RESEARCH ARTICLE

Cognitive Behavioral Therapy in Breast Cancer Patients - a Feasibility Study of an 8 Week Intervention for Tumor Associated Fatigue Treatment

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Abstract

Background: Tumor associated fatigue (TAF) or cancer related fatigue (CRF) is not a new concept. Nonetheless, no real headway has been made in the quantitative analysis of its successful treatment via cognitive behavioral therapy. Since 20 to 30% of all breast cancer patients suffer from anxiety and/or depression within the first year of their diagnosis, this issue needs to be addressed and a standard treatment protocol has to be developed. This study focused on developing a simple, reproducible and short (8 weeks) protocol for the cognitive behavioral therapy support of tumor associated fatigue patients. Materials and Methods: Between the year 2011 and 2012, 23 breast cancer patients fulfilled the diagnosis TAF requirements and were introduced into this study. Our method focused on a psycho-oncological support group using a predetermined, highly structured and reproducible, cognitive behavioral therapy treatment manual. Eight weekly, 90 minute sessions were conducted and patients were evaluated before and after this eight session block. Tumor fatigue specific questionnaires such as the multidimensional fatigue inventory (MFI) as well as the hospital anxiety and depression scale (HADS) were used in order to quantitatively evaluate patient TAF. Results: Of the 23 patients enrolled in the study, only 7 patients fulfilled the TAF diagnostic criteria after the psycho-oncological group treatment. This represents a 70% reduction in diagnosable tumor associated fatigue. The HADS analysis showed a 33% reduction in patient anxiety as well as a 57% reduction in patient depression levels. The MFI scores showed a significant reduction in 4 of the 5 evaluate categories. With the exception of the “mental fatigue” MFI category all results were statistically significant. Conclusions: This study showed that a highly structured, cognitive behavioral therapy group intervention will produce significant improvements in breast cancer patient tumor associated fatigue levels after only 8 weeks.

Keywords: Breast cancer - cognitive behavioral therapy - tumor associated fatigue - cancer related fatigue

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Introduction

Tumor associated fatigue (TAF), also known as cancer related fatigue (CRF), emerges to be a largely neglected factor in the treatment of breast cancer patients. While a plethora of diagnostic developments and treatment options are under continuous evaluation, the main focus remains on the somatic aspects of this disease, i.e. meeting the oncological challenge by medication, chemotherapy, radiation treatment, surgical intervention and standard clinical care (Schumacher, 2003). Largely due to this aggressive somatic treatment of breast cancer, tumor associated fatigue is often diagnosed too late in breast cancer therapy (Montazeri et al., 2001; Fafouti et al., 2010; Schleife et al., 2012; Kovacs et al., 2013; Abu-Helalah et al., 2014). Nonetheless an ever-growing body of literature is becoming available for breast cancer patients suffering from TAF/ CRF. Although psycho-oncological treatment plays an ever bigger role in modern breast cancer treatment, the subgroup suffering from tumor associated fatigue syndrome is a more recent area of investigation (Larkey et al., 2014; Schmidt et al., 2014). TAF is ambiguously defined and includes patients presenting with avolition and lethargy, as well as a plethora of nonspecific symptoms (Mitchell, 2010; Li et al., 2014). The national comprehensive cancer network (NCNN) recommends two general approaches: medication based and non-medication based therapy (Montazeri et al., 2001). While the differentiation between a cancer associated depression and a tumor associated fatigue is fluent, the prior should be treated using pharmaceuticals. The latter on the other
hand, may be treated by a non-medication based therapy which includes an increase in physical activity, stress- and problem management training, and psycho-oncological therapy/intervention. While it is difficult to determine whether patients suffers from a depression or a TAF, due to the lack of a quick diagnostic tool, it is paramount to determine which is the main source of patient discomfort (Bjorneklett et al., 2012) as the therapy may vary depending on the diagnosis.

20 to 30% of all breast cancer patients suffer from anxiety and/or depression within the first year of their diagnosis. The depression generally subsides after approximately one year. Fear and anxiety of a possible recurrence however remain. Several studies showed that both depression and anxiety contribute to and are related to cancer associated fatigue. (Vodermaier and Millman, 2011; Cosco et al., 2012). Differentiating fatigue and depression is difficult to the fact that both are associated with lethargy, tiredness and exhaustion. In order to distinguish the two, a two-question test was established (Roth et al., 2010).

These questions are: i) Within the last month, did you feel an increasing sadness, hopelessness, and/or gloominess? ii) Within the last month, did you experience less enthusiasm and joy within your daily activities?

If both of these questions are answered with yes, the diagnosis of a depression is more likely. If only one question is answered with “yes” a TAF is possible. This preliminary TAF diagnosis may then be supported by world health organization (WHO) criteria catalogue, determining the severity of a clinically relevant cancer/tumor associated fatigue. The differentiation between the two is important due to the fact that a fatigue treated with antidepressants does not show any improvement while patients still experience all the side effects of the medication (Minton et al., 2008). The role of a pharmaceutical intervention in TAF patients is currently unclear. While psycho-stimulating substances are being evaluated in studies, no fundamental breakthrough has been made in this area. The main focus of treatment thus lies on increased physical activity and individual psycho-oncological support and nutrition improvement. All three have shown to lead to a significant improvement in anxiety and depression symptoms in the tumor associated fatigue patients group (Campos et al., 2011; Chan, 2011; Fitch, 2012; Finneghan-John et al., 2013).

The focus of this paper is therefore: i) Is it possible to positively influence anxiety and depression components of TAF breast cancer patients by, cognitive behavioral therapy group intervention with an 8 week intervention? ii) Is it possible to develop a practical, simple and clearly structured protocol for a psycho-oncological intervention in TAF patients?

Materials and Methods

Between 2011 and 2012, 24 patients were recruited into the study. The average age was 45 years, one patient discontinued the study due to personal reasons. The remaining 23 patients were either currently in adjuvant breast cancer therapy or had recently finished therapy. The longest time interval between the last oncological therapy session and the beginning of the psycho-oncological intervention was 24 months; anti-hormonal therapy excluded. Tumor associated fatigue was diagnosed using the criteria catalog for a tumor associated fatigue syndrome. The world health organization (WHO) suggested a criteria catalogue which allows for the evaluation of the severity of a clinically relevant cancer associated fatigue. These criteria are also used by the fatigue coalition USA (Curt et al., 2000; Cellà et al., 2001) and consist of 11 patient evaluated questions as well as three physician evaluated categories. If six of these fourteen criteria are met, the patient presents with a clinically relevant tumor associated fatigue. Naturally, all 23 of the patients fulfilled this requirement before the first therapy session. Quantitative evaluation of patient TAF status was performed before and after the group sessions were completed. Evaluation tools were the HADS and MFI questionnaire. A study timetable is provided in Figure 1.

Hospital Anxiety and Depression Scale (HADS)

The HADS is a self-evaluation tool, quantitatively analyzing patient anxiety and depression symptoms. It was administered to patients before and after 8 therapy sessions. Both, anxiety and depression symptoms are scored separately. Each of the seven questions is answered on a scale from 0 to 3, yielding a total of 21 possible points. 0-7 points is regarded as normal while 8-10 points is a moderate, 11-14 points a serious and 15-21 points a very serious symptom constellation (Zigmond and Snaith, 1983).

Figure 1. Schematic Representation of the Study Timetable

Figure 2. Schematic Representation of Group Sessions
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Multidimensional Fatigue Inventory (MFI)

The MFI is a 20-item quantitative, self-reported questionnaire designed to evaluate TAF. The following dimensions are evaluated: general fatigue, physical fatigue, mental fatigue, reduced motivation and reduced activity (Smets et al., 1995). Evaluating these questions in a manner that questions directed towards a negative answer were scored on a positive scale when answered with a low numerical value, allowed for a simple compilation of the cumulative data. In order to evaluate the degree of fatigue, an absolute numerical value was assigned to all answers. For example: strong fatigue (5), little fatigue (1). The MFI therefore allows a quantitave correlation of patient fatigue before and after our intervention.

Group Intervention: The basis of this psycho-oncological intervention is the improvement of patient quality of life. Pre-defined patient goals are: i) Improvement in self-management: patients learn to actively control fatigue symptoms ii) Development of realistic goals iii) Discover and use of available resources iv) Information on healthy nutrition v) Learning to deal with tumor associated stress

Each group performed a total of eight sessions, 90 minutes each, once a week over an 8 week period. Each session was led by an experienced psycho-oncologist and was highly structured based on a predetermined manual.

Each session begins with a 10 to 15 minute relaxation exercise, during which self-awareness and relaxation techniques were addressed and practiced. This portion was followed by an educative session regarding several different topics. This lecture was no longer than 15 minutes. A 15 minute open discussion focusing on the subject of each session followed. Before ending each session, a 5 minutes break was introduced giving each patient the option to focus their thoughts on this particular subject. Closing every meeting, each patient was asked to, very briefly, summarize their feelings regarding this subject. Closing every meeting, each patient was asked to, very briefly, summarize their feelings regarding this topic in two sentences or less. This highly structured lesson scheme was applied to every one of the eight sessions, allowing each patient to prepare themselves for the next session, a 5 minutes break was introduced giving each patient the option to focus their thoughts on this particular subject. Closing every meeting, each patient was asked to, very briefly, summarize their feelings regarding this topic in two sentences or less. This highly structured lesson scheme was applied to every one of the eight sessions, allowing each patient to prepare themselves for the next session.

Statistics

Statistical analysis was performed using the VassarStats® (Vassar College, Poughkeepsie, NY, USA) statistics program. Pearson’s chi-squared tests and t-tests were used in order to evaluate significances when appropriate. A p-value <0.05 was considered significant.

Informed Consent

Written informed consent was obtained from all patients for publication of this manuscript. A copy of the written consent is available for review by the Editor-in-Chief of this journal. This study was conducted in accordance with institutional review board standard operating procedures.

Results

Before beginning therapy, 23 patients fulfilled six or more diagnosis criteria for tumor associated fatigue, thus qualifying them for this trial. As mentioned above, 11 patient self-scored categories were analyzed. The analysis of the diagnostic criteria showed that only 7 of the 23 initial patients would qualify for this trial after 8 therapy sessions, meaning that only seven patients met the criteria for tumor associated fatigue after intervention. This 70% reduction is statistically significant p<0.05.

MFI: The evaluation of the MFI questionnaire may be divided into five subgroups. These are: general fatigue, physical fatigue, reduced motivation, reduced activity and mental fatigue. Table 1 shows the MFI scores for all five categories before and after treatment. The improvement is significant in all categories after 8 sessions, with the exception of the mental fatigue category. The general fatigue category initially scored with an average value of 16 before intervention, but could be reduced to a value of 13 afterwards. Physical fatigue scores were reduced from 15 to 12, reduced motivation was decreased from 11 to 9 and reduced activity was scored with 15 before and 12 after therapy. These average scores all differed significantly. The mental fatigue category score was reduced from 14 to 12, though significance was not established here. A p-value of 0.146 allows for carefully assuming of a trend towards improvement.

HADS: This questionnaire evaluated both an anxiety and depression aspect. The results pertaining to the anxiety aspect showed 18 patients to have a pathological score before the first intervention (i.e. a score above 7). A significant reduction could be shown to 12 patients after 8 sessions. This represents a 33% reduction in pathological anxiety scores. The average HADS score for anxiety was 12 before and 8 after intervention. These results are statistically significant. Average scores are shown in Table 2.

Prior to the first session, 14 of the 23 patients showed pathological results on the depression scale

<table>
<thead>
<tr>
<th>Table 1. Multidimensional Fatigue Inventory (MFI) Scores for 5 Categories before and After Psycho-Oncological Intervention</th>
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<tbody>
<tr>
<td><strong>Category</strong></td>
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<tr>
<td>General fatigue</td>
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<tr>
<td>Physical fatigue</td>
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<tr>
<td>Reduced motivation</td>
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<td>Reduced activity</td>
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<td>Mental fatigue</td>
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* A=Average; SD=standard deviation

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<tr>
<th>Table 2. Depression and Anxiety (HADS) Scores before and After Psycho-Oncological Intervention. 0-7 Points=Normal, 8-10 Points=Moderate, 11-14 Points=Serious, 15-21 Points=Very Serious Symptom Constellation</th>
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</thead>
<tbody>
<tr>
<td><strong>Category</strong></td>
</tr>
<tr>
<td>Anxiety</td>
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<tr>
<td>Depression</td>
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* A=average; SD=standard deviation
This trial by the use of the fatigue coalition diagnosis
fatigue, depression and anxiety before and after eight
anxiety and depression scale as well as multidimensional
after picture of the patients state of mind. The hospital
scoring systems were used in order to paint a before and
group intervention, two well-established and validated
as it may be to quantify results of a psycho-oncological
allowing a very structured and reproducible approach to
tumor associated fatigue. Thus, a manual was developed,
techniques were taught, practiced and developed in a
outpatient procedure. During these sessions, relaxation
is the shortest time period of cognitive behavioral therapy
regimen consisting of eight sessions, 90 minutes each. This
study revisited this topic by giving 23 patients the option to
able to also show a positive effect (Duijts et al., 2011). Our
psychological measures onto fatigue symptoms and was
increase physical activity, relaxation techniques, psycho-
are statistically significant.

Discussion

Tumor associated fatigue or cancer related fatigue is a
problematic and common occurrence in cancer patients.
At least 20 to 30% of all breast cancer patients report
fatigue symptoms. These symptoms include tiredness,
episodes, weariness and exhaustion and may
result in severe somatic consequences. TAF diagnostic
criteria were established in 1998 by the fatigue coalition,
but unfortunately tumor associated fatigue is not yet
classified within the ICD 10 system (Curt et al., 2000;
Cella et al., 2001). This may be one of the reasons
why tumor associated fatigue is often undiagnosed
and untreated. Several studies investigated the effect
of non-pharmacological therapy options onto tumor
associated fatigue (Kangas et al., 2008). These included
increase physical activity, relaxation techniques, psycho-
therapeutic intervention, nutrition counseling as well as
counseled interaction with other patients. Especially data
presented at the San Antonio Breast Cancer Symposium
2013 strongly supports physical activity as a positive
prognostic factor (Irwin, 2013) in breast cancer patients.
A recent meta-analysis investigated the effect of behavioral
psychological measures onto fatigue symptoms and was
able to also show a positive effect (Duijts et al., 2011). Our
study revisited this topic by giving 23 patients the option to
undergo a highly structured, cognitive behavioral therapy
regimen consisting of eight sessions, 90 minutes each. This
is the shortest time period of cognitive behavioral therapy
reported in literature to this date (Bjorneklett et al., 2012;
Finnegan-John et al., 2013). All of it can be performed as
an outpatient procedure. During these sessions, relaxation
techniques as well as coping and resource management
Techniques were taught, practiced and developed in a
group setting. The base assumption was that through
these techniques, patients will experience a decrease in
anxiety and depression symptoms resulting in a patient
collective no longer fulfilling the diagnosis criteria for
tumor associated fatigue. Thus, a manual was developed,
allowing a very structured and reproducible approach to
this type of psycho-oncological intervention. Difficult
as it may be to quantify results of a psycho-oncological
group intervention, two well-established and validated
scoring systems were used in order to paint a before and
after picture of the patients state of mind. The hospital
anxiety and depression scale as well as multidimensional
fatigue inventory were utilized to quantify the levels of
fatigue, depression and anxiety before and after eight
therapy sessions. Initially, 23 patients qualified for
this trial by the use of the fatigue coalition diagnosis
questionnaire. After completing 8 group sessions only 7
patients retained a pathological TAF diagnosis score. This
is a reduction of approximately 70% and by itself shows
the benefit of this program. The quantitative analysis of
the MFI scoring system showed a significant reduction
in four of five investigated parameters. An improvement
in the mental fatigue category could only be shown as a
trend. Furthermore, a statistically significant decrease in
the average anxiety and average depression score could
be shown using the HADS questionnaire. This analysis
showed a 33% reduction in anxiety and 57% reduction
and depression symptoms. It may therefore be said that
this type of structured, reproducible, behavioral therapy
based psycho-oncological intervention proved useful and
adequate in all evaluated dimensions.

While common sense may dictate that teaching
relaxation-, coping- and resource management skills, as
well as allowing the patient to interact with other patients
may be beneficial in the treatment of tumor associated
fatigue, this study, although small in patient number,
quantitatively proves that this type of intervention should
be part of any cancer treatment. Due to the small sample
size this work may be prone to over interpretation and
larger trials must follow. An interesting question is
whether or not additional intervention, i.e. additional
sessions would allow for further improvement of these
results. While one therapy block showed a significant
decrease in anxiety, depression and fatigue symptoms,
to a point where 70% fewer patients would no longer
qualify for a TAF diagnosis, it would certainly be
interesting to determine if further intervention produces
additional benefit. Based on our results and the results of
current meta-analyses regarding physical activity plus
psycho-oncological intervention, we currently employ
a program which extends these interventions to three
sessions coupled with extensive physical therapy.

In conclusion, this feasibility study showed that,
even though evaluating a small collective, a highly structured,
cognitive behavioral therapy based psycho-oncological
intervention will produce significant improvement in
patient well-being. This may be achieved in a short period
of 8 weeks. A 70% reduction in overall TAF diagnoses, as
well as a 33% reduction in anxiety and a 57% reduction
and depression symptoms could be produced. This leads
to a strong recommendation of this study group to include
this type of psycho-oncological intervention into all
postoperative breast cancer care.

References

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