PROTOCOL FOR A 63197 (R50796) + PQ DOG STUDY

1. Objective

Assessment of the efficiency of 63197 as an emetic agent when in combination with PQ and the resultant change in simple toxicity.

2. Proposal

a) Selection of stable non-vomiting dogs by giving the carrier (complan) by oral gavage. (Estimated 22 dogs out of 25 stable).

b) Confirmation that selected dose of 63197 will induce vomiting when given to dogs orally in complan. 2 dogs

c) Establish the LD75/100 for PQ in complan in the absence of any vomiting. 2-4 dogs

d) Administer a 63197/PQ/complan mix to show that:

1. the emetic action of 63197 is obtained in the presence of PQ

2. sufficient PQ is removed to protect the dog (collect vomit and measure the PQ content; monitor blood levels of PQ) 10 dogs and repeat of (c) for positive control. 6-8 dogs

COST

1. 25 dogs
2. Husbandry etc for 25 dogs for (say) 4 weeks
3. Preparation of mixtures. 2.5 hrs for RA
4. Dose equivalent of about 47 dogs at 10 minutes per dog, ie 1 day RA and 1 day LA
5. Observation, taking notes and any necessary action - about 10 hours for RA £3,000
6. Blood samples etc and their analysis, if agreed, to be costed.

Monkey study being undertaken at Huntington.

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