# RENAL FAILURE Vol. 26, No. 1, pp. 55–57, 2004

CLINICAL STUDY

# Mild Renal Injury in Behçet's Disease#

Idris Sahin, 1,\* Tevfik Noyan, Ece Karaman, Ahmet Sadık Yalçınkaya, Ahmet Metin, and Tekin Akpolat 4

<sup>1</sup>Department of Internal Medicine and Nephrology, <sup>2</sup>Department of Biochemistry, and <sup>3</sup>Department of Dermatology, Yüzüncü Yıl University Medical School Training Hospital, Van, Turkey <sup>4</sup>Department of Internal Medicine and Nephrology, Ondokuz Mayıs University Medical School, Samsun, Turkey

#### **ABSTRACT**

Aim: The aim of this study is to investigate the frequency of microalbuminuria and abnormal urinary beta<sub>2</sub>-microglobulin excretion in patients with Behçet's disease (BD). Materials and Methods: Twenty-eight patients and 27 healthy controls were included in this study. Urine albumin/creatinine and beta<sub>2</sub>-microglobulin/creatinine ratios were calculated. Results: The frequency of microalbuminuria and abnormal urinary beta<sub>2</sub>-microglobulin excretion was higher among patients with BD than in control group, but this was not statistically significant (p>0.05). Conclusion: Microalbuminuria and abnormal beta<sub>2</sub>-microglobulin excretion are markers of renal injury, which have not been investigated in BD previously. Renal injury in BD is more frequent than has been recognized and it is most often in mild nature.

Key Words: Behçet's disease; Renal injury; Microalbuminuria; b-2-microglobulin.

### INTRODUCTION

Behçet's disease (BD) was first described by Hulusi Behçet in 1937 and consists of a triad of recurrent ulcers of the oral and genital mucosa with relapsing uveitis.<sup>[1,2]</sup> Since its first description, additional organ involvements have been documented. Skin, central nervous system, and pulmonary involvements are among those reported. Subcutaneous thrombophlebitis, deep vein thrombosis, epididymitis, arterial occlusion and/or aneurysms, arthralgia, arthritis, family history, renal problems and gastrointestinal features are

55

<sup>\*</sup>See Ref. [1].

<sup>\*</sup>Correspondence: Dr. Idris Sahin, Yüzüncü Yıl Universitesi Tip Fakultesi, Ic Hastalıkları AD, Maras Cad. 65200, Van, Turkey; Fax: 00 90 432 216 75 19; E-mail: idrissahin41@hotmail.com.

other manifestations of BD. [2-8] Cumulative analysis of renal involvement in BD demonstrated 159 cases with specific renal disease recently[9] and thereafter several more cases with specific renal disease have been reported. The frequency of renal problems associated with BD reported to vary between 0% to 55%. [9] The aim of this study is to investigate the frequency of microalbuminuria and abnormal urinary beta2-microglobulin excretion in patients with BD.

#### PATIENTS AND METHODS

Twenty-eight patients (9 male, 19 female, mean age 30 years, range 19-55) and 27 healthy controls (13 male, 14 female, mean age 37 years, range 22-65) were included in this study. All patients fulfilled the criteria of International Study Group for the diagnosis of BD.[16] Patients having genito-urinary infection or a history of urogenital disease or received cyclosporine previously were excluded.

Urine creatinine concentrations were measured by Hitachi modular PP autoanalyzer (Tokyo, Japan) using kit named Creat-Roche (Roche Diagnostics GmbH, Mannheim, Germany). Urinary albumin excretion was detected by immunoturbidimetric assay (Alb plus-Roche, Roche Diagnostics GmbH, Mannheim, Germany). Urine beta2-microglobulin was measured by chemiluminescence method (Immulite, BIODPC, Los Angeles, USA). Urine albumin/creatinine and beta2microglobulin/creatinine ratios were calculated.

Microalbuminuria was defined as albumin/creatinine ratio (mg/mmol) between 2.38 to 19 (men) and 2.96 to 20 (women).[17] Urinary beta2-microglobulin/ creatinine ratio higher than 300 µg/gr were accepted as abnormal.[18]

Table 1. Microalbuminuria and abnormal urinary beta2microglobulin excretion among the patients with Behçet's disease (BD) and controls.

Parameter	Patients with BD (n=28)	Control group (n=27)
Microalbuminuria		
Present	8 (29%)	2 (7%)
Absent	20	25
Abnormal urine		
Beta <sub>2</sub> -microglobulin excretion		
Present	7 (25%)	3 (11%)
Absent	21	24

Chi-square test with Yates' correction was used statistical analysis and a p value less than 0.05 was accepted statistically significant.

#### RESULTS

The number of patients and controls with microalbuminuria and abnormal urinary beta2-microglobulin excretion are presented in Table 1. Although the frequency of microalbuminuria and abnormal urinary betamicroglobulin excretion was higher among patients with BD than in control group, this was not statistically significant (p>0.05).

#### DISCUSSION

Renal problems associated with BD have been reported many times since 1963 and the frequency of renal involvement shows a wide variation. Oshima et al.[19] found abnormal urine findings in 13 of 65 (20%) patients with BD. Chaiek and Fainaru have published one of the striking articles regarding renal involvement in BD in 1975. [3] Based on the evaluation of the clinical findings of 683 previously published cases and their 41 cases, Chajek and Fainaru stated that 'BD does not affect the kidney'. Herreman et al. [20] have investigated urinary findings and kidney biopsies of 10 patients without any symptoms related to renal disease. They found proteinuria more than 100 mg/day in four patients and described several abnormalities by light microscopy and immunofluorescent staining even in patients without proteinuria. Rosenthal et al. [21] have evaluated urinalysis records of 77 patients retrospectively (urinalysis records were available only in 33 patients) and 51 patients prospectively. They found proteinuria and/or microhematuria in 25 of the 84 patients (30%).

Besides, in general, a control group was not present and renal problems were only a small and insignificant part in the studies reporting frequency of renal problems in BD. In addition, renal problems have not been addressed in many studies. [22,23] Although renal problems associated with BD are not common in clinical practice, it may progress to renal failure. Amyloidosis, glomerulonephritis and cyclosporine nephrotoxicity are the major causes of renal failure in BD.[10] Cyclosporine is used in the management of some mani-

Microalbuminuria and abnormal betaz-microglobulin excretion are markers of renal injury, which have not been investigated in BD previously. Although the



RIGHTS LINK()

## Mild Renal Injury in Behçet's Disease

number of patients included in this study is limited, the frequency of renal injury is higher compared to the control group. Our finding supports the previous data which shows renal involvement is not rare. In conclusion, renal injury in BD is more frequent than has been recognized and it is most often in mild nature.

#### REFERENCES

- Shimizu, T.; Ehrlich, G.E.; Inaba, G.; Hayashi, K. Behçet's disease (Behçet's syndrome). Semin. Arthritis Rheum. 1979, 8, 223-260.
- Kaklamani, V.G.; Vaiopoulos, G.; Kaklamanis, P.G. Behçet's disease. Semin. Arthritis Rheum. 1998, 27, 197–217.
- Chajek, T.; Fainaru, M. Behcet's disease. Report of 41 cases and a review of the literature. Medicine (Baltimore) 1975, 54, 179–196.
- Koç, Y.; Güllü, İ.; Akpek, G.; Akpolat, T.; Kansu, E.; Kiraz, S.; et al. Vascular involvement in Behçet's disease. J. Rheumatol. 1992, 19, 402–410.
- Akpolat, T.; Koç, Y.; Yeniay, İ.; Akpek, G.; Güllü, İ.; Kansu, E.; et al. Familial Behçet's disease. Eur. J. Med. 1992, 1, 391–395.
- Yazici, H.; Yurdakul, S.; Hamuryudan, V. Behcet's syndrome. Curr. Opin. Rheumatol. 1999, 11, 53– 57.
- Zouboulis, C.C. Epidemiology of Adamantiades Behçet's disease. Ann. Med. Interne 1999, 150, 488–498.
- Akpolat, T. Management of patient with Behçet's disease. Nephrol. Dial. Transplant. 1998, 13, 3002– 3004
- Akpolat, T.; Akkoyunlu, M.; Akpolat, I.; Dilek, M.; Odabas, A.R.; Ozen, S. Renal Behçet's disease: a cumulative analysis. Semin. Arthritis Rheum. 2002, 31, 317–337.
- Akpolat, T.; Diri, B.; Oguz, Y.; Yilmaz, E.; Yavuz, M.; Dilek, M. Behçet's disease and renal failure. Nephrol. Dial. Transplant., in press.
- Kaklamani, V.G.; Nikolopoulou, N.; Sotsiou, F.; Billis, A.; Kaklamanis, P. Renal involvement in Adamantiades-Behcet's disease. Case report and review of the literature. Clin. Exp. Rheumatol. 2001, 19 (Suppl). S55-S58.
- 12. Zahiri, K.; Hachim, K.; Zamd, A.; Fatihi, E.; Benghanem, M.G.; Ramdani, B.; Sqalli, S.; Zaid,

- D. Renal involvement in Behcet's disease. About six cases. Rev. Med. Interne **2003**, 24, 4-10.
- Altiparmak, M.R.; Tanverdi, M.; Pamuk, N.; Tunç, R.; Hamuryudan, V. Glomerulonephritis in Behcet's disease: report of seven cases and review of the literature. Clin. Rheumatol. 2002, 21, 14–18.
- Aksu, K.; Kocanaogullari, H.; Keser, G.; Inal, V.; Kabasakal, Y.; Oksel, F.; Gumusdis, G.; Doganavsargil, E. A case of Behcet's disease with pulmonary arterial aneurysm and secondary amyloidosis. Rheumatology (Oxford) 2002, 41, 831–832.
- Cengiz, N.; Karaoglanoglu, M.; Cengiz, K.; Akpolat, T. Behcet's disease and focal segmental glomerulosclerosis. Clin. Nephrol. 2003, 59, 234– 235
- International Study Group for Behçet's disease. Criteria for diagnosis of Behçet's disease. Lancet 1990, 335, 1078–1080.
- Pontremoli, R.; Sofia, A.; Ravera, M.; Nicolella, C.; Viazzi, F.; Tirotta, A.; Ruello, N.; Tomolillo, C.; Castello, C.; Grillo, G.; Sacchi, G.; Deferrari, G. Prevalence and clinical correlates of microalbuminuria in essential hypertension: the MAGIC Study. Hypertension 1997, 30, 1135-1143.
- Koh, K.T.; Chia, K.S.; Tan, C. Proteinuria and enzymuria in non-insulin-dependent diabetics. Diabetes Res. Clin. Pract. 1993, 20, 215–221.
- Oshima, Y.; Shimizu, T.; Yokohari, R.; Matsumoto, T.; Kano, K.; Kagami, T.; et al. Clinical studies on Behçet's syndrome. Ann. Rheum. Dis. 1963, 22, 36–45.
- Herreman, G.; Beaufils, H.; Godeau, P.; Cassou, B.; Wechsler, B.; Boujeau, J.; et al. Behcet's syndrome and renal involvement: a histological and immunofluorescent study of eleven renal biopsies. Am. J. Med. Sci. 1982, 284, 10-17.
- 21. Rosenthal, T.; Weiss, P.; Gafni, J. Renal involvement in Behcet's syndrome. Arch. Intern. Med. 1978, 138, 1122–1124.
- Ben Hmida, M.; Hachicha, J.; Kaddour, N.; Makni, H.; Aydel, F.Z.; Chakroun, N.; et al. ANCA in Behcet's disease. Nephrol. Dial. Transplant. 1997, 12, 2465–2466.
- Calamia, K.T.; Mazlumzadeh, M.; Balabanova, M.; Bagheri, M.; O'Duffy, J.D. Clinical characteristics of United States patients with Behçet's disease. IXth International Conference on Behçet's Disease. Yonsei Med. J. 2000, 41, 9S (abstract).