

http://www.jnephropharmacology.com

Journal of Nephropharmacology



Association of serum leptin with serum C-reactive protein in hemodialysis patients

Ebrahim Rastegari¹, Hamid Nasri^{1*}

¹Section of Hemodialysis, Shahrekord University of Medical Sciences, Shahrekord, Iran

ARTICLE INFO	A B S T R A C T				
Article Type:	Introduction: Recent investigations have shown that leptin is cleared principally by the kidney.				
Original	- Objectives: To examine whether and how in patients on hemodialysis the level of C-reactive protein leve correlate with serum leptin.				
Article History:	Patients and Methods: The total patients were 36. The mean patients' age were 46 (16) years. The median				
Received: 24 March 2012	length of the time patients were on hemodialysis were 19 months.				
Accepted: 22 June 2012	Results: The mean serum C-reactive protein was 8.7 (6.6) mg/l (median: 8 mg/l). The mean serum leptin				
ePublished: 1 July 2012	 was 9.4 (14) ng/ml (median: 5.75 ng/ml). In this study we found a significant inverse correlation of serun leptin with serum C-reactive protein (r= -0.57, p= 0.041) was seen. 				
Keywords:	Conclusion: Our data supports, the positive effect of leptin on nutrition and support the theory of				
Hemodialysis	protective effects (reverse epidemiology) of leptin in hemodialysis patients.				
End-stage renal failure					
Leptin					
C-reactive protein					

Implication for health policy/practice/research/medical education:

In a study on 36 stable hemodialysis patients, we found, the inverse correction of leptin with C-reactive protein level. Our data supports, the positive effect of leptin on nutrition and support the theory of protective effects (reverse epidemiology) of leptin in hemodialysis patients.

Please cite this paper as: Rastegari E, Nasri H. Association of serum leptin with serum C-reactive protein in hemodialysis patients. J Nephropharmacol 2012; 1(2): 19-21.

Introduction

Recent investigations have shown that leptin is cleared principally by the kidney. Serum leptin levels are increased in patients with chronic kidney failure and those undergoing hemodialysis (1,2) and it has been thought that high leptin value may contribute to uremic anorexia and malnutrition (4,5). In the general population leptin is considered an "appetite inhibitor", however in contrast to preliminary findings, its role in chronic renal disease and hemodialysis patients is not completely found. While serum leptin is generally elevated in chronic kidney failure and hemodialysis patients, however, some other studies have not been shown to be a cause of uremia-related anorexia (6,7). More recent investigations in hemodialysis patients suggest a paradoxically inverse correction between higher serum leptin and improved markers of nutritional condition (6,7), a finding that is indicative of reverse epidemiology in this population (8). In fact, leptin,

similar to serum albumin, has been reported to be a negative acute phase reactant in end-stage kidney failure (7). C-reactive protein (CRP) is an acute phase protein whose synthesis in the liver. CRP is regulated by different cytokines, especially interleukin 6 (IL-6). Plasma level of CRP in the absence of active disease are low, but can rise up to 1000-fold in patients with an inflammatory reaction. In addition to being a marker of inflammation, CRP itself may have pro-inflammatory properties while it can activate the complement system (8,9). Therefore elevated plasma concentrations of CRP, a sensitive marker of underlying systemic inflammation (10-12). Serum CRP value have also been shown to be significantly elevated in patients on hemodialysis (13,14) and suggests chronic inflammation. Serum CRP, is a sensitive and independent marker of malnutrition (15). According to the present data, investigations concerning the association of CRP with serum leptin needs further examination.

*Corresponding author: Prof. Hamid Nasri, Department of Internal Medicine, Shahrekord University of Medical Sciences, Iran. E-mail: hamidnasri@ yahoo.com

Objectives

The aim this study was to elucidate whether and how in patients on hemodialysis the serum value of CRP as the marker of inflammation correlate with serum leptin.

Patients and Methods

Patients

This cross-sectional study was conducted on patients with end-stage kidney disease, who were undergoing regular hemodialysis. The study was conducted in hemodialysis section of Shahrekord University of Medical Sciences.

Laboratory assessments

Blood samples were obtained after an overnight fast. For patients, Levels of serum CRP, leptin (normal range of values for males is 3.84 (1.79) and for females is 7.36 (3.73) ng/ml were measured by enzyme-linked immunosorbent assay (ELISA). Duration and doses of hemodialysis treatment were calculated from the patients' records. The duration of each hemodialysis session was 4 hours. For the efficacy of hemodialysis the urea reduction rate (URR) was calculated from pre- and postblood urea nitrogen (BUN) data (16). Body mass index (BMI) calculated using the standard formula (postdialyzed weight in kilograms/height in square meters; kg/m²) (17).

Ethical approval

All patients signed the consent form for participation in this study. Research study was approved by the ethics committee of Shahrekord University of Medical Sciences, Iran.

Statistical analysis

Results are expressed as the mean (SD) and median values. Statistical correlations were assessed using partial correlation test. For leptin correlation, the logarithm of serum leptin values was used. All statistical analyses were performed using SPSS (version 12.00). Statistical significance was determined at a p-value < 0.05.

Results

Study patients were 36. Table 1 shows patients' data. The mean patients' age were 46 (16) years. The mean length of the time patients had received hemodialysis was 32 (36) months. The mean serum CRP was 8.7 (6.6) mg/l. The mean serum leptin was 9.4 (14) ng/ml (median: 5.75 ng/ml). In this study a significant inverse correlation of serum leptin with serum CRP (r=-0.57, p=0.041) was seen.

Discussion

In this study we detected a significant inverse correlation of serum leptin with CRP value. The increased levels of leptin in hemodialysis patients are mainly due to retention of the hormone. Recent studies in maintenance dialysis patients suggest a paradoxically inverse association between higher

Table 1. Data of patients.

Total patients= 36	Minimum	Maximum	Mean (SD)	Median
Age (years)	16	80	46(16)	43
DH* (months)	2	156	32(36)	19
CRP mg/l	3	40	8.7(6.6)	8
Mg mg/dl	1.6	3.5	2.5±0.4	2.4

serum leptin and improved markers nutritional status (6,7), a finding that is in accord with the theory of reverse epidemiology (8,18-21). Indeed, leptin, similar to serum albumin, has been reported to be a negative acute phase reactant in hemodialysis patients (7). While, in the general population, leptin is considered an "appetite inhibitors", its role in hemodialysis patients is somewhat different. Serum leptin is generally elevated in hemodialysis, but this has not been shown to be related to anorexia. In contrast leptin has been shown to act synergistically with erythropoietin to stimulate the end-stage colony-forming-unit erythroid in humans (22).

Conclusion

Our data indirectly supports the some previous investigations regarding the hypothesis that in patients on hemodialysis, leptin is a negative acute phase reactant and leptin might have a positive effects on nutrition and support the theory of protective effects of leptin in hemodialysis patients.

Author's contributions

ER and HN wrote the manuscript equally.

Conflict of interests

The authors declared no competing interests.

Ethical considerations

Ethical issues (including plagiarism, misconduct, data fabrication, falsification, double publication or submission, redundancy) have been completely observed by the authors.

Funding/Support

This study was supported by a grant from Shahrekord University of Medical Sciences.

References

- 1. Zoccali C, Panuccio V, Tripepi G, Cutrupi S, Pizzini P, Mallamaci F. Leptin and biochemical markers of bone turnover in dialysis patients. J Nephrol 2004; 17: 253-260.
- 2. Wolf G, Chen S, Han DC, Ziyadeh FN.Leptin and renal disease. Am J Kidney Dis 2002; 39(1): 1-11.
- 3. Stamatiadis DN, Chan JL, Cogswell R, Stefanopoulou HC, Bullen J, Katsilambros N. et al. Elevated leptin fragments in renal failure correlate with BMI and haematopoiesis and are normalized by haemodialysis. Clin Endocrinol (Oxf) 2004; 60(4): 434-41.
- Stenvinkel P. Leptin and Its Clinical Implications in Chronic Renal Failure. Miner Electrolyte Metab 1999; 25: 298-302.
- Nakazono H, Nagake Y, Ichikawa H, Makino H. Serum Leptin Concentrations in Patients on Hemodialysis. Nephron 1998; 80: 35-40.
- 6. Pecoits-Filho R, Lindholm B, Stenvinkel P. End-stage renal disease: a state of chronic inflammation and hyperleptinemia. Eur J Clin Invest 2003; 33: 527-8.
- Don BR, Rosales LM, Levine NW, Mitch W, Kaysen GA. Leptin is a negative acute phase protein in chronic hemodialysis patients. Kidney Int 2001; 59: 1114-20.
- Wolbink GJ, Brouwer MC, Buysmann S, ten Berge IJ, Hack CE. CRP-mediated activation of complement in vivo: assessment by measuring circulating complement-C-reactive protein complexes. J Immunol 1996; 157: 473-9.

20

- 9. Morrow DA, Ridker PM. C-reactive protein, inflammation, and coronary risk. Med Clin North Am 2000; 84: 149-61.
- 10. Clyne B, Olshaker JS. The C-reactive protein. J Emerg Med 1999; 17: 1019-25.
- 11. McIntyre C, Harper I, Macdougall IC, Raine AE, Williams A, Baker LR. Serum C-reactive protein as a marker for infection and inflammation in regular dialysis patients. Clin Nephrol 1997; 48: 371-4.
- 12. Haubitz M, Brunkhorst R, Wrenger E, Froese P, Schulze M, Koch KM. Chronic induction of C-reactive protein by hemodialysis, but not by peritoneal dialysis therapy. Perit Dial Int 1996; 16: 158-62.
- 13. Stenvinkel P, Heimburger O, Lindholm B, Kaysen GA, Bergstrom J. Are there two types of malnutrition in chronic renal failure? Evidence for relationships between malnutrition, inflammation and atherosclerosis (MIA syndrome). Nephrol Dial Transplant 2000; 15: 953-60.
- Panichi V, Migliori M, De Pietro S, Metelli MR, Taccola D, Perez R, et al. Plasma C-Reactive Protein in Hemodialysis Patients: A Cross-Sectional, Longitudinal Clinical Survey. Blood Purif 2000; 18 :30-6.
- 15. Ortega O, Rodriguez I, Gallar P, Carreno A, Ortiz M, Espejo B, et al. Significance of high C-reactive protein levels in pre-dialysis patients. Nephrol Dial Transplant 2002; 17: 1105-9.
- 16. Nasri H, Baradaran A, Naderi AS. Close association

between parathyroid hormone and left ventricular function and structure in end-stage renal failure patients under maintenance hemodialysis. Acta Med Austriaca 2004; 31(3): 67-72.

- 17. Baradaran A, Nasri H. Association of Helicobacter pylori IgG antibody with various demographic and biochemical parameters in kidney transplant recipients. Saudi J Kidney Dis Transpl 2011; 22(6): 1115-20.
- Coen G. Leptin and bone metabolism. J Nephrol 2004; 17: 187-9.
- 19. Stenvinkel P, Lindholm B, Lonnqvist F, Katzarski K, Heimburger O. Increases in serum leptin levels during peritoneal dialysis are associated with inflammation and a decrease in lean body mass. J Am Soc Nephrol 2000; 11: 1303-9.
- 20. Odamaki M, Furuya R, Yoneyama T, Nishikino M, Hibi I, Miyaji K, et al. Association of the serum leptin concentration with weight loss in chronic hemodialysis patients. Am J Kidney Dis 1999; 33: 361-8.
- 21. Heimburger O, Lonnqvist F, Danielsson A, Nordenstrom J, Stenvinkel P. Serum immunoreactive leptin concentration and its relation to the body fat content in chronic renal failure. J Am Soc Nephrol 1997; 8: 1423-30.
- 22. Mikhail AA, Beck EX, Shafer A. Leptin stimulates fetal and adult erythroid and myeloid development. Blood 1997; 89: 1507-12.

Copyright © 2012 The Author(s); Published by Society of Diabetic Nephropathy Prevention. This is an open-access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.