Letters to the Editor: May neuroleptics maintain clozapine-induced gains?

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There is strong evidence that clozapine is superior to conventional neuroleptics in the management of treatment-resistant schizophrenia [1], [2]. However, clozapine has a broad spectrum of adverse effects, and additionally places the patient at risk for the development of a life-threatening disorder, agranulocytosis. While the risk is maximum during the first 4 months of therapy, about 25 per cent of the recorded cases of granulocytopenia have occurred at later time points. A patient continues to be at risk for clozapine-induced granulocytopenia and agranulocytosis even years after the initiation of treatment [2].

An issue that may therefore be considered is: once a patient shows clinical response to clozapine, is it a viable therapeutic strategy to replace clozapine with a conventional neuroleptic? Expressed otherwise, can the clinical gains with clozapine be maintained with conventional maintenance therapy?

This is an unresearched area, but there is a parallel of sorts both for and against the idea that successful therapeutic agent may be replaced by a different agent during the maintenance phase of treatment. On the one hand, once peak response is obtained, electroconvulsive therapy is usually discontinued and replaced by an appropriate drug regimen; the results are, in general, satisfactory. On the other hand, after the completion of a successful course of electroconvulsive therapy, maintenance therapy with antidepressant drugs may fail if such drug therapy had been unsuccessful during the acute phase of illness [3].

Considering that clozapine is most commonly prescribed for the treatment-resistant schizophrenic patient, the latter situation is a closer parallel. However, only a controlled trial of continuation therapy comparing clozapine with a conventional neuroleptic will resolve the issue. Since the findings will be of importance to decision-making in schizophrenic patients successfully treated with clozapine, such a trial is urgently required.

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