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Associations between gut microbiota and health behaviours in a cross-sectional sample of young

adult cancer survivors: Secondary analysis from the Chemo-Gut study

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Background: Health behaviours are a modifiable factor that can potentially impact the gut microbiota, physical and psychosocial health. Chemotherapy adversely affects the gut microbiota during treatment, but little is known about the long-term effects or how health behaviours are related.

Methods: This secondary analysis from a cross-sectional pilot study investigated relationships between chemotherapy treatment, gut microbiota, and patient reported outcomes for health behaviours (i.e. diet and exercise) and psychosocial symptoms in cancer survivors aged 18-39 years old, compared to healthy controls. Gut microbial diversity and composition was assessed from stool samples using 16S rRNA gene sequencing. Descriptive statistics, one-way ANOVA, and Spearman's correlation analyses are reported.

Results: Survivors (n=17) and healthy controls (n=18) participated. Mean age at diagnosis was 31 (\pm 5) years. Mean time off treatment was 17 (\pm 16) months. 23.5% of survivors and 38.9% of controls used probiotics in the last 2 years. Survivors tended to rate their diet as moderately healthy (58.8%) while controls rated their diet as moderately healthy (50%) or healthy (44.4%). Survivors (76.5%) reported engaging in less exercise (≤5 hours/week), compared to controls $(44.4\% \ge 6 \text{ hours/week})$. Both groups reported moderate exercise intensity (76.5% of survivors vs. 61.1% of controls). Survivors ≤ 6 months post-treatment had lower gut microbiota alpha diversity than both survivors >6 months post-treatment (p=0.04) and controls (p=0.19). In survivors, poorer self-rated diet quality correlated with greater anxiety (rho= -.52, p=.033), while greater depressive functional interference correlated with higher exercise intensity (rho= .57, p=.018). In survivors, higher abundance of Lachnospiraceae (rho=.51, p=.05) correlated with better diet health, but higher *Bacteroides* (rho=-.62, p=.023) abundance correlated with poorer diet health. Higher abundance of Lachnospiraceae (rho=.51, p=.035), Faecalibacterium (rho= .65, p= .013), Anaerostipes (rho= 53, p= .042), Subdoligranulum (rho= .59, p= .027), Alistipes (rho= .75, p= .013), and Bacteroides (rho= .66, p= .027) correlated with greater exercise frequency. Lower abundance of *Lachnospiraceae* correlated with higher exercise intensity (rho= -.63, p = .012).

Conclusion: This small exploratory study provides evidence of potential longer-term gut microbial dysbiosis in cancer survivors. Diet and exercise behaviours may be related to certain types of bacteria, but the direction of causality is unknown. The first-year post-treatment may be a key time to intervene with behavioural and dietary-based interventions. Larger trials are needed.

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