Review Article

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Multiparametric magnetic resonance imaging for prostate cancer: A review and update for urologists

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Recently, imaging of prostate cancer has greatly advanced since the introduction of multiparametric magnetic resonance imaging (mpMRI). mpMRI consists of T2-weighted sequences combined with several functional sequences including diffusion-weighted imaging, dynamic contrast-enhanced imaging, and/or magnetic resonance spectroscopy imaging. Interest has been growing in mpMRI because no single MRI sequence adequately detects and characterizes prostate cancer. During the last decade, the role of mpMRI has been expanded in prostate cancer detection, staging, and targeting or guiding prostate biopsy. Recently, mpMRI has been used to assess prostate cancer aggressiveness and to identify anteriorly located tumors before and during active surveillance. Moreover, recent studies have reported that mpMRI is a reliable imaging modality for detecting local recurrence after radical prostatectomy or external beam radiation therapy. In this regard, some urologic clinical practice guidelines recommended the use of mpMRI in the diagnosis and management of prostate cancer. Because mpMRI is the evolving reference standard imaging modality for prostate cancer, urologists should acquire cutting-edge knowledge about mpMRI. In this article, we review the literature on the use of mpMRI in urologic practice and provide a brief description of techniques. More specifically, we state the role of mpMRI in prostate biopsy, active surveillance, high-risk prostate cancer, and detection of recurrence after radical prostatectomy.

Keywords: Diagnosis; Magnetic resonance imaging; Prostatic neoplasms; Watchful waiting

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INTRODUCTION

Recently, interest has been growing in prostate magnetic resonance imaging (MRI) because of improvements in the accuracy of prostate cancer detection and characterization via the combination of conventional T1-weighted imaging and T2-weighted imaging (T2WI) with various functional imaging modalities [1]. These functional imaging modalities include diffusion-weighted MRI (DWI), dynamic contrast-

enhanced MRI (DCEI), and magnetic resonance spectroscopy imaging (MRSI). Combined conventional MRI with DWI, DCEI, and/or MRSI is known as multiparametric MRI (mpMRI) [2].

mpMRI is currently regarded as the most sensitive and specific imaging technique for prostate cancer evaluation, including detection, staging, localization, and aggressiveness measurement [3,4], mpMRI can also help to detect tumors missed on conventional biopsy and allow for targeted

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biopsies [5,6]. Furthermore, mpMRI has improved many aspects of prostate cancer management. mpMRI is used to monitor disease progression during active surveillance and recurrence after definite treatment [7,8]. Although mpMRI is not routinely recommended, recent guidelines have suggested that mpMRI can help in deciding on enrolment for active surveillance, in identifying anteriorly located prostate cancer in patients with suspicious negative biopsy results, and in deciding on nerve-sparing procedures in intermediate and high-risk disease [9-12].

This article presents an overview of mpMRI and the individual mpMRI techniques with their strong and weak points for clinical application. We also discuss the emerging role of MRI in detecting and managing prostate cancer.

TECHNIQUES AND DIAGNOSTIC ACCU-RACY OF mpMRI

Currently, mpMRI is regarded as the reference standard imaging modality for prostate cancer because a single MRI sequence cannot adequately detect and characterize prostate cancer. Although the ideal set of sequences for prostate mpMRI has not been determined, mpMRI is composed of high-resolution T2WI, DWI, and DCEI with optional MRSI. The principle, strong and weak points of each mpMRI sequence are summarized in Table 1. In addition, an image of a radical prostatectomy specimen and the corresponding mpMRI images are presented in Fig. 1 to aid in understanding the characteristics of each mpMRI sequence.

Table 1. Principles and characteristics of T2WI and each functional sequence

	'	'		
Sequence	Principle	Finding of prostate cancer	Advantages	Disadvantages
T2WI	Water content of tissue	Low signal intensity	High resolution Sharp demarcation of the prostate capsule	Central or transition zone tumor detection
DWI	Proton diffusion properties	High signal intensity on DWI Low signal intensity on ADC map	Central or transition zone tumor detection Assessment of tumor aggressiveness	Poor resolution and image distortion
DCEI	T1WI with contrast medium	Enhance and wash out rapidly	Local recurrence detection after defi- nite treatment	Long acquisition time
MRSI	Concentration of metabolites	Increased choline plus creati- nine/citrate	Assessment of tumor aggressiveness	Needs more expertise, long acquisition time

T2WI, T2-weighted imaging; DWI, diffusion-weighted imaging; ADC, apparent diffusion coefficient; DCEI, dynamic contrast-enhanced imaging; T1WI, T1-weighted imaging; MRSI, magnetic resonance spectroscopy imaging.

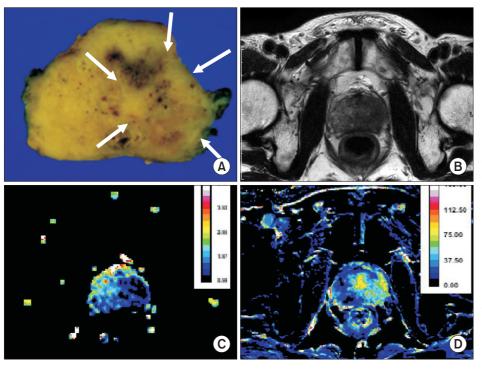


Fig. 1. A 72-year-old patient with prostate cancer. (A) On a picture of radical prostatectomy specimen, arrows indicate prostate cancer. (B) On T2-weighted imaging, prostate cancer shows slightly low signal intensity although the contrast between prostate cancer and adjacent normal tissue is not apparent. (C) On color-coded map of apparent diffusion coefficient, prostate cancer shows significantly decreased values, presented as dark blue color. (D) On initial area under the curve map derived from dynamic contrast-enhanced magnetic resonance imaging, prostate cancer shows increased vascularity, presented as yellow and green color.



T2WI, which reflects the water content of tissue, is the basis of mpMRI. Because of the high resolution and sharp demarcation of the prostate capsule, T2WI can be used to determine prostate zonal anatomy and prostate cancer staging. In contrast to cancer in other organs, prostate cancer presents low signal intensity compared with adjacent glandular tissue because the abundant amount of water in the normal gland demonstrates high signal intensity on T2WI. This signal difference between normal and cancer tissue helps in cancer detection in the gland-rich peripheral zone. However, cancer identification on T2WI may be limited in the transitional zone, which does not contain a large amount of water. Moreover, the T2 shortening effect by biopsy-induced hemorrhage decreases signal intensity even in noncancerous tissue. Therefore, despite satisfactory performance as reported by early studies, recent literature has demonstrated the limitations of T2WI for prostate cancer detection. Therefore, the sensitivity and specificity of T2WI show significant variation in studies, i.e., 55%-88% for sensitivity and 67%-82% for specificity [13]. Furthermore, such potential drawbacks of T2WI have introduced the need for mpMRI.

DWI, which quantifies the degree of water diffusing in the tissue, can be applied to prostate cancer detection because the high cellularity and abundant intra- and intercellular membranes in cancer tissue generally result in diffusion restriction. On an apparent diffusion coefficient (ADC) map, prostate cancer shows a lower signal intensity than do adjacent normal glands. Investigators have validated the usefulness of the ADC map for cancer detection, which has mostly demonstrated addition of the ADC map to T2WI to improve cancer detection. Moreover, assessment of tumor aggressiveness using DWI is attracting much attention, although accurate Gleason score prediction is difficult [14]. Although DWI can increase the sensitivity of prostate cancer detection, it is also limited for visualizing cancer tissue in the transitional zone because BPH-related nodules also demonstrate decreased diffusion. Moreover, tumor necrosis may manifest increased water diffusion, i.e., high signal intensity on an ADC map. More vulnerability to artifacts, relatively low signal-to-noise ratio, and low spatial resolution are also potential drawbacks of DWI. As a result, the sensitivity (65%–84%) and specificity (77%–87%) of the combined approach of DWI and T2WI has shown significant variability in different studies [13].

DCEI is an imaging modality that is designed to evaluate the status of tumor angiogenesis. DCEI requires the acquisition of repeat gradient echo images before and after injection of contrast materials such as chelated gadolinium. Owing to rapid imaging, DCEI provides the time-intensity curve in each voxel. Because tumors are evidently associated with neoangiogenesis that induces an increase in the blood volume and transvascular permeability, tracing the dynamic flux of the contrast agent with DCEI shows strong and rapid contrast enhancement. Therefore, DCEI helps to monitor treatment effects as well as cancer detection. Recently, studies have also reported that DCEI can improve diagnostic performance for detecting local recurrence in patients who undergo radical prostatectomy [15]. However, DCEI may cause false-positive diagnosis because inflammation is also accompanied by increased vascularity. Patient motion and peristalsis of the rectum during imaging may cause misregistration in imaging series, thereby disturbing the analysis of the time-intensity curve. The reported sensitivity and specificity of DCEI alone for prostate cancer detection also varies by reports (46%-90% for sensitivity and 74%-96% for specificity) [15].

MRSI depicts the relative concentration of certain metabolites. MRSI distinguishes prostate cancer with normal prostate tissue on the basis of the choline plus creatinine-to-citrate ratio. Increased choline and creatinine are observed in areas of proliferative prostate cancer. Citrate is produced by the normal prostate epithelium and is decreased in the area of prostate cancer. Thus, a higher choline plus creatinine-to-citrate ratio is observed in the areas of high tumor concentration. MRSI also provides information about the aggressiveness of prostate cancer. The sensitivity and specificity of MRSI alone are from 75% to 89% and from 77% to 91%, respectively [16]. MRSI has some limitations because the method takes additional time and requires more expertise than do other functional sequences.

CURRENT CLINICAL PRACTICE GUIDE-LINES ON mpMRI

The recommendation for mpMRI varies in different clinical practice guidelines. However, recently published guidelines from the European Association of Urology (EAU), the American Urological Association (AUA), and the National Comprehensive Cancer Network (NCCN) mention the potential role of mpMRI in prostate biopsy, active surveillance, and recurrent prostate cancer. Moreover, the EAU guideline recommended mpMRI in several fields of prostate cancer management.

For prostate cancer diagnosis, the EAU guideline suggested that mpMRI can be used to trigger a targeted repeat biopsy [9]. The EAU guideline suggested that mpMRI followed by transrectal ultrasonography (TRUS)-guided

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or MRI-guided biopsies might be particularly useful at identifying prostate cancer located in anterior regions, which is usually missed by conventional TRUS-guided biopsy. However, the EAU concluded that these results need to be verified and that the cost-effectiveness of mpMRI before confirmatory biopsy is questionable. In patients with low-risk prostate cancer, mpMRI is not recommended for staging purposes. The EAU suggested that mpMRI should be used in local staging only if its results change patient management. However, mpMRI can aid in deciding when to perform nerve-sparing procedures in intermediate and high-risk disease. In patients with biochemical recurrence who are candidates for salvage therapy, the EAU guideline suggested that mpMRI can guide biopsy [10]. However, the EAU guideline mentioned that it remains to be determined whether mpMRI will be able to correctly detect postprostatectomy local recurrences in patients with a prostatespecific antigen (PSA) level <0.5 ng/mL. When mpMRI is applied, a standardized scoring system, such as the Magnetic Resonance Prostate Imaging Reporting and Data System (MR PI-RADS) or Standards of Reporting for MRI-Targeted Biopsy Studies (START), is highly recommended.

The NCCN guideline suggested that mpMRI may be considered for patients who choose active surveillance to exclude the presence of anterior cancer if the PSA rises and systematic prostate biopsy results remain negative [11], mpMRI may provide additional information in certain clinical settings, such as rising PSA or a positive digital rectal exam result after radiotherapy in the setting of a negative prostate biopsy. mpMRI may be particularly useful in men being considered for local salvage therapy. However, they concluded that mpMRI is not recommended for routine use.

The AUA/ASTRO (American Society for Radiation Oncology) guidelines also suggested that mpMRI could identify sites of local recurrence and improve salvage radiation targeting [12]. Moreover, mpMRI could evaluate the need to add androgen deprivation in patients with massive recurrences not appropriate for conventional radiotherapy.

INTERPRETATION OF mpMRI

Although prostate mpMRI has been widely used, interpretation of mpMRI has not been standardized until quite recently. Before the introduction of the PI-RADS classification, a Likert scale of scores from 1 to 5 was used to characterize a radiologist's level of suspicion for focal lesions based on impression without fixed criteria. Lack of a standardized mpMRI reporting system led to substantial

variability in interpretation.

In 2012, the European Society of Urogenital Radiology (ESUR) introduced the PI-RADS classification for structured reporting of mpMRI [2]. The PI-RADS classification is the first and most widely accepted mpMRI scoring system. In PI-RADS, each suspicious lesion is scored on a 5-point scale for each sequence including T2WI, DWI, DCEI, and MRSI, with 1 being benign and 5 being highly suspicious of malignancy. The maximum score depends on the number of sequences performed. In addition, the scoring system recommends that a diagnosis of suspected prostate cancer be made if the PI-RADS score is 4 or higher (if 3 sequences were performed, sum of score ≥10, if 4 sequences were preformed, sum of score ≥13). Recent meta-analysis reported that PI-RADS appeared to have appropriate diagnostic accuracy with pooled sensitivity of 78% and specificity of 79% [17]. However, those authors suggested that the ESUR guideline should be improved in two aspects: the provision of clear instructions for overall score calculation (sum of score vs. overall 5-point scale) and recommendations regarding the use of a threshold for prostate biopsy. Because of heterogeneity among studies, a cutoff score for prostate biopsy cannot be provided.

Recently, PI-RADS version 2 dealt with these problems [18] PI-RADS version 2 assessed the probability of clinically significant prostate cancer by using a 5-point scale based on T2WI, DWI, and DCEI. In PI-RADS version 2, prostate biopsy should be considered for PI-RADS assessment category 4 or 5. For PI-RADS assessment category 2 or 3, biopsy may or may not be appropriate, depending on other clinical variables. In PI-RADS version 2, the primary determining sequence for a lesion located in the peripheral zone is DWI and that for a lesion located in the transition zone is T2WI. The score of the primary determining sequence and PI-RADS assessment category correspond closely regardless of the score of the other sequences. DCEI plays only a minor role in determining PI-RADS assessment category, and each lesion gets a (+) or (-) score based on DCEI.

Another mpMRI scoring system called the National Cancer Institute (NCI) MP-MRI evaluation score also exists, although this system has not been widely used [19]. The NCI MP-MRI scoring system is based on the number of positive sequences for prostate cancer. In this system, lesions are separated into 3 categories of low, moderate, and high risk. A lesion is considered low risk if positive on 1 or 2 of the 3 sequences. If all 3 sequences are positive, the lesion is considered moderate or high risk. A lesion is considered high risk if all 4 parameters are positive, including MRSI.

An international working group introduced standards of reporting MRI-targeted biopsy studies of the prostate, also



known as START [20]. They suggested the START checklist, which contain the panel's recommendations of statements to include in a report of MRI-targeted biopsy. According to this checklist, the field strength of the magnet, the specific coils being used, a brief description of the sequences, the reporting method used, and the experience of the reporting radiologist should be reported.

ROLE OF mpMRI IN PROSTATE BIOPSY

TRUS-guided 12-core biopsy is the generally recommended method for prostate biopsy and has been validated in enhanced cancer detection. However, about 20% of patients are reported to have prostate cancer on a repeat biopsy despite a previous negative biopsy result for cancer [21]. Moreover, a considerable number of patients experience an upgrade in postprostatectomy Gleason score [22]. This diagnostic uncertainty may lead patients to repeat prostate biopsy, delayed treatment, or overtreatment.

The use of mpMRI to improve the accuracy of prostate biopsy has increased because of several potential benefits of MRI-targeted or guided biopsy. One benefit is the precise localization of significant prostate cancer before biopsy, which may result in accurate pretreatment risk stratification. Recent studies have reported that localization of the index lesion by use of mpMRI is reliable [23,24]. In this regard, mpMRI is reported as an effective method for detecting and localizing clinically significant prostate cancer, especially in men with negative biopsy results [25]. Moreover, mpMRI increases repeat biopsy yields by identifying anteriorly located prostate cancer [26-29].

Several prostate biopsy methods are available depending on the means of applying MRI. The first method is visual estimation MRI-targeted TRUS-guided biopsy. This method carries a learning curve and lacks real-time feedback, although the additional cost is minimal. However, this method is limited because of inconsistencies in targeting precision. Another method is software coregistered MRI/ TRUS fusion biopsy, which overcomes the limitation of the previous method. MRI/TRUS fusion biopsy has greater reproducibility because of less operator dependence and real-time feedback. However, the additional cost of the device and software may be another problem. In-bore MRguided biopsy also has real-time feedback of biopsy needle placement in addition to increased accuracy. However, application of this method is limited because of economic feasibility and inability to routinely sample the remaining prostate. Recently, prostate biopsy using an MRI-compatible robot was introduced and reported as a feasible method [30,31].

Table 2. Recent articles reporting the role of mpMRI in prostate biopsy

Source Year Total MRI Previous biol Abdi et al. [32] 2015 192 86 (-) Results Kim et al. [33] 2015 68 34 Not applicable Jambor et al. [43] 2014 55 55 No previous bio Puech et al. [35] 2013 95 95 Not applicable Javali et al. [34] 2014 278 140 Not applicable Brock et al. [44] 2015 1,003 1,003 Not applicable Brock et al. [44] 2014 121 (-) Results Mozer et al. [37] 2015 152 Not applicable Rastinehad et al. [38] 2014 105 Not applicable Sonn et al. [45] 2014 105 Not applicable Ukimura et al. [40] 2015 177 107			Blobsy core (n)	ore (n)		מור במווריו	r Iostate calicel detection late	ומוב
Total MRI 2015 192 86 2015 68 34 2014 55 55 2013 95 95 2014 278 140 2015 1,003 1,003 2014 121 121 2015 152 152 2014 105 105 2013 649 649 2014 105 105 2015 127 127	Previous biopsy Sequence	Navigation	-		Overall	all	Significant cancer	nt cancer
2015 192 86 2015 68 34 2014 55 55 2013 95 95 2014 278 140 2015 1,003 1,003 2014 121 121 2015 152 152 2014 105 105 2013 649 649 2014 105 105 2015 127 127			MRI	TRUS	MRI	TRUS	MRI	TRUS
2015 68 34 2014 55 55 2013 95 95 2014 278 140 2015 1,003 1,003 2014 121 121 2015 152 152 2014 105 105 2013 649 649 2014 105 105 2015 127 127	esults DWI, DCEI	MRI-targeted	14	12	42%	19%	35%	14%
2014 55 55 2013 95 95 2014 278 140 2015 1,003 1,003 2014 121 121 2015 152 152 2014 105 105 2013 649 649 2014 105 105 2015 127 127	applicable DWI, DCEI	MRI-targeted	12	12	54%	21%	25%	%/
2013 95 95 2014 278 140 2015 1,003 1,003 2014 121 121 2015 152 152 2014 105 105 2013 649 649 2014 105 105 2014 105 105 2014 105 105 2015 127 127	No previous biopsy DWI, DCEI, MRSI	RSI MRI-targeted	1–5	12	%89	%16	%09	74%
2014 278 140 2015 1,003 1,003 2014 121 121 2015 152 152 2014 105 105 2013 649 649 2014 105 105 2015 105 105 2017 127 127	applicable DWI, DCEI	MRI-targeted	4	12	%69	%65	%29	52%
2015 1,003 1,003 2014 121 121 2015 152 152 2014 105 105 2013 649 649 2014 105 105 2015 127 127	applicable MRSI	MRI-targeted	6 or 12	6 or 12	24%	10%	NA	NA
2014 121 121 2015 152 152 2014 105 105 2013 649 649 2014 105 105 2015 127 127	applicable DWI, DCEI, MRSI	RSI MR/TRUS fusion	2	12	46%	47%	17%	12%
2015 152 152 2014 105 105 2013 649 649 2014 105 105 2015 127 127	esults DWI, DCEI	MR/TRUS fusion	4	12	79%	38%	24%	78%
2014 105 105 2013 649 649 2014 105 105 2015 127 127	No previous biopsy DWI, DCEI	MR/TRUS fusion	2-3	12	54%	21%	43%	37%
2013 649 649 2014 105 105 2015 127 127	applicable DWI, DCEI	MR/TRUS fusion	4	12	51%	46%	45%	32%
2014 105 105 40] 2015 127 127	applicable DWI, DCEI, MRSI	RSI MR/TRUS fusion	2/lesion	12	22%	NA	ΝΑ	NA
2015 127 127	esults DWI, DCEI	MR/TRUS fusion	4	12	34%	27%	22%	15%
	applicable DWI, DCEI	MR/TRUS fusion	æ	11	%19	41%	43%	23%
Quentin et al. [41] 2014 128 128 No previous	No previous biopsy DWI, DCEI	MRI-guided	2/lesion	12	23%	23%	45%	45%
Pokorny et al. [42] 2014 223 223 No previous	No previous biopsy DWI, DCEI	MRI-guided	m	12	%02	21%	%29	35%

mpikiki, multiparametric magnetic resonance imaging; i RUS, transrectai ultrasonography; DWI, dirfusion-weighted imaging; DCEI, dynamic contrast-enhanced imaging; MRSI, magnetic resonance spectroscopy imaging; NA, not applicable.

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The disadvantages of manually performed MRI-guided prostate biopsy might be eliminated with robotic biopsy.

Recent studies that have reported the accuracy of mpMRI targeted or guided biopsy are summarized in Table 2. MRI-targeted biopsy was reported to improve the detection rate of prostate cancer compared with conventional biopsy when a similar number of cores is biopsied [32-34]. Moreover, most studies reported that MRI-targeted or guided biopsy yielded a comparable prostate cancer detection rate with a decreased number of cores compared with conventional 12-core biopsy [35-42]. Although two studies reported a decreased detection rate, significant prostate cancer was comparably detected despite the lower number of biopsied cores [43,44]. The prostate cancer detection rate with the use of MRI targeted or guided 1- to 5-core biopsy as a first-round prostate biopsy method was comparable with that of 12-core TRUS-guided biopsy, although MRI-targeted biopsy showed slightly inferior results compared with MRI/TRUS fusion or MRI-guided biopsy [37,41-43]. Moreover, in men with negative biopsy results, the number of cores biopsied could be reduced with mpMRI instead of the standard 12-core biopsy [39]. In most studies, mpMRI consisted of DWI and DCEI, although we cannot determine the most appropriate set of sequences for prostate biopsy. The combination of DWI and DCEI showed a prostate cancer detection rate similar to that of mpMRI consisting of DWI, DCEI, and MRSI [36,43,45].

ROLE OF mpMRI IN ACTIVE SURVEIL-LANCE

Active surveillance is regarded as a standard treatment method for patients with low-risk and very-low-risk prostate cancer, which has indolent characteristics. Patients who undergo active surveillance are followed up with a regular PSA level test, digital rectal examination, and repeat prostate biopsy. Current practice guidelines do not recommend mpMRI for prostate cancer patients eligible for active surveillance. However, mpMRI has an emerging potential role in detecting clinically significant prostate cancer before enrollment in active surveillance and in reducing the number of repeat biopsies during active surveillance follow-up.

One of the most important aspects of active surveillance is selection of appropriate patients. In this regard, risk misclassification can be a huge problem. Some recent studies have suggested the promising detection rate of clinically significant prostate cancer in an active surveillance cohort (Table 3). However, the detection rate of significant prostate cancer varied widely within the studies if the definition of significant prostate cancer was changed [19,46-48]. Nevertheless, at least a quarter of active surveillance candidates with a suspicious lesion on mpMRI are reclassified as having clinically significant prostate cancer. Moreover, more than one-half of patients are identified as patients with significant prostate cancer if studies using prostatectomy specimen pathology as a standard reference are selected [47,49-51]. In other words, significant prostate cancer could be more precisely excluded before active surveillance enrollment if a lesion is not seen on mpMRI.

mpMRI during active surveillance was recently reported as a promising follow-up modality, although more studies on these fields should be done to gain a better understanding. Among functional sequences, DWI was reported as a potential monitoring sequence for patients with early prostate cancer who opt for active surveillance [52]. In that study, there was a significant decrease in ADC values over time in patients who progressed to radical treatment, although ADC values were stable in patients who did not progress to radical treatment. Moreover, Siddiqui et al. suggested that the active surveillance nomogram using

Table 3. Recent studies of the role of mpMRI in determining active surveillance eligibility

Source	Year	Patients No.	Sequence	Standard reference	Definition of sPCa	sPCa detection rate
Kim et al. [49]	2015	287	DWI	Prostatectomy specimen	Stage≥pT3 or TV≥0.5 mL or GS pattern ≥4	75%
Abd-Alazeez et al. [46]	2014	194	DWI, DCEI	Biopsy	Multiple definitions	25%-41%
Park et al. [50]	2014	298	DWI, DCEI	Prostatectomy specimen	Pathologic stage \geq T3 or GS \geq 7	52%
Stamatakis et al. [19]	2013	85	DWI, DCEI, MRSI	Biopsy	Multiple definitions	38%-56%
Turkbey et al. [51]	2013	133	DWI, DCEI, MRSI	Prostatectomy specimen	Stage≥pT3 or TV≥0.5 mL or GS pattern ≥4	99%
Shukla-Dave et al. [47]	2012	181	MRSI	Prostatectomy specimen	Multiple definitions	66%-77%
Thompson et al. [48]	2014	150	DWI, DCEI, MRSI	Biopsy or prostatectomy	Multiple definitions	30%-41%
				specimen		

mpMRI, multiparametric magnetic resonance imaging; sPCa, significant prostate cancer; DWI, diffusion-weighted imaging; TV, tumor volume; GS, Gleason score; DCEI, dynamic contrast-enhanced imaging; MRSI, magnetic resonance spectroscopy imaging.



mpMRI might reasonably decrease the number of repeat biopsies in patients undergoing active surveillance by as much as 68%, which agrees with other articles [53-55].

ROLE OF mpMRI IN HIGH-RISK PROS-TATE CANCER

High-risk prostate cancer has been defined differently in several guidelines and articles. However, the most generally accepted definition of high-risk prostate cancer is a clinical T stage≥T2c or T3a in addition to Gleason score≥8 or a PSA level>20 ng/mL [11,56]. Recently, some articles have reported mpMRI as a useful modality for predicting pathological outcomes in patients with high-risk prostate cancer (Table 4).

mpMRI has shown high specificity for detecting extracapsular extension (ECE), although sensitivity varied according to the study [57-60]. Additional mpMRI combined with conventional clinical variables increased prediction accuracy for recurrence after prostatectomy in high-risk disease [57]. Moreover, mpMRI improved decision-making to preserve the neurovascular bundle [60]. However, the negative predictive value for predicting ECE, which is important for deciding on nerve-sparing radical prostatectomy, was not sufficiently high in other studies [57-59]. Moreover, mpMRI was limited because it was unable to identify and localize focal ECE, which is a large majority of ECE [61]. Seminal vesicle invasion could be also predicted by using mpMRI with a positive predictive value of 62% to 95% and a negative predictive value of 73% to 83% [57.59].

The sensitivity and specificity of mpMRI for detecting lymph node invasion was 14% to 33% and 91% to 97%, respectively [57,59]. Because of low sensitivity, mpMRI was not routinely indicated for lymph node workup in patients without suspicious lymph nodes on computed tomography (CT) [62]. Moreover, positron emission tomography-CT show-

ed higher sensitivity and specificity for detecting lymph node invasion compared with mpMRI [59]. Despite these limitations, whole-body mpMRI as a one-stop imaging modality was recently proposed instead of prostate mpMRI plus bone scan plus CT in high-risk patients [63]. Although whole-body mpMRI was reported to be superior to a combination of imaging modalities, these results should be verified.

ROLE OF mpMRI IN DETECTION OF RE-CURRENCE AFTER RADICAL PROSTA-TECTOMY

Most postprostatectomy recurrent prostate cancer is diagnosed by PSA elevation. Once PSA increment is detected, it is essential to identify whether prostate cancer has recurred at a local or a distant site to determine the treatment modalities. In current practice, imaging or pathological evidence of local recurrence is not necessary to initiate local salvage treatment because current imaging techniques cannot adequately detect small-sized local recurrence.

In a recent meta-analysis, mpMRI was reported to have sufficient accuracy for detecting local recurrence in patients with low PSA and small-sized recurrence [64]. Recently, an increasing number of studies have been published reporting the acceptable diagnostic accuracy of mpMRI for detecting local recurrence (Table 5). Among the functional sequences, DCEI has been regarded as the most reliable sequence in detecting local recurrence after prostatectomy [65,66]. However, it must be taken into account that vascularity and contrast enhancement can be reduced in patients who have received androgen deprivation therapy. In this regard, the accuracy of DCEI might be reduced in patients who undergo androgen deprivation therapy. Sensitivity and specificity

Table 4. Recent studies of the role of mpMRI in high-risk prostate cancer

Source	Year	Patients No.	Sequence	Definition of high-risk prostate cancer	Factor	Sen (%)	Spe (%)	PPV (%)	NPV (%)
Jeong et al. [57]	2013	922	DWI	PSA≥20 ng/mL, GS≥8, clinical stage≥T2c	ECE	43	84	79	52
					SVI	35	94	62	83
					LNI	14	97	23	95
Pinaquy et al. [59]	2015	47	DWI	PSA≥20 ng/mL, GS≥8, clinical stage≥T2c	ECE	72	77	86	59
					SVI	73	95	95	73
					LNI	33	91	50	84
Somford et al. [58]	2013	183	DWI, DCEI	PSA≥20 ng/mL, GS≥8, clinical stage≥T2c	ECE	65	73	89	38
Park et al. [60]	2014	67	DWI, DCEI	PSA≥20 ng/mL, GS≥8, clinical stage≥T2c	ECE	80	85	89	74

mpMRI, multiparametric magnetic resonance imaging; Sen, sensitivity; Spe, specificity; PPV, positive predictive value; NPV, negative predictive value; DWI, diffusion-weighted imaging; PSA, prostate-specific antigen; GS, Gleason score; ECE; extracapsular extension; SVI, seminal vesicle invasion; LNI, lymph node invasion; DCEI, dynamic contrast-enhanced imaging.



Table 5. Recent studies of the role of mpMRI in detecting local recurrence after radical prostatectomy

Source	Year	Patients No.	Sequence	Standard reference	Sensitivity	Specificity
Wassberg et al. [67]	2012	52	DCEI	Biopsy	Reader 1: 88%	Reader 1: 74%
					Reader 2: 100%	Reader 2: 58%
Panebianco et al. [68]	2013	242	DWI, DCEI	Biopsy, PSA reduction after RT	DCEI: 98%-100%	DCEI: 94%-97%
					DWI: 97%-98%	DWI: 95%-96%
Roy et al. [66]	2013	28	DWI, DCEI, MRSI	Biopsy	DCEI: 97%	
					DWI: 65%	
					MRSI: 53%	
					DWI+DCEI: 94%	
					All sequences: 74%	
Kitajima et al. [70]	2014	115	DWI, DCEI	Biopsy, PSA reduction after RT	89%	85%
Linder et al. [65]	2014	187	DCEI	Biopsy, PSA reduction after RT, increased size on imaging study	91%	45%
Cha et al. [69]	2014	57	DWI, DCEI	Biopsy	Reader 1: 79%	Reader 1: 87%
					Reader 2: 90%	Reader 2: 82%

mpMRI, multiparametric magnetic resonance imaging; DCEI, dynamic contrast-enhanced imaging; DWI, diffusion-weighted imaging; PSA, prostate-specific antigen; RT, radiotherapy; MRSI, magnetic resonance spectroscopy imaging.

of DCEI alone for detecting local recurrence after radical prostatectomy range from 88% to 100% and from 45% to 97%, respectively [65-68]. Moreover, DCEI increased interobserver agreement and addition of DCEI to T2WI significantly increased accuracy for detecting local recurrence [67].

Recently, some studies have shown that DWI is also a reliable sequence in detecting local recurrence [68,69]. Moreover, the combination of DWI and DCEI seemed to have more consistent specificity of 82% to 87% compared with DCEI alone [66,69,70]. Accuracy of combined functional sequences has not been sufficiently reported. According to a recent study, T2WI plus DCEI has the highest sensitivity of 97% followed by DCEI alone and T2WI plus DWI plus DCEI [66].

CONCLUSIONS

mpMRI, which is composed of T2WI and several functional sequences, is regarded as the single most accurate imaging modality for characterizing prostate cancer. Recently, the role of mpMRI has been expanded to prostate biopsy, active surveillance, advanced disease detection, and local recurrence detection after radical prostatectomy. In this regard, urologists should acquire cutting-edge knowledge about mpMRI because it is a rapidly evolving imaging modality.

CONFLICTS OF INTEREST

The authors have nothing to disclose.

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