

Received 4.14.11  
First Name: David  
Last Name: Hodgson  
Institution: Princess Margaret Hospital  
Address 1: 610 University Avenue  
Address 2:  
City: Toronto  
State/Province: Ontario  
Country: Canada  
Zip: M5G 2M9  
Phone: 416-946-2919  
Alternate Phone:  
Email: david.hodgson@rmp.uhn.on.ca

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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: Identification of susceptibility loci in radiation-induced breast cancer by exome and CNV analysis  
Planned research population (eligibility criteria): Patients with breast cancer following thoracic radiation (eg. Hodgkin lymphoma, Wilms tumor whole lung radiation, sarcoma, etc).  
Proposed specific aims: 1) Exome sequencing on peripheral blood lymphocytes or buccal swab epithelial cells to identify candidate gene loci/mutations. 2) Copy number variation analysis using Affymetrix 6.0GW SNP/CNV arrays to identify candidate susceptibility loci from peripheral blood lymphocytes/buccal swabs. 3) Examine LOH in radiation-induced tumors based on locus detection from Aims 1 and 2. 4) Validate identification of candidate loci through analysis of germline sequences against patients who received thoracic radiation but did not develop breast cancer (if available). 4) Similar analysis to Aim 3 using patients with sporadic breast cancer as comparison population.  
Will the project require non-CCSS funding to complete?: Yes  
If yes, what would be the anticipated source(s) and timeline(s) for securing funding?:  
This is one of three projects that are part of an application to the Canadian Institutes for Health Research (deadline June 1, 2011).

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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: Yes  
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: Secondary  
Chronic Disease:  
Psychology / Neuropsychology:  
Genetics: Primary  
Cancer Control:  
Epidemiology / Biostatistics:

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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: Primary

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Health Behaviors

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Tobacco:  
Alcohol:  
Physical activity:  
Medical screening: Correlative Factors  
Other:  
If other, please specify:

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Psychosocial

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Insurance:  
Marriage:  
Education:  
Employment:  
Other:  
If other, please specify:

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Medical conditions

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Hearing/Vision/Speech:  
Hormonal systems:  
Heart and vascular:  
Respiratory:  
Digestive:

Surgical procedures: Correlative Factors

Brain and nervous system:

Other:

If other, please specify:

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Medications

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Describe medications:

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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):

Health status:

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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.: Dr. Mohammed Akbari TEL: 416-351-3800 x 6353 Women's College Hospital Research Institute, Toronto, ON. Canada.

Will this project utilize CCSS biologic samples?: Yes

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If yes, which of the following?

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Buccal cell DNA: Yes

Peripheral blood: Yes

Lymphoblastoid cell lines:

Second malignancy pathology samples: Yes

Other requiring collection of samples:

If other, please explain:

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Other general comments: