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TOXICITY OF TEFLON DISPERSING AGENTS

A brief summary of our toxicity work on AHT and other Tefloo dispersing agents with emphasis on liver enlargement which seems to be the mest sensitive sign of toxicity is given below. The detailed reports of work completed to date will be available within a few days.

ART - (Ammonium 3,6 dioxa 2,5 di(trifluoro methyl undecafluoronousmosts)

The oral ALD for tate was found to be 60 mg/kg. Survivors showed definite liver enlargement in doses down to 1.5 mg/kg and with possible changes at 0.45 and 0.13 mg/kg. Single doses of 12 mg/kg produced liver enlargement which tended to increase during the two months following the dose. One one-hundradthof the lethal dose or 0.6 mg/kg given daily 5 times a week for 2 weeks produced enlargement which was significant in those rats killed on the day of final treatment and in those killed 14 days later. Mistological examination of the livers indicated that the colorgement was due to increase in cell size rather than an increase in the number of calls.

The lethal dose by okin absorption in rabbits was 130 mg/kg. Although the changes in liver weight in these rabbits are wore difficult to evaluate, there was a tendency toward enlargement and similar signs of liver injury.

A 25% aqueous solution in contact with the eye caused damage which persisted through 8 days. Washing with water 20 seconds after instillation provented permanent damage. Ten and twenty-five persont solutions were also irritating to guinea pig skin but did not cause skin sensitizetion,

CH-APFC - (Ammunium perfluorocaprylate)

JAZ

The oral ALD for rate was 670 mg/kg. Liver enlargement was definite down to a dose of 200 mg/kg with possible early signs down to 1.5 mg/kg.

Co-AFC - (Ameonium W-hydrohexa decafluoronouanoare)

The oral ALD was 1500 mg/kg. Survivors showed enlargement which appears evident in doses as low as 12 mg/kg.

"Teflon" Feeding Tests with "Teflon" 7, "Teflon" 6 made with Cg-AFFC, "Teflon" 6Cmade with Cg and "Teflon" 6C made with AHT.

The compounds were fed at a level of 25% in the diet of rate for 3 weeks. Rate were sucrificed 2, 3 and 5 weeks after feeding of test materials started.

Livers of rats sacrificed after two and three weeks of continuous feeding showed slight enlargement only in the group fad "Teflon" 5C with ART. After a two-week rest period the remaining rats were killed and those fed "Teflon" 5C with ART and "Taflon" Cg AFFC showed liver weights significantly different from the controls and those fed "Teflon" 7. The values of those fed "Teflon" 6C with Cg-AFC fell midway between the controls and the others. Although the numbers of animals used were small and the time offseding relatively short, the trend observed confirms the carlier liver enlargement observed in rats fed 25% "Teflon" 6 remin in the diet for 90 days (H. Report No. 49-60). A direct comparison among these compounds is difficult to make in these feeding tests because we do not know the concentrations of the fluoro acid dispersing agents present.

Cooclusions:

ART is a very toxic compound. Not only does it have a low lethal does but a single dose of 1/5 the lathal dose produced liver enlargement which increased with time. And 1/100 of the lethal dose fed 10 times produced definite liver enlargement. In addition, it was easily absorbed through the okin and produced liver damage in a second species. When 'Teflon' containing less than 5 ppm ART was fed to rate, it still produced enlargement which was apparent after 2 weeks.

The C, and C, aside have much lower acute toxicity, but they too have the ability to increase the size of the liver of rate at low doses. These short experiments may indicate differences in rate of development rather than qualitative differences but completion of microscopic examination of mimals in the current series as well as dosing of greater numbers of rate at the critical levels and holding them for longer periods would be asseded to establish the lowest effect level for each compound.

It is recommended that all of these materials, copecially AHT, be bounded with extreme care. Contact with the skin should be strictly avoided. Tasts on a third species, e.g. dogs, should be carried out where changes in liver function could be studied over a long period of time. The results of such tests might also throw some light on any possible species differences in \$AZ musceptibility.

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