

Beck Depression Inventory as a Predictor of Long-term Outcome Among Patients Admitted to the Breast Cancer Diagnosis Unit: A 25-year Cohort Study in Finland

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Abstract. *Aim: The Beck Depression Inventory (BDI) is one of the most widely used instruments for measuring the severity of depression. However, there has been no prospective study to investigate the long-term outcome in patients admitted to Breast Cancer Diagnosis Units. Patients and Methods: In the Kuopio Breast Cancer Study, women with breast symptoms were evaluated for total BDI score before any diagnostic procedures were carried out. The relapse-free survival (RFS) was calculated from the time of diagnosis to the time of first relapse including local relapse, contralateral breast cancer (BC) or metastatic disease. The overall survival (OS) was assessed as the time from the date of diagnosis to the date of last follow-up or death of the patient. The effect of the BDI on the RFS and on the OS were calculated by the Kaplan–Meier survival analysis and the difference between the groups was assessed by the log-rank test. The RFS and OS was estimated for the study groups with a low BDI score (<8) versus those with a high BDI score (≥8). The end-point of our study was to determine differences in long-term outcome and in BDI score in individuals with BC, benign breast disease (BBD) and in healthy study subjects (HSS). Results: In the Cox proportional hazard model, the total BDI score significantly predicted the 25-year RFS and OS in the HSS, BBD and BC groups combined (Hazard Ratio=1.87, $p=0.039$; Hazard Ratio=1.98, $p=0.048$,*

respectively), and in the Kaplan–Meier survival analysis with the log-rank test, the total BDI score predicted the 25-year RFS and OS in the HSS, BBD and BC groups combined ($p=0.043$; $p=0.036$, respectively). Conclusion: The BDI is a significant predictor of long-term outcome among patients admitted to the Breast Cancer Diagnosis Unit in Finland.

Breast cancer (BC) is the most common cancer in females, with approximately 1.7 million new BC cases diagnosed globally per year. In addition, over a half million women die every year from BC. In Finland 5008 women were diagnosed with BC in 2014 and 5-year and 20-year survival was reported to be 90% and 62%, respectively (1, 2). Many patients with BC experience recurrence and therefore it is of great importance to identify predictive factors to find the most effective treatment for each patient.

The Beck Depression Inventory (BDI) has become one of the most widely used psychometric methods detecting depression in normal populations and in different psychiatric patient cohorts (3). It has been translated into multiple languages including Arabian, Chinese, Japanese and Persian (4). Earlier, we assessed the psychometric tools BDI, Montgomery-Asberg depression rating scale and hopelessness/helplessness in healthy study subjects (HSS), benign breast disease (BBD) and BC groups and the results indicated a highly significant agreement between different psychometric inventories (5-9).

Because BC is a hormonally responsive neoplasm with great psychological impact, it is the tumour type most extensively investigated for possible psychological variables associated with risk and survival (10). Hormonal factors, such as early age at menarche, later age at menopause, later age at first full-term pregnancy and hormone replacement therapy, are known to be the main risk factors for sporadic BC (11-17). In addition, life-style factors, such as obesity, smoking, alcohol consumption and lack of physical activity, appear to contribute to an increased risk for this malignancy, although the results concerning such factors are inconsistent (11-17).

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Key Words: BDI, breast disease, breast cancer, 25-year outcome.

Table I. Characteristics of the study participants. Results are shown for patients with breast cancer (BC), those with benign breast disease (BBD) and for the healthy study participants (HSS).

Variable	HSS (n=28)	BBD (n=53)	BC (n=34)	p-Value
Age (mean, years)	45.7	47.6	51.6	0.12
Height (mean), cm	160.8	162.3	164.4	0.75
Body weight (mean), kg	68.3	67.8	72.5	0.25
Age at menarche (mean), years	13.4	13.4	13.4	0.99
Age at birth of I child (mean), years	25.0	25.0	25.2	0.92
Age at menopause (mean), years	50.0	48.9	47.9	0.53
No. of children (mean)	2.5	2.4	2.6	0.27
Parous, n (%)	23 (82%)	44 (83%)	31 (91%)	0.50
Breast feeding (mean), months	3.9	3.4	3.6	0.77
Use of oral contraceptives, n (%)	18 (64%)	25 (47%)	13 (38%)	0.12
HRT, n (%)	14 (50%)	36 (68%)	27 (79%)	0.44
Premenopausal, n (%)	18 (64%)	28 (53%)	13 (38%)	0.10
Postmenopausal, n (%)	10 (36%)	25 (47%)	21 (62%)	0.12
History of previous BBD, n (%)	10 (36%)	22 (42%)	18 (53%)	0.37
Family history of BC, n (%)	5 (18%)	5 (9%)	1 (3%)	0.21
Use of alcohol, n (%)	13 (46%)	31 (58%)	21 (62%)	0.44
Smoking, n (%)	10 (36%)	21 (40%)	15 (44%)	0.80
BDI (mean score)	7.8	8.5	8.9	0.70

HRT, Use of hormonal replacement therapy; BDI: Beck Depression Inventory.

Psychological factors, such as stressful and adverse life events, are widely thought to play a role in the aetiology of BC (18-39). There has been no prospective study to investigate the predictive value of the BDI in long-term outcome among patients admitted to the Breast Cancer Diagnosis Unit. Therefore, we carried out a prospective study to examine the association between BDI, and 25-year relapse-free (RFS) and overall (OS) survival in HSS, and patients with BBD and BC in a cohort in Finland.

Patients and Methods

The Kuopio BC Study was a multidisciplinary cooperative project conducted by different departments of the University of Kuopio and Kuopio University Hospital, and included all women who were referred to the hospital for breast examination between April 1990 and December 1995. The Kuopio BC Study followed the protocol of the International Collaborative Study of Breast and Colorectal Cancer coordinated by the European Institute of Oncology in Milan, and was initiated as a SEARCH program of the International Agency for Research on Cancer. The collaborative study is based on the assumption that BC and colorectal cancer may have common risk factors. Study centres for the BC study are situated in Canada, Finland, Greece, Ireland, Italy, Russia, Slovakia, Spain and Switzerland (40). The study participants showed BC symptoms (a lump in the breast or in the axilla, pain in the breast, bleeding from the nipple, nipple discharge or skin dimpling), or an abnormality of the breast and the indications for referral in this study were in line with our previous investigations in a BC Diagnostic Unit in Finland (41, 42).

This prospective case-control study was approved by the Kuopio University Hospital Board on Research Ethics (approval number 14/12/1989) and was conducted in accordance with the Declaration

of Helsinki. All study participants gave their written informed consent to participate in this study (43). One hundred and fifteen women participated and were interviewed (to determine the level of emotional depression) by a psychiatrist (P.O.) before any diagnostic procedures, so neither the interviewer nor the patient knew the diagnosis at the time of the interview. The interviews were recorded and the ratings were completed before the final diagnosis. The clinical examination, mammography and biopsy showed BC in 34 (29.6%) patients, BBD in 53 (46.1%) patients and 28 (23.4%) HSS (Table I).

Beck Depression Inventory (BDI). The women completed the 21-item BDI; the items of BDI contain four statements each, and describe the intensity of a particular depressive symptom (44-46). The total BDI score was rated as follows: grade I, score 0-3 (n=39), no depression; grade II, score 4-7 (n=22), little depression; grade III, score 8-11 (n=22), mild depression; grade IV, score 12-15 (n=16), moderate depression and grade V, score 16-30 (n=16), severe depression. In the present study, the total BDI score was used as a continuous variable with a cut-off of 8 for the total BDI score.

Statistical analysis. Significance of the results was calculated with the SPSS/PC statistical package (SPSS Inc., Chicago, IL, USA). Correlations and differences between the study groups were measured with the two-sided chi-square test and non-parametric Kruskal-Wallis variance analyses. The data on RFS and OS were collected and inspected from Kuopio University Hospital registry. The RFS was calculated from the time of diagnosis to the time of first relapse including the local relapse, the contralateral BC or the metastatic disease or the death. The OS was assessed as the time from the date of diagnosis to the date of last follow-up or death of the patient. The effect of the BDI on the RFS and on the OS were calculated by the Kaplan-Meier survival analysis and the difference between the groups was assessed by the log-rank test. The RFS and

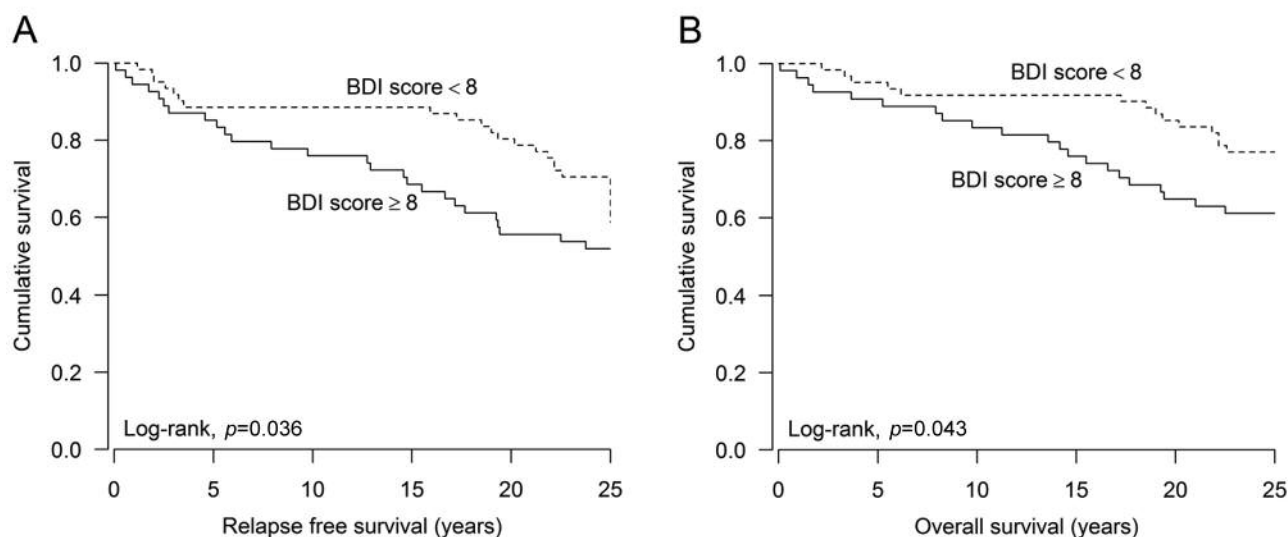


Figure 1. Kaplan–Meier survival curves for relapse-free (RFS) (1A) and overall (OS) survival (1B) for patients with breast cancer (BC), those with benign breast disease (BBD) and for the healthy study participants (HSS) groups combined ($n=115$) according to total Beck Depression Inventory (BDI) score. The total BDI score was a continuous variable for the study patients. The BDI score had a statistically significant effect on both RFS ($p=0.036$) and OS ($p=0.036$) by the log-rank test.

OS was estimated for the groups with a low BDI score (<8) versus groups with a high BDI score (≥ 8). The p -values and the hazard ratios (HRs) and their 95% confidence intervals (CI) were calculated from the Cox proportional hazard models. p -Values of 0.05 or less were considered to be statistically significant.

Results

Although the patients in the BC group were older than those in the BBD and HSS groups (51.5 versus 47.5 and 45.7 years, respectively), the age difference was not statistically significant ($p=0.12$). The majority of the patients (85/115, 74%) were married or living in a steady relationship. The groups differed only slightly from each other as to the factors of the reproductive life of the women and the mean BDI score values were quite similar in the HSS, BBD and BC groups ($p=0.0702$, Table I).

In the Kaplan–Meier survival analysis of the difference between the groups assessed by the log-rank test showed a low (<8) total BDI score to be a statistically significant favourable predictor of RFS (log-rank $p=0.36$, Figure 1A) and OS (the log-rank $p=0.43$, Figure 1B). In the Cox regression analysis, a low (<8) total BDI score was a statistically significant favourable predictor of the RFS (HR=1.87, 95% CI=1.03-3.38, $p=0.039$, Table II) and the OS (HR=1.98, 95% CI=1.00-3.90, $p=0.048$, Table III) in the HSS, BBD and BC groups combined. A similar although statistically insignificant pattern was seen in the HSS and the BBD groups (Table II). The 25-year RFS rate in the group with a low total BDI score (<8) versus those with a high total

Table II. Analysis of 25-year relapse-free survival (RFS) for the healthy study subjects (HSS) ($n=28$), benign breast disease (BBD) ($n=53$) and breast cancer (BC) ($n=34$) groups and for the study groups combined according to total Beck Depression Inventory (BDI) score. The p -values and hazard ratios (HRs) and their 95% confidence intervals (CI) were calculated from the Cox proportional hazard models.

Group	RFS (%)		HR	95% CI	p -Value
	BDI<8	BDI≥8			
All	31.1	48.1	1.87	1.03-3.38	0.039
HSS	36.8	20.0	0.53	0.11-2.56	0.430
BBD	17.9	33.3	2.08	0.68-6.36	0.199
BC	50.0	80.0	2.17	0.88-5.36	0.092

Table III. Analysis of 25-year overall survival (OS) for the healthy study subjects (HSS) ($n=28$), benign breast disease (BBD) ($n=53$) and breast cancer (BC) ($n=34$) groups and for the study groups combined according to total Beck Depression Inventory (BDI) score. The p -values and hazard ratios (HRs) and their 95% confidence intervals (CI) were calculated from the Cox proportional hazard models.

Group	OS(%)		HR	95% CI	p -Value
	BDI<8	BDI≥8			
All	77.1	61.1	1.98	1.00-3.90	0.048
HSS	84.2	80.0	1.34	0.22-7.99	0.752
BBD	85.7	79.2	1.59	0.43-5.91	0.492
BC	50.0	30.0	1.67	0.67-4.16	0.273

BDI score (≥ 8) differed significantly (31.1% *versus* 48.1%, respectively, Table II) and in the 25-year OS rate (77.1% *versus* 61.1%, respectively, Table III).

Discussion

The BDI, created by Aaron Beck, is a 21-question self-report inventory and marked a shift among health professionals, who had until then viewed depression from a Freudian psychodynamic perspective, instead of it being linked to the patient's own thoughts and cognitions. The BDI variables are based on Beck's records of the symptoms and signs of depressed patients in psychotherapy (44-46). A group of such symptoms and signs that were specific for the depressed patients were chosen for the BDI rating scale. There are three versions of the BDI: the original BDI, first published in 1961, then later revised in 1978 as BDI-1A, and the BDI-II, published in 1996. The current BDI version, the BDI-II, is designed for individuals aged 13 years and over, and is composed of items relating to symptoms of depression such as hopelessness and irritability, cognitions such as guilt or feelings of being punished, as well as physical symptoms such as fatigue, weight loss, and lack of interest in sex (47,48). The BDI was originally developed to provide a quantitative assessment of the intensity of depression. Because it is designed to reflect the depth of depression, it can monitor changes over time and provide an objective measure for judging improvement and the effectiveness or otherwise of treatment methods. The BDI is widely used as an assessment tool by health care professionals and researchers in a variety of settings.

The BDI suffers from the same bias as other self-report psychometric questionnaires, in that scores can be easily minimized or exaggerated by the test subject. Like all inventories, the way the questionnaire is administered can have an effect on the final score. If a test subject is asked to fill-out the questionnaire in front of other people in a clinical environment, for instance, social expectations have been shown to elicit a different response compared to questionnaire administration *via* a postal survey (49).

In study participants with concomitant physical illness, the BDI's reliance on physical symptoms such as fatigue may artificially inflate scores due to symptoms of the illness, rather than of depression (50). In an effort to deal with this bias, Beck and colleagues developed a short screening scale consisting of seven items from the BDI-II considered to be independent of physical function (Beck Depression Inventory for Primary Care, BDI-PC). The BDI-PC produces only a binary outcome of depressed or not depressed for study participants above a cutoff score of 4 (51). Although the BDI is designed as a screening tool rather than a diagnostic instrument, the BDI is often used by general practitioners to reach a quick diagnosis (52, 53).

In the Cox proportional hazard model, the total BDI score significantly predicted the 25-year RFS and OS in the HSS, BBD and BC groups combined, and in the Kaplan-Meier survival analysis with the log-rank test, the total BDI score predicted the 25-year RFS and OS in the HSS, BBD and BC groups combined.

Conclusion

The BDI is a significant predictor of long-term outcome among patients admitted to the Breast Cancer Diagnosis Unit in Finland.

Conflicts of Interest

No conflict of interest exists. The Authors alone are responsible for the content and writing of this article.

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