous plexus of the foot, is also crucial [20]. Increased local blood flow intensifies the shear forces and therefore improves the reactivity of the vascular endothelium.

In comparison with baseline values, PAmpl increased in all three groups: in Group 1 by 4.2%, in Group 2 by 17% and in Group 3 by 2.5%. Likewise, there were increases in PSlope in all three groups: in Group 1 by 2%, in Group 2 by 19% and in Group 3 by 3%. The CT on the other hand was reduced by 1.5% in Group 1, by 8% in Group 2 and by 5% in Group 3. The PT also decreased, by 1.2% in Group 1, by 6.5% in Group 2 and by 1% in Group 3. These changes indicate improvement in the vascular perfusion of the lower extremities. Only the results for Group 2 and the CT in the control group were significant and highly statistically significant.

In the present study, the authors used impedance plethysmography to evaluate local blood flow in the calf. Barcroft and Dornhorst started studying this issue in the 1940s by introducing wrap plethysmography as a research technique, analyzing blood flow in the soleus and gastrocnemius muscles of the calf during physical exercise involving contraction and relaxation of posterior calf muscle groups. During alternating dorsal and plantar flexion against resistance, Barcroft and Dornhorst observed a 40% increase in blood flow in the calf area [21]. Reading et al. confirmed a 20% increase in vascular perfusion in the calf area in patients participating in training sessions with a bicycle ergometer with a 25-Watt load [22]. Goodman et al. analyzed the work of the central and peripheral circulatory system in 31 patients who had undergone coronary artery bypass grafting surgery. The patients took part in a 12-week workout session that included walking and jogging, and the researchers observed an 18% increase in vascular perfusion in the calf area. Strain gauge plethysmography was used as a research method [23].

Possible reasons for the increase in local blood flow may be reflected in the changes taking place in the structure and function of the blood vessels. Post-workout changes in peripheral circulation may be due to the development of new collateral blood vessels, or increases in the diameter of existing ones. The increase in local blood flow causes higher shear force influence on the vessels, which is a physiological agent of improving the reactivity of the vascular endothelium. Improvement in endothelial vasodilator function is caused by increases in the blood flow directly related to the reduction in vascular resistance. It is also suggested that chronic increased shear forces on the vessel wall may stimulate the release of growth factors, resulting in the stimulation of vascular adaptation processes and, consequently, an increase in vessel diameter [20].

On the basis of the results of the present authors’ own previous research [13] it can be concluded that increases in PAmpl and PSlope values indicate improvement in the elasticity of blood vessel walls. This may lead to a reduction in vascular resistance and an increase in blood flow, according to Poiseuille’s law. This law explains the relationship between the size of the vessel wall, blood viscosity and flow rate. The flow rate, expressed as the volume of blood flowing through the blood ves-
Changes in the vascular perfusion in lower limbs as a result of physical training may be associated with blood flow redistribution during exercise from less working to more involved muscles. This view was presented by Tan et al., highlighting the reduced oxygen saturation measured in the femoral veins after training [26]. Better use of oxygen by the muscles of the lower limbs is not necessarily associated with better distribution of the circulating blood. This phenomenon can be the result of metabolic changes occurring due to an increase in mitochondrial enzymes and increased oxidative muscle fibers after a training program [27].

Summing up, the present study confirmed that physical activity leads to positive changes in blood flow in the peripheral vessels of the lower limbs in patients with ACS, which is shown by the significant improvement in the plethysmographic parameters in Group 2 after four weeks of controlled physical exercise. Impedance plethysmography allows non-invasive monitoring of local blood flow and is a precise and repeatable method for the assessment of peripheral circulation in patients during cardiac rehabilitation.

References

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