

Received: 11.19.2010  
First Name: A Blythe  
Last Name: Ryerson  
Institution: Rollins School of Public Health, Emory University  
Address 1: 1518 Clifton Rd, NE CNR 3rd floor  
Address 2:  
City: Atlanta  
State/Province: GA  
Country: United States  
Zip: 30322  
Phone: 404-272-4248  
Alternate Phone:  
Email: blytheryerson@gmail.com

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Requirements to submit AOI:

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A comprehensive review of previously published data has been completed.: Yes  
The specific aims are clear and focused.: Yes  
The investigator has appropriate experience and expertise to develop the concept proposal; if not, has identified a mentor or senior co-investigator.: Yes  
The investigator agrees to develop an initial draft of the concept proposal within 6 weeks of approval of the AOI and to finalize the concept proposal within 6 months.: Yes

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Project Title: The effect exercise has on preventing late cardiac outcomes among childhood cancer survivors treated with anthracyclines  
Planned research population (eligibility criteria): Survivors previously treated with anthracyclines; no history of radiation with potential impact to the heart; answered the 2007 follow-up survey; \*note that sibling controls will also be used for this analysis  
Proposed specific aims: We are interested in determining what fraction of early cardiac outcomes among childhood cancer survivors treated with anthracyclines could be prevented by improving physical fitness among this population. Using data from the CCSS, we will attempt to decompose the total, direct, and indirect cardiac effects of anthracycline use among long term childhood cancer survivors. Anthracyclines are cardiotoxic, but part of their effect may occur because a cancer diagnosis also influences exercise deconditioning, a known risk factor for many cardiac outcomes. Within this framework, the effect of anthracyclines on the outcome, excluding all effects of exercise deconditioning, is the direct effect. The indirect effect is the effect of anthracycline exposure explained by subsequent exercise deconditioning. The indirect and direct effects together form the total effect of anthracycline exposure on the outcome. The aim is to estimate how important the mechanism of exercise deconditioning is relative to the total effect of anthracycline cardiotoxicity in order to determine the relative importance exercise interventions may be in survivors previously exposed to anthracyclines.  
Will the project require non-CCSS funding to complete?: No  
If yes, what would be the anticipated source(s) and timeline(s) for securing funding?:

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Does this project require contact of CCSS study subjects for . . .

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Additional self-reported information: No

Biological Samples: No

Medical record data: No

If yes to any of the above, please briefly describe.:

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What CCSS Working Group(s) would likely be involved? (Check all that apply)

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Second Malignancy:

Chronic Disease: Secondary

Psychology / Neuropsychology:

Genetics:

Cancer Control:

Epidemiology / Biostatistics: Primary

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To describe the anticipated scope of the study, please indicate the specific CCSS data to be included as outcome (primary or secondary) or correlative factors. (Check all that apply)

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Late mortality:

Second Malignancy:

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Health Behaviors

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Tobacco:

Alcohol:

Physical activity: Primary

Medical screening: Correlative Factors

Other:

If other, please specify:

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Psychosocial

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Insurance: Correlative Factors

Marriage: Correlative Factors

Education: Correlative Factors

Employment: Correlative Factors

Other:

If other, please specify:

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Medical conditions

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Hearing/Vision/Speech:

Hormonal systems:

Heart and vascular: Primary

Respiratory:

Digestive:

Surgical procedures:

Brain and nervous system:

Other:

If other, please specify:

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Medications

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Describe medications: Heart medications will be used in defining outcome

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Pregnancy and offspring: Correlative Factors

Family History: Correlative Factors

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Psychologic/Quality of Life

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BSI-18:

SF-36:

CCSS-NCQ:

PTS:

PTG:

Other:

If other, please specify:

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Chronic conditions (CTCAE v3): Correlative Factors

Health status: Correlative Factors

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Demographic

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Age: Correlative Factors

Race: Correlative Factors

Sex: Correlative Factors

Others:

If others, please specify:

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Cancer treatment

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Chemotherapy: Correlative Factors

Radiation therapy: Correlative Factors

Surgery: Correlative Factors

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Anticipated sources of statistical support

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CCSS Statistical Center:

Local institutional statistician: Yes

If local, please provide the name(s) and contact information of the statistician(s) to be involved.: Ann Mertens, PhD - Ann.Mertens@choa.org Harland Austin, DSc - haustin@emory.edu

Will this project utilize CCSS biologic samples?: No

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If yes, which of the following?

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Buccal cell DNA:

Peripheral blood:

Lymphoblastoid cell lines:

Second malignancy pathology samples:

Other requiring collection of samples:

If other, please explain:

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Other general comments: