

HIGH SERUM LEPTIN LEVELS ASSOCIATE WITH GOOD NUTRITIONAL STATUS IN NON-OBESE CHRONIC HAEMODIALYSIS PATIENTS

Ekrem Kara, Elbis Ahabap, Taner Basturk, Yener Koc, Tamer Sakaci, Tuncay Sahutoglu, Cuneyt Akgol, Mustafa Sevinc, Zuhale Atan Ucar, Arzu Ozdemir Kayalar, Abdulkadir Unsal

Sisli Etfal Research and Educational Hospital, Department of Nephrology, Istanbul, Turkey

Objectives:

Leptin is a adipocyte-derived peptide hormone and a member of proinflammatory cytokines which functions to suppress appetite and increase energy expenditure in the healthy population. Serum leptin levels are significantly elevated in patients with end stage renal disease (ESRD) primarily due to decreased clearance by kidneys. The role of hyperleptinemia and the influence of serum leptin levels on nutrition-inflammation status in chronic hemodialysis (HD) patients remained to be elucidated. Thus, we aimed to determine the association between serum leptin levels and malnutrition-inflammation status in non-obese chronic hemodialysis patients.

Methods:

Sixty-five patients who were on dialysis between January 1987-January 2013 were included in this cross-sectional study. Patients who had been undergoing HD treatment for at least 3 months, age ≥ 18 years, BMI <25 kg/m² were selected. To minimize the confounding effects of residual renal function on leptin and nutritional markers, we studied only anuric patients. The exclusion criterias were hospitalizations, major surgery, obvious infections or inflammatory disease within the preceding 3 months, end stage liver disease, metastatic malignancies, malabsorption syndromes. To determine nutrition-inflammation status; body mass index (BMI), dry weight, triceps scinfold thickness (mm), malnutrition inflammation score (MIS), serum albumin, prealbumin, hs-CRP and TNF- α levels were obtained from all patients. Patients were classified into 3 groups according to the serum leptin levels which adjusted by BMI and gender: Group 1 (low leptin), group 2 (normal leptin), group 3 (high leptin).

Table 1. The demographic features and laboratory findings of the patients according to the three groups of serum leptin levels.

Demographics	Low Leptin (n=9)	Normal Leptin (n=31)	High Leptin (n=25)	P
Age (y)	46.2 \pm 16.4	52.7 \pm 18.8	52.1 \pm 17.4	N.S.
Sex (male:female)	9/0	18/13	7/18	0.001
Duration on HD (months)	76.7 \pm 66.0	94.8 \pm 74.5	57.7 \pm 55.7	N.S.
Dry weight (kg)	49.6 \pm 10.8	55.8 \pm 8.5	54.1 \pm 8.5	N.S.
Body mass index (kg/m ²)	18.1 \pm 1.9	21.3 \pm 2.0	21.3 \pm 2.8	0.003
Delivered dose of dialysis				
KtV	1.50 \pm 0.20	1.56 \pm 0.35	1.76 \pm 0.38	N.S.
Anthropometry				
Triceps skinfold thickness (mm)	9.1 \pm 3.2	11.1 \pm 4.3	15.4 \pm 6.8	0.003
Laboratory				
Predialysis urea (mg/dl)	117.0 \pm 39.6	135.9 \pm 25.1	135.9 \pm 28.0	N.S.
Predialysis creatinine (mg/dl)	7.6 \pm 2.7	8.8 \pm 2.0	8.4 \pm 1.9	N.S.
Hemoglobin (g/dl)	10.0 \pm 2.7	10.7 \pm 1.6	10.3 \pm 1.7	N.S.
Uric acid (mmol/L)	5.0 \pm 0.6	5.7 \pm 1.0	6.0 \pm 1.0	N.S.
Na (mmol/L)	138.1 \pm 2.7	138.2 \pm 3.0	138.0 \pm 2.2	N.S.
K (meq/L)	4.9 \pm 0.8	5.1 \pm 0.6	5.0 \pm 0.7	N.S.
Ca (mg/dl)	8.6 \pm 0.7	8.7 \pm 0.8	8.6 \pm 1.1	N.S.
P (mg/dl)	5.1 \pm 2.0	5.4 \pm 1.3	5.3 \pm 1.3	N.S.
CacP	46.2 \pm 19.3	48.3 \pm 14.6	47.4 \pm 15.1	N.S.
Intact PTH (pg/ml)	598.2 \pm 864.1	770.8 \pm 824.8	424.5 \pm 486.2	N.S.
Total cholesterol (mmol/L)	153.8 \pm 47.9	157.5 \pm 43.6	183.4 \pm 32.7	0.041
LDL (mmol/L)	83.0 \pm 28.4	92.7 \pm 37.0	106.8 \pm 35.6	N.S.
Triglyceride (mmol/L)	128.8 \pm 129.2	148.1 \pm 64.0	154.8 \pm 63.2	N.S.
Bicarbonate (mEq/L)	23.1 \pm 2.0	22.2 \pm 2.2	22.4 \pm 2.0	N.S.
Nutrition				
Albumin (g/L)	3.3 \pm 0.6	3.8 \pm 0.2	3.8 \pm 0.3	0.001
Prealbumin (mg/dl)	25.1 \pm 7.0	26.1 \pm 7.0	30.9 \pm 8.0	0.033
TIBC (g/L)	175.2 \pm 26.7	207.9 \pm 46.9	218.7 \pm 44.4	0.045
Malnutrition Inflammation Score (MIS)	10.0 \pm 4.3	6.2 \pm 2.6	5.8 \pm 2.8	0.002
Inflammation				
hs-CRP (mg/L)	25.5 \pm 52.4	20.1 \pm 26.1	9.6 \pm 18.0	N.S.
TNF- α (pg/mL)	22.5 \pm 6.7	25.3 \pm 9.9	20.5 \pm 7.3	N.S.
Ferritin (ng/ml)	817.7 \pm 613.9	632.1 \pm 461.2	794.4 \pm 457.4	N.S.

Table 2. Factors correlates with serum leptin levels.

Leptin (ng/L)	P	r
Age (y)	0.028	-0.273
Gender	0.001	-0.665
Body mass index (kg/m ²)	0.001	-0.489
Triceps skinfold thickness (mm)	0.001	-0.606
KtV	0.028	-0.273
Albumin (g/L)	0.055	-0.263
Prealbumin (mg/dl)	0.031	-0.267
Total cholesterol (mmol/L)	0.002	-0.374
LDL (mmol/L)	0.038	-0.258
Triglyceride (mmol/L)	0.005	-0.339
TIBC (g/L)	0.041	-0.254
Malnutrition Inflammation Score	0.028	-0.272
hs-CRP (mg/L)	N.S.	-0.215
TNF- α (pg/mL)	N.S.	-0.141

Figure 1. Correlation graphic of leptin and MIS.

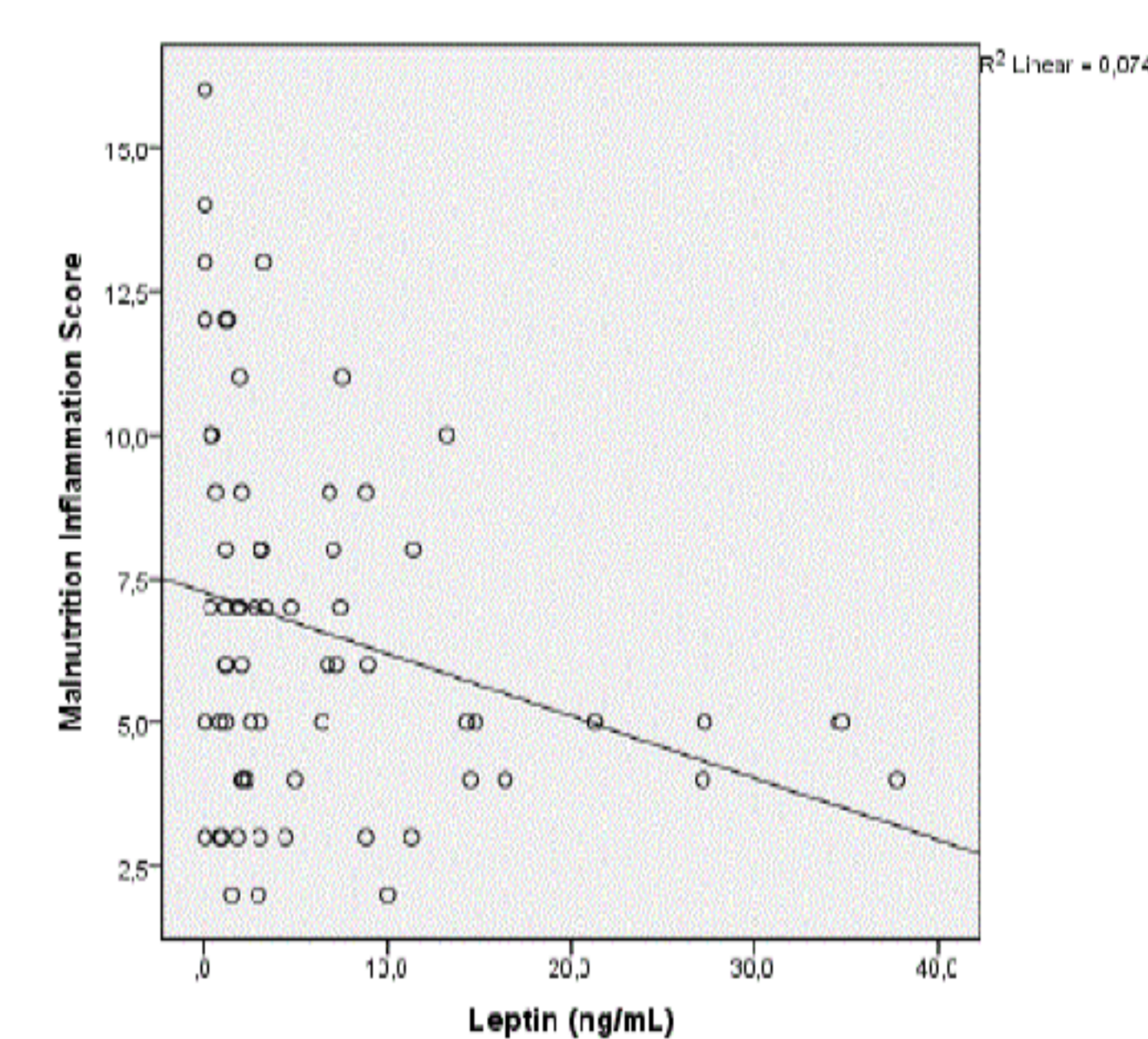


Table 3. Linear regression analyse of leptin and BMI as a predictor of MIS.

Predictor	r	Adj R ²	%95 Confidence Interval		F	P
			Lower	Upper		
Leptin	-0.272	0.059	-0.204	-0.012	5.037	0.028
BMI	0.533	0.273	-0.943	-0.405	25.04	0.001

Results:

Mean age and duration on dialysis of 65 patients (Male/Female: 34/31) were 51.6 \pm 17.8 years and 78.0 \pm 67.9 months, respectively. Mean leptin levels and malnutrition inflammation scores were 6.2 \pm 8.1 ng/mL and 6.6 \pm 3.2, respectively. As expected, leptin levels increased with older age, female gender, higher BMI and triceps skinfold thickness. There were no correlation between leptin and duration on haemodialysis. Elevated serum leptin levels were significantly correlated good nutritional status parameters including higher albumin (p= 0.001), prealbumin (p= 0.033), total iron binding capacity (p= 0.045), total cholesterol (p= 0.041) and lower malnutrition inflammation score (p= 0.002). In linear regression analyse, significantly negative correlation between serum leptin levels and malnutrition inflammation score remained after adjustment for BMI. No correlation was established between leptin and inflammation parameters including ferritin, hs-CRP and TNF- α .

Conclusions:

Elevated serum leptin levels seems to be associated with good nutritional status. However, the lack of correlation between leptin and inflammatory markers suggests that hyperleptinemia and inflammation are independent events in this population.

References:

- Sharma K, Considine RV, Michael B, et al. Plasma leptin is partly cleared by the kidney and is elevated in hemodialysis patients. *Kidney Int* 1997; 51:1980-1985.
- Kalantar-Zadeh K, Kopple JD, Block G, et al. A malnutrition-inflammation score is correlated with morbidity and mortality in maintenance hemodialysis patients. *Am J Kidney Dis* 38:1251-1263, 2001.
- Svobodova J, Haluzik J, Bednarova V, et al. Relation between serum leptin levels and selected nutritional parameters in hemodialysed patients. *Vnitr Lek* 2001;47:594-8.

