Factors associated with hip pain in end-stage renal disease patients on prevalent hemodialysis: a cross-sectional study

Hüma Bölük Şenlikci¹*, Sevgi İkbali Afsar¹, Selin Özen¹ and Cihat Burak Sayın²

Abstract

Background: Hemodialysis (HD) patients suffer from musculoskeletal disorders. The most reported musculoskeletal problem is arthralgia. Hip arthralgia has been commonly reported in patients undergoing HD. Hip pain can lead to a decrease in levels of physical activity, limitation in joint range of motion, and consequently difficulties in performing activities of daily living (ADL) and impair the quality of life (QoL). The aim of the study is to reveal the prevalence of hip pain and related factors in HD patients. This cross-sectional study included 73 patients on prevalent HD whose ages ranged from 25 to 65 years and who were on HD for more than 6 months. Physical examination and radiological imaging were done to every patient. Visual analog scale, Barthel Index, and Short Form-36 were used to evaluate pain, ADL, and QoL, respectively.

Results: Hip arthralgia was detected in 32 patients. Around 43% of which were diagnosed hip osteoarthritis, 34% greater trochanteric pain syndrome, 15% femoroacetabular impingement, and 6% soft tissue calcifications. Diabetes mellitus and hemodialysis duration were found to be significantly different between the groups of hip pain and without hip pain. Diabetes mellitus was identified as an independent risk factor for hip pain in hemodialysis patients. ADL and QoL were significantly lower in patients with hip pain compared to those without (p < 0.01; p < 0.05, respectively).

Conclusions: The results of our research show that HD patients should be screened for the presence of hip pain and other musculoskeletal disorders and that this is an area which requires further consideration and medical research.

Keywords: Renal dialysis, Musculoskeletal pain, Osteoarthritis, Chronic kidney failure

Background

End-stage renal disease patients undergoing hemodialysis (HD) suffer from various symptoms related to many organ systems. Some of these are musculoskeletal dysfunction and pain. As much as 80% of patients undergoing HD have chronic musculoskeletal pain, particularly arthralgia [1]. Approximately 40% of HD patients have previously reported lower extremity pain [2]. Hip pain is one of the causes of musculoskeletal pain in patients undergoing HD. Osteoarthritis, inflammatory arthritis, greater trochanteric pain syndrome (GTPS), avascular necrosis, femoroacetabular impingement (FAI), and osteoporosis resulting in fractures are among the most common causes of hip pain in the general population [3, 4]. Hip pain can lead to a decrease in levels of physical activity, muscle weakness, limitation in joint range of motion, and consequently difficulties in performing activities of daily living (ADL) and quality of life (QoL) [5].

Hip pain and other musculoskeletal symptoms are overlooked as serious problems of HD patients. The aim
of the present study is to evaluate the prevalence and features of hip pain and to reveal related factors and effects on health-related quality of life in hemodialysis patients.

**Method**

**Study design and participants**

A cross-sectional study of patients with end-stage renal disease receiving HD at the Baskent University Faculty of Medicine, Hemodialysis Units, Ankara, Turkey. The study was conducted between 2019 January and 2020 January. Patients aged between 25 and 65 years undergoing HD for more than 6 months, three times a week, who did not have acute infection and cognitive impairment and had stable general condition were included in the study. Patients were questioned regarding symptoms of anterior and lateral hip pain or pain in the groin. Patients who had a history of rheumatologic disorders, cerebrovascular disease, total hip replacement, uncontrollable systemic disease, active cancer, and cognitive impairment and those who did not agree to participate in the trial were excluded from the study. As a result, a total of 73 patients (32 patients with hip pain and 41 without) were included in the study.

The study was performed in accordance with the Declaration of Helsinki. It was approved by the Ethics committee of Baskent University Faculty of Medicine, Hemodialysis Units, Ankara, Turkey. The study participants beginning with an inspection followed by palpation of the greater trochanter and iliotibial band, joint range of motion (ROM), and gait evaluation. Specific tests such as the flexion abduction external rotation (FABER) test and flexion adduction internal rotation (FADIR) test were performed. Gait abnormalities, such as the presence of the Trendelenburg gait or an antalgic gait, were recorded. Neurological examination including sensory-motor examination and eliciting lower extremity deep tendon reflexes was performed. The straight leg raise test and femoral nerve stretch test were performed to rule out lumbar pathologies.

All patients with hip pain were evaluated radiologically. For a definitive diagnosis, X-ray and magnetic resonance imaging (MRI) were performed to identify the etiology of hip pain.

The diagnosis of hip osteoarthritis (OA) was made according to The American College of Rheumatology classification criteria [10]. The X-ray of each hip was scored using the Kellgren and Lawrence scale which is used to grade the stage of osteoarthritis: grade 0 refers to a normal joint, grade 1 refers to possible osteophytes with normal joint space, grade 2 refers to a possible narrowing of the joint space with definite osteophytes, grade 3 refers to a definite narrowing of the joint space, and finally grade 4 refers to a marked narrowing of the joint space, possible cysts or sclerosis and usually osteophytes [11].

A patient received a diagnosis of greater trochanteric pain syndrome (GTPS) in the presence of lateral hip pain and/or distinct tenderness around the greater

**Data collection**

Demographic and clinical data such as age, gender, body mass index (BMI), HD duration, and co-morbidities (hypertension (HT), diabetes mellitus (DM), and cardiac disease (CD)) were collected during a face-to-face interview conducted by a physician. At the time of the interview, patients were also questioned on the presence of hip pain. In those with complaints of hip pain, duration, characteristics, localization, and spread of the pain were questioned. Pain severity was assessed using the VAS and classified as mild (0–4), moderate (5–7), and severe (8–10). Pain lasting more than 3 months was classified as chronic pain and pain of less than 3-month duration was classified as acute pain.

All patients then underwent a detailed physical examination. Laboratory parameters as serum calcium, phosphate, alkaline phosphatase (ALP), parathyroid hormone, and 25-cholecalciferol levels were recorded.

**Outcome measures**

**Short Form-36**

The Short Form-36 (SF-36) is used to assess health-related QoL. The SF-36 consists of 36 items in 2 parts: mental and physical. The physical role, emotional role, physical function, energy/vitality, mental health, social function, pain, and general health are subgroups of assessment. Scores range from 0 to 100, with higher scores indicating a better health-related quality of life. The validity and reliability of the SF-36 has previously been studied [6].

**Barthel Index**

The Barthel Index is used to assess performance in ADL. Higher scores indicate greater ability of functional independence. The Barthel Index consists of ten personal activities: feeding, toileting, dressing and undressing, getting on and off the toilet, bladder-bowel control, transfers from bed to wheelchair and back, walking or using a wheelchair on a flat ground, and ascending-descending stairs [7].

**Visual analog scale**

A 10-cm visual analog scale (VAS) was used to assess the pain. Patients were asked to score the intensity of their pain [8, 9].

**Examination of hip pain**

Physical examination of the hip was performed in all study participants beginning with an inspection followed by palpation of the greater trochanter and iliotibial band, joint range of motion (ROM), and gait evaluation. Specific tests such as the flexion abduction external rotation (FABER) test and flexion adduction internal rotation (FADIR) test were performed. Gait abnormalities, such as the presence of the Trendelenburg gait or an antalgic gait, were recorded. Neurological examination including sensory-motor examination and eliciting lower extremity deep tendon reflexes was performed. The straight leg raise test and femoral nerve stretch test were performed to rule out lumbar pathologies.

A patient received a diagnosis of greater trochanteric pain syndrome (GTPS) in the presence of lateral hip pain and/or distinct tenderness around the greater
trochanter plus at least one of the following criteria: (1) pain at the extremes of hip rotation, abduction, or adduction; (2) pain on hip abduction against resistance; (3) tenderness over the gluteus medius muscle; (4) a positive FABER test; and (5) pain radiating down the lateral aspect of the thigh (pseudoradiculopathy). X-ray imaging was also used for differential diagnosis of these patients [12, 13].

Femoroacetabular impingement is also one of the most commonly occurring causes of hip pain. FAI was diagnosed in patients who described chronic, deep, and anterior groin pain while sitting or during activity and had positive FABER and FADIR tests. Over 90° hip flexion was limited or painful. A definitive diagnosis was made using X-ray imaging and MRI of the hip. On anteroposterior radiography, increased acetabular depth and/or presence of the crossover sign for focal acetabular retroversion and/or > 40° lateral center edge angle were detected. MRI was used in cases of doubt [14].

Statistical analysis

Statistical analyses were carried out using the SPSS version 20.0 software for Windows (IBM SPSS Inc., Chicago, IL). Conformity of the data to the normal distribution was assessed using the Kolmogorov-Smirnov test. The mean ± standard deviation (SD) of normally distributed data was provided. Median (minimum-maximum) values of non-normally distributed data were provided. Categorical data was shown in numbers (n) and percentages (%). The chi-square test and Fischer’s exact test were applied to categorical data. Between-group evaluation of normally distributed data was performed using the t-test. Non-normally distributed data was compared using the Mann-Whitney U test. Variables with a p value less than 0.25 were included in the multivariable regression model. The predictive value of the Barthel Index score was determined by the Youden index method of ROC curve analysis. A value of p ≤ 0.05 was accepted as statistically significant. Spearman’s rho correlation test was used for categorical non-parametric evaluations with the Pearson correlation test.

Results

In total, 41 male (56.2%) and 32 female (43.8%) patients were included in the study (mean age 54.4 ± 11.3). The median HD duration was 5 years (interquartile range 0.5–30 years). Approximately 45% (n = 33) of the patients had hypertension, 24.7% (n = 18) had diabetes mellitus, and 16.4% (n = 12) had coronary artery disease. A total of 32 patients undergoing HD (43.2%) had hip pain. Eighteen patients had previously been seen by a physical medicine and rehabilitation, rheumatology, and orthopedics specialist for the hip pain; the remaining 14 patients had never been evaluated by a physician due to hip pain. Other sociodemographic and clinical characteristics of the study participants are shown in Table 1. In the group comparisons, the number of patients with diabetes mellitus and those who had been receiving HD for more than 10 years was significantly greater in the hip pain group (37.5% vs 14.6%, p = 0.031; 43.8% vs 17.1%, p = 0.019, respectively). There was no significant difference between group difference in other clinical characteristics, demographic findings, and laboratory parameters (Table 1).

The outcomes of patients undergoing HD with hip pain are presented in Table 2. According to the VAS scores for hip pain, 12 patients had mild pain (37.5%), 14 patients had moderate pain (43.7%), and 6 patients had severe pain (18.7%). The majority of the patients had chronic hip pain (90.6%). Nearly half of the participants reported bilateral hip pain (46.9%), and 31.3% of the patients reported pain in the right hip. On hip joint examination, FADIR and FABER test positivity was detected in twenty and twenty-seven of the 73 patients, respectively. Tenderness by palpation was found in 14 patients. Limitation in hip range of motion was present in 11 patients. The etiology of the hip pain was osteoarthritis in 14 patients (43.6%) and GTPS in 11 patients (34.3%), 5 patients were diagnosed with FAI (15.6%), and the remaining 2 (6.2%) had soft tissue calcification. According to the Kellgren and Lawrence scale, the majority of the patients had grade 3 osteoarthritis (57.1%) of the hip (Table 2).

The Barthel Index scores were significantly lower in patients with hip pain than in those without (88.1 ± 14.5 vs 98.5 ± 3.0; p < 0.001). All SF-36 subgroup scores were significantly lower in patients with hip pain (Table 3).

Variables that may be of clinical significance in patients with hip pain are shown in Table 4 (p < 0.25). Sociodemographic and clinical characteristics of the patients were included in the multivariate logistic regression; diabetes mellitus was identified as an independent risk factor for hip pain in HD patients (OR = 3.91; p = 0.030) (Table 4).

The predictive value of Barthel Index at 95% CI showed sensitivity 68.8% and specificity 78.1% for hip pain (+LR 5.9; −LR 0.6; +PV 71%; −165 PV 76.2%) (Fig. 1).

There was no statistically significant correlation between hip pain VAS scores and vitamin D and PTH levels (r = −0.013, p = 0.916; r = 0.091, p = 0.442, respectively).

Discussion

The aim of this study was to evaluate the prevalence of hip pain in hemodialysis patients and investigate its effects on ADL and health-related QoL. The frequency of OA and GTPS was higher in hemodialysis patients when compared to previous studies of the general population. The frequency of FAI was similar to that of the general
population (10–15%) [15–18]. ADL and health-related QoL scores were significantly lower in those with hip pain.

Diabetes mellitus and hemodialysis duration were significant factors affecting the presence of hip pain. Moreover, diabetes mellitus was identified as an independent risk factor for hip pain.

Chronic pain is a common problem in HD patients and may occur in the form of headaches, arteriovenous fistula-related pain, widespread body pain, and musculoskeletal pain [19]. Musculoskeletal pain is the main cause of chronic pain in HD patients [20]; arthralgia secondary to osteoarthritis has been reported in 83% of patients [1, 20]. Hip pain occurs less commonly and is present in 7.5–18% of HD patients [1, 20–22]. In our study, the prevalence of hip pain was higher than previously reported in HD patients; this may be due to several reasons. Firstly, the mean age of the patient population of our study was greater than that in previous studies [22]. Secondly, in our study, the duration of dialysis was longer in patients with hip pain when compared to those without. Similarly, dialysis duration in the patients of our study was longer than that in previous studies [20]. In addition, the patients in the aforementioned study were evaluated only in terms of hip OA; 53.9% of those with hip pain had OA. Hence, this finding is in line with the prevalence of hip OA in our study.

Greater trochanteric pain syndrome is one of the common causes of hip pain. Patients mostly complain of pain radiating from the hip to the upper lateral aspect of the leg; however, the presence of tenderness around the greater trochanter without lateral leg pain may also be a sign of GTPS. In a study in which stroke patients with normal cognitive function were investigated in terms of GTPS, pain emerged with maneuvers that constitute the diagnostic criteria of GTPS in asymptomatic patients, and a diagnosis of GTPS was made based on these findings at a rate of 19% [15]. In our study, patients were evaluated for the presence of GTPS, and 11 GTPS diagnoses were made using direct radiography, excluding other diagnoses.

Other possible reasons for the high prevalence of hip pain in our study may be that all study participants were examined for signs of GTPS and FAI, rather than just those with symptoms of hip pain, and unlike in some other studies, all diagnoses were supported by imaging methods [1, 16, 20].

In our study, DM was identified as an independent risk factor for hip pain in HD patients. DM has previously been defined as a risk factor for chronic musculoskeletal

### Table 1 Demographic and clinical characteristics of the hemodialysis patients

|                      | All population n = 73 | Hip pain in HD patients | Without hip pain n = 41 | P     |
|----------------------|-----------------------|-------------------------|-------------------------|-------|
| **Age, years**       | 54.4 ± 11.3           | 54.0 ± 11.5             | 54.8 ± 11.2             | 0.771*|
| **Gender, n (%)**    |                       |                         |                         |       |
| Female               | 32 (43.8)             | 18 (56.3)               | 14 (34.1)               | 0.095**|
| Male                 | 41 (56.2)             | 14 (43.8)               | 27 (65.9)               |       |
| **BMI, kg/m²**       | 26.0 ± 6.2            | 26.5 ± 7                | 25.6 ± 5.6              | 0.553*|
| **HD duration, years** | 5 (0.5–30.0)       | 9 (0.5–30)              | 4 (0.5–24)              | 0.171**|
| < 10 years           | 52 (71.2)             | 18 (56.3)               | 34 (82.9)               | 0.019**|
| > 10 years           | 21 (28.8)             | 14 (43.8)               | 7 (17.1)                |       |
| **Co-morbidities**   |                       |                         |                         |       |
| HT, n (%)            | 33 (45.2)             | 13 (40.6)               | 20 (48.8)               | 0.636**|
| DM, n (%)            | 18 (24.7)             | 12 (37.5)               | 6 (14.6)                | 0.031**|
| CD, n (%)            | 12 (16.4)             | 5 (15.6)                | 7 (17.1)                | 0.999**|
| **Laboratory parameters** |                 |                         |                         |       |
| Calcium (mg/dL)      | 8.6 ± 0.8             | 8.5 ± 0.7               | 8.7 ± 1.0               | 0.414*|
| Phosphate (mg/dL)    | 5.2 ± 1.3             | 5.1 ± 1.1               | 5.3 ± 1.4               | 0.589*|
| PTH (pg/mL)          | 461 (1.5–2460)        | 493.2 (1.5–2000)        | 427 (8.5–2460)          | 0.443*|
| Vitamin D (ng/mL)    | 13 (1.0–62.7)         | 14.3 (3.5–49.2)         | 12.7 (1–62.7)           | 0.831*|
| < 20 ng/mL           | 54 (74.0)             | 24 (75.0)               | 30 (73.2)               | 0.999*|
| ALP (IU/mL)          | 118 (39–950)          | 140.5 (63–880)          | 99 (39–950)             | 0.059*|

**BMI** body mass index, **DM** diabetes mellitus, **HT** hypertension, **CD** cardiac disease, **HD** hemodialysis, **PTH** parathyroid hormone, **ALP** alkaline phosphatase

*Chi-square test, p ≤ 0.05 was accepted as statistically significant; **Mann-Whitney U test
pain in HD patients [23] and has been associated with tendinopathy, arthralgia, septic arthritis, and bone loss resulting in fractures of the hip joint [16, 23–25]. Previous studies have also shown that HD patients who have arthralgia have a poor QoL [26]; a relationship between many musculoskeletal system findings, poor physical function, and QoL has been identified [22, 26]. In our study, both mental and physical QoL scores of HD patients with hip pain were lower than those without hip pain.

When we compared the two groups in terms of laboratory parameters, there was no correlation between parathyroid hormone (PTH) levels and hip pain severity in HD patients. Similarly, no relationship was found between PTH level and chronic musculoskeletal pain in a study involving HD patients [16]. However, there are also studies claiming the opposite [27]. In the aforementioned study, bone pain was found to be associated with secondary hyperparathyroidism; however, a higher PTH level is not the only indicator of secondary hyperparathyroidism [27]. In our study, patients were not examined for secondary hyperparathyroidism. Therefore, it is not surprising that there was no direct relationship between hip pain and PTH level.

Another finding in our study was that there was no correlation between vitamin D levels and hip pain severity in HD patients. Additionally, when the groups with and without hip pain were compared in terms of vitamin D levels, no significant difference was found between them. In previous studies, a relationship was found between vitamin D level and musculoskeletal pain in healthy individuals [28, 29]. No such relationship was found in HD patients [30, 31]. Our findings appear to be in accordance with the literature. Half of the patients with hip pain in our study consisted of patients with hip OA. The relationship between vitamin D and osteoarthritis has been the subject of many previous studies; no correlation has been found between OA and vitamin D levels [32, 33], but some studies have shown a relationship between OA progression and vitamin D levels [34]. To date, studies in this area have not been conducted on HD patients.

In our study, the most common etiology of hip pain was OA, followed by GTPS and FAI. There are no studies evaluating the relationship between GTPS, FAI, and vitamin D in hemodialysis patients in the literature.

| Clinical characteristics of the hemodialysis patients with hip pain | HD patients with hip pain (n = 32) |
|--------------------------|-------------------------------|
| VAS value | n (%) |
| 0–4: mild | 12 (37.5) |
| 5–7: moderate | 14 (43.7) |
| 8–10: severe | 6 (18.7) |
| Duration of pain | |
| Acute < 3 months | 3 (9.4) |
| Chronic > 3 months | 29 (90.6) |
| Side of pain | |
| Right | 10 (31.3) |
| Left | 7 (21.9) |
| Bilaterally | 15 (46.9) |
| Etiology of pain, n (%) | |
| Osteoarthritis | 14 (43.8) |
| GTPS | 11 (34.3) |
| FAI | 5 (15.6) |
| Soft tissue or tendon calcification | 2 (6.2) |
| Kellgren and Lawrence scores | |
| Grade 1 | 2 (14.3) |
| Grade 2 | 2 (14.3) |
| Grade 3 | 8 (57.1) |
| Grade 4 | 2 (14.3) |

| Table 3 Comparison of Barthel Index scores and SF-36 subgroup scores between HD patients with and without hip pain | |
|--------------------------|--------------------------|
| Hip pain in HD patients | p |
| With hip pain | Without hip pain |
| n = 32 | n = 41 |
| Barthel Index scores | 88.1 ± 14.5 | 98.5 ± 3.0 | < 0.001* |
| SF-36 subgroup scores | | | |
| Physical function | 42.5 (0–90) | 80 (15–100) | < 0.001* |
| Physical role | 75 (0–100) | 75 (0–100) | 0.004* |
| Emotional role | 66.7 (0–100) | 100 (0–100) | 0.004* |
| Energy/vitality | 48.9 ± 13.2 | 57.2 ± 11.5 | 0.005* |
| Mental health | 61.1 ± 8.5 | 66.6 ± 9.8 | 0.014* |
| Social function | 56.6 ± 17.7 | 73.8 ± 16.5 | < 0.001* |
| General pain | 56.3 (0–100) | 77.5 (22.5–100) | < 0.001* |
| General health | 45 (10–65) | 55 (20–80) | < 0.001* |

| Table 4 Multivariate analysis of risk factors for hip pain in HD patients | OR | 95% CI | p |
|--------------------------|--------------------------|
| Diabetes mellitus | 3.91 | 1.17–13.05 | 0.030* |

OR odds ratio, CI confidence interval
*Logistic regression analysis

HD: hemodialysis, VAS: visual analog scale, GTPS: greater trochanteric pain syndrome, FAI: femoroacetabular impingement

*Chi-square test, p ≤ 0.05 was accepted as statistically significant
To the best of our knowledge, there are no studies to date which have evaluated hip pain alone in HD patients. The main limitations of the study are its cross-sectional design which did not allow the causality of the associations to be examined and the possibility that musculoskeletal pain due to polyneuropathy may have been confused with the musculoskeletal diagnosis. Another limitation is that patients were not investigated in terms of osteoporosis; this can be investigated using dual-energy X-ray absorptiometry in future studies. Finally, another limitation of this study included the small number of patients and not measuring the role of vitamin D deficiency on hip pain.

However, the use of radiological imaging further supported the diagnosis in those with GTPS, FAI, and OA. Conducting MRI of the hip in all patients to support the clinical diagnosis of the cause of hip pain may be considered in future studies. Although expensive, limiting the use of MRI may have overlooked diagnoses such as early avascular necrosis, amyloid deposits, crystal-induced arthropathies, deformities, and insufficiency fractures related to bone and mineral disturbance commonly encountered in patients on dialysis.

**Conclusion**
The findings of this study indicate that hip pain is more common in HD patients compared to the general population and that DM is an independent risk factor for hip pain in this patient group. The presence of hip pain also had a negative impact on ADL and QoL. The results of our research show that HD patients should be screened for the presence of hip pain and other musculoskeletal disorders and that this is an area which requires further consideration and medical research.

**Abbreviations**
CD: Cardiac disease; DM: Diabetes mellitus; FABER: Flexion abduction external rotation; FADIR: Flexion adduction internal rotation; FAI: Femoroacetabular impingement; GTPS: Great trochanteric pain syndrome; HD: Hemodialysis; HT: Hypertension; MRI: Magnetic resonance imaging; OA: Osteoarthritis; PTH: Parathyroid hormone; QoL: Quality of life; ROM: Range of motion; SF-36: Short Form-36; VAS: Visual analog scale

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Authors’ contributions
HBŞ and SIA: data collection, writing, and supervision. SO and CBS: statistical analyses, supervision, critical analyses, and data collection. All authors have read and approved the final manuscript.

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Availability of data and materials
Data is available.

Declarations

Ethics approval and consent to participate
The study was approved by the Ethical Committee of Başkent University Medical School (Project number: KA18/248).

Written informed consent was given from all participants.

Consent for publication
Not applicable

Competing interests
The authors declare that they have no competing interests.

Author details
1 Department of Physical Medicine and Rehabilitation, Başkent University Faculty of Medicine, Ankara, Turkey. 2 Department of Nephrology, Başkent University Faculty of Medicine, Ankara, Turkey.

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