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Advances in MR imaging for cervical spondylotic myelopathy

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Journal

European Spine Journal, 24(Suppl 2)

ISSN

0940-6719

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Publication Date

2015-04-01

DOI

10.1007/s00586-013-2915-1

Peer reviewed

Advances in MR imaging for cervical spondylotic myelopathy

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Received: 9 December 2012 / Revised: 22 May 2013 / Accepted: 12 July 2013 / Published online: 6 August 2013
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Abstract

Purpose To outline the pathogenesis of cervical spondylotic myelopathy (CSM), the correlative abnormalities observed on standard magnetic resonance imaging (MRI), the biological implications and current status of diffusion tensor imaging (DTI), and MR spectroscopy (MRS) as clinical tools, and future directions of MR technology in the management of CSM patients.

Methods A systematic review of the pathogenesis and current state-of-the-art in MR imaging technology for CSM was performed.

Results CSM is caused by progressive, degenerative, vertebral column abnormalities that result in spinal cord damage related to both primary mechanical and secondary biological injuries. The T2 signal change on conventional MRI is most commonly associated with neurological deficits, but tends not to be a sensitive predictor of recovery of function. DTI and MRS show altered microstructure and biochemistry that reflect patient-specific pathogenesis.

Conclusion Advanced imaging techniques, including DTI and MRS, show higher sensitivity to microstructural and biochemical changes within the cord, and may aid in management of CSM patients.

Keywords Cervical · Spinal cord · DTI · MRI · Myelopathy

Introduction

Since its advent, magnetic resonance imaging (MRI) has played an indispensable role in the management of patients with cervical spondylotic myelopathy (CSM). There has been tremendous advancement in MR technology over the past several decades, resulting in enhanced resolution and image quality that is manifold greater than previous generations. As the image quality has increased, the application of MRI to CSM has progressed in parallel, evolving from purely a diagnostic modality to a non-invasive tool that can be used to potentially predict neurological outcome and response to intervention.

In addition to conventional MRI, recent application of novel imaging techniques such as diffusion tensor imaging (DTI) [1–5] and magnetic resonance spectroscopy (MRS) [6] to CSM further highlights the potential impact of MR technology on this debilitating disease process. By providing pertinent information regarding the spinal cord microstructure and metabolism, these novel techniques provide increased sensitivity to the spinal cord injury and derangement of cellular function that ubiquitously occurs during CSM pathogenesis.

This manuscript seeks to examine the application of MR technology to the management of CSM patients. Recent and future advances in both conventional and novel MR techniques will be discussed.

CSM pathophysiology

In order to fully appreciate the importance of MRI in the management of CSM, as well as the significance of its radiographical findings, an understanding of the complex CSM pathophysiology is necessary. CSM is caused by

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progressive degenerative vertebral column abnormalities that result in spinal cord damage related to both primary mechanical and secondary biological injury. Primary mechanical spinal cord injury can be initiated by static and dynamic forces including compression, distraction, and shear [7] (Fig. 1). Direct spinal cord compression is the most frequently encountered mechanism, and it has been theorized that myelopathic symptoms appear after the

spinal cord has been reduced in size by 30 % or to a transverse area of less than 60 mm² [8]. Secondary spinal cord injury in CSM is likely related to a variety of mechanisms such as glutamergic toxicity, free radical-mediated cell injury, cationic-mediated cell injury, and apoptosis [9–11]. Ischemia is considered to be a significant contributor to the pathophysiology of CSM. Ventral compression impairs perfusion via the transverse arterioles originating

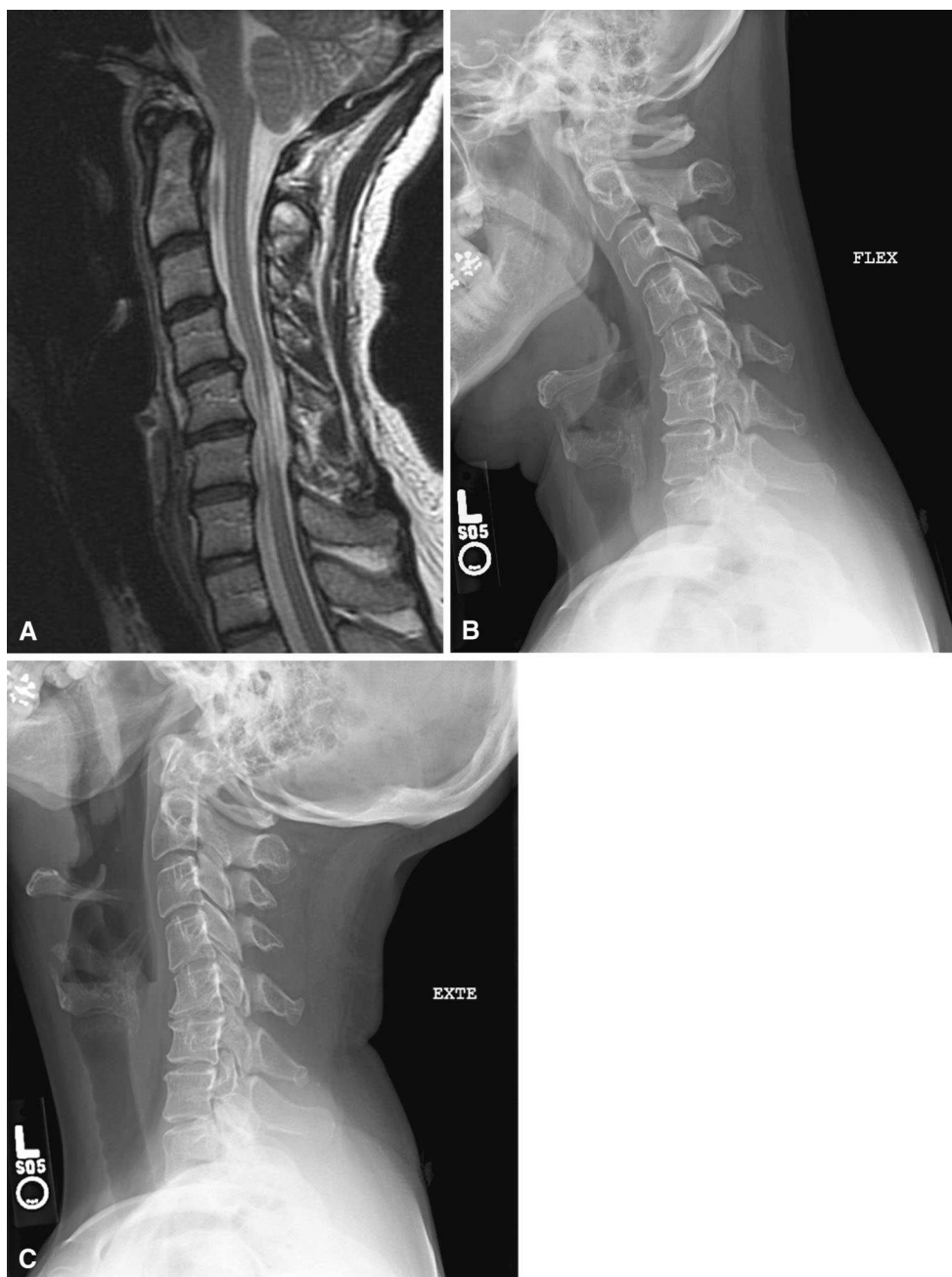


Fig. 1 Cervical spondylotic myelopathy. **a** Cervical spine MRI of a 45-year-old woman with CSM demonstrating extensive spinal cord atrophy and T2 weighted signal change at the C5 and C6 levels

despite the absence of frank spinal cord compression. This spinal cord damage was caused by repetitive shear injury across the ventral disc bulges during flexion (**b**) and extension (**c**)

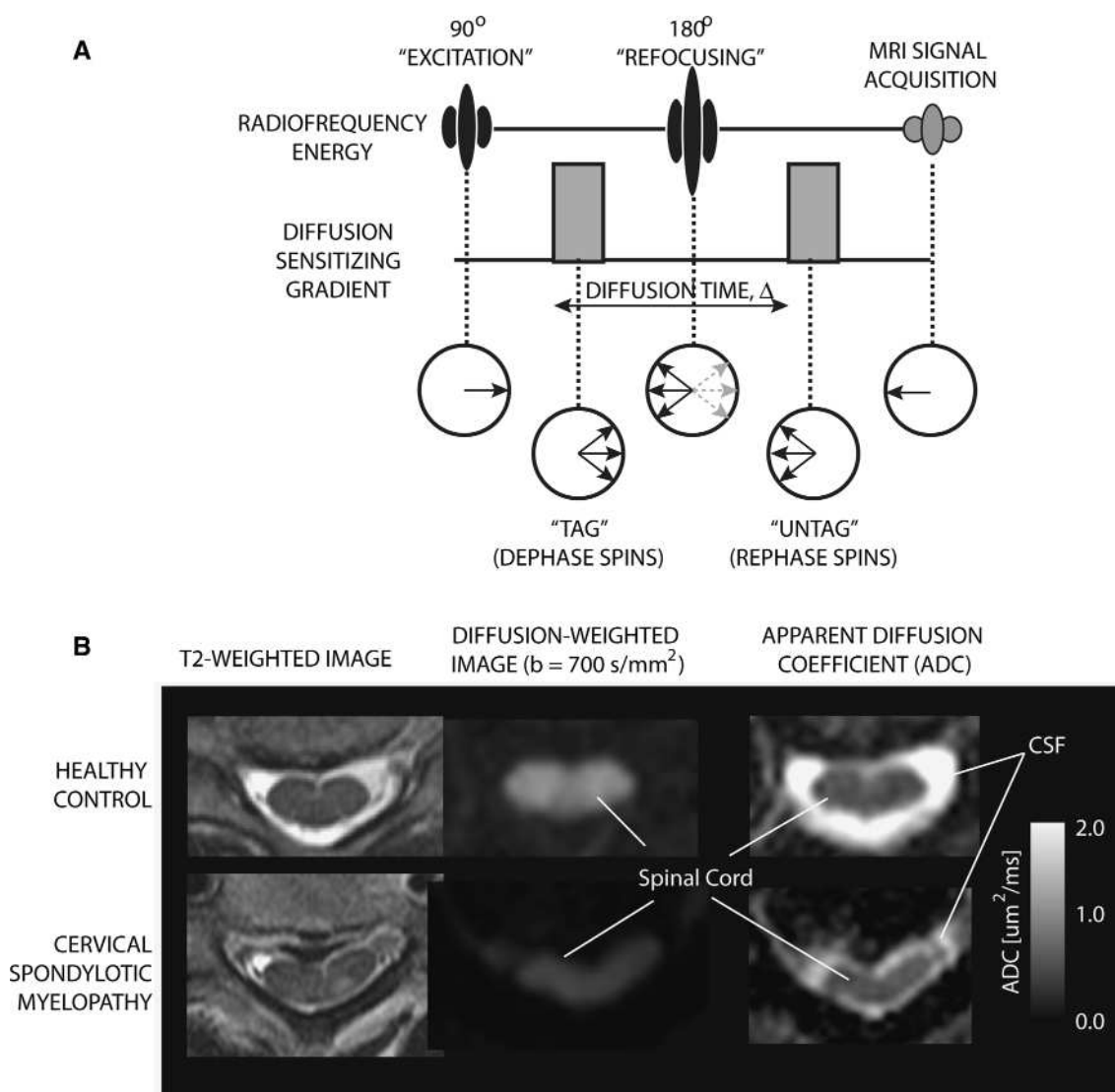


Fig. 2 Diffusion-weighted imaging (DWI) of the spinal cord. **a** The pulse sequence diagram (*top*) and water hydrogen spin dynamics (*bottom*) during a one-dimensional DI experiment. The use of pulsed “diffusion sensitizing” magnetic field gradients tags the water hydrogens by dephasing them at one time point and then untags them by rephrasing them at a later time point. Water hydrogens that are stationary will completely rephase and lead to maximum signal intensity on DWIs. Water hydrogens that are moving (i.e. diffusion) will have incomplete rephasing resulting in loss of signal amplitude by $S = S_0 e^{-bADC}$, where S is the MRI signal amplitude (image

intensity) with diffusion weighting, S_0 is the MRI signal amplitude (image intensity) without diffusion weighting, b is the “ b value” or the level of diffusion weighting (dependent on the gradient amplitude, duration, etc.), and ADC is the apparent diffusion coefficient. **b** Example T2-weighted (*left*), diffusion weighted (*middle*), and ADC maps (*right*) for a healthy control volunteer (*top*) and patient with cervical spondylotic myelopathy (*bottom*). Note that at the site of chronic compression, the signal intensity on DWIs is decreased and ADC is increased relative to the normal spinal cord

from the anterior sulcal arteries, whereas posterior vascular compromise can occur in the intramedullary branches of the central grey matter [12].

Conventional MRI

Prior to the development of MRI, computed tomography (CT) with or without myelography was the mainstay of the

radiographical evaluation of CSM patients. Although these modalities offered useful anatomical information regarding the osseous vertebral column and degree of spinal canal stenosis, only limited information could be directly ascertained about the condition of the spinal cord. As such, there were definite limitations in the ability for imaging to adequately assess the spinal cord damage in this patient population.

Through its ability to provide high resolution imaging of soft tissue anatomy, MRI bridged this knowledge gap,

providing excellent anatomical detail of the spinal cord macrostructure. MRI has granted insight into the aforementioned structural histopathological changes that occur during CSM pathogenesis. The elucidation of T2 and/or T1 weighted signal change by MRI technology has spawned a plethora of clinical research studies that have intensively investigated their significance. These findings are attributed to myelomalacia, edema, gliosis, and ischemic white matter changes. Some authors suggested that these observations result in irreversible spinal cord injury [13], whereas others feel that these regions represent a wide spectrum of recuperative potential [14]. There have even been several grading systems proposed to classify the spinal cord signal change subtypes [15, 16]; however, this remains one of the most controversial and actively debated topics in the area of degenerative spine disease.

There is great interest in assessing the utility of MRI to predict neurological outcome following decompression surgery for CSM. The reasons for this are many. Operative management may result in clinical improvement, worsened clinical function, or preservation of function without clear improvement or decline. Patients and their families commonly inquire about whether surgery will alleviate their symptoms and/or return lost neurological function. Surgery can involve technically challenging decompression and stabilization procedures, and the complication rate is not insignificant, particularly in the elderly. This, combined with the fact that a subset of CSM patients are likely to stabilize clinically without surgery [17], increases the importance of prospectively determining which patients are most likely to benefit from surgical intervention.

An expert group representing the Joint Section on Disorders of the Spine and Peripheral Nerves of the American Association of Neurological Surgeons and Congress of Neurological Surgeons published the Guidelines for the Surgical Management of Cervical Degenerative Disease in 2009 [18]. One of the highlights of this seminal work was their findings regarding the relationship between spinal cord signal change and clinical outcome in CSM patients. Based on their thorough and exhaustive literature review, they concluded that multilevel T2 hyperintensity, T1 focal hypointensity combined with T2 focal hyperintensity, and spinal cord atrophy each convey a poor prognosis following surgical intervention.

In addition to these factors, there is also some evidence that regression of T2 signal change following surgical intervention is associated with a better prognosis [19, 20]. From a theoretical standpoint this makes sense, as this phenomenon implies a healing of the injured spinal cord. This concept is at least in part supported by histopathological studies in which denuded axons and numerous thin myelinated fibers have been found in the damaged spinal cord segments in patients with CSM, suggesting that focal

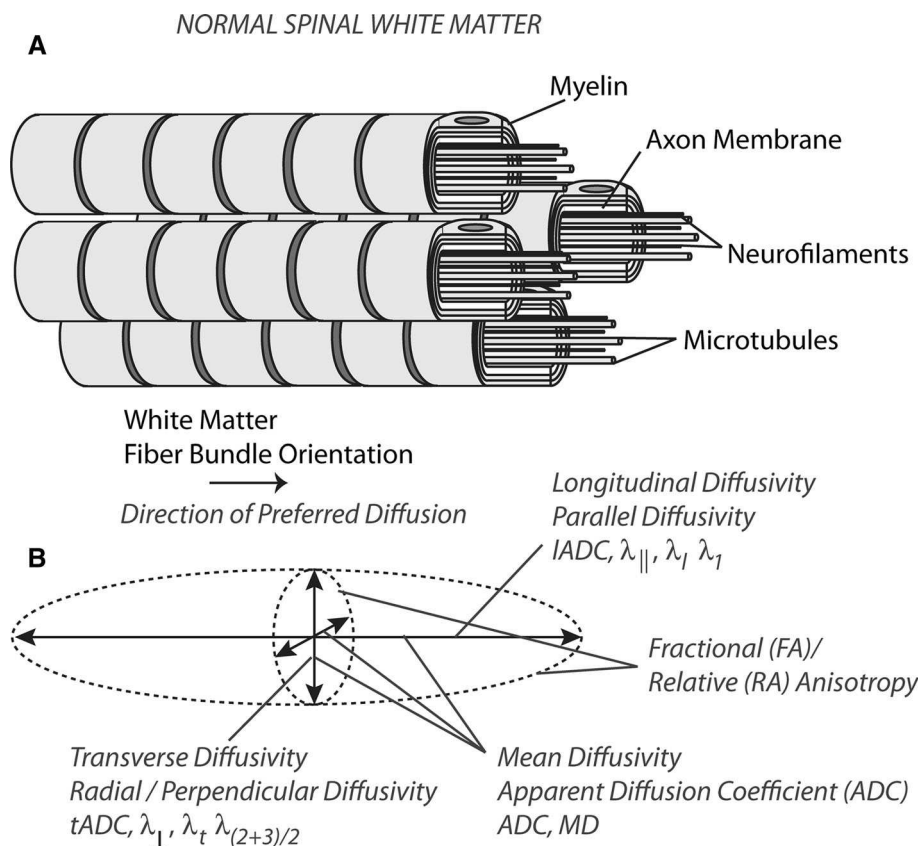
demyelination and remyelination can occur within the white matter [21]. It has been postulated that this process may be an important mechanism in the recovery of lost neurological function following therapeutic management of this disorder.

Diffusion tensor imaging (DTI)

Diffusion-weighted imaging (DWI) is a magnetic resonance imaging (MRI) technique that can be used to measure microscopic random movement of water molecules (i.e. diffusion). Specifically, the same magnetic field gradients used to create the MR images can be implemented in such a way that water molecules are “tagged” (or “dephased”) at one time point, then “untagged” (or “rephased”) at a slightly later time point (Fig. 2a). The random movement of water molecules during the time of tagging and untagging, termed the “diffusion time”, results in signal attenuation in the MR image that is proportional to the apparent diffusion coefficient (ADC), a measure of water diffusion magnitude or simply the “rate” of water diffusion. In other words, the faster water molecules are diffusing within a tissue, the larger the signal attenuation in DWIs, and the larger the ADC. Hence, tissues with high water mobility and few boundaries to water motion, such as regions containing cerebrospinal fluid (CSF) and vasogenic edema, have high ADC values whereas tissues with a high degree of complexity and boundaries to diffusion, such as white matter fiber bundles or tumors, have a relatively lower ADC (Fig. 2b). It is the sensitivity to microstructural complexity below the resolution of traditional MRI techniques that makes diffusion MRI particularly attractive for interrogating the health of the spinal cord in patients with cervical myelopathy.

The standard DWI experiment involves measuring the ADC in a single direction (e.g. from left-to-right or from anterior-to-posterior), or averaging ADC measured in three orthogonal directions. In tissues with a preferred orientation, such as muscle or nerves where fibers are oriented in a similar direction, ADC varies according to the direction of measurement. In the central nervous system (CNS), diffusion measurements perpendicular or transverse to the direction of white matter tracts are lower than diffusion measurements that are parallel to the orientation of white matter tracts, partially due to a larger number of boundaries to diffusion in the transverse orientation including myelin, the axon membrane, neurofilaments, and microtubules (Fig. 3). In addition, extracellular water is also confined to diffusing parallel to the orientation of the fiber bundles. To describe the diffusion characteristics of directionally dependent tissues, diffusion tensor imaging (DTI) was developed. DTI is a

Fig. 3 Microstructure and diffusion characteristics of healthy spinal white matter. **a** Model of axon bundles showing boundaries to diffusion in the transverse orientation, including myelin, axon membranes, neurofilaments, and microtubules. **b** The diffusion ellipsoid model derived from the traditional diffusion tensor shows anisotropic diffusion with preferred diffusion in the direction of white matter fiber bundles, with lower ADC in the transverse orientation and higher ADC in the longitudinal orientation



diffusion MRI technique that can be used to quantify the magnitude and direction of water diffusion with respect to a pre-defined coordinate system [22, 23] (Table 1). More specifically, if diffusion MRI measurements are made in various directions (i.e. more than six directions), then a mathematical “diffusion tensor” can be constructed, which consists of a diffusion matrix defining the tensor field of ADC values. The preferred magnitude and direction of diffusion are then found from decomposing the diffusion tensor into its eigenvalues and eigenvectors, respectively. The largest eigenvalue, termed the “primary eigenvalue”, “longitudinal ADC”, IADC, λ_1 , or $\lambda_{||}$, forms the long axis of a “diffusion ellipsoid” used to model the diffusivity and is typically oriented parallel to the direction of white matter fibers (Fig. 3). The two smallest eigenvalues, termed the “secondary and tertiary eigenvalues”, “transverse ADC”, tADC, or λ_{\perp} form the short axes of the “diffusion ellipsoid” and are typically oriented perpendicular to the direction of white matter fibers (Fig. 3). Thus, the degree of difference between longitudinal and transverse ADC measurements determines the degree of diffusion anisotropy within an image voxel or tissue (e.g. CSF has isotropic diffusivity, thus IADC = tADC; whereas white matter has a high degree of diffusion anisotropy thus IADC > tADC). It is these features of

the diffusion tensor that allow inference as to the microstructural integrity of various tissues [24, 25].

Various diffusion tensor “stains”, or image contrasts, have been developed to simplify the information present in the diffusion tensor. The “fractional anisotropy”, or FA, ranges from 0 (isotropic) to approximately 1 (anisotropic) and is the most common diffusion tensor index used for quantifying the degree of diffusion anisotropy [26]. Other lesser used anisotropy metrics include the relative anisotropy (RA), volume ratio (VR), measured anisotropy (MA) [27], along with circular or spherical invariants [26, 28]. Thus, changes in overall water diffusivity (e.g. mean ADC or mean diffusivity, MD), changes in directionally dependent ADC measurement (e.g. IADC and tADC), and/or changes in water diffusion anisotropy (e.g. FA) all provide information regarding the microstructural integrity within the spinal cord [24, 25].

DTI in cervical spondylotic myelopathy

Cervical spondylotic myelopathy (CSM) is a chronic progressive disorder of the spinal cord with a relatively ill-defined onset of pathogenesis when compared with other spinal disorders. Although little is known about the diffusion characteristics within human spinal cord during the

Table 1 Clinical and biological interpretation of quantitative DTI metrics

DTI metric	What it measures/clinical interpretation
ADC (apparent diffusion coefficient), MD (mean diffusivity)	Average magnitude of water mobility in all directions General size of extracellular space Increases with increased vasogenic edema Decreases under ischemic events Decreases during acute compression Increases during chronic compression due to axon degeneration
FA (fractional anisotropy)	Diffusion anisotropy (i.e. the degree to which ADC varies with respect to orientation) High in normal spinal cord white matter due to tightly packed axons in a specific orientation Increased slightly during acute compression Decreased during chronic compression due to axon degeneration
tADC (transverse ADC), λ_{\perp} , RD (radial diffusivity)	ADC magnitude perpendicular to spinal cord orientation (i.e. transverse orientation) Low in normal spinal cord white matter due to transverse boundaries to water diffusion (e.g. myelin, axonal membranes, neurofilaments) Decreases during acute compression Increases during axonal degeneration and demyelination
lADC (longitudinal ADC), λ_{\parallel} , AD (axial diffusivity)	ADC magnitude parallel to spinal cord orientation (i.e. rostral-caudal orientation) High in normal spinal cord white matter Decreases with decreasing neurological function
RA (relative anisotropy) VR (volume ratio) MA (measured anisotropy)	Other (less common) measures of diffusion anisotropy

early phases of spinal cord compression prior to progressive myelopathy, the literature suggests that acute compression of spinal cord tissue may result in a focal decrease in ADC as well as a focal increase in FA (Fig. 4). Assuming spinal cord axons can be modeled as infinite cylinders, diffusion MR simulations by Ford et al. [29, 30] suggest that compression axon fibers will result in a decrease in transverse ADC, which may lead to a slight increase in FA. Additional diffusion MR simulations by Nilsson et al. [31] clearly demonstrate the decreased mean ADC with increasing compression of spinal cord white matter. Facon et al. [3] explored acute spinal cord compression in two patients and noted a slightly elevated FA at the level of compression as compared to normal controls (0.80–0.83 vs. 0.75 in healthy volunteers). Similarly, Hernandez et al. [4] examined 12 patients with CSM and noted a focal increase in FA and decrease in ADC at the site of compression, which was also observed in a pilot study by Benae et al. [32] and attributed to compression of spinal tissue resulting in a focal increase in the boundaries to diffusion in the transverse direction. In perhaps, one of the largest studies of DTI in CSM, Mamata et al. [33] described a subset of CSM patients (46 %) having either normal-to-decreased ADC, or normal-to-elevated FA, at the site of compression and described these patients as having few clinical symptoms other than arm or neck pain, likely related to focal compression of the spinal cord.

Progressive, chronic compression of the spinal cord results in many pathological changes that are detectable using diffusion MRI (Fig. 5). Specifically, clinical studies have clearly documented a significant increase in ADC and decrease in FA in occur the late stages of chronic compression of the spinal cord [1–3, 33–46]. These changes have also been verified using an animal model of chronic compression [47], illustrating the characteristic increase in ADC and decrease in FA as late as 9 months after the start of compression. The specific changes in diffusion MR characteristics observed at this alter stage of injury closely resemble those observed in chronic spinal cord injury (SCI) from trauma, albeit to a lesser degree [48, 49]. Specifically, the changes in diffusion characteristics are likely to be a result of chronic, repeated ischemic insults to the spinal cord leading to downstream histopathological changes (Fig. 5) including gliosis, loss of motoneuron function, vasogenic edema, and ultimately necrosis and cavitation [50–55]. These pathological changes, together, result in elevation of ADC due to the increase in extracellular water and suppression of FA due to lack of directional organization within the cord.

An increase in mean ADC typically accompanies a decrease in FA at the site of chronic stenosis, however, empirical evidence suggests that mean ADC and FA may vary in their clinical and biological significance. For example, Uda et al. [34] implemented receiver-operator

Fig. 4 Microstructure and diffusion characteristics of acute spinal cord compression.

a During acute compression of spinal cord white matter fibers are compressed and blood flow to these regions is reduced, resulting in transient ischemia. **b** Transient, acute compression of the spinal cord results in compression of the diffusion ellipsoid, resulting in lower ADC and, in some cases, localized increases in FA values relative to adjacent spinal segments

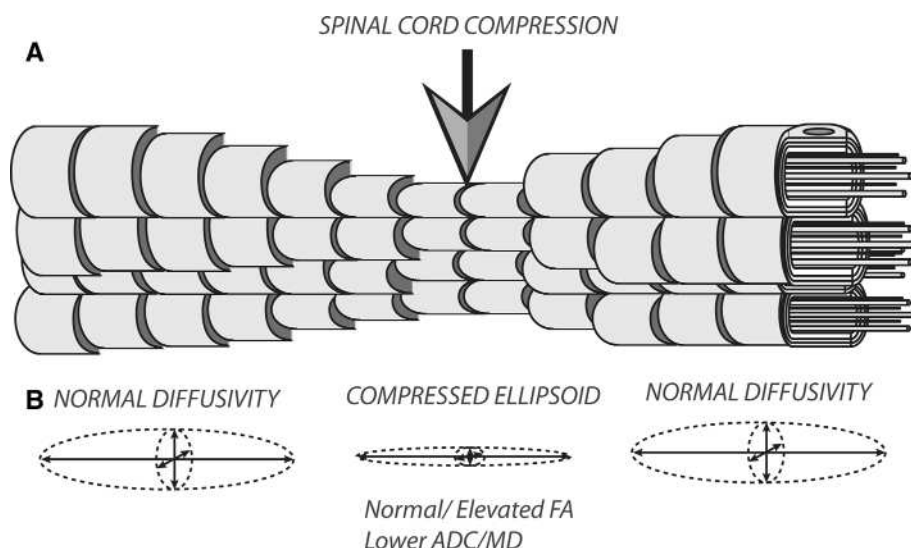
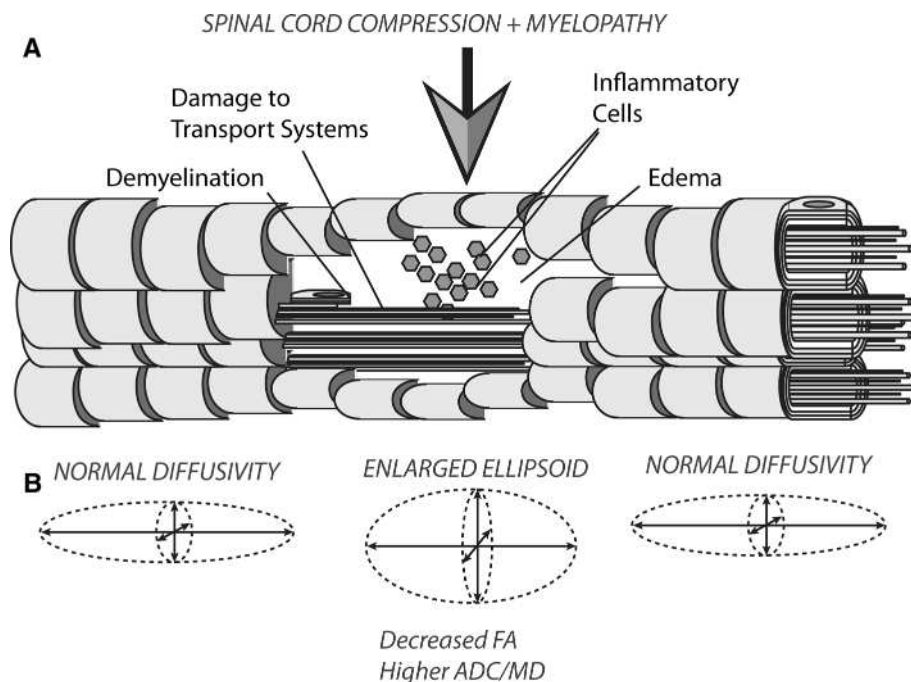


Fig. 5 Pathological changes and diffusion characteristics in cervical spondylotic myelopathy.

a After chronic compression of the spinal cord resulting in sustained ischemia, white matter tracks eventually degenerate, resulting in demyelination, damage to axon transport systems, the influx of inflammatory cells, and vasogenic edema, which eventually leads to functional impairment. **b** ADC increases and FA decreases in the site of chronic compression due to the increased extracellular water concentration and decreased fiber tract density, respectively



characteristic (ROC) analysis and determined the mean ADC as the best predictor of myelopathy when compared with FA (ROC area under the curve, AUC = 0.903 for mean ADC and AUC = 0.760 for FA). Similarly, data from Demir et al. [2] suggested nearly an 80 % sensitivity and 53 % specificity for detecting myelopathy in patients with spinal cord compression. On the other hand, FA appears to be significantly correlated with specific clinical outcomes, including a positive linear correlation with the modified Japanese Orthopedic Association score (mJOA) [5] and a clinical pain assessment via self questionnaire [38], and a negative linear correlation with Nurick score [5]. A recent study by Jones et al. [5, 56] also showed that patients with a higher pre-operative FA at the site of compression appeared to have

better functional recovery when compared to patients with lower pre-operative FA measurements, further implicating mean FA at the site of compression as being a potential biomarker for determining favorable surgical candidates. Consistent with this hypothesis, other studies have demonstrated that a significantly higher ADC and significantly lower FA occur in symptomatic when compared to asymptomatic patients with spinal cord compression [37, 39]. Comparatively, the presence and degree of T2 hyperintensity observed within the spinal cord was not shown to be predictive of neurological impairment [5, 38], demonstrating a mere 60 % sensitivity but more than 90 % specificity for detecting myelopathy in patients with spinal cord compression [2].

Deflection of the primary eigenvector orientation derived from DTI data has also been examined as a potential biomarker for clinical impairment. A study by Song et al. [1] illustrated that nearly 75 % of patients with CSM had disruption in the orientation of the primary eigenvalue as illustrated with FA color maps (Fig. 6). Similarly, Cui et al. [57] noted an increase in primary eigenvector orientational entropy, or randomness, in patients with CSM when compared with healthy controls. Consistent with this observation, DTI tractography, a method that uses the direction of the primary eigenvector to create “pseudo”-axonal tracts [58, 59], CSM clearly defines the site of compression in patients with CSM (Fig. 6).

MR spectroscopy

In comparison to DTI, the application of MRS to CSM has been significantly understudied, but has some intriguing

potential. MRS offers metabolic information regarding cellular biochemistry and function of the neural structures within the cervical spine. MRS can be used to assay a number of pertinent biochemical markers, including *N*-acetyl aspartate (NAA), lactate, choline (CHO), and creatine (Cr). Of particular interest are NAA and lactate. NAA is found almost exclusively in axons and neurons, and is considered an indicator of axonal integrity. Although not completely understood, lactate is considered to play a central role in metabolic dysfunction after central nervous system injury, presumably related to ischemia and mitochondrial dysfunction.

Presently, spinal cord MRS is most frequently used to investigate multiple sclerosis (MS) lesions. These studies have demonstrated the suppressed levels of NAA in MS patients when compared with normal volunteers [60, 61], and some correlation between NAA levels and clinical

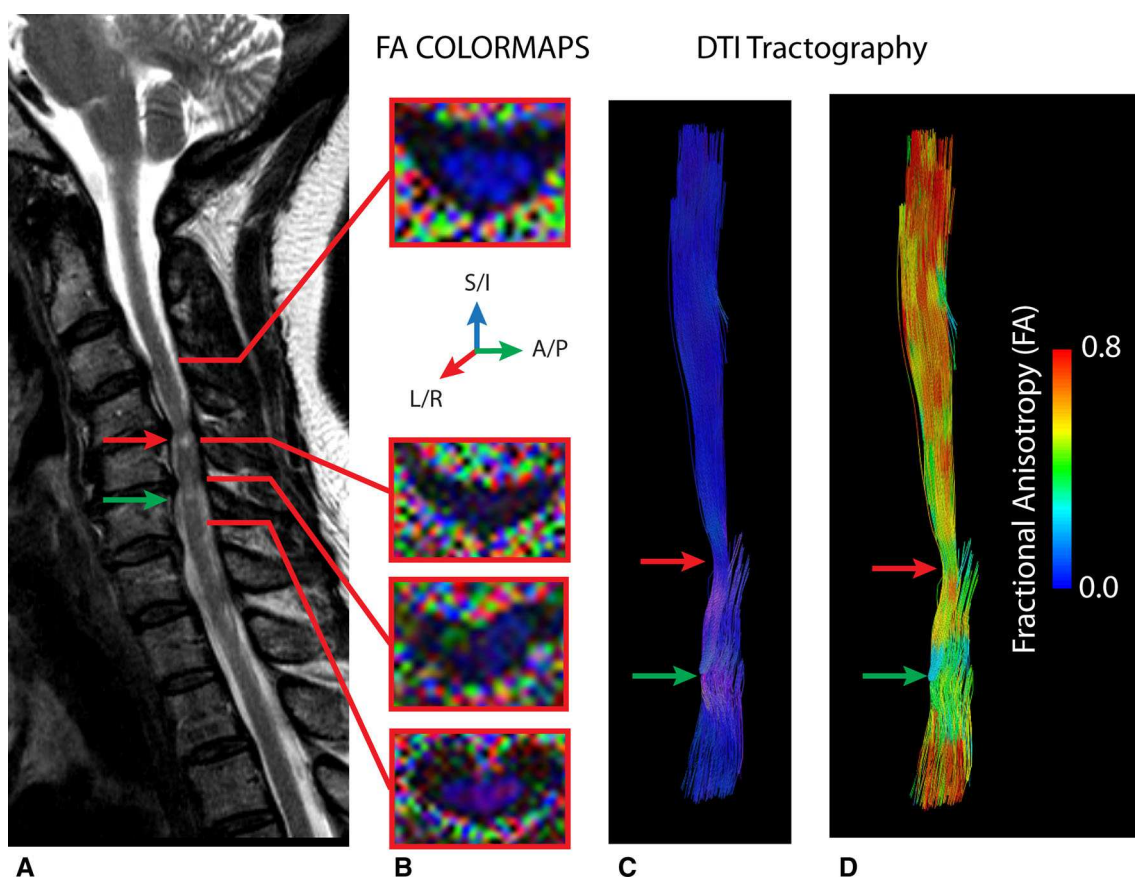
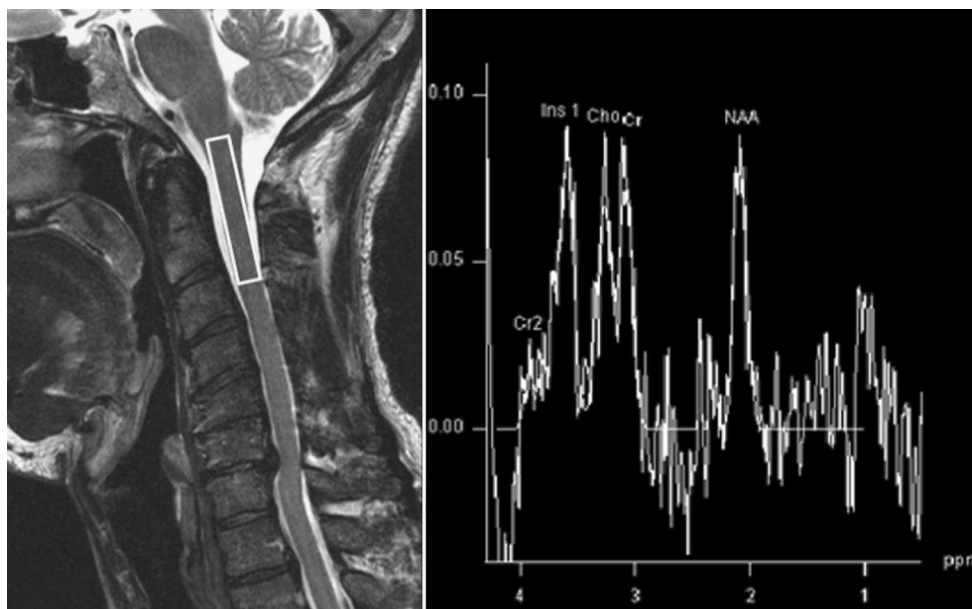


Fig. 6 Disruption of fiber orientation in CSM. **a** Sagittal T2-weighted image of a 65-year-old-female patient with CSM showing T2 hyperintensity at the site of most severe spinal cord compression. **b** FA colormaps showing the orientation of the primary eigenvector (blue superior/inferior, green anterior/posterior, red left/right) in the normal-appearing spinal cord (top image at C3–4) and regions of compression (bottom images). Note that in the areas of compression, the FA is low (dark) and the orientation of the primary eigenvector is deflected from the superior/inferior (blue) orientation, resulting in

other colors appearing within the spinal cord. **c** DTI tractography labeled (colored) with respect to the primary eigenvector orientation shows deflection (pink regions) through the area of compression. **d** DTI tractography labeled (colored) with respect to fractional anisotropy (FA) shows a decrease in FA at the site of primary eigenvector deflection. Red arrows showing area of significant compression resulting in T2 hyperintensity. Green arrow showing second area of compression caudal from the first lesion

Fig. 7 MRS obtained with voxel at the C2 level in a 60-year-old gentleman with CSM. The MRS spectra demonstrate a low NAA/Cr ratio indicative of axonal and neuronal injury



status. Although less studied, cervical spine MRS has recently been applied to CSM. In a cohort of 22 CSM patients, Holly et al. [6] found that the NAA/Cr ratio was significantly lower in CSM patients than in normal volunteers (Fig. 7), suggesting the increased axonal and neuronal injury in these patients. An abnormal lactate signal was present in nearly one-third CSM patients and none of the control subjects, further supporting the role of ischemia in the pathogenesis of CSM.

Limitations of DTI and MRS

Although these advanced imaging techniques offer novel insights into CSM, there are some limitations. The spinal cord is relatively small, and differences in magnetic susceptibility of adjacent tissues can cause contamination. The cross-sectional area of the spinal cord is only slightly larger than the minimum voxel size commonly used for MRS, and a suboptimally placed voxel can cause a significant decrease in signal-to-noise ratio [62]. MRS and DTI are very sensitive to patient or structural movement during scan acquisition. The physiological rostral-caudal movement of the spinal cord in response to cardiac pulsations and the respiratory cycle is fairly significant, and even more marked than in the brain [62]. Thus, cardiac gating, specialized radiofrequency coils, and MR signal suppression bands are frequently used to enhance the quality of both MRS and DTI in the spinal cord.

Future directions

In addition to predicting outcome following surgical intervention, there are several other potential future

applications for MR technology in the management of CSM patients. The medical literature is clear that some patients with mild CSM can be successfully treated non-operatively. Advanced MRI techniques such as DTI or MRS could potentially serve as a non-invasive method to monitor asymptomatic or mildly affected patients treated nonoperatively for impending neurological deterioration. Such an early warning system does not currently exist, as serial standard MR imaging is likely to demonstrate a stable radiographical appearance despite the progression of cellular spinal cord injury and subsequent neurological symptomatology. As such, in conjunction with other non-operative treatment techniques such as physical therapy and cervical collar placement, asymptomatic and mildly affected CSM patients under observation could undergo serial DTI and/or MRS to assess for subclinical disease progression.

There has been an increasing interest in the investigation of novel molecular and biochemical therapies to treat the secondary biological injury that occurs during CSM pathogenesis. Some of the laboratory investigations include the inhibition of cell apoptosis with a Fas ligand blocking antibody [63], the administration of neurotrophins either through genetically altered fibroblasts [64] or adenovirus-mediated retrograde spinal cord delivery [65], and dietary therapy to repair injured plasma membranes and cellular oxidative damage [66]. Once translated to clinical use, irrespective of the exact neural repair strategy, these therapies will require a non-invasive modality to ascertain cellular response to intervention. DTI and/or MRS could potentially serve such a modality, as both techniques can identify subtle changes in spinal cord microarchitecture and biochemistry, unlike conventional MRI. In fact, DTI

has been used to assess spinal cord regeneration following epidermal neural crest stem cell grafting in an SCI model [67]. In this study, DTI demonstrated decreased mean diffusivity and increased diffusion anisotropy in study animals, indicating functional and structural improvement of the spinal cord.

In addition to DTI and MRS, other novel MR technologies may provide further insight into spinal cord health. Techniques such as q-space imaging, chemical exchange saturation transfer (CEST) imaging and magnetization transfer imaging (MTI), dynamic susceptibility contrast (DSC) perfusion MRI, dynamic contrast enhanced (DCE) perfusion MRI, arterial spin labeling (ASL) and spinal cord fMRI all hold promise for providing important physiological information about the status of the spinal cord. Q-space imaging, a diffusion MRI technique that provides quantitative information about the physical size of microstructural compartments, has been successfully performed *ex vivo* in spinal cord tissues to study the evolution of traumatic injury [68–71] and recently been explored in human CSM patients [35]. CEST, amide proton transfer (APT), and MT imaging are quantitative spectroscopic molecular imaging techniques that can be used to study the exchange rates between various biochemical species [72]. Specifically, CEST involves quantification of biochemical species by taking advantage of off-resonance excitation of hydrogen atoms on exchangeable proteins or lipids. Only a few studies have been performed using CEST or APT within the spinal cord [73], but MTI has been very useful for studying MS and amyotrophic lateral sclerosis (ALS) since it can be optimized to quantify myelin integrity [74, 75]. Perfusion imaging techniques such as DSC, DCE, and ASL have yet to gain traction as a tool for assessing spinal cord health; however, they are likely to play an important role in assessing CSM in the future, since spinal cord ischemia and hypoxia are thought to exacerbate spinal cord injury. In addition, future studies will likely also focus on quantification of neuronal function using spinal cord fMRI to detect functional impairment and to localize regions of injury [76]. This technique, however, has yet to be tested in CSM patients.

Conclusion

MRI plays an integral part in the management of CSM patients, and has evolved from primarily a diagnostic modality to one that can help prognosticate patient outcome following surgical intervention. Advanced imaging techniques such as DTI and MRS provide novel information regarding the spinal cord microstructure and biochemistry, which have further enhanced our knowledge base regarding the pathogenesis and cellular derangement encountered in

CSM. It is likely that in the future, these MR techniques and others will play an expanded role in the management of CSM patients treated with and without surgery.

Acknowledgments Research support: NIH 1R21NS065419-01A1 and 1R01NS078494-01A1.

Conflict of interest None.

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