INTRODUCTION

Bone is a dynamic organ which provides structural support for the body, protection of vital organs, storage of crucial nutrients, minerals, and lipids and production of blood cells. It is made of a relatively hard composite material that is constantly remodeling to adapt to the daily forces placed upon it, when an imbalance between bone resorption and bone formation happens, osteoporosis occurs. QUS was proposed as an indirect assessment of bone quality, as the modality has the advantages of being portable, with no ionizing radiation used, in addition to comparable cost and time effectiveness in relation to the gold standards for measuring bone density; DXA and Quantitative computed tomography (QCT).1,2

As DXA and QCT provide limited information on bone structure, the QUS stiffness index were suggested to assess dimensional structure and strength.3 The possible clinical applications of QUS include diagnosis of osteoporosis, monitoring of skeletal changes associated with disease prognosis and treatment, and fracture risk assessment.4 Previous studies within the region examined BMD and QUS separately, no studies where done to examine the ability of QUS to evaluate bone density in the young Saudi population.5-7

The aim of this study was to examine whether QUS stiffness index measured at the calcaneus area that can reflect the status of bone at the area most likely to endure fractures (spine and neck of
femur), or whether a combined assessment of QUS and BMD should be used to assess bone loss at different areas.

**METHODS**

**Study Design:** One hundred and one young adult Saudi females attending King Saud University were enrolled in the study (age range 20–24.9 years). The exclusion criteria consisted of pregnancy, diseases or medication affecting bones metabolism, a previous history of fracture, and any terminal illness. Informed consent was obtained from all subjects, and the study was approved by local research ethics committee (CAMS 18/3536). Age, body weight (kg) and height (m) were recorded, body mass index (BMI) (kg/m²) was calculated. Both QUS in the calcaneus area and BMD in the lumbar spine at the area of L2-L4 (Spine) and femoral neck area (N.femur) were obtained for each subject.

**QUS measurements:** A water-bath ultrasound system which generates a band of frequencies from 200 to 600 kHz (Lunar Achilles Insight™ - GE Healthcare) was used for measurements of SI in the calcaneal region of the independent foot. Both SI (automatically calculated from broadband ultrasound attenuation and the speed of sound) and T-score were recorded using a standard protocol supplied by the manufacturer. Data collection was obtained after quality control was carried out using the quality phantom. Precision and reliability of the technique were examined by repeating the measurements on 20 subjects in three consecutive days.

**BMD measurements:** DXA scans (Lunar iDXATM - GE Healthcare) were utilized to measure BMD (g/m²) in the lumbar spine (L2-L4) and the femoral neck area. A standard protocol supplied by the manufacturer was used, which included a quality control test using a standard phantom.

**Statistical Analysis:** The variability within (MSw) and between (MSb) subjects for SI were examined using analysis of variance test (Anova test), precision (expressed as the coefficient of variation), and reliability (expressed as the intraclass correlation coefficient of reliability) for the SI were calculated using Minitab™ software. Using SPSS™ software, Pearson Correlation Coefficient was used to examine the presence of an association between SI, T-score values in the calcaneus and the BMD and T-score values in the lumbar spine and the femoral neck area.

**RESULTS**

Variability between subjects was greater than that within subjects, the measurements of precision and reliability of the Achilles densitometer parameters are tabulated in Table-I. The precision varied from 1.77–1.78, whereas the reliability ranged from 92.2%–98.6%. Age, anthropometric data, QUS and BMD values in the spine and neck of the femur are presented in Table-II.

The association between QUS parameters (stiffness index and T-score calcaneus) and DXA parameters (BMD spine, T-score spine, BMD N.femur and T-score N.femur) were investigated and tabulated in Table-III. Positive correlation were found between stiffness index and BMD spine and BMD N.femur (r= 0.29 & r=0.25 respectively, P < 0.05) and a strong positive correlation between T-score calcaneus and both T-score Spine and T-score N.femur (r = 0.5 & r=0.58 respectively, P < 0.01).

**DISCUSSION**

The presence of portable and cost effective measuring tools for the diagnosis of osteoporosis is essential; QUS presents itself as a non-destructive, non-ionizing, inexpensive technique for the assessment of the mechanical properties of bone, which would contribute to the detection of osteoporosis, and would also allow the formulation of a fracture threshold.8

SI, a parameter derived from speed of sound (SOS) and broadband ultrasound attenuation (BUA), was suggested as a variable to discriminate women with...
low bone density from healthy postmenopausal controls. The ability to use QUS as a screening tool to determine those who might be most beneficial from preventive measures will be of benefit.

The presence of a relationship between stiffness index in the calcaneus and BMD in the spine and N.femur has not been examined, previous studies were conducted to establish reference values for SI and BMD values within the region in separate studies. Whether QUS results alone or a combined assessment of QUS and bone density can be used to predict future fracture risk in females is not clear, further studies are needed to determine if such possibilities can be achieved.

The aim of the study was to measure the precision and reliability of the QUS densitometer, to examine whether QUS parameters could reflect the status of bone at the area most likely to endure fractures, and to investigate the ability to use the technique for screening of osteoporosis and osteopenia.

High precision (≈ 1.77) and reliability (from 95% to 98%) were found in the two parameters measured by QUS, demonstrated by the small coefficient of variation and the variability between subjects being greater than within subjects, which mean that QUS enabled the measurements of significant differences between subjects for both SI and T-score (Table-I).

Mean values of SI in our study demonstrate similar values when compared to measurements obtained previously in the Eastern region of Saudi Arabia, Lebanon, China and Britain, but lower than that of American young females (Table-II). Although ethnic differences which affect bone density is ruled out when data from the Saudi community is compared to data obtained from other countries of the gulf, the reduction in BMD values in the Saudi females could be attributed to many factors including differences in the subjects selection protocol, possible inadequate accrual of bone during childhood and adolescence, lack of exercise and sedentary lifestyle with few outdoor activities, which may result in the possibility of an increase in the incidence of osteoporosis later in life. Such findings invite more research on the endogenic and exogenic factors affecting peak bone mass in the Saudi population.

A positive correlation was found between both QUS and DXA parameters with BMI attributed to the effect of weight only (results published previously). Similar results were reported previously, as increasing weight is known to be associated with higher bone density. Such findings in the Saudi population suggest the importance of considering body weight in the evaluation of patients in relation to the diagnosis of osteoporosis.

The ability to examine the presence of a relationship between stiffness index in the calcaneus and BMD in the spine and femur is needed. Nevertheless, when QUS and BMD were measured along a single direction and at the same previous location, results were highly correlated in both in-vitro and in-vivo studies. Such correlation is not known in the literature, where measurements are obtained from different sites, as previous studies conducted to establish reference values for SI and BMD values were done separately. The presence of a positive relationship between SI in the calcaneus area and BMD both in the spine and femur region suggests that the mechanisms that effect bone density affects as well bone structure. This mechanism seems to vary at different body site hence the association being positive but not relatively strong (r = 0.29 & 0.25 respectively). These findings could be attributed to differences in bone structure at different sites in

| Table-III: Pearson correlation coefficient of QUS and DXA parameters. |
|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|
|                 | SI              | T-score * calcaneus | BMD * Spine     | T-score * Spine | BMD * N.femur   | T-score * N.femur |
| T-score Calcaneus | 0.79**          |                  |                  |                  |                  |                  |
| BMD Spine        | 0.29**          | 0.29**           |                  |                  |                  |                  |
| T-score Spine    | 0.49**          | 0.5**            | 0.78**           |                  |                  |                  |
| BMD * N.femur    | 0.25*           | 0.32**           | 0.47**           | 0.55**           |                  |                  |
| T-score * N.femur| 0.5**           | 0.58**           | 0.63**           | 0.77**           | 0.63**           |                  |

*, Correlation is significant at the 0.05 level.  
** Correlation is significant at the 0.01 level.
addition to other factors. Previous in-vitro studies found that ultrasound properties are dependent on the microstructure of the bone and the trabecular orientation with the QUS being affected by the microstructure of the bone more than bone mass, which resulted in the QUS parameter unlike BMD exhibiting anisotropic effect dependent on the direction of propagation.

**Limitations of this study** include the inability to extrapolate our results to other ultrasound and DXA machines, middle-aged, pre and post-menopausal females were not included in the study, these areas should be investigated in the future. In conclusion this study has found that the QUS is a reliable technique to be used with DXA for the investigation of osteopenia and osteoporosis.

**CONCLUSIONS**

Nevertheless, results demonstrate that the association between T-score in the calcaneus and T-score in the spine and femur are stronger \( r=0.57 & 0.59 \) respectively. Such findings suggest that during the screening, it is possible to use T-score data obtained from QUS for referral to BMD measurements test, although QUS parameters on its own should not be used as a screening tool, or for the assessment of fracture risk.

**ACKNOWLEDGMENT**

This research project was supported by a grant from the "Research Center of the Female Scientific and Medical Colleges", Deanship of Scientific Research, King Saud University.

**Declaration of interest:** The Author reports no conflicts of interest and is responsible for the content and writing of the paper.

**REFERENCES**

1. Gregg EW, Kriska AM, Salamone LM, Roberts MM, Anderson SJ, Ferrell RE, et al. The epidemiology of quantitative ultrasound: a review of the relationships with bone mass, osteoporosis and fracture risk. Osteoporosis Int. 1997;7(2):89-99.
2. Njeh CF, Boivin CM, Langton CM. The role of ultrasound in the assessment of osteoporosis: a review. Osteoporosis Int. 1997;7(1):7-22.
3. Gluer CC. Quantitative ultrasound techniques for the assessment of osteoporosis: expert agreement on current status. J Bone Mineral Res. 1997;12(8):1280-1288. DOI: 10.1007/bf01659128
4. Hans D, Arlot ME, Schott AM, Roux JP, Meunier PJ, editors. Histochemical and biochemical methods of bone analysis. New York: Plenum Publishing Corporation; 1983. p. 83-92.
5. Ardawi MSM, Maimany AA, Bahish TM, Nasrat HAN, Milaat WA, Al-Raddadi RM. Bone mineral density of the spine and femur in healthy Saudis. Osteoporosis Int. 2005;16(1):43-55. DOI: 10.1007/s00198-004-1639-9
6. Hammad LF. Quantitative ultrasound measurements of stiffness index in young adult females. Clin Pract. 2013;5(1). DOI: 10.3823/107
7. Hammad LF. Bone mineral density in University aged Saudi females. Pak J Med Sci. 2015;32(1):556-560. DOI: 10.12669/pjms.313.7264
8. Langton CM, Palmer SB, Porter RW. The measurement of broadband ultrasonic attenuation in cancellous bone. Engineering Med. 1984;13(2):89-91. DOI: 10.1243/EMED.1984.016.02.02
9. Hadji P, Hars O, Wuster C, Bock K, Alberts US, Bohnet HG, et al. Stiffness index identifies patients with osteoporotic fractures better than ultrasound velocity or attenuation alone. Maturitas. 1999;31(3):221-226. DOI: 10.1016/s0378-5122(99)00001-1
10. Yoshimi I, Aoyagi K, Okano K, Yahata Y, Kusano Y, Moji K, et al. Stiffness index of the calcaneus measured by quantitative ultrasound and menopause among Japanese women: The Hizen-Oshima Study. Tohoku J Experimental Med. 2001;195(2):93-99. DOI: 10.1620/tjem.195.93
11. Sadat-All M, Al-Elq A, Al-Habdan I, Al-Mohanna FA, Al-Mulhim AA. Quantitative ultrasound (QUS) of the os calcis in Saudis women: defining Saudi reference value for the diagnosis of low bone mass. Arch Osteoporosis. 2010;5(1-2):139-144. DOI: 10.1007/s11657-010-0047-3
12. Barrett-Connor E, Siris ES, Wehren LE, Miller PD, Abbott TA, Berger ML, et al. Osteoporosis and fracture risk in women of different ethnic groups. J Bone Mineral Res. 2005;20(2):185-194. DOI: 10.1359/JBMR.041007
13. El-Dessouki MI. Osteoporosis in postmenopausal Saudi women using dual x-ray bone densitometry. Saudi Med J. 2003;24(9):953-956.
14. Kang C, Speller R. Comparison of ultrasound and dual energy X-ray absorptiometry measurements in the calcaneus. Br J Radiol. 1998;71(848):861-867.
15. Liu W, Xu CL, Zhu ZQ, Wang W, Han SM, Zu SY, et al. Characteristics of calcaneal quantitative ultrasound normative data in Chinese mainland men and women. Osteoporosis Int. 2006;17(8):1216-1224. DOI: 10.1007/s00198-006-0816-6
16. Wehbe J, Cortibaou C, Chidiac RM, Nehme A, Melki R, Bedran F, et al. Age-associated changes in Quantitative Ultrasoundometry (QUS) of the os calcis in Lebanese women-assessment of a Lebanese reference population. J Musculoskeletal Neuronal Interactions. 2003;3(3):232-239.
17. Dougherty G, Al-Marzouk N. Bone density measured by dual-energy X-ray absorptiometry in healthy Kuwaiti women. Calcified Tissue Int. 2001;68(4):225-229. DOI: 10.1007/s002230010315
18. Hammadouche M, Al-Khayarin M, Zirie M, Bener A. Bone density measured by dual energy X-ray absorptiometry in Qatari women. Maturitas. 2005;52(3):319-327. DOI: 10.1016/j.maturitas.2005.05.011
19. El Maghraoui A, Guerboub AA, Achemlal L, Nouijai AA, Ghazi M, et al. Bone mineral density of the spine and femur in healthy Moroccan women. J Clin Densitometry. 2006;9(4):454-460. DOI: 10.18563/jocld.2006.07.001
20. Maalouf G, Salem S, Sandid M, Attallah P, Eid J, Saliba N, et al. Bone mineral density of the Lebanese reference population. Osteoporosis Int. 2000;11(9):756-764.
21. Kudlacek S, Schneider B, Peterlik M, Leb G, Klaushofer K, Weber K, et al. Normative data of bone mineral density in an unselected adult Austrian population. Euro J Clin Invest. 2003;33(4):332-339. DOI: 10.1046/j.1365-2362.2003.01128.x
22. McKelvie ML, Fordham J, Clifford C, Palmer SB. In vitro comparison of quantitative computed tomography and broadband ultrasonic attenuation of trabecular bone. Bone. 1989;10(2):101-104. DOI: 10.1016/8756-3282(89)90005-7
23. Waud CE, Lew R, Baran DT. The relationship between ultrasound and densitometric measurements of bone mass at the calcaneus in women. Calcified Tissue Int. 1992;51(6):415-418.
24. Faulkner KG, McClung MR, Coleman LJ, Kingston-Sandah E. Quantitative ultrasound of the heel: correlation with densitometric measurements at different skeletal sites. Osteoporosis Int. 1994;4(1):42-47.
25. Gluer CC, Wu CY, Genant HK. Broadband ultrasound attenuation signals depend on trabecular orientation: an in vitro study. Osteoporosis Int. 1993;3(4):185-191.