Perceived positive impact of cancer among long-term survivors of childhood cancer: a report from the childhood cancer survivor study

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Abstract

Objective: Investigations examining psychosocial adjustment among childhood cancer survivors have focused primarily on negative effects and psychopathology. Emergent literature suggests the existence of positive impact or adjustment experienced after cancer, as well. The purpose of this study is to examine the distribution of Perceived Positive Impact (PPI) and its correlates in young adult survivors of childhood cancer.

Methods: 6425 survivors and 360 siblings completed a comprehensive health survey, inclusive of a modified version of the Post-traumatic Growth Inventory (PTGI) as a measure of PPI. Linear regression models were used to examine demographic, disease and treatment characteristics associated with PPI.

Results: Survivors were significantly more likely than siblings to report PPI. Endorsement of PPI was significantly greater among female and non-white survivors, and among survivors exposed to at least one intense therapy, a second malignancy or cancer recurrence. Survivors diagnosed at older ages and fewer years since diagnosis were more likely to report PPI. Income, education and marital/relationship status appeared to have varied relationships to PPI depending upon the subscale being evaluated.

Conclusions: The existence and variability of PPI in survivors in this study suggest that individual characteristics, inclusive of race, gender, cancer type, intensity of treatment, age at diagnosis and time since diagnosis, have unique and specific associations with different aspects of perceived positive outcomes of childhood cancer.

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Background

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Research has tended to focus on the ways in which serious life-threatening conditions negatively influence health and functioning. Yet, empirical evidence also suggests that some individuals perceive personal growth or benefits as a result of exposure to traumatic experiences [1–4]. In the case of pediatric oncology, published investigations examining psychosocial adjustment among childhood cancer survivors have focused primarily on negative effects and psychopathology [5–7]. However, while limited, emergent research also suggests the possibility that survivors are resilient and report positive life circumstances which they attribute to having had cancer.

Positive outcomes assumedly attributable to cancer are conceptualized in the psycho-oncology literature as benefit finding or perceived benefits [8–10], resilience and thriving [11–13], and post-traumatic growth [14–19]. They are defined as cognitive processes by which those who have experienced life changing or traumatic events apply positive interpretations to and find meaning in the

event [14], and occur when individuals formulate adaptive interpretations or worldviews as a result of experiencing these events [20]. Findings from these investigations are consistent with the theoretical notion of 'post-traumatic growth' [21], in that they offer evidence that cancer changes or influences survivors' lives in some or all of the following domains: (1) life perspective (e.g. altered priorities, greater joy or appreciation for one's life, greater sense of meaning, enhanced religious or spiritual beliefs); (2) relationships (e.g. greater appreciation for one's relationships, greater sense of intimacy, enhanced emotional expressiveness, increased sensitivity to others); and (3) self-perception (e.g. sense of emotional growth, strength, self-reliance).

Studies suggest that positive life changes attributable to cancer are reported by a majority of adult cancer patients and survivors [10,17]. Sumalla et al. [18] estimated that in most cases 80% or more of adult cancer survivors regarded themselves as having benefited in some way from the experience. Indeed, young survivors also are capable of describing subjective perceptions of how they believe having had cancer has affected their lives. For example, 304 childhood cancer survivors (aged 14-25 years) were asked about their experience of positive and negative life changes after their illness [22], and over half of the sample reported themselves as having enhanced concern for others (60%), ability to cope with tragedy (54%), sense of identity (52%), and spiritual wellbeing (52%). In accompanying qualitative interviews with 50 young adult survivors of childhood cancer (aged 17-29 years at interview, 1 month-17 years of age at diagnosis, at least 3 years posttreatment), a majority of study participants reported positive effects attributable to having had cancer, including increased psychological maturity (65%), greater compassion and empathy (61%), new values and priorities (57%), new strengths (48%), and increased recognition of vulnerability and struggle, with a deeper appreciation for life (44%) [23]. A recent investigation of 150 adolescent survivors of childhood cancer (aged 11–19 years, at least 1 year post-treatment) indicated that 84.7% of survivors identified at least one positive consequence of having had cancer, and 32% identified four or more positive consequences [14]. Positive influence was associated with older age at diagnosis, greater treatment severity and life threat, and less time since diagnosis. In another study of 199 child and adolescent cancer patients (aged 7-18 years), 'benefit-finding' was associated with older age at diagnosis, less time since diagnosis, and being African American [9].

For many childhood cancer survivors, particularly those who were very young at the time of treatment, their memories of cancer are often created through stories passed on to them by family members. Thus, for a young adult cancer survivor population, perceived positive impact (PPI) of cancer may be less a reflection of change attributable to cancer and more a function of 'event centrality', the extent to which a stressful or life-threatening event (such as cancer) becomes a salient organizing principle for the individual's growing sense of self and view of the world [20,24].

Assessing the PPI of cancer in childhood survivors is instructive for several reasons. Recent research suggests that subjective perceptions of conditions or phenomenon are better predictors of distress and well-being than are objective health status measures [25-29]. Thus, studies of PPIs and their correlates or predictors will achieve a more comprehensive and balanced understanding of psychosocial adjustment in long-term survivors of childhood cancer. Such understanding can guide the development of interventions that not only prevent or minimize cancer's negative effects but also promote self-esteem and adjustment, and subsequently facilitate successful achievement of developmental tasks typical of adolescence and young adulthood, such as establishing employment or career paths, forming a family, and achieving autonomy. Furthermore, theoretical models of benefit-finding, PPI and post-traumatic growth in this field are evolving. Findings from investigations such as this may help to advance theoretical suppositions as they apply to the impact of chronic or serious life-threatening illnesses in young people whose psychosocial and cognitive development are incomplete at the time of exposure to these medical conditions.

Given recent interest in examining potential positive impacts of cancer, this study aims to (1) examine the distribution of perceived positive impact (PPI) of cancer in young adults who are childhood cancer survivors, and compare it to that of siblings; and (2) examine the extent to which cancer-related factors (e.g. age at diagnosis, time since diagnosis, cancer type) and key sociodemographic variables predict the likelihood of young adult survivors attributing positive outcomes to having had cancer. This study overcomes many of the limitations of current research in the field of psychosocial oncology in that it involves a large multi-institutional sample, includes a comparison group, and is powered to examine the simultaneous influences of sociodemographic and cancer-related risk factors on outcomes.

Methods

Sample and data collection

The Childhood Cancer Survivor Study (CCSS) is a large, multi-institutional cohort study that tracks the health status of survivors of childhood cancer diagnosed between 1970 and 1986 who survived at

least 5 years post diagnosis. The institutional review board at each participating center reviewed and approved the CCSS protocol and documents sent to participants. Study participants provided informed consent for participation in the study and for release of medical-record information. Detailed descriptions of the study design and characteristics of the cohort are reported elsewhere [30,31].

In 1995–1996, eligible study participants were mailed a 24-page baseline questionnaire. All subjects who completed that baseline survey received a subsequent follow-up questionnaire sometime in 2002–2004, in which were included standardized measures of health-related quality of life, psychological distress, and the outcomes and correlates reported here. Of 20 691 survivors of childhood cancer identified for the original cohort, 3058 (14.8%) were lost to follow-up despite extensive efforts to locate them. Among the remaining 17 633 survivors, 14 358 (81.4%) completed the baseline questionnaire. In addition, a sample of the participating survivor population were randomly selected

and asked to nominate their nearest age sibling to be a part of the comparison group. Of the 4782 siblings nominated, 3899 siblings (81.5%) participated in the baseline survey. In 2003, 11576 (80.6%) of the original cohort were located and requested to participate in a follow-up survey. Of these, 9308 (80.4%) completed and returned a survey, with 6425 being over 18 years of age and having completed all survey items required for this study by themselves and thus included in the analysis reported here. Also, a sub-sample of 500 siblings over age 18 was randomly selected to survey. Of these, 360 (72.0%) completed all of the items of interest to this study. Figure 1 offers a graphic representation of subject recruitment and retention of the CCSS survivor group.

Although siblings of childhood cancer survivors are also influenced by the life disruption associated with cancer in the family, they can still serve as an adequate same-aged peer comparison group [32]. Recent research suggests that siblings approximate a psychologically healthy comparison group, in

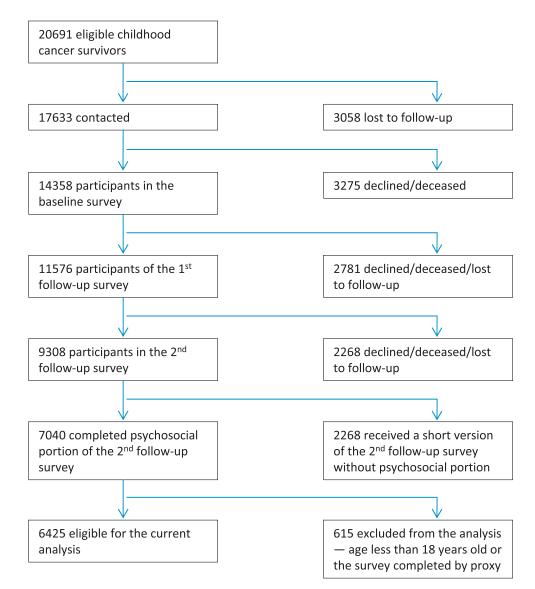


Figure I. Subject recruitment

that rates of post-traumatic stress and emotional distress in adult siblings of adult survivors of childhood cancer are similar to those found in population norms [33,34].

Measures

PPI, the primary outcome variable, was assessed using a modified version of the Post-traumatic Growth Inventory (PTGI), a 21-item scale comprising subscales suggestive of positive growth in five domains: Relating to Others, New Possibilities, Personal Strength, Spiritual Change, and Appreciation for Life [21]. Modified wording of the question stem instructed survivor respondents to indicate how much their cancer experience has influenced their life. Siblings were instructed to indicate how much their life was influenced by their sibling's cancer experience. Each item was rated on a 6-point scale, with values ranging from 0 ('I am *NOT influenced by my/my sibling's experience'*) to 5 ('I am influenced to a VERY GREAT degree as a result of my/my sibling's experience'). Items are summed to derive five subscale scores. Higher scores suggest increasing levels of PPI.

Sociodemographic and cancer-specific variables were analyzed as potential correlates of PPI in the survivors. Self-reported sociodemographic variables included age at study, gender, employment status, marital/relationship status, education, race, and household income. Cancer-related variables were derived via medical records abstraction [30,31], and included type of cancer, age at diagnosis, relapse or second malignancy (Yes/ No), and years since diagnosis. A dichotomous 'intensity of treatment' variable (no intense therapy; at least one intense therapy) was created based on high-risk chemotherapy exposures (anthracyclines/ alkalating agents, other) and/or exposures to high-risk types and locations of radiotherapy (to brain or spine,), as compared to no chemotherapy or radiation [34].

Statistical analyses

Exploratory data analysis was performed to examine the distributions of variables of interest. To test the psychometric properties of the PTGI (as a measure of PPI) in a childhood cancer survivor population, we conducted a confirmatory factor analysis (CFA) in the eligible survivor population (i.e. aged 18 years or older at the survey and completed all 21 PTGI items, n = 6425) using a five-factor solution suggested by prior administrations of the PTGI instrument [1,35]. The weighted least-squares method was used, as the multivariate normality assumption was in doubt. To statistically test the fit of the five-factor model with that of a single-factor model, bootstrap was used, assessing the significance of the differences in three measures of goodness of fit—(1) the goodness of fit index (GFI), (2) the adjusted goodness of fit index (AGFI), and (3) the root mean square error of approximation (RMSEA), between the two models. In addition, we assessed internal consistency of each of the five factors by Cronbach's alpha.

Demographic factors were compared between the survivors and siblings with statistical significance evaluated using bootstrap, which takes into account potential within family correlation between a survivor and his/her sibling. The continuous scores of the five PTGI subscales among survivors were compared with those among siblings, using the modification of linear regression by Generalized Estimating Equations to account for the potential within-family correlation [36], adjusting for gender and two variables that differ significantly between the survivors and siblings in this sample: age at study and race. Effect size and confidence intervals are reported. Effect size is defined as the difference of a score between the survivor group and the sibling group divided by standard deviation of the latter. This approach is justified based on viewing the sibling group as an appropriate reference population. Adjusted differences in scores were similarly expressed as adjusted effect sizes. Similarly, relationships between the five PTGI subscales and selected correlates were assessed using linear regression models. A backward variable-selection method was employed to build a summary model that describes the independent and simultaneous associations of each subscale with demographic and clinical factors. In these models, age at diagnosis and years since cancer diagnosis were assumed to have linear effects. This linear assumption was examined using natural cubic splines [37], given the possibility that the relationships of these variables with the PTGI might be curvilinear [38]; however, results indicated that the linear assumption was reasonable and, therefore, used in the analysis. We considered treatment exposures within the first 5 years from the original diagnosis of cancer in defining treatment variables. All statistical analyses were performed using SAS Version 9.2 [39] and two-sided statistical inferences were employed throughout the analyses.

Results

Demographic characteristics of the survivors and siblings participating in this study are summarized in Table 1, along with cancer-related descriptive data of participating survivors. Cancer survivors were similar to siblings in gender and education level, but were more likely to be younger at interview (mean 32.3 years versus 33.9 years among siblings, p < 0.001), non-white (p < 0.001), not employed (p = 0.04), not married (p < 0.001) and

Table I. Descriptive statisti	cs
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	Siblings	Survivors
	n = 360(%)	n = 6425(%)
Age at interview**		
18–29	132 (36.7)	2598 (40.4)
30–39	135 (37.5)	2718 (42.3)
40+	93 (25.8)	1109 (17.3)
Race**	()	()
White non-Hispanic	322 (93.6)	5603 (87.5)
All others	22 (6.4)	798 (12.5)
Gender	22 (011)	,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,
Male	172 (47.8)	3064 (47.7)
Female	188 (52.2)	3361 (52.3)
Education	100 (0212)	55501 (5215)
≤High school graduate	52 (14.5)	953 (14.9)
Some college	127 (35.5)	2311 (36.2)
≥College graduate	179 (50.0)	3118 (48.9)
Employed*	177 (30.0)	5110 (10.7)
Yes	298 (83.0)	4999 (78.4)
No	61 (17.0)	1381 (21.6)
Personal income**	01 (17.0)	1301 (21.0)
<\$20 000	109 (34.8)	2658 (42.6)
<pre>\$20 000 +</pre>	204 (65.2)	3588 (57.4)
Ever married ^{**}	201 (05.2)	5500 (57.1)
Yes	256 (71.5)	3731 (58.6)
No	102 (28.5)	2635 (41.4)
Survivor diagnosis	102 (20.5)	2000 (11.1)
Leukemia		2123 (33.0)
CNS		669 (10.4)
HD		912 (14.2)
NHL		507 (7.9)
Kidney (Wilms)		620 (9.6)
Neuroblastoma		405 (6.3)
Soft tissue sarcoma		593 (9.2)
Bone cancer		596 (9.3)
Survivor age at diagnosis		570 (7.5)
0-4		2355 (36.7)
5-9		1446 (22.5)
10–14		1378 (21.4)
15–20		1246 (19.4)
Years since survivor diagnosis		1210(17.1)
15–19		1753 (27.3)
20–24		2296 (35.7)
25–29		1649 (25.7)
30–34		727 (11.3)
2nd malignancy or recurrence in		/2/ (11.3)
survivor		
Yes		1150 (17.9)
No		5275 (82.1)
At least one intense therapy		52/5 (02.1)
Yes		4686 (78.6)
No		· · ·
INU		1275 (21.4)

 $^{*}p < 0.05, \, ^{**}p < 0.001.$

to have a lower income (p = 0.009). On average, survivors were 8.7 years of age at time of diagnosis (SD = 5.9), and 23.6 years beyond their diagnosis (SD = 4.5). As reported elsewhere [34], descriptive demographic and cancer-related characteristics of survivors and siblings who completed the current survey of interest to this study were compared with those who did not complete it (but did complete the original CCSS baseline survey) using chi-square tests. Similarly, siblings included in this study were compared to siblings who completed the baseline questionnaire but not the survey of interest to this study. Compared to survivors who completed the baseline survey, follow-up survey respondents for this study were significantly more likely to be older, female, white, employed, married/partnered, and older age at diagnosis and fewer years since diagnosis. Survivor participants did not differ from non-participants by cancer diagnosis, survival time, or on a standardized measure of psychological distress assessed at baseline. As expected, based on their random selection for participation in the psychosocial portion of the questionnaire, sibling participants did not differ from non-participants by gender, age, race, educational attainment, employment, marital status, or baseline psychological distress.

Confirmatory factor analysis

The GFI derived from the five-factor CFA was 0.88, the AGFI was 0.85, and the RMSEA was 0.05. RMSEA is an index of fit that is less influenced by sample size, with recommended levels being 0.05 or below [40]. Cronbach's alpha was 0.92, 0.89, 0.86, 0.89, 0.84 for subscales indicating 'new possibilities', 'relating to others', 'personal strength', 'spiritual change', and 'appreciation for life', respectively. These indices suggest a reasonable fit of the five-factor structure to this survivor sample. CFA was repeated with a single factor to determine whether the five-factor model explained the variability in responses significantly better than the single-factor model. The single-factor model had significantly reduced fit indices (GFI = 0.84, AGFI = 0.80, RMSEA = 0.06; p < 0.001). Thus, the five-factor model accounted for the variance of responses significantly better than the singlefactor model.

Survivor-sibling comparisons

A comparison of survivor and sibling scores for PPI is summarized in Table 2. Mean scores on all subscales, adjusted for differences in age, gender, and race, were significantly higher among survivors than siblings. In all instances, survivors were more likely than siblings to indicate that they felt that their lives had been influenced by cancer. Effect sizes were generally small to moderate.

Multivariable modeling

Table 3 presents results of multivariable modeling (adjusted effect sizes) for survivors. As evidenced in the five subscale scores, endorsement of PPI was significantly more evident among female and nonwhite survivors, as well as among survivors exposed to at least one intense therapy or a second malignant neoplasm or recurrence of cancer. The

Subscale Score	Survivor mean score	Sibling mean score	Difference	Effect size ^a (95% CI)	p-Value
Relating to Others	18.59	6.4	2.18 (1.12–3.25)	0.22 (0.11-0.32)	< 0.001
New Possibilities	10.75	8.35	2.40 (1.70–3.11)	0.37 (0.26-0.48)	< 0.00
Personal Strength	11.87	9.65	2.23 (1.59-2.86)	0.37 (0.27–0.48)	< 0.001
Spiritual Change	5.30	4.34	0.96 (0.61-1.31)	0.28 (0.18–0.38)	< 0.001
Appreciation of Life	9.58	8.56	1.02 (0.55–1.49)	0.23 (0.12–0.34)	< 0.001

Table 2. Age-, gender-, and race-adjusted scores and score difference between survivors and siblings

^aEffect size = Standardized difference (i.e. Difference between groups divided by standard deviation of siblings).

likelihood of reporting higher PPI increased significantly with age at diagnosis such that childhood survivors diagnosed at older ages were more likely to report PPI than those diagnosed at younger ages. In most cases, PPI decreased with years since diagnosis. Respondents with fewer years since diagnosis were more likely to endorse PPI. While the effect sizes appear small for each additional year, over several years the magnitude of change becomes more clinically meaningful. When compared to survivors of childhood leukemia, survivors of Hodgkin's disease, brain and kidney (Wilm's) tumors and neuroblastoma reported significantly less PPI. In contrast, survivors of bone cancer reported significantly more PPI when compared to leukemia survivors.

Sociodemographic and cancer-related variables appeared to have varied relationships to PPI depending upon the subscale domain, although the small effect sizes suggest that these significant relationships may simply be a function of the power of the analysis and not clinically meaningful, thus subject to further research. For example, those without a college degree reported significantly greater PPI with regard to relating to others and perceiving new life possibilities. In contrast, college graduates were more likely than those without a college degree to report greater appreciation for life. Survivors who were married or in long-term relationships reported more spiritual change and appreciation of life than those not married or in committed relationships. Survivors reporting personal incomes of <\$20000/year reported less personal strength but more spiritual change compared to those earning more. Employment status was not observed to be associated with any subscale.

Discussion

The findings reported here suggest that long-term post-cancer perceptions of positive impact are greater among survivors than siblings. It may be that childhood survivors' experiences of exposures to invasive medical procedures, treatments, and long-term complications contribute to a greater likelihood of reporting PPI when compared to siblings, for whom the experience of childhood cancer in the family often involves disruptions in daily routines and relationships with parents and not necessarily physical debilitation or trauma [41,42]. The experience of physical symptoms, debilitation, or therapy-related late effects may be of relatively greater salience, in contrast to social life disruption, when it comes to predicting PPI.

The existence and variability of PPI in young adult survivors in this study suggests that survivor characteristics, inclusive of race, gender, cancer type, intensity of treatment, age at diagnosis, and time since diagnosis, have unique and specific associations with different aspects of PPI. For instance, the positive relationship observed between age at diagnosis and PPI suggests that the cognitive capacity to acknowledge the severity of the life disruption caused by cancer may be a necessary antecedent to later perceiving some positive effect. This finding supports theories of post-traumatic growth and resilience that are predicated on a subject perceiving an experience as traumatic in order to derive positive meaning or growth [2,43]. However, relevant and competing theories (e.g. event centrality) do not require a subject to acknowledge an experience as traumatic in order to grow from it [9,20].

Educational attainment and marital/relationship status also appeared to be positively associated with reporting PPI. The successful attainment of life goals and dreams, such as finishing school, having a significant, meaningful and intimate relationship, or starting a family, may contribute to survivors feeling like their lives are normal after having had cancer. These achievements perhaps forge a more positive conception of benefits derived from once having had cancer. Provocative is the converse notion that acknowledgement or recognition of positive impacts of cancer may somehow increase the likelihood of achieving these or other normative developmental tasks. This cross-sectional study cannot determine the direction of this relationship; however, the question of whether or not cognitive behavioral therapies can incur a reframing of one's cancer experience and subsequently improve mental or occupational health outcomes for childhood cancer survivors is worthy of future investigation.

The gender and race differences observed here suggest that PPI may be a socially and culturally influenced coping process whereby men and women, and Whites and non-Whites, differ. These

$ \begin{array}{cccccc} \mbox{were reduction} & \label{eq:constraint} & eq:constra$	Variables	Relating to others subscale Diff (95% CI)	New possibilities subscale Diff (95% CI)	Personal strength subscale Diff (95% CI)	Spiritual change subscale Diff (95% CI)	Appreciation of life subscale Diff (95% CI)
$ \begin{array}{cccccc} \mbox{thermatic} & -0.0 (-0.18 \times -0.03)^{**} & -0.18 (-0.24 \times -0.03)^{**} & Ref & -0.13 (-0.21 \times -0.05)^{**} & Ref & -0.13 (-0.21 \times -0.05)^{**} & Ref & -0.22 (-0.23 \times -0.13)^{**} & Ref & -0.22 (-0.23 \times -0.13)^{**} & Ref & -0.22 (-0.23 \times -0.13)^{**} & Ref & -0.22 (-0.23 \times -0.20)^{**} & -0.22 (-0.23 \times -0.20$	Years since diagnosis Age at diagnosis	0.01 (0.01 to 0.02)**	-0.01 (-0.02 to -0.01)**	-0.01 (-0.02 to -0.01)** 0.02 (0.01 to 0.02)**	0.01 (0.01 to 0.02)**	-0.01 (-0.02 to -0.01)** 0.01 (0.01 to 0.02)**
$ \begin{array}{cccccccccccccccccccccccccccccccccccc$	Race White non-Hispanic All others	-0.10 (-0.18 to -0.03)** Ref	18 (-0.26 to	-0.13 (-0.20 to -0.05)** Ref	-0.13 (-0.21 to -0.05)** Ref	−0.09 (−0.17 to −0.02)* Ref
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$ \begin{array}{c} \ \ \ \ \ \ \ \ \ \ \ \ \ $	iducation ≲ High school grad iome college ≥ College graduate	0.20 (0.12 to 0.27)** 0.06 (0.01 to 0.11)* Ref	0.09 (0.01 to 0.17)* 0.02 (-0.04 to 0.08) Ref			-0.09 (-0.17 to -0.02)* -0.06 (-0.11 to -0.01)* Ref
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$ \begin{array}{cccc} \mbox{ed} & & & & & & & & & & & & & & & & & & &$	ersonal income ¢\$20 000 20 000			-0.09 (-0.14 to -0.04)** Ref	0.10 (0.04 to 0.16)** Ref	
$ \begin{array}{ccccc} \mbox{goos} & -0.15 (-0.22 \mbox{to} -007)^{**} & \mbox{fef} & fe$	ver married es Jo				0.12 (0.06 to 0.18)** Ref	0.09 (0.04 to 0.15) ** Ref
$ \begin{array}{c} \mbox{currence} \\ \mbox{ lef } \mbox{ Ref } & 0.25 (0.18 to 0.31)^{**} & 0.25 (0.18 to 0.33)^{**} & 0.22 (0.15 to 0.28)^{**} & Ref \\ \mbox{ Ref } & \mbox{ Ref } & \mbox{ 0.22 (0.15 to 0.29)^{**} & Ref } & \mbox{ 0.15 (0.15 to 0.27)^{**} & \mbox{ Ref } & \mbox{ Ref } & \mbox{ Ref } & \mbox{ Ref } & \mbox{ 0.15 (0.08 to 0.22)^{**} & \mbox{ Ref } & \mbox{ Ref } & \mbox{ Ref } & \mbox{ 0.15 (0.08 to 0.22)^{**} & \mbox{ Ref } & \mbox{ Ref } & \mbox{ Ref } & \mbox{ Ref } & \mbox{ 0.15 (0.15 to 0.27)^{**} & \mbox{ Ref } & \mbox{ Ref } & \mbox{ Ref } & \mbox{ 0.15 (0.08 to 0.22)^{**} & \mbox{ Ref } & \mbox{ Ref } & \mbox{ Ref } & \mbox{ Ref } & \mbox{ 0.15 (0.08 to 0.22)^{**} & \mbox{ Ref } & \mbox{ Ref } & \mbox{ Ref } & \mbox{ 0.15 (0.08 to 0.22)^{**} & \mbox{ Ref } & \mbox{ 0.16 (-0.14 to 0.03)^{**} & -0.18 (-0.22 to -0.008)^{**} & -0.018 (-0.22 to -0.008)^{**} & -0.02 (-0.12 to 0.08) & \mbox{ 0.01 (-0.08 to 0.11) } & \mbox{ 0.07 (-0.04 to 0.18) } & 0.$	ear of diagnosis 970–1973 974–1978 979–1986	-0.15 (-0.22 to -0.07)** -0.04 (-0.10 to 0.01) Ref				
interse therapy $0.20 (0.14 \text{ to } 0.26)^{**}$ $0.22 (0.15 \text{ to } 0.29)^{**}$ $0.21 (0.15 \text{ to } 0.27)^{**}$ $0.15 (0.08 \text{ to } 0.22)^{**}$ RefRefRefRef Ref Ref Ref -0.12 (-0.20 \text{ to } -0.03)^{**} $-0.14 (-0.23 \text{ to } -0.04)^{**}$ $-0.19 (-0.28 \text{ to } -0.10)^{**}$ $-0.18 (-0.27 \text{ to } -0.08)^{**}$ -0.11 (-0.20 \text{ to } -0.02)^{**} $-0.06 (-0.14 \text{ to } 0.03)$ $-0.02 (-0.12 \text{ to } 0.08)^{**}$ $-0.02 (-0.12 \text{ to } 0.08)^{**}$ -0.02 (-0.12 \text{ to } 0.08) $-0.03 (-0.14 \text{ to } 0.08)$ $0.01 (-0.08 \text{ to } 0.11)$ $0.07 (-0.04 \text{ to } 0.18)$	MN or Recurrence es Jo	0.25 (0.18 to 0.31)** Ref	0.25 (0.18 to 0.33)** Ref	0.22 (0.15 to 0.28)** Ref	0.23 (0.16 to 0.30)** Ref	0.20 (0.13 to 0.26) ** Ref
$ \begin{array}{llllllllllllllllllllllllllllllllllll$	t least one intense therapy es Jo		0.22 (0.15 to 0.29)** Ref	0.21 (0.15 to 0.27)** Ref	0.15 (0.08 to 0.22)** Ref	0.18 (0.11 to 0.24) ** Ref
	liagnosis CNS HL	-0.12 (-0.20 to -0.03)** -0.11 (-0.20 to -0.02)* -0.02 (-0.12 to 0.08)	-0.14 (-0.23 to -0.04)** -0.06 (-0.15 to 0.02) -0.03 (-0.14 to 0.08)	-0.19 (-0.28 to -0.10)** -0.06 (-0.14 to 0.03) 0.01 (-0.08 to 0.11)	-0.18 (-0.27 to -0.08)** -0.12 (-0.22 to -0.02)* 0.07 (-0.04 to 0.18)	-0.30 (-0.39 to -0.21)** -0.04 (-0.13 to 0.05) 0.03 (-0.07 to 0.13)

Variables	Relating to others subscale	New possibilities subscale	Personal strength subscale	Spiritual change subscale	Appreciation of life subscale
	Diff (95% CI)	Diff (95% CI)	Diff (95% Cl)	Diff (95% CI)	Diff (95% CI)
Kidney (Wilms)	-0.21 (-0.30 to -0.12)**	-0.29 (-0.39 to -0.19)**	-0.21 (-0.30 to -0.12)**	-0.21 (-0.30 to -0.11)**	-0.17 (-0.26 to -0.08)**
Neuroblastoma	-0.17 (-0.28 to -0.06)**	-0.23 (-0.35 to -0.11)**	-0.13 (-0.24 to -0.02)*	-0.10 (-0.22 to 0.02)	-0.04 (-0.14 to 0.07)
Soft tissue sarcoma	-0.03 (-0.12 to 0.07)	-0.08 (-0.18 to 0.02)	0.03 (-0.06 to 0.12)	-0.01 (-0.11 to 0.09)	-0.01 (-0.11 to 0.08)
Bone cancer	0.14 (0.05 to 0.24)**	0.28 (0.18 to 0.38)**	0.23 (0.13 to 0.32)**	0.09 (-0.02 to 0.20)	0.16 (0.06 to 0.25)**
Leukemia	Ref	Ref	Ref	Ref	Ref
*p<0.05, ***p<0.01; blank c *Effect size = Standardized 1 ^b For each subscale, all variabl ^c Employment status ('Employ	* $p < 0.05$, ** $p < 0.01$; blank cells indicate that the standardized mean difference was not statistically significant at $p < 0.05$. *Effect size = Standardized mean difference (i.e. Differences in mean compared to referent, divided by standard deviation). *Pror each subscale, all variables whose effect-size estimates are shown in the table were included in the model and adjusted for each other. *Employment status ('Employed') was one of the candidate variables in the backward selection, but it was not selected in any of the models.	rence was not statistically significant at $p < 0$ npared to referent, divided by standard devis the table were included in the model and as e backward selection, but it was not selected	ly significant at p < 0.05. led by standard deviation). in the model and adjusted for each other. Negative scores reflect relatively less growth, while positive scores reflect more growth. ut it was not selected in any of the models.	t relatively less growth, while positive scor	ss reflect more growth.

differences may reflect a greater coping ability among women and individuals from racial minority groups that results from being primed to deal with adversity related to the subtle but sometimes explicit challenges of experiencing raceor gender-based discrimination. This speculative interpretation of race and gender-based differences may be examined further in the context of stress-based theories of allostasis, which suggest that there exists a threshold up to which people are resilient and can manage stress [44,45]. However, over-exposures to stress in general, and discrimination in particular, prohibit possibilities of resilience or growth, and instead result in multiple and varied deleterious health outcomes [46-48].

If we are to understand PPI in the context of event centrality, than we must look for evidence to the effect that a cancer history has somehow shaped and/or continues to shape individuals' senses of themselves or the world around them. For example, survivors of bone cancers were significantly more likely to report PPI than survivors of any other cancer types (Table 3). It is possible that bone cancer, with its attendant loss of limb or body disfigurement, is perceived as a most salient aspect of a childhood cancer survivor's identity due to its constant physical reminders (e.g. amputation, limb salvage). In turn, PPI was significantly greater among leukemia survivors compared to all other cancer types (except bone cancers), perhaps due to the relatively longer length of time spent in treatment (upward of 2–3 years). Survivors who reported more intense therapies or a recurrence of cancer or second malignancy also were more likely to report PPI. One can assume that more intense treatments are accompanied by more severe, sickening or debilitating side effects (e.g. nausea, vomiting, hair loss, diarrhea, constipation, mouth sores) and thus may contribute to a greater likelihood that survivors recall the cancer experience as traumatic, or at least as having a greater impact on their life at the time. Also, consistent with prior research [14], reporting PPI in this study sample decreased over time. Perhaps in the absence of reminders, any perceived growth or potential positive life influence attributable to cancer is forgotten, or becomes less central to the identity of the survivor over time. Without cognitive or behavioral reinforcements, survivors may be less likely to maintain new values, attitudes, beliefs, or behaviors that may have emerged in the first years following cancer treatment.

CNS/brain tumor survivors may be an exception, in that the centrality of on-going cognitive, behavioral, and physically debilitating effects contribute to a greater likelihood of perceiving a more negative long-term impact of cancer. Indeed, brain tumor survivors experience disproportionate

Table 3. Continued

rates of learning disabilities, unemployment and other social life disruptions when compared to survivors of other childhood cancer types [49,50]. Thus, cognitive and behavioral limitations may preclude PPI.

A lack of empirical data and underdeveloped theories of post-traumatic growth, thriving, and PPI limits our ability to fully understand the findings reported here. Evaluating the influence of cancer in a cross-sectional study involving subjects who, on average, are 24 years beyond their cancer therapy potentially confounds our ability to definitively conclude that reports of PPI are actually attributable to cancer. Yet, they do not negate the possibility that one's cancer experience can and does become central to one's identity and sense of the world, at least for some period of time in one's life. By changing the question stem of the PTGI to assess 'influence' of cancer on one's life as opposed to change attributable to cancer, we may have altered the validity and reliability of the PTGI as a measure of 'post-traumatic growth', although psychometric analyses supported the instrument's theoretically derived factor structure in this study sample.

Despite the large sample size and the large geographic representation of survivors, there are other limitations to this study. The survey was completed by mail, and thus is self-report and expected to be less sensitive than an interview. Additionally, not all of those contacted completed this survey, which raises the potential for some selfselection in the respondents. Although there could be a number of reasons for such self-selection, it is possible that those who were most distressed by the questions failed to complete the forms.

Future research is needed to examine the extent to which PPI is related to, and distinguishable from, health-related quality of life and psychological health and well-being. Future challenges also will involve differentiating the experiences of cancer survivors from variously affected (siblings, survivors of other medical, physical, or sexual trauma) or non-affected (e.g same age peers, population norms) populations. In addition, future studies utilizing longitudinal study designs would have the potential for assessing cancer-related life changes within the context of rapid social and emotional changes that commonly occur in this younger-aged population. Finally, investigation into the relationship of PPI to post-traumatic stress symptoms will be an important effort in understanding better the long-term affect of cancer on young adult survivors. This information can guide the development of future interventions that aim not only to prevent or treat distress reactions among some survivors but also promote survivors' interpretations of their cancer experience in a manner that gives meaning and continuity to their sense of self and life story [51].

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