

1 Sex-related differences in gout: a comparative study in a Moroccan cohort.

2

3 **ABSTRACT**

4 **Background:**

5 The aim of this study was to characterize sex-related differences in the clinical,
6 biological, and therapeutic features of gout in a Moroccan cohort.

7 **Methods:**

8 We conducted a cross-sectional study including 147 patients with gout (92 men
9 and 55 women). Epidemiological, clinical, biological, radiological, and
10 therapeutic characteristics were analyzed and compared according to sex.

11 **Results:**

12 Compared with men, women with gout were older (64.9 vs. 60.85 years) and
13 had a higher prevalence of cardiometabolic comorbidities, particularly
14 hypertension and diabetes ($p < 0.05$). Renal involvement was significantly more
15 frequent and more severe in women ($p = 0.003$). In contrast, men exhibited a
16 more inflammatory and severe disease profile, with higher pain scores and
17 elevated C-reactive protein levels ($p = 0.003$), as well as a higher frequency of
18 tophi and structural joint damage. No significant differences were observed
19 regarding monoarticular versus polyarticular presentation. Treatment patterns
20 were broadly similar between sexes, although colchicine use was significantly
21 more frequent in men ($p = 0.001$).

22 **Conclusion:**

23 Men and women with gout exhibit distinct clinical phenotypes. Female gout is
24 characterized by a predominantly metabolic and renal profile, whereas male

25 gout is associated with a more inflammatory and structural disease pattern.
26 These findings highlight the need for sex-specific management strategies.

27 **Keywords:**

28 Gout;Sex; Comorbidities; Renalimpairment; Inflammation

29 **INTRODUCTION**

30 Gout is the most common inflammatory arthritis, with a prevalence estimated
31 between 1% and 4% in Western countries, and has been steadily increasing
32 over recent decades [1–3]. This trend is largely attributed to population aging,
33 the rising prevalence of cardiometabolic comorbidities, and the increased use
34 of medications promoting hyperuricemia, particularly diuretics [2,3]. Gout is
35 associated with impaired quality of life, functional limitations, and increased
36 cardiovascular morbidity and mortality [1,23].

37 Historically considered a predominantly male disease, gout in women has
38 recently gained increasing attention. Its prevalence rises significantly after
39 menopause, likely due to the loss of the uricosuric effect of estrogens [4,10–
40 13]. Women with gout are generally older at diagnosis and present with a
41 higher prevalence of comorbidities, particularly hypertension, diabetes, and
42 chronic kidney disease, compared with men [5–8,14]. In contrast, men more
43 frequently exhibit lifestyle-related risk factors, such as alcohol consumption
44 and dietary habits, as well as a more inflammatory and structurally severe
45 disease pattern [2,4,8,9].

46 Despite these findings, gout in women remains under-recognized and
47 insufficiently studied. Many studies include relatively small numbers of women
48 or lack detailed sex-based analyses [5–7]. Moreover, treatment strategies are

49 often not sex-specific, despite the existence of distinct clinical profiles
50 suggesting the need for individualized management [15,16,24].

51 In this context, and given the increasing burden of gout, it is essential to better
52 characterize sex-related differences across diverse populations. Data from
53 North African countries remain scarce.

54 The aim of our study was therefore to compare epidemiological, clinical,
55 biological, radiological, and therapeutic characteristics of gout between men
56 and women in a Moroccan cohort. We hypothesized that female gout is
57 characterized by a predominantly metabolic and renal profile, whereas male
58 gout presents a more inflammatory and structural phenotype.

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60 **MATERIALS AND METHODS**

61 We conducted a cross-sectional study including 147 patients with gout (92 men
62 and 55 women), followed in a rheumatology setting. The diagnosis of gout was
63 based on clinical and biological findings and/or the identification of
64 monosodium urate crystals in synovial fluid, in accordance with the 2015
65 ACR/EULAR classification criteria.

66 Data were collected retrospectively from medical records and completed
67 during clinical visits. Variables included sociodemographic characteristics,
68 comorbidities (hypertension, diabetes, dyslipidemia, renal impairment), and
69 risk factors such as smoking, alcohol consumption, diuretic use, and family
70 history of gout.

71 Clinical assessment included age at disease onset, pattern of joint involvement,
72 number of flares, joint distribution, pain intensity assessed using a visual analog
73 scale (VAS), presence of tophi, and chronicity of gout. Biological parameters
74 included serum uric acid, C-reactive protein (CRP), and renal function assessed
75 by creatinine clearance.

76 Radiological evaluation included standard radiographs assessing structural
77 damage and ultrasonography identifying the double contour sign. Synovial fluid
78 analysis was performed when available to detect monosodium urate crystals.

79 Treatments were analyzed by distinguishing symptomatic therapy (colchicine,
80 nonsteroidal anti-inflammatory drugs, corticosteroids) and urate-lowering
81 therapy (allopurinol, febuxostat).

82 Quantitative variables were expressed as mean \pm standard deviation and
83 qualitative variables as frequencies and percentages. A bivariate analysis was
84 performed to compare variables according to sex, with a significance threshold
85 set at $p < 0.05$. No multivariate analysis was performed.

86 The study was conducted in accordance with the Declaration of Helsinki.

87 Approval was obtained from the local ethics committee, and informed consent
88 was obtained from all patients.

89 **Table 1. Sociodemographic characteristics, comorbidities, and risk factors**
90 **according to sex**

Variables	Men (n=92)	Women (n=55)	p-value
Meanage (years)	60.85 \pm 13	64.9 \pm 11	0.122
Diabetes	18.47%	38.18%	0.025

Hypertension	34.7%	58.18%	0.006
Dyslipidemia	18.4%	21.8%	0.622
Diuretic use	23.9%	18.9%	0.43
Smoking	13.04%	1.8%	0.032
Alcoholconsumption	9.2%	0%	0.02
Family history of gout	8.7%	0%	0.025

91 **Table 2. Clinical and biological characteristics according to sex**

Variables	Men	Women	p-value
Monoarticular	67.3%	74.5%	0.35
Polyarticular	32.6%	25.4%	0.36
VAS	49.89 ± 17	41.09 ± 18	0.05
Renalinvolvement	23.9%	31.6%	0.023
Tophi	10.8%	3.6%	0.04
Chronic gout	35.4%	29.3%	>0.05
Serumuricacid	81.23 ± 24	77.28 ± 24	>0.05
CRP	71.20 ± 19	46.77 ± 21	0.003
Creatinine clearance	51.4 ± 22	36.2 ± 21	0.003
MSU crystals	15.21%	10.9%	>0.05

93 **Table 3. Radiological and therapeutic data**

Variables	Men (n=92)	Women (n=55)	p-value
Radiographic erosions	34.5%	18.9%	>0.05
Double contour sign (ultrasound)	70.5%	88.8%	>0.05
Colchicine	88%	62%	0.001
NSAIDs	16.3%	9.09%	0.22
Corticosteroids	16.3%	11%	0.37
Allopurinol	78.26%	69.09%	0.32
Febuxostat	9.8%	12.7%	0.71

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95 **DISCUSSION**

96 In our study, we compared men and women with gout and identified significant
 97 differences in clinical, biological, and therapeutic profiles. Women were older
 98 and had a higher prevalence of cardiometabolic and renal comorbidities,
 99 whereas men exhibited a more inflammatory and structurally severe disease
 100 pattern.

101 Our findings are consistent with the literature, which reports a male
 102 predominance of gout but an increasing frequency in women, particularly after
 103 menopause [1–4]. Women in our cohort were older at diagnosis, in agreement
 104 with observational studies showing a delay of several years compared to men

105 [5–9]. This difference is largely attributed to the protective effect of estrogens
106 on urate metabolism, which declines after menopause [10–13].

107 Comorbidity profiles also differed by sex. Women more frequently had
108 diabetes, hypertension, and chronic kidney disease, as reported in several
109 international cohorts [5–8,14]. These findings likely reflect older age and a
110 higher burden of cardiovascular risk factors. In contrast, men more frequently
111 exhibited lifestyle-related risk factors, including smoking and alcohol
112 consumption [2,4,8,9,17,18].

113 Clinically, joint distribution did not differ significantly between sexes, consistent
114 with previous studies [19,20]. However, disease severity was greater in men,
115 with higher pain levels, more frequent tophi, and more pronounced structural
116 damage, in line with previous reports [2,3,8].

117 Biological findings reinforced this distinction. Serum uric acid levels were
118 similar, but men had higher inflammatory markers, whereas women had more
119 impaired renal function, suggesting different underlying mechanisms [7,21].

120 Therapeutic approaches were broadly similar. However, higher colchicine use
121 in men may reflect greater inflammatory activity. In contrast, more cautious
122 management is often adopted in women due to age and comorbidities,
123 consistent with current recommendations [15,16,24]. Previous studies have
124 also shown that women less frequently achieve target urate levels [4,6,8,23–
125 26].

126 These findings have important clinical implications. In women, management
127 should focus on controlling metabolic comorbidities and renal function. In men,
128 lifestyle modifications remain essential to prevent gout flares.

129 Our study has limitations, including its monocentric design, relatively small
130 sample size, and retrospective data collection. Additionally, some factors such
131 as dietary and hormonal variables were not systematically assessed.

132 Nevertheless, this study provides original data from a Moroccan population
133 and confirms clinically relevant sex-related differences in gout. These findings
134 support the need to integrate sex as a key determinant in gout management.

135 CONCLUSION

136 Men and women with gout exhibit distinct clinical phenotypes. Female gout is
137 characterized by a predominance of metabolic and renal comorbidities,
138 whereas male gout is associated with a more inflammatory and structurally
139 severe disease pattern.

140 These findings highlight the importance of considering sex-specific differences
141 in the clinical assessment and management of gout. Tailored therapeutic
142 strategies that account for comorbidities, renal function, and lifestyle factors
143 may improve patient outcomes.

144 Further studies, particularly prospective and multicenter investigations, are
145 needed to better understand these differences and optimize personalized
146 management approaches.

147 Conflict of interest

148 The authors declare no conflicts of interest.

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