### **Contact Information**

Name Tom Lumbers

**Institution** University College London

Address 222 Euston Road

London, NW12DA United Kingdom

Phone Number +447980692582

**Alternate Phone Number** 

Email Address t.lumbers@ucl.ac.uk

# **Project Requirements and Description**

Requirements to submit AOI (all answers must be "yes" to proceed)

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 coinvestigator.	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 Genomic Prediction of Cancer Treatment-Related

Cardiotoxicity in Long-Term Survivors of Childhood Cancer

### Planned research population (eligibility criteria)

Inclusion criteria:

- 1. Survivors with at least 5 years survival
- 2. CCSS Participants with Genotype Data

#### Exclusion criteria:

1. History of heart failure, atrial fibrillation, cardiomyopathy, or congenital heart disease prior to cancer treatment

#### Proposed specific aims

1. Evaluate the predictive performance of a clinical risk score for cardio-toxicity (CTCAE cardiac disorders) based on baseline risk factors (sex, age at cancer diagnosis, and anthracycline and chest radiotherapy doses).

- 2. Integrate polygenic scores for dilated cardiomyopathy and heart failure, developed using unpublished GWAS base data (DCM 15,000 cases; HF 150,000 cases), and evaluate the incremental AUC for events over the clinical risk score.
- 3. Validation of the integrated score, combining clinical risk predictors with polygenic scores, in a second patient cohort from the Genomics England Cancer study.

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	No
Medical record data	No

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

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

Second Malignancy	
Chronic Disease	Secondary
Psychology/Neuropsychology	
Genetics	Primary
Cancer Control	
Epidemiology/Biostatistics	Secondary

### **Outcomes or Correlative Factors**

Late Mortality	
Second Malignancy	

### **Health Behaviors**

Tobacco	Correlative Factors
Alcohol	Correlative Factors
Physical Activity	Correlative Factors
Medical Screening	
Other	

## If other, please specify

## **Psychosocial**

Insurance	
Marriage	
Education	
Employment	
Other	

## If other, please specify

### **Medical Conditions**

Hearing/Vision/Speech	
Hormonal Systems	
Heart and Vascular	Primary
Respiratory	
Digestive	
Surgical Procedures	
Brain and Nervous System	
Other	Primary

# If other, please specify

Cancer Treatment-Related Cardiotoxicity

# **Medications**

### **Describe medications**

# Psychologic/Quality of Life

BSI-18	
SF-36	
CCSS-NCQ	
PTS	
PTG	
Other	

# If other, please specify

### Other

Pregnancy and Offspring	
Family History	
Chronic Conditions (CTCAE v3)	Correlative Factors
Health Status	

# Demographic

Age	Correlative Factors
Race	Correlative Factors
Sex	Correlative Factors
Other	

# If other, please specify

# **Cancer Treatment**

Chemotherapy	Correlative Factors
Radiation Therapy	Correlative Factors
Surgery	

### **Anticipated Sources of Statistical Support**

CCSS Statistical Center	No
Local Institutional Statistician	Yes

If local, please provide the name(s) and contact information of the statistician(s) to be involved.

Will this project utilize CCSS biologic samples?

No

If yes, which of the following?

If other, please explain

## **Other General Comments**

#### Agree

I agree to share this information with St. Jude

This Service is governed by and operated in accordance with US law. If you are located outside of the US, you use this Service voluntarily and at your own risk. If you choose to submit personal data like your name and email address, please note that your Information will be transferred to and processed in the United States. By checking this box while using this Service, you acknowledge that the data protection and other laws of other countries, such as the United States, may provide a less comprehensive or protective standard of protection than those in your country, and consent to your Information being collected, processed and transferred as set forth in the Privacy Policy and US law.