**Cumulative effect of psychotherapy in remission of symptomatology of major depressive disorder**

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**ABSTRACT**

**Background:** Major depressive disorder constitutes the most commonly appeared psychological disorder. The aim of this study was to investigate the cumulative effect of psychotherapy over drug therapy in remitting depressive symptoms was examined.

**Method-materials:** The sample consisted of 62 patients given the diagnosis of major depressive disorder who suffered in the past two years with at least 2 major depressive episodes. Additional inclusion criteria were, age of patients to be between 26-60 years old, onset of disorder ≥4 years, and weight of symptomatology having scores in MADRS ≥34 and in CGI ≥5. All patients were assigned randomly in four groups of combined therapy. The progress of depressive symptoms was measured with taxonomic manuals of psychological diseases MADRS from 1rst week of hospitalization, and every 3 weeks, until the 24th week of treatment.

**Results:** Of the 42 participants, 24 were male and 38 female with mean age 42.3±8.8 years. The statistical significant improvement on depressive symptoms (p<0.001) occurred from induction week (MADRS=41) to the 6th week (MADRS=32) of hospitalization defined the level of remission of depressive symptoms. This remission mostly was a mixed effect of drug therapy and psychotherapy, since all 4 groups of patients showed similar decline in MADRS scores. Maintenance phase occurred about the 3rd month of treatment (MADRS=14), at which point the greater cumulative effect over pharmacotherapy was that of psychodynamic-analytical therapy at 6th month with 52% remission (p<0.001) from baseline, while that of cognitive-behaviorial therapy was 40%, and respectively only 22% for the supportive therapy.

**Conclusions:** Remission rate of depressive symptoms occurred about the 6th week of hospitalization, with maintenance phase to occur at 12th week of treatment, time point at which psychodynamic and cognitive-behaviorial psychotherapy had a statistical and clinical greater effect over drug therapy on remission maintenance of depressive symptoms compared to supportive therapy.

**Keywords:** Major depressive disorder, psychotherapy, depression rating scale MADRS.
INTRODUCTION

Major depressive disorder (MDD) constitutes the most commonly appeared psychological disorder. Epidemiological studies demonstrate that it evolves to a chronic disease, with high rate of mortality. It is estimated that the 30% of patients with major depression continue to be depressive at the following year, the 18% for the next two years, while the 12% for the 5 following years. The 15% of patients appears remaining symptoms which constitute indications, although they lead to a bad result, frequent relapses, attempts of suicide and decreased psychosocial functionality, by increasing simultaneously the cost of nursing. The 50% of patients with major depressive disorder appears relapse within the next year, while the 30% is leaded to remaining elements, frequent relapses, mortality and psychosocial reduce of functionality and after the 5 years of the first nursing.

Most of the people that suffer from reactionary type of depressive disorders can be confronted as external patients, but those who suffer from MDD, according to the classifying manuals of mental diseases DSM-IV and ICD-10, it has to be nursed when there is high suicidal ideation and productive ideas with or without psychotic elements, whether when there is depressive co-morbidity with other organic or pathological situations, for which is needed follow-up at a hospital, or at cases that control of antidepressant cure or antipsychotic treatment has to take place and to begin a selected type of psychotherapy for the patient.

At detailed psychodynamic approach aim of the therapist is the guidance of patient in order to recognize and discover these recitals which were created inside him and the feelings of disappointment, disadvantage, guilt or unworthiness, anger or pain, which cause him the depression. Another aim is the revealing and the expression of negative feelings of pain and anger that remain entrapped in psychologically. In alternative, a new psychotherapeutic view sorts the depression as symptom and the anger or
pain as a deeper disorder. The therapist, working with patient’s pain or anger under the “transfer-antitransition”, tries with remedial process to placate the anxiety, by remove those defense mechanisms, which shelter the pain or anger like a vicious system, which restores the negative thoughts and the depressing feeling, essentially to replace gradually the patient’s former defense mechanisms with other more healthy mechanisms. The beginning of psychotherapy sessions does not recant the granting of antidepressants or antipsychotic substances, which are essential for the acute of depression to be encountered.

At cognitive-behavioral approach aim of the patient is to learn how to recognize and alter the dysfunctional automatic thoughts, the cognitive malformations and the dysadaptive cognitive shapes.

In order to achieve this, there are two center technical categories that aim the first one at changes of behavior, while the second one at alteration of cognitive process at patient’s thoughts.

The therapist begins with the interview by collecting relative information of depressive person’s childhood’s experiences, which contributed at the development and maintenance of nuclear conviction that is now problematic and brings misrepresentations in relation with himself. The modification of the above cognitive figures takes places by a lot of techniques. For example, by the technique of “understanding of trait meaning” the therapist tries by questions to the depressive patient to understand the reasons that conducted him to depression. By the technique of self-guided disclosure tries to guide the patient, after understanding his problem, to develop ideas, behaviors and ways to encounter the problems that are related with depression. By the technique of “chances control” the therapist helps the depressive patient to see how possible is for the worst that he fears to happen. Other noted techniques that are used is, the technique of “avoiding the use of absolute expressions” and the technique of “direct doubt”, that is being applied at suicidal patients, with the therapist to opposite directly patient’s views on suicide or the end of life.

At the terms of six months supportive therapy of this investigation, the application of supportive intervention of the therapeutic team, consistent by clinical psychologist, psychiatrist, nurses, social worker and occupational therapist, was to localize through the mental structures of the depressive patient the pain and the anger that stay...
trapped inside him and to help him express them, emerge the mental and psychological reasons that hide behind the depressive symptoms, to mental-educate and support the depressive patients, by cognitive techniques or expressive type techniques.\textsuperscript{15,17} Moreover, localizes the anger and pain of patient and connects those feelings at conscious level, with “reference persons” or past or present situations which exist in subconscious level and were important for the birth of depressive structure of patient. Only at this time, the important steps for the removal and remission of depressive symptoms take place.

Aim of the study was to explore the cumulative effect of psychotherapy over pharmaceutical therapy and the recession of symptoms of patients hospitalized with MDD. It was searched the period of recession of symptomatology and the holding time of recession up to 6 months of surveillance after introductory hospitalization.

Methodology

The sample constituted 62 patients with average age 42.3\textpm{}8.9 years (24 males and 38 females), with at least 2 depressive episodes and corresponding hospitalizations the last 2 years and presence of symptoms of MDD, according the taxonomic manuals of mental illness DSM-IV and ICD-10 and they are analyzed at the following clinical signs and symptoms: continuous depression, lack of energy and burnout, reduced appetite and mood for sleep, isolation trends, social withdrawal, anxiety, concerns and indecision, disappointment and indifference about life, people and future, lack of interests, decrease of sexual energy, feelings of guilt, unworthiness, bad mood at the biggest part of day and almost every day, decrease of pleasure, medium or high suicidal ideation and suicidal danger.\textsuperscript{6-8,11,18}

The symptoms have to stay for at least 6 months. At some patients depressive feeling was accompanied by psychotic elements, tingling, and ideas of self accusation or destruction of “Ego”. Patients’ admission requirements at the sample of this study, except those which defines the DSM-IV, was the age space from 26 to 60 years, the time that disease started >=4 years and the gravity of symptoms with prices at scales of depression of Montgomery Asberg MADRS>=34 and the clinical global assessment, CGI>=5. We recorded the progress of symptomatology from the first week of introduction and every 3 weeks until the 24\textsuperscript{th} week of hospitalization. All the patients
pharmaceutical therapy, 18 of them were subjected to combined therapy of psychodynamic detailed psychotherapy (PDP) and pharmaceutical therapy, 12 to combined therapy with cognitive behavioral psychotherapy (CBP) and pharmaceutical therapy, 16 at combined therapy of simple supportive and pharmaceutical therapy, while the other 16 were objected only to pharmaceutical therapy.

Patients were in random classified in 4 groups of combined therapy (specifically 3 groups of combined psychotherapy with pharmaceutical treatment and 1 group of only-therapy with pharmaceutical therapy), after primary being equated in age and in depressive’s symptomatology gravity. The progress of their depressive symptomatology was recorded by use of the scale of depression of Montgomery and Asberg from the introduction week and every third week, until the 6th month of therapy. Patients got antidepressants (venlafaxine, fluoxetine, fluvoxamine, citalopram, paroxetine), antipsychotic (risperidone, aripiprazole) and benzodiazepines.

The study was double-blind, namely the patients were in random classified in these 4 therapy groups without knowing about the type of therapy that they got, while in the other hand therapists didn’t know about the recording of therapeutical progress and the progress of patient’s depressive symptoms, while they were evaluated from independent judge-therapist.

The sessions of detailed psychodynamic therapy were hourly with frequency 3 times a week and during all the semester of patients’ surveillance. The sessions of cognitive-behavioral were hourly, they were taking place once a week and their total number was 20 sessions. The sessions of detailed psychodynamic and cognitive-behavioral were conducted by clinician psychologist and psychiatrist. The supportive was taking place 2 or 3 times a week for the semester that the investigation lasted, the intervention was of half an hour till three quarters of hour, and was conducted from nurses and psychologist. The patients were informed that they were going to participate in an research study, but their personal data wouldn’t be published but only the total data of the progress of all patients that take part to the study, by maintaining the medical and psychological confidentiality.

**Results**

The first sign of recession is being observed to appear already from the 6th or 7th week of hospitalization, with the
reduce of depressive symptomatology of patients to be recorded at the total prices of MARDS from 41 at the introduction week, to MADRS=39 the 3rd week and MARDS=32 the 6th week of hospitalization, regardless therapy, as it appears in figure 1 [F(1,30)=12.68,p<0.01]. The statistically significant improvement of depressive symptomatology of patients already from the 6th week of treatment (MARDS=32, p<0.01) is being signed as the first level of depressive symptoms’ recession, with rate of improvement 22% from the levels according to the introduction week (MARDS=41)(figures 1 and 2). The conservation level of the improvement is observed to begin about the 12th week of hospitalization (MADRS=14)(figure 1), with rate of improvement 66% form the levels according to the introduction week (MARDS=41)(figures 1 and 2) and composes the conservation beginning of improvement, which maintains since the sixth month of observation (24th week). The size of effect of the improvement rate from the first level of recession of the 6th week till the conservation level of the 3rd month (12th week) was found to be 44%. The total rate of improvement at MARDS prices of depressive symptomatology, from the levels according to the introduction week since the 6th month of observation appears at figure 2 for the total sample of patients, independently of the kind of treatment.

With more details, for the effect of the sorts of treatment at the reduce of depressive symptoms it was found that the sorts of treatment that affected statistically more at the conservation of recession of MDD were the following, as they resulted by the statistical analysis with multiple regression analysis (multiple regression analysis, parameter estimates) [F (3,123)=25.34, p,0.001]: the psychodynamic–pharmaceutical therapy combination with 71% of improvement from prices’ levels according to MARDS (p<0.001), the combination of cognitive-pharmaceutical therapy with 63%, while less improvement appeared either the combination of supportive-pharmaceutical therapy (41.5%) or the pharmaceutical therapy alone (39%), as well as the rate of recession started to appear from the 5th month of therapy, when the therapy included only medicines in combination of support with medicines (figure 3). It is observed that the combined therapy either psychodynamic-pharmaceutical therapy, or cognitive-pharmaceutical therapy lasted for more time, till the 6th month, the conservation period of recession of patients’ symptoms with major depression at acute phase. The combined
therapy of psychodynamic-pharmaceutical therapy maintained the depressive symptomatology’s conservation improvement at 32% over the levels of pharmaceutical therapy’s effect, while the combination of cognitive-pharmaceutical therapy maintained the recession at 42% over the activity levels of pharmaceutical therapy. The cumulative rate of effect of the three types of psychotherapy (psychodynamic, cognitive, supportive) over pharmaceutical therapy, at the improvement of depression symptoms from the base’s levels of introduction week, every week of following until the 24\textsuperscript{th} week, appears at figure 4.

Specifically, for the PDP the recession level is installed about 6\textsuperscript{th} week of therapy with percentage of cumulative effect 12,5\% over pharmaceutical therapy, while for CBP the recession level is observed three weeks later, i.e. the 9\textsuperscript{th} week with 16\% of improvement.

The conservation phase of the improvement of depressive symptoms, concerning the PDP happens at the 12\textsuperscript{th} week, with 16, 7\% rate of reduction of the symptoms and increases with rates 32\%, 48\% and 52\% each at the 16\textsuperscript{th}, 20\textsuperscript{th} and 24\textsuperscript{th} week of hospitalization and therapeutic monitoring. Regarding the CBP, the conservation of the improvement of depressive symptoms appears three weeks later of psychodynamic approach, at the 12\textsuperscript{th} week of therapy, i.e. with 16\% cumulative effect over pharmaceutical therapy by increasing to 32\%, 39\% and 40\% each, at the 36\textsuperscript{th}, 20\textsuperscript{th} and 24\textsuperscript{th} week of hospitalization (figure 4). In contrast the supportive therapy shows a slightly and rather transient cumulative effect over the conservation of the improvement of depressive symptoms, since it is observed at the 16\textsuperscript{th} and 20\textsuperscript{th} week of hospitalization a transient change over the supportive and against the pharmaceutical therapy as monotherapy, with rates 8\% and 22\% each, but in 3 weeks were disappeared.

**Discussion**

Recession at depressive symptoms from the introduction week was observed at the 6\textsuperscript{th} week of hospitalization of patients’ total independently of the therapeutic choice they had, by signing the first level of symptomatology’s recession of major depressive disorder (MDD), firstly due to the effect of pharmaceutical treatment, as well the 4 patient groups appeared similar decrease at MARDS prices. The beginning of the phase of conservation was observed about the 3\textsuperscript{rd} month of therapy, where was found the kind of therapies to effect statistically more at the recession.
The conservation of MDD with the greater improvement to take place at the 6th month, whether it concerned the combination of PDP-pharmaceutical therapy, or the combination of GBP-pharmaceutical therapy, while fewer effects presented the combination of supportive-pharmaceutical therapy or the pharmaceutical therapy alone. The pure clean cumulative effect of PDP, as recession phase of depressive disorders, takes place about the 6th week of therapy, while the conservation phase of recession is observed at the 9th week of therapy. For the CBP the consolidating of recession level comes three weeks later of PDP, while the conservation of recession shifts at the 12th week of therapy.

The data of this study come to agreement with the reports of Vittengl et al.19, who at a meta-analysis with sample of 1800 patients with MDD found that the GBP leaded to the prolongation of the recession period at 29%. Another meta-analysis showed that the combination of psychotherapy with pharmaceutical therapy was an effective intervention at patients with MDD in relationship with other therapies.20 At research studies at patients with childhood traumas with and without MDD, appeared that the patients with childhood traumas and MDD had a very good response to the combination of anti-depressive treatment and psychotherapy.21,22

At previous studies, appeared that at the phase of continuing therapy at patients who received GBP the period of recession was decreased at 12% in comparison with those who received only pharmaceutical treatment or supportive therapy.10,23 Study of Maina et al.,9 showed that the recession periods of MDD symptoms were bigger than 75% when short PDP in combination with pharmaceutical therapy was applied (paretoxine or siloprame), and no when applied only pharmaceutical therapy or combination of pharmaceutical and supportive therapy.

The PDP and CBP prolong the recession conservation of depressive symptoms, as they act at pro-conscious and unconscious level, touching the unsolved conflicts at the patient’s shelf psychological structures. At the therapeutic sessions the person comes to contact and faces the conflicts of pain and grieves, of anger and guilt. The therapist through the process of transition-antitransition restores corrigenda these conflicts by teach the patient to use secondary mechanisms of defense, more mature than the primary ones, that are involved in depression.
The feelings of despair, pain, anger, grieve, melancholy, indisposition and unpleasure exist on the patient’s face and his emotional non-verbal expressions of every depressive patient. The purpose of all therapists and nurses is to detect and decode at patient’s verbal and non-verbal expressions these feelings. The therapist needs to know for the “transition” of elements of anger or pain of the depressive patients to the face of nurses, psychiatrists or psychologists, and the reversal as well for the “anti-transition” of therapists’ feelings to the patients due to behaviors or patients’ verbal comments. The transition of pain or anger to the therapists or other person of the environment takes place from the depressive patient by the defense mechanism of “depressive identification of endobolis” to beloved persons from the past. It needs then in the appropriate time to be clear to the patient, that the unpleasant moments belong to the experiences, negative and positive, from the past, which in the therapeutic relationship we will try to emerge and release.

The continuous body care and supportive therapy is provided by the nurses to the depressive patient with aim to partially activate emotional reactions in his emotional poorness.24 As negative the patient is, transiently in time, opposite life of even by thinking suicide, the therapists and nurses remind him with empathy that depression is something that you are going through “at this phase of your life, and it's totally normal to feel bad, it is sure that in a while you are going to feel better”. When patient starts to get out of depressive phase then he is going to be grateful to the support that was given to him, and it will stay strong through the dynamic evolution of his full recovery. The cooperation of nursing and medical staff exists and at the case of suicidal danger patients, so as the proper meters to be applied for the removal of attempt danger, in combination with emotional support.24–26

Conclusion
The cumulative effect of psychotherapies, as the findings of this study show, lies to the decrease of depression’s relapse chance or at least to prolong for a longer period a possible disease’s relapse that would anyway take place in a period of 5 or 6 months from the prior hospitalization or depressive episode, if the patient had been subjected only in pharmaceutical therapy without any other psychotherapeutic intervention.

BIBLIOGRAPHY


Figure 1. Phase of recession and improvement for the total sample of patients ($N=62$), as it appears at the phase of recession and conservation of the whole of the sample, as it is represented at the moderate general price MARDS every week of therapy.

Figure 2. Total rate of improvement of depressive symptomatology, of the whole of the sample, at Mards prices from the base levels, independently of therapy.
Figure 3. Effect of the kind of therapies (psychodynamic, analytic and pharmaceutical confrontation, cognitive behavioral psychotherapy and pharmaceutical confrontation, pharmaceutical therapy, supportive therapy and pharmaceutical confrontation) at the improvement effect of depressive symptomatology from the introduction week till the 24th week of hospitalization.
Figure 4. Cumulative rate of effect of psychotherapy species (psychodynamic analytical psychotherapy and pharmaceutical confrontation, cognitive behavioral psychotherapy and pharmaceutical confrontation, supportive therapy and pharmaceutical confrontation) over pharmaceutical therapy.