Title: Characterization of drinking in childhood cancer survivors compared to general population

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Background and Rationale: During the past 40 years survival rates from childhood cancer have increased dramatically and now 70% of children diagnosed with cancer are expected to survive long-term. Research has focused on the many physical health obstacles experienced by survivors of childhood cancer as a result of their cancer or treatments. Psychological and social consequences of childhood cancer have been less of a focus and research has yielded contradictory findings. Descriptions of health risk behaviors such as alcohol consumption among childhood cancer survivors are few and conflicting, with some authors reporting heavy use[1] and others reporting little use[2, 3] or comparable use to peers[4]. An understanding of drinking behaviors in this population is important since childhood cancer survivors may be more vulnerable to adverse health consequences of drinking compared to the general population.

In general populations, alcohol consumption, particularly in higher quantity, is associated with a number of health [5, 6] and social harms[7, 8]. Drinking among survivors is more risky than in general populations since alcohol exposes survivors to further toxins and mutagenic agents which may place individuals at risk for secondary cancers or organ complications[9, 10]. Furthermore alcohol use may interact with prescribed medications, such as those used to treat cardiac problems from chemotherapy [8]. As a result of these late effects it has been recommended by some that childhood cancer survivors abstain from alcohol use[10].

Of particular concern are those diagnosed with cancer during adolescence and young adulthood, while facing developmental issues relating to identity, social acceptance and achievement. A period of serious illness during this time can disrupt these developmental tasks, leaving survivors feeling alone, uncertain about their identity and future. This developmental period coincides with the peak time for alcohol use in the lives of adolescents and young adults.[11, 12] Previous research has described the use of alcohol to reduce stress and anxiety [13] and the correlation between higher stress and adolescent alcohol use[14].

Little research has characterized the quantity and frequency of alcohol consumption and risk factors for potentially harmful drinking among this population or compared cancer survivors drinking patterns to the general population. A clear epidemiological description of alcohol use and risk factors will better characterize the scope of the issue and facilitate the development of targeted recommendations related to screening, education and treatment of alcohol problems in the long-term follow-up of cancer survivors.

Specific Aims/Research Hypotheses:

The proposed study will utilize data from two sources: the Long-Term Follow-Up Study using participants in the Childhood Cancer Survivor's Study (CCSS) (N=14,054) and the National Alcohol Survey (NAS-2000) which was carried out in the year 2000 and assessed alcohol use among a sample of 8,054 English and Spanish speaking men and women in all 50 states of the U.S and Washington D.C.. Participants matching the age range of the CCSS data, ages 18-48 (N=6,065) will be used for this study. The NAS-2000 employed a sample collected through random digit dial techniques and Computer Assisted Telephone Interview. Alcohol variables between the CCSS and the NAS-2000 are closely comparable and allow for the assessment of beverage specific drinking frequency and quantity across both populations. The prevalence of drinking behaviors among childhood cancer survivors will be compared with similar age, racial/ethnic and gender groups in the national population. Predictors for risky and heavy drinking will be examined among childhood cancer survivors especially among the hypothesized highest risk groups; young adults and survivors diagnosed during adolescence. The study has the following specific aims:

Aim 1: To describe drinking patterns among survivors of childhood cancer and compare them to national norms by age, gender and race/ethnicity.

Aim 2: To determine factors associated with risky and heavy drinking. Being a young adult (ages 18-29) and cancer diagnosis as a teen will be associated with heavier drinking controlling for previously identified risk factors or concerns including; adverse health or mental health status, anxieties or fears relating to the cancer or its treatment, having a history of cognitive compromising treatments such as cranial radiotherapy and intrathecal methotrexate, age of initiating drinking, race/ethnicity, income and education, type of cancer.

Analysis Framework: (for variables description, see Appendix A, page 4)

Aim 1: Outcomes of interest

A. Current drinking; (past year)

B. Risky drinking; no more than one drink per day for women and no more than two drinks per day for men [15].

C. Heavy drinking; five or more drinks per day

Predictor Variable:

not applicable, descriptive data

Aim 2: Outcomes of interest:

Risky drinking Heavy drinking **Predictor Variables**:

current age, (young adult vs. other)

age of cancer diagnosis

Controlling for the effects of previously identified risk factors: adverse health or mental health status, fears/anxieties relating to the cancer or its treatment, treatments such as cranial radiotherapy and intrathecal methotrexate, age of initiating drinking, race/ethnicity, income and education, type of cancer

Subject population: Long-Term Follow-Up Study participants in the (CCSS) (N=14,054) and the National Alcohol Survey-2000 (N=6,065).

Specific Tables and Figures: (See Appendix B)

Table 1: Prevalence of current drinking, risky drinking and heavy drinking described in CCSS and NAS-2000. Table will include prevalence described by gender, age and racial/ethnic groups.

Table 2: Factors associated with risky drinking and heavy drinking; bivariate associations.

Table 3: Factors associated with risky drinking and heavy drinking; multivariate models examining the role of age of diagnosis, current age, adverse health or mental health status and cognitive compromising treatments controlling for demographic and personal variables.

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ENDNOTES

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Alcohol Questions Alcohol variable definitions Predictor/control variables	p 8 p 9
Health Status	р9
Demographic Variables	р10
Cancer Variables	р10

Equivalency of Alcohol Questions between the Childhood Cancer Survivor Study and the National Alcohol Survey-2000

Appendix A

Childhood Cancer Survivor Study	National Alcohol Survey-2000
N.8 Have you had at least one drink of beer,	Use B4 (below), but for those who answered,
wine, or liquor during the past year?	less than once a month, then ask:
No	B5. Think back over the last year, since
Yes	(current date last year). Did you have a whole
	drink of any alcoholic beverage like wine, beer,
	or liquor in these last twelve months?
N.6 During the past 2 years, on average how	B4. How often do you have any kind of
many times per month did you drink the	beverage containing alcohol—whether it is
following: (listed separately by wine, beer and	wine, beer, whiskey, or any other drink? Is it
mixed drink)	(average over past 12 months)
Responses:	Responses:
0-999	3+ daily
	2+ daily
	once a day
	nearly every day
	three or four times a week
	once or twice a week
	Two or three times a month
	About once a month
	Less than once a month
N.7 On the days that you drink, on average,	(Skips lifelong abstainer or ex-drinker)
how many drinks do you have?	L15. How many drinks did you have at that
No drinks in past 2 years	time? (By "drink", we mean one can or bottle
One drink/day	of beer, one bottle or wine cooler, one glass of
Two drinks/day	wine or a drink with a shot of hard liquor.
Three drinks/day	
Four drinks/day	# of drinks
Five drinks/day	
six or more drinks/day	
N.4 How old were you when you stated	B31. About how old were you when you first
drinking?	started drinking alcoholic beverages, not
Years old	including small tastes?
	AGE

Risky drinking is a variable which we can construct from existing questions about quantity and frequency of alcohol consumption^{*}. Risky drinking is defined by the United States Department of Health and Human Services and published in *The Physicians' Guide to Helping Patients with Alcohol Problems*, (1995) and specifies drinking more than three drinks per day <u>or</u> seven drinks per week for women and more than four drinks per day <u>or</u> more than fourteen drinks per week for men—at least one day in the last year, to be risky drinking.[United States Department of Health and Human Services, 1995 #462] This definition for risky drinking was chosen since it balances both daily and weekly drinking guidelines and best reflects the health impact of drinking patterns.[Dawson, 2000 #460] Specifically, this measure best balanced the need for high sensitivity to outcomes such as alcohol dependence, impaired driving, liver disease, peptic ulcer and hypertension without decreasing specificity to an unacceptable standard.[Dawson, 2000 #460] Heavy drinking is defined as five or more drinks per day at least once a month and can also be constructed from existing questions in the CCSS and the NAS.

PREDICTOR VARIABLES

Data will be collected in the Baseline survey or the Medical Abstraction Form

1.) HEALTH STATUS: Questions from CCSS

Six domains of Health Status will be employed including: general health, mental health, functional impairment, activity limitations, pain, or anxiety/fears as a result of the cancer or its treatment. Scoring for adverse health status for each domain will be constructed to be consistent with the use of these measure in Hudson et al, 2003.[Hudson, 2003 #461]

Domaine	Item#	Item Wording
Mental Health-based on Brief	J.16-J.35	Includes a global measure, (global severity index),
Symptom Inventory-18		and sub-scales for depression, anxiety and
		somatization
Pain as a result of cancer and	J.36	"Do you currently have pain as a result of our
its treatment		cancer, leukemia, tumor or similar illness, or its
		treatment?"
Anxiety/fears as a result of the	J.37	"Do you currently have anxieties/fears as a result of
cancer or its treatment		your cancer, leukemia, tumor or similar illness, or
		its treatment?"
Functional status (from	N.10-	"Because of any impairment or health problems, do
National Health Interview	N.12	you need the help of other persons with personal
Survey)		care needs, such as eating, bathing, dressing, or
		getting around your home?"
		"Because of any impairment or health problems, do
		you need the help of other persons in handling
		routine needs, such as everyday household chores,
		doing necessary business, shopping, or getting around for other purposes."
		"Does any impairment or health problem keep you
		from holding a job or attending school?"
Limitations of activity, (from	N.14	See instrument for exact question wording
Behavioral Risk Factor		
Surveillance System Survey		
Questionnaire)		
Self-assessed health	N.15	Would you say that your health is: Excellent, very
		good, good, fair, poor.

^{*} Description of survey questions for both instruments is available in Appendix A.

Question	Item #	Wording
Age	A.1	
Gender	A.2	What is your sex?
Race/ethnicity	A.4, A.4a	
Income	Q.8, Q.9	Over the last year, what is the total income of the
		household you live in?
		Over the last year, what is your personal income?
Education	0.1	What is the highest grade or level of schooling that
		you have completed? (made dichotomous, high
Special educational programs	0.3	school or less and more than high school)
Learning disabled		O.3 In elementary, junior, or high school were you
Advance placement		ever in any of the following programs?
Homebound education		(learning disabled or special education, advanced
		placement or homebound education > 1 year)
Cancer		Information collected from each institution
Age of diagnosis		Information collected from each institution (not in
		baseline survey)
Methotrexate, IT	Chemo II	Medical Abstraction Form, Chemotherapy,
		Question II.
Cranial radiation		Medical Abstraction Form, Radiation Therapy,
		(p. 10-11)

Additional control variables

APPENDIX B

PROPOSED TABLES FOR Characterization of drinking behaviors in survivors of childhood cancer compared to a general population

Table 1: Prevalence of current drinking, risky drinking and heavy drinking described in CCSS and NAS-2000. Table will include prevalence described by survivor, sibling and general population and subdivided by gender, age and racial/ethnic groups.

Variable	-	-Survivo	rs N=	CCS	S-Sibling	s N=		NAS N=	
	% rep	orting dr	rinking % reporting drin		ing drinki	ing types			ng types
	types								
	Current	Risky	Heavy	Current	Risky	Heavy	Current	Risky	Heavy
Total Population									
Gender									
Female									
Male									
Race/ethnicity									
White, non-									
Hispanic									
Black, non-									
Hispanic									
Hispanic									
Other									
Age at interview									
18-20									
20-29									
30-39									
40+									
Age at diagnosis									
0-4									
5-9									
10-14									
15-21									

A: statistically significant differences ($p\leq.05$) between survivors and siblings.

B: statistically significant differences ($p \le .05$) between survivors and NAS.

C: statistically significant differences ($p\leq.05$) between siblings and NAS.

Variable	CCSS-Survivors N= OR (95% CI)		
	Risky	Heavy	
Gender			
Female (ref)			
Male			
Race/ethnicity			
White, non-Hispanic (ref)			
Black, non-Hispanic			
Hispanic			
Other			
Age at interview			
18-20			
20-29			
30-39			
40+ (ref)			
Household Income			
<\$9,999			
\$10,000-19,999			
\$20,000-39,000			
\$40,000-59,999			
over \$60,000			
Age at diagnosis			
0-4			
5-9 (ref)			
10-14			
15-21			
Education			
High school or less			
More than High school			
Educational programs			
Special Education			
Advanced Placement			
Homebound ≥ 1 year			
IT Methotrexate or Cranial radiation			
Adverse Health, Any Domain			
General Health			
Mental Health			
Functional Impairment			
Activity Limitations			
Pain			
Anxiety about Cancer			
Cancer Diagnosis			
Leukemia			
Central Nervous System			
Hodgkin disease			
Non-hodgkin Lymphoma			
Wilms Tumor			
Neuroblastoma			

Table 2: Factors associated with risky drinking and heavy drinking; bivariate associations; OR (95% CI).

Sarcoma	
Bone	
Household Income	
<\$9,999	
\$10,000-19,999	
\$20,000-39,000	
\$40,000-59,999	
over \$60,000	

Table 3: Two models examining factors associated with risky drinking and heavy drinking. Model 1 examines known risk factors in general populations. Model 2 examines cancer specific variables. Model 2 is created using items starting with and below "age at diagnosis." Variables will be added one at a time using forward stepwise regression techniques and retaining those variables that make a significant contribution to the model.

Variable	CCSS-Survivors N= OR (95% CI)			CCSS-Survivors N= OR (95% CI)		
		Risky Drinking		Heavy Drinking		
Total Population	Model 1	Model 2	Model 1	Model 2		
Age at interview						
18-20	Xx	Xx	Xx	Xx		
20-29	Xx	Xx	Xx	Xx		
30-39	Xx	Xx	Xx	Xx		
40+	ref	ref	ref	ref		
Race/ethnicity						
White, non-Hispanic	ref	ref	ref	ref		
Black, non-Hispanic	Xx	Xx	Xx	Xx		
Hispanic	Xx	Xx	Xx	Xx		
Other	XX	XX	XX	XX		
Gender						
Female	ref	ref	ref	ref		
Male	XX	XX	XX	XX		
Household Income						
<\$9,999	Xx	Xx	Xx	Xx		
\$10,000-19,999	Xx	Xx	Xx	Xx		
\$20,000-39,000	Xx	Xx	Xx	Xx		
\$40,000-59,999	Xx	Xx	Xx	Xx		
over \$60,000	ref	ref	ref	ref		
Education						
High school or less	Xx	Xx	Xx	Xx		
More than High school	ref	ref	ref	ref		
Educational programs						
Special Education	Xx	Xx	Xx	Xx		
Advanced Placement	Ref	Ref	Ref	Ref		
Homebound ≥ 1 year	XX	XX	XX	XX		
Age at diagnosis						
0-4		Xx		Xx		
5-9 (ref)		Ref		Ref		
10-14		Xx		Xx		
15-21		XX		XX		
Adverse Health, Any Domain (vs	8					

better health-ref)		
General Health	Xx	Xx
Mental Health	Xx	Xx
Depression	Xx	Xx
Anxiety	Xx	Xx
Somatization	Xx	Xx
Functional Impairment	Xx	Xx
Activity Limitations	Xx	Xx
Pain	Xx	Xx
Anxiety about Cancer	XX	XX
IT Methotrexate or Cranial radiation	XX	XX
Cancer Diagnosis		
Leukemia	Ref	Ref
Central Nervous System	Xx	Xx
Hodgkin disease	Xx	Xx
Non-hodgkin Lymphoma	Xx	Xx
Wilms Tumor	Xx	Xx
Neuroblastoma	Xx	Xx
Sarcoma	Xx	Xx
Bone	XX	XX