

would tell us the normal 4-hour urinary excretion of cyclic AMP in control subjects and if they could compare this with data collected from sufficient patients who exhibit a rapid change in mood to make a sample large enough from which to draw a statistically valid conclusion.

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EMOTIONAL ILLNESS IN PSYCHIATRIC TRAINEES

DEAR SIR,

Dr. Waring's study (*Journal*, July, 125, 10-11), contains, I believe, two serious methodological problems which may largely invalidate his findings. First, the General Health Questionnaire, as the author correctly states, is a '... reliable, valid and sensitive screening schedule for detecting emotional illness in general practice'. The same cannot, unfortunately, be assumed for the population to which it was applied. Secondly, as I am informed by Dr. Waring, the covering letter to those doctors in the control group began as follows:

'Dear Doctor:

I have been carrying out a study at the Institute of Psychiatry on attitudes, personality features and emotional factors in doctors training in psychiatry. Over 86 per cent of the doctors in the survey at the Institute have kindly responded. I am in need of a control group of doctors in training, but not training in psychiatry, etc.' Thus these respondents knew they were to be a control group and further that the study concerned assessment of their colleagues in psychiatry. One can only speculate on how this may have biased the outcome.

I believe we must consider the questions posed by Dr. Waring as still unanswered.

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COMBINED ANTIDEPRESSANT MEDICATION

DEAR SIR,

There is still much controversy concerning the relative benefits and potential hazards of using

combined antidepressant medication, namely the concurrent administration of a tricyclic antidepressant and a monoamine oxidase inhibitor (MAOI) (1). Winston (2), Schuckit *et al.* (3) and Sethna (4) have put forward the case for combined antidepressant treatment in refractory cases that do not respond to one or other treatment alone, and psychiatrists at St. Thomas' Hospital are well known for their views that many more depressed patients could be successfully treated if doctors were less fearful of the hazards of combined antidepressant medication.

Successful results were obtained by Winston (2) using 25 to 100 mg. amitriptyline daily combined with isocarboxazid, 10 to 20 mg. daily, whereas Sethna (4) used 50 to 75 mg. amitriptyline daily with phenelzine 15 mg. three times a day. Further, Pollitt (5) reports good results using only small daily doses of amitriptyline (50 mg.) combined with isocarboxazid or phenelzine. If combined antidepressant medication is more effective than either antidepressant alone it is of importance to discover the pharmacological basis for this apparent synergism.

The effect of many drugs is said to be potentiated by the concurrent administration of MAO inhibitors (including tricyclic antidepressants), narcotics, barbiturates, tranquilizers, anaesthetics and alcohol (1, 6). Because of their 'enzyme-inhibiting' properties, the MAOI drugs have been reported by some workers to interfere with the hepatic microsomal enzyme system which is responsible for the metabolism of many of the above mentioned drugs, including the tricyclic antidepressants (6-10).

We wish to report our observations of the effect of MAOI (isocarboxazid) administration upon plasma concentrations of a tricyclic antidepressant (amitriptyline). Eight patients from an out-patient clinic were selected for the study; six were receiving 50 mg. amitriptyline at night and two 25 mg. nightly. One patient was also receiving diazepam, but the dose was not changed throughout the duration of the study. Isocarboxazid in daily divided doses of 15 to 20 mg. was given to or withdrawn from patients who were already receiving amitriptyline. Heparinized venous blood samples were obtained from each patient before, during, and/or after at least one month of combined tricyclic/MAOI administration, at the same time of day (3.00 p.m.) on each occasion. Plasma samples were analysed for amitriptyline and its major active metabolite, nortriptyline, using a gas-chromatographic technique (11) which has a lower limit of sensitivity for both drugs of approximately 20 ng./ml.

Results in all eight patients showed that plasma levels of both amitriptyline and nortriptyline did not exceed 30 ng./ml. before, after or during combined antidepressant administration. Even though it was

difficult to be exact in quantitating plasma levels below 30 ng./ml., close inspection of the results obtained indicated that in no case had MAOI administration produced a significant increase in amitriptyline or nortriptyline plasma concentrations.

It would appear from these preliminary results that the synergic effect obtained with combined antidepressant medication is due to a potentiation of effect at central noradrenergic and serotonergic neurones, rather than through any inhibition of tricyclic drug metabolism.

In this study we did not try to assess changes in the psychiatric state of patients. It would be interesting to repeat the study using either larger doses of tricyclic drug or more sensitive analytical methods, and to correlate tricyclic antidepressant plasma levels with clinical response. This matter is now under consideration. We wish to thank Dr. Pollitt for his help in this study.

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DEPENDENCY AND OVERPROTECTION IN BOYS WHO STEAL

DEAR SIR,

We would like to report the results of giving the Self Administered Dependency Questionnaire (SADQ) to the mothers of 42 boys coming to a child guidance clinic in Yorkshire with a main complaint of stealing. Thirty-eight other boys referred for psychiatric treatment and thirty-four boys referred for psychological testing because of educational problems were used for purposes of comparison. The questionnaire has been described previously (Berg, 1974). The SADQ was employed by one of us (H.S.) in this connection as part of a larger research project. The three groups of boys did not differ significantly on age (combined total mean age 9.5 years), social class distribution, IQ (combined total mean full scale IQ on the WISC 95.2) or RQ (combined total mean on the Schonell 78).

The group of stealers had a significantly lower mean actual communication score (2.5) than either the psychiatric (4.2) or the educational (5.3) controls. Previously 64 primary school children from the general population were found to have a mean of 6.0; S.D. = 3 (Berg, 1974). Actual affection, assistance and travel mean scores did not differ significantly. The four types of preference score (Berg and McGuire, 1974) compared across the three groups for three levels of actual score, using two-way analysis of variance, failed to show any significant differences.

The stealers were significantly antisocial in other ways and were considered to be suffering from conduct disorders. The SADQ findings suggest that conduct-disordered boys who steal may be unusually uncommunicative and their mothers unconcerned about this state of affairs.

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