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Early Clinical Experiences with Sulpiride

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Sulpiride is a neuroleptic, recently introduced into the UK, that appears to have interesting pharmacological properties. It has been available in France for over a decade and has been shown to be singularly free of such severe side-effects as tardive dyskinesia. Its mechanism of action has only recently come under systematic scrutiny; in small doses, it appears to block the pre-synaptic receptors, whereas at larger doses, it acts on the postsynaptic receptors. There should therefore be a differential clinical effect at different dose ranges. It has been suggested from experience in both France and Scandinavia that it has both anti-psychotic and anti-depressant properties. This action, coupled with the absence up to now of any reported case of tardive dyskinesia in association with it, makes sulpiride an important drug to study in clinical psychiatry.

Over the past year we have treated 45 patients with sulpiride; clinical records were carefully maintained so that the data could be looked at as objectively as an open study would allow. No ratings on scales were performed, except that the diagnosis of schizophrenia was cross-checked with DSM-III criteria and a global rating of the clinical state was recorded in the notes at frequent intervals. Patients were prescribed sulpiride only if clinical symptoms warranted its use, and in a significant number the drug was started because other neuroleptics had failed to produce the necessary clinical response.

Data on 37 patients (27 males and 10 females) are presented: three patients had been on this drug for too short a period to make any clinical judgement about its effect, there were four patients who had complicating organic illnesses, and one whose diagnosis was in doubt.

Eighteen patients were treated with sulpiride

alone, while the remaining nineteen received the drug in addition to other psychotropic medication. Of the latter, eight patients received sulpiride and conventional neuroleptics; the rest received a variety of other drugs including benzodiazepines, antidepressants, etc. No evidence of adverse interaction with other psychotropics was detected. The dose range varied from 400 mg to 2400 mg daily.

Of the sixteen patients in whom sulpiride was prescribed for negative symptoms ten showed significant improvement, in some cases quite dramatic and unexpected. Many of these also showed a global change for the better. There were twelve patients treated for affective symptoms and ten of these showed a positive response. The five patients with tardive dyskinesia showed improvement not only in the clinical state but also an alleviation of the abnormal movements. Results were less promising in those ten patients in whom sulpiride was used specifically for acute psychotic symptoms. Less than half improved, and in two it produced a state of confusion with catatonia-like features early after the start of therapy. These quickly disappeared after the drug was stopped. In three other patients also, lack of improvement or worsening of symptoms led to the drug being withdrawn.

These early experiences with sulpiride suggest that it is a potent aid to the management of psychotic states. We have found it to have some unique properties, which need to be explored further by carefully controlled studies, especially since the results suggest that it is useful in states where conventional neuroleptics are known to be relatively ineffective, eg, schizophrenic deficit states and in patients who have affective symptomatology. The feeling of increased awareness and drive, and the improvement in concentration noted by several patients during treatment with sulpiride after they had been transferred from conventional neuroleptics, could be an important advantage with respect to social functioning in the community, and may themselves be factors in the beneficial effect on negative and depressive symptoms. Its lack of association with tardive dyskinesia makes sulpiride particularly suitable for managing patients who have developed the condition during prolonged use of conventional neuroleptics; this latter group of patients are also particularly likely to have pronounced negative symptoms. Our colleagues and I are proceeding to examine the role of this drug by controlled trials.

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