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Role of lumbar spine signal intensity measurement by MRI in the diagnosis of osteoporosis in post-menopausal women



Maha M. Saad¹, Ahmed Tohamy Ahmed^{1*}, Khaled E. Mohamed² and Mohamed R. Habba¹

Abstract

Background: Osteoporosis is a worldwide health problem and a common cause of bone fractures; the most common type of osteoporosis is post-menopausal type. MRI has a role in the diagnosis of osteoporosis and can be used as a screening tool, so the purpose of our study was to define a quantitative MRI-based score (M-score) for the detection of lumbar spine osteoporosis and to evaluate the correlation between lumbar spine signal intensity measured by MRI and BMD (bone mineral density) in post-menopausal women. Our case-control study involved 100 cases (50 old post-menopausal females as a case group and 50 healthy females as a control group of matched age). Both groups were subjected to history taking, dual-energy X-ray absorptiometry (DEXA), and conventional lumbar MRI. DEXA was performed for the lumbar spine and all scores (T-score, Z-score, BMD) were calculated. Lumbar MRI was performed (sagittal T1WI and T2WI) from L1–L4 levels. SNR_{L1–L4} and M-score were calculated from T1W images.

Results: All DEXA scores were significantly lower in post-menopausal females compared to the control group (P < 0.0001). Meanwhile, SNR_{L1-L4} and M-score were significantly higher among cases than controls (P < 0.0001). The diagnostic threshold of SNR_{L1-L4} and M-score for distinguishing osteoporotic from non-osteoporotic females was 104.5 for SNR_{L1-L4} with a sensitivity of 94%, specificity 60%, positive predictive values (PPV) 31%, and negative predictive values (NPV) 98%, and 3.5 for M-score with a sensitivity of 93.3%, specificity 83.5%, PPV 50%, and NPV 98.6%. SNR_{L1-L4} was negatively correlated with M-score (r = -0.74, P < 0.0001), T-score (r = -0.42, P < 0.0001), Z-score (r = -0.32, P < 0.0001), and BMD (r = -0.31, P < 0.0001). M-score was negatively correlated with T-score (r = -0.48, P < 0.0001), Z-score (r = -0.36, P < 0.0001), and BMD (r = -0.37, P < 0.0001). M-score moderately agreed with T-score (r = -0.5, P < 0.0001).

Conclusion: The MRI-based score (M-score) of the lumbar spine is an accurate quantitative method for distinguishing osteoporotic from non-osteoporotic females. M-score and SNR_{L1-L4} are significantly correlated with osteoporotic indices (T-score and BMD).

Keywords: Osteoporosis, Lumbar spine, Magnetic resonance imaging, Dual-energy X-ray absorptiometry

Background

Osteoporosis is a worldwide health problem and is considered as the most common bone disease. It is proved that it is the main cause of bone fractures in elderly population. It is defined as a decreased mass per unit volume of normally mineralized bone [1, 2].

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The most common type of osteoporosis is postmenopausal type (estrogen-deficient osteoporosis). About 15% of white people in their 50s and 70% of those over 80 are affected [3]. Women are affected more than men [1]. In the developed world, depending on the method of diagnosis, 2–8% of males and 9–38% of females are affected [4, 5].

Dual-energy X-ray absorptiometry (DEXA) is considered as the gold standard for the diagnosis of osteoporosis as it uses negligible radiation dose and easy to use quantitative imaging modality that has high precision



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[6]. According to the World Health Organization guidelines, T-score helps to classify bone mineral density (BMD) [7]. Other methods rather than DEXA as quantitative computed tomography and quantitative ultrasound also have a role [8].

MRI has a role in detecting osteoporosis that the appearance of bone marrow is determined by its relative amount of protein, fat, water, and cells on MRI pulse sequence. The age-related changes of bone marrow revealed decreased cellularity and increased fat content, so this fact can be used as helpful information to detect the role of MRI in the diagnosis of osteoporosis in association with DEXA scan. T1-weighted spin-echo images are the best to evaluate the cellular content of the bone marrow [9].

So the purpose of our study is to assess the utility of quantitative MRI-based score (M-score) to detect osteoporosis in correlation with BMD measured by DEXA.

Methods

This is a case-control single-center study approved by the local institutional ethics committee; written informed consent was obtained from all patients.

Study population

Our case-control study was done over a period of time of 1 year from June 2016 to June 2017 and involved 50 patients and 50 normal volunteers who underwent lumbar MRI. *Inclusion criteria* included post-menopausal females > 50 years who had undergone lumbar MRI for low back pain, and the reference control group of healthy females of matched age with normal body mass index (BMI = 19-25 kg/m²) to calculate the MRI-based score equivalent to the T-score used in DEXA. Exclusion criteria were (a) MRI of the lumbar spine with contrast, (b) known patients with neoplastic disease, (c) traumatic dorso-lumbar vertebral injuries, (d) time elapsed between MRI examination and DEXA longer than 6 months, (e) absolute contraindication to MR imaging (claustrophobia, implanted defibrillator or pacemaker, cochlear implant, some types of clips used on brain aneurysms), and (f) patients refusing participation in the study.

All patients were subjected to history taking (age, menstrual, medical), DEXA scan, and conventional MRI assessment of the lumbar spine. Then the calculated quantitative MRI-based score (M-score) to detect osteoporosis was done and correlated with BMD measured by DEXA.

DEXA scan

DEXA scan for the lumbar spine was done for all participants by using DEXA GE model (DPx-pro). BMD, Z-score, and T-score were calculated according to International Society for Clinical Densitometry (ISCD) guidelines (a minimum of two consecutive levels are used for lumbar T-score evaluation, after exclusion of vertebrae

with focal structural abnormalities, degenerative changes, tubing artifact). The calculated values for the post-menopausal women (> 50 years) are compared with those of the healthy control group reference population values. T-scores are complicated statistical scores: T-score > 1.0 SD (normal), between -1.0 and -2.5 SD (osteopenia), ≤ -2.5 SD (osteopenis), and ≤ -2.5 SD with 1 or more fragility fractures (severe osteoporosis).

Magnetic resonance imaging methods and image analysis

All patients underwent lumbar MRI by using 1.5 Tesla MR scanner (Philips Medical Systems, Achieva) and post-processing was done by using a dedicated Philips workstation (Extended workspace, version 2.6).

Patient's preparation

The ferromagnetic materials were taken off. The procedure was explained to the patient, and then the patient was asked to lie supine and instructed not to move during the study.

MRI protocol and technique

Routine lumbar spine assessment from L1–L4 was performed usually by obtaining sagittal T1W spin-echo sequence (TR = 7, TE = 400–600, slice thickness = 4 mm, fov = 280 mm, matrix = 320×320), the most useful sequence for the assessment of bone marrow. Region of interest (ROI) was manually placed as a circle in the vertebral body excluding the cortical bone, subchondral abnormalities, focal lesions (e.g., hemangioma), and posterior venous plexus. Three ROIs were used for each vertebra, and each of them acquired on a different slice (with their mean used for analysis; that signal-to-noise ratio is not equivalent within vertebrae). Another ROI was placed in an artifact-free site outside the patient to measure the noise.

MRI image analysis

All MR images were reviewed by a consensus of two experienced radiologists blinded to the clinical information and DEXA scan results. Images were transferred to an offline workstation for post-processing. Signal-to-noise ratio (SNR) was obtained by dividing the intra-vertebral intensity by the standard deviation of the noise.

The diagnostic performance of SNR_{L1-L4} was estimated for each patient and used to obtain M-score (MRI-based score) for the diagnosis of osteoporosis. The SNR_{L1-L4} of the control group was also used in the equation by using their mean (SNR_{ref}) and standard deviation (SD_{ref}) [10].

The M-score was defined according to the formula as follows:

$$M\text{-score} = \frac{\text{SNR}_{\text{L1-L4}} - \text{SNR}_{\text{ref}}}{\text{SD}_{\text{ref}}}$$

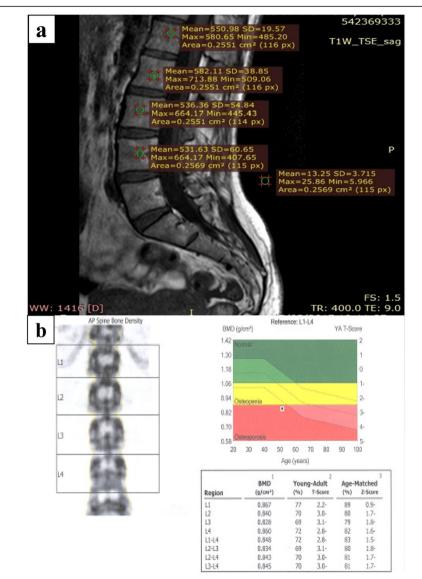


Fig. 1 Lumbar MRI of a 55-year-old post-menopausal woman complained from low back pain, no history of any chronic illness, spinal trauma, family history of osteoporosis, history of taking any medications, or taking calcium and Vit. D supplementation. MRI of lumbar spine T1W (L1–L4) revealed that the $SNR_{L1-L4} = 148.5$ and M-score within abnormal values = 4.1093, DEXA scan of the lumbar spine, PA (L1–L4) revealed osteoporosis according to T-score results. The mean BMD of the lumbar spine is 0.847 g/cm^2 , equivalent to a T score of -2.8

Statistical analysis

Statistical analysis and tests were specified according to the variable type. A commercially available PC-based software package (SPSS) was used.

Results were compared with the final diagnosis and the predictive values of MRI sequences were calculated by obtaining positive predictive values (PPV), negative predictive values (NPV), sensitivity, specificity, and total accuracy in the detection of osteoporosis.

Receiver operating curve (ROC) analysis was used to define the cutoff values of M-score and ${\rm SNR_{L1-L4}}$ for the discrimination of osteoporotic from non-osteoporotic females.

Chi-squared test was used for analyzing differences between the results of MRI (calculated M-score) and other imaging modalities (DEXA scan). The sensitivity and specificity of MRI and other imaging modalities were calculated and compared.

Results

The study involved 50 post-menopausal women with a mean age of 59.4 + 4.1 years (age range 46-68 years) and 50 healthy females of matched ages. BMI was significantly higher among post-menopausal women in comparison to healthy controls (33.8 kg/m^2 versus 22.01 kg/m^2 , respectively; P value < 0.0001).

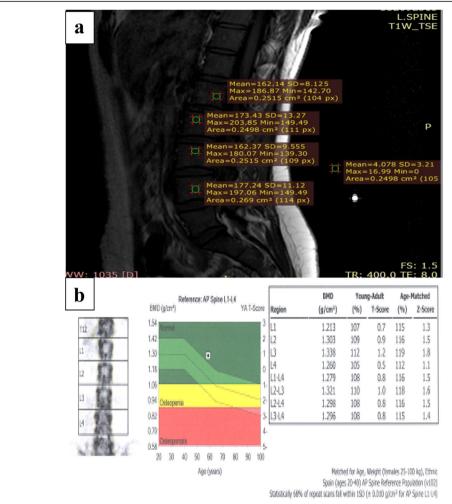


Fig. 2 Lumbar MRI of a 58-year-old post-menopausal woman with a history of low back pain. No history of any chronic illness, spinal trauma, family history of osteoporosis, history of taking any medications, or taking calcium and vit. D supplementation. MRI of lumbar spine T1W (L1–L4) revealed that the $SNR_{L1-L4} = 53.93$, and M-score = 0.045 within normal values, DEXA scan of the lumbar spine, and PA (L1–L4) revealed normal T-score results. The mean BMD of the lumbar spine is 1.27 g/cm², equivalent to a T-score of 0.8

All DEXA scores including T-score, Z-score, and BMD were significantly lower in post-menopausal females compared to the control group (P value < 0.0001). Meanwhile, SNR_{L1-L4} and M-score were significantly higher among cases than controls (P value < 0.0001) (Figs. 1 and 2). According to WHO criteria, of the case group, it was evoked that 11 patients (22%) had a normal T-score, 24 patients (48%) had osteopenia, and 15 patients (30%) had osteoporosis. Obesity was representing the most co-morbid condition related to osteoporosis that was present in the majority of patients (82%).

The ROC curve assessment of SNR_{L1-L4} for the discrimination of osteoporotic from non-osteoporotic females revealed that AUC = 0.696 at 95% confidence interval 0.636–0.981 and the diagnostic capability of using a threshold of 104.5 as a cutoff had a sensitivity of 94%, specificity 60%, PPV 30.6%, and NPV 98% (P value = 0.0043; Fig. 3).

The ROC curve assessment of M-score for the discrimination of osteoporotic from non-osteoporotic females revealed that AUC = 0.904 at 95% confidence interval 0.636–0.981 and the diagnostic capability of using a threshold of 3.5 as a cutoff had a sensitivity of 93.3%, specificity 83.5%, PPV 50%, and NPV 98.6%; *P* value < 0.0001 (Fig. 4). Meanwhile, the ROC curve assessment of M-score for the discrimination of postmenopausal from healthy control females revealed that AUC = 0.975 at 95% confidence interval 0.636–0.981 and the diagnostic capability of using a threshold of 2.2 as a cutoff had a sensitivity of 92%, specificity 98%, PPV 97.9%, and NPV 92.5% (*P* value < 0.0001; Fig. 5).

The study showed that SNR_{L1-L4} was negatively correlated with M-score (r = -0.74, P value < 0.0001), T-score (r = -0.42, P value < 0.0001), Z-score (r = -0.32, P value < 0.0001), and BMD (r = -0.31, P value < 0.0001) (Table 1).

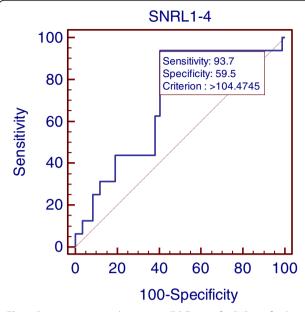


Fig. 3 Receiver operating characteristic (ROC) curve for SNR_{L1-L4} for the discrimination of osteoporotic from non-osteoporotic females revealed that by using a threshold of 104.5 as a cutoff value had a sensitivity of 94%, specificity 60%, PPV 30.6%, and NPV 98% at a *P* value = 0.0043

The ROC curve assessment of M-score for the discrimination of normal from low BMD revealed that AUC = 0.696 at 95% confidence interval 0.636-0.981 and the diagnostic capability of using a threshold of 2.8 as a cutoff had a sensitivity of 53.8%, specificity 88.6%, PPV 89.7%, and NPV 50.8% (P value = 0.0007; Fig. 6).

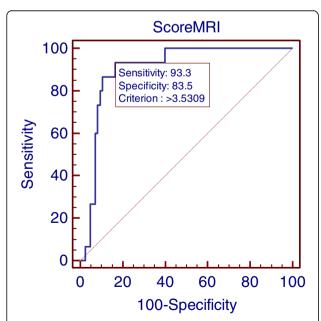


Fig. 4 Receiver operating characteristic (ROC) curve for M-score for the discrimination of osteoporotic from non-osteoporotic females revealed that by using a threshold of 3.5 as a cutoff value had a sensitivity of 93.3%, specificity 83.5%, PPV 50%, and NPV 98.6% at a *P* value < 0.0001

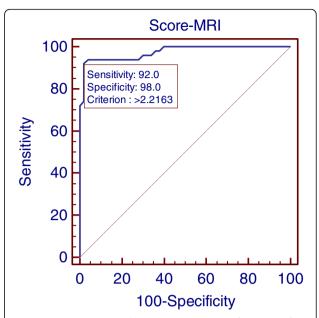


Fig. 5 Receiver operating characteristic (ROC) curve for M-score for the discrimination of post-menopausal from healthy females revealed that by using a threshold of 2.2 as cutoff value had a sensitivity of 92% and a specificity of 98% at a *P* value of < 0.0001

The study revealed that M-score was negatively correlated with SNR_{L1-L4} (r=-0.74, P value < 0.0001), T-score (r=-0.48, P value < 0.0001), Z-score (r=-0.36, P value < 0.0001), and BMD (r=-0.37, P value < 0.0001) (Table 2).

Discussion

Several studies had used CT modality for evaluating osteoporosis and had been performed in the last decade which demonstrated how routine abdominal CT obtained for other clinical indications could be used for quantitative diagnosis of osteoporosis [11], and also, demonstrated the possibility of accurately detecting osteoporosis-related fragility fractures by CT [12, 13]. As there are more than 80 million CT examinations performed each year in the USA, the idea of extracting

Table 1 Correlations between signal-to-noise ratio (SNR)_{L1-L4} and radiological indices using magnetic resonance imaging (MRI) and dual-energy X-ray absorptiometry (DEXA)

	SNR _{L1-L4}	
	Pearson's correlation coefficient (r)	*P value (two-tailed)
M-score	- 0.743	< 0.0001**
T-score	- 0.418	< 0.0001**
Z-score	- 0.315	0.001**
BMD	-0.312	0.002**

^{*}Correlation was significant at p-value < 0.05

^{**}Correlation was highly significant at P value < 0.01

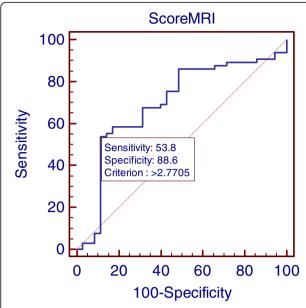


Fig. 6 Receiver operating characteristic (ROC) curve for M-score for the discrimination of normal from low BMD revealed that by using a threshold of 2.8 as a cutoff value had a sensitivity of 53.8% and a specificity of 88.6% at a P value = 0.0007

more information from imaging data collected for other purposes holds merit [14].

Regarding MRI, previous studies demonstrated that the bone marrow signal intensity on the standard T1-weighted images is negatively correlated with BMD and osteoporosis as shown by Shen et al .[15, 16] and Shah and Hanrahan [9]. This correlation may open a window for the possibility to detect osteoporosis using MRI. Moreover, osteoporosis-related fragility fractures may be accurately detected by MRI as shown by Bazzocchi et al. [12]. Only one study by Bandirali et al. [10] had previously aimed to define a new quantitative lumbar spine MRI-based method on standard T1-weighted images for diagnosing osteoporosis.

The MRI-based score (M-score) can be calculated easily that lumbar spine MRI is the routine and commonest examination for low back pain assessment. From the aspect of being a common examination for low back pain,

Table 2 Correlations between M-score and radiological indices using magnetic resonance imaging (MRI) and dual-energy X-ray absorptiometry (DEXA)

	M-score	
	Pearson's correlation coefficient (r)	*P value (two-tailed)
SNR _{L1-L4}	- 0.743	< 0.0001**
T-score	- 0.484	< 0.0001**
Z-score	- 0.356	< 0.0001**
BMD	- 0.368	< 0.0001**

^{*}Correlation was significant at P value < 0.05

so it can be used as a screening tool for the detection or suspicion of osteoporosis in patients.

We agreed with a study done by Bandirali et al. [10] in the main concept and high sensitivity and specificity of M-score with its high utility as a screening tool. Our study revealed sensitivity and specificity values around 93% and 84%, respectively, of M-score correlated with DEXA examination as a reference gold standard. Based on our sample which involved 100 subjects, it was found that SNR_{L1-L4} was negatively correlated with BMD (r = -0.3, P < 0.0001), compared to (r = -0.67, P < 0.001) in Bandirali et al.'s study.

It was found in our study that the threshold of Mscore was 3.5 with a sensitivity of 93.3%, specificity 83.5%, PPV 50%, and NPV 98.6%, and this means that for every 100 patients with M-score ≥ 3.5, 93 patients will have a true positive rate or really identified as to have osteoporosis. This also means that for every 100 patients with M-score < 3.5, 84 subjects will have a true negative rate or correctly identified as not having osteoporosis. In Bandirali et al.'s study, the demonstrated threshold of M-score by them was 5.5 with a sensitivity of 54%. The difference in the threshold of M-score could be due to the sample size and sample type, as our sample intently included post-menopausal women which are at high risk of development of osteopenia or osteoporosis due to hormonal changes and obesity as a common co-morbid finding as described below. In obese patients and patients with spinal degenerative disease, lumbar BMD may be increased and gives the false impression that these patients have a low risk of fractures [17, 18].

But with DEXA scan, the diagnosis of osteoporosis depends on T-score and not BMD which gives more easy assessment away from calibration difficulties in the case of using BMD [6]. So M-score measured by MRI was used as a predictor of osteoporosis and a parameter to be correlated with T-score calculated by DEXA with good diagnostic performance.

Our study has several strengths that may be deserved to be mentioned. First, MRI and DEXA were simultaneously performed in two-thirds of the patients (66%). In the rest of the patients (34%), the median time elapsed between the two examinations was low (about 33 days) and it is unlikely to have influenced the correlation between DEXA and MRI.

Second, regarding the reference group of a healthy woman undergoing MRI, bone densitometry data were also available because they undergo DEXA as well. Moreover, we excluded the healthy females with any abnormalities considering that they must have healthy bone status.

Third, we excluded post-menopausal women with vertebral osteoporotic fractures. That on DEXA, patients with low-trauma fractures have non-reliable non-osteoporotic BMD.

^{**}Correlation was highly significant at P value < 0.0001

This study had also some limitations; firstly, the use of M-score threshold values will be different for each single-center according to different SNRs values provided by each MRI system. Secondly, the signal-to-noise ratio assessment by the placement of the region of interest (ROI) placed within the vertebral body excluding the cortical bone, subchondral abnormalities, focal lesions (e.g., hemangiomas), and posterior venous plexus and outside of the patient are simple methods of measurement as compared to spectroscopic measurement [19, 20]. This study is specific for post-menopausal women; however, other multicenter studies are mandatory for both males and females as also males > 50 years have a higher risk for osteoporosis.

Conclusion

In conclusion, we recommended to include the measurement of M-score and SNR_{L1-L4} as part of the routine evaluation of the lumbar spine, that it is considered an easy, reliable, and reproducible screening tool for osteoporosis without any additional imaging, radiation exposure, especially in post-menopausal women or patients at risk for osteoporosis that will help to early detect persons who need medical advice, prevent the disease and improve patient's outcome.

Abbreviations

BMD: Bone mineral density; DEXA: Dual-energy X-ray absorptiometry

Acknowledgements

Not applicable

Ethical approval and consent to participate

Approved by the local institutional ethics committee (Committee of Scientific Research Ethics (CSRE), Suez Canal University, Egypt); written informed consent was obtained from all patients.

Availability of data and materials

The dataset used and/or analyzed during the current study are available from the corresponding author on reasonable request.

Authors contributions

MM helped in the preparation of the study methodology, data collection, and analysis of data. AT participated in the formulation of the study, preparation of methodology, data collection, analysis and revision of the collected data, and writing the paper. KE participated in the supervision of the study methodology and collected the data. MR participated in the revision of the study methodology and collected the data. All authors have read and approved the final manuscript.

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Consent for publication

Consent for publication was obtained from the patients.

Competing interests

The authors declare that they have no competing interests.

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