Effect of Liv.52 on Growth and Alcohol-induced Hepatic Dysfunction in Rats

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Our earlier work on mice and rats has shown encouraging protective effects of Liv.52 against carbon tetrachloride induced hepatic toxicity (Joglekar *et al.* 1963; Karandikar *et al.* 1963). Hence the present work was undertaken on rats to study the effect of Liv.52 on growth and alcohol induced hepatic dysfunction.

MATERIALS AND METHODS

Part A: Effect on Growth:

We used 54 female albino rats of similar age and of weight range 90-100 gm divided into equal groups; one of them acted as a control. The other group received 4.0 ml/100 gm of Liv.52 paediatric drops daily by intragastric tube for seven weeks. Both groups were given the same diets (15 gms/rat/day) consisting of wheat flour, wheat bran, milk powder, calcium carbonate, table salt and kardai oil. Gain in weight was recorded every milk.

Part B: Effect of Alcohol-induced Hepatic Dysfunction:

We used 14 albino rats of either sex and weight range 100-150 gm; divided them into two equal groups. One group was given orally 1.2 ml/100 gm body weight of 25% ethyl alcohol daily while the other group received Liv.52 paediatric drops 0.4 ml/100 gm along with the same dose of alcohol. All the rats were given the usual standard diet mentioned previously. At the end of 4 months, 2 rats from each group were sacrificed for ascertaining histological damage to the liver.

RESULTSResults are given in Tables 1 and 2.

Table 1: Effect on growth				
Period	Control – Gain in wt. in gms/100 gms	Liv.52 treated - gain in wt. in gms/100 gms.	Significance	
1 week	13.8	17.3	<i>p</i> >0.1	
2 weeks	6.5	8.9	p>0.1	

3 weeks	1.0	7.7	p<0.05
4 weeks	0.2	1.7	p<0.05
5 weeks	5.2	6.8	p<0.05
6 weeks	5.3	13.8	<i>p</i> <0.05
7 weeks	2.3	2.9	<i>p</i> >0.1

Table 2: Part B			
Sleeping time in	Sleeping time in		
minutes	minutes		
Alcohol group	Alcohol + Liv.52 group		
37	27		
71	31		
52	32		
68	27		
69	36		
Mean 59.4 ± 3.3	Mean 30.6 ± 3.3 p < 0.01		

Part A: Table 1 (See Photographs)

Liv.52 has caused significant gain in weight as compared with control during 3rd to 6th weeks.

Part B: Table 2

Alcohol group showed a significant prolongation of hexobarbitone sleeping time as compared with the group receiving alcohol and Liv.52

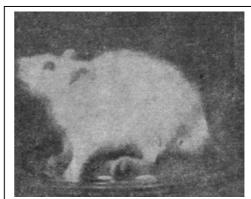
Histopathology of Livers:

Initial fibrotic changes and suggestion of lobulation were observed in the livers from the alcohol group when the rats were sacrificed after 4 months of alcohol therapy. In the other group no such changes were noticed.

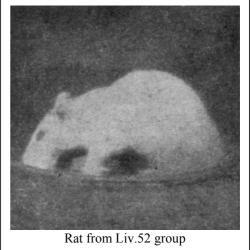
COMMENTS

Kopera and Armitage (1954) studied the effect of chlorpromazine and other drugs on growth by noting the increase or decrease in weight in order to have an approximate estimate of chronic toxicity. On a similar basis in this work the effect of Liv.52

as compared with the control rats.



Rat from Control group



administration (for seven weeks) was studied in rats. From the results (Table 1) it appears that Liv.52 causes significant gain in weight especially during 3rd and 6th weeks of administration

Liv.52 has shown protective effects against liver toxicity in both clinical and experimental work (Patrao *et al.* 1957; Sheth *et al.* 1960). Liv.52 has shown promising results against carbon tetrachloride-induced liver damage (Joglekar *et al.* 1963, Karandikar *et al.* 1963). In this work ethyl alcohol – an alleged hepatotoxic agent was used. Ashworth (1947) fed 2.4 ml/100 gm of 25% ethyl alcohol to rats for 8 weeks to cause fatty infiltration of liver. This change was unaffected by various diets used. However, Best *et al.* (1949) could show that large doses of choline can prevent the fatty change and prehepatic fibrosis induced by 15% ethyl alcohol given as drinking water to rats for 7 months. Klatskin and Krehl (1954) got

similar results. We administered 1.2 ml of 25% ethyl alcohol group sacrificed after 4 months did show early fibrosis and suggestion of lobulation. These changes were totally absent in the rat livers from alcohol with Liv.52 group. From the results of Hexobarbitone induced sleeping time (Table 2) it appears that Liv.52 has significantly curtailed the prolongation of sleeping time observed in the alcohol group.

Kutob (1962) has shown that toxicity of ethanol can be increased when chloroform is administered subcutaneously. Attempt to produce significant liver damage with combined use of alcohol and chloroform is in progress.

SUMMARY

Liv.52, an indigenous proprietary medicine has beneficial effect on growth. Chronic alcohol administration for four months produced marked hepatic damage in rats, which could be protected by Liv.52.

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REFERENCES

- 1. Ashworth, C.T.: Production of fatty infiltration of liver in rats in spite of adequate diet. Proc. Soc. Exp. Biol. Med. 1947, 66, 382-385.
- 2. Best, C.H., W.S. Hartroft, C.C. Lucas and J.H. Ridout: Liver damage produced by feeding alcohol or sugar and its prevention by choline. Brit. Med. J. 1949, II, 1001-1006.
- 3. Joglekar, G.V., G.K. Chitale and J.H. Balwani: Protection by indigenous drugs against hepatotoxic effects of carbon tetrachloride in mice. Acta Pharmacol. et Toxicol. 1963, 20, 73-79.
- 4. Karandikar, S.V., G.V. Joglekar, G.K. Chitale and J.H. Balwani: Protection by indigenous drugs against hepatotoxic effects of carbon tetrachloride a long-term study. Acta Pharmacol. et Toxicol. 1963, 20, 274-280.
- 5. Klatskin, G.: Effects of alcohol on the liver. J.A.M.A. 1950, 170, 1671-76.
- 6. Klatskin, G., W.A. Krehl and H.O. Conn.: The effect of alcohol on the choline requirement. J. Exp. Med. 1954, 100, 605-614.
- 7. Kopera, J. and A.K. Armitage: Comparison of some pharmacological properties of chlorpromazine, promethazine and pethidine, Brit. J. Pharmacol, 1954, 9, 392-401.
- 8. Kutob, S.D. and G.L. Plaa: The effect of acute ethanol intoxication on chloroform-induced liver damage. J. Pharmacol. Exp. Ther. 1962, 135, 245-251.
- 9. Patrao, S.: Observations on Liv.52. Journal of Indian Medical Profession, 1957, 4, 1878-1879.

- 10. Roe, J.H. and C.A. Kuether: The determination of ascorbic acid in whole blood and urine through the 2-4 dinitrophenylhydrazine derivative of dehydroascorbic acid. J. Biol. Chem. 1943, 147, 399-407.
- 11. Sheth, S.C., B.J. Northover, N.S., Tibrewala, U.R., Warerkar and V.S. Karande: Therapy of cirrhosis of liver and liver damage with indigenous drugs experimental and clinical studies. Indian J. Paediat. 1960, 27, 204-211.