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Predictors of Marriage and Divorce in Adult Survivors of Childhood Cancers: A Report from the Childhood Cancer Survivor Study

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Abstract

Background/Objective—Adult survivors of childhood cancer can have altered social functioning. We sought to identify factors that predict marriage and divorce outcomes in this growing population.

Methods—Retrospective cohort study of 8,928 ≥ five-year adult survivors of childhood malignancy and 2,879 random sibling controls participating in the Childhood Cancer Survivor Study. Marital status, current health, psychological status, and neurocognitive functioning were determined from surveys and validated instruments.

Results—Survivors were more likely to be never-married than siblings (relative risk (RR) = 1.21; 95% confidence interval (CI) 1.15–1.26) and the U.S. population (RR=1.25; 95% CI= 1.21 – 1.29), after adjusting for age, gender, and race. Patients with central nervous system (CNS) tumors were at greatest risk for not marrying (RR=1.50; 95% CI= 1.41–1.59). Married survivors divorced at frequencies similar to controls. In multivariable regression analysis, non-marriage was most associated with cranial radiation (RR=1.15; 95% CI=1.02–1.31 for >2400 centigray). In analysis of neurobehavioral functioning, non-marriage was associated with worse task efficiency (RR=1.27;

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95% CI=1.20–1.35), but not with emotional distress, or problems with emotional regulation, memory, or organization. Physical conditions predictive of non-marriage included short stature (RR=1.27; 95% CI=1.20–1.34) and poor physical function (RR=1.08; 95% CI=1.00–1.18). Structural equation modeling suggested that cranial radiation influenced marriage status through short stature, cognitive problems, and poor physical function.

Conclusions—Childhood cancer survivors married at lower frequencies compared to peers. Patients with CNS tumors, cranial radiation, impaired processing efficiency, and short stature were more likely to never marry. Divorce patterns in survivors were similar to peers.

Keywords

Survivorship; Cancer; Predictors; Marriage; Divorce

Introduction

Approximately 80% of children with cancer will survive five or more years from diagnosis of their disease (1). The impact of cancer treatment on physical health, during the first several decades following diagnosis and treatment, has been well-characterized, and survivors are known to be at increased risk for second neoplasms (2–4), cardiovascular disease (5,6), endocrine dysfunction (6), and early death (7,8). Several psychological sequelae have been described as well, with sub-groups of survivors reporting depression (9–11), anxiety (10), and post-traumatic stress symptoms (12–14).

In addition to physical and mental health, attention must be given to the overall functioning of survivors in society. For instance, survivors have been shown to experience lower educational attainment (15), higher rates of unemployment (16,17), and difficulty obtaining health insurance (16,18). Although the institution of marriage has undergone many changes in modern times, it represents another social outcome that can be used to gauge the adaptation of survivors to life after cancer since it represents an aspiration for the majority of young adults in today's society (19). Relationships are challenging for all adults, but may be especially difficult for survivors, who struggle with the burdens of past disease. In one study, 29% of childhood cancer survivors cited disability or prior illness as a barrier to marriage (20). Uncertainty about future health may also impact survivor relationships (21–23).

The available literature on marriage outcomes after childhood cancer is characterized by inconsistent findings, likely resulting from the limited size and/or distinct composition of the study populations (24–29). Moreover, many of the earlier reports did not assess the underlying causes of observed patterns or have appropriate comparisons to non-cancer populations. Most recently in 2007, Frobisher et al. reported reduced marriage frequencies in 9,954 British childhood cancer survivors diagnosed from 1940 to 1991 compared to those expected from the general population and concluded that survivors were less likely to get married (30). While this study was large, there was limited measurement of emotional and cognitive functioning as potential mediators of decreased marriage frequencies. Also, a key marital outcome, divorce, was not examined.

The Childhood Cancer Survivor Study (CCSS) provides a unique opportunity to add to our understanding of marriage outcomes because of the size and characterization of the cohort, as well as the availability of a sibling comparison group. In this paper, we 1) describe marriage and divorce frequencies in childhood cancer survivors from the CCSS cohort, with comparison to both a sibling cohort and data from the U.S. Census; and 2) identify patient and treatment factors that predict marital status, including psychosocial distress and neurocognitive impairment.

Methods

Study Population

CCSS Cohort—The CCSS is a 26-institution retrospective cohort of survivors of childhood cancer designed to study the late effects of cancer therapy. Eligibility criteria included: 1) diagnosis of leukemia, central nervous system (CNS) tumor, Hodgkin's lymphoma, non-Hodgkin Disease (HD), Wilms tumor, neuroblastoma, soft tissue sarcoma, or bone tumor; 2) diagnosis and initial treatment at a participating center; 3) diagnosis between January 1, 1970, and December 31, 1986; 4) age <21 years at diagnosis; and 5) survival of ≥ 5 years after diagnosis. The methodology has been previously described (31) and study documents are available.¹ Each participating center's institutional review board reviewed and approved the CCSS protocol and contact documents.

Starting in August, 1994, participants completed an extensive baseline questionnaire which included demographic characteristics marital status, and health history. Two subsequent surveys were administered (2000 Survey: beginning in May, 2000, and 2003 Survey: beginning in November, 2002) to obtain updated information. Trained data abstractors reviewed participants' medical records for detailed cancer diagnosis and treatment information.

Of the 20,691 patients eligible for participation, 14,363 completed the baseline questionnaire; 3,058 were lost to follow-up; and 3,205 refused participation. Of the 14,363 initial participants, 10,366 completed the first follow-up questionnaire (2000 Survey), and 9,308 completed the second follow-up questionnaire (2003 Survey). Cases were excluded from the current analysis if they were younger than 15 years ($n=3$) or if they were married prior to diagnosis of malignancy ($n=75$), yielding 9,230 individuals, of whom 8,928 had known marital status.

Siblings—A random sample of participating survivors ($n= 6,005$) was asked to contact their sibling closest in age for participation in the study. Of these, 3,839 siblings completed the baseline (enrollment) survey, 2,540 completed the 2000 Survey, and 2,951 completed the 2003 Survey. For the current analysis, siblings were restricted to those subjects age 15 years and older, alive, and in follow-up as of 2003 Survey, resulting in 2,789 siblings with known marital status. See Table 1 for case characteristics compared to siblings.

U.S. Population—Data on marital status of the U.S. population were obtained from the 2002 Current Population Survey (CPS), as issued by the Bureau of Census. The report includes marital status, stratified by gender, current age (15 years and older), education and race.²

Measures

On each CCSS survey questionnaire, participants categorized themselves as “single/never married,” “married,” “living as married,” “widowed,” “divorced,” or “separated/no longer living as married.” Responses were grouped into three outcomes: “never-married,” “currently-divorced,” and “ever-divorced.” “Never-married” was available from the 2003 Survey. Subjects responding “divorced” or “separated” on 2003 Survey were defined as “currently-divorced,” consistent with past studies (24,32). Cases who reported “divorced” or “separated” on any survey were classified as “ever-divorced.” It is possible that an individual responding “married” on consecutive surveys may in fact be divorced and remarried. We anticipate that the number of divorce cases missed in this manner will be negligible, given a median time of 5 years between the baseline and 2000 Survey, and about 2 years between the 2000 and 2003 surveys.

¹<http://www.stjude.org/ccss>.

²<http://www.census.gov/population/www/socdemo/hh-fam/cps2002.html>.

In the 2002 CPS, “never-married” and “currently-divorced” were clearly defined; “ever-divorced” was not available. Also, the CPS did not include a “living with partner as married” category. Therefore, when drawing comparison to the general population, cohort members in the “living with partner as married” category as of 2003 Survey were considered “never-married.”

Data from the 2003 Survey were used for variables that change with time including education, income, employment status, and height. Diminished height was defined as height below the tenth percentile for age, gender, and ethnicity, as reported by the Centers for Disease Control and Prevention (CDC).³ Perceived infertility was defined as “yes” to the question, “Has a doctor ever told you that you might have trouble having children?”

Psychological health was evaluated on the baseline and the 2003 Survey with the Brief Symptom Inventory-18 (BSI-18), an 18-item checklist that measures symptoms of anxiety, depression, and somatic distress (33). Responses were scored to generate a Global Severity Index (GSI) score (34). In our analysis, subjects with GSI elevations ≥ 50 on either of two BSI-18 administrations were classified as having a positive history of psychological distress, consistent with a previous validation study in cancer survivors by Recklitis et al.(35).

Neurocognitive functioning, including executive skills, was evaluated with the Childhood Cancer Survivorship Study Neurocognitive Questionnaire (CCSS-NCQ), a 25-item instrument that is predominantly a subset of items from an early investigational version of the Behavior Rating Inventory of Executive Functioning-Adult version. Krull et al identified four domains that demonstrated good internal consistency: task efficiency, emotional regulation, organization, and memory skills.(36) Subjects were classified as “high risk” for neurocognitive dysfunction if the response on any of the questions for the respective factor was “often a problem” consistent with validation studies of this instrument.

Analyses

Frequencies of “never-married” and “currently-divorced” were described in CCSS cases and compared to frequencies for siblings and the U.S. population (as of the 2002 CPS), overall and in a stratified fashion, by age and gender. The “currently-divorced” proportion was calculated as the number “divorced” or “separated,” divided by the total number “married,” “widowed,” “divorced,” or “separated.” Likelihood ratio tests were used to determine the statistical significance of differences between groups. Survivors were compared to U.S. population data and to the sibling comparison group. Generalized estimating equation formulations of the model and significance tests were utilized to account for the intra-family correlation between survivors and siblings (37)

Among survivors, case-case comparisons were conducted with respect to the outcomes, “never-married” and “ever-divorced.” The analysis of “ever-divorced” was restricted to those subjects who had been married at least once and who had reported marital status on all three surveys. Log-binomial regression models were used to evaluate associations between explanatory variables and each outcome. These models allow direct calculation of age-adjusted RRs with 95% confidence intervals to compare the probability of outcomes between survivor sub-groups and were selected over logistic regression due to the high prevalence of the outcome (38). Multivariable regression models, including factors marginally significant in the unadjusted analysis ($p < 0.2$), were created to determine the independent role of each variable, adjusted for age at diagnosis, gender, and educational status. Potential confounders and interactions were also evaluated

³<http://www.cdc.gov/nchs/data/ad/ad361.pdf>

Structural equation models of the observed data (weighted least-square parameter estimates [delta parameterization]) were analyzed using Mplus 5.2 software (39). All variables were directly observed measures; there were no latent variables. Never married (N=2616) and Ever Married (N=3924) sub-samples at the Follow-up 2 survey with complete data comprised the final sample for the SEM analysis. We chose to use samples with complete data rather than to use data imputation in order to avoid potentially distorting coefficients of association and correlation relating variables (40). The best-fitting model was determined according to the following criteria: 1) conceptually sound; 2) statistically significant parameter estimates (PE) that represent the strength of the path between two variables (read as standardized regression coefficients); 3) meets the established SEM fit criteria (non-significant χ^2 statistic ($P > 0.05$); 4) root mean square error of approximation (RMSEA) ≤ 0.05 ; 5) weighted root mean square residual (WRMR) < 1.0 (41,42)]; 6) comparative fit index (CFI) and Tucker Lewis index (TLI) ≥ 0.90 (43); and 7) the highest percentage of explained variance for the outcome.

Results

Marital Status of CCSS Cohort at Last Contact

At last contact, 42.4 % (n = 3,783) of survivors were currently married, 7.3 % (n = 654) were divorced or separated, 0.2 % (n = 20) were widowed, and 46.4 % (n = 4,141) had never been married. Of those never-married (n=3698), 90% were living as single and 10% lived with a partner outside of marriage.

Comparison of Survivor Marital Status with Siblings and the U.S. Population

Survivors were significantly more likely to never have married than siblings (RR= 1.21; 95% CI 1.15–1.26) and the U.S. population. (RR=1.25; 95% CI= 1.21 – 1.29), after adjusting for age, gender, and race (See Table 2 and Table 3). The trend was apparent across all age groups 25 years old and older. It was particularly marked for those in the 35–44 year and 45+ year age groups, where survivors were 1.90 (95% CI 1.55 – 2.32) and 2.35 (95% CI 1.29 – 4.28) times more likely than siblings to be never-married, after adjusting for gender and race (data not shown). Cases with a history of CNS tumor (RR=1.49; 95% CI= 1.40 – 1.58) and leukemia (RR=1.19; 95% CI=1.12 – 1.25) had the greatest likelihood of never marrying. Upon further stratification, the probability of never marrying remained elevated in leukemia patients who received cranial radiation (RR 1.25; 95% CI= 1.18–1.32), but not in those treated with chemotherapy only (RR=1.03; 95% CI=0.96–1.10).

Survivors divorced at similar frequencies to siblings (RR= 1.08; 95% CI= 0.96–1.21) and to population controls (RR = 0.96; 95% CI 0.89 – 1.03, p=0.23). No cancer diagnosis group had an elevated risk of divorce (Table 3). No statistically significant differences in divorce frequencies were observed across age or gender groups (data not shown).

Predictors of Never-Married Status in Survivors

Univariate analysis adjusted for gender and age at last contact and gender indicated that age < 13 years at diagnosis (RR=1.52; 1.34–1.72) and cranial radiation > 2400 centigray (RR=1.28 compared to no cranial radiation; 95% CI=1.14–1.43) were the strongest predictors of non-marriage among treatment factors (Table 4). The following medical and neuropsychological conditions were significantly associated with never marrying (Table 4): short stature, history of tumor recurrence, poor self-reported physical functioning, emotional distress, problems with task efficiency, problems with organization, and problems with memory. Report of a perceived fertility problem was associated with a lower likelihood of not marrying (RR=0.91; 95% CI= 0.87–0.95).

Table 5–Table 7 displays the results of three separate multivariable models (that all included gender, age at last contact, age at diagnosis, and educational attainment) divided into disease and treatment factors, neurobehavioral functioning, and physical functioning factors. Significant disease and treatment predictors of non-marriage included cranial radiation >2400 centigray (RR=1.15 compared to no radiation; 95% CI=1.02–1.31) and history of recurrence (RR=1.10; 95% CI=1.00–1.20). Impaired task efficiency was the only neurobehavioral condition significantly associated with not being married in adjusted analysis (RR= 1.27; 95% =1.20–1.35). Problems with emotional regulation were associated with a greater likelihood of getting married. In terms of physical conditions, short stature (RR=1.27; 95% CI=1.2–1.34) and poor self-reported physical functioning (RR=1.08; 95% CI=1.00–1.18) were associated with not ever marrying. Perceived fertility problem was not included in the adjusted model because the direction of the association suggested that fertility status likely was determined after marriage.

Male gender and younger age at diagnosis were consistently associated with greater likelihood of not getting married, in adjusted analyses. No differences were noted upon further stratification of the significant factors identified in multivariable analysis by gender or cranial radiation.

Table 8 and the Figure collectively present the results of the structural equation modeling. Table 8 provides a description of all significant variables and their contribution to the model, including the estimated regression coefficients (EST) for each parameter, the standard error of the parameter estimates (SE), the coefficient divided by the standard error (EST/SE, or z-score), the standardized coefficients (STDYX), and the p-value for the path between the two variables. For binary dependent variables, the regression coefficients produced are logistic regression coefficients. Figure 1 represents a simplified graphic version of the complete SEM results. A well fitting model ($\chi^2=21.91$, $df=14$, $P=0.08$; CFI=0.999; TLI=0.998; RMSEA=0.009; WRMR = 0.557) explained 45.6% of the variance in survivors' never having been married. The strongest predictor of never having married, based on the weight of the standardized coefficients, was younger current age followed by short stature, poor task efficiency, male gender, history of CNS radiation, better memory, poor physical function, and poor emotional functioning.

History of CNS radiation was an indirect influence on never having married through 1) short stature ($P<0.0001$), 2) poor memory ($P<0.0001$), 3) poor physical function ($P<0.0001$); and 4) poor task efficiency ($P<0.001$). History of CNS radiation also was a direct influence on never having married, presumably through factors that we did not measure in this study. Short stature was an indirect influence on never having married through poor task efficiency ($P<0.0001$), and poor physical function ($P<0.0001$). The indirect impact of poor task efficiency on never having married through by poor memory; the indirect impact of poor physical function was through by poor task efficiency.

Predictors of Ever-Divorced Status in Survivors

Among ever-married survivors, after adjusting for gender and age at last contact, factors found to be significantly associated with history of divorce were poor physical functioning (RR=1.52; 95% CI=1.30–1.78), perceived fertility problem (RR=1.14; 95% CI=1.01–1.27), emotional distress (RR=1.40; 95% CI= 1.25–1.57), problems with task efficiency (RR=1.37; 95% CI=1.20–1.58), impaired working memory (RR=1.34; 95% CI=1.16–1.56), and problems with emotional regulation (RR=1.32; 1.15–1.52) as displayed in Table 4. No significant treatment factor was identified.

Multivariable models were examined for divorce (Table 5–Table 7). An age younger than 13 years at diagnosis (RR=1.28; 95% CI=1.03–1.60), emotional distress (RR=1.33; 95%

CI=1.15–1.54), and self-report of poor physical functioning (RR=1.40; 95% CI=1.18–1.67) were independently predictive of divorce. Interactions were examined; there were no difference in the association between risk factors and divorce status between males and females and by cranial radiation status.

Discussion

In this large, multi-site cohort of adult survivors of childhood cancer, we concluded that survivors were 1.21 times more likely to be unmarried than the sibling comparison group and 1.25 times more likely to be unmarried than the U.S. Census population, after adjusting for age, gender, and race. Our risk estimates are similar to that of the 2007 report by Frobisher et al. based on the 9,954 member British Cancer Survivor Study (BCCSS) (30). Younger age at diagnosis and history of cranial radiation were the most important predictors of never getting married among cases. From structural equation modeling, we found that cranial radiation exposure was an indirect influence on never having married mediated by short stature, impaired memory, worse processing speed, and poor physical function. Emotional distress among survivors is a direct influence of never getting married, separate from cranial radiation exposure. Our other major finding was that divorce patterns among childhood cancer survivors are similar to that of the general population and a sibling comparison group. This reassuring conclusion is contrary to an older report by Byrne et al. in 1989 (24). Ours is the largest study to our knowledge that examines divorce outcomes.

Our results should be further compared and contrasted with that of the other large, recent cohort study by Frobisher et al. in the BCSS. The BCCSS study only compared cases to population data and no summary relative risk statistic was reported. However, marriage frequencies stratified by age and gender from the Frobisher publication suggested that survivors were 1.1–1.6 times more likely to be unmarried. These estimates are similar to our own verified with both sibling and general population comparison groups. Both the CCSS and the BCSS studies identified males, history of CNS tumor, exposure to CNS radiation, and poor physical function as predictors of non-marriage.

Our CCSS study of marriage was unique in that we also included standardized measures specific to emotional and cognitive functioning to understand why certain patient groups were less likely to marry. In the CCSS cohort, structural equation modeling helped to elucidate that cranial radiation *indirectly* influenced never getting married through worse cognitive processing difficulties and short stature, as well as poor physical function. In the childhood cancer survivor population, short stature is usually due to decreased pituitary function as a result of CNS radiation. In the general population, diminished height is known to be associated with lower marriage rates (44), and bachelors are significantly shorter (45). In 1996, a meta-analytic review concluded that females are more romantically attracted to taller males (46). In a more recent large study of responses to personal advertisements, males with higher education and taller height had significantly more responses (47). Pawlowski speculates that “male height is an important trait on the mate market” because it is an indicator of reproductive potential, while education and intelligence are proxies for economic status (47). There is evidence that taller males father more children (45) and are perceived as healthier (48).

Structural equation analysis suggests that cranial radiation also has a *direct* influence on non-marriage, presumably mediated through some factor that we did not measure in this study. Future studies should examine the potential role of factors such as social intelligence, attractiveness to the other sex, altered sexual maturation, and libido. Emotional distress and male gender were other factors directly associated with never getting married.

Cranial radiation has been associated with social difficulties in past studies. Pui et al. found cranial radiation to predict non-marriage in female survivors of acute lymphoblastic leukemia (ALL) (49). In a study of adolescent survivors, Barrera et al. concluded that those treated with cranial radiation were less likely to have close friends than survivors treated without cranial radiation (50). Thus, it seems that the negative effects of cranial radiation on social integration begin at an early age and persist into adulthood.

The current study has some methodological characteristics that should be considered in the interpretation of the results. Due to the time elapsed between surveys and the nature of the question about marital status, it is possible that some cases of divorce were missed. As a result, we may have under-estimated the risk of being ever-divorced. The CCSS participants were diagnosed between 1970 and 1986 in an earlier era, and thus may not be directly generalizable to more recently treated cohorts of pediatric cancer survivors. Finally, although the size of the CCSS cohort is a strength, it also limits the nature of contact with participants to standardized questionnaires. Thus, while we can state that survivors marry less frequently than controls of similar age and gender, we do not have data directly relating to the thoughts, desires, and motivations underlying this behavior.

The CCSS is a valuable resource for survivorship studies because of the multi-site design, large sample size, and high participation rates (51)]. For the baseline CCSS survey, 69% of the total eligible population participated (15% could not be located and 15% declined participation). Participation rates on the follow-up surveys have ranged from 77–81%. Comparisons of available demographic and cancer-related characteristics between participants and non-participants at the initial baseline questionnaire showed that the only significant difference between these groups was vital status. That is, the next-of-kin relatives of patients who died more than 5 years after diagnosis were less likely to have participated than patients who were still alive. Comparisons have also been done between participants and non-participant at subsequent questionnaires (52)]. While differences are moderate in size (<10% differences), the study retains more female, white race, college-educated, higher-income, and older participants. In our current analysis, we adjust for gender, race, age, and socio-economic status.

Marriage and divorce patterns are objective measures that can be used to gauge social integration and success of intimate relationships among childhood cancer survivors. While it can be debated whether marriage is a desirable outcome, marriage is generally an expected developmental goal in our society to the extent that most adults in the U.S. are married by the age of 30 years. Our large cohort study confirms that childhood cancer survivors are less likely to be married compared to their non-cancer peers. Among survivors, patients with CNS tumors or a history of cranial radiation were most likely not to marry. Cranial radiation influenced marriage status through short stature, cognitive processing difficulties, and poor physical function. Except for those with reduced physical function, there was no increased risk of divorce among survivors who did marry. Studies such as ours are important to understand how the growing population of childhood cancer survivors functions in our society. Separate analyses are underway in the CCSS to better understand factors that contribute to other adult benchmarks such as living independently, achieving higher education, and personal income.

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APPENDIX

The **Childhood Cancer Survivor Study (CCSS)** is a collaborative, multi-institutional project, funded as a resource by the National Cancer Institute, of individuals who survived five or more years after diagnosis of childhood cancer. **CCSS** is a retrospectively ascertained cohort of 20,346 childhood cancer survivors diagnosed before age 21 between 1970 and 1986 and approximately 4,000 siblings of survivors, who serve as a control group. The cohort was assembled through the efforts of 26 participating clinical research centers in the United States and Canada. The study is currently funded by a U24 resource grant (NCI grant # U24 CA55727) awarded to St. Jude Children's Research Hospital. Currently, we are in the process of expanding the cohort to include an additional 14,000 childhood cancer survivors diagnosed before age 21 between 1987 and 1999. For information on how to access and utilize the CCSS resource, visit www.stjude.org/ccss

APPENDIX

CCSS Institutions and Investigators

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CCSS Institutions and Investigators

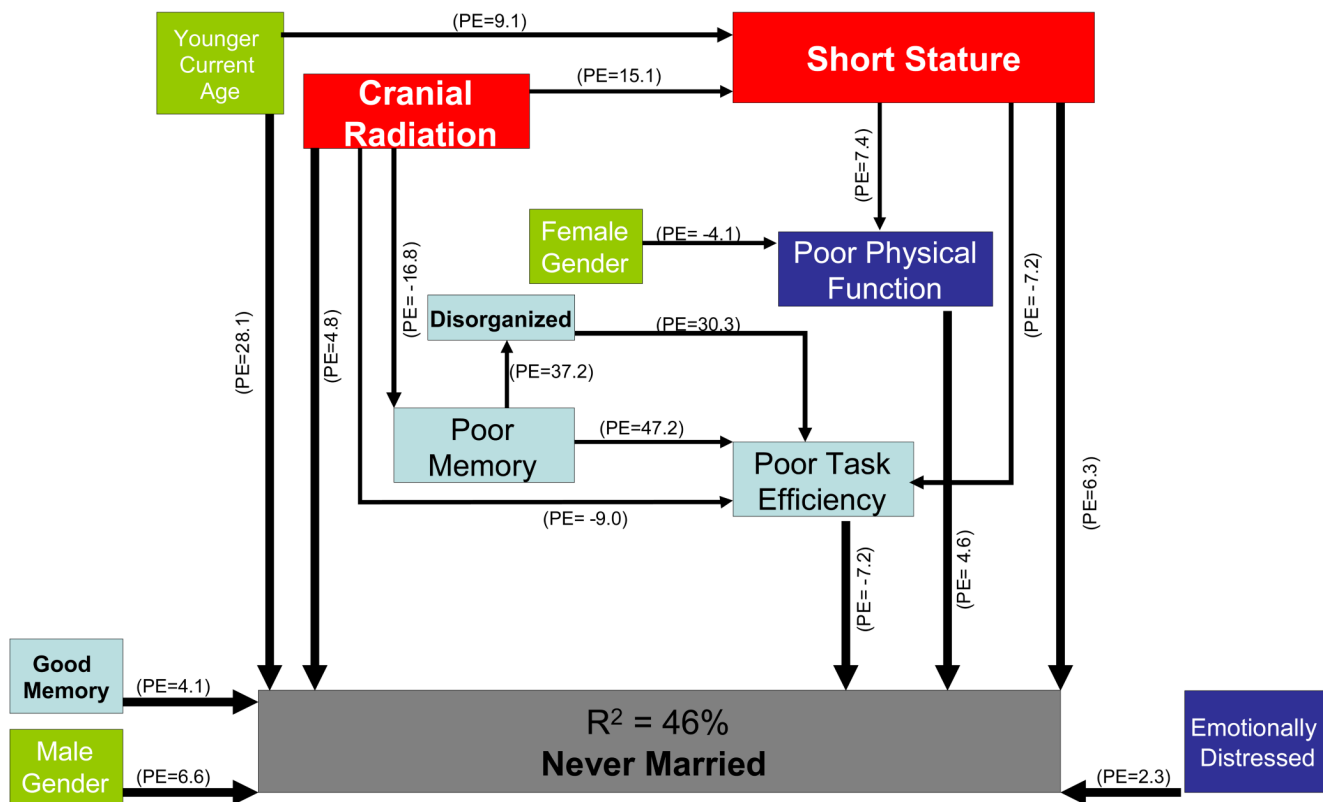
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[‡] Member CCSS Steering Committee

[#] Project Principal Investigator (U24 CA55727)



PE= standardized parameter estimate (EST/SE)

Figure 1. Graphic representation of structural equation modeling of predictors of never-married status in CCSS cases

Table 1

Characteristics of Childhood Cancer Survivor Study (CCSS) Cases and Siblings

	CCSS Cohort		Sibling Cohort		p (chi-sq)
	N	%	N	%	
Diagnosis					
Leukemia	3159	84.2			N/A*
Central Nervous Tumor	1171	12.7			
Hodgkin Disease	1151	12.5			
Non-Hodgkin Lymphoma	697	7.6			
Kidney (Wilms)	868	9.4			
Neuroblastoma	629	6.8			
Soft tissue sarcoma	809	8.8			
Bone cancer	746	8.1			
Sex					
Male	4695	50.9	1315	46.1	<.0001
Female	4535	49.1	1537	53.9	
Age at last contact (years)					
15-19	338	3.7	147	5.2	<.0001
20-24	1742	18.9	415	14.6	
25-29	1982	21.5	522	18.3	
30-34	2131	23.1	531	18.6	
35-39	1640	17.8	510	17.9	
40+	1397	15.1	727	25.5	
Race/ethnicity					
White non-Hispanic	7895	85.9	2532	91.9	<.0001
Other	1299	14.1	222	8.1	
Education					
Did not complete high school	458	5.0	122	4.3	<.0001
Completed high school	4814	52.7	1313	46.2	
College graduate	3859	42.3	1408	49.5	
Household income					
< \$40,000	2905	32.4	637	22.9	<.0001
>= \$40,000	6062	67.6	2149	77.1	
Personal income					
< \$40,000	6625	75.0	1574	63.1	<.0001
>= \$40,000	2205	25.0	921	36.9	
Employment status					
Unemployed	428	4.7	69	2.4	<.0001
Disabled	708	7.8	37	1.3	
Employed or retired	7895	87.4	2718	96.2	

* N/A= Not applicable

Table 2

Frequencies of Never-Married Status – Comparison of Survivors with Siblings and the U.S. Census Population, adjusted for race

	Survivors		Siblings		U.S. Census	
	N	%	N	%	N	%
All ages 18–54 years						
Total	3856	46.1	821	31.7 [†]		32.4 [*]
Male	2083	49.2	421	35.3 [†]		36.5 [*]
Female	1773	43.0	400	28.6 [†]		24.7 [*]
Age 18–24 years						
Total	1444	85.3	371	83.7		84.4
Male	726	89.9	183	88.4		86.5
Female	718	81.1	188	79.7		79.7
Age 25–34 years						
Total	1841	48.2	338	34.7 [†]		37.2 [*]
Male	1028	52.4	186	39.9 [†]		41.8 [*]
Female	813	43.7	152	29.9 [†]		27.5 [*]
Age 35–44 years						
Total	524	21.5	99	11.0 [†]		15.9 [*]
Male	304	24.1	46	11.2 [†]		19.4 [*]
Female	220	18.7	53	10.9 [†]		9.7 [*]
Age 45–54 years						
Total	47	11.5	13	4.8 [†]		10.0 [*]
Male	25	12.0	6	5.6		11.9
Female	22	11.1	7	4.3 [†]		6.9 [*]

[†] Indicates p < 0.05, survivors vs. siblings

^{*} Indicates p < 0.05, survivors vs. U.S. Census

Table 3

Relative risk of Never-Married and Ever-Divorced in CCSS cases compared to sibling comparison group, overall and stratified by cancer diagnosis, adjusted by age at evaluation, gender, and race (please note, there is no referent within the diagnostic category: every row is compared to the sibling group)

	Never-married			Ever Divorced				
	N	%	RR (95%CI)	p-value	N	%	RR (95%CI)	p-value
All Cancers	3698	41.4	1.21 (1.15–1.26)	<0.0001	981	21.4	1.08 (0.96–1.21)	0.20
Leukemia	1508	49.2	1.19 (1.12–1.25)	<0.0001	282	20.8	1.09 (0.94–1.27)	0.26
Central Nervous Tumor	698	62.5	1.49 (1.40–1.58)	<0.0001	78	21.5	1.07 (0.86–1.34)	0.553
Non-Hodgkin Lymphoma	204	29.6	1.09 (0.98–1.22)	0.13	104	24.4	1.19 (0.97–1.46)	0.10
Kidney (Wilms)	377	46.2	0.97 (0.90–1.04)	0.41	56	14.4	0.80 (0.60–1.07)	0.14
Neuroblastoma	333	59.9	1.14 (1.06–1.22)	0.0002	38	19.8	1.17 (0.86–1.60)	0.32
Soft tissue sarcoma	270	33.9	1.11 (1.02–1.21)	0.02	87	19.4	0.93 (0.75–1.15)	0.50
Bone cancer	149	20.1	1.02 (0.87–1.19)	0.02	120	22.6	1.03 (0.85–1.24)	0.78
Hodgkin Disease	159	13.9	1.05 (0.91–1.21)	0.54	216	24.4	1.13 (0.96–1.33)	0.13
Sibling Comparison Group	776	27.8		Ref	N/A	304	19.9	Ref

Univariate analysis of the association of patient factors, cancer treatment, and medical conditions with marital status, adjusted for gender and age at last contact

Table 4

Characteristic	Never-married N	%	RR (95%CI)	p-value	Ever-divorced N	%	RR (95%CI)	p-value
Age at diagnosis (years)								
<13	3338	50.6	1.52 (1.34–1.72)	<.0001	580	20.5	1.12 (0.98–1.28)	0.10
13–20	360	15.4	1.00		401	22.7	1.00	
Gender								
Female	1692	38.5	1.00		524	22.0	1.00	
Male	2006	44.2	1.16 (1.12–1.21)	<.0001	457	20.7	0.92 (0.82–1.03)	0.13
Diagnosis								
Leukemia	1508	49.2	1.87 (1.60–2.17)	<.0001	282	20.8	0.97 (0.82–1.15)	0.73
Central Nervous Tumor	698	62.5	2.22 (1.90–2.59)	<.0001	78	21.5	0.96 (0.76–1.20)	0.70
Non-Hodgkin Lymphoma	204	29.6	1.56 (1.31–1.86)	<.0001	104	24.4	1.07 (0.87–1.31)	0.55
Kidney (Wilms)	377	46.2	1.59 (1.35–1.88)	<.0001	56	14.4	0.73 (0.55–0.97)	0.03
Neuroblastoma	333	59.9	1.96 (1.67–2.30)	<.0001	38	19.8	1.00 (0.73–1.38)	0.99
Soft tissue sarcoma	270	33.9	1.69 (1.42–2.00)	<.0001	87	19.4	0.84 (0.68–1.05)	0.14
Bone cancer	149	20.1	1.33 (1.09–1.62)	0.005	120	22.6	0.94 (0.77–1.14)	0.54
Hodgkin Disease	159	13.9	1.00		216	24.4	1.00	
Cranial radiation								
>2400	89	54.6	1.28 (1.14–1.43)	<.0001	9	14.5	0.68 (0.37–1.26)	0.23
>0 and ≤2400	726	49.6	1.10 (1.04–1.17)	0.001	128	19.5	1.01 (0.83–1.22)	0.92
0	1150	42.5	1.00		273	19.8	1.00	
Stem cell Transplant								
Yes	37	44.0	0.96 (0.81–1.15)	0.70	7	16.7	0.81 (0.41–1.59)	0.53
No	3355	41.6	1.00		867	20.8	1.00	
Treatment duration								
≥2 years	1884	45.1	1.06 (1.01–1.10)	0.01	409	20.1	0.98 (0.87–1.10)	0.69
<2 years	1386	37.6	1.00		434	21.2	1.00	
Perceived fertility problem								
Yes	1208	33.8	0.91 (0.87–0.95)	<.0001	481	23.2	1.14 (1.01–1.27)	0.03
No	2262	45.8	1.00		466	19.7	1.00	
Short stature								
Yes	978	57.2	1.20 (1.16–1.24)	<.0001	150	23.7	1.16 (0.99–1.35)	0.07
No	2571	37.0	1.00		810	20.9	1.00	
Subsequent Malignant Neoplasm								
Yes	220	27.8	1.05 (0.97–1.15)	0.24	114	21.6	0.91 (0.77–1.09)	0.31
No	3478	42.7	1.00		867	21.3	1.00	
Recurrence								
Yes	415	45.8	1.11 (1.06–1.17)	<.0001	93	21.7	0.98 (0.82–1.19)	0.87
No	3283	40.9	1.00		888	21.3	1.00	
Poor physical function (SF-36 T-score <40)								
Yes	375	42.9	1.19 (1.14–1.24)	<.0001	141	31.7	1.52 (1.30–1.78)	<.0001
No	2717	40.9	1.00		687	19.6	1.00	

Characteristic	Never-married		Ever-divorced		RR (95%CI)	p-value
	N	%	N	%		
Emotional Distress (GSI T-score \geq 50)						
Yes	1613	39.9	544	25.3	1.40 (1.25–1.57)	<.0001
No	1513	36.8	412	18.0	1.00	
NCQ: problems with task efficiency						
Yes	935	49.4	220	25.4	1.37 (1.20–1.58)	<.0001
No	1637	35.6	500	18.5	1.00	
NCQ: problems with organization						
Yes	550	44.4	141	22.7	1.13 (0.96–1.32)	0.15
No	2022	38.5	579	19.7	1.00	
NCQ: problems with memory						
Yes	581	43.5	174	25.9	1.34 (1.16–1.56)	0.0001
No	1991	38.6	546	18.9	1.00	
NCQ: problems with emotional regulation						
Yes	666	41.8	205	24.6	1.32 (1.15–1.52)	0.0001
No	1906	38.9	515	18.9	1.00	
Educational Attainment						
Did not complete high school	193	50.4	45	31.7	2.15 (1.66–2.78)	<.0001
Completed high school	2139	46.2	579	26.8	1.80 (1.60–2.03)	<.0001
College graduate	1323	34.4	352	15.5	1.00	

Table 5

Multivariable regression model of the association between disease and treatment factors with marital status, adjusted for gender and age at last contact

Characteristic	Never-married			Ever-divorced				
	N	%	RR (95%CI) p-value	N	%	RR (95%CI) p-value		
Age at diagnosis								
<13 (years)	3338	50.6	1.38 (1.14-1.67)	580	20.5	1.28 (1.03-1.60)	0.03	
Gender								
Female	1692	38.5	1.00	524	22.0	1.00	0.04	
Male	2006	44.2	1.13 (1.06-1.19)	457	20.7	0.83 (0.70-0.99)		
Educational Attainment								
Did not complete high school	193	50.4	1.14 (1.00-1.30)	0.04	45	31.7	2.58 (1.80-3.70)	<.0001
Completed high school	2139	46.2	1.12 (1.05-1.20)	0.0003	579	26.8	1.79 (1.49-2.17)	<.0001
College graduate	1323	34.4	1.00	352	15.5	1.00		
Cranial radiation (Centigrav)								
>2400	89	54.6	1.15 (1.02-1.31)	0.03	9	14.5	0.60 (0.32-1.11)	0.10
>0 and <2400	726	49.6	1.03 (0.97-1.11)	0.33	128	19.5	0.91 (0.73-1.13)	0.40
0	1150	42.5	1.00	273	19.8	1.00		
Sperm cell Transplant								
Yes	37	44.0	1.05 (0.82-1.34)	0.71	7	16.7	1.6 (0.61-4.19)	0.33
No	3355	41.6	1.00	867	20.8	1.00		
Treatment duration (years)								
≥ 2 years	1884	45.1	1.04 (0.97-1.12)	0.29	409	20.1	1.05 (0.85-1.30)	0.67
< 2 years	1386	37.6	1.00	434	21.2	1.00		
Recurrence								
Yes	415	45.8	1.10 (1.01-1.2)	0.04	93	21.7	0.96 (0.70-1.32)	0.82
No	3283	40.9	1.00	888	21.3	1.00		

Table 6

Multivariable regression model of the association between neurobehavioral conditions with marital status, adjusted for age at last contact

Characteristic	Never-married			Ever-divorced			
	N	%	RR (95%CI) p-value	N	%	RR (95%CI) p-value	
Age at diagnosis (years)	3338	50.61	1.44 (1.24–1.66)	580	20.5	1.06 (0.9–1.24)	0.50
Gender							
13–20	360	15.4	1.00	401	22.7	1.00	
Female	1692	38.51	1.15 (1.09–1.22)	524	22.0	0.93 (0.81–1.07)	0.31
Male	2006	44.2	1.00	457	20.7	1.00	
Educational Attainment							
Did not complete high school	193	50.41	1.06 (0.92–1.22)	41	531.71	1.86 (1.34–2.59)	0.0002
Completed high school	2139	46.21	1.11 (1.05–1.18)	579	26.81	1.67 (1.45–1.91)	<.0001
College graduate	1323	34.4	1.00	352	15.5	1.00	
Emotional Distress (GSI T-score ≥ 50)							
Yes	1613	39.91	1.03 (0.97–1.08)	395	44.25	1.33 (1.15–1.54)	0.0001
No	1513	36.8	1.00	412	18.0	1.00	
NCQ: problems task efficiency							
Yes	935	49.4	1.27 (1.2–1.35)	220	25.41	1.14 (0.97–1.35)	0.12
No	1637	35.6	1.00	500	18.5	1.00	
NCQ: problems organization							
Yes	550	44.41	1.05 (0.98–1.12)	171	41.22	0.95 (0.8–1.14)	0.60
No	2022	38.5	1.00	579	19.7	1.00	
NCQ: problems with memory							
Yes	581	43.50	0.97 (0.91–1.04)	174	25.91	1.12 (0.94–1.33)	0.20
No	1991	38.6	1.00	546	18.9	1.00	
NCQ: problems emotional regulation							
Yes	666	41.8	0.9 (0.85–0.97)	205	24.61	1.11 (0.95–1.31)	0.19
No	1906	38.9	1.00	515	18.9	1.00	

Table 7

Multivariable regression model of the association between physical conditions with marital status, adjusted for age at last contact

Characteristic	Never-married			Ever-divorced		
	N	%	RR (95%CI) p-value	N	%	RR (95%CI) p-value
Age at diagnosis (years)	3338	50.6	1.40 (1.21-1.62) <.0001	580	20.5	1.08 (0.91-1.26) 0.38
Gender						
13-20	360	15.4	1.00	401	22.7	1.00
Female	1692	38.5	1.20 (1.13-1.26) <.0001	524	22.0	0.95 (0.83-1.09) 0.45
Male	2006	44.2	1.00	457	20.7	1.00
Educational Attainment						
Did not complete high school	193	50.4	1.10 (1.04-1.17) 0.0009	45	31.7	1.68 (1.46-1.93) <.0001
Completed high school	2139	46.2	1.00	579	26.8	1.00
College graduate	1523	34.4	1.40 (1.21-1.62) <.0001	352	15.5	1.08 (0.91-1.26) 0.38
Short stature						
Yes	978	57.2	1.27 (1.2-1.34) <.0001	150	23.7	1.13 (0.94-1.36) 0.18
No	2571	37.0	1.00	810	20.9	1.00
Poor physical function (SF-36 T-score <40)						
Yes	375	42.9	1.08 (1.00-1.18) 0.05	141	31.7	1.4 (1.18-1.67) 0.0001
No	2717	40.9	1.00	687	19.6	1.00

Table 8

Structural Equation Model for Predictors of Never-Married Status in CCSS Cases (corresponding to Figure)

	Estimate (EST)	Standard Error (SE)	EST/SE	STD YX Estimate	P-Value
Younger Current Age	0.091	0.003	28.06	0.518	<0.0001
Short Stature	0.225	0.035	6.34	0.198	<0.0001
Poor Task Efficiency	-0.047	0.007	-7.17	-0.154	<0.0001
Male Gender	0.233	0.035	6.62	0.090	<0.0001
CNS Radiation	0.219	0.045	4.83	0.079	<0.0001
Better Memory	0.046	0.011	4.06	0.078	<0.0001
Poor Physical Function	0.004	0.001	4.63	0.074	<0.0001
Poor Emotional Function	0.003	0.001	2.32	0.040	0.020