INCORPORATION IN VIVO OF C14 FROM LABELED METHANOL INTO THE METHYL GROUPS OF CHOLINE

Sir:

An investigation has been undertaken to test whether methyl groups supplied in the form of methanol can enter into transmethylation reactions of the body. An earlier test of whether methanol could support the growth of animals on a diet free of "biologically labile" methyl groups and containing homocystine was negative.¹ In the present experiment the more sensitive tracer technique, utilizing C14-labeled methanol, has been employed. The amount of radioactivity in the methyl groups of the choline isolated from the tissues of the rat after the administration of the labeled methanol was such as to indicate that methanol made available appreciable amounts of methyl groups which could be used in the transmethylation reactions of the body. The possible significance of this finding to the mechanism of transmethylation and even to the biological synthesis of "labile" methyl groups becomes of considerable importance and is being further investigated. In this connection the interesting observation of Binkley and Watson² may be pointed out, that methyl phosphate appears to be utilized in the formation of creatine from guanidoacetic acid by rat liver homogenates.

A total of 9 ml. of a 2.4% aqueous solution of C¹⁴-labeled methanol with an activity of 5.33×10^6 counts per minute per ml. was injected subcutaneously in 1-ml. portions twice daily into a 161-g. rat over a five-day period. During this time the animal was kept in an open-circuit metabolism apparatus for the collection of the expired carbon dioxide. For fifteen days prior to injection and for the duration of the experiment the rat was allowed free access to a diet of the following composition (in g.): sucrose 54.85, vitamin-free casein 20, pL-methionine 0.15, fat (Covo) 19, Osborne and Mendel salt mixture 4,

(1) du Vigneaud, Chandler, Moyer and Keppel, J. Biol. Chem., 131, 57 (1939).

(2) Binkley and Watson, ibid., 180, 971 (1949).

corn oil (Mazola) 1, containing 4.0 mg. of α tocopherol acetate, 0.1 mg. of 2-methyl-1,4naphthoquinone, 750 I. U. of vitamin A and 125 I. U. of vitamin D; water-soluble vitamins, administered *per os* twice daily, in the following amounts (mg. per day): thiamine hydrochloride, riboflavin, pyridoxine hydrochloride, nicotinic acid and *p*-aminobenzoic acid, 0.08 mg. each; calcium *d*-pantothenate 0.4, inositol 0.8, folic acid 0.02 and biotin 0.0008; 2 micrograms of vitamin B₁₂ every other day.

During the five-day period a radioactivity of 22×10^6 counts per minute, out of the total injected radioactivity of 48×10^6 counts per minute, appeared in the expired carbon dioxide. The animal was then sacrificed; choline was isolated from the carcass as the chloroplatinate (*Anal.* Calcd. for C₁₀H₂₈N₂O₂·PtCl₆: Pt, 31.68. Found: Pt, 31.14), and creatine as the creatinine potassium picrate (purity determined by the Jaffe reaction, 100%). The choline was then degraded to trimethylamine, which was isolated as the chloroplatinate and recrystallized from waterethanol (*Anal.* Calcd. for C₆H₂₀N₂·PtCl₆: Pt, 36.96. Found: Pt, 37.06). The specific activities of these compounds, determined after combustion and isolation of the carbon dioxide as barium carbonate, are given in the table, in terms of counts per minute per millimole of compound.

Compound	Specific activity
C ¹⁴ -Labeled methanol injected	ca. 7×10^6
Choline chloroplatinate	$7.18 imes 10^5$
Trimethylamine chloroplatinate	$6.45 imes10^{5}$
Creatinine potassium picrate	$1.11 imes10^5$

No exchange of methyl groups was found to occur between choline and C^{14} -labeled methanol, allowed to stand together for several days.

This work has been confirmed with another animal. Complete details of these experiments and related ones will be forthcoming shortly.

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NEW BOOKS

Ion Exchange, Theory and Application. Edited by FREDERICK C. NACHOD, Sterling-Winthrop Research Institute, Rensselaer, New York. Academic Press, Inc., Publishers, 125 East 23d Street, New York, N. Y. 1949. xxii + 411 pp. Illustrated. 16.5 × 24 cm. Price, \$8.50.

Since the publication of methods for manufacturing synthetic ion exchange resins by Adams and Holmes, the ion exchange process has spread from a few applications to many applications in almost all fields of chemistry and related fields. The large volume of published literature in this field is so diversified in nature and contained in so many journals that it has been extremely difficult for anyone interested in the field to survey it. This volume is very timely and will be of great value to those desiring to apply ion exchange techniques to specific problems in a large variety of fields of study, as well as a reference book for courses in chemistry and chemical engineering.

In addition to a very short introduction, the subject matter of this volume is divided into sixteen chapters: Ion Exchange Equilibria, The Kinetics of Fixed-Bed Ion Exchange, Fundamental Properties of Ion Exchange Resins, Ion Exchange Equipment Design, Ion Exchange in Water Treatment, Multistage Systems in Ion Exchange, Desalting Sea Water, Applications of Ion Exchange to the Separation of Inorganic Cations, Ion Exchange as a Tool in Analytical Chemistry, Metal Concentration and Recov-