From Worn Out to Energized: Tackling Fatigue After Transplant and CAR T-cell Therapy

Celebrating a Second Chance at Life Survivorship Symposium

May 3-9, 2025



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# Background



# What is fatigue?





# According to the dictionary:

• "A weariness or exhaustion from labor, exertion, or stress"







# What about <u>cancer-related</u> fatigue (CRF)?

"Cancer-related fatigue is a distressing, persistent, subjective sense of physical, emotional, and/or cognitive tiredness or exhaustion related to cancer treatment that is not proportional to recent activity and interferes with usual functioning"

-National Comprehensive Cancer Network





## Let's break that down...

Cancer-related fatigue is a

- 1. <u>distressing, persistent</u>, subjective sense of
- <u>physical, emotional, and/or cognitive</u> <u>tiredness or exhaustion</u> related to cancer treatment that is
- 3. <u>not proportional to recent activity</u> and
- 4. interferes with usual functioning



# **Different beasts**



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# How is CRF experienced?

Meta-ethnography of 16 qualitative studies

Six overarching themes



Bootsma et al., *Psychooncology*, 2020



# How common is CRF?

- Short answer: <u>very</u> common
- About 52% of patients across 84 studies

- Most common side effect of cancer treatment
- Can also be a symptom of cancer itself
- Often rated as most distressing

Ma et al., Int J Nurs Stud, 2020



# How common is CRF?









Ma et al., Int J Nurs Stud, 2020



# Why does CRF happen?

## Bottom line: we don't really know for sure

#### Some things that may contribute:

Systemic inflammation

Anemia Depression Sleep disturbance Physical inactivity Higher BMI Pain Type of Transplant (autoSCT vs. alloSCT) Infection Difficulty breathing Problems with organs Coping strategies and expectations Cancer treatments

Sidana et al., 2019



# When does CRF happen?





# How long does CRF last?





# How long does CRF last?

% of patients who ranked each symptom as lasting the longest





#### 2025 SURVIVORSHIP SYMPOSIUM

Curt et al., Oncologist, 2000

# What impact does CRF have?

% of patients who said the symptom impacts everyday life the most 10% Fatigue Nausea Depression 22% 60% Pain



#### 2025 SURVIVORSHIP SYMPOSIUM

Curt et al., Oncologist, 2000

# What impact does CRF have?

#### % of CAR-T patients who said the symptom impacts everyday life the most



■ Fatigue ■ Loss of Appetite ■ Headache ■ Chills ■ Confusion



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Whisenant et al., Seminars in Oncology Nursing, 2021

# So, what can be done about CRF?



# Patient/Family Education and Counseling



- Education about fatigue, especially if undergoing fatigue-inducing treatments (e.g., RT, CT)
- Fatigue not necessarily sign that treatment not working or that disease progressing
- Fatigue can be a consequence of treatment
- May benefit from nutrition consult with a registered dietician



# **General Strategies**

- Energy conservation
- Maintain a diary
- Distraction
- Daytime naps < 30 minutes, in-bed, before 3pm
  - "Nappuccino" Drink a small coffee prior to napping

# **Pharmacologic Interventions**



#### Some evidence psychostimulants and corticosteroids can help



# Non-pharmacologic interventions



#### 1. Physical Activity



#### 2. Psychosocial Interventions



## 3. Bright light therapy



# **Physical Activity**

- 72 studies (5367 patients) in active treatment or follow-up.
- Moderate effect of exercise in reducing CRF compared to control group.
- Exercise type does not matter.
- Tailored may benefit from physical therapy or other exercise program.





# **Bright Light Therapy**

- Commonly used to treat seasonal affective disorder
- Systematic exposure to bright light may normalize circadian rhythms
- Pilot studies show that it may prevent/treat fatigue in cancer patients and survivors.



# What does bright light therapy look like?

- 30 minutes each morning upon awakening
- 4 to 8 weeks
- Bright light (broad spectrum)
- Small lamp, 45-degree angle, arm length away
- Light goggles are another option





# The Bottom Line

- CRF is real and is different from regular fatigue
- Common, severe, impactful, and long-lasting
- Causes and mechanisms are unknown
- There are a variety of tools available to treat or help with the management of CRF



# CBT-I: The Gold Standard





# Penn Sleep & Cancer Clinic/Pilot Study



# Penn Sleep & Cancer Clinic

- Collaboration with Perelman Center and Dept of Psychiatry
  - Began in February of 2024.
- Provided CBT-I and CRF management to patients diagnosed with cancer and cancer survivors.
  - Treatment outcomes were assessed with the ISI, BFI, & SE%
- Provided information and education to oncology teams on identifying insomnia and fatigue in the context of cancer.
- Providers were encouraged to refer patients with insomnia and/or CRF
- Referrals were sent via the EMR.
- Majority of sessions were conducted via telehealth.



#### **Reach and Retention**

- Reach, defined as the number of patients referred/the number of patients scheduled for an intake.
- Retention, defined as the number of patients who completed treatment/the total number of scheduled patients.

March 2024-February 2025	Total Number of Patients Referred	Total Number of Patients Who Completed Intake	Percent	
Reach	78	44	56%	
	Number of Patients Who Completed Treatment	Total Number of Scheduled Patients		
Retention	26	44	59%	



### **Pre/Post Data**

- Of the 44 patients who completed treatment, 13 subjects are included in the present descriptive analysis (means, SD):
  - Patients who were not included did not have pre/post data or received fatigue management (1-2 sessions), not CBT-I.
- The subsequent slides include data on:
  - Demographic Information
  - Pre/Post ISI, BFI, and SE% data



### **Demographics**



Mage = 61.7 years, SDage = 14.7



# **Pre/Post Data**

Pre/Post CBT-I	Pre (x̄)	SD	Post (x)	SD	Percent Change	Effect Size
in Cancer						
(N=13)						
ISI	14.3	2.4	6.2	2.9	56%	3.0
BFI	4.4	2.1	2.2	2.1	50%	1.0
SE%	75.7	10.7	91.9	4.8	21%	-1.9

Mean number of sessions of CBT-I was 6.8.









#### Brief Fatigue Inventory





#### **Sleep Efficency**



Pre/Post



#### Insomnia & Fatigue in Breast Cancer



- This preliminary analyses suggest that clinical gains via CBT-I are robust and sustained for both insomnia & fatigue outcomes.
- Will be replicated in other cancers, such as leukemias, lymphomas, and multiple myeloma.



# **Future Directions**

- Expansion of Services
  - Increase accessibility of services, including to underserved populations
- Research & Innovation
  - More research is needed on novel treatment approaches to CRF, especially in transplant and CAR-T
- Training & Education
  - Enhance provider training in BSM to improve integration in oncology care
- Long-Term Impact
  - Assess the sustained benefits of treatment and explore the role of sleep interventions in cancer survivorship and recurrence prevention



# **Thank You**



# Penn Medicine





Gary Freedman, MD



Michael Perlis, PhD

Deseray McCoy, Admin

Mark Seewald, CRC

Perelman Oncology Services

All the amazing patients!





Thank you for listening! Contact: <u>amuench@pennmedicine.upenn.edu</u>



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# **Questions?**



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