

1995

- CCSS had 0 publication
- Internet access by Dial-up
- Amazon.com sold its 1st book online
- I was a new PhD & got the great opportunity to work with CCSS founders



2001

- CCSS had published 5 papers
- Amazon.com turned its first profit

Norm Breslow

A CCSS founder who, with John Potter,
recruited me to work on CCSS in **1995**

“He really did lay the foundation for all modern statistical methods in epidemiology and public health,”
said Ron Brookmeyer, a UCLA biostatistics professor.



Second Malignant Neoplasms in Five-Year Survivors of Childhood Cancer: Childhood Cancer Survivor Study

Joseph P. Neglia, Debra L. Friedman, Yutaka Yasui, Ann C. Mertens, Sue Hammond, Marilyn Stovall, Sarah S. Donaldson, Anna T. Meadows, Leslie L. Robison



**If I am the reviewer, I
would reject this paper.**



Working Group => Gate of entry to CCSS

**PLEASE CONTACT ME IF YOU HAVE
ANY INTEREST OR IDEA ON POTENTIAL
EPI/BIOSTAT PROJECTS**

Yutaka.Yasui@stjude.org

Epidemiology/Biostatistics Working Group Report

**CCSS Investigators Meeting
June, 2017**

- 1. Scope of your Working Group**
- 2. Publications since the 2015 mtg**
- 3. Ongoing work**
- 4. Future focus**

- 1. Scope of your Working Group**
2. Publications since the 2015 mtg
3. Ongoing work
4. Future focus

- **Methodological and “other” studies**
- **Analysis of CCSS mortality data**
- **Integrity/innovation of population-science methodology (with Wendy, Ann, Les, ...), working with other WGs**

1. Scope of your Working Group
- 2. Publications since the 2015 mtg**
3. Ongoing work
4. Future focus

Publications 264

The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

Reduction in Late Mortality among 5-Year Survivors of Childhood Cancer

Gregory T. Armstrong, M.D., M.S.C.E., Yan Chen, M.M., Yutaka Yasui, Ph.D., Wendy Leisenring, Sc.D., Todd M. Gibson, Ph.D., Ann C. Mertens, Ph.D., Marilyn Stovall, Ph.D., Kevin C. Oeffinger, M.D., Smita Bhatia, M.D., M.P.H., Kevin R. Krull, Ph.D., Paul C. Nathan, M.D., Joseph P. Neglia, M.D., M.P.H., Daniel M. Green, M.D., Melissa M. Hudson, M.D., and Leslie L. Robison, Ph.D.

ASSOCIATED PRESENTATION

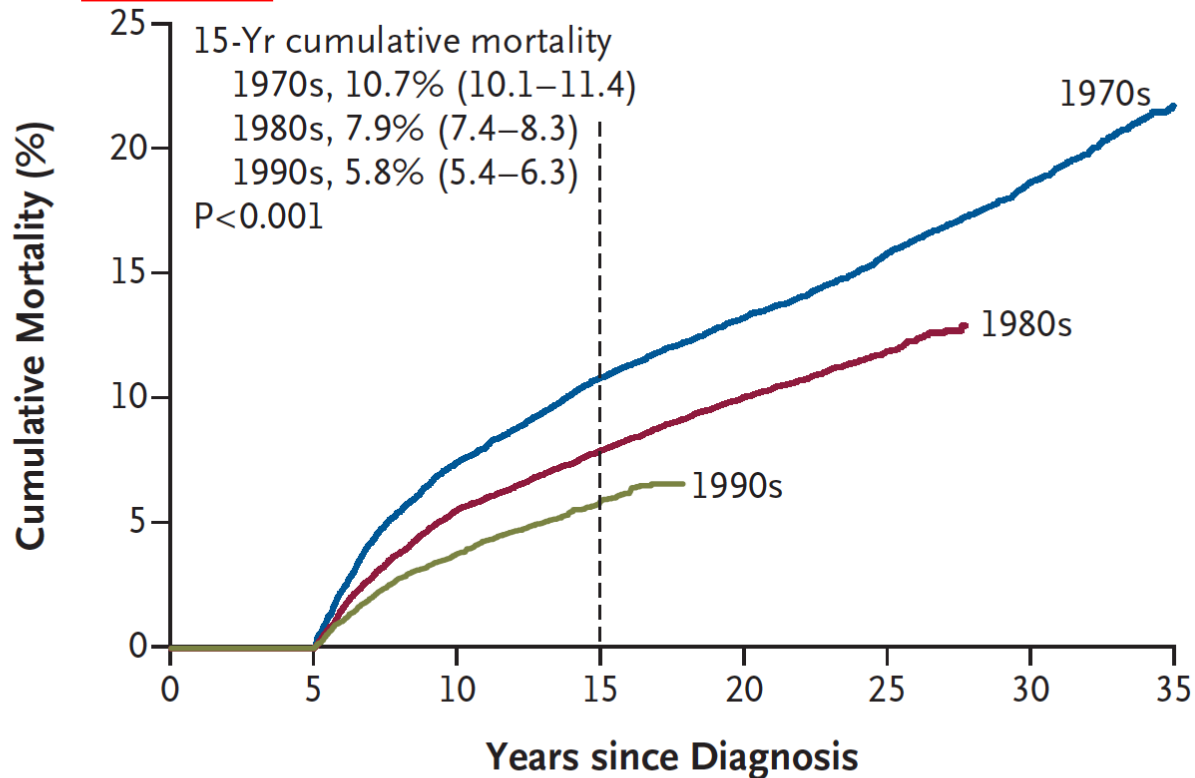


Meeting: **2015 ASCO
Annual Meeting**
Presenter: **Gregory T.
Armstrong**

[View Video](#)

Childhood Cancer Survivor Study Epidemiology/Biostatistics Working Group

A Death from Any Cause

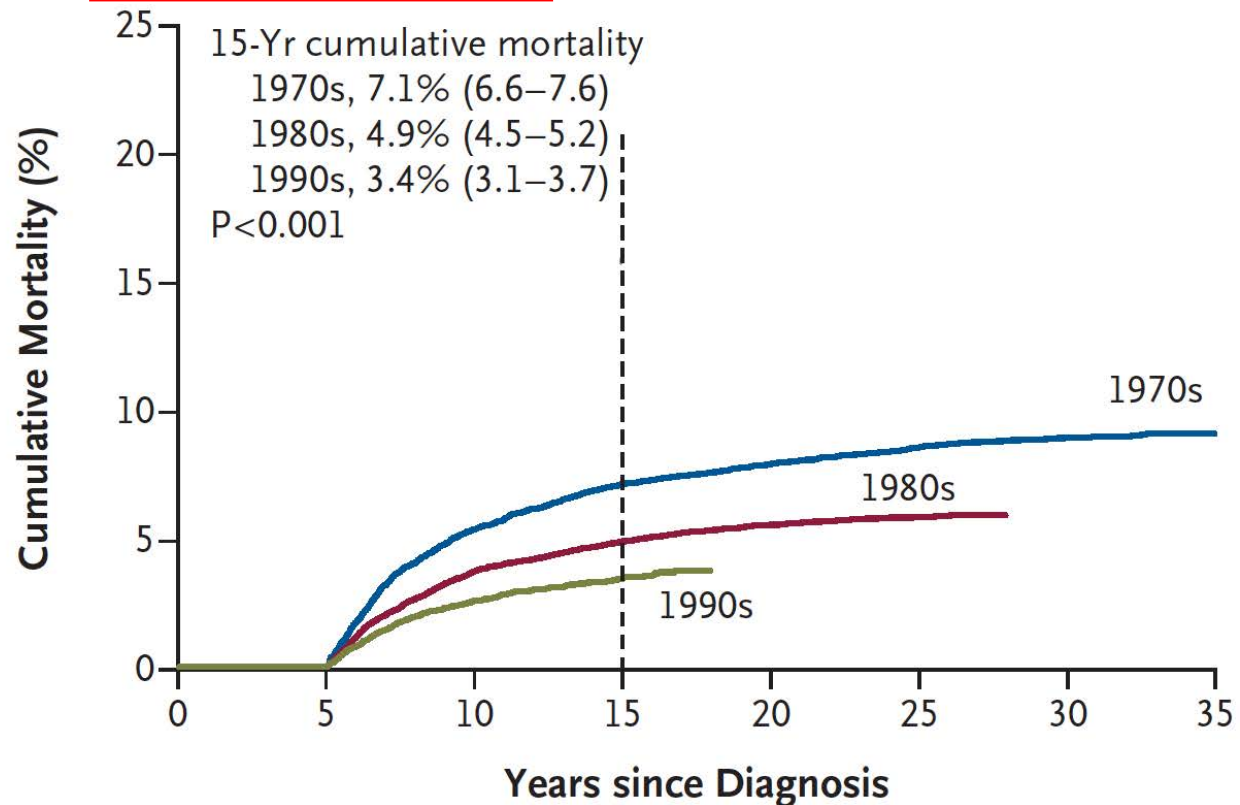


No. at Risk

1970s	9,416	8,722	8,406	8,182	7942	5556	1506
1980s	13,181	13,443	13,105	10,389	3583		
1990s	11,436	11,411	3,924				

Childhood Cancer Survivor Study Epidemiology/Biostatistics Working Group

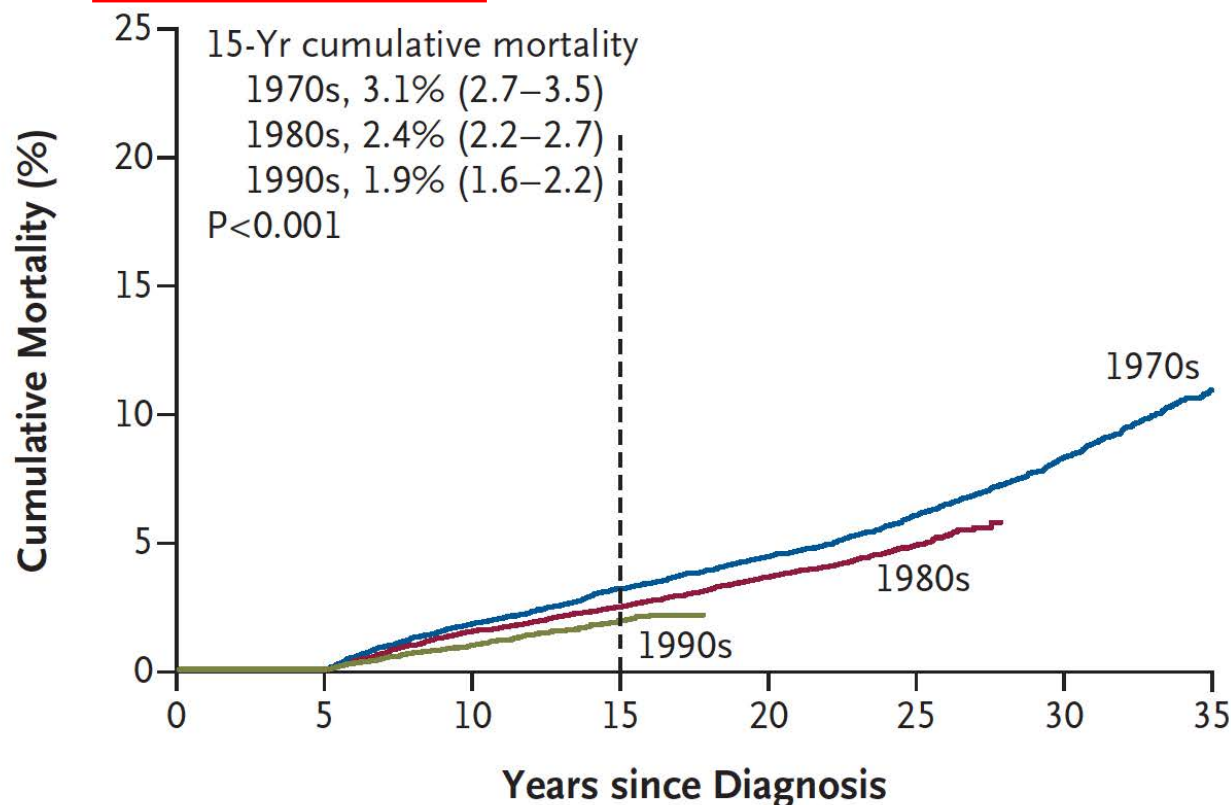
B Death from Recurrence or Progression



No. at Risk

1970s	9,416	8,722	8,406	8,182	7942	5556	1506
1980s	13,181	13,443	13,105	10,389	3583		
1990s	11,436	11,411	3,924				

C Death from Health-Related Cause



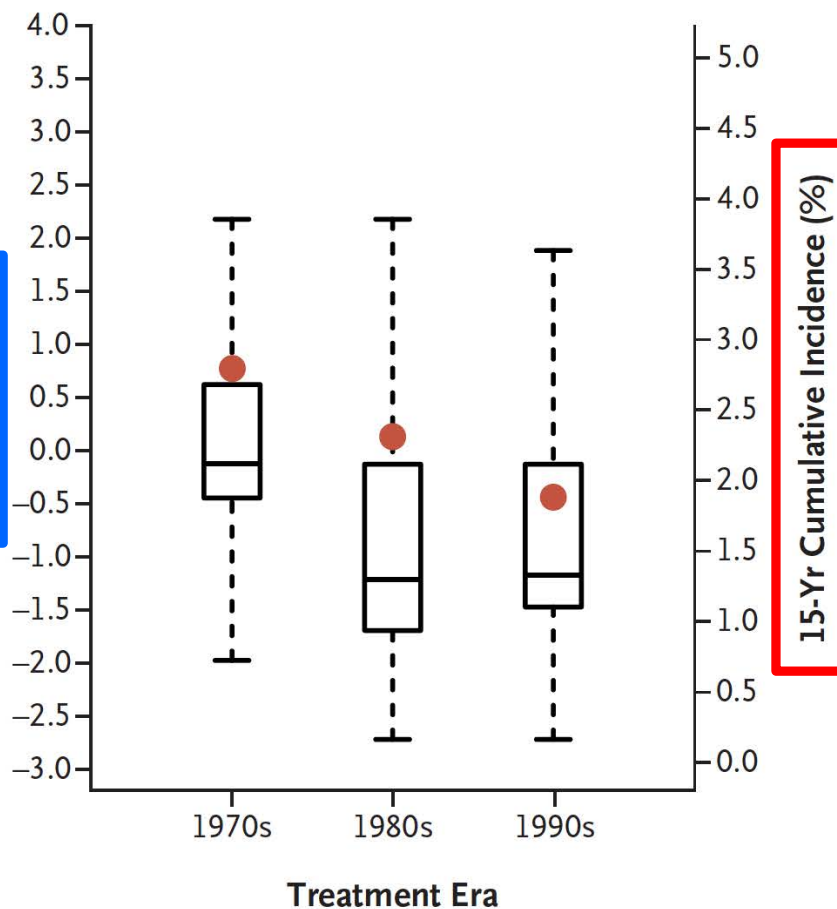
No. at Risk

1970s	9,416	8,722	8,406	8,182	7942	5556	1506
1980s	13,181	13,443	13,105	10,389	3583		
1990s	11,436	11,411	3,924				

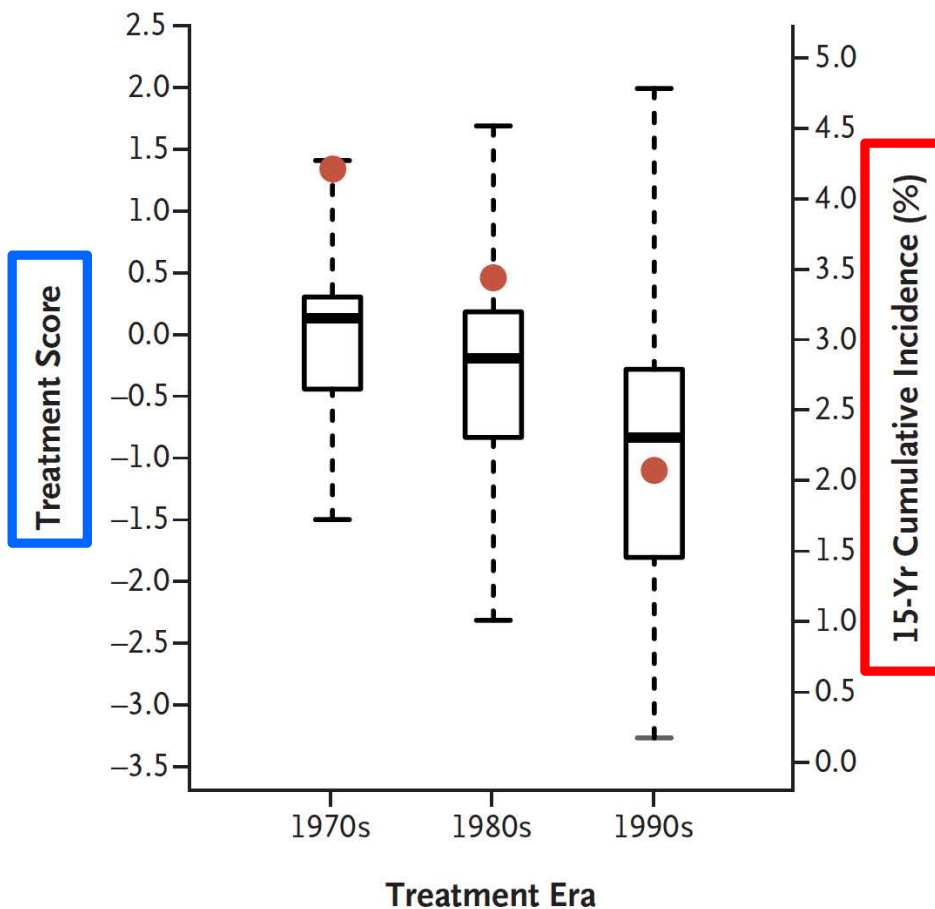
- **Temporal trends papers** (Armstrong *et al.* NEJM 2016, Turcotte *et al.* JAMA 2017, Ness *et al.* Ann Intern Med 2017)
- **Show trends in the outcome of interest**
- **Show trends in relevant treatment**
- **“Is it really the therapy changes that led to the outcome changes?”**

Childhood Cancer Survivor Study Epidemiology/Biostatistics Working Group

A Acute Lymphoblastic Leukemia

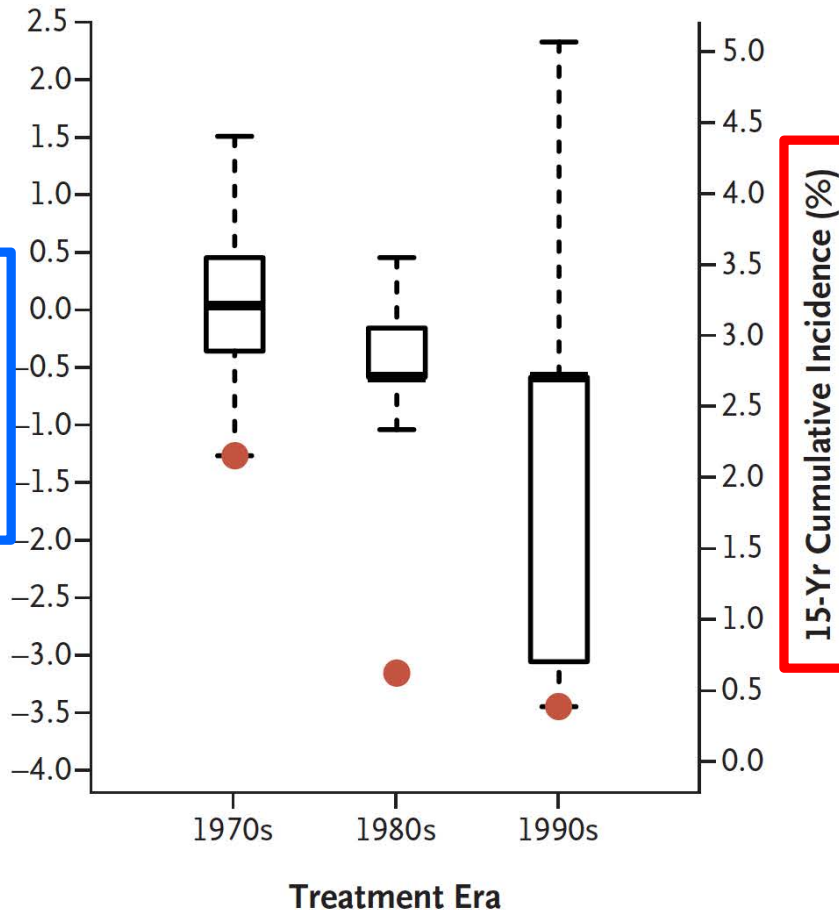


B Hodgkin's Lymphoma

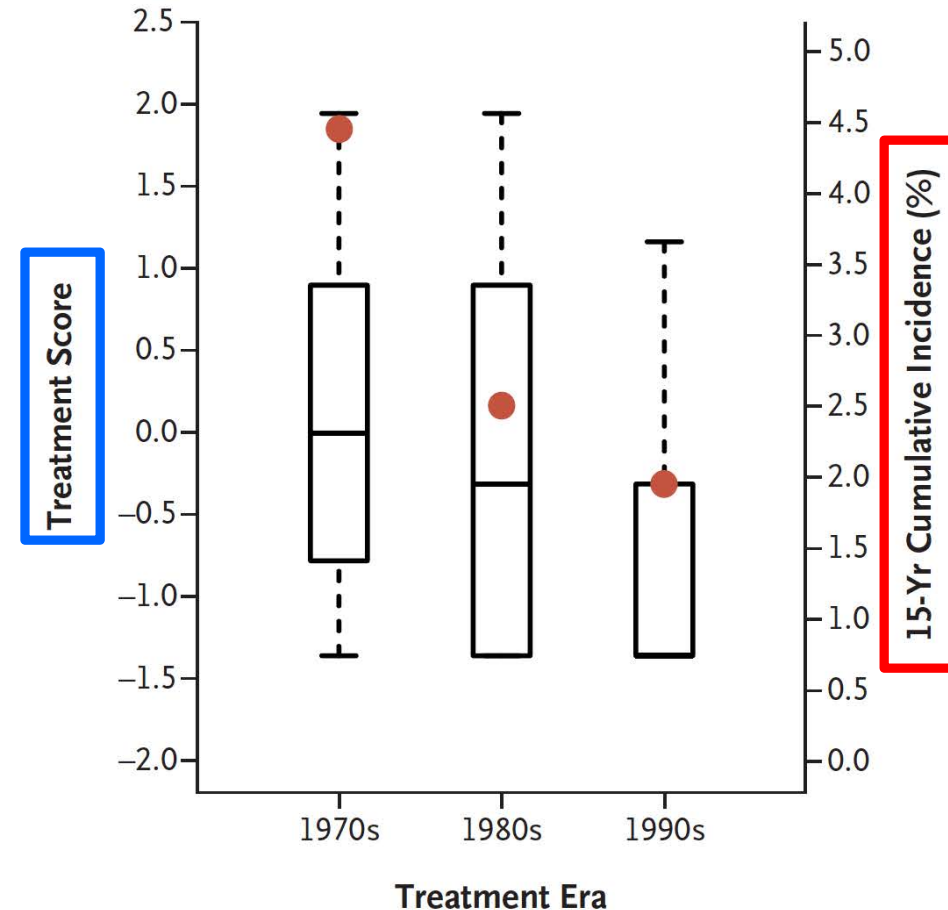


Childhood Cancer Survivor Study Epidemiology/Biostatistics Working Group

C Wilms' Tumor



D Astrocytoma



**Acute
Lymphoblastic
Leukemia**

Relative rate of mortality

No adjustment for therapy	0.88 (0.81– 0.95)
---------------------------	-------------------

**Every 5 years of treatment era:
12% lower mortality rate**

**Acute
Lymphoblastic
Leukemia**

No adjustment for therapy	0.88 (0.81– 0.95)
Adjustment for therapy	1.02 (0.83–1.24)†

† Data were adjusted for cranial radiotherapy dose, anthracycline dose, and exposure to epipodophyllotoxins and glucocorticoids.

Publications 274

Communication

Childhood Cancer Survivorship Research in Minority Populations: A Position Paper From the Childhood Cancer Survivor Study

Smita Bhatia, MD, MPH¹; Todd M. Gibson, PhD²; Kirsten K. Ness, PhD²; Qi Liu, MS³; Kevin C. Oeffinger, MD^{4,5};
Kevin R. Krull, PhD²; Paul C. Nathan, MD, MSc⁶; Joseph P. Neglia, MD, MPH⁷; Wendy Leisenring, ScD⁸; Yutaka Yasui, PhD^{2,3};
Leslie L. Robison, PhD²; and Gregory T. Armstrong, MD, MSCE²

Publications 268

VOLUME 34 • NUMBER 14 • MAY 10, 2016

JOURNAL OF CLINICAL ONCOLOGY

O R I G I N A L R E P O R T

Racial/Ethnic Differences in Adverse Outcomes Among Childhood Cancer Survivors: The Childhood Cancer Survivor Study

*Qi Liu, Wendy M. Leisenring, Kirsten K. Ness, Leslie L. Robison, Gregory T. Armstrong, Yutaka Yasui, and
Smita Bhatia*

**Hispanic
vs.
NHW**

**African American
vs.
NHW**

Socioeconomic status

Annual household income <20,000

Education <high school

With health insurance

3.5 (3.1–3.9); $P < .001$

1.5 (1.3–1.7); $P < .001$

0.7 (0.6–0.8); $P < .001$

0.5 (0.5–0.6); $P < .001$

0.5 (0.5–0.6); $P = .001$

1.6 (1.3–2.0); $P < .001$

	Hispanic vs. NHW	African American vs. NHW
Socioeconomic status		
Annual household income <20,000	1.6 (1.5–1.8); $P < .001$	3.5 (3.1–3.9); $P < .001$
Education <high school	1.3 (1.1–1.5); $P < .001$	1.5 (1.3–1.7); $P < .001$
With health insurance	0.5 (0.5–0.6); $P < .001$	0.7 (0.6–0.8); $P < .001$
Risky health behaviors		
Current smokers	0.6 (0.6–0.8); $P < .001$	0.5 (0.5–0.6); $P < .001$
Alcohol consumption (binge drinking)	1.1 (1.0–1.2); $P = .07$	0.5 (0.5–0.6); $P = .001$
Physically inactive ^b	1.0 (0.8–1.2); $P = .8$	1.6 (1.3–2.0); $P < .001$

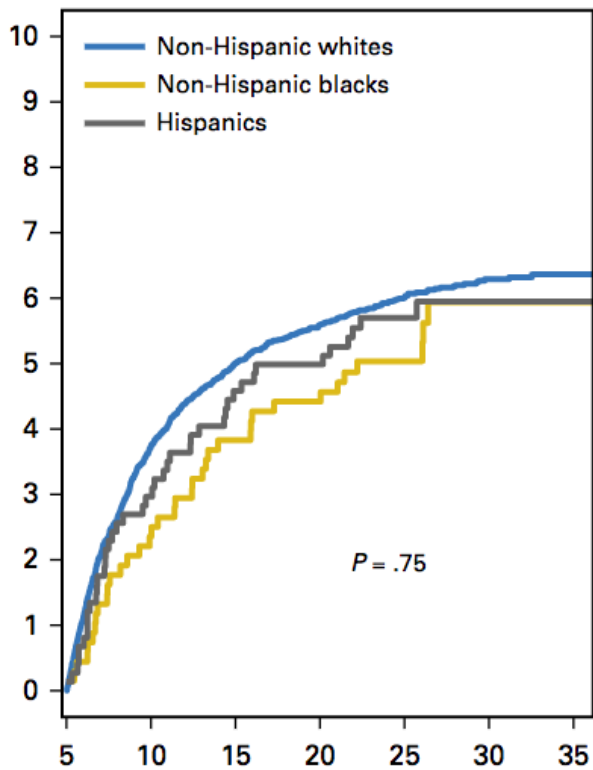
	Hispanic vs. NHW	African American vs. NHW
Socioeconomic status		
Annual household income <20,000	1.6 (1.5–1.8); $P < .001$	3.5 (3.1–3.9); $P < .001$
Education <high school	1.3 (1.1–1.5); $P < .001$	1.5 (1.3–1.7); $P < .001$
With health insurance	0.5 (0.5–0.6); $P < .001$	0.7 (0.6–0.8); $P < .001$
Risky health behaviors		
Current smokers	0.6 (0.6–0.8); $P < .001$	0.5 (0.5–0.6); $P < .001$
Alcohol consumption (binge drinking)	1.1 (1.0–1.2); $P = .07$	0.5 (0.5–0.6); $P = .001$
Physically inactive ^b	1.0 (0.8–1.2); $P = .8$	1.6 (1.3–2.0); $P < .001$
Obesity	1.6 (1.4–1.7); $P < .001$	1.6 (1.4–1.8); $P < .001$
Hypertension	1.1 (0.9–1.2); $P = .3$	1.3 (1.2–1.5); $P < .001$
Diabetes	1.7 (1.4–2.1); $P < .001$	2.3 (1.9–2.9); $P < .001$
	1.4 (1.1–1.8); $P = .003$	1.9 (1.5–2.3); $P < .001$
	0.9 (0.6–1.3); $P = .6$	1.9 (1.4–2.5); $P < .01$
	0.8 (0.5–1.1); $P = .2$	1.5 (1.1–2.0); $P = .01$

	Hispanic vs. NHW	African American vs. NHW
Socioeconomic status		
Annual household income <20,000	1.6 (1.5–1.8); $P < .001$	3.5 (3.1–3.9); $P < .001$
Education <high school	1.3 (1.1–1.5); $P < .001$	1.5 (1.3–1.7); $P < .001$
With health insurance	0.5 (0.5–0.6); $P < .001$	0.7 (0.6–0.8); $P < .001$
Risky health behaviors		
Current smokers	0.6 (0.6–0.8); $P < .001$	0.5 (0.5–0.6); $P < .001$
Alcohol consumption (binge drinking)	1.1 (1.0–1.2); $P = .07$	0.5 (0.5–0.6); $P = .001$
Physically inactive ^b	1.0 (0.8–1.2); $P = .8$	1.6 (1.3–2.0); $P < .001$
Obesity	1.6 (1.4–1.7); $P < .001$	1.6 (1.4–1.8); $P < .001$
Hypertension	1.1 (0.9–1.2); $P = .3$	1.3 (1.2–1.5); $P < .001$
Diabetes	<u>1.7 (1.4–2.1); $P < .001$</u>	<u>2.3 (1.9–2.9); $P < .001$</u>
Diabetes ^c	<u>1.4 (1.1–1.8); $P = .003$</u>	<u>1.9 (1.5–2.3); $P < .001$</u>
	0.9 (0.6–1.3); $P = .6$	1.9 (1.4–2.5); $P < .01$
	0.8 (0.5–1.1); $P = .2$	1.5 (1.1–2.0); $P = .01$

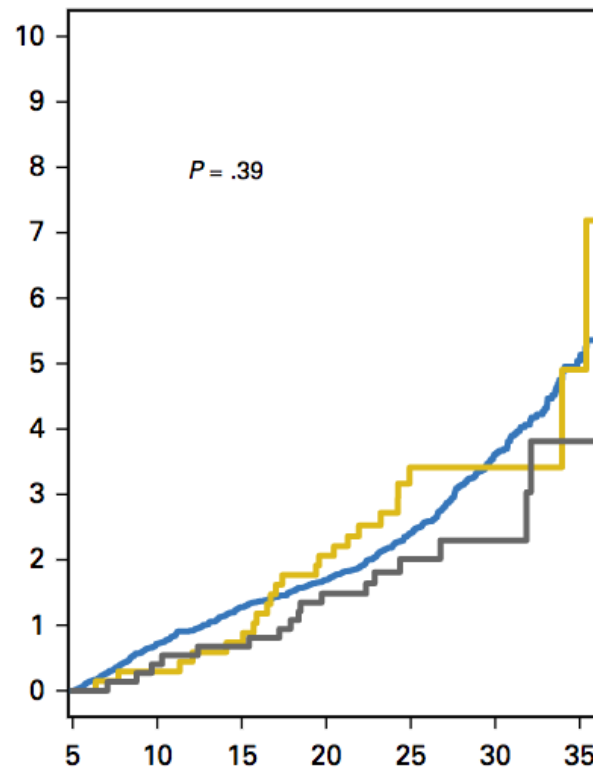
	Hispanic vs. NHW	African American vs. NHW
Socioeconomic status		
Annual household income <20,000	1.6 (1.5–1.8); $P < .001$	3.5 (3.1–3.9); $P < .001$
Education <high school	1.3 (1.1–1.5); $P < .001$	1.5 (1.3–1.7); $P < .001$
With health insurance	0.5 (0.5–0.6); $P < .001$	0.7 (0.6–0.8); $P < .001$
Risky health behaviors		
Current smokers	0.6 (0.6–0.8); $P < .001$	0.5 (0.5–0.6); $P < .001$
Alcohol consumption (binge drinking)	1.1 (1.0–1.2); $P = .07$	0.5 (0.5–0.6); $P = .001$
Physically inactive ^b	1.0 (0.8–1.2); $P = .8$	1.6 (1.3–2.0); $P < .001$
Chronic conditions		
Obesity	1.6 (1.4–1.7); $P < .001$	1.6 (1.4–1.8); $P < .001$
Hypertension	1.1 (0.9–1.2); $P = .3$	1.3 (1.2–1.5); $P < .001$
Diabetes	1.7 (1.4–2.1); $P < .001$	2.3 (1.9–2.9); $P < .001$
Diabetes ^c	1.4 (1.1–1.8); $P = .003$	1.9 (1.5–2.3); $P < .001$
Stroke	0.9 (0.6–1.3); $P = .6$	1.9 (1.4–2.5); $P < .01$
Stroke ^c	0.8 (0.5–1.1); $P = .2$	1.5 (1.1–2.0); $P = .01$

Cumulative cause-specific mortality

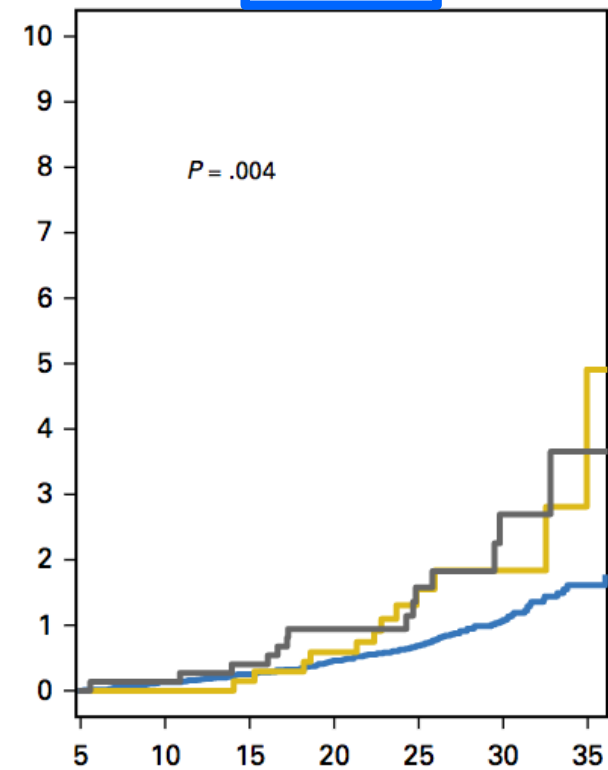
Recurrence



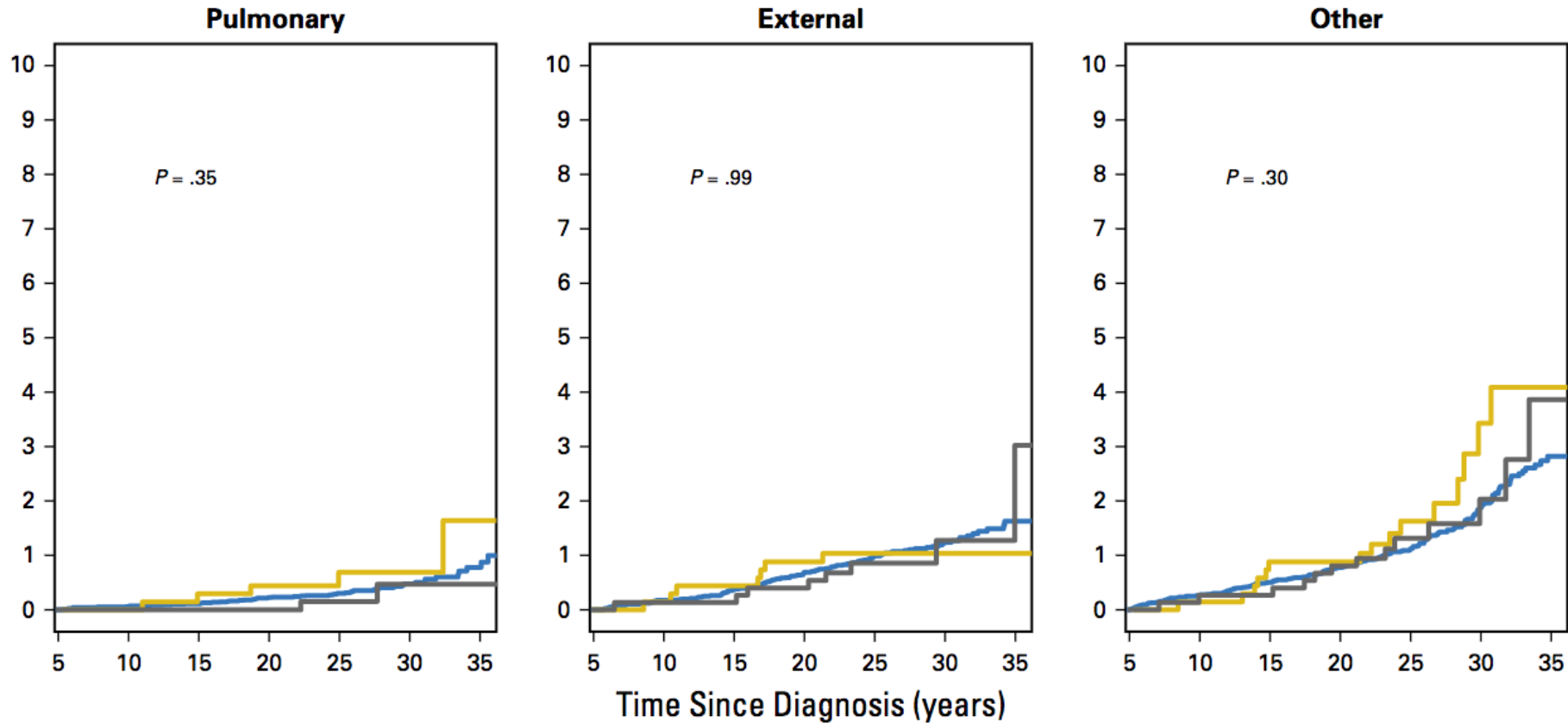
SN



Cardiac



Cumulative cause-specific mortality



RR of Mortality	All-Cause	
	RR (95% CI)	<i>P</i>
Adjusted for clinical/demographic variables†		

- By and large, **comparable burden** of morbidity and mortality
- A few differences in risk were **explained by differences in socioeconomic status**

clinical/demographic variables, treatment, and <u>SES</u> §		
NHW	Ref	
NHB	1.0 (0.8 to 1.4)	.88
Hispanic	0.9 (0.7 to 1.2)	.46

Health care use, adjusted for education, income, and insurance

General physical examination

Cancer-related care

Surveillance for long-term toxicities, adjusted for education, income, and insurance

Mammography in women treated with chest irradiation

Echocardiogram in anthracycline-exposed survivors

**Hispanic
vs.
NHW**

1.1 (1.0–1.2); $P = .1$

1.0 (0.9–1.1); $P = .8$

0.9 (0.5–1.4); $P = .6$

1.0 (0.8–1.1); $P = .6$

Health care use, adjusted for education, income, and insurance

General physical examination

Cancer-related care

Surveillance for long-term toxicities, adjusted for education, income, and insurance

Mammography in women treated with chest irradiation

Echocardiogram in anthracycline-exposed survivors

Hispanic
vs.
NHW

1.1 (1.0–1.2); $P = .1$

1.0 (0.9–1.1); $P = .8$

0.9 (0.5–1.4); $P = .6$

1.0 (0.8–1.1); $P = .6$

African American
vs.
NHW

1.2 (1.0–1.4); $P = .02$

0.7 (0.6–0.8); $P < .001$

0.6 (0.3–1.1); $P = .09$

0.8 (0.6–0.9); $P = .01$

Publications 254

VOLUME 33 • NUMBER 32 • NOVEMBER 10 2015

JOURNAL OF CLINICAL ONCOLOGY

O R I G I N A L R E P O R T

Equivalence Ratio for Daunorubicin to Doxorubicin in Relation to Late Heart Failure in Survivors of Childhood Cancer

Elizabeth A.M. Feijen, Wendy M. Leisenring, Kayla L. Stratton, Kirsten K. Ness, Helena J.H. van der Pal, Huib N. Caron, Gregory T. Armstrong, Daniel M. Green, Melissa M. Hudson, Kevin C. Oeffinger, Leslie L. Robison, Marilyn Stovall, Leontien C.M. Kremer, and Eric J. Chow

Childhood Cancer Survivor Study Epidemiology/Biostatistics Working Group

Table 3. HRs for Heart Failure Based on Doxorubicin and Daunorubicin Dose Categories

		Daunorubicin		
Model and Dose Category	Dose, mg/m ²		HR*	95% CI
	Median	IQR		
Primary model*				
None			Reference	
≤ 0.1 to < 200 mg/m ²	100	75-118	1.09	0.57 to 2.08
≤ 200 to < 300 mg/m ²	246	221-270	3.16	1.16 to 8.61
≤ 300 to < 400 mg/m ²	350	328-371	4.33	1.73 to 10.84
≤ 400 mg/m ²	480	432-545	10.72	5.13 to 22.42
Secondary model†				
None			Reference	
≤ 0.1 to < 150 mg/m ²	99	51-103	1.35	0.18 to 10.42
≤ 150 to < 300 mg/m ²	213	183-251	2.83	0.37 to 21.57
≤ 300 mg/m ²	379	346-449	20.17	8.83 to 46.06

Abbreviations: HR, hazard ratio; IQR, interquartile range.

*Model was adjusted for sex; age at diagnosis; chest radiotherapy dose; and exposure to another anthracycline besides doxorubicin or daunorubicin, such as epirubicin, idarubicin, or mitoxantrone. It was also stratified by cohort.

†Model was adjusted for sex and age at diagnosis and was also stratified by cohort.

‡The CIs of the ratios are the 2.5th and 97.5th percentiles of the bootstraps.

Childhood Cancer Survivor Study

Epidemiology/Biostatistics Working Group

Table 3. HRs for Heart Failure Based on Doxorubicin and Daunorubicin Dose Categories

Model and Dose Category	Daunorubicin				Doxorubicin			
	Dose, mg/m ²		HR*	95% CI	Dose, mg/m ²		HR	95% CI
	Median	IQR			Median	IQR		
Primary model*								
None			Reference				Reference	
≤ 0.1 to < 200 mg/m ²	100	75-118	1.09	0.57 to 2.08	122	80-169	2.80	1.75 to 4.49
≤ 200 to < 300 mg/m ²	246	221-270	3.16	1.16 to 8.61	253	226-278	6.31	4.11 to 9.69
≤ 300 to < 400 mg/m ²	350	328-371	4.33	1.73 to 10.84	347	318-370	13.19	9.04 to 19.25
≤ 400 mg/m ²	480	432-545	10.72	5.13 to 22.42	459	430-505	18.43	12.82 to 26.50
Secondary model†								
None			Reference				Reference	
≤ 0.1 to < 150 mg/m ²	99	51-103	1.35	0.18 to 10.42	102	71-122	3.97	1.14 to 13.76
≤ 150 to < 300 mg/m ²	213	183-251	2.83	0.37 to 21.57	211	180-258	9.29	4.58 to 18.86
≤ 300 mg/m ²	379	346-449	20.17	8.83 to 46.06	391	345-455	34.74	19.24 to 62.73

Abbreviations: HR, hazard ratio; IQR, interquartile range.

*Model was adjusted for sex; age at diagnosis; chest radiotherapy dose; and exposure to another anthracycline besides doxorubicin or daunorubicin, such as epirubicin, idarubicin, or mitoxantrone. It was also stratified by cohort.

†Model was adjusted for sex and age at diagnosis and was also stratified by cohort.

‡The CIs of the ratios are the 2.5th and 97.5th percentiles of the bootstraps.

Childhood Cancer Survivor Study Epidemiology/Biostatistics Working Group

Table 3. HRs for Heart Failure Based on Doxorubicin and Daunorubicin Dose Categories

Model and Dose Category	Daunorubicin				Doxorubicin				Daunorubicin-to-Doxorubicin Ratio			
	Dose, mg/m ²		HR*	95% CI	Dose, mg/m ²		HR	95% CI				
	Median	IQR			Median	IQR			Ratio	95% CI	Mean	95% CI
Primary model*											0.45	0.23 to 0.73
None			Reference				Reference		—			
≤ 0.1 to < 200 mg/m ²	100	75-118	1.09	0.57 to 2.08	122	80-169	2.80	1.75 to 4.49	0.39	0.04 to 0.78		
≤ 200 to < 300 mg/m ²	246	221-270	3.16	1.16 to 8.61	253	226-278	6.31	4.11 to 9.69	0.50	0.00 to 1.12		
≤ 300 to < 400 mg/m ²	350	328-371	4.33	1.73 to 10.84	347	318-370	13.19	9.04 to 19.25	0.33	0.03 to 0.62		
≤ 400 mg/m ²	480	432-545	10.72	5.13 to 22.42	459	430-505	18.43	12.82 to 26.50	0.58	0.09 to 1.12		
Secondary model†											0.41	0.29 to 1.28‡
None			Reference				Reference		—			
≤ 0.1 to < 150 mg/m ²	99	51-103	1.35	0.18 to 10.42	102	71-122	3.97	1.14 to 13.76	0.34	0.14 to 2.60‡		

Abb
*Mo
epiru
†Mc
‡Th

**Daunorubicin was less cardiotoxic than doxorubicin;
the daunorubicin-to-doxorubicin cardiotoxicity equivalence
ratio was between 0.4 and 0.5.**

ch as

Publications 293

Research Article

Statistics
in Medicine

Received 22 January 2016, Accepted 12 December 2016 Published online 18 January 2017 in Wiley Online Library

(wileyonlinelibrary.com) DOI: 10.1002/sim.7217

Regression analysis of mixed panel count data with dependent terminal events

Guanglei Yu,^a Liang Zhu,^b Yang Li,^c Jianguo Sun^{a*†}  and
Leslie L. Robison^d

$$\hat{\mu}_0(t; \beta, \hat{\omega}(t)) = \frac{\sum_{i=1}^n r_i(t) \hat{\omega}_i(t) \tilde{N}_i(t)}{n \tilde{S}_r^{(0)}(t; \beta, \hat{\omega}(t))}$$

and

$$\hat{\mu}_0(t; \beta, \hat{\omega}(t)) d\hat{\Gamma}_0(t; \beta, \hat{\omega}(t)) = \frac{\sum_{i=1}^n (1 - r_i(t)) \tilde{N}_i(t) d\tilde{O}_i(t)}{n \tilde{S}_p^{(0)}(t; \beta, \hat{\omega}(t))},$$

where

$$\begin{aligned} \tilde{S}_r^{(d)}(t; \beta, \hat{\omega}(t)) &= \frac{1}{n} \sum_{i=1}^n r_i(t) \hat{\omega}_i(t) \exp\{\beta^T Z_i\} Z_i^{\otimes d}, \\ \tilde{S}_p^{(d)}(t; \beta, \hat{\omega}(t)) &= \frac{1}{n} \sum_{i=1}^n (1 - r_i(t)) \hat{\omega}_i(t) \exp\{\beta^T Z_i\} Z_i^{\otimes d}, \quad d = 0, 1, 2, \end{aligned}$$

$\mathbf{a}^{\otimes 0} = 1, \mathbf{a}^{\otimes 1} = \mathbf{a}, \mathbf{a}^{\otimes 2} = \mathbf{a}\mathbf{a}'$, and $\boldsymbol{\omega}(t) = (\omega_1(t), \dots, \omega_n(t))'$. By plugging the two estimators earlier into Equation (5), we obtain

$$U(\beta) = \sum_{i=1}^n \int_0^\tau r_i(t) \hat{\omega}_i(t) (Z_i - \bar{Z}_r(t; \beta, \hat{\omega}(t))) \tilde{N}_i(t) dH(t) + (1 - r_i(t)) (Z_i - \bar{Z}_p(t; \beta, \hat{\omega}(t))) \tilde{N}_i(t) d\tilde{O}_i(t) = 0, \quad (6)$$

where $\bar{Z}_r(t; \beta, \hat{\omega}(t)) = \tilde{S}_r^{(1)}(t; \beta, \hat{\omega}(t)) / \tilde{S}_r^{(0)}(t; \beta, \hat{\omega}(t))$ and $\bar{Z}_p(t; \beta, \hat{\omega}(t)) = \tilde{S}_p^{(1)}(t; \beta, \hat{\omega}(t)) / \tilde{S}_p^{(0)}(t; \beta, \hat{\omega}(t))$.

Let $\hat{\beta}$ denote the estimator of β_0 given by solving Equation (6). To establish the asymptotic properties of $\hat{\beta}$, define $N_i^d(t) = I(D_i \leq t, D_i \leq C_i)$,

$$M_i^d(t; \delta) = N_i^d(t) - \int_0^t I(T_i^* \geq s) \exp\{\delta^T Z_i\} d\Delta_0(s),$$

and $\bar{Z}_d(t; \hat{\delta}) = \tilde{S}_d^{(1)}(t; \hat{\delta}) / \tilde{S}_d^{(0)}(t; \hat{\delta})$, where $\Delta_0(t) = \int_0^t \lambda_0^d(s) ds$ and

$$\tilde{S}_d^{(d)}(t; \delta) = \frac{1}{n} \sum_{i=1}^n I(T_i^* \geq t) \exp\{\delta^T Z_i\} Z_i^{\otimes d}, \quad d = 0, 1, 2.$$

Publications 293

- Regression analysis methodology
- Recurrent event data (e.g., BCC, pregnancy)
+
- Panel count data: observed at discrete time points

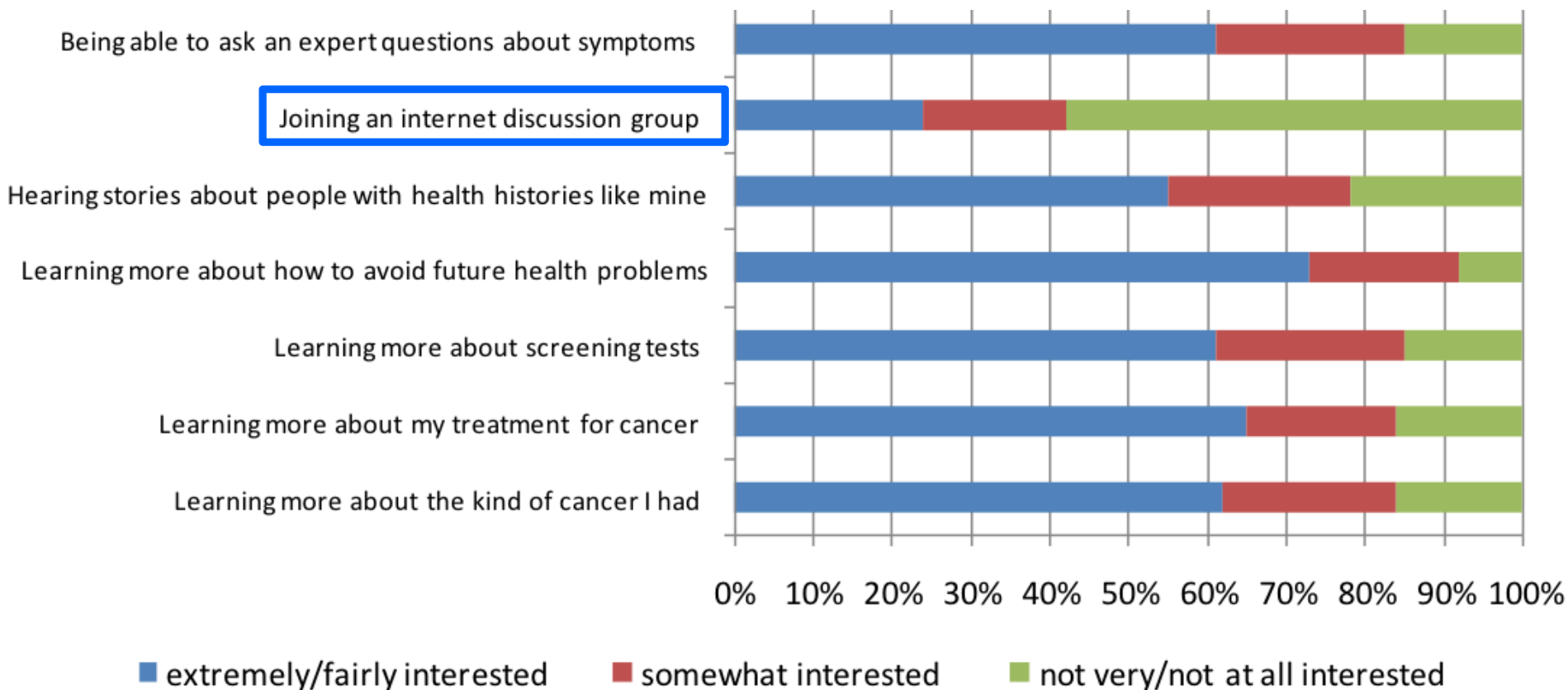
Submitted manuscript 1

|Where, When, and Why Adult Survivors of Childhood Cancer Seek Internet-based Health Information

Mechelle D Claridy¹, Melissa M Hudson^{2, 3}, Jeanne Steele⁴, Lee Caplan¹, Pauline A Mitby⁵,
Wendy Leisenring⁶, Selina A Smith⁷, Leslie L Robison², Ann C Mertens⁸

Childhood Cancer Survivor Study

Epidemiology/Biostatistics Working Group



Childhood Cancer Survivor Study Epidemiology/Biostatistics Working Group

In the past 12 months, have you looked for health or medical information for yourself while using the internet?	CCSS (n=1386) OR (95% CI)	HINTS (n=2385) OR (95% CI)
Overall	2.76 (2.40-3.19)	1.0

How much do you trust information about health or medical topics from sources listed below?	Doctor or Healthcare Professional	
	CCSS Mean (SD)	HINTS Mean (SD)
Total	3.77 (0.50)	3.63 (0.62)
	*p-value	
	< 0.01	

Submitted manuscript 2

Yuan Y, Zhou QM, Li B, Cai H, Chow EJ, Armstrong GT.

**A Threshold-free Prospective Prediction Accuracy
Measure for Censored Time to Event Data**

(invited for revision by Statistics in Medicine)

1. Scope of your Working Group
2. Publications since the 2015 mtg
- 3. Ongoing work**
4. Future focus

- **Address bias concern due to non-participation (FU Survey 5)**
- **Inverse Probability Weighting (IPW)** to boost contribution for survivors with characteristics of low participation
- Methodological paper that describes the application of IPW in CCSS

- **Tiled study design: using temporal overlap as a method to extend longitudinal follow-up among carefully selected time-limited cohorts ([Chow](#))**
- **Use of an incentive to increase biologic sample (Oragene) return rate ([McDonald](#))**
- **Radiation dose reconstruction methods for intensity modulated radiation therapy ([Howell](#))**
- **Handling Missing Data due to No Consent for Medical Record Abstraction by Multiple Imputation ([Martin/Liu](#))**

Funded Ancillary Projects

Principal Investigator: Liang Zhu (St. Jude Children's Research Hospital)

Title: New Methods to Address Dilemmas in Mixed Recurrent-event and

Dates of Funding: 7/16 - 6/18

Funding Source: National Institutes of Health (R21)

Award: \$501,312

Study Aims: To develop semi-parametric methods for regression analysis of recurrent event and compare them with alternative methods by simulation studies

Principal Investigator: Yan Yuan (University of Alberta)

Title: Risk Prediction Model of Premature Menopause in Childhood Cancer

Dates of Funding: 7/16 - 12/18

Funding Source: Canadian Institutes of Health Research

Award: \$179,858

Study Aims: To develop a prediction model for early menopause

Childhood Cancer Survivor Study **Epidemiology/Biostatistics Working Group**

Principal Investigator: Lennie Wong (City of Hope)

Title: Cost effectiveness of breast cancer screening guidelines for female survivors

Dates of Funding: 7/17 - 6/20

Funding Source: American Cancer Society

Award: \$527,000

Study Aims: 1) Examine the cost-effectiveness of 1) annual clinical breast exam vs. MRI as adjunct to mammography.

Principal Investigator: Yutaka Yasui, Jinghui Zhang (St. Jude Children's Research Hospital)

Title: Late Effects Prediction using Clinical Phenotypes and Whole Genome Sequencing

Dates of Funding: 4/17 - 3/22

Funding Source: National Institutes of Health, RO1

Award: \$3,457,455

Study Aims: 1) Build individual risk prediction models with the SJLIFE cohort for basal cell carcinoma, and multiple subsequent neoplasms, 2) Validate the risk prediction models in higher SN counts (CCSS)

Submitted Ancillary Projects

Principal Investigator: Liang Zhu (University of Texas MD Anderson)

Title: Statistical Analysis for Mixed Outcome Measures in Recurrent Event

Proposed Funding Source: American Cancer Society

Study Aims: Develop a likelihood-based semiparametric estimation method for recurrent event and panel-count data.

Status: Submitted October 2016, scored 20, awaiting funding decision

1. Scope of your Working Group
2. Publications since the 2015 mtg
3. Ongoing work
- 4. Future focus**

- **Temporal trends**
- **Prediction modeling and model evaluation**
- **Cost-effectiveness analysis**
- **GWAS analysis**

G x Tx interaction / Tx-stratified analysis

Alternative analytic methodologies

Working Group => Your entry to CCSS

**PLEASE CONTACT ME IF YOU HAVE
ANY INTEREST OR IDEA ON
POTENTIAL EPI/BIOSTAT PROJECTS**

Yutaka.Yasui@stjude.org