### Diffusion Weighted Image MRI in Assessment of Patients with Multiple Myeloma

### Thesis

Submitted for Partial Fulfilment of Master Degree in Radio-diagnosis

Presented by

Arwa Adel Mahmoud Hendy

M.B.B.Ch, Ain Shams University

### Under supervision of Dr. Reem Hassan Basiouny

Assistant Professor of Radiodiology Faculty of Medicine Ain Shams University

### Dr. Rasha Tolba Khattab

Lecturer of Radiodiology Faculty of Medicine Ain Shams University

Faculty of Medicine Ain Shams University 2018



سورة البقرة الآية: ٣٢

# Acknowledgment

First and foremost, I feel always indebted to AUAH, the Most Kind and Most Merciful.

I'd like to express my respectful thanks and profound gratitude to **Dr. Reem Hassan**Basiouny, Assistant Professor of Radiodiology - Faculty of Medicine- Ain Shams University for his keen guidance, kind supervision, valuable advice and continuous encouragement, which made possible the completion of this work.

I am also delighted to express my deepest gratitude and thanks to **Dr. Rasha Tolba Khattab**, Lecturer of Radiodiology, Faculty of Medicine, Ain Shams University, for his kind care, continuous supervision, valuable instructions, constant help and great assistance throughout this work.

I would like to express my hearty thanks to all my family for their support till this work was completed.

Last but not least my sincere thanks and appreciation to all patients participated in this study.

Arwa Adel Mahmoud Hendy

# List of Contents

Title	Page No.
List of Tables	i
List of Figures	ii
List of Abbreviations	v
Introduction	1
Aim of the Work	3
Review of Literature	
Anatomy of The Spine & bone marrow	4
MR Imaging of the Normal Bone Marrow	13
Pathology of Multiple Myeloma	22
MRI Imaging of Bone Marrow In Multiple Myelo	ma29
Diffusion Weighted Imaging Principles	38
The Application of DWI in Multiple Myeloma	47
Patients and Methods	54
Results	61
Case presentation	68
Discussion	76
Summary	85
Conclusion	86
References	87
Arabic Summary	

# List of Tables

Table No.	Title	Page No.
Table (1):	Comparison between the studied regarding ADC value	· -
<b>Table (2):</b>	Comparison between the subt	v <b>-</b>

# List of Figures

Fig. No.	Title	Page No.
Fig. (1-1):	Vertebral body	5
Fig. (1-2):	A: Sagittal CT MIP of the lumbar sp	
	demonstrating the vertebral body	
	pedicle (P), superior articular prod	
	(SAP), inferior articular process (LA	
	and zygapophyseal joint (Z)	
Fig. (1-3):	Intervertebral disc	
Fig. (1-4):	Ligaments	
Fig. (1-5):	Sagittal T2 fat-saturated (A) and axial	
	weighted (B) MRIs of the lumbar sp	
	demonstrating high signal intensity in	
	nucleus pulposus (NP) and inner annu	
	fibro-sis and decreased signal in the or	
<b>T</b> ! (1.0)	annulus fibrosis (AF)	
Fig. (1-6):	Spines	
Fig. (2-1):	Normal bone marrow in an 8-year	
F! (0.0)	child	
Fig. (2-2):	Normal appearance of spinal b	
F! (0.0)	marrow in a 45-year-old woman	
Fig. (2-3):	Bone marrow conversion and re-convers	
Fig. (2-4):	Normal appearance of adult bone mar	
T' (0.5)	on MRI	
Fig. (2-5):	Sagittal in-phase (A) and out-of-phase	
E' - (9.1)	T1-weighted images	
Fig. (3-1):	Normal immunoglobulin molec	
	containing paired heavy chains with	
Fig (9.9).	smaller light chain attached to each	
Fig. (3-2):	Origins of plasma cells	
Fig. (3-3):	· ·	
Fig. (4-1):	Appearance of MM lesions	
Fig. (4-2):	Bone marrow infiltration patterns	34

# List of Figures (Cont...)

Fig. No.	Title	Page No.
Fig. (4-3): Fig. (4-4):	suppressed T2-weighted (right) image the spine displaying a diffuse be marrow infiltration of the cervithoracic, lumbar and sacral spine velow signal intensity on T1-intermediate to high signal intensity fat-suppressed T2-weighted images	fat- s of one cal, with and on
	T2-weighted images of the sacrum with new focal lesion in the vertebral body S2, hypointense on T1- and hyperinter on T2- weighted images with a different hypointense signal on a T1-weight image of the surrounding bone marrow	y of ense fuse ated
Fig. (5-1):	Diffusion of water molecules	
Fig. (5-2):	a)DWI (b value = $500 \text{ sec/mm}^2$ ) sh	ows
	liver metastasis (arrow), renal cyst	
	gallbladder all of which exhibit h	O
	signal intensity. (b) Corresponding A	
	map, the metastasis shows low diffusive reflected by the low ADC, while the	-
	and gallbladder show high ADC, beca	•
	of the long T2 relaxation time of the fl	
	9	ine-
	through)	42
Fig. (6-1):	Example of sequence images belongs	
	patient with multiple myeloma	
Fig. (6-2):	Another Example of diffusion-weighted	
	value images b0, b200, b400, b600,	
	b1,000 in a patient with mult	_
	myeloma	_

# List of Figures (Cont...)

Fig. No.	Title	Page No.
Fig. (6-3):	Diffusion-weighted b0 and b1,000 im with a region-of-interest in the vertex body of T10 and L3 for the calculation the apparent diffusion coefficient (Al in this patient with multiple myeld and typical focal hyperintense lesi with a slightly diffusely increased significant.	oral n of DC) oma ons
Fig. (6-4):	intensity of the entire vertebral body  Diffusion-weighted b1,000 images o patient with monoclonal gammopathy undetermined significance demonstrat hypointense signal intensity, indicat	f a y of ing
Fig. (8-1):	normal fatty marrow Pie chart demonstrating the gender of	
<b>8</b> (- )	study group	
Fig. (8-2):	Comparison between the studied groregarding ADC value	_
Fig. (8-3):	Comparison between the subtypes multiple myloma regarding ADC value	of
Fig. (8-4):	Receiver operating characteristic cu (ROC) for ADC value in differentiat between focal group and normal in fo	rve cion
Fig. (8-5):	Receiver operating characteristic cu (ROC) for ADC value in differentiat between diffuse group and normal I group	rve cion MM

# List of Abbreviations

Abb.	Full term
ADC	A , 1:00 : 00 : ,
ADC	Apparent diffusion coefficient
DCE	Dynamic contrast-enhanced
DWI	Diffusion weighted imaging
Fs	Fat-suppressed / fat-saturated
<i>MDCT</i>	$ Multidetector\ computed\ tomography$
MGUS	Monoclonal gammopathy of undetermined significance
<i>MM</i>	$ Multiple\ myeloma$
M-protein	$ Monoclonal\ protein$
<i>MRI</i>	Magnetic resonance imaging
SE	Spin-echo
<i>SMM</i>	Smouldering multiple myeloma
T1-weighted	T1-weighted spin-echo
T2-weighted	T2-weighted spin-echo

### INTRODUCTION

(MM) is a hematologic malignancy characterized by abnormal plasma cells in the bone marrow and/or in extramedullary sites, urinary and/or serum monoclonal immunoglobulin, and osteolytic lesions in most patients. The Durie and Salmon staging system for MM, which was introduced in 1975, is still used to assess tumor burden and prognosis. The plain radiographic skeletal survey is an important part of this system, since multiple osteolytic lesions define stage III disease (Palumbo and Anderson, 2011; *Rajkumar et al.*, 2014).

The prognostic implications of magnetic resonance (MR) imaging of the bone marrow for MM have already been established for abnormal versus normal MR imaging results and for individual MR imaging patterns of involvement *Song* et al., 2014) For example, abnormal spinal MR imaging results have been shown to help identify patients with asymptomatic (smoldering) myeloma who are likely to benefit from early treatment (Moulopoulos et al., 1995).

More recently, it was shown that patients with smoldering myeloma and more than one unequivocal focal lesion on whole-body MR images have an increased risk of developing myelomarelated symptoms and should receive treatment (Fechtner et al., 2010; Kastritis et al., 2014). Accordingly, the most recent International Myeloma Working



Group criteria for MM incorporate MR imaging findings in the definition of symptomatic disease (Rajkumar et al., 2014; Dimopoulos et al., 2015).

Diffusion-weighted imaging (DWI) with calculation of apparent diffusion coefficients (ADCs) may be used as an adjunct method to increase diagnostic confidence and to better distinguish a diffuse MR imaging pattern from a normal MR imaging pattern. So far, this technique has been applied to a relatively small number of patients with myeloma, with promising results for initial assessment and prognosis (Messiou et al., 2011a; Padhani et al., 2013).

### **AIM OF THE WORK**

To evaluate the apparent diffusion coefficients (ADCs) of magnetic resonance (MR) imaging patterns in the bone marrow of patients with multiple myeloma (MM) and to determine a threshold ADC that may help distinguish a diffuse from a normal pattern with high accuracy.

### Chapter 1

# ANATOMY OF THE SPINE & BONE MARROW

The spine is a vital functional unit that protects the spinal cord and supports the head, thorax, abdomen, and pelvis. Together, the rigid osseous structures combined with ligaments, disks, spinal cord, and spinal nerves facilitate complex motor and sensory functions required for life, be it work or play (Harnsberger et al., 2006).

### **Vertebral Body Anatomy**

Except in special circumstances, the vertebrae assume a relatively constant anatomic theme and consist of a vertebral body and a vertebral (neural) arch. The vertebral body serves as a pillar, supporting the head and trunk. The posterior margin of the bodies, along with the vertebral arches, forms the bony spinal canal, protecting the spinal cord. The neural arch is formed by paired pedicles and paired lamina. Seven processes arise from the vertebral arch: paired superior and inferior articular processes, paired transverse processes, and one spinous process. The intervertebral foramen, formed by two adjacent vertebrae, allows for passage of spinal nerves and vessels (Fig. 1-1 & 1-2).

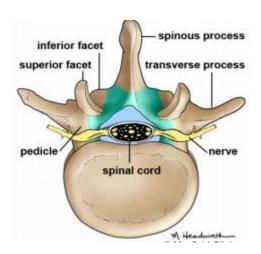


Fig. (1-1): Vertebral body.

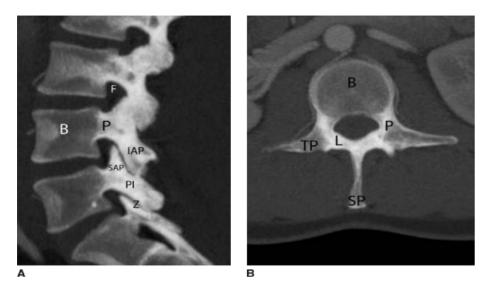


Fig. (1-2): A: Sagittal CT MIP of the lumbar spine demonstrating the vertebral body (B), pedicle (P), superior articular process (SAP), inferior articular process (IAP), and zygapophyseal joint (Z). The superior articular processes extend superiorly from the junction of the laminae and pedicles, while the inferior articular processes extend inferiorly from the undersurface of each lamina. The zygapophyseal joints are formed by the superior articular facet and the inferior articular facet. The pars interarticularis (PI) is the portion of the lamina between the superior and inferior articular processes. The intervertebral foramen (F) allows for the passage of spinal nerves and vessels. :Axial CT MIP of the lumbar spine demonstrating the vertebral body (B) and posterior neural arch. (P,pedicle; TP, transverse proocess; L, lamina; SP, spinous process).

The intervertebral discs consist of the inner nucleus pulposus and the outer annulus fibrosis. The main function of the disks is to distribute load and to allow flexion/extension and lateral bending. A remnant of the notochord, the nucleus pulposus, is usually located more dorsally compared to the center of the vertebral body. The annulus fibrosis attaches to the anterior longitudinal ligament (ALL) and posterior longitudinal ligament (PLL) and is also fused to the epiphyseal ring of the vertebral bodies. MRI of a normal disk demonstrates high signal on T2-weighted imaging related to the water content of the nucleus pulposus and inner annulus fibrosis. The outer annulus fibrosis demonstrates low signal on T1- and T2-weighted imaging (Fig 1-3) (Harnsberger et al., 2006).

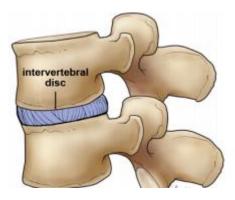


Fig. (1-3): Intervertebral disc