SPECIAL ARTICLES

SCIENCE

THE DESTRUCTION OF COENZYME I AND COCARBOXYLASE IN SKELETAL AND CARDIAC MUSCLE AFTER DEATH

In the course of studies of coenzyme breakdown in heart muscle after experimental coronary ligation in dogs, the idea was entertained that the amount of coenzyme breakdown after one hour of death might be considered the maximum breakdown that could be expected in that length of time following any experimental procedure. In order to test this hypothesis, six dogs (previously on a standard diet of "Pard" dog food) were anesthetized with pentobarbital sodium, the chests were opened and the hearts removed. Immediately on removal of the heart, a sample of left ventricle and a similar sample of deltoid muscle were removed, heated in boiling water for five minutes, homogenized and subsequently analyzed for cocarboxylase and coenzyme I. One hour later a second sample of each tissue was removed and treated in a similar manner. The dog and excised heart were allowed to remain at room temperature during the hour. Coenzyme I was determined by the method of Axelrod and Elvehjem,¹ as modified by Greig.² Cocarboxylase was determined manometrically, using split carboxylase prepared as described by Green et al.³

The results of these analyses are submitted in Tables 1 and 2. TABLE 1

| Exp. No. | Skeletal muscle control | Skeletal muscle 1 hr. post- mortem | Per cent. change | Heart muscle control | Heart muscle 1 hr. post- mortem | Per cent. change |
|---|---|---|--|--|--|--|
| II 1. II 2. II 3. II 4. II 5. II 6. Average | $2730 \\ 2430 \\ 1649 \\ 2640 \\ 924 \\ 1173$ | $2650 \\ 3380 \\ 0 \\ 2235 \\ 783 \\ 1725$ | $\begin{array}{r} - 2.9 \\ + 28.1 \\ - 100.0 \\ - 15.1 \\ - 15.3 \\ + 47.0 \\ - 9.7 \end{array}$ | 3120 3180 881 3920 642 2955 | $2150 \\ 2540 \\ 155 \\ 2375 \\ 826 \\ 2610$ | $\begin{array}{r} -30.9 \\ -20.1 \\ -82.5 \\ -39.3 \\ +28.7 \\ -11.7 \\ -25.9 \end{array}$ |

COENZYME I IN γ /GM OF DRY MUSCLE

One may be impressed by the surprising lack of destruction under these conditions. Instead of finding nearly complete breakdown of cocarboxylase and coenzyme I, in the case of cocarboxylase in skeletal muscle at least, there is frequently less breakdown than was found previously in shock.⁴

¹ A. E. Axelrod and C. A. Elvehjem, Jour. Biol. Chem., 131: 77, 1939.

² Margaret E. Greig, personal communication. ³ D. E. Green, D. Herbert, V. Subrahmanyan, Jour. Biol. Chem., 138: 327, 1941.

4 Margaret E. Greig and Wm. M. Govier, Jour. Pharmacol. and Exp. Therapy, 79: 169, 1943.

TABLE 2

| | COCARBOXY | LASE IN | ү/бм` о | F DRY | MUSCLE | |
|---|--|---|---|--|--|--|
| Exp. No. | Skeletal muscle control | Skeletal muscle 1 hr. post- mortem | Per cent. change | Heart muscle control | Heart muscle 1 hr. post- mortem | Per cent. change |
| II 1. II 2. II 3. II 4. II 5. II 6. Average | 23.7 22.8 28.3 21.1 21.3 18.4 | 22.520.318.814.220.920.4 | $ \begin{array}{r} - 3.4 \\ - 10.9 \\ - 33.6 \\ - 32.7 \\ - 1.4 \\ + 9.8 \\ - 12.0 \\ \end{array} $ | 55.0 68.6 63.9 48.6 71.0 65.8 | 61.2 53.3 55.0 35.4 53.0 63.3 | $ \begin{array}{r} + 10.1 \\ - 22.3 \\ - 13.9 \\ - 27.1 \\ - 25.4 \\ - 3.8 \\ - 13.7 \end{array} $ |

By way of explanation, one may suggest that after death the breakdown products of the coenzymes are not removed from the tissue by the circulation, and consequently by their accumulation may serve to inhibit catabolic enzymes. This explanation may be supported by work showing that thiamin inhibits yeast cocarboxylase phosphatase⁵ and that nicotinamide inhibits coenzyme I nucleotidase.⁶

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AN EXPERIMENTAL METHOD FOR EVALU-ATING BLOOD SUBSTITUTES

REPORT ON SALINE, PLASMA, POLYVINYL ALCOHOL AND ISINGLASS

THE following conditions are desirable in an experiment designed to evaluate substances which may be used for the intravenous treatment of traumatic shock:

(1) The group of animals which is used for the test should be as nearly homogeneous as possible.

(2) The trauma should be of the same degree in each animal and easy to apply. Complicating factors such as anesthesia and hemorrhage into the injured tissues should be reduced to a minimum.

(3) The injury should be highly fatal in the control animals but mild enough to allow a proportion of the treated animals to recover. Percentage survival is considered to be the best single index of the effectiveness of treatment.

(4) The experiments should be carried out with the animals kept at a constant environmental temperature.1

(5) If a large number of animals can be handled at one time, there are the obvious advantages of con-

⁵ H. G. K. Westenbrink, D. A. vanDorp, M. Gruber, H. Veldman, *Enzymologia*, 9: 73, 1940.

⁶ P. J. G. Mann and J. H. Quastel, Biochem. Jour., 35: 502, 1941.

¹F. M. Allen, Arch. Surg., 41: 155, 1939.