

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

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**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 (T<sub>1</sub>), during (T<sub>2</sub>) and after chemotherapy (T<sub>3</sub>). 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/m<sup>2</sup>; 69.1% receiving anthracycline-based chemotherapy). More patients experienced CRF at T<sub>3</sub> (22.1%) than at T<sub>2</sub> (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 ( $\beta$  = -0.56,  $p$  < 0.01), general ( $\beta$  = -0.16,  $p$  < 0.01), emotional ( $\beta$  = -0.11,  $p$  < 0.01) and mental sub-domains ( $\beta$  = -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 ( $\beta$  = -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.