EFFECT OF PARTIAL AND TOTAL CORONARY ARTERY OCCLUSION IN RATS ON

ACTIVITY OF CREATINE KINASE AND ITS MB ISOZYME

É. I. Krasnova, * P. P. Minin, and I. S. Zavodskaya

UDC 616.132.2-007.271-092.9-07:[616.127-008.931+616.153.1]:577.152.273

KEY WORDS: rat heart; arterial occlusion; myocardial infarction; creatine kinase.

To make a comprehensive study of the initial stages of myocardial infarction the writers have previously used with success models of lesions of heart muscle in small animals involving "extraordinary" stimulation of reflexogenic zones [1, 3]. To continue the study of the stages of myocardial infarction and changes in myocardial metabolism in chronic coronary insufficiency the writers have developed a model providing various degrees of occlusion of the left coronary artery (LCA) in rats.

In the investigation described below, activity of creatine kinase (CK) and its MB isozyme (CK-MB) was studied in the heart muscle and blood serum of rats at different times after partial and total occlusion of LCA.

EXPERIMENTAL METHOD

Experiments were carried out on 62 non-inbred male rats weighing 180-200 g. LCA was occluded in anesthetized (amobarbital with chlorpromazine, intraperitoneally) with artificial ventilation of the lungs, by occluding the artery totally or partially by introducing a smooth stilet measuring 1.25 mm in section [2].

CK and CK-MB activity was studied at different times of occlusion of LCA in parallel spectrophotometric tests on blood serum and heart muscle extract [6]. The CK isozymes were fractioned by microcolumn chromatography [5] with certain modifications [3]. CK activity was expressed in international units (IU) and CK-MB activity was calculated as a percentage of total CK activity.

EXPERIMENTAL RESULTS

Levels of CK and CK-MB activity in the myocardial and blood serum of intact rats (Table 1) agreed with data in the literature [4]. Total serum CK activity in animals undergoing a mock operation was moderately raised compared with that in intact rats. Relative CK-MB activity was not significantly changed, evidence that the heart muscle is not the principal source of creatine kinase activity in the blood when it is raised as a result of a mock operation.

After partial occlusion of LPA total CK and CK-MB activity decreased in the heart; the greatest decrease in CK and CK-MB activity was observed 24 h after the operation (from 1233.8 \pm 47.9 to 343.3 \pm 21.6 IU/g and from 23.3 \pm 1.0 to 20.7 \pm 1.1%, respectively). Conversely, CK and CK-MB activity in the blood serum increased. Blood enzyme activity was raised as early as 15 min after coronary arterial occlusion, and its highest level was reached after 24 h (49.0 \pm 3.7 and 324.2 \pm 20.4 IU/ml and 5.5 \pm 0.6 and 9.5 \pm 0.4%, respectively).

Total occlusion of LCA led to a more marked fall of CK and CK-MB activity in the heart muscle and to an increase in their activity in the blood compared with values obtained in rats subjected to incomplete occlusion of the artery. For instance, CK and CK-MB activity 1 h after total occlusion of LCA was 4-5 times higher than in the control groups. Relative CK-MB activity was higher after complete than after incomplete occlusion, evidence of the

*Corresponding member of the Academy of Medical Sciences of the USSR.

Department of Pharmacology, Institute of Experimental Medicine, Academy of Medical Sciences of the USSR. No. 2 Chair of Internal Medicine, S. M. Kirov Military Medical Academy, Leningrad. Translated from Byulleten' Éksperimental'noi Biologii i Meditsiny, Vol. 102, No. 11, pp. 554-556, November, 1986. Original article submitted February 14, 1986.

TABLE 1. Changes in CK and CK-MB Activity during Coronary Arterial Occlusion in Rats

Group of animals	Number of animals	Heart		Blood	
		CK, IU/g	CK-MB, %	CK, IU/m1	CK-MB, %
Intact	8	1233,8±47,9	23,3±1,0	49,0±3,7	5,5±0,6
Mock operation (after 1 h)	6	1228,3 \pm 44,9 >0,05	20.7 ± 1.1 >0.05	75,3±4,8 <0,001	$4,9\pm0,6$ >0,05
With partial occlusion after 15 min	8	$1001,1\pm44,3$ <0.01	$16,5\pm0,5$ <0.001	110,5±5,8 <0,001	$7,0\pm0,7 > 0.05$
after 1 h	8	847,5±30,6 <0,001	15,3±0,4 <0,001	188,3±6,7 <0,001	$ \begin{array}{c c} 10,0\pm0,8 \\ < 0,001 \end{array} $
after 24 h P	6	$343,1\pm21,6$ < 0,001	$\begin{array}{c} 20,7 \pm 1,1 \\ > 0,05 \end{array}$	$324,2\pm20,4$ $<0,001$	9.5 ± 0.4 <0.001
after 14 days With total occlusion	6	868,3±19,0	20,3±1,0	45,0±2,8	5,0±0,7
after 15 min	12	<0.001 887.5 ± 17.5 <0.001	>0,05 16,6±0,6 <0.001	>0,05 179,0±4,5 <0,001	>0.05 10.2 ± 0.6 <0.001
after 1 h	8	$\begin{array}{c} <0,001\\ 216,3\pm10,1\\ <0,001 \end{array}$	$ \begin{array}{c c} & <0,001 \\ & 13,0\pm1,1 \\ & <0,001 \end{array} $	261,8±9,3 <0,001	$\begin{array}{c} <0,001\\ 22,7\pm2,4\\ <0,001 \end{array}$

important role of the myocardium as the source of the raised blood enzyme level associated with total OCA occlusion.

The results are thus evidence that the intensity of the rise in the blood enzyme levels is directly proportional to the degree of occlusion of LCA, and levels of CK activity and activity of its cardiospecific MB isozyme in the blood and myocardium are inversely proportional to one another: A greater fall of myocardial enzyme activity corresponds to a higher rise in its level in the blood serum.

It can be concluded from a comparison of these results that the model of chronic myocardial ischemia involving partial occlusion of LCA in rats can be used with advantage to study various forms of disturbance of the commany circulation and ways of correcting metabolism in heart muscle.

LITERATURE CITED

- 1. S. V. Anichkov, I. S. Zavodskaya, E. V. Moreva, et al., Neurogenetic Dystrophies and Pharmacotherapies, Leningrad (1969).
- 2. É. I. Krasnova, Abstracts of Proceedings of the 11th All-Union Conference on the Physiology and Pathology of Corticovisceral Relations [in Russian], Leningrad (1981), p. 240
- 3. P. P. Minin, Byull. Éksp. Biol. Med., No. 7, 51 (1980).
- 4. D. W. Mercer and M. A. Varat, J. Clin. Chem., 21, 1088 (1975).
- 5. S. B. Rosalki, J. Lab. Clin. Med., 69, 696 (1967).