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Coexistence of Urinary Incontinence and Major Depressive Disorder with Health-Related Quality of Life in Older Americans with and without Cancer

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Abstract

Purpose—This study evaluates the prevalence and factors associated with major depressive disorder (MDD) in a population of cancer survivors and the impact of co-occurring MDD and urinary incontinence (UI) on health-related quality of life (HRQOL).

Methods—The prevalence of MDD risk among cancer survivors (breast, prostate, bladder, colorectal, lung and endometrial/uterine cancers) and those without cancer was estimated using the Surveillance, Epidemiology and End Results Program-Medicare Health Outcomes Survey (SEER-MHOS) linked database (n=9,282 with cancer/n=289,744 without cancer). Risk for MDD was measured using 3 items from the Diagnostic Interview Schedule and HRQOL was measured by the SF-36. UI was defined as self-reported leakage of urine causing a problem in previous 6 months. Factors associated with MDD were investigated using logistic regression and the impact of co-occurring MDD and UI on HRQOL scores was determined using linear regression.

Results—The prevalence of MDD risk ranged from 19.2% for prostate-34.1% for lung. Lung cancer diagnosis was associated with risk of MDD. Being 5 years from diagnosis was associated with decreased risk of MDD (Prevalence Odds Ratio (POR)=0.82, 95% Confidence Interval (95% CI): 0.71, 0.95). The coexistence of both UI and MDD was associated with a decrease across HRQOL subscales; including 40-points on role emotional (RE) score.

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Competing Interests

The authors have no conflicts of interest or financial disclosures.

Conclusions—Cancer survivors reporting co-occurrence of UI and MDD experienced significant decrements in HRQOL.

Implications of cancer survivors—Understanding the combined effect of UI and MDD may help clinicians to better recognize and alleviate their effects on cancer survivors' HRQOL.

Keywords

urinary incontinence; major depressive disorder; health related quality of life; cancer; survivor

Introduction

By the year 2030, an estimated 71 million Americans will be 65 years of age and older, accounting for approximately 20% of the U.S. population [1]. This large-scale population shift will result in an increase in cancer incidence as more than 60% of new cancers occur in adults 65 years of age and older [2]. Approximately 13.7 million people in the U.S. are currently cancer survivors and this number is expected to increase to 18 million by the year 2022, representing a 31% increase in cancer survivors [3, 4]. Increasing age has been previously associated with more frequent health problems which can result in negative impacts on health-related quality of life (HRQOL) [5, 6].

There is increasing awareness of the importance of the overall health of cancer survivors, and in research to identify potential ways to improve the health care of cancer survivors. Major depressive disorder (MDD) and urinary incontinence (UI) are both prevalent, non-cancer issues in the elderly and in cancer survivors, but have been understudied among cancer survivors. The Center for Disease Control (CDC) and population-based studies estimate that approximately 1 in 10 adults in the U.S. are currently experiencing major depression [7]. In addition, the lifetime prevalence of depression is estimated to be 16% in the general population [8] and is estimated to be even higher in elderly populations, possibly up to 30% [9]. The rate of MDD is particularly high in those recently diagnosed or being treated for cancer, with a prevalence ranging from 10% to 25.5% [10]. MDD is a key cause of disability [11] and has been associated with higher mortality and a higher number of comorbid conditions [9] among the elderly.

Urinary symptoms are also highly prevalent in the general population, with estimates of about 17% among men and 38% among women who are above the age of 60 years [12–14]. It has been shown that UI impacts many facets of an individual's life, including social, physical, psychological, work productivity and sexual health [15–19]. In addition, urinary symptoms are associated with certain cancers (for example: bladder, prostate, gynecological cancers) as well as a side effect of cancer treatments [20–23]. Previous research found that reporting UI symptoms was independently associated with decreased HRQOL scores in a population of older Americans across cancer diagnoses [24]. UI is a potentially treatable and reversible condition; in fact, multiple treatment options exist [25, 26]. For cancer survivors, who experienced persistent UI symptoms after a prostatectomy or hysterectomy, some surgical treatments have been effective in improving their symptoms [26–28].

The link between UI and depression has been described, and the co-occurrence of these two conditions has been demonstrated to have an additive impact on decrements of HRQOL, in the general population [29]. However, there has been very little, if any, research investigating the co-occurrence of MDD and urinary incontinence and the association with HRQOL among cancer survivors. To help better understand the issues of MDD and urinary incontinence in cancer survivors, this study aims to evaluate the prevalence of MDD among older Americans with and without cancer; to determine factors associated with MDD such as cancer diagnoses (bladder, breast, colorectal, endometrial/uterine, prostate or lung), demographics, comorbidities, and urinary incontinence symptoms; and to investigate the combined impact of UI and MDD on HRQOL.

Methods

SEER-MHOS data linkage

This study utilizes the population-based SEER-MHOS database, which links the Surveillance, Epidemiology and End Results [SEER] program of cancer registries with the Medicare Health Outcomes Survey [MHOS] to provide detailed information about Medicare beneficiaries who have been diagnosed with cancer. Extensive details of the SEER-MHOS data linkage have been published previously [30]. The National Cancer Institute's SEER program collects and publishes cancer incidence and survival data from population-based cancer registries for approximately 28% of the U.S. population [31]. Additionally, SEER collects demographic, tumor characteristic and selected cancer care data for individuals. The MHOS, which includes self-reported information on socioeconomic, demographic, co-morbidity, health and functional status, is given to randomly selected Medicare beneficiaries from each managed care plan under contract with the Centers for Medicare & Medicaid Services (CMS) each year. These selected Medicare beneficiaries are given a baseline survey, and if they are still in the same managed care plan two years later they also receive a follow-up survey.

IRB exemption was obtained from University of North Carolina at Chapel Hill.

Cohort and participant selection

This cross-sectional analysis draws from the baseline and follow-up data from five SEER-MHOS cohorts (2001 and 2003, 2002 and 2004, 2003 and 2005, 2004 and 2006, 2005 and 2007). These cohorts were selected as these survey years include measures of both UI and the depressive symptom questions used in the MDD algorithm. For participants diagnosed with cancer and who have completed two or more SEER-MHOS surveys, the responses to the first MHOS survey after the most recent cancer diagnoses were used. For those without cancer diagnoses, the first survey was used. If the respondent had missing data on any of the questions used to determine MDD or UI then the follow-up survey was used. Study participants were excluded for the following reasons: (1) if missing data on MDD or UI (n=635 for those with cancer diagnosis, n=16,321 for those without a cancer diagnosis); (2) men with breast cancer diagnosis (n=7) which had too small of a sample size to produce stable estimates, and (3) participants under the age of 65 years (n=209 for those with cancer diagnosis, n=20,833 for those without a cancer diagnosis).

The final sample sizes included in this analysis for 6 different cancer types were as follows: (1) prostate (n=3,133), (2) female breast (n=2,649), (3) colorectal (n=1,645), (4) bladder (n=710), (5) lung (n=607) and (6) endometrial and uterine (n=538). 289,744 individuals who had never been diagnosed with cancer were selected.

SEER-MHOS Measures

In our study, MHOS administered the 3 questions described above from the Diagnostic Interview Schedule (DIS). Being at “risk of major depressive disorder” (MDD) was defined using an algorithm by Rost *et al.*, 1993[32]. The algorithm includes two methods in which to screen positive for MDD. The first is a positive screen for risk of MDD was defined by an affirmative response to the question ‘In the past year, have you had 2 weeks or more during which you felt sad, blue or depressed; or when you lost interest or pleasure in things that you usually cared about or enjoyed? (yes/no)’. The second way to screen positive for risk of MDD, included an affirmative response to *both* of the following questions: (1) ‘In the past year, have you felt depressed or sad much of the time? (yes/no)’; and (2) ‘Have you ever had 2 years or more in your life when you felt depressed or sad most days, even if you felt okay sometimes? (yes/no)’ in addition to responding to at least ‘some of the time’ to the question ‘how much of the time during the past 4 weeks have you felt downhearted and blue? (none of the time/some of the time/most of the time/all of the time)’. This last question is a replacement for a similar question used in the methods proposed by Rost *et al.*, asking how much of the time in the past week the participant felt depressed [32].

No information is available on the sensitivity and specificity of these 3 items in a Medicare population. However, Whooley and colleagues (1997) examined the sensitivity and specificity of a two-item self-report depression screener from the DIS (“During the past month, have you often been bothered by feeling down, depressed, or hopeless?” and (2) “During the past month, have you often been bothered by little interest or pleasure in doing things?”) in an older study population with an average age of 53 [33]. For Whooley’s two-item depression screener, a sensitivity of 96% (95% CI: 90–99), specificity of 57% (95% CI: 53–62), and area under the ROC curve of .82 (95% CI: .78–.86) were found [33]. Due to the similarities in the study populations and the questions administered, it is likely that the sensitivity and specificity of the 3 DIS items in our Medicare population would be comparable. The gold standard for identifying those at risk of MDD would be a diagnosis by a trained physician, which would be immensely difficult to obtain for a sample size as large as the one utilized here.

UI was defined by an affirmative to the following question: ‘Many people experience problems with urinary incontinence, the leakage of urine. In the previous six months, have you accidentally leaked urine? (yes/no)’ in conjunction with indicating that the urine leakage was either ‘a big problem’ or a ‘small problem’ via the question ‘How much of a problem, if any, was the urine leakage to you?’ (a big problem/a small problem/not a problem).

HRQOL scores were assessed by the Short-Form 36 (SF-36, version 1) and, subsequently the Veterans Rand-12 (VR-12). The MHOS switched from the SF-36 to the VR-12 in 2006, thus affecting the surveys from the last two cohorts included in this analysis. SF-36 and VR-12 Physical Component Summary (PCS) scores have been linked using a published

algorithm [34]. The PCS T-score metric was normed, with the average in the U.S. population being 50 and a standard deviation of 10 [35]. Higher scores reflect better HRQOL. All available physical subscales were used, including bodily pain (BP), role-physical (RP), physical functioning (PF) and general health (GH) [36]. The mental subscales utilized (social functioning (SF), role-emotional (RE) and vitality (VT)) did not include the Mental Component Summary (MCS) or mental health (MH) subscale as both scales relied on the one of the questions used to screen for risk of MDD [36]. The subscales were defined by items that were transformed on a 0–100 scale. The subscales, unlike PCS, are not normed to the U.S. population average.

Key variables investigated in this analysis included: cancer diagnosis, age at survey, current smoking status, race, gender, marital status, education, comorbid conditions, time since diagnosis, cancer severity and difficulty with activities of daily living (defined as responding affirmatively to one or more of the following: difficulty getting out of a chair, using a toilet, walking, dressing, eating, or bathing). Comorbid conditions were defined as a dichotomous variable (yes/no) and were self-reported on the MHOS survey by participants.

Statistical Analysis

Univariate distributions of demographic and clinical covariates of the study population were characterized by cancer type (bladder, breast, colorectal, endometrial/uterine, lung, and prostate) and variables were defined as follows (index/referent): MDD (yes/no), UI (yes/no), age (65–74, 75–84, 85), smoking status (yes/no), race (other/non-Hispanic white), gender (male/female), marital status (other/married), education (> high school, high school), high blood pressure (yes/no), stroke (yes/no), chronic lung disease (including COPD, asthma and emphysema) (yes/no), gastrointestinal (including Crohn's disease, ulcerative colitis, and inflammatory bowel disease) (yes/no), diabetes (yes/no), difficulty completing one or more activities (getting out of a chair, using the toilet, walking, bathing, dressing, eating) (yes/no), joint pain (yes/no), any cardiovascular disease (one or more of chronic heart failure, myocardial infarction, angina or coronary artery disease, or other heart condition) (yes/no), cancer severity (distant or regional/in situ or localized), and time since cancer diagnosis to time of survey (defined as: survey within 2 years after diagnosis, survey within 2–5 years after diagnosis, and survey >5 years after cancer diagnosis). Time since diagnosis categorizations were selected to approximate active or early post treatment, short-term survival, and long-term cancer survival periods, respectively. A chi-square was used to test the difference in the distribution of the categorical demographic and clinical covariates between each cancer group compared to those without cancer. The variable for time since diagnosis and cancer severity did not have a clear referent group (as the referent group in this model is those without cancer) and thus was not tested for significance.

Due to the very large sample size of this study, a minimally important difference (MID) equal to 1/3 of the standard deviation was used to determine statistical significance for continuous HRQOL measures (PCS is normed to a standard deviation of 10; for subscales, the standard deviation of those without cancer was used) [37]. Unless noted, all future references to statistical significance for HRQOL scales are based on their respective MID.

Logistic regression models were used to examine demographic and clinical factors associated with MDD by calculating Prevalence Odds Ratios (PORs) and 95% Confidence Intervals (95% CIs). The variables in the model included all cancer types and other demographic and clinical factors used in univariate analyses. Another logistic model was performed limited to cancer survivors (excluding non-cancer individuals) and included variables for cancer severity and time since cancer diagnosis. Cancer survivors with colorectal cancer were chosen as the referent group in this second model due to a similar prevalence of MDD and UI compared to the no cancer group.

Linear regression models were used to estimate the adjusted means on HRQOL, by cancer site, combined MDD and UI status, adjusting for relevant covariates listed above for the logistic regression models. A multiplicative interaction was assessed between MDD and UI for HRQOL using cross-product terms in the linear regression model. For this analysis, Medicare beneficiaries were defined based on their joint reporting of MDD and UI by creating 3 dummy variables for the regression models. This resulted in them being categorized into four different groups: (1) no MDD, no UI; (2) no MDD, UI; (3) MDD, no UI; and, (4) MDD, UI.

All statistical analysis was completed using SAS 9.3 (Cary, NC).

Results

The distributions of demographic and clinical covariates by cancer type are shown in Table I. The mean (SD) of PCS in the no cancer population was 39.8 (12.0). The mean score for many HRQOL scales (PCS, BP, PF, GH, SF, RE and VT) for those diagnosed with lung cancer was significantly lower than the mean HRQOL score for participants without cancer. The prevalence of MDD risk among cancer patients ranged from 19.2% in prostate cancer patients to 34.1% in lung cancer; compared to a prevalence of 20.5% in the no cancer population. Reporting UI was more common across all cancer types than in the no cancer group (27.1%) but was particularly high among older Americans with a history of endometrial/uterine, breast and prostate cancers. In general, the percentage of the no cancer group suffering from comorbid health conditions tended to be lower than the proportions among cancer survivors. This may be due, in part, to a higher percentage of individuals of an older age at the time of survey completion in the cancer group compared to those with no cancer.

The prevalence of co-occurring UI and MDD risk was highest in those with lung cancer (14.2%) and endometrial/uterine cancer (12.0%) whereas among those without cancer the prevalence was 8.6%. The prevalence ranged from 9.2% to 10.8% in all other cancer types.

Factors associated with MDD risk among Medicare beneficiaries with and without cancer are presented in Table II. Being diagnosed with lung cancer (POR= 1.66, 95% CI: 1.37, 2.01) was associated with the largest increase in odds of risk for MDD across cancer types; whereas being diagnosed with other cancer diagnoses (breast, colorectal, endometrial/uterine and prostate) was not associated with elevated MDD risk. Another significant factor associated with MDD was having difficulty completing daily activities (POR= 2.55, 95%

CI: 2.50, 2.61). UI was in particular significantly associated with MDD (POR=1.68, 95% CI: 1.63, 1.70), as was being a current smoker (POR= 1.40, 95% CI: 1.35, 1.44). Other variables with significant POR above 1.2 included being not married, having less education, stroke, chronic lung disease, gastrointestinal problems, joint pain and cardiovascular disease.

The estimates for factors associated with MDD risk among only the cancer survivors and when taking into account time since diagnosis and cancer severity are similar in magnitude to the results presented in Table II. When fitting the same model only among cancer patients, a slight decrease in odds of MDD (Table III; POR=0.83, 95% CI 0.72, 0.97) was observed for cancer survivors who were more than 5 years from diagnosis relative to those less than 2 years from diagnosis. Cancer severity was not significantly related with MDD risk.

Results from linear regression of co-occurrence of MDD and UI and cancer types associated with PCS and physical HRQOL subscales (BP, RP, PF and GH) adjusted for demographic factors and comorbid conditions are presented in Table IVa. Decrements in PCS and physical subscales associated with UI, MDD and coexisting UI and MDD are illustrated in Figure 1. The estimate for the intercept in this model represents the adjusted mean PCS or physical subdomain score for the non-cancer group without any comorbid conditions. A statistical interaction ($p < 0.001$) was observed between UI and MDD with PCS, BP, RP and PF, and therefore the results are presented by the combined prevalence of these two conditions. In this analysis, the co-occurrence of MDD and UI was also associated with an almost 3-point decrease in PCS (adjusted mean score= 47.36, 95% CI: 47.22, 47.51), which was not significant based on the MID criteria. However, the co-occurrence of MDD and UI was significantly associated with decreased HRQOL as defined by the MID criterion for BP (adjusted mean score=73.48, 95% CI: 74.12, 74.82), RP (adjusted mean score=68.81, 95% CI: 68.10, 69.52) and GH (adjusted mean score=62.31, 95% CI: 62.00, 62.62). For PCS and GH, lung cancer diagnosis was also significantly associated with decreased scores.

The results for the co-occurrence of MDD and UI and cancer types associated with mental HRQOL subscales (SF, RE, and VT) are presented in Table IVb. Decrements in mental subscales associated with UI, MDD and coexisting UI and MDD are also illustrated in Figure 1. For the three mental subdomains included in this analysis, the co-occurrence of MDD and UI was significantly associated with decreased HRQOL scores. This was especially prominent for RE, with an average decrease of 40 points for those with MDD and UI (adjusted mean score=55.33, 95% CI: 54.65, 56.02). Similarly for SF scores, the co-occurrence of both MDD and UI was associated with a large observed decrease in SF (adjusted mean score=71.5, 95% CI: 71.14, 71.87). Similarly to PCS and GH, Medicare beneficiaries with lung cancer had the lowest SF (adjusted mean score=88.06, 95% CI: 86.14, 89.98) and VT score across cancer types (adjusted mean score=65.33, 95% CI: 63.85, 66.80).

Discussion

There has been considerable previous research that has demonstrated that UI has been associated with higher prevalence of depression and anxiety in the general population [15,

38, 39]. However, this is the first study, to our knowledge, to assess the co-occurrence of MDD risk and UI in a population of older Americans with and without cancer. The results of this study have high public health impact potential due to the substantial prevalence of MDD risk. In this population of Medicare beneficiaries, MDD risk was prevalent, ranging between 20–34% among cancer survivors and older Americans without cancer. This study found that the co-occurrence of MDD risk and UI had a large negative impact on RE and SF scores, as well as a similar decrease among other HRQOL scales although to a lesser extent. We did not observe significant changes in PCS meeting our MID criteria except for a 4-point decrement among lung cancer participants compared to those without cancer; however, the decrease observed may not have clinical relevance when considering other MIDs proposed for PCS in the literature [40, 41].

While high prevalence of MDD risk was observed in this study population, only a history of a lung cancer diagnosis had a strong association with reporting symptoms associated with MDD. This could be due to the poor prognosis typical of lung cancer diagnoses, as well as the fact that approximately half of lung cancer patients were within 2 years of diagnosis with more severe disease and thus, were likely in active treatment [42]. Reporting MDD was associated with many demographic and clinical factors, with the strongest factors being UI and having physical difficulty completing activities. Due to the fact that MDD is often underdiagnosed, the results of this study highlight particular factors that may be useful in identifying MDD and improving diagnosis and treatment of it among older Americans with and without cancer [43]. A longitudinal study among men found that untreated depressive symptoms at baseline to be associated with an almost 3-fold increased risk of nocturia at follow-up [44]. Therefore, identification and subsequent treatment of MDD may aid in the prevention of future UI symptoms. [45]

While cancer severity was not found to be associated with MDD, the results do suggest that increasing years from time of diagnoses among cancer survivors was associated with decreased odds of reporting MDD. Prevalence of depression being the highest immediately after diagnosis and decreasing over time has been previously reported among breast cancer survivors [46, 47] but was not observed in a New Zealand cohort of cancer survivors [48]. These differences could be due to geographic differences in the study populations or differing medical and treatment experiences. A population-based study of long-term cancer survivors did not find an increased risk of MDD compared to the general population [49], which supports the finding reported here that suggests MDD risk might be more relevant in the first two years after a diagnosis.

We also found age at time of survey to be inversely associated with risk of MDD, although this effect was not strong in magnitude. Previous reports investigating the effect of aging on risk of MDD or depressive symptoms have been inconsistent [50–52]. Although not established, it has been hypothesized that increased age after a certain threshold may be associated with a decreased susceptibility to depressive symptoms or anxiety due to decreased emotional responsiveness [51].

The prevalence of MDD reported in this study is comparable to depression estimates previously reported among elderly populations [9]. Our method of defining MDD risk was

based on a previous published algorithm for determining risk MDD based on a similar questionnaire [32]. A sensitivity analysis defining ‘depressive symptoms’ as an affirmative response to any one of the three MDD questions increased the prevalence but did not significantly alter results of conclusions for the predictors of MDD or the associations with HRQOL scales.

Decrements across multiple HRQOL scales (PCS, GH, SF and VT) were associated with lung cancer diagnoses. However, for many of the HRQOL subscales the lowest average adjusted mean score was observed for the coexistence of both MDD and UI; this decrement was particularly strong for SF and RE. The decrements observed ranged in magnitude from almost 10 points on the PF to almost 40 on RE; changes this large in magnitude are considered moderate to large clinically important differences [40]. This suggests that these two conditions together have a considerable impact on the emotional and social lives of Medicare beneficiaries, more so than the cancer diagnoses investigated in this analysis. Improved awareness of the emotional and social impact of these conditions, among other HRQOL decrements, may promote the advancement of strategies for detection and treatment of both MDD and UI.

This study is limited by the cross-sectional nature, and thus we cannot draw conclusions about temporality between MDD, UI and HRQOL. In addition, there were predictors of MDD or UI that could not be included in this analysis such as history of MDD, treatment of MDD, cognitive impairment and social contacts [53]. There were also unmeasured risk factors for UI such as other urological conditions and reproductive history [54, 55] that could be relevant to prevalent UI and thus, HRQOL scales. In addition, we could not take into account cancer treatments due to large amounts of missing data which could have impacted the relationship between MDD, UI and HRQOL scores.

This analysis has many strengths, in particular it benefits from the large sample size and the extensive information available in the linked SEER-MHOS database and the ability to compare across cancer types as well as between cancer survivors and those without cancer. Many previous reports about MDD in cancer survivors have been limited to investigating the association in one cancer diagnosis type (eg: breast) [56, 57] or have been conducted in younger populations [58]. We were also able to adjust for relevant covariates, such as demographics, comorbid conditions and clinical characteristics that are associated with MDD and HRQOL scales.

As the population of both cancer survivors and the elderly continue to grow in the U.S., it will be important to identify health concerns that contribute to quality of life. In particular, this report highlights two conditions, MDD and UI, which are prevalent in elderly populations and the co-occurrence result in significant decrements in HRQOL. Both MDD and UI are often underdiagnosed, but treatable conditions [38, 44]. The predictors of MDD identified in this paper could be useful in screening for these conditions in older Americans with and without cancer, a population already susceptible for lower HRQOL. By better understanding the predictors of MDD and the joint impact of MDD and UI among cancer survivors, survivorship efforts can be focused to address these conditions and improve the HRQOL of this growing population.

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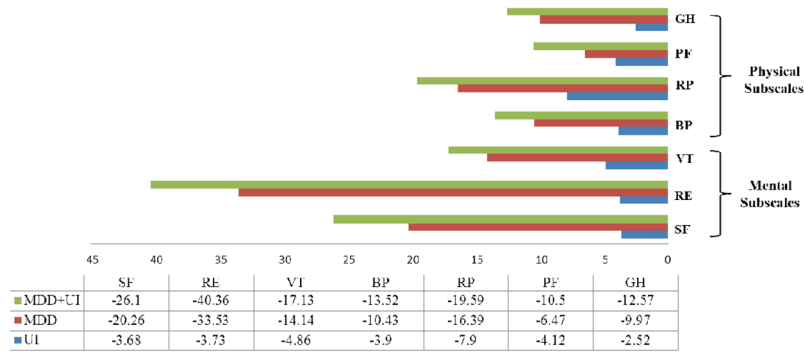


Figure 1. Decrements in physical and mental HRQOL subscales^a associated with urinary incontinence (UI), major depressive disorder (MDD) and coexisting MDD and UI among Medicare beneficiaries, adjusting for demographics and comorbid conditions^b.

^aBP= bodily pain; RP= role-physical; PF= physical functioning; GH= general health; SF= social functioning; RE= role-emotional; VI=vitality

^bDemographics and comorbidities adjustment set (index, referent): age at survey (65–74, 75–84, 85+), smoker (yes, no), race (other, white), gender (male, female), marital status (other, married), education (> high school, high school), high blood pressure, stroke, chronic lung disease (includes COPD, emphysema or asthma), gastrointestinal problems (including Crohn’s, ulcerative colitis, or inflammatory bowel disease) diabetes, difficulty completing one or more activities (getting out of chair, use of toilet, walking, eating, dressing, bathing), joint pain, cardiovascular disease

Table 1

Distribution of Demographic and Clinic Characteristics by Cancer Type^a

	Bladder	Breast	Colorectal	Endometrial/Uterine	Lung	Prostate	No cancer
	N (%)	N (%)	N (%)	N (%)	N (%)	N (%)	N (%)
Age							
65-74	240 (33.8)	1,112 (42.0)	557 (33.9)	196 (36.4)	284 (46.8)	1,202 (38.4)	147,521 (50.9)
75-84	362 (51.0)	1,118 (44.6)	793 (48.2)	258 (48.0)	276 (45.5)	1,572 (50.1)	113,178 (39.1)
85	108 (15.2)**	356 (13.4)**	295 (17.9)**	84 (15.6)**	47 (7.7)**	359 (11.5)**	29,045 (10.0)
Current smoker							
Yes	105 (15.0)**	201 (7.7)**	123 (7.6)**	18 (3.4)**	107 (17.9)**	281 (9.1)	28,662 (10.0)
Race							
White	616 (89.0)	2,122 (82.2)**	1283 (80.8)**	429 (82.3)**	484 (82.2)**	2413 (79.3)**	245,074 (86.5)
Gender							
Female	220 (31.0)**	2,649 (100)	836 (50.8)**	538 (100)	317 (52.2)**	0	175,531 (60.6)
Male	490 (69.0)	0 (0)	809 (49.2)	0	290 (47.8)	3,133 (100)	114,213 (39.4)
Marital status							
Not married	259 (38.3)**	1,441 (56.9)**	706 (44.9)	283 (55.6)**	266 (46.0)	720 (24.2)**	120,063 (43.3)
Education							
> High school	271 (38.8)**	990 (37.8)**	579 (35.8)*	208 (39.0)**	209 (35.3)	1,380 (44.8)**	93,698 (32.9)
Comorbid Conditions							
High blood pressure	444 (63.3)	1,699 (64.9)**	1031 (63.1)	346 (64.8)	355 (59.2)	1,813 (58.4)**	175,848 (61.3)
Stroke	82 (11.8)**	197 (7.5)*	173 (10.7)**	38 (7.1)	75 (12.5)**	329 (10.6)**	25,155 (8.8)
Cardiovascular disease	320 (45.4)**	772 (29.3)**	578 (35.4)	156 (29.1)*	237 (39.4)**	1,090 (35.0)	96,350 (33.4)
Chronic lung disease	114 (16.4)*	349 (13.4)	218 (13.5)	52 (9.8)*	260 (43.7)**	431 (13.9)	37,678 (13.2)
Gastrointestinal problem	37 (5.3)	138 (5.3)	161 (10.1)**	38 (7.2)**	28 (4.8)	123 (4.0)	13,323 (4.7)
Diabetes	150 (21.5)	526 (20.1)	364 (22.5)**	115 (21.7)	123 (20.5)	630 (20.3)	56,387 (19.6)

	Bladder	Breast	Colorectal	Endometrial/Uterine	Lung	Prostate	No cancer
	N (%)	N (%)	N (%)	N (%)	N (%)	N (%)	N (%)
Joint pain	361 (51.3)	1,495 (56.7)**	782 (47.8)**	306 (57.4)**	280 (46.5)*	1,434 (45.9)**	147,409 (51.1)
Difficulty completing activities ^b	357 (50.4)**	1,217 (46.2)**	764 (47.0)**	286 (53.6)**	355 (59.2)**	1,336 (43.1)	121,847 (42.3)
Time since cancer diagnosis							
<2 years	203 (28.6)	533 (20.1)	366 (22.3)	59 (11.0)	316 (52.1)	679 (21.7)	N/A
2-5 years	162 (22.8)	565 (21.3)	385 (23.4)	57 (10.6)	101 (16.6)	771 (24.6)	N/A
5 years	345 (48.6)	1551 (58.6)	894 (54.4)	422 (78.4)	190 (31.3)	1,683 (53.7)	N/A
Cancer Severity							
In Situ/Localized	516 (86.3)	1,802 (79.6)	859 (61.1)	390 (86.5)	243 (46.3)	2366 (78.3)	N/A
Regional/Distant	82 (13.7)	462 (20.4)	546 (38.9)	61 (13.5)	282 (53.7)	657 (21.7)	N/A
Urinary Incontinence (UI)							
Yes	202 (28.8)	921 (35.2)**	461 (28.5)	214 (40.2)**	177 (29.7)	1,024 (33.1)**	77,725 (27.1)
Major Depressive Disorder (MDD)							
Yes	173 (24.4)*	587 (22.2)*	362 (22.0)	127 (23.6)	207 (34.1)**	602 (19.2)	59,465 (20.5)
Coexisting UI and MDD							
Yes	71 (10.1)	283 (10.8)**	148 (9.1)	64 (12.0)**	85 (14.2)**	286 (9.2)	24,727 (8.6)
Physical Component Score (PCS)							
Mean (SD)	37.3 (11.6)	37.7 (12.0)	37.8 (11.7)	36.9 (12.0)	32.2 (11.4)***	39.7 (11.8)	39.8 (12.0)
Bodily Pain (BP)							
Mean (SD)	62.8 (28.0)	64.0 (27.7)	64.8 (27.3)	63.4 (28.2)	55.9 (30.0)***	67.9 (27.3)	66.5 (27.4)
Role-physical (RP)							
Mean (SD)	45.2 (43.4)	48.5 (31.5)	46.4 (43.7)	47.3 (42.3)	33.2 (38.9)	51.0 (43.3)	53.9 (43.8)
Physical Functioning (PF)							
Mean (SD)	54.7 (32.0)	52.4 (31.5)	52.9 (32.0)	49.0 (31.5)	40.0 (30.4)***	59.8 (31.7)	58.3 (31.8)
General Health (GH)							
Mean (SD)	51.6 (21.5)	55.7 (22.4)	53.7 (23.0)	56.5 (22.7)	40.4 (23.1)***	54.9 (22.6)	57.9 (22.4)
Social Functioning (SF)							

	Bladder	Breast	Colorectal	Endometrial/Uterine	Lung	Prostate	No cancer
	N=710	N=2,649	N=1,645	N=538	N=607	N=3,133	N=289,744
	N (%)	N (%)	N (%)	N (%)	N (%)	N (%)	N (%)
Mean (SD)	73.2 (28.2)	76.3 (27.8)	73.7 (28.5)	75.6 (27.8)	61.6 (31.9)***	76.7 (27.1)	78.5 (26.9)
Role-emotional (RE)							
Mean (SD)	69.0 (40.1)	72.1 (39.7)	67.6 (41.5)	71.9 (42.3)	57.4 (43.8)***	72.0 (40.1)	73.3 (39.5)
Vitality (VT)							
Mean (SD)	51.4 (22.4)	53.3 (22.4)	53.2 (22.4)	52.9 (21.4)	42.2 (24.6)***	55.9 (22.8)	55.9 (22.4)

^aTwo-tailed p-value for chi-squared test, comparing distribution of covariate of cancer group to no cancer group where applicable, or t-test for HRQoL scores.

^bDifficulty defined as difficulty completing one or more activities (getting out of a chair, using the toilet, walking, bathing, dressing, eating)

* p <0.05

** p <0.01

*** Significance for PCS and SF defined using a minimally important difference (MID) equal to 1/3SD

Table II

Logistic regression results for associations with major depressive disorder (MDD) compared to older Americans without cancer.

	Variable	POR	95% CI	P
Urinary Leakage				
	UI	1.68	(1.63, 1.70)	<.0001
Cancer type ^a				
	Bladder	1.21	(1.00, 1.47)	0.05
	Breast	1.00	(0.90, 1.12)	0.9
	Colorectal	1.06	(0.92, 1.21)	0.4
	Endometrial/Uterine	0.99	(0.79, 1.24)	0.9
	Lung	1.66	(1.37, 2.01)	<.0001
	Prostate	1.03	(0.93, 1.14)	0.6
Demographics ^b				
	Age (65–74)	1.19	(1.15, 1.23)	<.0001
	Age (75–84)	1.03	(0.99, 1.06)	<.0001
	Current Smoker	1.40	(1.35, 1.44)	<.0001
	Race (non-white)	1.17	(1.14, 1.20)	<.0001
	Gender (male)	0.96	(0.94, 0.98)	0.001
	Marital Status (not married)	1.47	(1.44, 1.50)	<.0001
	Education (> high school)	0.70	(0.69, 0.72)	<.0001
Comorbid conditions ^b				
	High blood pressure	1.06	(1.04, 1.09)	<.0001
	Stroke	1.45	(1.40, 1.50)	<.0001
	Chronic lung disease	1.39	(1.35, 1.42)	<.0001
	Gastrointestinal problems	1.80	(1.72, 1.87)	<.0001
	Diabetes	1.14	(1.11, 1.16)	<.0001
	Joint Pain	1.32	(1.29, 1.35)	<.0001
	Cardiovascular disease	1.31	(1.29, 1.34)	<.0001
Activities of daily living				
	Difficulty	2.55	(2.50, 2.61)	<.0001

^aNoncancer participants are referent for cancer type

^bDemographics and comorbidities adjustment set (index, referent): age at survey (65–74, 75–84, 85+), smoker (yes, no), race (other, white), gender (male, female), marital status (other, married), education (> high school, high school), high blood pressure, stroke, chronic lung disease (includes COPD, emphysema or asthma), gastrointestinal problems (including Crohn's, ulcerative colitis, or inflammatory bowel disease) diabetes, joint pain, cardiovascular disease, and difficulty completing one or more activities (getting out of chair, use of toilet, walking, eating, dressing, bathing)

Table III

Logistic regression results for associations with major depressive disorder (MDD) among cancer cases, controlling for cancer severity and time since diagnosis.

	Variable	POR	95% CI	P
Urinary Leakage				
	UI	1.62	(1.43, 1.83)	<.0001
Cancer type ^a				
	Bladder	1.20	(0.92, 1.56)	0.2
	Breast	0.95	(0.77, 1.18)	0.7
	Colorectal	1.00 (ref)	--	--
	Endometrial/Uterine	0.98	(0.72, 1.19)	0.9
	Lung	1.36	(1.04, 1.77)	0.02
	Prostate	0.96	(0.78, 1.19)	0.7
Demographics ^b				
	Age (65–74)	1.35	(1.10, 1.64)	0.001
	Age (75–84)	1.13	(0.94, 1.36)	0.7
	Current Smoker	1.37	(1.12, 1.67)	0.002
	Race (non-white)	1.04	(0.89, 1.21)	0.6
	Gender (male)	1.02	(0.82, 1.27)	0.9
	Marital Status (not married)	1.54	(1.35, 1.75)	<.0001
	Education (> high school)	0.68	(0.60, 0.80)	<.0001
Comorbid conditions ^b				
	High blood pressure	1.10	(0.97, 1.26)	0.1
	Stroke	1.38	(1.15, 1.67)	0.007
	Chronic lung disease	1.46	(1.25, 1.71)	<.0001
	Gastrointestinal problems	1.93	(1.54, 2.42)	<.0001
	Diabetes	0.96	(0.82, 1.11)	0.5
	Joint Pain	1.31	(1.15, 1.49)	<.0001
	Cardiovascular disease	1.24	(1.09, 1.41)	0.0009
Time from cancer to survey				
	<2 years	1.00 (ref)	--	--
	2–5 years	0.86	(0.72, 1.02)	0.08
	5 years	0.83	(0.72, 0.97)	0.02
Cancer Severity				
	In situ/Localized	1.00 (ref)	--	--
	Regional/Distant	1.07	(0.93, 1.22)	0.4
Activities of daily living				
	Difficulty	2.80	(2.44, 3.20)	<.0001

^a Colorectal cancer patients within 2 years of treatment, with *in situ*/localized cancer and those in the referent group of all demographic and comorbid conditions are referent group in model.

^b Demographics and comorbidities adjustment set (index, referent): age at survey (65–74, 75–84, 85+), smoker (yes, no), race (other, white), gender (male, female), marital status (other, married), education (> high school, high school), high blood pressure, stroke, chronic lung disease (includes COPD, emphysema or asthma), gastrointestinal problems (including Crohn's, ulcerative colitis, or inflammatory bowel disease) diabetes, joint pain, cardiovascular disease, and difficulty completing one or more activities (getting out of chair, use of toilet, walking, eating, dressing, bathing)

Table IVa

Adjusted mean scores for HRQOL subscales for combined major depressive disorder (MDD) and urinary incontinence (UI) among older Americans diagnosed with cancer.^a

Variable	PCS		BP		RP		PF		GH	
	Mean	95% CI	Mean	95% CI	Mean	95% CI	Mean	95% CI	Mean	95% CI
Intercept	50.19	(50.11, 50.28)	87.00	(86.92, 87.37)	88.40	(87.95, 88.81)	83.85	(83.60, 84.08)	74.88	(74.70, 75.06)
Major Depressive Disorder and Urinary Incontinence										
No MDD + UI	48.32	(48.21, 48.44)	83.10	(84.11, 84.63)	80.50	(79.95, 81.05)	79.73	(79.43, 80.04)	72.36	(72.12, 72.59)
MDD + No UI	48.44	(48.31, 48.57)	76.57*	(76.23, 76.90)	72.01*	(71.38, 72.64)	77.38	(77.02, 77.73)	64.91*	(64.63, 65.18)
MDD + UI	47.36	(47.22, 47.51)	73.48*	(74.12, 74.82)	68.81*	(68.10, 69.52)	73.35	(72.95, 73.75)	62.31*	(62.00, 62.62)
Cancer type ^b										
Bladder	49.07	(48.40, 49.74)	85.09	(83.61, 86.94)	84.37	(81.24, 87.50)	83.23	(81.41, 85.05)	71.47	(70.06, 72.88)
Breast	49.34	(49.00, 49.70)	86.79	(86.09, 87.84)	85.63	(83.95, 87.30)	81.92	(80.97, 82.87)	73.28	(72.58, 74.01)
Colorectal	49.28	(48.83, 49.72)	86.32	(85.38, 87.60)	85.90	(83.77, 88.03)	81.57	(80.37, 82.78)	72.77	(71.83, 73.70)
Endometrial/Uterine	49.85	(49.08, 50.61)	88.38	(86.59, 90.04)	89.11	(85.43, 92.78)	81.68	(79.61, 83.76)	75.28	(73.67, 76.89)
Lung	45.80*	(45.08, 46.52)	81.97	(80.33, 83.93)	77.53	(74.28, 80.78)	74.32	(72.35, 76.28)	64.20*	(62.68, 65.72)
Prostate	49.82	(49.48, 50.15)	86.88	(86.15, 87.81)	85.97	(84.40, 87.54)	83.73	(82.82, 84.63)	73.06	(72.36, 73.76)

^a PCS= Physical component score; BP= bodily pain; RP= role-physical; PF= physical functioning; GH= general health.

^b Demographics and comorbidities adjustment set (index, referent): age at survey (65–74, 75–84, 85+), smoker (yes, no), race (other, white), gender (male, female), marital status (other, married), education (> high school, high school), high blood pressure, stroke, chronic lung disease (includes COPD, emphysema or asthma), gastrointestinal problems (including Crohn's, ulcerative colitis, or inflammatory bowel disease) diabetes, difficulty completing one or more activities (getting out of chair, use of toilet, walking, eating, dressing, bathing), joint pain, cardiovascular disease

^c Noncancer participants are referent for cancer type

* Clinical significance for HRQOL scales defined using a minimally important difference (MID) equal to 1/5SD

Table IVb

Adjusted mean scores for HRQOL subscales^a for combined major depressive disorder (MDD) and urinary incontinence (UI) among older Americans diagnosed with cancer^b

Variable	SF		RE		VT		
	Mean	95% CI	Mean	95% CI	Mean	95% CI	
Intercept	97.60	(97.39, 97.82)	95.69	(95.28, 96.10)	72.90	(72.73, 73.08)	
No MDD + UI	93.92	(93.64, 94.20)	91.96	(91.42, 92.49)	68.04	(67.81, 68.27)	
MDD + No UI	77.34*	(77.02, 77.67)	62.16*	(61.55, 62.77)	58.76*	(58.49, 59.02)	
MDD + UI	71.50*	(71.14, 71.87)	55.33*	(54.65, 56.02)	55.77*	(55.47, 56.07)	
Cancer type ^c	Bladder	95.88	(94.16, 97.60)	95.20	(92.19, 98.21)	71.50	(70.13, 72.86)
	Breast	96.82	(95.93, 97.70)	96.16	(94.55, 97.77)	71.26	(70.54, 71.96)
	Colorectal	95.29	(94.16, 96.41)	93.71	(91.65, 95.77)	71.92	(71.02, 72.83)
	Endometrial/Uterine	98.78	(96.85, 100.70)	99.11	(95.56, 102.65)	73.12	(71.56, 74.68)
	Lung	88.06*	(86.14, 89.98)	88.85	(85.71, 91.98)	65.33*	(63.85, 66.80)
	Prostate	96.85	(96.01, 97.69)	94.65	(93.14, 96.17)	73.14	(72.47, 73.83)

^aSF= social functioning; RE= role-emotional; VT=vitality.

^bDemographics and comorbidities adjustment set (index, referent): age at survey (65–74, 75–84, 85+), smoker (yes, no), race (other, white), gender (male, female), marital status (other, married), education (> high school, high school), high blood pressure, stroke, chronic lung disease (includes COPD, emphysema or asthma), gastrointestinal problems (including Crone's, ulcerative colitis, or inflammatory bowel disease) diabetes, difficulty completing one or more activities (getting out of chair, use of toilet, walking, eating, dressing, bathing), joint pain, cardiovascular disease

^cNoncancer participants are referent for cancer type

* Clinical significance for HRQOL scales defined using a minimally important difference (MID) equal to 1/3SD