

group, has received a combination of antipsychotics with anxiolytics (benzodiazepine group). Control group was treated only with classic antipsychotics (fluphenazine, haloperidol). Testing of psychotic symptoms was realised with the Brief Psychiatric Rating Scale (BPRS) and anxiety was measured by Spielberger's State-Trait Anxiety Index (STAI). Obtained data were pondered and presented numerically from 1.00 (without anxiety) to 3.00 (very high anxiety). Psychotic behavior was also presented numerically as psychotic index.

In both groups anxiety was at level of 2.96 index points before treatment (very high anxiety). In control group there was no significant improvement (2.38 at the end of examination) which indicates minimal improvement ($p = 0.10$). In experimental group which was treated with combine therapy (antipsychotics and benzodiazepines) there was significant improvement of anxiety, specially psychotic one ($r = 0.059$). This fact indicates very low anxiolytic potential of phenothiazines. Reduction of psychotic features in sense of partial remission of sch phenomenons is significant in both groups (from 178.4 to 46.3 index points, $p = 0.05$).

Hence anxiety in main dynamic force which has great influence at beginning and a development of sch process, polytherapy is necessary in treatment of these patients. It is consisted, in first place, of antipsychotics and anxiolytics, and very often antidepressive because of postschizophrenic depressive syndrome which occurs very often. In that way, it is possible to reach significant reduction of, not only psychotic anxiety but also free floating anxiety which always is present in pre psychotic and post psychotic period in most schizophrenic patients.

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WEIGHT GAIN ASSOCIATED WITH CONVENTIONAL AND NEWER ANTIPSYCHOTICS: A META-ANALYSIS

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A comprehensive literature search of English and non-English articles identified 78 studies which included data on weight change in more than one patient treated with a specific antipsychotic. For each agent, a meta-analysis estimated the effects of 10 weeks of treatment on body weight. The degree of weight change at 10 weeks was estimated by random effects regression. Except for molindone, antipsychotic treatment was associated with weight gain. Placebo was associated with a mean reduction of 1.68 kg. Among conventional agents, mean weight change ranged from a reduction of 0.41 kg with molindone to an increase of 3.25 kg with thioridazine, with an intermediate effect observed for haloperidol, an increase of 1.06 kg. Among newer antipsychotics, mean increases were as follows: clozapine 4.46 kg, olanzapine 4.15 kg, sertindole 2.92 kg, risperidone 2.10 kg and ziprasidone 0.87 kg. Insufficient data were available to evaluate quetiapine. Pairwise tests of differences between newer antipsychotics showed no significant difference between olanzapine and clozapine. Weight gain was significantly lower with ziprasidone compared with clozapine, olanzapine, risperidone and sertindole. There were also significant differences between clozapine and risperidone and olanzapine vs risperidone and vs sertindole. Both conventional and newer antipsychotics are associated with weight gain. Among newer agents, clozapine appears to have the greatest potential to induce weight gain and ziprasidone has the least. The differences among newer agents may impact upon the choice of treatment for some patients. Not only is weight gain undesirable because of associated health

risks, but it may also cause non-compliance with antipsychotic therapy which predisposes patients to relapse.

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THE EFFECT OF TREATMENT WITH TYPICAL AND ATYPICAL NEUROLEPTIC DRUGS ON NEUROPSYCHOLOGICAL MEASUREMENTS IN PATIENTS WITH SCHIZOPHRENIA

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Neuropsychological measures (frontal lobe tests, conjugate lateral eye movements) were performed in sixteen patients with paranoid schizophrenia 1) during acute episode, before starting pharmacological treatment (PANSS 126 ± 24) and 2) during improvement, on maintenance dose of neuroleptic drugs (PANSS 62 ± 24). Eight patients were treated with typical neuroleptic (chlorpromazine, levomepromazine, fluphenazine, perphenazine) and eight - with atypical one (clozapine, olanzapine, ziprasidone).

In whole group, after alleviation of psychotic symptoms, the results of frontal lobe tests improved, and in case of Stroop test A and B, significantly so. There were no differences between typical and atypical neuroleptic drug as to the magnitude of such improvement.

During acute episode, schizophrenic patients exhibited excessive activation of left hemisphere in response to emotional and spatial questions (i.e. directed to right hemisphere), measured by CLEM method. During improvement, an increase of activation of right hemisphere was observed in response to these questions, at the expense of left hemisphere activation. Such increase, reflecting a regulatory action on hemispheric activation was significantly greater in patients treated with atypical neuroleptic drugs.

The results obtained suggest a possibility of improvement of some cognitive functions as well as regulation of activation asymmetry in schizophrenic patients with neuroleptic treatment, the latter may be more marked with atypical than typical neuroleptics.

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ZIPRASIDONE: *IN VIVO* EVIDENCE OF CENTRAL 5HT_{1A} AGONIST ACTIVITY

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Ziprasidone is a novel antipsychotic with a unique specificity for dopaminergic and serotonergic receptors. *In vitro* studies of cAMP accumulation indicate that ziprasidone is an agonist at 5HT_{1A} receptors. Since 5HT_{1A} receptor agonism is thought to contribute to reduced extrapyramidal side-effect (EPS) liability and enhanced efficacy against both negative and affective symptoms of schizophrenia, we investigated the *in vivo* 5HT_{1A} agonist activity of ziprasidone by measuring its effects on dorsal raphe cell firing and cortical dopamine release in the rat. Ziprasidone inhibited dorsal raphe firing with an ED₅₀ of 300 µg/kg intravenous (IV) and its inhibitory effect was blocked by pre-treatment with the 5HT_{1A} antagonist WAY-100635 (10 µg/kg IV). Although the 5HT_{2/D₂} antagonist olanzapine also slowed unit activity (ED₅₀ = 1000 µg/kg IV), this effect was not attenuated by WAY-100635, but was reversed by pre-treatment with the norepinephrine re-uptake inhibitor, desipramine (5 µg/kg IV). This is consistent with olanzapine's α₁ antagonist activity, low affinity for 5HT_{1A} receptors, and