## LETTER TO THE EDITOR

# INFLUENCE OF METHANOL ON THE RELEASE OF DOPAMINE FROM RAT BRAIN SLICES - AN IN VITRO STUDY

(Received on June 15, 1993)

Sir,

Methanol, the simplest of the alcohols, produces severe metabolic toxicity in man. It forms part of the adulterant of the illicit liquors made by bootleggers and also used extensively in industries as a useful solvent. The toxic syndrome due to methanol poisoning includes severe metabolic acidosis, ocular toxicity leading to blindness and CNS deterioration ultimately resulting in high rate of mortality. 90% of methanol is metabolised in the liver by alcohol dehyrogenase to produce formaldehyde and formic acid which is excreted through kidneys. The remaining 10% is excreted unchanged through the lungs and kidney. The half-life of methanol is about 20-24 hrs.

Ethanol is known to alter the biogenic amine release from the brain tissues. It has been shown that methanol change the brain content of biogenic amines when administered in vivo (1). However, it is not known whether methanol alters the release of biogenic amines from the brain slices in vitro also. Hence this preliminary work was undertaken. In the present study, it was proposed to study only dopamine release as a model.

Healthy adult male Wistar strain of albino rats (150-180 g) were used for this study. The animals were housed under standard laboratory conditions and fed with standard rat feed and water ad libitum. The experimental animals were decapitated by rapid guillotione method at 8 AM and hypothalamus dissected out on pre-cooled glass plate (2), weighed and immediately transferred to 6 ml of incubation medium (Krebs bicarbonate buffer). The medium with brain slices was incubated at 37°C for 30 min (Group I) and 60 min (Group II) respectively. 10 control slices and 20 test slices (10 for each time period) were used. The

test slices were incubated with medium containing methanol (250 mg%). After incubation, the medium was contrifuged at 1000 RPM for 5 min and supernatant was removed for fluorometric analysis of dopamine (3).

It was found that the dopamine levels increased significantly (P < 0.001) in both 30 min and 60 min incubation with methanol. They are shown in Fig. 1.

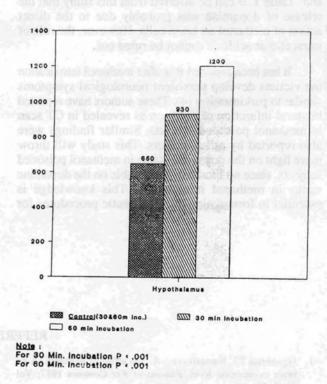


Fig. 1: Dopamine levels in the incubating medium. (ng/g of hypothalamus)

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TABLE I: Dopamine levels in the incubating medium (ng/g of hypothalamus).

| S. No. | Control<br>hypothanalamus<br>slices | Hypothalamic slices with methanol in the medium |                         |
|--------|-------------------------------------|---|-------------------------|
|        |                                     | 30 min of incubation                            | 60 min of in cubation   |
| 1.     | 686                                 | 940   | 1000                    |
| 2.     | 555                                 | 714   | 958                     |
| 3.     | 925                                 | 1078  | 1332                    |
| 4.     | 480                                 | 727   | 1416                    |
| 5.     | 761                                 | 933   | 1250                    |
| 6.     | 665                                 | 999   | 1050                    |
| 7.     | 553                                 | 800   | 999                     |
| 8.     | 623                                 | 1249  | 1605                    |
| 9.     | 666                                 | 810   | 965                     |
| 10.    | 669                                 | 1050  | 1435                    |
| N = 10 | $\Sigma x = 6574$                   | $\Sigma x = 9300$                               | $\Sigma x = 12010$      |
|        | x = 657.4                           | x = 930   | x = 1201                |
|        | S.D. = 123.53                       | S.D. = 170.65                                   | S.D. = 236.29           |
|        | S.E. = 39.09                        | S.E. = 54.00                                    | S.E. = 74.77            |
|        |                                     | 'T' Value 4.092 (P<.001)                        | 'T' Value 6.447 (P<001) |

and Table 1. It can be inferred from this study that the release of dopamine was probably due to the direct action of methanol on brain cells. However, the role of intracellular acidosis cannot be ruled out.

It has been reported that after methanol intoxication the victims develop persistent neurological symptoms similar to parkinsonism (4). These authors have reported bilateral infarction of putamen as revealed in CT scan in methanol poisoned subjects. Similar findings were also reported by other workers. This study will throw more light on the dopamine status in methanol poisoned subjects, since no literature is available on the dopamine status in methanol intoxication. This knowledge is essential in formulating the therapeutic procedures for

these patients in future.

This study will also form the basic model to explore further the mechanism of action of methanol. It is proposed to use various blockers which blocks the synthesis and break down of the monoamines, to study the biodynamics of monoamines under the influence of methanol, at a later date. Further this study raises the question: does methanol increase the release of dopamine by altering the fluidity of cell membranes in the brain slices?

#### ACKNOWLEDGEMENTS

Authors thank Indian Council of Medical Research for the financial support for this study.

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