Contact Information

Name Rawan Hammoud

Institution St. Jude Children's Research Hospital

Address 262 Danny Thomas Place, MS 735

Memphis, TN, 38105

United States

Phone Number 9015956091

Alternate Phone Number

Email Address rawan.hammoud@stjude.org

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

Comparison of excess risk of late mortality and subsequent malignant neoplasms after Hodgkin lymphoma: a Childhood Cancer Survivor Study and Dutch Hodgkin lymphoma Survivor Study collaboration

Planned research population (eligibility criteria)

- Survivors eligible for CCSS diagnosed with Hodgkin lymphoma at ages 15-<21 years between 1970-1999.
- Participants from the Dutch Hodgkin lymphoma Survivor Study diagnosed at ages 15-<21 years between 1970-1999.

Proposed specific aims

I. Aim 1.1: To compare the absolute excess risk (AER) and standardized mortality ratios (SMRs) for all-cause and cause-specific late-mortality among adult survivors of Hodgkin lymphoma diagnosed ages 15-<21 years in the CCSS and Dutch cohorts.

Hypothesis 1.1: Survivors from North America will experience a higher risk of death from late effects when compared to the risk of the Dutch cohort survivors.

Aim 1.2: To identify treatment and lifestyle-related factors associated with the observed differences in late mortality between the CCSS and Dutch cohorts.

Hypothesis 1.2: We hypothesize that survivors from both cohorts may have received similar treatment exposures. However, survivors treated in North America will have a greater burden of modifiable lifestyle risk factors compared with the Dutch cohort, contributing to the increased risk of death from late effects.

II. Aim 2.1: To compare the AER and standardized incidence ratios (SIRs) of subsequent malignant neoplasms (SMNs; overall and type-specific) among survivors of Hodgkin lymphoma diagnosed between the age of 15-<21 years in the CCSS and the Dutch cohorts.

Hypothesis 2.1: Survivors from North America will have an increased risk of developing SMNs when compared to the risk of the Dutch cohort survivors.

Aim 2.2: To identify treatment, lifestyle-related factors, and for breast cancer, differential use of hormone replacement therapy, associated with the observed differences in risk of developing SMNs between the CCSS and Dutch cohorts.

Hypothesis 2.2: We hypothesize that survivors from both cohorts may have received similar treatment exposures. However, survivors treated in North America will have a greater burden of modifiable lifestyle risk factors compared with the Dutch cohort, contributing to the increased risk of developing SMNs.

Exploratory aim 1.1: Acknowledging the lack of a population-based registry for myocardial infarction (MI) in the US, we aim to compare the cumulative incidence and rate of MI among survivors of Hodgkin lymphoma diagnosed between the age of 15-<21 years in the CCSS and the Dutch cohorts.

Hypothesis: Survivors from North America will have an increased risk of MI when compared to the risk of the Dutch cohort survivors. -

Exploratory aim 1.1: To identify treatment and lifestyle-related factors associated with the observed differences in risk of developing an MI between the CCSS and Dutch cohorts.

Hypothesis: We hypothesize that although survivors from both cohorts may have received similar treatment exposures. However, survivors treated in North America will have a greater burden of modifiable lifestyle risk factors compared with the Dutch cohort, contributing to the increased risk of developing an MI.

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 | Secondary |
|----------------------------|-----------|
| Chronic Disease | Secondary |
| Psychology/Neuropsychology | |
| Genetics | |
| Cancer Control | |
| Epidemiology/Biostatistics | Primary |

Outcomes or Correlative Factors

| Late Mortality | Primary |
|-------------------|-----------|
| Second Malignancy | Secondary |

Health Behaviors

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

If other, please specify

Psychosocial

| Insurance | Correlative Factors |
|-----------|------------------------|
|-----------|------------------------|

| Marriage | |
|------------|------------------------|
| Education | Correlative Factors |
| Employment | Correlative Factors |
| Other | |

If other, please specify

Medical Conditions

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

If other, please specify

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 | Correlative Factors |

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 | Correlative Factors |

Anticipated Sources of Statistical Support

| CCSS Statistical Center | Yes |
|----------------------------------|-----|
| Local Institutional Statistician | No |

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?

Other General Comments

I plan to submit this proposal for the career development trainee award.

Agree

I agree to share this information with St. Jude

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