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Radiographic Analysis of Cervical and Spinal Alignment in Multilevel ACDF with Lordotic Interbody Device

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Abstract

Background
Restoration and maintenance of cervical lordosis is an important clinical parameter in spine surgery. The purpose of this study was to determine the extent to which a multilevel anterior cervical discectomy and fusion (ACDF: greater than 3 levels) procedure restores cervical lordosis and the affect of increasing lordosis on sagittal vertical axis.

Methods
We performed a retrospective radiographic analysis of 69 patients who underwent multilevel ACDF by 2 surgeons between 2013 and 2014. We measured the global and segmental sagittal alignment of the cervical spine using the Cobb method at 4 time intervals (preop, post op 4wks, 10wks and 6 months) as well as the sagittal vertical axis (SVA) using both a C1-S1 and C7-S1 plumb line methods at 2 time intervals (preop and post op 4wks). Radiographs were measured by three reviewers.

Results
Interrater reliability was good to excellent for all measurements. Cervical lordosis significantly increased from pre-op 10.26° to 4 weeks postop 19.44° and was maintained up to 6 months 19.34° (p<0.0005). Segmental cervical lordosis was also significantly increased from preop 8.22° to post op at 4 weeks (20.26°) and was maintained at post op 10weeks 20.30° and post op 6 months 19.56° (p<0.0005). C7-S1 SVA and C1-S1 SVA also significantly increased from 12.04mm preop to 27.49mm post op 4 wks (p<0.0005) and -1.93mm preop to 8.67mm post op (p<0.0005) respectively. A change in C2-C7 lordosis positively correlated with a change in C7-SVA and C1-SVA (r=0.37, P<0.005, and r=0.312, p<0.05 respectively).

Conclusions
Multilevel ACDF significantly increases and maintains both segmental and global cervical lordosis up to 6 months after surgery. Increasing C2-C7 global lordosis is correlated with increasing positive sagittal vertical axis. Level of evidence: IV.

Introduction
Overall spinal sagittal alignment—spinal tilt in the anterior posterior dimension—is determined by the delicate configuration of the four sagittal plane curves: the two kyphotic thoracic and sacral curves, and the two lordotic lumbar and cervical curves.1 While our understanding of the mechanical interplay between these spinal segments is still growing, alterations in the alignment or relationship between these curves can have profound impact on load biomechanics, balance, and pain. Overall spinal sagittal alignment is represented by the sagittal vertical axis (SVA), which is the distance from the posterior superior corner of the sacrum to a plumb line from either C1 (centroid of the C2 vertebra) or C7 on a lateral full-length radiograph. Neutral sagittal alignment C7-SVA is highly conserved in normal adults within ±2.5cm of the posterior superior corner of the sacrum.2 Anterior offset of the C7-SVA >5cm has
been linked with increasing pain and disability.\textsuperscript{3,5}

In the cervical spine normal sagittal alignment is variable, but typically maintained at 20-35° lordosis as measured by the Cobb angle between the superior endplate of C2 to the inferior endplate of C7.\textsuperscript{4} Abnormal kyphosis in the cervical spine increases cantilever loads from the center of gravity of the head resulting in an abnormal stress environment within the spine.\textsuperscript{5,8} Chavanne et al. showed in their cadaveric study that increasing kyphosis results in increased intramedullary pressure in the spinal cord.\textsuperscript{9} Moreover, Shimizu et al showed that progressive kyphosis resulted in demyelination and neuronal loss in a animal model.\textsuperscript{10} In clinical studies, increasing kyphotic alignment of the cervical spine has been correlated with worse patient outcomes.\textsuperscript{11-13} For this reason, restoration of neutral cervical lordosis is desired in cervical degenerative surgery.\textsuperscript{14}

The interplay between spinal segments is complex, but there is increasing agreement that fusion of one segment of the spine leads to reciprocal changes in other segments.\textsuperscript{15} Klineberg et al reported reciprocal changes between the thoracic and lumbar spine with increasing lumbar lordosis corresponding with increased thoracic kyphosis and in a separate paper that reduction of thoracic kyphosis reduces lordosis.\textsuperscript{16} While anterior cervical plating has been previously shown to increase and maintain cervical lordosis in single and multilevel degenerative disease,\textsuperscript{17,18} there have been no studies assessing the impact of increasing cervical lordosis on overall spinal alignment. In the present study we add to the body of evidence supporting the restoration and maintenance of cervical lordosis by using a multilevel ACDF procedure and analyze the impact of increasing cervical lordosis on overall spinal sagittal alignment.

\section*{Methods}

Institutional ethical board review was granted and the study initiated in May of 2014. Patients who had undergone multilevel ACDF (≥3 levels) by senior authors (J.O. and G. C.) between 2013 and 2014 were reviewed retrospectively. Patients who were adults (≥18 years age) and had a diagnosis of central stenosis with myelopathy or myeloradiculopathy who had failed conservative treatment were included. Patients were excluded if they lacked radiographic follow up (4 weeks, 10 weeks and 6 months).

\subsection*{Procedure}

Two ACGME accredited spine fellowship-trained orthopaedic surgeons performed multilevel ACDF consistent with Smith-Robinson techniques. Anterior decompression was performed removing all disc material and posterior osteophytes followed by the insertion of a trapezoidal interbody allograft device. This process was repeated for subsequent levels. Finally a lordotic anterior plate was then selected and fixed to the cervical spine with screws. Implants consisted of: lordotic cortico-cancellous interbody devices produced by MTF/DePuy Synthes (West Chester, PA) with rigid anterior plate fixation by Globus Medical (Audubon, PA).

\subsection*{Measurement technique}

Lateral cervical radiographs were all taken in identical settings with a set source-to-image-distance (SID) of 72”. The patient stood in a comfortable position resting the left shoulder against the x-ray bucky and looking ahead at a marker, while keeping their chin parallel to the floor.

Lateral full length 36” scoliosis films were obtained again at 72” SID with the arms crossed in front of the patient (elbows flexed to 45° and fingers on the opposite elbow) and the left shoulder resting on the x-ray bucky. The patient looked ahead at a marker and kept their chin parallel to the floor.

Patient radiographs were reviewed from the following time points: preoperative and postoperative 4 weeks, 10 weeks and 6 months.

Regional cervical sagittal alignment was measured using the widely used Cobb C2-C7 method, which has been shown to have excellent intra and interobserver reliability.\textsuperscript{19,20} We measured the angle formed between the inferior end plate of C2 and the superior end plate of C7 for global analysis of cervical alignment. Local sagittal alignment was measured from the superior endplate of the highest to the inferior endplate of the lowest surgical vertebrae. Sagittal vertical axis (SVA) was measured at two time points:
preoperative and postoperative 4 weeks. The SVA was performed by taking the distance of a plumb line dropped from the centroid of C2 (C1-level equivalent) and the center of the C7 vertebral body to the most posterior superior aspect of the sacral promontory (Figure 1 and Figure 2).

Analysis
Measurements were performed by two orthopaedic residents (Y. K. and A. L.) as well as a fellowship-trained musculoskeletal radiologist (E. L.). All measurements were performed twice and the mean of the two readings was used as a final measurement. For correlation and regression analysis, all readings from the three reviewers were averaged.

Statistics
All statistics were performed using SPSS version 23.0 (IBM). Summary statistics were calculated for demographic and radiographic measurements. Inter-observer reliability was verified using the interclass correlation statistic. For the analysis of global and segmental cervical lordosis, means from each time point were compared using a repeated-measures ANOVA. For analysis of sagittal vertical axis, a paired sample t-test was selected to compare pre- and post-operative means. P-values <0.05 were considered statistically significant. The Pearson correlation coefficient was used to assess the relationship between global cervical lordosis and sagittal vertical axis. Linear regression was used to predict the change in SVA based on change in cervical lordosis.

Results
Demographic Data
We identified 69 patients meeting our inclusion criteria (47 female). The average age was 57 years (range 33-81). A summary of patient demographics is available in Table 1. The majority of patients underwent four level ACDF (n=46), followed by three level ACDF (n=23). For analysis of sagittal vertical axis, 58 patients within the previous group who had also had full-length 36” scoliosis films available for review. The average age was 56.72 (range 33-81). The summary of patient demographics is available in Table 2. The majority of patients underwent four level ACDF (n=39), followed by three level ACDF (n=19).

Reliability analysis
We demonstrated good (ICC >0.8) to excellent (ICC >0.9) interrater reliability for all measured radiographic variables. For measurement of global lordosis ICC was 0.96 (95% CI 0.96, 0.97). For measurement of segmental lordosis interrater reliability was good with ICC of 0.89 (95% CI 0.85, 0.93). Interrater reliability was excellent for C1-SVA measurement with an ICC of 0.91 (95% CI 0.90, 0.93). Interrater reliability was excellent for C2-C7 SVA measurement with an...
ICC of 0.94 (95% CI 0.92, 0.95).

**Cervical Lordosis**

The mean global lordosis measured at each time point was as follows: preoperative 10.26˚ (95% CI 9.16˚, 11.37˚), 4 weeks 19.45˚ (95% CI 18.53˚, 20.35˚), 10 weeks 19.81 (95% CI 19.96, 20.65˚) and 6 months 19.34 (95% CI 18.47˚, 20.21˚) (Figure 3). A repeated measures ANOVA showed a statistically significant difference in the means of the pre-op time point and all post operative time points, F=316, p<0.0005. Pairwise analysis showed the postoperative time points were not significantly different between 4 wks to 10wks and 10 wks to 6 months (p=0.143, and p=0.113 respectively). Thus the null hypothesis was rejected.

The mean segmental lordosis measured at each time point was as follows: preoperative 8.22˚ (95% CI 7.2˚, 9.23˚), 4 weeks 20.35˚ (95% CI 19.5˚, 21.21˚), 10 weeks 20.3˚ (95% CI 19.46˚, 21.12˚), and 6 months 19.57˚ (95% CI 18.68˚, 20.45˚). A repeated measures ANOVA showed a statistically significant difference in the means of the preoperative time point and all post operative time points, F=454, P<0.0005. Pair wise analysis showed the postoperative time points were not significantly different between 4 weeks to 10 weeks, and 10 weeks to 6 months (p=1.00 and p=0.077 respectively). Thus the null hypothesis was rejected.

**Table 1. Study demographics for cervical alignment.**

<table>
<thead>
<tr>
<th>Demographic</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Mean Age</td>
<td>56 yrs</td>
</tr>
<tr>
<td>Sex</td>
<td>47 female</td>
</tr>
<tr>
<td>Smoking</td>
<td>N=32</td>
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<tr>
<td>Mean Weight</td>
<td>185.1 Lbs</td>
</tr>
<tr>
<td>Mean Height</td>
<td>64.9 inches</td>
</tr>
<tr>
<td>Levels Fused</td>
<td>4 levels, N=46, 3 Levels, N=23</td>
</tr>
</tbody>
</table>

**Table 2. Study demographics for full-length sagittal vertical axis.**

<table>
<thead>
<tr>
<th>Demographic</th>
<th></th>
</tr>
</thead>
<tbody>
<tr>
<td>Mean Age</td>
<td>56 yrs</td>
</tr>
<tr>
<td>Sex</td>
<td>39 female, 19 Male</td>
</tr>
<tr>
<td>Smoking</td>
<td>N=28</td>
</tr>
<tr>
<td>Mean Weight</td>
<td>182.1 Lbs</td>
</tr>
<tr>
<td>Mean Height</td>
<td>65.1 inches</td>
</tr>
<tr>
<td>Levels Fused</td>
<td>4 levels, N=39, 3 Levels, N=19</td>
</tr>
</tbody>
</table>

**Sagittal Vertical Axis**

The mean preoperative C2-C7 SVA was 12.04mm (±2.55mm) and the mean postoperative C2-C7 SVA was 27.49mm (±2.24mm), and the mean difference was 15.49 (±1.62mm) (Figure 4). A paired sample t-test showed the two means were significantly different (t=-9.48, p<0.0005). Thus the conclusion was made that there was a significant increase in C2-C7 SVA.

The mean preoperative C7-SVA was -1.93mm (±2.19mm) and the mean postoperative C7-SVA was 8.67mm (±2.04mm), and the mean difference was 10.59(±1.45mm). A paired sample t-test showed the two means were significantly different (t=-7.27, p<0.0005). Thus, there was a significant increase in

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Fig. 3. Line chart of the average degree of cervical lordosis measured at four time points (preoperative, 4 weeks, 10 weeks and 6 months postoperative).

Fig. 4. Bar graph showing the pre- and postoperative C1-SVA means.

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Global Lordosis to Sagittal Vertical Axis Correlation

For the subset of patients who had full length scoliosis films correlation analysis was performed. The mean change in global lordosis from pre- to postoperative was 9.28° (±0.96). The mean change in C7-SVA from pre- to postoperative was 10.59mm (±3.28). The mean change in C2-C7 SVA from pre- to postoperative was 15.44mm (±3.67). The change in C2-C7 lordosis had a positive correlation with both the change in C7-SVA (r=0.37, P<0.005) and the change in C2-C7 SVA (r=0.312, p<0.05). Linear regression analysis with change in C2-C7 lordosis as the independent variable and change in C7-SVA and C2-C7 SVA as the dependent variables revealed significant association (r² = 0.135 (p<0.005) and r² = 0.10 (p<0.05) respectively (Figure 5).

Discussion

Cervical lordosis is derived from both primary (developed in utero) and secondary (resulting from environmental forces) factors. Loss of natural lordosis has been related to pathological changes in the spinal cord. The largest amount of regional variation in tolerated sagittal alignment occurs in the cervical spine. Gore et al reported the C2-C7 angle in 200 asymptomatic patients as measuring between 16° and 27° depending on age. In an elegant synthesis of 12 studies of cervical alignment, Kuntz et al reported a pooled estimate of C2-C7 cervical lordosis as 17° with a variability of -11° to +45° across studies. Several studies have correlated both pre- and postoperative cervical lordosis with improved clinical outcome scores. Tang et al. showed that patients who had loss of lordosis as measured by increasing C2-C7 SVA had worse outcomes on Neck Disability Index (NDI) and SF-36 instruments. In their prospective randomized study comparing parallel to lordotic interbody devices in ACDF, Villavicencio et al showed that patients who had increased lordotic segmental sagittal alignment had a higher degree of clinical improvement as measured by the NDI and SF-36 instruments. Recently in their retrospective study, Iyer et al showed that negative correlation between C2-C7 SVA and NDI. While not directly correlated with clinical outcome, Okada et al showed that non-lordotic configuration of the cervical spine was associated with accelerated degenerative changes (i.e. posterior disk protrusion and disc space narrowing).

While there is significant variability in what can be considered normal cervical alignment, standing sagittal vertical axis is maintained within a much more narrow physiologic range. Deviations from this can inhibit individuals ability for upright posture and ambulation. Typically the normal C7-SVA is maintained within ±2.5cm to maintain balance and horizontal gaze. The reciprocal changes in one spinal region with the alteration of another have long been recognized. Previous studies have correlated increasing SVA with increases in cervical lordosis. In their multicenter retrospective study Smith et al. studied patients with C7-SVA >5cm who underwent lumbar pedicle subtraction osteotomy and showed that as SVA decreased there was a relaxation in cervical hyperlordosis. Moreover, increasing SVA has been correlated with poor clinical outcomes in multiple studies.

In the present study we showed significant increases in cervical lordosis after a multilevel ACDF procedure(Figure 6). In addition, we showed a weak (r²=0.1) but statistically significant (p<0.05) correlation between increasing cervical lordosis and a positive shift in the vertical alignment of the spine (SVA). Based on this analysis, roughly 10% of sagittal vertical axis change can be reliably explained by the change in cervical lordosis. While the governance of sagittal

Fig. 5. Scatter plot and regression analysis of change in C2-C7 Lordosis vs. change in C7-SVA in pre- to postoperative measurements.
vertical balance is complex and likely multifactorial, it seems reasonable to conclude from this study that the change in cervical lordosis plays at least a partial role in the alteration of sagittal alignment. As this study is purely radiographic, the clinical implications of this relationship were not determined. Future studies should examine outcome measures with relation to overall change in cervical lordosis and its relationship with SVA.

In the present study we hypothesized that multilevel ACDF would increase cervical lordotic alignment but the increase in SVA was surprising. This increase also seems to fit the model of the positive correlation between cervical lordosis and SVA to maintain physiologic horizontal gaze discussed in previous studies. Although analysis of the lumbar segment was not the purpose of this study, we suspect the compensatory increase in SVA resulted from a relaxation in lumbar lordosis and is reciprocal to the model proposed by Smith.

Limitations to this study include the following. First, this study is retrospective in nature and no comparison groups were used. Furthermore there are potential differences in surgical technique performed between the two surgeons, although both surgeons performed each surgery jointly. Additionally it should be noted that this study had a high predominance of female patients (2:1), which is contradictory to reported sex ratios for spondylolytic myelopathy (typically male to female 3:2). Finally, this study offers only 6 months of clinical follow up with no clinical outcomes to gauge treatment efficacy. While we show a significant increase in lordosis, follow up studies should be directed at measuring clinical outcome scores and fusion rates.

**Conclusions**

In this paper we show that multilevel ACDF with lordotic interbody devices significantly increases and maintains both global and segmental cervical lordosis. In addition we show that increasing cervical lordosis can have the secondary effect of increasing sagittal vertical balance, the clinical ramifications of which are currently unknown. Future studies should examine the clinical implications of increasing lordosis and if there is an optimal amount of correction for best results.

**References**


COI & Disclosures
The authors declare no relevant financial disclosures.

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