

00418 **Leptin as a Potential Fatigue Biomarker for the Onset of Cancer-related Fatigue (CRF): A Prospective Cohort Study**

Toh Yi Long¹, Tan Chia Jie¹, Shwe Maung¹, Yeo Hui Ling¹, Ho Han Kiat¹, Gan Yan Xiang², Rebecca Dent², Ng Chee Hui², Yap Yoon Sim², Kiley Loh², Lee Guek Eng², Beh Sok Yuen², Tan Jing Ying Tira², Mabel Wong², Alexandre Chan²

¹National University of Singapore, ²National Cancer Centre Singapore

Aims: While leptin is known for its metabolic effects ranging from appetite suppression to increasing energy expenditure, it has also been characterized as an adipokine. Being pro-inflammatory, leptin is postulated to have a role in predicting onset of Cancer-Related Fatigue (CRF). This study aims to evaluate the correlation between leptin and CRF in early-stage breast cancer patients receiving chemotherapy.

Methodology: In a prospective cohort study, patients completed assessments at baseline (T1), during (T2) and after chemotherapy (T3). At each time point, plasma leptin level was measured using Luminex bead-immunoassay and the validated Multi-Dimensional Fatigue Symptom Inventory-Short form (MFSI-SF) was utilized to measure CRF. Data was longitudinally analysed using a Generalized Estimating Equation model, incorporating clinically relevant parameters and pro-inflammatory cytokines that were statistically associated with total MFSI-SF score.

Result: 136 patients were recruited (mean age \pm SD = 51.5 \pm 8.8 years; mean BMI of 23.9 kg/m²; 69.1% receiving anthracycline-based chemotherapy). More patients experienced CRF at T3 (22.1%) than at T2 (13.2%) compared to baseline. Median (\pm inter-quartile range) leptin level was 4.33 \pm 2.82 ng/mL. Leptin inversely correlated with total MFSI-SF score (β = -0.56, p < 0.01), general (β = -0.16, p < 0.01), emotional (β = -0.11, p < 0.01) and mental sub-domains (β = -0.06, p = 0.025). In the final model, plasma leptin levels showed a statistically significant inverse correlation with total MFSI-SF score over time (β = -0.22, p < 0.01) after adjusting for anxiety, depression, insomnia, age, menopausal status and chemotherapy.

Conclusion: This is the first study to demonstrate leptin as a potential fatigue biomarker to predict the onset of CRF over time. The inverse correlation observed suggests that the underlying mechanism linking leptin to fatigue may be preceded by other non-inflammatory factors. Future studies are required to validate the findings and to evaluate the sensitivity and specificity to confer clinical utility.