Osteoporosis is a major public health problem worldwide, with increased morbidity and mortality due to its complications.\(^1\) Postmenopausal osteoporosis has attained the status of a major epidemic and most women with it present with a fracture as the first indication of the disease.\(^2\) In the USA, osteoporosis and the treatment of osteoporosis-related fractures costs are estimated to be in excess of $18 billion a year.\(^3,5\) Bubshait and Sadat-Ali\(^4\) estimated the yearly cost of treating osteoporosis-related femoral fractures in Saudi Arabia as SAR4.27 billion a year. Proper interventions could reduce this cost tremendously.

Glucocorticoids are considered an important component of therapy for a variety of medical conditions including autoimmune, rheumatic, pulmonary and gastrointestinal disorders. Patients treated with glucocorticoids are at risk for many adverse complications.\(^5\)

One of the most important complications associated with long-term use of glucocorticoids is glucocorticoid-induced osteoporosis (GIOP).\(^6,7\) Prolonged use of glucocorticoids causes osteocyte apoptosis with an increase in bone resorption and a decrease in bone formation at both the local and systemic levels, leading to rapid weakening of bone architecture and an increase in fracture risk.\(^8,9\) Evidence indicates that GIOP is the most common cause of secondary osteoporosis, leading to fractures in 30% to 50% of patients taking chronic glucocorticoids.\(^10\) However, studies showed that many patients treated with glucocorticoids are not properly evaluated and do not receive prophylaxis or treatment to prevent bone loss.\(^11,12\)

Osteoporosis secondary to sickle cell disease\(^13\) and cancer chemotherapy\(^14\) were reported among the Saudi Arab population. A review of the medical literature found no studies on GIOP in Saudis. We hypoth-
esized that evaluation and management of GIOP at our institution is below the standards. This study was carried out to answer the following questions: 1) To what extent is DEXA scanning performed for patients taking glucocorticoids? 2) What percentage of patients taking long-term glucocorticoids are simultaneously prescribed medications to prevent or treat GIOP?

**METHODS**

This retrospective study was approved by the research committee of King Faisal University and was carried out at King Fahd Hospital of the University, Al Khobar. We used the local electronic pharmacy prescription monitoring system to identify patients older than 18 years of age, who were prescribed ≥7.5 mg of prednisolone per day for 6 months or longer during the period 1 July 2007 through 31 December 2007. Patients who were prescribed other glucocorticoids apart from oral prednisolone or who took the steroids intermittently were not included in the study. Subsequently, each patient chart was reviewed for clinical data including age, sex, dose and duration of glucocorticoids therapy. A mean prednisolone dose was taken for patients receiving the minimum daily dose of ≥7.5 milligrams or more for the 6-month period. Dual-energy x-ray absorptiometry (DEXA) scan imaging reports were reviewed when available and data on BMD, T-score and Z-score were documented. Patients with a T score of ≤2.5 SD were defined as osteoporotic and those between –1 to –2.5 SD were defined as osteopenic for analysis, as defined per the WHO criteria.16 Patients who had a DEXA scan were divided into those younger than the age of 35 years, as they had achieved peak bone mass (PBM), and 35 years of age or older. The patient was considered to be screened if he had a BMD measurement while taking glucocorticoid therapy. The concomitant use of osteoporosis prophylaxis or treatment such as calcium, vitamin D, bisphosphonate, calcitonin, and hormonal therapy were noted.

Data was analyzed using a t test to compare means between the non-osteoporotic, osteopenic and osteoporotic patients. All tests were performed using SPSS (Statistical Package for the Social Sciences), version 14.0, Chicago, Illinois15 a P value of <.05 considered statistically significant.

**RESULTS**

During the period 1 July 2007 through 31 December 2007, 516 patients received oral corticosteroid therapy according to the local electronic pharmacy prescription monitoring system. One hundred and sixty-five patients (32.0%) received ≥7.5 mg of oral prednisolone per day for 6 months or longer, including 100 males and 65 female patients with a mean (SD) age and range of 37 (12.7) years (range, 7.5–60 years) for males and 40.8 (15) years (range, 18–62 years) for females (Table 1, 2). Only 26 patients (15.8%) had BMD measured by DEXA scan during glucocorticoid therapy who had normal BMD. Fourteen patients were <35 years and 12 were >35 years. In the former group, 10 (71.5%) were osteoporotic and 11 females and 2 males were osteopenic (Table 3). None of the patients who had a DEXA scan during glucocorticoid therapy had normal BMD. Fourteen patients were <35 years and 12 were >35 years. In the former group, 10 (71.5%) were osteoporotic compared to 5 (42%) in the latter group. Calcium supplementation was prescribed for 81 (81.0%) males compared to 40 (61.5%) females (P=.05). There was no significant difference in the frequency of prescription of vitamin D supplementation between males and females. None of the 165 patients who received ≥7.5

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**Table 1.** Primary diseases in patients (n=165) receiving ≥7.5 mg oral prednisolone daily for 6 months or longer.

| Primary disease                        | Number (%) |
|----------------------------------------|------------|
| Asthma                                 | 22 (13.3)  |
| Inflammatory bowel disease             | 15 (9)     |
| Idiopathic thrombocytopenic purpura    | 13 (7.9)   |
| Chronic nephritis                      | 16 (9.7)   |
| Neuropathies                           | 5 (3)      |
| Rheumatoid arthritis                   | 45 (27.3)  |
| Renal transplant                       | 33 (20)    |
| Systemic lupus erythematosus           | 16 (9.8)   |

**Table 2.** Prednisolone therapy and anti-osteoporotic prevention and treatment in 165 patients receiving daily prednisolone for 6 months or longer.

|                          | Male (n=100) | Female (n=65) | P (males vs. females) |
|--------------------------|--------------|---------------|-----------------------|
| Mean (SD) dose and range of prednisolone (mg) | 15.9 (12.7) (7.5–60) | 21.5 (19.5) (7.5–100) | .05 |
| Mean (SD) duration and range of prednisolone (mo) | 40.4 (29.9) (6–108) | 41.2 (36.4) (6–121) | .2 |
| Treatment with calcium   | 81 (81%)     | 40 (61.5%)    | .05                   |
| Treatment with vitamin D | 71 (71%)     | 38 (58.5%)    | .1                    |
| Antiresorptives/anabolics| 0            | 0             |                       |
Secondary osteoporosis is common but still neglected. It is the cause for osteoporosis in almost two-thirds of males, more than half of pre-menopausal and peri-menopausal females, and about one-fifth of post-menopausal females. Etiologies of secondary osteoporosis are many; however, GIOP is one of the leading causes. GIOP and its risk of fragility fractures are well recognized. The combined effect of higher dose, longer duration and a continuous pattern of glucocorticoid therapy could increase vertebral fracture risk by 17-fold and hip fracture risk by 7-fold. For this reason, many international guidelines regarding evaluation and treatment of GIOP have been developed. Those guidelines indicate that the use of more ≥5 milligrams for 3 months or longer warrants a DEXA scan to diagnose bone loss and to start prophylactic anti-resorptive therapy using a bisphosphonate or other drugs to prevent osteoporosis.

Our study showed that during a 6-month period a total of 516 patients were receiving oral prednisolone and 165 patients (32%) were taking ≥7.5 milligrams of steroid per day for 6 months or longer. According to international guidelines, all patients in the later group should have been screened and received effective prophylaxis or therapy for osteoporosis. Despite the fact that we used a higher dose and longer duration of therapy (≥7.5 milligrams for ≥6 months) than what is recommended in recent guidelines, only 26 (15.8%) of our patients were evaluated by DEXA scan and all of them had either osteopenia or osteoporosis. A DEXA scan was ordered for only 5 of 100 men. This indicates that physicians are less concerned about the development of osteoporosis in male patients despite the fact that men older than the age of 50 years were found by our group to have osteoporosis of the hip in 24.3% and osteoporosis of the spine in 37.4% in a previous study. The majority of our patients were taking calcium and vitamin D supplements, which is a higher rate than in other studies, but unfortunately, none of our patients, including those who were documented to have low bone mass, were prescribed bisphosphonates or any other anti-resorptive or anabolic therapy to prevent or treat osteoporosis. Neglect in providing adequate and recommended prophylaxis or therapy for patients on long-term glucocorticoids appears to be universal. In the study of Gudbjornsson et al (2002), only 9% of patients were taking a bisphosphonate.

Hart and Green (2002) found that 64.7% of their patients did not receive the proper prophylaxis for GIOP. Ungprasert et al (2007) believed that neglect by internal medicine physicians is quite common even in teaching institutions, as in their hospital only 5.8% of patients were evaluated for GIOP while Guzman-Clark and associates (2007) found that the common barrier for GIOP management was the physician lack of awareness and knowledge.

The mean age of our patients was lower than that reported in other studies. Our study has several shortcomings including all those associated with being retrospective. In addition, we did not have vitamin D levels and bone markers available for analysis. Lastly, we did not evaluate the indications of glucocorticoid therapy, which can be a contributing factor to low bone mass.

In conclusion, this study shows that GIOP is neither diagnosed early nor properly managed at our teaching institution. The fact that none of our patients who were on long-term glucocorticoid therapy had been prescribed anti-resorptive drugs raises a concern about the development of osteoporosis in male patients.
the lack of knowledge of international guidelines regarding GIOP. Awareness programs appear to be urgently needed for the physicians prescribing glucocorticoids in order to prevent GIOP and its complications.

REFERENCES

1. Nichols KJ. Evaluation of osteoporosis. J Am Osteopath Assoc. 2000;100:54-7.
2. Chopra A. Osteoporosis: a new understanding of its impact and pathogenesis. J Am Osteopath Assoc. 2000;100:51-4.
3. Iqbal MM. Osteoporosis: Epidemiology, diagnosis and treatment. South Med J. 2000;93:2-8.
4. Bulbsha D, Sadat-Ali M. Economic implications of osteoporosis-related femoral fractures in Saudi Arabian society. Calcif Tissue Int. 2007;81:459-66.
5. McDonough AK, Curtis JR, Saag KG. The epidemiology of glucocorticoid-associated adverse events. Curr Opin Rheumatol. 2008;20:131-137.
6. Mazziotti G, Angelini A, Bleizikian JP, Canalis E, Giustina A. Glucocorticoid-induced osteoporosis: an update. Trends Endocrinol Metab. 2006;17:144-9.
7. van Staa TP, Leufkens HG, Cooper C. The epidemiology of cortico-steroid-induced osteoporosis: a meta-analysis. Osteoporosis Int. 2002;13:777-87.
8. van Staa TP. The pathogenesis, epidemiology and management of glucocorticoid-induced osteoporosis. Calcif Tissue Int. 2006;79:126-137.
9. Canalis E, Mazziotti G, Giustina A, Bleizikian JP. Glucocorticoid-induced osteoporosis: pathophysiology and therapy. Osteoporosis Int. 2007;18:1319-1328.
10. Shaker JL, Lukert BP. Osteoporosis associated with excess glucocorticoids. Endocrinol Metab. Clin North Am. 2005;34:341-356.
11. Walsh IJ, Wong CA, Pringle M, Tattersfield AE. Use of oral corticosteroids in community and the prevention of secondary osteoporosis: a cross-sectional study. BMJ. 1996;313:344-346.
12. Gudbjornsson B, Juliusson UI, Gudjonsson FV. Prevalence of long-term steroid treatment and the frequency of decision making to prevent steroid-induced osteoporosis in daily clinical practice. Ann Rheum Dis. 2002;61:32-36.
13. Sadat-Ali M, Al-Eq A. Sickle cell disease: is it a cause for secondary osteoporosis? West Afr J Med. 2001;20:134-137.
14. Al-Amri A, Sadat-Ali M. Cancer chemotherapy induced osteoporosis: how common is it among Saudi Arabian cancer survivors. Ind J Cancer. Forthcoming 2009.
15. Morgan GA, Gregio <SPAN>OV</SPAN>. Statistical Package for the Social Sciences for Windows: An Introduction To Use and interpretation in Research [CD-ROM], version 14.0. SPSS Inc. Chicago, Illinois, 2007.
16. Painter SE, Kleerekoper M, Camacho PM. Secondary osteoporosis: a review of the recent evidence. Endocr Pract. 2006;12:338-45.
17. Steinbuch M, Youket TE, Cohen S. Oral glucocorticoid use is associated with an increased risk of fracture. Osteoporosis Int. 2004;15:323-8.
18. Bennis KK, Repp AL, Kleerekoper M. Glucocorticoid-induced osteoporosis. Curr Opin Endocrinol Diabetes Obes. 2007;14:446-450.
19. American College of Rheumatology Ad Hoc Committee on Glucocorticoid-Induced Osteoporosis. Recommendation for the prevention and treatment of glucocorticoid-induced osteoporosis. 2001 Update. Arthritis and Rheum. 2001;44:1496-1503.
20. Adler RA, Hochberg MC. Suggested guidelines for evaluation and treatment of glucocorticoid-induced osteoporosis for the Department of Veterans Affairs. Arch Intern Med. 2002;162:2619-2624.
21. National Osteoporosis Society Guidelines on the prevention and management of corticosteroid-induced osteoporosis, London NOS 1998.
22. Eastell R, Reid DM, Compton J, Cooper C, Fogelman I, Francis RM, et al. A UK consensus group on management of glucocorticoid-induced osteoporosis. An update. J Intern Med. 1998;244:271-92.
23. Sadat-Ali M, Al Elq A. Osteoporosis among male Saudi Arabs. A pilot study. Ann Saudi Med. 2006;26:450-454.
24. Guzman-Clark JR, Fang MA, Sehl ME, Traylor L, Hahn TJ. Barrier in the management of glucocorticoid-induced osteoporosis. Arthritis Rheum. 2007;57:140-146.
25. Hart SR, Green B. Osteoporosis prophylaxis during corticosteroid treatment: failure to prescribe. Postgrad Med J. 2002;78:242-43.
26. Ungprasert S, Wangkaew S, Louthrenoo W. Physicians awareness of the prevention of corticosteroid induced osteoporosis. J Med Assoc Thai. 2007;90:59-64.