



Best Practice and Interventions in Survivorship

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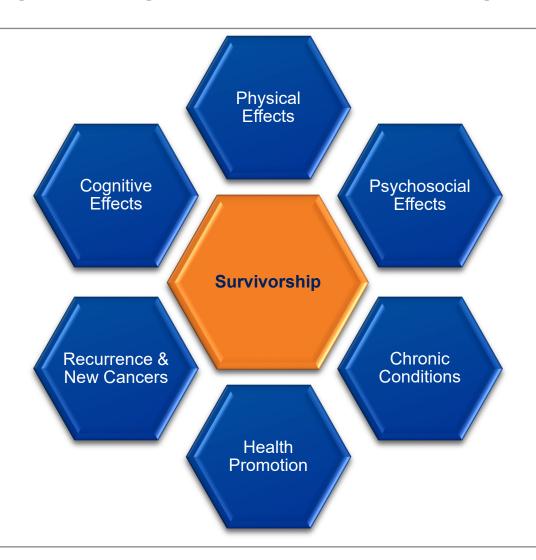
Outline

- Complexity of survivorship care in hematopoietic stem cell transplantation (HSCT)
- Interventions to promote physical functioning post HSCT
- Interventions to promote mental functioning post HSCT
- Future directions





Complexity of Survivorship Care







Complexity of Survivorship Care

Prevention

of recurrent and new cancers and late effects

Surveillance

for metastasis,
recurrence, or
secondary
cancers;
assessment of
medical and
psychosocial late
effects

Intervention

for impacts of cancer and its treatment

Coordination

between specialists and primary care providers





Complexity of Survivorship Care

- Multidisciplinary long-term follow-up and survivorship clinics to address the unique needs of HSCT recipients:
 - –Dermatology
 - -Ophthalmology
 - -Oral health
 - -Cardiology
 - –Pulmonary
 - -Endocrine
 - -Sexual health and fertility

- –Physical therapy and rehabilitation
- Occupational therapy
- –Psychology
- —Infectious disease
- –Neurology
- –Gastroenterology





Barriers to Survivorship Care

- Approximately 50% of transplant programs have long-term follow up clinics
- Clinics vary widely in services provided
- Logistics and costs
- Lack of expertise
- Sense of ownership over addressing the survivorship care needs of HSCT recipients

Hashmi SK, et al. BBMT 2018 Dignan FL, et al BMT 2021





Interventions to Promote Physical Functioning

- Multiple randomized clinical trials tested exercise intervention during HSCT
- Most studies: interventions started before or during HSCT, duration 4 weeks – 6 months
- Physical exercise: both aerobic and muscle strengthening interventions
- Supervised or partially supervised interventions, variable intensity
- Outcomes: cardiorespiratory fitness, quality of life, fatigue

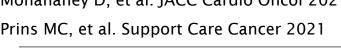




Interventions to Promote Physical **Functioning**

- Moderate positive effects on cardiorespiratory fitness
- Moderate positive effects on lower extremity strength
- Small to moderate positive effects on global quality of life (QOL)
- Small to moderate positive effects on fatigue
- Numerous methodological limitations
- Variability across the HSCT recovery

Mohananey D, et al. JACC Cardio Oncol 2021 Prins MC, et al. Support Care Cancer 2021







Interventions to Promote Physical Functioning

- We need physical exercise interventions in patients with chronic graft-versus-host disease (GVHD)
 - Major cause of disability
 - Reduced functional capacity
 - -Muscular and functional limitations
 - -Inflammatory disease
 - -Sarcopenic obesity
 - Preclinical evidence to support its promise



Fiuza-Luces C, et al. BMT 2016





Type of Intervention	Intervention intensity	Timing Intervention	Outcomes	Limitations
Mind-body & stress management	Relaxation guided imagery, breathing exercises, progressive muscle relaxation.	During HSCT Short follow-up	Small effects on anxiety, depression, and fatigue. QOL improvement seen in more comprehensive coping programs. Small effect sizes compared to cognitive behavioral therapy	Poor adherence in self- directed interventions Includes autologous and allogeneic HSCT collectively
Cognitive and behavioral therapy (CBT)	CBT methods Some relaxation in addition to CBT Face-to-face interventions.	During or immediately following HSCT	Some benefits with lower distress, improved emotional functioning. Enduring benefits at 3 and 12 months post-HSCT. More intensive interventions yielded larger and more significant effects.	Mostly White participants Includes autologous and allogeneic HSCT collectively Did not target high-risk patients

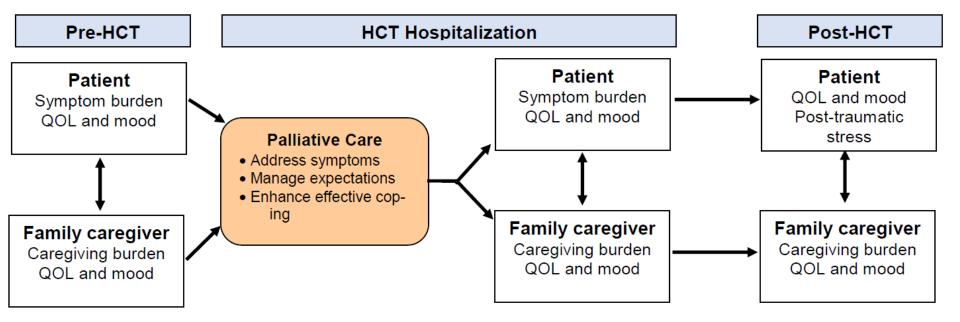
Bevans M, et al. BBMT 2017; Baliousis M, et al. Psychooncology 2016





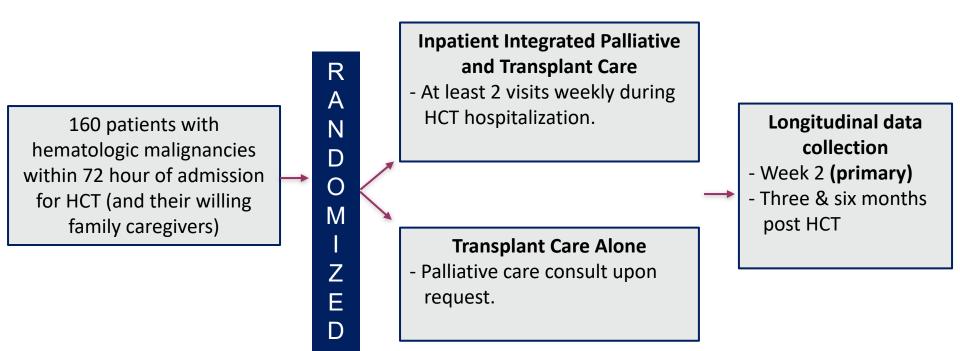
Conceptual Model:

Figure 1









El-Jawahri, et al. JAMA 2016



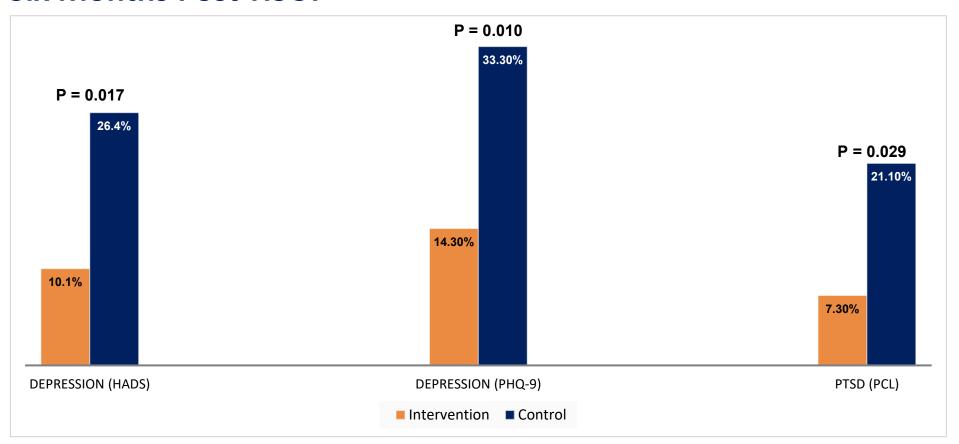


Week-2 Outcomes	Adjusted Mean Difference	95% CI	P- Value
QOL: FACT – BMT	7.73	1.27 to 14.19	0.019
Fatigue: FACT – Fatigue	3.88	0.21 to 7.54	0.038
Symptom burden: ESAS	-6.26	-11.46 to -1.05	0.019
Depression symptoms: HADS-D	-1.74	-3.01 to -0.47	0.008
Anxiety symptoms: HADS-A	-2.26	-3.22 to -1.29	<0.001
Depression: PHQ-9	-1.28	-2.82 to 0.27	0.104
El-Jawahri JAMA 316(20) 2016			





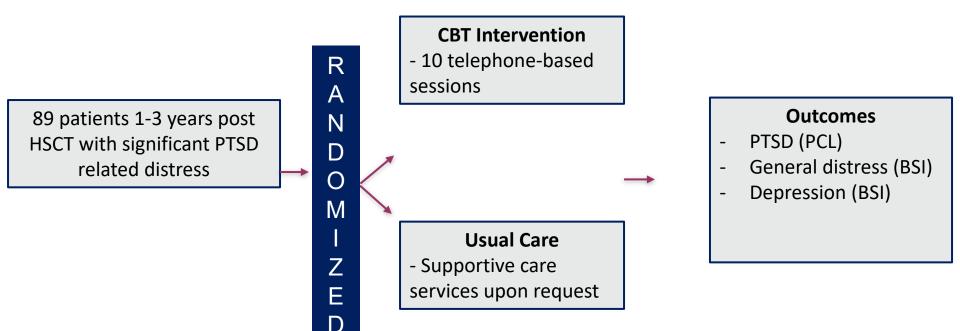
Six Months Post-HSCT



El-Jawahri, et al JCO 2017







Du Hamel, JCO 2010

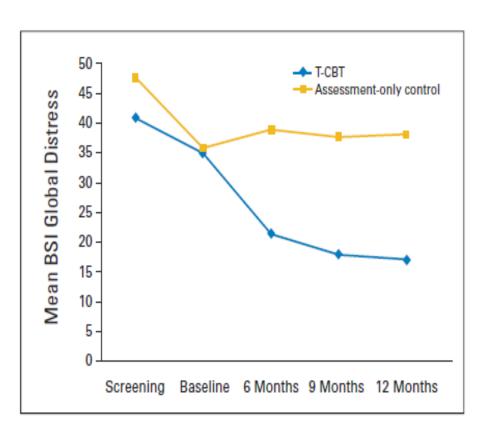


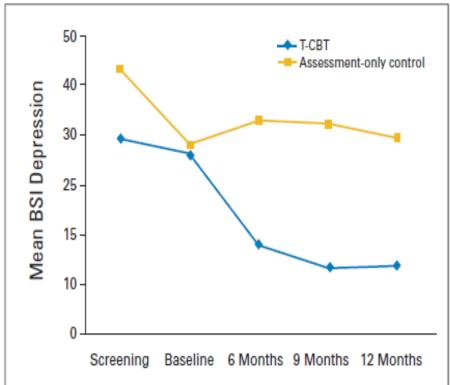


	T-CBT Group		Assessment-Only Group	
Outcome	Mean	95% CI	Mean	95% CI
Total PCL-C				
Screening	34.01	30.79 to 37.24	38.23	35.27 to 41.20
Baseline	32.05	28.60 to 35.50	33.97	30.18 to 37.76
6 months postbaseline	25.38	21.69 to 29.07	32.05	27.18 to 36.93
9 months postbaseline	24.63	21.08 to 28.18	31.99	27.42 to 36.56
12 months postbaseline	24.00	19.20 to 28.01	30.89	26.33 to 35.45













Why is Mental Functioning Important?

Allogenic HSCT	N	HR	95% CI	P-value
Overall survival Depression	1095	1.13	1.04-1.23	p=0.003
Acute GVHD, II-IV Depression	1115	1.25	1.14-1.37	p<0.0001
Chronic GVHD Depression	1109	1.06	0.96-1.16	p=0.26
Length of stay Depression	1093	0.97	0.95-0.99	p=0.002
El-Jawahri, Cancer 2016				





Future Directions

- Physical Medicine and Rehabilitation (PM&R) interventions for HSCT recipients
- Strategies to tailor and enhance physical exercise and recovery interventions in HSCT recipients
- Need to focus on patients with chronic GVHD: physical and mental functioning interventions
- Strategies to increase access and scalability of these interventions – Digital Therapeutics





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