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Requirements to submit AOI:

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

Project Title: Evaluation of the intrinsic molecular profiles of radiation-preceded breast cancer Planned research population (eligibility criteria): Childhood survivors of malignancy treated with radiation including to the chest, who have subsequently developed breast cancer Proposed specific aims: The objective of this study is to obtain gene expression profiles of breast cancers arising in women treated with radiotherapy as children. 1. The specific aim is to ascertain the intrinsic molecular breast cancer subtype among childhood cancer survivors treated with radiation. 2. Compare the frequency of basal-like intrinsic subtype to that observed in publically available cohorts of sporadic breast cancer. 3.An exploratory subaim will compare gene expression profiles between radiation-preceded CCSS breast cancers and breast cancers from women in databases for differences beyond intrinsic subtype, with particular reference to enrichment of mammary stem cell signatures and the metaprofiles identified from our radiation mammary chimera mouse model.

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?:

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

Additional self-reported information: No

Biological Samples: Yes Medical record data: Yes

If yes to any of the above, please briefly describe.: Slides of breast tumor specimens

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

Second Malignancy: Primary		
Chronic Disease:		
Psychology / Neuropsychology: Genetics: Secondary		
Cancer Control:		
Epidemiology / Biostatistics:		
To describe the anticipated scope of the study, please indicate the specific CCSS data to be included as <u>outcome</u> (primary or secondary) or <u>correlative factors</u> . (Check all that apply)		
Late mortality: Secondary		
Second Malignancy: Primary		
Health Behaviors		
Tobacco: Correlative Factors		
Alcohol: Correlative Factors		
Physical activity: Correlative Factors		
Medical screening: Correlative Factors		
Other:		
If other, please specify:		
Psychosocial		
Insurance:		
Marriage:		
Education: Correlative Factors		
Employment:		
Other:		
If other, please specify:		
Medical conditions		
Hearing/Vision/Speech:		
Hormonal systems:		
Heart and vascular:		
Respiratory:		
Digestive:		
Surgical procedures:		
Brain and nervous system:		
Other:		
If other, please specify:		
Medications		

Describe medications:		
Pregnancy and offspring: Family History: Correlative Factors		
Psychologic/Quality of Life		
BSI-18: SF-36: CCSS-NCQ: PTS:		
PTG: Other: If other, please specify:		
Chronic conditions (CTCAE v3): Correlative Factors Health status: Correlative Factors		
Demographic		
Age: Correlative Factors Race: Correlative Factors Sex: Correlative Factors Others: If others, please specify:		
Cancer treatment		
Chemotherapy: Correlative Factors Radiation therapy: Correlative Factors Surgery: Correlative Factors		
Anticipated sources of statistical support		
CCSS Statistical Center: Yes Local institutional statistician: Yes If local, please provide the name(s) and contact information of the statistician(s) to be involved.: Statistician working with Dr. Charles Perou (chuck perou@med.unc.edu) at the University of North Carolina School of Medicine		

Will this project utilize CCSS biologic samples?: Yes

Buccal cell DNA:	
Peripheral blood:	
Lymphoblastoid cell lines:	
Second malignancy pathology samples: Yes	
Other requiring collection of samples:	
If other, please explain:	

Other general comments: