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## Social support and outcomes in older adults with lung cancer

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#### **Abstract**

**Background:** Insufficient social support is associated with increased mortality among older adults. Lung cancer is primarily a disease of older adults and is the leading cause of all cancer deaths. We assessed the association of social support with outcomes among older adults with lung cancer.

**Materials and Methods:** Adults age 65 and older with lung cancer with a completed geriatric assessment (GA) were assessed. Emotional social support (ES) and tangible (material, instrumental) support (TS) measures and patient characteristics were obtained from the GA. The electronic health record was used to extract clinical variables. Simple linear regression models evaluated the association between social support scales with patient and clinical factors.

**Results:** 79 adults were assessed. White race was positively associated with ES score (p=.04), while higher BMI (p=.03), depression (p=.03) and anxiety (p=.02) were associated with worse ES. Higher BMI was associated with higher/better TS score (p=.02) while living alone was associated with lower/worse TS score (p=.03). Completion of platinum-based doublet chemotherapy with immunotherapy as planned was associated with higher ES scores (p=.02) and higher TS scores (p=.02). Disease progression was associated with lower ES scores (p=.03).

**Conclusion:** Social support may influence clinical outcomes in older adults with lung cancer. As lung cancer often portends to poor prognosis, social support may be an important prognostic indicator.

#### Keywords

social support; geriatric assessment; older adults; lung cancer

### INTRODUCTION

The increased risk of mortality from insufficient social support is comparable to that of smoking or excessive alcohol consumption and potentially more troubling than inadequate levels of physical activity and obesity. Older adults are particularly susceptible to the higher mortality-low social support association, as shown in a study of 1,288 patients over the age of 65 that reported a 23% higher 12-year mortality risk associated with decreasing social support levels<sup>2</sup>. Studies in adults with cancer have shown that patients at high risk for low social support include those who are unmarried<sup>3</sup>, have depression,<sup>3</sup> and have chronic diseases.<sup>4</sup> In a systematic review of social influences on clinical outcomes in breast cancer, five out of seven articles reported an association between low social support and cancer progression.<sup>5</sup> For example, in a study of 2,264 women with breast cancer who were followed for ten years after they had completed their cancer treatment, women with lower social support had higher overall mortality. Similar findings were found in 168 patients with ovarian cancer who were followed from surgery to time of death. Patients with ovarian cancer in the top 75<sup>th</sup> percentile for social support scores had a 34% lower hazard ratio for death compared to patients in the bottom 25<sup>th</sup> percentile.<sup>7</sup> In a Japanese study following 44,152 people from cancer onset to disease progression, men with the lowest level of social support (baseline questionnaire) had 3.07 times higher risk of colorectal mortality.<sup>8</sup> To our knowledge, less is known about the association of social support with clinical outcomes among older adults with lung cancer.

Numerous validated scales exist for measuring social support, including the Medical Outcomes Survey (MOS) Social Support Survey which is focused on perceived quality of social support in two domains: tangible and emotional. Tangible social support pertains to the provision of basic needs (assistance with getting to the doctor, preparation of meals, etc.) while emotional support includes the presence of someone who encourages the patient to express their emotions in a meaningful way and offers empathy and comfort. Because the consequences of low social support and social isolation can be especially dire in older persons, to it is important to assess and consider the unique aspects of this variable in the treatment of older persons with cancer. In older patients with cancer who have low social support, studies have shown higher rates of unplanned hosital admissions and have pointed to the importance of having social networks.

The median age of cancer diagnosis is 66, with 54% of all new cancer cases diagnosed after the age of 65. <sup>14</sup> Thus, it is fair to characterize cancer as a disease of aging. The median age at diagnosis for lung cancer is 70 years and 70% of new diagnoses are in individuals over the age of 65. <sup>14</sup> Social support may be particularly challenging in this older patient population,

as reflected in studies showing particularly high levels of psychological stress associated with this diagnosis. <sup>15</sup> Older patients with lung cancer often have decreased mobility as well as comorbid chronic diseases such as Chronic Obstructive Pulmonary Disease(COPD), which may further increase their need for social support. <sup>4,16</sup> The positive impact of social support in patients with lung cancer was shown in a study of 289 adults that documented a positive relationship between high quality social support and better psychological function as well as increased patient resilience. <sup>17</sup> A second study of 218 patients with lung cancer showed a positive association between their perceived social support and their perception of their health at twelve months follow up. <sup>18</sup>

An important gap in lung cancer research is the investigation of potential associations between social support, measures of function and quality of life, and clinical outcomes. Within this area of investigation, it is important to consider the unique social support circumstances of older persons with lung cancer. <sup>19</sup> In our study of older adults with lung cancer (age 65 and older), we first investigated the association of emotional and tangible social support with measures included in a brief geriatric assessment (GA) that has been validated in clinical trials<sup>20</sup> and clinical practice settings.<sup>21</sup> We then investigated the association of social support with clinical outcomes.

#### **MATERIALS AND METHODS**

#### **Study Participants**

The sample includes adults age 65 or older diagnosed with Small Cell Lung Cancer (SCLC) or Non-Small Cell Lung Cancer (NSCLC) who completed the Cancer and Aging Group (CARG) geriatric assessment tool. The sample came from three studies, all of which included a brief geriatric assessment. NCT01752751 included adults age 65 or older with head/neck or lung cancer and conducted the GA prior to initiation radiotherapy or chemoradiotherapy (2012–2018). NCT01702844 included adults with NSCLC age 65 of older and conducted the GA prior to initiation of nab-paclitaxel treatment; patients had to have completed a single prior treatment regimen (2012–2015). NCT01137825 includes adults diagnosed with any cancer including lung age 65 or older with the GA administered at any time during cancer care – pre-treatment, during, or post-treatment. All participants provided written informed consent (2009–2019). Of the initial sample of 99 patients with lung cancer, eleven were excluded because they had an incomplete oncology history in the electronic medical record (EMR) and nine were excluded because they did not have complete responses for either emotional or tangible social support, leaving a final sample of 79.

#### Measures

**Brief Geriatric Assessment.**—The GA assessment used in this study <sup>20</sup> developed by Dr. Hurria and colleagues is available through the Cancer and Aging Research Group (CARG) website (http://www.mycarg.org/). All participants had completed the GA either pre-treatment (N=48, 61%), during treatment (N=17, 22%), or post-treatment (N=14, 17%). Geriatric Assessments were conducted within ten years of our final data collection cut-off

date of July 2019. Table 1 provides an overview of GA domains, measures, scoring range, and validated cut-points.

Social Support Scale.—The primary variable of interest is the Social Support Scale<sup>9</sup> which is comprised of two subscales, Emotional Support (ES) and Tangible Support (TS). Scoring for each subscale ranges from 0 to 100, and is dichotomized at 50, signifying lower support (<50) as compared to higher support (>=50). The ES subscale inquires about having someone to (1) listen to you when you need to talk, (2) give you good advice about a crisis, (3) give information to help you understand a situation, (4) confide in or talk to about yourself or your problem, (5) whose advice you really want, (6) share your most private worries and fears with, (7) turn to for suggestions about how to deal with a personal problem, and (8) who understands your problems. The TS subscale inquires about someone (1) who can help when you are confined to bed, (2) to take you to the doctor if needed, (3) to prepare your meals if you were unable to do it yourself, and (4) to help you with daily chores if you were sick.

**Function and quality of life.**—The GA includes self-reported demographic variables – age, race, gender, education, marital status, and living alone. Research staff assessed the Karnofsky Performance Status (KPS) and Timed Up and Go (TUG). All other measures were patient-reported.

Clinical diagnosis, treatment, and outcomes.—Electronic medical records data were extracted retrospectively from lung cancer diagnosis through end of treatment, departure from UNC Health Care, or death. Electronic medical record data included information on lung cancer diagnosis, treatment plan, side effects, adverse events, and treatment response.

#### Statistical Analysis

Simple linear regression models were used to evaluate the associations between the patient characteristics and the ES and TS subscales. Parameter estimates, standard errors, and p-values are reported in Table 3. The same analyses were conducted for associations of ES and TS with clinical variables (Table 4). When considering complement of regimens as planned, the associations of ES and TS were estimated to compare patients who completed regimens as planned to patients who received but did not complete treatment. All analyses were performed using SAS statistical software v9.4 (Cary, NC). Significance level was set *a priori* at 0.05.

#### **RESULTS**

#### **Sample Characteristics**

The final study sample (Table 2) includes 79 patients ranging in age from 65 to 90 years who had completed Social Support scales for at least one type of social support (77 completed ES scales and 78 completed TS scales). GAs were administered between 2009 and 2019. The sample was 87% white, 52% male, 34% high school graduate or less, 51% married, and 32% living alone. Mean Body Mass Index (BMI) was 26 (range 16–49), with 40% overweight (BMI 25–29) and 22% obese (BMI 30 plus).

#### **Geriatric Assessment Scores**

For the primary variable of Social Support, both ES and TS subscale scores were very high (range 0–100 with higher scores indicating greater social support). The mean ES score was 83.4 95% CI (78.95–87.85), with only 6% reporting "lower" emotional support. The mean TS score was 85.4 (95% CI: 80.84–89.99), with only 6% reported "lower" tangible support. Figure 1 illustrates the percent of patients within each decade of subscale units. For example, 22% and 28% of study participants scored TS and ES, respectively, in the 70–80 range.

Twenty-four percent of study participants had research staff-assessed Karnofsky Performance Status (KPS) scores of less than 80, 43% required more than fourteen seconds to complete the Timed Up and Go test, and 23% had fallen at least once in the last six months. Ninety-six percent scored "lower" function on the Physical Function score, 54% reported some limitations in Instrumental Activities of Daily Living (IADL), and 32% scored "lower activity" on the Social Activities Limitation scale. On the Mental Health Index (MHI), 28% were depressed and 35% were anxious.

#### **Clinical Characteristics and Outcomes**

Eighty-five percent of the study sample were diagnosed with NSCLC and 6% with SCLC (Table 2). Almost half of the study sample (47%) had Stage IV cancer, 94% received chemotherapy, 8% immunotherapy, and 43% oral targeted therapy.

Adverse events during the patient's treatment course included hospitalizations (62%), dose reductions (43%), dose delays (44%), and early treatment discontinuation (61%). Patient status at the time of data entry was deceased (71%), no evidence of disease (12%), and disease progression (17%).

# Univariate Associations of Social Support Subscales with Patient Characteristics and Geriatric Assessment Measures

Univariate associations of social support subscales with patient characteristics and geriatric assessment measures are presented in Table 3.

**Emotional Support (ES).**—Mean ES score was 13.96 points higher (better) in white patients compared to patients of color (p=.04). A one-unit increase in BMI was associated with a .93 increase in the ES score (p=.03). Scoring <50 on the Tangible Support subscale (signifying "lower" support) was associated with 52.64 points lower on the Emotional Support scale (p<.001). MHI Depression was associated with 11.65 lower score on the ES subscale (p<.001), as was MHI Anxiety (decrease of 11.91, p=.02).

**Tangible Support (TS).**—Living alone was associated with 11.00 points lower TS score (p=.03), and a one-unit increase in BMI was associated with a 1.02 increase in TS (p=.02). There was also borderline negative association with MHI-Anxiety (patients with anxiety had lower TS) (p=.06).

#### Univariate Associations of Social Support Subscales with Clinical Data

Univariate analysis of associations between social support subscales and clinical data are shown in Table 4. Completion of platinum-based doublet chemotherapy as planned with a biologic was associated with 14.61 points lower TS (p=.05). Completion of platinum-based doublet chemotherapy with immunotherapy as planned was associated with 14.3 points higher ES (p=.02) and 13.73 points higher TS (p=.02). (Among those with metastatic disease, disease progression was associated with 17.8 points lower ES (p=.03).

#### DISCUSSION

In this study, the social support scale commonly used in a validated geriatric assessment was explored for associations with patient characteristics, measures of function and quality of life, and clinical outcomes among older adults with lung cancer. In univariate analysis, we found a significant association between higher emotional support (ES) with white race, higher BMI, greater tangible support (TS), and less depression and anxiety. Living alone and lower BMI were both significantly associated with lower patient-reported TS. We found a significant association between lower ES and disease progression, and higher ES and TS with the completion of platinum-based doublet with immunotherapy as planned.

A notable finding from this study was that patients with higher social support were more likely to have higher BMI. A serious complication of lung cancer is cancer cachexia. Cachexia is associated with muscle and sometimes fat loss due to a decrease in appetite and the production of inflammatory mediators. These changes lead to a decreased BMI in patients who present with cachexia at the time of diagnosis.<sup>22</sup> Recent studies suggest that up to 46% of patients with non-small cell lung cancer already had cachexia at the time of diagnosis and treatment initiation.<sup>23</sup> A possible explanation for this difference in BMI is that patients with better social support may have presented earlier in the course of their diagnosis and more likely to present with less cachexia. The higher social support may also reflect having someone to help the patients with cooking and maintaining a healthy diet.

Another noteworthy finding of our study is that patients with lower ES scores were more likely to experience disease progression. As noted earlier, the association of low quality social support with increased risk of mortality has been well studied in other cancers including breast, colon, and ovarian cancer and given that lung cancer is the leading cause of all cancer deaths<sup>24</sup> understanding the mechanism by which poor social support leads to an increase in lung cancer progression and mortality is important. One potential molecular mechanism for social support's influence on cancer progression is the finding of elevated inflammatory mediators in patients with lower levels of social support.<sup>25</sup> The same inflammatory mediators (combined with low social support) have been associated with increased cancer-related mortality.<sup>25</sup> Other studies have suggested the mechanism of clinical influence of social support may be more multifactorial, focusing on social factors effect on behavioral, treatment, and physiological aspects of cancer.<sup>26</sup>

Our study has some important limitations. Relatively few patients in our sample reported low levels of emotional or tangible social support. Only 6% of the study sample scored within the range of "lower support" on these subscales. It is unclear if these findings are

unique to our specific population of older adults with lung cancer receiving care at our academic cancer center although similar findings of higher mean levels of self-reported social support have been observed in other studies, as well.<sup>3,11</sup> GAs were conducted at varying time points during the cancer care continuum as outlined in our methods. This contributed to the heterogeneity of the sample. The GA timeframe was 2009 to 2019, which include a period of substantial advances in the treatment of lung cancer. A combination of a small sample size and low prevalence of "lower support" did not allow multivariate analyses to assess the relative contributions of variables that were significant in our univariate analysis. Further evaluation in a larger sample with a more racially and ethnically diverse patient population is warranted.

Despite these limitations, our findings highlight the presence of significant associations between clinical endpoints (such as disease progression and completion of planned number of cycles) and emotional social support in a sample of older adults with lung cancer. Further research on the role and impact of social support among older patients with lung cancer may ultimately inform interventions that can address the impact of low social support. These interventions may in turn improve cancer outcomes, quality of life, and overall patient satisfaction with care.

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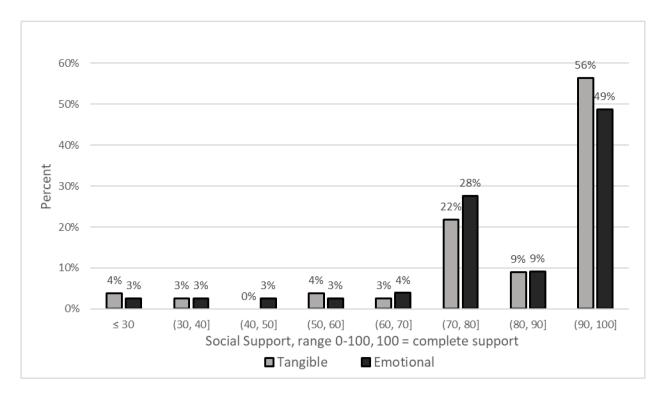
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**Figure 1.** Social Support Scores

#### Table 1.

#### Brief Geriatric Assessment

Domains	Measures	Score Range	Dichotomized		
Research Staff Assessed					
Function	Timed Up and Go (TUG)	Greater number of seconds → Lower function	Time>14 seconds = lower function <sup>27,28</sup>		
	Karnofsky Performance Status (KPS)	Range: 0–100 Lower value → lower function	<80 = lower function <sup>29,30</sup>		
Body Composition	Body Mass Index (BMI)	No range limits	Normal (18.5–24.9), overweight (25–29.9), obese ( 30) <sup>31</sup>		
	•	Patient Reported	•		
Function	Patient-reported KPS	Range: 30–100 Higher score → better functioning	<80 = lower functioning <sup>30</sup>		
	Physical function	Range: 0–20, 20= not limited	<20 = limited <sup>32</sup>		
	Instrumental activities of daily living (IADL)	Range: 0–14 14= no limitations	<14 = limitations <sup>33</sup>		
	Number of falls in last 6 months <sup>34</sup>	No range limits	1 or more		
Psychological	Mental Health Index (MHI): Depression	Range 0-43	Depression score >/= 12 <sup>35</sup>		
	Mental Health Index (MHI): Anxiety	Range 0–20	Anxiety score >/=6 <sup>35</sup>		
Social	Social Activity Limitation	Range: 0–100 Higher score = more activity	<50 = less activity <sup>36</sup>		
	Social support-emotional	Range: 0–100 Higher score = greater support	<50 = lower support <sup>9</sup>		
	Social support-tangible	Range: 0–100 Higher score= greater support	<50 = lower support <sup>9</sup>		

Table 2.

### Patient Characteristics (N=79)

Variable	Value
Age – mean (range)	73 (65–90)
Race	
White	69(87%)
Black or African-American	8 (10%)
Native Hawaiian or Pacific Islander	1 (1%)
Native Indian or Alaskan Native	1 (1%)
Gender	
Male	41 (52%)
Female	38 (48%)
Education	
More than high school	27 (34%)
High school graduate or less	52 (66%)
Marital Status	
Not Married	38 (49%)
Married	40 (51%)
Unknow	1
Living Alone	
No	49 (68%)
Yes	23 (32%)
Unknown	7
Body Mass Index (BMI) – mean (range)	27 (15–49)
Underweight (BMI <18.5)	1 (1%)
Normal weight (BMI 18.5 to <25)	28 (35%)
Overweight (BMI 25 to <30)	31 (39%)
Obese (BMI 30 or higher)	17 (22%)
Unknown	2 (3%)
Clinical Diagnosis	1
Small Cell Lung Cancer	5 (6%)
Limited	3
Extensive	2
Non-Small Cell Lung Cancer	66 (84%)
Adenocarcinoma	47
Squamous Cell Cancer	16

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Variable	Value
Large Cell Carcinoma	1
Mixed Histology	2
Missing histology/unspecified histology	8 (10%
Stage at diagnosis	
1	11 (14%
2	6 (8%)
3	19 (24%
4	32 (41%
Unknown	11 (14%
Metastasis sites	
Bone	11 (14%
Brain	13 (16%
Both	4 (5%)
Assessments and Questionnaires	
Social Support-Emotional Score (range 0–100) – score <50 ("lower" support)	5 (6%)
Social Support-Tangible Score (range 0–100) score <50 ("lower" support)	5 (6%)
Treatment stage when the Geriatric Assessment was conducted	
Pre-treatment Pre-treatment	48 (61%
During treatment	17 (22%
Post treatment	14(18%
Karnofsky Performance Status (KPS <80)	19 (24%
Timed Up and Go test more than 14 seconds	34 (43%
Patient-Reported Karnofsky Performance Status (<80)	25 (32%
Falls in past 6 months 1 or more	18 (23%
Physical Function <20 ("lower" function)	73 (96%
Instrumental Activities of Daily Living/IADL <14 (some limitations)	42 (54%
Social Activity Limit Score (range 0–100) <50 ("lower" activity)	25 (32%
Mental Health Index (MHI)-Depressed (range 0–43) – Depressed (score >12)	19 (28%
Mental Health Index (MHI)-Anxious (range 0–20) – Anxious (score >=6)	26 (35%
Treatment	
Surgery: yes	34 (43%
Unknown	8 (10%

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Variable	Value
Radiation: yes	54 (68%)
Unknown	5 (6%)
Clinical Trial: Yes	26 (33%)
Unknown	19 (24%)
Treatment Goal	
Palliative Care	44 (56%)
Curative	27 (20%)
Unknown	8 (10%)
Number of Prior Regimens Started	
0	17 (22%)
1	36 (46%)
2	9 (11%)
3	8 (10%)
4	9 (11%)
Chemotherapy treatment ever received – can be more than one treatment	
Single-Agent Chemotherapy	26 (42%)
Platinum-Based Doublet	37 (47%)
Doublet Therapy without Platinum	2 (3%)
Platinum-Based Doublet with Biologic	8 (10%)
Platinum Doublet with Immunotherapy	13 (16%)
Chemotherapy – yes	62 (78%)
Unknown	5 (6%)
Completed Planned Cycles of Chemotherapy – yes	22 (28%)
Unknown	23 (29%)
Treated with Immunotherapy – yes	6 (8%)
Unknown	17 (22%)
Completed Planned Cycles of Immunotherapy yes	3 (50%)
Given Oral Targeted Therapy – yes	34 (43%)
Unknown	7 (9%)
Adverse Events During Treatment	
Hospitalized yes	36 (46%)
Dose reduction – yes	23 (29%)
Treatment Delay – yes	19 (24%)
Early treatment Discontinuation yes	32 (41%)
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Variable	Value
Patient Status at the time of Data Entry	
Deceased	48 (61%)
No Evidence of Disease	8 (10%)
Disease Progression	7 (9%)
Unknown	16 (20%)

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Table 3.

Univariate associations of Social Support (Emotional and Tangible) with patient characteristics and Geriatric Assessment measures

	Social Support-Emotional Score		Social Support-Tangible Score	
Variable	Estimate	p value	Estimate	p value
Age	-0.29	.45	03	.94
Race (white)	13.96	.04	11.95	.08
Gender (male)	-1.71	.71	.86	.85
Education (high school graduate or less)	-4.88	.31	-2.83	.56
Marital status (married)	7.35	.11	8.46	.07
Living Alone – yes	-8.44	.08	-11.00	.03
Body Mass Index (BMI)	.93	.03	1.02	.02
Karnofsky Performance Status score <80	55	.92	2.15	.69
Patient-Reported Karnofsky Performance Status score <80	-2.68	.59	2.33	.64
Timed Up and Go test > 14 seconds	-5.26	.25	22	.96
One or more falls in last 6 months	-5.08	.37	1.44	.80
Physical Function – low function score <20	6.64	.57	-15.36	.21
Instrumental Activities of Daily Living (IADL) – limitations score <14	-3.26	.48	1.73	.71
Social Activity Limit Score – lower social activity score <50	-5.30	.28	-1.45	.77
Social Support-Tangible Score – less tangible support score <50	-52.64	<.001		
Mental Health Index (MHI) – Depressed	-11.65	.03	-8.63	.14
Mental Health Index (MHI) – Anxious	-11.91	.02	-9.66	.06

Social Support subscales Emotional and Tangible range from 0–100, with higher scores signifying higher social support. Positive estimate implies positive association with social support; negative estimate implies inverse association with social support.

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 Table 4.

 Univariate associations of Social Support (Emotional and Tangible) with treatment and outcomes

	Social Support-F	Social Support-Emotional N= 77		Social Support-Tangible N= 78	
Variable	Estimate	p value	Estimate	p value	
Number of regimens started	0.38	.83	1.89	.30	
Received single-agent	-0.01	1.00	2.49	.63	
Received platinum-based doublet	-9.35	.06	-0.68	.90	
Received doublet w/out platinum	4.22	.77	8.69	.54	
Received platinum-based doublet chemotherapy with biologic	-6.97	.35	-14.61	.05	
Received platinum-based doublet chemotherapy with immunotherapy	14.30	.02	13.73	.02	
Completed planned cycles of chemotherapy	4.79	.34	2.56	.60	
Completed planned cycles of oral targeted therapy	-6.67	.26	-6.35	.25	
Hospitalization – yes	2.82	.60	2.40	.68	
Dose Delay- yes	-1.95	.78	-4.51	.56	
Dose Reductions- yes	-2.30	.72	-6.21	.37	
Deceased	3.90	.53	4.61	.47	
No evidence of disease	9.46	.23	1.41	.86	
Disease progression	-17.80	.03	-10.04	.25	

 $<sup>^*</sup>$ Positive estimate implies positive association between the clinical variables and the outcome – Social Support score.