Predictors of poor outcome in cervical spondylotic myelopathy patients underwent anterior hybrid approach: focusing on change of local kyphosis

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Abstract
Object: This study was a retrospective multivariable analysis for risk factors of poor outcome in patients underwent anterior hybrid approach, and discussed the causes of worsening of postoperative local alignment.
Methods: A total of 86 patients with progressive spinal cord compression and local kyphosis underwent an anterior hybrid approach (ACDF + ACCF), between June 2011 and June 2017. We evaluated clinical outcome by the Japanese Orthopaedic Association (JOA) score and recovery rate. Patients were divided into two groups according to worsening and improving of postoperative local alignment. Multivariate logistic regression analysis and receiver operating characteristic (ROC) curve were applied to the evaluation of risk factors. Mann-Whitney U-test, Independent t-test and Chi-squared test was performed for the comparison of local kyphosis between postoperative and last follow-up.
Results: There were twenty patients who had a recovery rate of less than 50%. Advance age, longer duration of symptoms, bigger T1 slope angle and lower change of local kyphosis angle was significantly associated with a poor clinical outcome by multivariate logistic regression analysis. The ROC curve showed that the cutoff values of change of local angle between preoperative and last follow-up was 10.2 angle. The cause of worsening of postoperative local alignment had T1 slope, C2-7 SVA, adjacent segment degeneration (ASD), and implant subsidence.
Conclusions: The change of local kyphosis was a predictor of clinical outcome after the hybrid approach. Furthermore, postoperative ASD, implant subsidence, T1 slope and C2-7 Cobb were associated with recurrence of postoperative cervical kyphosis.
Introduction
Cervical spondylotic myelopathy (CSM) as a result of cervical stepwise degeneration usually has a deterioration of neurological function. In some patients with CSM, the quite formation of cervical kyphosis (CK) owed to biomechanical forces change and tonic muscular activity.[1] Progressive cervical kyphosis could lead to cervical pain, lower quality of life and decrease of cord volume in spinal canal.[2–5] Although conservative treatment could improve this disease to a certain extent,
surgery remained the key to decompression of the spinal cord and correction of kyphosis.[6-8] The approaches of treatment of multilevel CSM (m-CSM) with CK included anterior cervical discectomy and fusion (ACDF), anterior cervical corpectomy and fusion (ACCF) and posterior laminectomy, laminoplasty. Both anterior and posterior approaches were able to improve the clinical recover of neurological function, but the selection of approaches remained controversial.[9, 10]

With the improvement of medical apparatus and surgeon’s skill level, the difficulty of anterior approaches and postoperative complications is gradually decreasing. Because of the efficacy of anterior approaches with directly decompress spinal cord and great correction of CK,[11] a hybrid approach (ACDF + ACCF) was applied to the treatment of patients with m-CSM and CK. Nevertheless, few studies have focused on the analysis of risk factors of surgical outcomes in patients after correction of kyphotic alignment by the hybrid approach. The aim of our study was to assess the relationship between improvement of kyphotic deformity and recover of neurological function, particularly the changes of local kyphotic angle. Furthermore, the causes of worsening of postoperative local alignment were investigated deeply using statistical analyses. Our findings could offer advices to surgeons, thereby achieve optimal surgical outcome.

Material And Methods

Patients population

Between June 2011 and June 2017, eighty-six patients with m-CSM and cervical kyphosis underwent a hybrid approach which includes one level ACDF and one level ACCF. All procedures were approved by the Ethics Committee of The Third Hospital of Hebei Medical University, and the written informed consent had acquired from each patient. The STROBE statement was strictly observed over the course of this study.

The criteria were defined as following: (1) patients who were diagnosis with m-CSM and cervical kyphotic deformity; (2) Ossification of the posterior vertebral body, disc herniation and kyphotic alignment contributed to the spinal cord compression; (3) patients had progressive clinical symptoms with or without increased signal intensity of the spinal cord on T2-weighted MRI; (4) each case underwent the hybrid approach for decompression of spinal cord and correction of CK. Some
exclusion criteria were defined as following: (1) patients had cervical surgery before; (2) patients with spinal tumors or congenital cervical deformity or peripheral nerve disease; (3) patients with incomplete imaging data.

Neurological assessment

In this study, each individual patient had a follow-up over 12 months, who was evaluated neurological function by the Japanese Orthopedic Association (JOA) score before surgery and last follow-up. The time period of optimum recovery usually occurred within 12 months after surgery.[12] The surgical outcome was based on the following formula: the recovery rate = (postoperative JOA score - preoperative JOA score)/(17 - preoperative JOA score) × 100%. A score ≥ 75% was defined as excellent, ≥ 50% but < 75% as good, ≥ 25% but < 50% as fair, and < 25% as poor. Therefore, we divided recovery rates < 50% and ≥ 50% into poor group and good group, respectively.

Radiologic assessment

We had evaluated the radiological material on lateral radiographs and magnetic resonance imaging (MRI) before and after surgery. The cervical alignment defined as follows: positive = lordotic and negative = kyphotic. The T1 slope was defined as the angle between the superior endplate of T1 and a horizontal line on standing lateral radiograph. The C2-7 angle was defined as the angle of two lines of parallel to the inferior end plates of C2 and C7. (Fig. 1) The local kyphotic angle (CKA) was measured between the superior endplate of upper end vertebrae and the inferior endplate of lower end vertebrae. The C2-C7 SVA was defined as the distance between the plumb line of C2 centroid and posterior superior corner of C7. (Fig. 2) The loss of intervertebral height more than 3 mm was defined as cage subsidence between postoperative and final follow-up.[13] The C2-7 ROM was defined as the change in the maximal extension and flexion of cervical lateral radiographs. The worsening of postoperative local alignment was defined as recurrence of kyphotic deformity after surgical correction. Moreover, the increased single of spinal cord on T2-weighted images (T2WIs) and postoperative adjacent segment degeneration (ASD) was evaluated by MRI.

Surgical procedure

The hybrid approach (ACDF + ACCF) was determined by patient’s physical condition, clinical signs and
symptoms, and the compression location or levels of spinal cord on MRI or computed tomography (CT). Patients with ossification of the posterior vertebral body, kyphotic deformity and multilevel spinal cord compression would be performed an anterior hybrid approach. The corpectomy was performed on the vertebral with posterior ossification, and the discectomy was performed on the segment of disc herniation. Blunt dissection was carried out in cervical muscles and reached responsible segments from inner side of the sternocleidomastoid. The anterior corpectomy bounded by the uncovertebral joint on both sides. We adequately removed the ossification of the posterior vertebral body for decompression of spinal cord. An optimal polyether-ether-ketone (PEEK) cage and a titanium box cage with autogenous bone were implant in an appropriate location with anterior fixation of plate system. The rupture bleeding of venous plexus had to be highly-regarded. All patients required the utilization of a cervical collar for 3 or 4 weeks after surgery.

Statistical Analyses
Patient’s continuous variables and categorical variables were described as means and standard deviations and frequencies and percentages, respectively. We used univariate analyses to assess the correlations between factors and postoperative outcome. Multivariate logistic regression analysis was performed to the variables with a p-value less than 0.05 in univariate analysis. The receiver operating characteristic (ROC) curve was constructed to assess the cutoff values for analysis of risk factors. Mann–Whitney U-test, independent t-test and Chi-square test were performed for a comparison between improving of postoperative local alignment group and worsening of postoperative local alignment group. P-value < 0.05 represents a statistically significant difference. All analyses were performed using version 22.0 of SPSS software and the ROC curves were plotted with GraphPad Prism 8.

Results
In the Table 1, a series of variables about patient characteristics was showed. The good group had 66 patients with the recovery rates of JOA score greater than or equal to 50%, while the poor group had 20 patients with the recovery rates of JOA score greater less than 50%. Compared with the preoperative JOA score, there was a statistically significant improvement at last follow-up (p < 0.05).
However, a preoperative JOA score wasn’t correlate to surgical outcome (p = 0.641). With the univariate analyses, poor group had a significantly difference in age (p < 0.001), duration of symptoms (p = 0.002), preoperative T1 slope angle (p = 0.001), CLA (pre- and last) (p < 0.001), LKA at last follow-up (p = 0.003) and WPLA (p = 0.032). Furthermore, the Table 2 showed that age (OR = 1.09, 95% CI = 1.03-1.21, P = 0.017), duration of symptoms (OR = 4.64, 95% CI = 1.22-3.75, P = 0.001), T1 slope (OR = 1.02, 95% CI = 1.07-1.93, P < 0.001) and change of local kyphosis angle (OR = 10.67, 95% CI = 1.35-32.18, P = 0.003) were the risk factors of poor outcome after surgery.

### Table 1
Comparison of patient characteristics between good group and poor group

| Variables                  | Good (n = 66) | Poor (n = 20) | P-value |
|----------------------------|---------------|---------------|---------|
| Age at surgery (y)         | 57.1 ± 8.8    | 63.4 ± 9.2    | < 0.001 |
| Female, n (%)              | 35 (53%)      | 11 (55%)      | 0.877   |
| BMI (kg m$^{-2}$)          | 23.5 ± 3.6    | 24.6 ± 4.1    | 0.309   |
| Duration of symptoms (m)   | 11.6 ± 7.5    | 16.3 ± 11.8   | 0.002   |
| T1 slope (°)               | 14.3 ± 2.4    | 17.8 ± 3.2    | 0.001   |
| C2-7 SVA (mm)              | 17.6 ± 10.2   | 19.1 ± 9.4    | 0.266   |
| Involved levels, n (%)     |               |               | 0.762   |
| C2-5                       | 13 (19.7%)    | 4 (20%)       |         |
| C3-6                       | 31 (47%)      | 11 (55%)      |         |
| C4-7                       | 22 (33.3%)    | 5 (25%)       |         |
| JOA score                  |               |               |         |
| Preoperative               | 8.1 ± 2.2     | 7.5 ± 2.6     | 0.641   |
| Last follow-up*            | 14.3 ± 2.9    | 10.1 ± 1.5    | < 0.001 |
| Recovery rate, %           | 64.7 ± 18.3   | 34.9 ± 10.5   | < 0.001 |
| C2-7 ROM (°)               | 38.2 ± 12.7   | 39.5 ± 13.0   | 0.891   |
| C2-7 Cobb (°)              |               |               |         |
| Preoperative               | -10.7 ± 5.5   | -11.3 ± 6.2   | 0.219   |
| Postoperative              | 7.1 ± 4.4     | 6.9 ± 5.0     | 0.774   |
| Last follow-up             | 6.3 ± 5.7     | 5.6 ± 4.9     | 0.428   |
| Change of C2-7 (pre- and last) | 12.4 ± 8.7   | 13.1 ± 9.5   | 0.138   |
| LKA (°)                    |               |               |         |
| Preoperative               | -13.7 ± 10.5  | -11.3 ± 9.2   | 0.736   |
| Postoperative              | 5.1 ± 7.2     | 4.8 ± 6.3     | 0.552   |
| Last follow-up             | 5.6 ± 3.9     | 0.3 ± 5.4     | 0.003   |
| CLA (pre- and last)        | 12.1 ± 7.7    | 8.7 ± 6.5     | < 0.001 |
| WPLA, n (%)                | 10 (15.1%)    | 7 (35%)       | 0.032   |
| Implant subsidence > 3 mm, n (%) | 6 (9%)    | 2 (10%)      | 0.817   |

Notes: * Significantly different from preoperative JOA score (p<0.05)
Abbreviations: BMI, body mass index; T1, Thoracic 1; SVA, sagittal vertical axis; JOA, Japanese Orthopaedic Association; ROM, range of motion; LKA: Local kyphosis angle; CLA, change of local angle; WPLA, worsening of postoperative local alignment

### Table 2
Multiple logistic regression analysis forecasted risk factors for the postoperative outcome

| Variables                  | OR   | 95% CI        | P-value |
|----------------------------|------|---------------|---------|
| Age at surgery (y)         | 1.09 | (1.03-1.21)   | 0.017   |
| T1 slope                   | 1.02 | (1.07-1.93)   | < 0.001 |
| Duration of symptoms (m)   | 4.64 | (2.22-3.75)   | 0.001   |
| LKA at last follow-up      | 1.73 | (0.88-2.69)   | 0.118   |
| CLA                        | 10.67 | (1.35-32.18) | 0.003   |
| WPLA                       | 3.94 | (0.65-3.44)   | 0.122   |

Abbreviations: LKA: Local kyphosis angle; CLA, Change of local angle; WPLA, worsening of postoperative local alignment

The ROC curve illustrated the degree of correlation between predictive factors and postoperative poor outcome according to area under the curve (AUC). A larger AUC mean closer relationship between risk
factors and poor outcome. The change of local kyphosis angle (AUC = 0.829) between before surgery and last follow-up was a reliable predictor. The maximum of Youden index was defined as cutoff value which was the best compromise between sensitivity and specificity. Therefore, the cutoff values could forecast chances of developing poor outcome. The cutoff values of age, T1 slope, duration of symptoms and change of local kyphosis angle were 61.3 years, 16.7 angle, 9.5 months and 10.2 angle, respectively. (Table 3 and Fig. 3)

| Variables                      | Sensitivity (SN) | Specificity (SP) | AUC     | Cutoff  | P-value |
|--------------------------------|------------------|------------------|---------|---------|---------|
| Age at surgery (y)             | 0.876            | 0.634            | 0.748   | 61.3    | 0.001   |
| T1 slope                       | 0.617            | 0.812            | 0.763   | 16.7    | < 0.001 |
| Duration of symptoms (m)       | 0.876            | 0.584            | 0.763   | 9.5     | 0.002   |
| CLA                            | 0.578            | 0.903            | 0.829   | 10.2    | < 0.001 |

Abbreviations: AUC, area under the curve; SN, sensitivity; SP, specificity; CLA, Change of local angle

In this study, 17 patients (19.8%) occurred the worsening of postoperative local alignment.

Comparison of improving group and worsening group showed that T1 slope (p < 0.001), preoperative C2-7 SVA (p < 0.001), adjacent segment degeneration (p = 0.01) and implant subsidence (p = 0.024) were significantly related to change of postoperative curvature. (Table 4)

| Variables                      | Improving (n = 69) | Worsening (n = 17) | P-value |
|--------------------------------|--------------------|--------------------|---------|
| Age, (years)                   | 62.5 ± 9.0         | 60.9 ± 10.2        | 0.332a  |
| T1 slope, (°)                  | 18.7 ± 5.3         | 11.3 ± 4.2         | < 0.001b|
| Preoperative C2-7 SVA, (mm)    | 10.6 ± 8.8         | 24.9 ± 12.6        | < 0.001a|
| ASD, n (%)                     | 7(10.1%)           | 6(35.3%)           | 0.01c   |
| Implant subsidence > 3 mm, n (%)| 4(5.8%)            | 4(23.5%)           | 0.024c  |
| Involved levels, n (%)         |                    |                    | 0.921c  |
| C2-5                            | 14                 | 3                  |         |
| C3-6                            | 34                 | 8                  |         |
| C4-7                            | 21                 | 6                  |         |

Notes: a Mann-Whitney U-test; b Independent t-test; c Chi-square test; Abbreviations: ASD, Adjacent segment degeneration

Discussion

Historically, early clinical intervention and optimal surgical approach have been recognized as the prerequisites of a good surgical outcome. All kinds of risk factors affected surgical outcome in patients with CSM and CK. Several articles have been published that predictors of postoperative outcome in patients with CSM included age at operation, duration of symptoms and signal intensity of spinal cord.[14, 15] However, few studies paid close attention to clinical outcome after anterior hybrid
surgery for patient with multilevel CSM and kyphosis deformity. Did cervical kyphosis impede the recovery of neurological function after surgery? Chavanne[16] found that change of cervical alignment would lead to the increasing of intramedullary pressure, especially cervical kyphotic deformity exceeded 21 degrees. Whereas, the correlation between correction of kyphosis alignment and postoperative outcome remained controversial.[10, 17, 18]

An appropriate surgical approach was one of the important factors to achieve good postoperative outcome. However, the selection of surgical approach was the once controversial notion. Anterior approach could directly remove the lesion of compression of spinal cord and correct more kyphotic angle. Unfortunately, anterior approaches were more vulnerable to complications which were C5 palsy, cerebrospinal fluid leakage and temporary dysphagia.[19] Posterior approach relieved spinal cord compression according to the drift backward of spinal cord and the enlarging of vertebral canal volume. Posterior approach would be an optimum selection when patients were accompanied by a consecutive ossification of posterior longitudinal ligament.[20] Whereas, cervical kyphotic alignment reduced the space of drift backward of spinal cord, which would influence clinical outcome. In previous study, Cabrera[11] showed that anterior approach had a better restored of cervical kyphosis than the posterior approach, and Suda[21] showed that posterior laminoplasty was best indication for patients with local kyphosis less than 13 degrees. Furthermore, Yang[22] showed a new-style: anterior controllable ante-displacement fusion (ACAF), which could enlarge the volume of the spinal canal and simultaneously correct cervical kyphosis for patients with m-CSM and stenosis. In this study, an anterior hybrid approach was selected to remove ossification of the posterior vertebral body and correct kyphotic deformity. This method could not only directly decompress spinal cord, but also avoid the occurrence of low fusion rate and implant translocation after multi-ACCF. Therefore, the selection of surgical approaches should be planned on an individual basis.

In this study, the change of local kyphotic angle between preoperative and last-follow was associated with surgical outcome according to the JOA score, particularly corrective angle more than 10.2 degree. However, the change of C2-7 Cobb angle was not statistically significant difference between good group and poor group. We considered that the cause was the occurrence of cervical “S” or
reverse “S” type that lordosis and kyphosis existing side-by-side on the lateral films of cervical vertebra. Hence, the measurement of local kyphosis angle had greater meaningful to the prevention of poor postoperative outcome. In previous study, Uchida[10] explained also that correction of local cervical kyphosis was beneficial to improve the recovery of neurological function. Whereas, excessive correction of cervical kyphosis would give rise to short-term axial pain after surgery.[23]

Not all patients could maintain the effect of surgical correction with passage of time. In this study, there were seventeen patients with the worsening of postoperative local alignment as a result of each kind of suggestion factor. Although our research didn’t show that the worsening of postoperative local alignment was associated with poor outcome after surgery, the causes of worsening were necessary to study further. Lee[24] found that one of key factors of affecting cervical spine sagittal balance was T1 slope. Patients with cervical kyphotic deformity required low T1 slope angle to compensate for maintaining the sagittal balance. In addition, Katsuura[25] suggest that adjacent segment degeneration after anterior cervical fusion was also a key factor determining postoperative kyphotic change in the cervical fused segment. Our finding confirmed that the T1 slope and adjacent segment degeneration were associated with change of postoperative local alignment. The results expounded that surgical correction of cervical local kyphosis couldn’t effectively improve the T1 slope angle, so recurrence of kyphotic deformity aimed to compensate for the balance of cervical alignment.

Furthermore, we founded also that implant subsidence and large preoperative C2-7 SVA were the important factors of worsening of postoperative local alignment. The implant subsidence would lead to the height difference of anterior and posterior margin in surgical segments. Frequent head-down tilt after surgery was crucial reason of the subsidence in front of implant. Understanding and preventing risk factors could obtain maximum improvement in quality of life after surgery.

In our study, several limitations were worth mentioning. Firstly, this was a retrospective study with relatively small number of patients and lacked long-term follow-up data for some patients. Secondly, the evaluation of surgical outcome only focused on improvement of JOA score and recovery rate, neglected the evaluation according to the SF-36 scale and the HR-QOL scale. Thirdly, the relationship between the change of cervical kyphotic alignment and thoracolumbar or spino-pelvic parameters
were not considered. Finally, the adjacent segment degeneration was evaluated only at last follow-up, which may continue to occur in the future.

**Conclusion**

Age at operation, duration of symptoms, T1 slope angle and change of local kyphosis were the risk factors of clinical outcome in patients with CSM and cervical kyphosis. Furthermore, postoperative ASD, T1 slope, implant subsidence and C2-7 Cobb were associated with recurrence of postoperative cervical kyphosis.

**Declarations**

**Competing interests**

The authors declare that they have no competing interests.

**Author’s contributions**

Conceived and designed the study: Y.S. and X.J.C. Collected data: X.J.C. X.W. and L.J. Analyzed the data: L.J. and W.Z. Wrote the paper: X.J.C.

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Figures
Figure 1

A schematic drawing: patients with Cervical spondylotic myelopathy and cervical kyphosis;

Radiographic measurements: T1 slope and C2-7 Cobb angle.
A schematic drawing: patients underwent the hybrid approach (ACDF+ACCF); Radiographic measurements: C2-7 SVA.

Figure 3

Age at operation, T1 slope, duration of symptoms and change of local angle were shown by the ROC, respectively.