

LETTER TO THE EDITOR

PROLONGATION OF PENTOBARBITONE INDUCED ANAESTHESIA BY AZATHIOPRINE

Sir,

(Received on November 12, 1991)

A study was carried out on albino rabbits to find out the possible interaction between azathioprine (AZT), a widely used immunosuppressant and pentobarbitone which is commonly used for induction of anaesthesia. 10 albino rabbits (average wt 1.5 kg) of either sex were used. 5 of them were pretreated with AZT (12.5 mg/kg/day orally through a feeding tube for 5 days) and the rest 5 served as control. All of them were given pentobarbitone (30 mg/kg as IV bolus in 6% solution). Appearance of corneal reflex, pain sensation and righting reflex were noted (Table I).

TABLE I : Time taken (in min, Mean \pm SD) for the appearance of various parameters.

Rabbit	Corneal reflex	Pain sensation	Righting reflex
Control	21.6 \pm 2.073	89.2 \pm 2.280	120.0 \pm 3.162
AZT treated	36.6 \pm 3.714	124.0 \pm 3.162	208.6 \pm 9.813

(P < 0.001 for all the three parameters)

The delay in appearance of all these parameters in AZT treated group was found to be statistically significant (P<0.001).

AZT causes significant interactions with many drugs (1). Marked prolongation of anaesthetic effect of pentobarbitone in AZT treated rabbits as reflected by these parameters may result either due to direct interaction between the two drugs or due to delayed pentobarbitone metabolism consequent to hepatocellular suppression caused by AZT. Hepatotoxicity of AZT is well documented (1-4). The features of which are centrilobular congestion, fatty change, hemosiderin deposition, liver cell necrosis, cholestasis, bile duct proliferation and portal infiltrates of mononuclear cells and neutrophils. Before clinically detectable hepatic damage is produced AZT causes suppression of microsomal enzyme system (4) which is perhaps responsible for the prolonged pentobarbitone activity.

PUSHPENDRA SHARMA*, R. C. GUPTA**
AND A. K. MATHUR*

Departments of *Medicine and Pharmacology,
Jawahar Lal Nehru Medical College &
Associated Group of Hospitals,
Ajmer - 305 001 (Rajasthan)

REFERENCES

1. Davis M, Eddleston ALWF, Williams R : Hypersensitivity and jaundice due to azathioprine. *Post Graduate Medical Journal* 1980; 56 : 274-275.
2. Marubio AT, Danielson B. Hepatic veno-occlusive disease in a renal transplant patient receiving azathioprine. *Gastroenterology* 1975; 69 : 739-745.
3. Millard PR, Herbertson BM, Evans DB, Calne RY. Azathioprine hepatotoxicity in renal transplantation. *Transplantation* 1973; 16 :527-530.
4. Zaraday Z, Veith FJ, Gliedman ML, Soberman R. Irreversible liver damage after azathioprine. *Journal of American Medical Association* 1972; 222 : 690-691.