# **MAJOR PAPER**

# Robustness of a Combined Modified Dixon and PROPELLER Sequence with Two Interleaved Echoes in Clinical Head and Neck MRI

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**Purpose:** The combination of modified Dixon (mDixon) and periodically rotated overlapping parallel lines with enhanced reconstruction sequence with two interleaved echoes, which promotes uniform fat-suppression and motion insensitivity, has recently become available for commercial magnetic resonance imaging (MRI) scanners. To compare the robustness of this combination sequence with that of standard Cartesian mDixon sequence for fat-suppressed  $T_2$ -weighted imaging in clinical head and neck MRI.

**Methods:** Fifty patients with head and neck tumors were involved this study. All patients underwent MRI using both the combination and standard sequences. Two radiologists independently scored motion artifacts and water-fat separation error using a 4-point scale (1, unacceptable; 4, excellent). Furthermore, comprehensive comparative evaluation was performed using a 5-point scale (1, substantially inferior; 5, substantially superior). Data were statistically analyzed using the Wilcoxon signed-rank test.

**Results:** In the motion artifact assessment, ratings of 3 or 4 points were assigned to 45% (observer-1, 58.0%; observer-2, 32.0%) and 97% (100%; 94.0%) of images for the standard and combination sequences, respectively (P < 0.001). For the water–fat separation error assessment, ratings of 3 or 4 points were assigned to 100% (100%; 100%) and 85% (84.0%; 86.0%) of images, respectively (P < 0.001). In the comprehensive evaluation, of the 100 cases (observer-1, 50; observer-2, 50), 96 were rated at four or five points. In cases with slight or no motion artifacts and water–fat separation errors, the combination sequence was superior to the standard sequence in term of noise and sharpness, and equal in terms of contrast.

**Conclusion:** Although water-fat separation errors increased significantly in the combination sequence, most of these were acceptable. The significantly decreased motion artifacts in the combination sequence significantly improved image quality overall.

**Keywords:** Dixon, fat suppression, head and neck, magnetic resonance imaging, periodically rotated overlapping parallel lines with enhanced reconstruction

# Introduction

In the diagnosis of head and neck cancer, magnetic resonance imaging (MRI) is useful for evaluation of local invasion and lymph node metastases.<sup>1–3</sup> In particular, fat-suppressed  $T_2$ -weighted images ( $T_2$ WI) are essential for diagnosing the

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Received: November 6, 2019 | Accepted: March 3, 2020

extent of inflammation and edema.<sup>4</sup> The standard Cartesian modified Dixon (mDixon) sequence, termed the TSE mDixon XD sequence (dedicated software developed by Philips Medical Systems, Best, The Netherlands) is a uniform water-fat separated MRI method that uses two flexible echoes, a 7-peak fat model, and water-fat shift (WFS) correction algorism.5 It is able to acquire uniform fat-suppressed images even in high B<sub>0</sub> inhomogeneity regions, such as the oral and nasal cavities and the pharynx, compared with other fat suppression techniques.<sup>6-8</sup> However, this sequence is more sensitive to patient motion and pulsatile blood flow than are other fat suppression techniques, because it requires two echo times (TE) and repetition time (TR).9 As a measure against motion artifacts, the periodically rotated overlapping parallel lines with enhanced reconstruction (PROPELLER) method has been reported.<sup>10–12</sup> Recently, several methods for combining Dixon

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water–fat separation and PROPELLER motion correction have been reported.<sup>13–15</sup> At last, the combination of the mDixon and PROPELLER sequences with two interleaved echoes, proposed by Schär et al.,<sup>15</sup> has become available for commercial MRI scanners, and is termed the TSE\_mDixon\_ MultiVane\_XD sequence (dedicated software developed by Philips Medical Systems, Best, The Netherlands). However, to our knowledge, the robustness and usefulness of this sequence have not yet been proven in a clinical setting.

According to previous study by Schär et al.,<sup>15</sup> the TSE mDixon MultiVane XD sequence is designed to achieve both uniform water-fat separation and robustness to motion. Unlike the TSE mDixon XD sequence, two flexible echoes are acquired by shifting the acquisition window between odd and even echoes within one TR. Thus, both TE<sub>1</sub> and TE<sub>2</sub> blades are acquired simultaneously by a single shot, making it insensitive to motion. The Dixon water-fat separation is performed for each  $TE_1$  and  $TE_2$  pair with a  $B_0$  map; which acquire wateronly and fat-only blades, and correct the WFS. The motion correction algorism, which was developed based on the modified PROPELLER method,<sup>11</sup> is then performed. In addition, use of a parallel imaging technique, sensitivity encoding (SENSE),<sup>16</sup> leads to shorter scanning time and wider blade width.<sup>12</sup> Therefore, the blade width is equal to the TSE factor times the SENSE factor divided by two. When setting scan parameters, the shot per blade and TSE factor are limited to only 1 and 16 or more, respectively, meaning that only T<sub>2</sub>WI can be acquired.

We hypothesized that, in clinical cases with head and neck tumors, the TSE\_mDixon\_MultiVane\_XD sequence would yield decreased motion artifacts due to swallowing, breathing, etc., at a uniform fat-suppression, as compared with the TSE\_mDixon\_XD sequence. Thus, we here investigated whether the TSE\_mDixon\_MultiVane\_XD sequence was more robust in fat-suppressed  $T_2WI$  of the head and neck than the TSE\_mDixon\_XD in clinical cases.

# **Materials and Methods**

### Study patients

This study population consisted of 50 consecutive patients (5/2017-10/2017) who underwent head and neck MRI for diagnosis of tumor and follow-up in our institution and who agreed to participate in this study. Patient characteristics are presented in Table 1.

The prospective cross-sectional study was approved by the local Institutional Review Board, and written informed consent was obtained from all subjects.

### Magnetic resonance imaging

In this study, a 3T MR system (Ingenia Release 5.3; Philips Medical Systems, Best, The Netherlands), equipped with a 20-channel head and neck coil, was used. All patients underwent conventional routine MRI, including the  $T_2W_TSE_mDixon_XD$  ( $T_2W_mD$ ) sequence, to which the  $T_2W_TSE_mDixon_MultiVane_XD$  ( $T_2W_mD_MV$ ) sequence

Table 1 Patient characteristics

Characteristics	N = 50
Sex, n (%)	
Men	34 (68)
Women	16 (32)
Age, years	
Median (interquartile range)	68.5 (61–76)
Body mass index, kg/m <sup>2</sup>	
Median (interquartile range)	20.9 (19.0-22.7)
Tumor location, n (%)	
Oral cavity	33 (66)
Pharynx	11 (22)
Larynx	3 (6)
Salivary gland	2 (4)
Thyroid	1 (2)

was added consecutively. In order to exclude fatigue and familiarity bias, patients were randomly assigned to the  $T_2W_mD$  sequence-first group or  $T_2W_mD_MV$  sequence-first group, using computer-based randomization in advance. Before each scan, patients were given a short break of approximately 15 s, were told next scan duration, and were instructed not to move.

The detailed typical scan parameters are summarized in Table 2. In the T<sub>2</sub>W mD MV sequence, the TSE factor and SENSE factor were set to 32 and 2.5 respectively in order to widen the blade width and shorten the scan time. The increased TSE factor expanded the shot duration time, extended the TE, and caused increasing blur. To improve these problems, the echo spaces were reduced by widen bandwidth, and the TE-equivalent value was made equal to the  $T_2W$  mD sequence by setting the refocusing angle to 90°. The lower refocusing angle establishes pseudo steady state rapidly, which is able to shorten the TE-equivalent value and suppress blur.<sup>17</sup> The regional saturation band was placed on the inferior side to minimize pulsatile blood-flow artifacts. Depending on the tumor size and the extent of inflammation, the slice thickness, inter-slice gap, and number of slices were appropriately changed (from 3 to 6 mm, 10%) to 20%, and 20 to 50 slices). Finally, the SENSE factor was adjusted in the T<sub>2</sub>W mD sequence and MultiVane% was adjusted in the T<sub>2</sub>W mD MV sequence so that the scan time remained the same. Because we thought extending the scan time can increase the risk of exposure to patient movement.

### Image quality evaluation

The  $T_2W_mD_water$  and  $T_2W_mD_MV_water$  data sets were visually assessed in a random order on a workstation (SYNAPSE VINCENT; Fujifilm Co., Tokyo, Japan) by two experienced radiologists (observer-1, 11 years; observer-2, 30 years), independently. Both observers were unaware of patient information and sequence parameters. Adjustment of the

#### Table 2 Scan parameters

	T <sub>2</sub> W_mD sequence	T <sub>2</sub> W_mD_MV sequence	
Slice orientation	Transverse	Transverse	
FOV (mm)	240	240	
Acquisition voxel size (mm)	0.88/0.89/5.00	0.88/0.88/5.00	
Reconstruction voxel size (mm)	0.47/0.47/5.00	0.47/0.47/5.00	
Slice thickness (mm)	5	5	
Slice gap	0.5	0.5	
Slices	30	30	
Fold-over direction	AP	-	
SENSE reduction factor	1.5	2.5	
TSE factor	18	32	
Startup echoes	1	0	
Profile order	linear	linear	
TSE echo space/shot duration time (ms)	8.6/163	6.5/208	
Water-fat shift (pix)/ Bandwidth (Hz/pixel)	0.709/612.7	0.614/707.0	
Flip angle (°)	90	90	
Refocusing control	constant constant		
Angle (°)	120	90	
Act. TR (ms)	3500	3500	
Act. TE (ms)	90	90 107 (104/111)	
TE-equivalent (ms)	80	80 80	
REST slabs	1(inferior) 1(inferior)		
MultiVane percentage (%)	-	300	
Shots per blade	_	1	
MultiVane gross motion correction	_	yes	
NSA	1	1	
Packages	2	2	
Total scan time	02:27 02:34		

TSE, turbo spin echo; TR, repetition time; TE, echo time; FA, flip angle; FOV, field-of-view; NSA, number of signals averaged; SENSE, sensitivity encoding; mD, mDixon\_XD; MV, MultiVane\_XD; REST, regional saturation.

window-level and -width was allowed during their assessments. The range of assessment was from below the basicranium to the hypopharynx level. Brain and aliasing artifacts from the shoulder, a PROPELLER-specific error,<sup>18</sup> were not evaluated.

First, the motion artifacts and water–fat separation errors were assessed using a 4-point scale: 1 = unacceptable, diagnosis impossible due to severe artifact; 2 = poor, diagnosis possible although artifacts are present; 3 = good, artifacts slightly discernible; 4 = excellent, no artifacts. Second, for image quality evaluation in cases without motion artifacts and water–fat separation error, assessments of noise, sharpness, and contrast were added for cases that both observers assessed scored 3 or 4 points in both above artifact assessments. Two data sets were randomly toggled on the left or right side for blind review, and evaluated using the following 5-point scale: 1 = left images are substantially better than the right images; 2 = left images are better than the right images; 3 = left images are the same as the right images; 4 = left images are worse than the right images; 5 = left images are substantially worse than the right images. Finally, in all cases, it was evaluated which images were preferable overall for diagnosis using this 5-point scale.

### Statistical analysis

Normality of the distribution of the tested variables was analyzed with the Shapiro–Wilk test. Continuous variables are presented as median and quartiles for non-normally distributed variables, and categorical variables are expressed as absolute and relative frequencies (%). The motion artifact and water– fat separation error ratings were compared using the Wilcoxon signed-rank test. P < 0.05 was considered statistically significant. These statistical analyses were performed with SPSS software, release 23.0 (SPSS Inc., Chicago, IL, USA).

Linear weighted  $\kappa$ -coefficients were calculated using Excel to assess the inter-observer agreement. The  $\kappa$ -value is typically interpreted as an indication of no agreement when it is <0, as slight agreement when  $\kappa$  is 0–0.2, as fair agreement when  $\kappa$  is 0.2–0.4, as moderate agreement when  $\kappa$  is 0.4–0.6, substantial agreement when  $\kappa$  is 0.6–0.8, and almost perfect agreement when  $\kappa$  is 0.8–1.0.

# Results

Table 3 summarizes the results of assessment for motion artifact and water-fat separation error. In the motion artifact

Table 3 Artifact score by observer and technique

	Observer-1		Observer-2		
	mD	mD_MV	mD	mD_MV	
Motion artifact					
Excellent	6 (12%)	45 (90%)	2 (4%)	33 (66%)	
Good	23 (46%)	5 (10%)	14 (28%)	14 (28%)	
Poor	14 (28%)	0	17 (34%)	3 (6%)	
Unacceptable	7 (14%)	0	17 (34%)	0	
P-value	< 0.001		< 0.001		
$\kappa$ -coefficients	0.83				
Water–fat separation error					
Excellent	48 (96%)	21 (42%)	48 (96%)	30 (60%)	
Good	2 (4%)	21 (42%)	2 (4%)	13 (26%)	
Poor	0	5 (10%)	0	2 (4%)	
Unacceptable	0	3 (6%)	0	5 (10%)	
P-value	< 0.001		< 0.001		
<i>ĸ</i> -coefficients	0.92				

mD, modified Dixon\_XD; MV, MultiVane\_XD.

score, good or excellent ratings (scores of 3 or 4) were assigned to a mean of 45% of the  $T_2W_mD$  sequence images (observer-1, 58.0%; observer-2, 32.0%). In contrast, of those obtained with the  $T_2W_mD_MV$  sequence, on average 97% (observer-1, 100%; observer-2, 94.0%) had good or excellent ratings. The motion artifact score was rated significantly higher on  $T_2W_mD_MV$  sequence images than on  $T_2W_mD$ sequence images (P < 0.001). The water–fat separation error score from the  $T_2W_mD$  sequence was good or excellent in all cases. On the other hand, those from the  $T_2W_mD_MV$ sequence was good or excellent in approximately 85% of images (observer-1, 84.0%; observer-2, 86.0%). There was a significant difference between the  $T_2W_mD$  sequence and T<sub>2</sub>W\_mD\_MV sequence in terms of water–fat separation error score (P < 0.001). For inter-observer agreement, the calculated linear weighted  $\kappa$ -coefficients were 0.83 and 0.92 for motion artifact and water–fat separation error scores, respectively, indicating almost perfect agreement.

Representative cases of the  $T_2W_mD$  sequence and the  $T_2W_mD_MV$  sequence are shown in Fig. 1. Motion artifact was substantially improved, although a slight water–fat separation error is observed for subcutaneous fat. In addition, typical cases with unacceptable water–fat separation error in the  $T_2W_mD_MV$  sequence are shown in Fig. 2. All severe water–fat separation errors were observed in images of a relatively raised jaw.



**Fig. 1** Comparison between  $T_2W_mD$  sequence and  $T_2W_mD_MV$  sequence. Upper panel: The pharynx in a 63-year-old male patient with oropharynx squamous cell carcinoma before treatment. Lower panel: The oral cavity in a 68-year-old male patient with oral tongue squamous cell carcinoma before treatment. The  $T_2W_mD$  sequence shows severe motion artifacts which cover signal changes [(**a** and **c**) dotted arrows]. The  $T_2W_mD_MV$  sequence shows no motion artifact; thus, the tumor signals are observed clearly [(**b** and **d**) arrow]. However, in the  $T_2W_mD_MV$  sequence, water–fat separation errors occurred in a portion of subcutaneous fat [(**d**), arrowhead].  $T_2W, T_2$ -weighted image; mD, modified Dixon\_XD; MV, MultiVane\_XD.



Fig. 2 Unacceptable water-fat separation error in the T<sub>2</sub>W\_mD\_MV sequence. Upper panel: An 88-yearold female patient with postoperative recurrence of buccal mucosa squamous cell carcinoma. Lower panel: A 47-year-old male patient with nasopharynx squamous cell carcinoma after radiation-chemotherapy. (a and d) Sagittal image from  $T_2W_mD$  sequence. (**b** and **e**) Transverse image at the level of the jaw from T<sub>2</sub>W mD sequence. (c and f) Transverse image at the level of the jaw from T<sub>2</sub>W\_mD\_MV sequence. A severe water-fat separation error is observed on not only subcutaneous fat [(**c** and **f**) arrows], but also mandibular marrow [(**c** and **f**) arrowheads].  $T_2W$ ,  $T_2$ weighted image; mD, modified Dixon\_XD; MV, MultiVane\_XD.

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**Fig. 3** Image quality assessment between  $T_2W_mD$  sequence and  $T_2W_mD_MV$  sequence in cases with slight or no motion artifact and water–fat separation error. (a) Noise assessment. (b) Sharpness assessment. (c) Contrast assessment. Each bar graph shows a comparison of 16 cases by the two respective observers. The five categories correspond to the  $T_2W_mD_MV$  sequence being substantially better or worse than (>>), better or worse than (>), or equal to (=) the  $T_2W_mD_sequence.$  Grey and black colors represent the two radiologists' comparisons, respectively.  $T_2W$ ,  $T_2$ -weighted image; mD, modified Dixon\_XD; MV, MultiVane\_XD.



**Fig. 4** Comprehensive comparative assessment between the  $T_2W_mD$  sequence and  $T_2W_mD_MV$  sequence. Bar graph shows a comparison of all cases by the two respective observers. The five categories correspond to the  $T_2W_mD_MV$  sequence being substantially better or worse than (>>), better or worse than (>>), or equal to (=) the  $T_2W_mD$  sequence. Grey and black colors represent the two radiologists' comparisons, respectively.  $T_2W$ ,  $T_2$ -weighted image; mD, modified Dixon\_XD; MV, MultiVane\_XD.

Figure 3 shows the results of comparative assessments of noise, sharpness, and contrast in the cases with slight or no motion artifact and water–fat separation error. In terms of noise, the  $T_2W_mD_MV$  sequence was superior to the  $T_2W_mD_MV$  sequence (Fig. 3a). In terms of sharpness, the  $T_2W_mD_MV$  sequence was slightly superior to the  $T_2W_mD_MV$  sequence (Fig. 3b). In terms of contrast, these sequences were almost equivalent (Fig. 3c).

Figure 4 shows the results of the overall comparative evaluation. The T<sub>2</sub>W\_mD\_MV sequence was superior to the T<sub>2</sub>W\_mD sequence in almost cases (observer-1, 94.0%; observer-2, 98.0%;  $\kappa = 0.92$ ).

### Discussion

In this study, we compared the robustness of the newly available combination sequence with that of standard Cartesian mDixon sequence for fat-suppressed  $T_2WI$  of the head and neck in a clinical context. Our main finding was that the  $T_2W_mD_MV$  sequence improved motion artifacts and deteriorated water-fat separation error compared with the  $T_2W_mD$  sequence.

In this study, both observers scored some cases as having unacceptable motion artifacts for diagnosis on the  $T_2W_mD$ 

sequence. The T<sub>2</sub>W mD sequence is sensitive to patient motion as it requires two flexible echoes across TR. Additionally, a higher magnetic field tends to deteriorate the motion artifacts.<sup>19</sup> These drawbacks are critical in head and neck MRI because motion in this region occurs frequently due to swallowing, breathing, etc. In contrast, neither observer found any cases with unacceptable motion artifact for diagnosis on the  $T_2W$  mD MV sequence images. The  $T_2W$  mD MV sequence does not require multiple TRs as it acquires two echoes interleaved in each shot.15 Furthermore, the modified PROPELLER algorism makes this approach more insensitive to patient motion. While, with the T<sub>2</sub>W mD MV sequence, water-fat separation error was more frequently observed than with the T<sub>2</sub>W mD sequence. Although most of them were confined to a portion of subcutaneous fat and were acceptable for diagnosis, in a few cases, an unacceptable water-fat separation error was observed. All severe water-fat separation errors were observed on a relatively raised jaw, suggesting that they were caused by large B<sub>0</sub> inhomogeneities. The Dixon water-fat separation needs precision  $B_0$  map. In the  $T_2W_mD$  sequence, the  $B_0$  map is produced in each slice. On the other hand, in the  $T_2W$  mD MV sequence, the  $B_0$ map is produced in each low-resolution blade. Schär et al.<sup>15</sup> described that water-fat swaps may occur in some of blades partly due to the low resolution in the phase encoding direction, and level of artifacts depend on how many affected blades. Therefore, we consider that severe water-fat separation errors in raised jaw result from all or almost blades water-fat swaps. Schär et al.<sup>15</sup> have shown that combination of the mDixon and PROPELLER sequence with two interleaved echoes is superior for fat-suppressed T<sub>2</sub>WI as compared with the spectral fat-saturated standard PROPELLER sequence. Our study showed that the combination of mDixon and PROPELLER sequences is inferior to the standard mDixon sequence in terms of fat suppression. However, this unacceptable water-fat separation error may be resolved by making  $B_0$  inhomogeneities smaller with a simple method, such as ensuring that the patient's chin is lowered.

We had originally hypothesized that, in cases without motion artifact and water-fat separation error, the T<sub>2</sub>W mD MV sequence would lead to a slight blur, low noise, and equivalent contrast compared with the T<sub>2</sub>W mD sequence. The  $T_2W$  mD MV sequence has non-filled k-space corners and an oversampled k-space center,<sup>10,11</sup> and the TE-equivalent value was adjusted to equal by lowering the refocusing pulse. These hypotheses were consistent with the study results in terms of noise and contrast. However, in terms of sharpness, the T<sub>2</sub>W mD MV sequence was slightly superior to the  $T_2W$  mD sequence. This may be due to the use of a criterion of "motion artifact score of 3 or 4 points," because deteriorated sharpness was attributed to even a slight motion artifact. Although it might have been better to assess images using only a motion artifact score of 4 points, the number of cases assigned a score of 4 was too small to allow comparison [observer-1, six cases (12%); observer-2, two cases (4%)]. If the influence of motion artifacts was completely excluded, result of sharpness assessment may reverse since the  $T_2W_mD$  sequence has filled *k*-space corners compared with the  $T_2W_mD_MV$  sequence. However, in these three assessments, there was no case that  $T_2W_mD_MV$  sequence were substantially worse than the  $T_2W_mD$  sequence. These results convince us that  $T_2W_mD_MV$  sequence does not critically deteriorate image quality to undiagnosable level.

In the comprehensive comparative evaluation, both observers assessed that the  $T_2W_mD_MV$  sequence was preferable for diagnosis compared with the  $T_2W_mD$  sequence in almost cases. This result suggests that the degree of motion artifact is particularly important among various image quality assessment factors. Therefore, the comprehensive evaluation was influenced substantially by the robustness to patient motion.

Our study had some limitations. First, although image assessments were performed in blinded manner, experienced radiologists might recognize what sequence was applied. Second, only visual assessment was performed, and physical assessment was not performed. In this study, noise measurement was difficult, since many of the acquired clinical images had motion artifacts; furthermore, using parallel imaging and a radial sampling technique complicated the appearance of noise. Therefore, it was impossible to compare image quality accurately using the signal-to-noise ratio or contrast-to-noise ratio. Third, in assessments of noise, sharpness, and contrast, the influence of motion artifacts could not be completely excluded. The possibility that the findings of those assessments would differ if sample size was larger and the influence of motion artifacts was completely excluded cannot be ruled out. Especially, result of sharpness assessment will almost certainly reverse. Finally, sensitivity and specificity were not calculated by receiver operating characteristic (ROC) curve analysis. It is difficult to perform ROC analysis in this context, because there are few opportunities to scan normal patients due to our institution's policies. Thus, these points may need to be studied further. However, the observed significant reduction in motion artifacts suggests that we accomplished the purpose of this study, which was to investigate the robustness of the new combined sequence.

### Conclusion

For acquiring fat-suppressed  $T_2WI$  of the head and neck in clinical cases, the combination of mDixon and PROPELLER sequences with two interleaved echoes increased the water–fat separation error, as compared with the standard Cartesian mDixon sequence. However, this sequence decreased motion artifacts significantly and improved overall image quality, suggesting its potential for enhancing diagnostic performance.

# **Conflicts of Interest**

None of the authors has any conflicts of interest regarding this study.

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