The Reliability of Reduced Field-of-view DTI for Highly Accurate Quantitative Assessment of Cervical Spinal Cord Tracts

Takumi Yokohama¹, Motoyuki Iwasaki², Daisuke Oura¹, Sho Furuya², and Tomoyuki Okuaki³

Purpose: To compare the accuracy of fractional anisotropy (FA) and apparent diffusion coefficient (ADC) values between reduced FOV or so-called zonally oblique multislice (ZOOM) and conventional diffusion tensor imaging (DTI) in the cervical spinal cord.

Methods: Both ZOOM and conventional DTI were performed on 10 healthy volunteers. Intraclass correlation coefficient (ICC) was used to evaluate the reliability of the measurements obtained. Four radiologists evaluated the FA and ADC values at each cervical cord level and classified the visibility by 4 ranks. The geometric distortion ratios of the long axis and short axis were compared between ZOOM and conventional DTI. The imaging parameters were as follows: b-value = 600 s/mm²; TR = 4500 ms; TE = 81 ms; FOV = 70 × 47 mm² / 200 × 200 mm²; matrix = 80 × 51 / 128 × 126 (ZOOM and conventional DTI, respectively). The region of interest was carefully drawn inside the spinal cord margin to exclude the spinal cord component, without excluding the white matter fiber tracts.

Results: The average FA value decreased in both ZOOM and conventional DTI in lower spinal cord levels; in contrast, the ADC value increased in lower spinal cord levels. Zonally oblique multislice DTI was superior to conventional DTI with regard to inter-rater and intra-rater reliability; further, visibility was better and the standard deviation was smaller in ZOOM DTI. On both the long and short axis, the geometric distortion ratio was lower in ZOOM DTI at all cervical spinal cord levels compared with the conventional DTI. There was a significant difference in the distortion ratios of the long and short axis between ZOOM and conventional DTI.

Conclusion: Conventional DTI is unreliable owing to its susceptibility to the surrounding magnetic field. ZOOM DTI is reliable for performing highly accurate evaluations.

Keywords: fractional anisotropy, outer volume suppression, single-shot echo-planar imaging, small field-of-view
obtaining diffused images in at least one-dimensionally limited area. Field-of-view reduction is achieved by the application of non-coplanar excitation and refocusing pulses combined with outer volume suppression (OVS) for removal of unwanted transition zones.\textsuperscript{12} The outline of ZOOM DTI is shown in Fig. 1. Zonally oblique multislice DTI is based on a non-coplanar SE technique,\textsuperscript{13} which apply tilted radio frequency (RF) pulse between the excitation and the slice-selective region. As a result, a small, limited region is excited. Furthermore, the suppression pulse is imposed around the FOV, which is excited by non-coplanar SE to establish a rectangular FOV. This technique is called OVS.\textsuperscript{13} Small, non-distortion FOV is acquired by non-coplanar SE and OVS on ZOOM DTI. A diagram of ZOOM DTI is shown in Fig. 2. Zonally oblique multislice DTI is a diffusion-weighted (DW) SS-EPI sequence, and it is preceded by presaturation OVS sequence and the spectral presaturation with inversion recovery pulse. Suppression slabs are implemented by successive quadratic phase RF pulses\textsuperscript{14} (broad bandwidth, selective) followed by crusher gradients, and a small, non-distortion FOV image can be acquired by this technique. The advantages of cervical spinal DTI with ZOOM DTI are as follows: small FOV imaging reduces the readout scan time, so the artifact caused by off-resonance is also decreased.\textsuperscript{12} The phase encode step is reduced and no phase wrap is required owing to the limited FOV excited with non-coplanar SE and OVS.\textsuperscript{15} Hence, a fine cervical spinal cord DTI is obtained after a short scan time using ZOOM DTI.\textsuperscript{10} However, the differences in the reliability of the FA and ADC value measurements between ZOOM and conventional DTI have not been determined. The purpose of this study was to compare and establish the reliability of measured FA and ADC values and image quality using ZOOM DTI compared with conventional-DTI for the cervical spinal cord.

**Materials and Methods**

**Device and imaging target**

Imaging was performed on 10 healthy volunteers (mean age, 31.9 years [range, 23–57 years], 6 men and 4 women), using a 3T MR scanner (Ingenia; Koninklijke Philips N.V., Amsterdam, Netherlands). This study was approved by the Institutional Review Board of Otaru General Hospital.

**Imaging parameters**

Zonally oblique multislice and conventional DTI imaging sequences were used to obtain cervical spinal DTI images. Imaging settings are shown in Table 1.

Zonally oblique multislice DTI parameters were as follows: TR = 4500 ms; TE = 81 ms; FOV = 70 × 47 mm\textsuperscript{2};

![Fig. 1 Non-coplanar spin-echo (SE). (a) In general, the 90° radio frequency (RF) pulse is applied perpendicular to the imaging plane, but the 90° RF pulse is applied tilted by a small angle, \( a \), with respect to the imaging plane. This approach allows multiple-slice acquisition, but achieves an imperfect signal profile with an unwanted transition width \( d_{tr} \). (b) The 90° RF pulse is again tilted by angle \( a \), but additional outer volume suppression slabs are applied on both sides. OVS, outer volume suppression.](image1)

![Fig. 2 The location of outer volume suppression (OVS). The OVS is located, although no folding occurs in the phase direction.](image2)

| Sequence | ZOOM DTI | Conventional DTI |
|----------|----------|------------------|
| TR (ms)  | 4500     | 4500             |
| TE (ms)  | 81       | 81               |
| Matrix size (phase × read out) | 72 × 46 | 128 × 126 |
| FOV (mm) | 70 × 47  | 200 × 200        |
| Slice thickness (mm) | 5 | 5 |
| Slices | 24 | 24 |
| NSA | 10 | 5 |
| b-factor (s/mm\textsuperscript{2}) | 600 | 600 |
| Fat suppression | SPAIR | SPAIR |
| Scan time | 10 min 35 sec | 10 min 3 sec |

NSA, number of sample averages; SPAIR, spectral attenuated inversion recovery; ZOOM, zonally oblique multislice.
matrix = 80 × 51; number of sample averages (NSA) = 10; scan time = 10 min 35 sec. Conventional DTI parameters were as follows: TR = 4500 ms; TE = 81 ms; FOV = 200 × 200 mm²; matrix = 128 × 126; NSA = 5; scan time = 10 min 3 sec.

Scan time was set to the same interval in both settings, and the following parameters were the same: single-shot SE, b-value 600 s/mm²; 6 motion probing gradient (MPG) direction; slice thickness of 5 mm; and 24 slices. The acquired scan range was from the second cervical to the first thoracic vertebra. The slab center was placed at C4/5. The range of imaging for ZOOM and conventional DTI is shown in Fig. 3.

**Evaluation and analysis**

**Quantitative value by cervical spine level**
The FA map was calculated on ZOOM and conventional DTI. The FA value and apparent diffusion coefficient (ADC) of the spinal cord were measured. The ROI was placed on the spinal cord and enlarged maximally; however, the surrounding cerebrospinal fluid (CSF) was excluded by checking the 3D ROI square. The method used for setting the ROI is shown in Fig. 4. All operations were performed using Fiber Track (Philips Electronics Japan, Tokyo, Japan), the software in the MRI scanner. The mean and standard deviation (SD) of the FA value was calculated using scan data of 10 healthy volunteers. The Wilcoxon rank sum test was used for statistical analysis; analyses were performed using JMP version 12 (SAS Institute, Inc., Cary, NC, USA).

**Evaluation of measurements**
To examine the reliability of the measurements obtained from spinal cord imaging in 10 volunteers, intraclass correlation coefficient (ICC) was used to evaluate the inter-rater and intra-rater reliability of the FA and the ADC values. According to Shrout et al., ICC has three forms, namely Case 1, Case 2, and Case 3 (thesis) and can be expressed as ICC (n, k), where n refers to Case 1, 2, or 3, and k indicates number of people.

Intraclass correlation coefficient Case 1 is the intra-examiner confidence factor when one observer evaluates multiple subjects, and it is calculated by the following equation:

\[
ICC(1,1) = \frac{BMS - WMS}{BMS + (k-1)WMS}
\]

where BMS is between sum of mean square, WMS is within sum of mean square.

The difference between Case 2 and Case 3 is that the former is a random effect model and the latter is a fixed model. In this study, we adopted Case 2, which was calculated by the following formula:

\[
ICC(2,1) = \frac{BMS - EMS}{BMS + (k-1)EMS + k(JMS-EMS)} / n
\]

where EMS is the residual sum of mean square, and JMS is the judges sum of mean square.

The inter-rater reliability was evaluated by three radiologists (years of experience: 12, 5, and 3 years). The criteria of ICC are shown in Table 2.

**Image quality**
We compared the image qualities of ZOOM and conventional DTI using a 4-point scale visual evaluation. Four radiological technologists with 26, 14, 11, and 2 years of experience evaluated the distortion with b = 0 image as the visibility. Transverse images of each cervical spinal cord level evaluated are shown in Fig. 5. The scoring system was as follows: 4 = no image distortion, and spinal cord and CSF can be clearly discriminated; 3 = minimal image distortion, and the cord-CSF boundary line can be discriminated; 2 = considerable image

| Table 2 | The criteria of intraclass correlation coefficient (ICC) |
|---------|------------------------------------------------------|
| 0.9     | Great                                                |
| 0.8     | Good                                                 |
| 0.7     | Ok (fair)                                            |
| 0.6     | Possible                                              |
| 0.6     | Re-work                                               |

Re-work is necessary for ICC less than 0.6.

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**Fig. 3** The imaging range of zonally oblique multislice (ZOOM) diffusion tensor imaging (DTI) and conventional DTI.

**Fig. 4** Setting of the ROI for the cervical spinal cord. Notably, the entire cord was contoured as widely as possible while excluding the surrounding cerebrospinal fluid.
distortion, and it is difficult to discriminate the cord-CSF boundary line; 1 = poor image quality, and a cord-CSF boundary that cannot be completely differentiated.

Data were analyzed using the Tukey–Kramer method; a $P$-value < 0.05 was considered significant.

The distortion evaluation using the NEMA method
On a $b = 0$ image, we compared the geometric distortion ratio of the long and short axis between ZOOM DTI and conventional DTI by the National Electrical Manufacturers Association (NEMA) method. A transverse T$_2$-weighted image (T$_2$WI) obtained using a fast SE sequence was used as a standard image. Measurement of the long and short axis is shown in Fig. 6. The geometric distortion ratio was calculated using the following equation:

$$\text{Geometric distortion ratio} = \left( \frac{\text{Lm} - \text{La}}{\text{La}} \right) \times 100$$

where Lm = the length of the standard image and La = the length of the DTI image. Data were analyzed using the Wilcoxon rank sum test; a $P$-value < 0.05 was considered significant.

Measurement of SNR
The signal-to-noise ratio (SNR) for spinal cord imaging using the ZOOM and conventional DTI was calculated using ImageJ (National Institutes of Health, Bethesda, MD, USA), and the average value was obtained.

The ROI was a circle with a diameter of 30 mm and did not contain the CSF component. The setting of the ROI is shown in Fig. 7. The SNR can be obtained by the following equation:

$$\text{SNR} = \frac{\text{Signal intensity (SI)}}{\text{Standard deviation (SD)}}$$

Results

Quantitative values of the cervical spine
Fractional anisotropy and ADC values obtained using ZOOM and conventional DTI are shown in Table 3 and Figs. 8–11. The FA and ADC values of the cervical spinal cords were approximately 0.6 and 1.0, respectively. In both ZOOM and conventional DTI, the average FA value decreased in lower spinal cord levels; in contrast, the ADC value increased in lower spinal cord levels. The SD of the FA and ADC values were significantly higher in the lower cervical spine, especially at C5/6 and C6/7, in the conventional DTI compared with ZOOM DTI.

Reliability of the measurements
The intra- and inter-rater reliability at each cervical spinal cord level is shown in Tables 4 and 5. In terms of inter-rater reliability, for the FA value, ICC (2, 1) for ZOOM DTI was higher than 0.7 in all cervical spinal cords; however, in conventional DTI, ICC (2, 1) was less than 0.6 at C5/6. Similarly, for the ADC value, ICC (2, 1) for ZOOM DTI was larger than 0.6 in all the cervical spinal cords; in contrast, for the conventional DTI, ICC (2, 1) was less than 0.6 at C2/3, C4/5, and C5/6.

In terms of intra-rater reliability, for the FA value, ICC (1, 1) was higher than 0.6 in all cervical spinal cords in both ZOOM and conventional DTI. By contrast, for the ADC value, ICC (1, 1) for ZOOM DTI was less than 0.6 at C2/3.
Table 3  Mean and standard deviation of fractional anisotropy (FA) and apparent diffusion coefficient (ADC) at each cervical spinal level

|        | FA             | ADC             |
|--------|----------------|-----------------|
|        | ZOOM DTI \( n = 10 \) | Conventional DTI \( n = 10 \) | ZOOM DTI \( n = 10 \) | Conventional DTI \( n = 10 \) |
| C2/3   | 0.717 ± 0.0576 | 0.681 ± 0.0543  | 0.1303  | 0.9893 ± 0.1031 | 1.026 ± 0.9707 | 0.4497 |
| C3/4   | 0.690 ± 0.0613 | 0.684 ± 0.0858  | 0.5967  | 0.9610 ± 0.1013 | 1.027 ± 0.2040 | 0.5706 |
| C4/5   | 0.636 ± 0.0742 | 0.629 ± 0.123   | 0.7624  | 1.052 ± 0.2033 | 1.125 ± 0.3263 | 0.8798 |
| C5/6   | 0.594 ± 0.0698 | 0.632 ± 0.0879  | 0.1509  | 1.163 ± 0.1857 | 1.140 ± 0.2068 | 0.7335 |
| C6/7   | 0.554 ± 0.0854 | 0.569 ± 0.00804 | 0.7624  | 1.294 ± 0.1956 | 1.251 ± 0.2603 | 0.7054 |

*Wilcoxon rank sum test; values are presented as means ± standard deviations in a ROI. DTI, diffusion tensor imaging; ZOOM, zonally oblique multislice.

Fig. 8 Fractional anisotropy (FA) value at each cervical spinal level in zonally oblique multislice (ZOOM) diffusion tensor imaging. The FA value gradually decreased at the lower spinal cord levels.

Fig. 9 Fractional anisotropy (FA) value at each cervical spinal level in conventional diffusion tensor imaging (DTI). Note that the lower levels show a wider range of value than zonally oblique multislice (ZOOM) DTI.

Fig. 10 Apparent diffusion coefficient (ADC) value at each cervical spinal level on zonally oblique multislice (ZOOM) diffusion tensor imaging.

Fig. 11 Apparent diffusion coefficient (ADC) value at each cervical spinal level on conventional diffusion tensor imaging.

Additionally, ICC (1, 1) for conventional DTI was less than 0.6 at C2/3 and C6/7.

**Image quality**
The result of an image quality evaluation is displayed in Fig. 12. Zonally oblique multislice DTI was superior to conventional DTI at all cervical spinal levels using a 4-point scoring system. The scores at all cervical levels in ZOOM DTI were significantly higher than those of the conventional DTI.

**Distortion evaluation using the NEMA method**
The results of the geometric distortion ratio are shown in Table 6 and Figs. 13 and 14. On both the long and short axis, the geometric distortion ratio was lower in ZOOM DTI at all cervical spinal cord levels compared with the conventional
DTI. The geometric distortion ratio was higher on the short axis than in the long axis.

There was a significant difference in the distortion ratio of the long axis between ZOOM and conventional DTI at each spinal cord level; C2/3: 1.216% and 7.672% ($P < 0.05$); C3/4: 1.396% and 9.098% ($P < 0.01$); C4/5: 2.283% and 10.44% ($P < 0.05$); and C5/6: 8.347% and 21.55% ($P < 0.05$), in ZOOM and conventional DTI, respectively.

Similarly, the distortion ratio of the short axis was significantly different at C3/4: 0.3231% and 19.11% ($P < 0.01$), in ZOOM DTI and conventional DTI, respectively.

Signal-to-noise ratio
The results of SNR obtained at each spinal cord level are shown in Table 7 and Fig. 15. Although ZOOM DTI showed a larger variation in SNR, its SNR was larger than that of conventional DTI, except at C4/5.

Discussion
In this study, the acquired FA and ADC values at each cervical spinal level were approximately 0.6 and 1.0, respectively. The FA and ADC value were equivalent when compared with previous studies.\textsuperscript{18,19} However, we obtained a stable FA and ADC value in ZOOM DTI with regard to their SD at each

| Table 4 | Inter-rater reliability between ZOOM and conventional DTI |
|---------|-----------------|
|         | FA              | ADC              |
|         | ZOOM            | Conventional     | ZOOM            | Conventional     |
| C2/3    | 0.881           | 0.788            | 0.648           | 0.207            |
| C3/4    | 0.721           | 0.620            | 0.602           | 0.686            |
| C4/5    | 0.839           | 0.637            | 0.684           | 0.543            |
| C5/6    | 0.818           | 0.140            | 0.601           | 0.251            |
| C6/7    | 0.877           | 0.769            | 0.628           | 0.733            |

Intraclass correlation coefficient (2, 1) was performed; values are calculated from variance analysis. DTI, diffusion tensor imaging; FA, fractional anisotropy; ADC, apparent diffusion coefficient; ZOOM, zonally oblique multislice.

| Table 5 | Intra-rater reliability between ZOOM and conventional DTI |
|---------|-----------------|
|         | FA              | ADC              |
|         | ZOOM            | Conventional     | ZOOM            | Conventional     |
|         | ICC (1, 1)      | ICC (1, 3)       | ICC (1, 1)      | ICC (1, 3)       |
| C2/3    | 0.960           | 0.986            | 0.821           | 0.932            |
| C3/4    | 0.618           | 0.963            | 0.915           | 0.970            |
| C4/5    | 0.949           | 0.982            | 0.970           | 0.990            |
| C5/6    | 0.942           | 0.980            | 0.922           | 0.973            |
| C6/7    | 0.724           | 0.887            | 0.677           | 0.863            |

ICC (1, 1) and ICC (1, 3) were performed; values are calculated from variance analysis. ICC, Intraclass correlation coefficient; FA, fractional anisotropy; ADC, apparent diffusion coefficient; ZOOM, zonally oblique multislice.
spinal level. High accuracy of the estimated value is desired for the quantitative analysis of DTI when we assess the fiber tracts. However, quantitative values, which depend on the examiner, have poor repeatability. In our study, ICC (2, 1) of the FA and ADC values obtained for ZOOM DTI was higher than 0.6 in all cervical spinal cord levels compared to that for conventional DTI; this suggests the stability of the measurements. Additionally, for intra-rater reliability, it is considered that a more accurate quantitative value can be obtained by increasing the number of measurements taken.

Variable artifacts often occur on neck MRIs.\textsuperscript{20} In particular, these artifacts are evident at lower spinal levels (C5–Th1). These are mainly derived from magnetic susceptibility and motion during imaging. Magnetic susceptibility artifacts are caused by magnetic field nonuniformity due to air in the lung apex and surrounding the complex figure of the neck level. Motion artifacts also result from swallowing movements. These effects also reflect the result of the distortion ratio. In echo planar sequences such as DTI, the distortion generally occurs in the phase encode direction. Therefore, the geometrical distortion ratio of the short axis was higher than that of the long axis. Further, the difference in distortion between ZOOM and conventional DTI was smaller at the lower spinal cord levels.

In other words, despite ZOOM DTI being an EPI sequence, its rFOV, based on non-coplanar SE and OVS, can be useful in obtaining images with less distortion and higher accuracy.

In ZOOM DTI, it is difficult to distinguish between white matter and gray matter at sites that show high distortion such as the lower cervical spinal cord; however, they can be distinguished in the upper cervical spinal cord.

Zonally oblique multislice DTI showed a higher SNR compared to conventional DTI, but conventional DTI uses the sensitivity encoding (SENSE) technique, which seems to cause the increase in noise.

Additionally, the variation in SNR is larger in ZOOM DTI; this is attributed to the influence of breathing and body shape of the subjects.

We compared the image quality of DTI between the two techniques and determined that the visibility with ZOOM DTI was superior to that with conventional DTI. Notably, image quality was lower at a lower cervical level in both techniques. In particular, scores of conventional DTI at C5/6 and 6/7 were remarkably lower. There was a significant difference between C2/3 and C6/7 on ZOOM DTI.

Non-coplanar SE and OVS would help to overcome the limitations of ZOOM DTI despite the EPI sequences. Spatial resolutions of DTI images would be important for accurate measurements. A previous report mentioned that misleading results were caused by the partial volume effect between white matter and CSF.\textsuperscript{21} In particular, in a small organ such as the spinal cord, fine resolution that can depict a clear boundary between the subject and the surrounding tissue would be preferable. Of note, fluctuation in FA and ADC values was strongly influenced by CSF. Therefore, contouring of the ROI is very important to obtain highly accurate quantitative values. Region of interest should be set strictly inside the spinal cord and enlarged maximally, while excluding the possibility of encasement of CSF by checking 3D ROI squares. In this study, reduction of the FA value at the C6/7 level implies the difficulty of excluding the CSF influence when setting the ROI around the apex of the cervical cord sagittal curvature. Cerebrospinal fluid involvement, susceptibility, and motion artifacts at lower levels, especially in C5/6 and 6/7, cannot be ignored even in ZOOM DTI. However, SE imaging ordinarily reduces susceptibility and motion artifacts compared to EPI, so ZOOM DTI was superior for acquisition of less distorted DTI than the conventional method. Recently, new DTI imaging modalities based on turbo SE were developed.\textsuperscript{22} This technique can potentially reduce susceptibility artifacts and distortion at the lower level of the spinal cord. The in-plane resolution of ZOOM DTI was 1 mm\textsuperscript{2}, whereas that of conventional DTI was 1.56 mm\textsuperscript{2} for a similar scan time. Zonally oblique multislice DTI did not require phase over-sampling such as no-phase warp to establish a small FOV. High-resolution assessment within the available scan time in a clinical setting could be achieved by ZOOM DTI. In future studies, a time reduction technique, such as multi-band SENSE\textsuperscript{23} may contribute to the shorter scan time of DTI.

\textbf{Conclusions}

This study revealed the advantage of ZOOM. DTI for the quantitative estimation of cervical spinal cord tracts. Zonally

\begin{table}[h]
\centering
\begin{tabular}{|c|c|c|}
\hline
 & Zoom & Conventional & \textit{P} value* \\
\hline
C2/3 & 21.07 ± 13.98 & 15.81 ± 5.896 & 0.6501 \\
C3/4 & 29.42 ± 24.90 & 18.97 ± 7.152 & 1 \\
C4/5 & 17.39 ± 17.43 & 19.89 ± 11.48 & 0.1306 \\
C5/6 & 31.45 ± 24.09 & 15.75 ± 6.873 & 0.2899 \\
C6/7 & 26.32 ± 22.55 & 17.04 ± 13.08 & 0.3643 \\
\hline
\end{tabular}
\caption{Mean and standard deviation of signal-to-noise ratio (SNR) at each cervical spinal cord level}
\end{table}

\begin{figure}[h]
\centering
\includegraphics[width=0.5\textwidth]{fig15}
\caption{Signal-to-noise ratio (SNR) of zonally oblique multislice (ZOOM) and conventional diffusion tensor imaging in 10 healthy volunteers.}
\end{figure}
oblique multislice DTI provides better visibility and high accuracy using a small FOV and a shorter practical scan time compared with conventional DTI. The ROI contour should not include the surrounding CSF; however, susceptibility and motion artifacts at lower cervical levels cannot be completely avoided even in ZOOM DTI. To attain further rapid, high-resolution DTI sequences, combined ZOOM DTI and recently introduced techniques such as turbo spin-echo (TSE)-DWI and multi-band SENSE are desirable.

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Conflicts of Interest

Tomoyuki Okuaki is an employee of Philips Healthcare. The other authors declare that they have no conflicts of interest.

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