Features of vascular remodelling in the patients with arterial hypertension in a case of brain stroke

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Key words: stroke, hypertension, atherosclerosis.

Purpose of research – to determine the features and factors which influence the vascular remodelling in patients with arterial hypertension in the late restoration period of brain stroke, taking into account an intima-media complex thickness.

Materials and methods. The research enrolled 44 patients with stage III AH in the restoration period of a brain stroke. All participants underwent ultrasound examination of the brachycephal arteries, daily monitoring of arterial pressure, Doppler echocardiography. Patients were divided into 2 groups: the I group included 24 patients (54.5 %) with normal thickness of complex intima-media (CIM), the II group – 20 (45.5 %) patients with thickness of CIM >0.9 mm.

Results. In the patients with stage III AH in a case of brain stroke thickening of CIM was in 45.5 %, concentric hypertrophy – in 25.0 % and remodelling – in 10.0 %. Daily profiles of AP such as night-peaker were observed in 20.0 % of patients, an over-dipper – in 5.0 %, which indicated an unfavourable prognosis regarding repeated cases of the cardiovascular events. Structural and functional features in the vessels of patients with stage III AH were formed under an influence of both diastolic and systolic dysfunction of LV, hypertrophy and concentric remodelling, presence, duration and an inadequate treatment of AH, anthropometric and other risk factors in the patients with AH in the restoration period of a brain stroke, which requires a rigorous tactic of their correction.

Conclusions. Determination of the factors affecting structural and functional peculiarities of vessels in the patients with AH, together with commonly accepted risk factors, should be on a basis to assess their rehabilitation potential, stratification of a common cardiovascular risk, development of an individual rehabilitation program for disabled people with this pathology. Further, the role, character, prognosis and effectiveness of the revealed features correction in the patients with arterial hypertension in the restoration period of a brain stroke should be clarified.

Особливості ремоделювання судин у хворих на артеріальну гіпертензію з перенесеним інсультом

I. В. Дроздова

Артеріальна гіпертензія (АГ) та атеросклероз (особливо перенесений мозковий інсульт) формують структурно-функціональні зміни екстракраніальних артерій, але сьогодні особливості ремоделювання судин у хворих на АГ у віддаленій період мозкового інсулу (МІ) вивчені недостатньо.

Мета роботи – визначити особливості й чинники впливу на ремоделювання судин у хворих на артеріальну гіпертензію у пізному відновному періоді інсулу з урахуванням товщини комплексу інтима-медіа.

Матеріали та методи. 44 хворим на АГ III стадії в пізному відновному періоді МІ здійснили ультразвукове дослідження брахіцефалічних артерій, добове моніторування артеріального тиску та доплерехокардіографіче дослідження. Хворих поділили на 2 групи: перва включала 24 пацієнти (54,5 %) з нормальною товщиною комплексу інтима-медіа (КІМ), друга група – 20 (45,5 %) осіб з КІМ >0,9 мм.

Результати. У хворих на АГ III стадії з МІ потовщення КІМ виявили у 45,5 %, концентричну гіпертрофію – у 25,0 %, ренодевелопмент – у 10,0 %, добові профілі артеріального тиску night-peaker – у 20,0 %, over-dipper – у 5,0 % обстежених, що свідчило про несприятливий прогноз у них щодо повторних серцево-судинних подій. Структурно-функціональні особливості судин у хворих на АГ III стадії формувалися під впливом як діастоличної, так і систоличної дисфункції ЛШ, гіпертрофії ЛШ і концентричного ремоделювання, наявності, тривалості, неадекватного лікування АГ, антропометричних та інших факторів ризику хворих на АГ у віддаленому періоді МІ, що потребує жорсткої тактики їх корекції.

Висновки. Визначення чинників впливу на структурно-функціональні особливості судин у хворих на АГ, поряд із загально-прийнятими факторами ризику має бути в основі визначення їхнього реабілітаційного потенціалу, стратифікації загального серцево-судинного ризику, розробки індивідуальних програм реабілітації інвалідів із цією патологією. Надалі необхідно уточнити роль, характер, прогноз й ефективність корекції виявлених особливостей у хворих на АГ у віддаленому періоді МІ.
Материалы и методы. У 44 больных АГ III стадии в позднем восстановительном периоде МИ проводили ультразвуковое исследование брахиоцефальных артерий, суточное мониторирование артериального давления и допплерэхокардиографическое исследование. Больные были разделены на 2 группы: первая включала 24 пациента (54,5 %) с нормальной толщиной комплекса интима-медиа (КІМ), вторая группа – 20 (45,5 %) лиц с КІМ >0,9 мм.

Результаты. У больных АГ III стадии с МИ утолщение КІМ встречалось у 45,5 %, концентрическая гипертрофия – у 25,0 % и ремоделирование – у 10,0 %, суточные профили артериального давления night-peaker – у 20,0 % и over-dipper – у 5,0 % обследованных, что свидетельствовало о неблагоприятном прогнозе повторных сердечно-сосудистых событий. Структурно-функциональные особенности сосудов у больных АГ III стадии формировались под влиянием как диастолической, так и систолической дисфункции ЛЖ, концентрического ремоделирования и гипертрофии, наличия, длительности и неадекватного лечения АГ, андропометрических и других факторов риска у больных АГ отдленном периоде МИ, что требует жесткой тактики их коррекции.

Выводы. Изучение факторов, влияющих на структурно-функциональные особенности сосудов у больных АГ, наряду с общепринятыми факторами риска, должно быть в основе определения их реабилитационного потенциала, стратификация общего сердечно-сосудистого риска, разработки индивидуальных программ реабилитации инвалидов с этой патологией. В дальнейшем необходимо уточнить роль, характер, прогноз и эффективность коррекции выявленных особенностей у больных АГ в отдленном периоде МИ.

Arterial hypertension (AH), atherosclerosis and especially a brain stroke (BS) form numerous morphofunctional changes (remodelling) in the extracranial arteries, such as a common carotid artery (CCA), which result in increase of their diameter and decrease of vascular wall elastic properties [3,4,8,10,11]. Today, it is well known a relationship between complex of intima-media (CIM), the carotid arteries and risk factors for BS development [7,9,11,14]. One of the most sensitive determinants in possibility of cardiovascular events repeated cases occurrence in persons with AH is consider an intensity of the heart and blood vessels remodelling, size and severity of the endothelial dysfunction and the processes of atherosclerosis and arteriosclerosis [6,7,9,11,13]. Probability to assess the intensity of heart and blood vessels remodelling for a long period of time was focused on the serial measurement of heart, blood vessels walls and cavities thickness and kinetics, analysis of their relaxation characteristics on a background of physical loading, hyperventilation, reactive hyperemia or other pharmacological stress tests [3,5,8–11].

I. N. Dykan, L. V. Melnykova, T. S. Mishchenko, E. V. Pisotska, M. Tendera considered, that thickness of CIM is a specific sonographic marker of early atherosclerotic lesion of vascular wall, indicating about prevalence of atherosclerosis. Its increase varies depending on age, duration and stage of AH, body mass index and coronary arteries pathology, blood glucose concentration, presence of diabetes, smoking, etc. [5,8,9,11,16]. Thickening CIM of CCA is connected with a risk of cardiovascular disease, regardless of the lifestyle, social and other risk factors. Numerous epidemiological studies have been shown, that CIM thickness is associated with a risk of acute myocardial infarction and cerebral ischemic stroke [16,17].

Some of the heart and blood vessels remodelling indicators, which are traditionally assessed by using a duplex impulse or tissue Doppler echocardiography, have a great predictive meaning of the cardiovascular risk development [4,8,11]. To determine the character of blood pressure daily profile, correlation of deformation and frequency of the myocardium deformation with genetic factors, with comorbidity pathology and cognitive dysfunction in patients with AH, could have a significant, but limited value to assess occurrence of the repeated cardiovascular events [6,7,9,11,17]. Therefore, this article review sheds new light on the potential prognostic markers and factors, which influence the morphofunctional condition of the blood vessels in the patients with AH in case of brain stroke.

Purpose of research – to determine the features and factors which influence the vascular remodelling in the patients with arterial hypertension in the late restoration period of brain stroke, taking into account an intima-media complex thickness.

Materials and methods

An open controlled research was conducted on a basis of State Institution “Ukrainian State Research Institute of Medico-Social Problems of Disability Ministry of Health of Ukraine” in the framework of informed consent. Retrospective study enrolled 44 patients with stage III AH in the restoration period of a brain stroke. An inclusion criterion in the research was essential AH in patients aged from 30 to 59 years. Exclusion criteria: secondary forms of AH; renal arteries stenosis; persistent form of atrial fibrillation; severe kidney and liver functions violations; myocardial infarction, cardio-surgical interventions; decompensated diabetes; presence of the comorbidities, which could influence survival rate and quality of patients’ life within 3 months; presence of marked cognitive impairment and psychiatric pathology; treatment refusal. All patients received information and signed informed consent to participate in the research.

AH was diagnosed according to the requirements of National guidelines: “Guidance and Clinical Protocol of Medical Care: Arterial Hypertension”, recommendations of Ukrainian Association of Cardiologists (2013) and recommendations of the European Society of Cardiologists (2013), adapted clinical guidance “Modern Principles of Diagnostic and Treatment Patients with Acute Ischemic Stroke and TIA”, the Standard Clinical Protocol of Medical Care “Ischemic Stroke” [14].

Surveyed groups of patients included: 22 (50.0 %) persons with a delayed consequences of brain infarction (I.69.3); 9 (20.5 %) patients with delayed consequences of intra-cerebral hemorrhage (I.69.1); 3 (6.8 %) – with delayed effects of subarachnoid hemorrhage (I.69.0); 2 (4.5 %) – with unknown diagnoses of brain hemorrhage or infarction (I.69.4); 1 (2.3 %) – in the late restoration period of brain infarction or stroke, caused by thrombosis of brain arteries (I.63.3); 1 (2.3 %) – in the late restoration period of subarachnoid hemorrhage (I.60); 5 (11.3 %) – with delayed consequences of non – traumatic intracranial hemorrhages
Table 1. Factors which influence vascular remodelling in the patients with arterial hypertension in the late restoration period of a brain stroke depending on the intima-media complex thickness

<table>
<thead>
<tr>
<th>Indexes of vessels</th>
<th>Patients with arterial hypertension and ischemic stroke</th>
</tr>
</thead>
<tbody>
<tr>
<td>Remodelling, units of measurement</td>
<td>I group (n = 24)</td>
</tr>
<tr>
<td>D RCCA, cm</td>
<td>0.70 ± 0.01</td>
</tr>
<tr>
<td>Vs RCCA, cm/sec</td>
<td>89.77 ± 2.24</td>
</tr>
<tr>
<td>RI RCCA</td>
<td>0.72 ± 0.01</td>
</tr>
<tr>
<td>D LCCA, cm</td>
<td>0.69 ± 0.01</td>
</tr>
<tr>
<td>RI LCCA</td>
<td>0.72 ± 0.01</td>
</tr>
<tr>
<td>D RICA, cm</td>
<td>0.50 ± 0.01</td>
</tr>
<tr>
<td>Vs RICA, cm/sec</td>
<td>73.61 ± 1.23</td>
</tr>
<tr>
<td>RI RICA</td>
<td>0.71 ± 0.01</td>
</tr>
<tr>
<td>D LICA, cm</td>
<td>0.51 ± 0.01</td>
</tr>
<tr>
<td>Vs LICA, cm/sec</td>
<td>74.72 ± 1.24</td>
</tr>
<tr>
<td>RI LICA</td>
<td>0.71 ± 0.01</td>
</tr>
<tr>
<td>D RVA, cm</td>
<td>0.36 ± 0.01</td>
</tr>
<tr>
<td>Vs RVA, cm/sec</td>
<td>44.45 ± 1.34</td>
</tr>
<tr>
<td>RI RVA</td>
<td>0.71 ± 0.01</td>
</tr>
<tr>
<td>D RVA, cm</td>
<td>0.36 ± 0.01</td>
</tr>
<tr>
<td>Vs RVA, cm/sec</td>
<td>44.20 ± 1.48</td>
</tr>
<tr>
<td>RI RVA</td>
<td>0.71 ± 0.01</td>
</tr>
<tr>
<td>CIM, cm</td>
<td>0.07 ± 0.001</td>
</tr>
</tbody>
</table>

Results and discussion

Affection of carotid arteries (thickening of CIM >0.9 mm) in patients with BS has been found in 20 (45.5 %) persons (II group). On the other hand, 24 patients (54.5 %) have not had the damage of carotid arteries (I group). Majority of patients from the I and II groups had atherosclerotic plaques in the carotid arteries [14, 16, 17].

M. M. Dolzhenko considered, that all patients with AH, complicated by ischemic stroke, have a statistically significant higher thickness of CIM CCA, comparing with healthy people, regardless of the stroke substrate. It was corresponded to the presence of different localization hemodynamically insignificant (<50 %) stenosis in 50.3 % of cases without statistically significant asymmetry of blood flow [4].

Diameter of the right common carotid artery (D RCCA) in the II group of patients increased by 6.49 % (p < 0.01) (Table 1), which correlated with a numerous factors: anthropometric (waist circumference); heart remodelling (aortic root diameter, systolic valve opening, left atrial size, end-diastolic and end-systolic size of left ventricular (LV), end-diastolic volume of LV, its index, end-systolic and percussive volumes of LV, posterior wall of LV thickness, its relative thickness and interventricular septum thickness, myocardium mass of LV, its index) and vessels (diameters of the left common, right and left internal carotid arteries (ICA)). The following factors have been taken into account: loading by pressure (daily and right systolic (SAP) and diastolic (DAP) arterial pressure (AP), time indices, daily variability of DAP, daily minimum, average and maximum SAP and DAP, an average AP and pulse arterial pressure (PAP)).

Velocity of a blood flow in the right common carotid artery (Vs RCCA) in the II group of examined patients decreased by 7.97 % (p < 0.05), which was associated with the such factors as: anthropometric (waist circumference); the heart remodelling (aortic root diameter, systolic valve opening, left atrial size, end-diastolic and end-systolic size of left ventricular (LV), end-diastolic volume of LV, its index, end-systolic and percussive volumes of LV, posterior wall of LV thickness, its relative thickness and interventricular septum thickness, myocardium mass of LV, its index, area of right atrium and its index) and vessels (diameters of the right and left CCA, blood flow velocity in the left common and right and left internal carotid arteries (ICA), in the right and left vertebral arteries (VA)). It is also included loading by pressure (daily DAP, time indices SAP and DAP, daily average DAP and average AP, area index of SAP).

Resistance index of the right common carotid artery (RI RCCA) in the II group of patients increased by 3.19 %, which was associated with heart remodelling (thickness of the interventricular septum) and vessels (resistance index of left CCA and right ICA, right VA, diameter and resistance index of left VA).

Velocity of blood flow in the left common carotid artery (Vs LCCA) in the II group of patients decreased by 10.75 % (p < 0.001), which was associated with such factors as: anthropometric (body weight, waist circumference); heart remodelling (gradient of pressure on mitral valve, transverse flow velocity and in pulmonary artery, gradient of pressure in pulmonary artery, size, index and area index of left atrium, end-diastolic size and volume of LV, indexes of end-diastolic and percussive volumes of LV, posterior wall of LV thickness, its relative thickness and interventricular septum thickness).
tum thickness, myocardium mass of LV, its index, area of right ventricular) and vessels (blood flow velocity in the right and left ICA, in the right and left VA, in the right CCA, diameters of the right and left CCA).

The Vs LCCA could be also associated with the next factors: loading by pressure (daily SAP and DAP, their time indices, right SAP and DAP, daily average SAP and DAP, average AP, maximum DAP, the area index of DAP, average heart rate (HR)).

Comparing parameters of vascular remodelling in the patients with stages II and III AH O. O. Lisova revealed that vascular wall elasticity demonstrated statistically significant differences. Other vascular remodelling parameters in the patients with stage III AH were lower, than in the patients with stage II AH, but there were no significant differences between indicators of CIM CCA in both groups of patients [8]. M. M. Dolzhenko demonstrated, that patients with AH and ischemic stroke have statistically significant lower diameter of CCA and distensibility coefficient, a higher index of stiffness, compared with healthy people. This fact indicates a progressive remodelling of arterial blood flow in patients with AH, complicated by ischemic stroke [4].

Diameter of the right internal carotid artery (D RICA) in the II group of patients increased by 8.14 % (p < 0.01), which was associated with a heart remodelling (left ventricular index) and vessels (diameters of the right and left CCA, left ICA, blood flow velocity in the left ICA and right VA); with loading by pressure (time indices and daily variability of SAP, daily average and maximum DAP and average AP).

Complex of intima-media (CIM) in the II group of patients increased by 77.27 % (p < 0.001) and was varied in average 0.117 ± 0.04 mm. This one was correlated with heart remodelling (end-diastolic size of LV) and vessels (diameter of the left CCA).

M. M. Dolzhenko study demonstrated, that the worst indicators of extra cranial arteries remodelling in the patients with brain stroke by stiffness index were statistically significant correlated with systemic atherosclerosis progression in terms of CIM thickness of CCA [4].

According to the data of daily AP monitoring, among non-specific persons (the I group) and among persons with CIM thickening (the II group), there were the following patients with daily AP profile: non-dipper – 8 (33.3 %) and 6 (30.0 %), night-peaker – 2 (8.33 %) and 4 (20.0 %), dipper – 13 (54.17 %) and 9 (45.0 %), over-dipper – 1 (4.2 and 6.59 %) respectively.

It has been shown in the patients from the II group the increase in a daily SAP and DAP by 8.53 % (p < 0.001) and by 6.59 % (p < 0.05); time indices SAP and DAP – by 122.88 and 95.91 % (p < 0.05), variability of SAP – by 10.83. Similar trend has been observed for the same indicators in the I group of patients.

Myocardium posterior wall thickness, relative thickness of myocardium, interventricular septum thickness were increased by 5.65; 0.70; 5.60 % (p > 0.05) respectively, in the patients with stage III AH, i.e. the II group. LV myocardium mass and myocardium mass index were increased by 16.09 % (p < 0.05) and 10.35 %, respectively, in the II group of patients with stage III AH. In the II group of patients end-diastolic size of RV, area of a right atrium and the right atrium area index were increased by 8.59 % and 13.81 % (p < 0.01), and on 7.38 %, respectively, comparing with the group of patients with normal CIM thickness. In comparison with the I group, patients with stage III AH, who concerned to the II group, had the following changes: increasing in stroke volume by 2.30 %, decreasing in ejection fraction by 3.94 %, respectively.

It has been revealed that in the patients from the II group day SAP and DAP were significantly higher. Minimum, average and maximum day SAP and DAP in this group of patients increased by 2.11 and 3.36 % (p > 0.05); by 7.41 % (p < 0.05) and 5.25 %; by 8.96 % (p < 0.001) and 7.32 % (p < 0.05), respectively. Day minimum, average and maximum average and pulse AP have had a similar trend: in the II group of pa-
of CIM CCA and relative thickness of LV wall, presence of LV hypertrophy indicated that heart remodelling and systemic atherosclerosis progression in a case of AH could be the parallel process. On the other hand, they are the different links of common pathogenetic mechanism. They should be evaluated comprehensively in the patients with AH and ischemic stroke for the common cardiovascular risk stratification [4,6,7].

In the conclusion, it should be noted, that in the patients with stage III AH in the restoration period of a brain stroke CIM thickening was observed in half of the cases among examined patients. The atherosclerotic plaques of carotid arteries with a different localization hemodynamically insignificant stenosis without significant blood flow asymmetry were defined almost in all patients. Persons with CIM thickening were characterized by increased diameter by 8.49 % (p < 0.01) and decreased blood flow velocity by 7.97 % (p < 0.05) in the right CCA. Trend towards increasing of the diameters and indices of CCA and ICA resistance with blood flow velocity decreasing has shown a progressive arterial remodelling in the patients with AH in case of ischemic stroke. These features were formed under the influence of LV diastolic and systolic dysfunction (increase in size and index of left atrium, end-diastolic size and volume, end-systolic size and index of end-diastolic LV volume); LV concentric remodelling and hypertrophy (increase in thickness of myocardium posterior wall and interventricular septum, mass and myocardial mass index). It was also determined by the presence, duration and an inadequate treatment of AH (achievement of AP target level index). It was also determined by the presence, duration and an inadequate treatment of AH (achievement of AP target level index). It was also determined by the presence, duration and an inadequate treatment of AH (achievement of AP target level index).

Conclusions

1. In patients with stage III AH in the restoration period of a brain stroke CIM thickening was observed in half of cases among examined patients, and atherosclerotic plaques of carotid arteries with a different localization hemodynamically insignificant stenosis without significant asymmetry of blood flow were defined almost in all patients. This fact indicated a progressive arterial remodelling in the patients with AH in case of ischemic stroke.

2. Mainly among patients with stage III AH in case of brain stroke CIM thickening was in 45.5 %, the concentric hypertrophy – in 25.0 % and remodelling – in 10.0 %. Daily AP profiles were observed as well as night-peaker – in 20.0 % of patients, over-dipper – in 5.0 %, which indicated an unfavourable prognosis regarding repeated cases of the cardiovascular events.

3. Structural and functional features in the vessels of patients with stage III AH were formed under an influence of both LV diastolic and systolic dysfunction, LV hypertrophy and concentric remodelling, presence, duration and an inadequate treatment of AH, anthropometric and other risk factors in the patients with AH in the restoration period of a brain stroke. It requires a rigorous tactic of their correction.

4. Determination of factors which influence the structural and functional peculiarities of vessels in patients with AH, together with commonly accepted risk factors, should be on a basis to assess their rehabilitation potential, stratification of a common cardiovascular risk, development of an individual rehabilitation program for disabled people with this pathology.

Prospects of further research involve the clarification of role, character, prognosis and effectiveness of drug and non-pharmacologic correction of the revealed features in patients with arterial hypertension in the restoration period of a brain stroke.

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