

Hybrid decompression and reconstruction technique for cervical spondylotic myelopathy: case series and review of the literature

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Abstract: The primary treatment of choice for patients with cervical spondylotic myelopathy (CSM) is surgical decompression. The benefit of operative intervention has been well established but, the surgeons' decision of operative approach remains nuanced based on patient-specific variables and surgeon preference. Decompression can involve a cervical corpectomy or a discectomy. A hybrid construct is when both a cervical corpectomy and a discectomy are done in the same patient. The purpose of this study was to review the evidence on the clinical and biomechanical outcomes of hybrid decompression and reconstruction techniques in patients with multilevel CSM. A retrospective study was performed on consecutive patients who received hybrid anterior decompression and reconstruction at Rush University between 2013-2018. Preoperative clinical and radiographic variables were analyzed to characterize specific factors leading to the decision of the surgical approach. In addition, we performed a systematic review and meta-analysis to assess superiority in terms of operative time, blood loss, cervical lordosis, patient-reported outcomes (PRO), fusion rates, and complications. Hybrid surgery (HS) was utilized in cases where multilevel CSM was present in conjunction with stenosis posterior to the vertebral body or acute kyphotic deformity. Our meta-analysis highlighted comparable PRO, complications, and rate of success fusion between 3-level anterior cervical discectomy and fusion (ACDF) and hybrid technique. Furthermore, hybrid fusion led to increased postoperative cervical lordosis, higher fusion rate, lower total complication rate, lower implant failure/mesh subsidence rate, and lower blood loss than 2-level corpectomy. The cervical hybrid technique that combines cervical corpectomy and discectomy represents a balanced option with the benefits of two commonly utilized cervical spine procedures in patients with multilevel CSM. The literature on hybrid technique suggests in cases where multilevel ACDF is not feasible, combining discectomy and corpectomy is superior to two-level corpectomy with lower complication rates, improved clinical outcome, spinal alignment correction, and stronger biomechanical properties.

Keywords: Cervical spondylotic myelopathy (CSM); corpectomy; discectomy; hybrid technique

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Background

Cervical spondylotic myelopathy (CSM) is the most common cause of non-traumatic spinal cord dysfunction in adults (1). Based on the progressive natural history of this disorder, the primary management of patients with CSM remains prompt surgical intervention for neural decompression (2). While there is an abundance of evidence highlighting the effectiveness of surgery in improving patient outcomes, there remains considerable variability in the specific surgical approaches utilized. Fehlings et al. found a nearly two-fold difference in the frequency of anterior versus posterior cervical decompression surgeries utilized in Europe and Latin America (3). When dealing with multilevel CSM, the spectrum of operative considerations increases even further. Within the context of anterior cervical decompression, options have traditionally been between multiple anterior cervical discectomy and fusion (ACDF) versus anterior cervical corpectomy fusion (ACCF) depending on certain radiographic features (2). However, a growing number of studies have looked at various hybrid surgery (HS) models aimed at maximizing the benefits of these surgical techniques (4,5). Previous reviews of the literature on surgical management of multilevel CSM have been limited to clinical studies to reduce the heterogeneity of data. There are, however, biomechanical and radiographic studies that can provide additional insights for the decision-making process (6-8).

The purpose of this study was to review the evidence on the clinical and biomechanical outcomes of cervical fusion hybrid decompression and reconstruction techniques in patients with multilevel CSM and characterize preoperative patient variables that may benefit from a hybrid technique.

Methods

Case series

A retrospective review was performed on consecutive patients who received hybrid anterior decompression and reconstruction at Rush University between 2013–2018. The inclusion criteria were: (I) adult patients (\geq 18 years) with a new diagnosis of CSM; (II) preoperative MRI confirming multilevel disease; (III) underwent hybrid decompression and reconstruction surgery. Patients with other indications for surgery such as spine tumor, infection and trauma were excluded. This study was approved by the Rush University Institutional Review Board (19070801-IRB01-AM01).

Data abstraction was obtained from hospital and clinic records and included patient demographics, clinical, and radiographic variables. Preoperative consultation notes were examined to identify specific surgical indications and operative approaches considered by the surgeons. Imaging analysis consisted of preoperative cervical spine MRI and a postoperative cervical spine X-ray. Radiographs were digitally analyzed using PACS software for the sagittal Cobb angle (angle between inferior endplates of C2 and C7). Preoperative and postoperative patients' health status was measured using patient-reported outcomes (PRO) including a visual analogue scale for neck pain (VAS) and modified Japanese Orthopedic Association scale (mJOA). Lastly, the cases were individually reviewed with the primary surgeon to confirm their reasons for choosing the hybrid technique.

Surgical technique

The traditional anterior approach to the cervical spine was utilized for all of the cases. Patients were positioned supine with the head supported in a donut headrest and neck slightly extended. While not necessary in our case series, surgeons in our institution regularly utilize Gardner-Wells tongs connected to weights at the head of the bed for additional distraction. To improve intraoperative fluoroscopic visualization, patients' shoulders were taped and pulled caudally. For all of the cases, a transverse incision was planned centered at the corpectomy level using an intraoperative X-ray. The preferred side of surgery for the cervical spine remains a contentious topic with some surgeons emphasizing higher risk to the recurrent laryngeal nerve associated with the right-sided surgery. Anecdotal experience at our institution, where both right and leftsided surgeries are regularly performed, does not provide support for preferential use of left-sided surgery.

For all of the cases, a generous subplatysmal dissection prior to fascial dissection allowed adequate surgical exposure through a transverse incision without postoperative dysphagia. Blunt dissection through the superficial and deep cervical fascia was performed following the anterior border of the sternocleidomastoid muscle with the carotid sheath positioned laterally and trachea and esophagus medially. After identifying the prevertebral fascia, subperiosteal dissection of the longus colli bilaterally was performed prior to retractor positioning. With the index vertebra with the greatest canal narrowing confirmed on intraoperative fluoroscopy, standard discectomy at the cranial and caudal levels were performed with a combination of high-speed drill and Kerrison punches. Discectomy at the adjacent level for the ACDF was performed at the same time. With the uncinate processes exposed, corpectomy was then performed with removal of greater than 80% of the vertebral body leaving narrow columns of bone laterally to project the vertebral arteries. After completion of the bony resection, careful resection posterior longitudinal ligament was performed to ensure decompression of the neural elements. The anterior column reconstruction was performed using various strut graft options, including allograft, PEEK, titanium mesh cage, or carbon fiber cage. All of the cases were supplemented with an anterior cervical plate spanning both corpectomy and discectomy levels. Screws for the cervical plate were placed aiming for bicortical purchase to ensure maximal pullout strength. Postoperatively, cervical collars were utilized only in two cases with poor bone quality and screw purchase.

Systematic review and meta-analysis

We performed a systematic review of the literature using PubMed, the Cochrane Library, Scopus, and included citations to identify clinical studies on hybrid decompression and reconstruction techniques in patients with three-level CSM. The systematic review and metaanalysis were conducted based on the Preferred Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA) statement. Specific MeSH terms and keywords including "anterior cervical hybrid fusion", "hybrid surgery", "corpectomy combined with discectomy", "anterior cervical corpectomy and fusion", "vertebrectomy", "anterior cervical discectomy and fusion", "cervical spondylosis", and "cervical spondylotic myelopathy" were used to identify studies of interest. Additional manual searches through cited references were performed.

Studies were included in meta-analysis if they met the following criteria: (I) cohort or case-control comparative studies; (II) patients with 3-level CSM; (III) studies comparing the clinical outcomes of the hybrid fusion technique versus ACCF and/or versus ACDF; (IV) outcome assessments included duration of surgery, blood loss, surgical complications, fusion rate, radiographic outcomes including measured C2-7 lordosis, patient-related outcomes included mJOA score, Neck Disability Index (NDI). Studies were excluded if they involved (I) posterior surgical approaches, (II) inclusion of cervical arthroplasty within analysis, (III) 4 level CSM or surgery on noncontiguous levels, (IV) mean follow-up less than 1 year (V) if full-text articles could not be obtained, (VI) non-English publications, editorials, conference abstracts, errata, book chapters, and case reports were excluded.

Statistical analysis

Descriptive statistics was performed to characterize patient demographic, clinical, and surgical data in our case series using STATA 14.3 (StataCorp LLC, College Station, Texas, 2018). Statistical analysis for the systematic review and meta-analysis was performed using RevMan 5.3 (Copenhagen: The Nordic Cochrane Centre, The Cochrane Collaboration, 2014). An I² test was performed for each study to test statistical heterogeneity. An I²>50% was considered to indicate substantial heterogeneity and a random-effects model was used for meta-analysis. When I² was less than 50% a fixed-effects model was used. An odds ratio and 95% confidence interval were calculated for dichotomous outcomes whereas a standardized mean difference and 95% confidence interval were calculated for continuous outcomes. A P value <0.5 was considered to be statistically significant.

Results

Case series

After reviewing the charts of patients who received corpectomy between 2013 to 2018, a total of 10 patients met the inclusion criteria. The mean age of the patients was 66 (range, 57–90) years old consisting of 4 females and 6 males. Their preoperative BMI ranged from 21.5 to 38.9 with an average of 28. Of the 10 patients, the most frequent medical comorbidities were hypertension (n=10), dyslipidemia (n=6), and diabetes (n=4). Two patients were active smokers at the time of the procedure.

In determining the specific surgical approach, the most common indication for corpectomy over multiple discectomies was the presence of stenosis posterior to the vertebral body due to caudally or cranially herniated disc or thickened/calcified posterior longitudinal ligament (Figure 1). Other reasons included short segment kyphotic deformity and spondylolisthesis (Figure 2). In addition, these patients underwent a hybrid technique of corpectomy and discectomy over two-level corpectomies to allow three levels of fixation. There were two cases where the hybrid technique was used to address a single degenerated mobile segment between the corpectomy level and auto-fused vertebrae two levels adjacent to it (Figure 3). Two out of 10 patients were placed in cervical collars after surgery. The duration of postoperative cervical immobilization of these patients was between 6 and 12 weeks.

Clinical data and details of the surgical intervention are summarized in *Table 1*. The average surgical blood loss was 107.5 mL (range, 50–200 mL). The average postoperative sagittal Cobb angle was 11.8 degrees lordotic (range, 2.5–29.6). The overall complication rate was 40% with 2/10 patients experiencing transient postoperative dysphagia,



Figure 1 A 67 years old male with multilevel disc disease with significant retrovertebral compression at C4. He underwent C4 corpectomy and C5-6 ACDF, ACDF, anterior cervical discectomy and fusion.

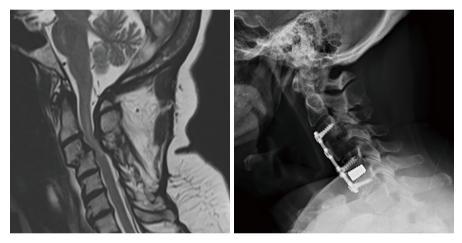


Figure 2 A 57 years old female with preoperative MRI showing acute kyphotic deformity at C2 and C3 in setting of multilevel disc disease. She underwent C4 corpectomy and C5-6 ACDF, ACDF, anterior cervical discectomy and fusion.

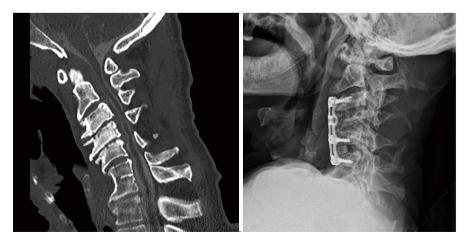


Figure 3 A 90 years old male with multilevel disc disease with retrovertebral compression at C4 along with autofusion of C6 and C7. He underwent C4 corpectomy and C5-6 ACDF. ACDF, anterior cervical discectomy and fusion.

 Table 1 Operative and postoperative clinical data

Procedure	Implant-corpectomy	Implant-discectomy	Plate	Complications	EBL (mL)	LOS (days)
C4 corpectomy, C5/6 ACDF	Allograft	Titanium mesh cage	Yes	Nil	100	3
C4 corpectomy, C5/6 ACDF	Allograft	Allograft	Yes	Epidural fluid collection requiring surgical decompression	150	2
C4 corpectomy, C6/7 ACDF	Allograft	Allograft	Yes	Nil	200	3
C4 corpectomy, C5/6 ACDF	Allograft	Allograft	Yes	Nil	50	5
C4 corpectomy, C5/6 ACDF	Allograft	Allograft	Yes	Dysphagia	100	5
C5 corpectomy, C3/4 ACDF	Carbon fiber cage	Carbon fiber	Yes	Nil	75	3
C5 corpectomy, C3/4 ACDF	PEEK	PEEK	Yes	Dysphagia	100	10
C5 corpectomy, C3/4 ACDF	Allograft	Allograft	Yes	Nil	100	7
C6 corpectomy, C4/5 ACDF	PEEK	PEEK	Yes	Myocardial infarction	100	36
C7 corpectomy, C5/6 ACDF	Carbon fiber cage	Allograft	Yes	Nil	100	2

ACDF, anterior cervical discectomy and fusion; PEEK, polyetheretherketone; EBL, estimated blood loss; LOS, length of stay.

1/10 patients experiencing a myocardial infarction during hospitalization, and 1/10 patients developing an epidural fluid collection requiring surgical decompression. One patient required additional posterior decompression and instrumentation due to persistent stenosis during initial hospitalization. Five patients had a postoperative followup longer than 1 year. At the end of the follow-up period, no one developed symptomatic pseudoarthrosis requiring additional surgical intervention. The postoperative PRO showed an average VAS of 1.8 (range, 0–4) and average mJOA of 11 (range, 6–14).

Systematic review

In total, 171 abstracts were reviewed of which 93 were excluded. Seventy-eight full-text articles were assessed of which 24 were excluded. Several articles were excluded for including arthroplasty within the HS group (9-28). Articles were also excluded if they included HS for 4 level CSM, or if they included non-contiguous constructs (29-32). Fourteen manuscripts including both case series of hybrid fusion constructs as well as retrospective cohorts comparing hybrid fusion to ACDF and/or ACCF are summarized in *Table 2* with operative data regarding hybrid fusion summarized in *Table 3* (5,8,33-44). Of these, six manuscripts met the predefined inclusion criteria for meta-analysis with data summarized in *Table S1* (5,34,36,37,42,43). A flow chart of study inclusion and exclusion is shown in *Figure 4* (45).

Meta-analysis

Of the six retrospective cohorts included in meta-analysis, only four studies included all three surgical arms of threelevel ACDF, hybrid fusion, and two-level ACCF (5,36,37,43). Liu et al. 2009 compared patients undergoing hybrid fusion and patients undergoing two-level ACCF whereas Xu et al. 2019 compared patients undergoing hybrid fusion and patients undergoing three-level ACDF (34,42). The six eligible studies were published between 2009 and 2019 and included 293 patients undergoing three-level ACDF, 299 patients undergoing hybrid fusion, and 202 patients undergoing two-level ACCF. The average follow-up ranged from 17.3 to 43.2 months. The average age of patients undergoing each procedure ranged from 46.1 to 57.1 years, 46.9 to 55.9 years and 47.8 to 55.2 years for ACDF, hybrid fusion, and ACCF respectively. All patients undergoing corpectomies in the included manuscripts had placement of titanium mesh grafts except for 2 patients in Liu et al. 2009 who underwent placement of autologous iliac crest. Polyetheretherketone (PEEK) grafts were used in all patients during ACDF except for in Xu et al. 2019 where carbon fiber grafts were favored. All patients had complete coverage of their construct with a titanium plate; no stand-alone grafts were included in the meta-analysis. All studies reported collar duration which ranged from 3 weeks postoperatively in Liu, Qi, et al. 2012 to 3 months in Guo et al. 2011. Liu et al. 2009 was the only study to discuss preoperative collar placement which ranged from 8-12 months.

Year	Year Author	Design	Comparators	Sample size	Average age (years)	Follow-up duration average (months)
2005	2005 Ashkenazi <i>et al.</i>	Retrospective case series	None	12	60.91	29 includes 4 level patients
6003	2009 Wei-bing <i>et al.</i>	Retrospective cohort	Corpectomy + ACDF vs. 2 level ACCF	HS: 20; ACCF: 39	HS: 58.75; ACCF: 61.82	Median 20 months, (range, 18–24)
6003	2009 Liu <i>et al.</i>	Retrospective cohort	Corpectomy + ACDF vs. 2 level ACCF	HS: 12; ACCF: 16	53.5	17.3
2011	2011 Guo <i>et al.</i>	Retrospective cohort	3 level ACDF vs. corpectomy + ACDF vs. 2 level ACCF	ACDF: 43; hybrid: 53; ACCF: 24	ACDF: 52.7; hybrid: 53.4; ACCF: 55.2	ACDF: 37.7; hybrid: 37.3; ACCF: 37.3
012	2012 Liu Hou <i>et al.</i>	Retrospective cohort	3 level ACDF vs. corpectomy + ACDF vs. 2 level ACCF	ACDF: 69; hybrid: 72 ACCF: 39	ACDF: 46.1; hybrid: 46.9; ACCF: 47.8	ACDF: 26.8; hybrid: 25.6 ACCF: 26.4
012	2012 Liu Qi <i>et al.</i>	Retrospective cohort	3 level ACDF vs. corpectomy + ACDF vs. 2 level ACCF	ACDF: 103 HS: 96; ACCF: 87	ACDF: 53.48; HS: 54.36; ACCF: 53.68	43.2
2016	2016 Zarate-Kalfopulos Retrospective et al. Case series	Retrospective Case series	Corpectomy + ACDF	15	64.8	29.6
017	2017 Zhou <i>et al.</i>	Retrospective Case series	ries No comparators; corpectomy + ACDF	28	64.2	22.8
017	2017 Kratzig e <i>t al.</i>	Retrospective cohort	Corpectomy + ACDF vs. 2 level ACCF	HS: 56; ACCF: 76	NR	NR
2017	Liu <i>et al.</i>	Retrospective cohort	Corpectomy + ACDF with two types of titanium cage	New cage: 45; traditional cage: 49	New cage: 57.1; traditional cage: 56.7	New cage: 74.4; traditional cage: 77.3
018	2018 Sun <i>et al.</i>	Retrospective cohort	HS with corpectomy cephalad vs. caudal position	48	Cephalad: 53.95; caudal: 54.11	Cephalad: 4.38; caudal: 4.81
019	2019 Xu <i>et al.</i>	Retrospective cohort	3 level ACDF vs. corpectomy + ACDF	ACDF: 40; hybrid: 42	ACDF: 55.3; hybrid: 55.9	35.5
019	2019 Huang <i>et al.</i>	Retrospective case series	No comparators; corpectomy + ACDF	42	55.9	20.3
019	2019 Wei e <i>t al.</i>	Retrospective cohort	3 level ACDF vs. corpectomy + ACDF vs. 2 level ACCF	ACDF: 38; hybrid: 24; ACDF: 57.1; hybrid: ACCF: 36 55.4; ACCF: 51.6	ACDF: 57.1; hybrid: 55.4; ACCF: 51.6	ACDF: 26.2; hybrid: 28.5; ACCF: 27.7

186

Sun 2018

Xu 2019

Wei 2019

Huang 2019

Table 3 Summa	ry of clinical and operativ	e data of the included studies			
Study	Levels	ACCF graft material	ACDF graft material	Plate placed over entire construct?	Collar duration
Ashkenazi 2005	5 NR	Titanium mesh	Titanium or carbon fiber	Yes	8–10 weeks, limited to night and during car travel
Wei-bing 2009	NR	Titanium mesh	Titanium	No, standalone ACDF included	3 months
Liu 2009	C3–6: 7; C4–7: 5	Titanium mesh; 2 patients in ACCF group had tricortical autologous iliac crest	PEEK	Yes	8–12 months preoperatively
Guo 2011	C3–6: 29; C4–7: 24	Titanium mesh	PEEK	Yes	3 months
Liu Qi 2012	C2–5: 25; C3–6: 32; C4–7: 39	Titanium mesh	PEEK	Yes	3 weeks
Liu Hou 2012	C3–C6: 32; C4–C7: 40	Titanium mesh	PEEK	Yes	6–8 weeks
Zarate- Kalfopulos 2016	NR S	Fibular graft	NR	Yes	NR
Zhou 2017	C3–6: 13; C4–7: 15	Titanium mesh	PEEK	No, standalone ACDF included	3 months
Kratzig 2017	C3–6: 18; C4–7: 38	Titanium mesh	Titanium	Yes	NR
Liu 2017	C3–6: 59; C4–7: 35	Titanium mesh: traditional and new version with curved superior endcap	PEEK	Yes	2 months

and angled inferior endcap

Titanium mesh

Titanium mesh

Titanium mesh

NR

Table 3 Summary	of clinical	and operati	ve data of	the included	studie
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Operative time

Four studies compared operative time in ACDF and hybrid fusion showing no significant difference in operative time between the two groups (SMD 0.22; 95% CI, -0.57, 1.01; P=0.59). Similarly, three studies comparing operative time in hybrid fusion and ACCF found no significant difference in operative time between the two groups (SMD 0.35; 95% CI, -0.71, 1.4; P=0.52).

C3-6: 21; C4-7: 17

C3-6: 20; C4-7: 22

C3-6: 18; C4-7: 24

C3-6: 11; C4-7: 13

polyetheretherketone; NR, not reported.

Blood loss

Four studies compared blood loss in ACDF and hybrid fusion with significantly less estimated blood loss ranged in the ACDF group (102.3-172.1 mL) than in the hybrid group (141.3-193.2 mL) (SMD -1.34; 95% CI, -2.39, -0.30; P=0.01). Three studies compared blood loss in hybrid fusion and ACCF showing a statistically significant reduction in blood loss in the hybrid group than in the ACCF group (SMD -0.61; 95% CI, -0.88, -0.34; P<0.00001).

2 months

6-8 weeks

6-8 weeks

NR

Cervical lordosis

PEEK

NR

ACDF, anterior cervical discectomy fusion; ACCF, anterior cervical corpectomy fusion; CDA, cervical disc arthroplasty; PEEK,

PEEK

Carbon Fiber Yes

Yes

NR

Yes

Four studies compared C2-7 cervical lordosis at followup in ACDF and hybrid fusion. The mean cervical lordosis in the ACDF group was similar to the hybrid group (14.9-24.27 degrees vs. 12.8-23.21 degree, respectively; SMD 0.26; 95% CI, -0.23, 0.75; P=0.31). Three studies compared C2-7 cervical lordosis at follow-up in hybrid fusion and ACCF. The mean cervical lordosis ranged from 17.3 degrees to 23.21 degrees in the hybrid group and 11.4 degrees to 15.63 degrees in the ACCF group. With an I^2 of 0%, a fixed-effects model meta-analysis was used and there was a statistically significant increased C2-7 cervical lordosis

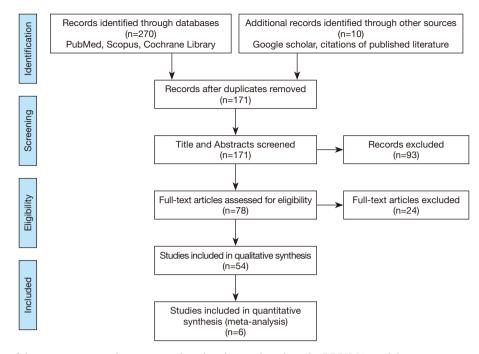


Figure 4 Flow chart of the systematic search strategy and study selection based on the PRISMA guideline.

at follow-up in the hybrid fusion group compared to the ACCF group (SMD 0.76; 95% CI, 0.49, 1.03; P<0.00001).

Change in cervical lordosis from preoperative to followup in ACDF and hybrid fusion was compared in three studies. The mean change in cervical lordosis in the ACDF group was statistically higher than in the hybrid group (4.9–15.1 vs. 1.8–8.1 degrees, respectively; SMD 0.56; 95% CI, 0.14, 0.99; P=0.009). Change in cervical lordosis from preoperative to follow-up in hybrid fusion and ACCF was compared in two studies. The mean change in cervical lordosis ranged from 4.5 to 8.1 degrees in the hybrid group and 1.7 to 2.7 degrees in the ACCF group. There was a statistically significant increased change in C2–7 cervical lordosis in the hybrid fusion group compared to the ACCF group (SMD 1.09; 95% CI, 0.71, 1.46; P<0.00001).

Patient reported outcomes

Five separate studies compared JOA score at follow-up in ACDF and hybrid fusion with comparable mean JOA score at follow-up ranged from 13.6 to 14.8 in the ACDF group and 13.1 to 14.9 in the hybrid group (SMD –0.10; 95% CI, –0.65, 0.45; P=0.72) (*Figure 5A*). Five separate studies compared JOA score at follow-up in hybrid fusion and

ACCF also showed comparable postoperative JOA scores between groups (SMD 0.03; 95% CI, -0.36, 0.43; P=0.88) (*Figure 5B*).

Three separate studies compared NDI at follow-up in ACDF and hybrid fusion with similar mean NDI score at follow-up ranged from 5.6 to 14.7 for ACDF and 6 to 14.3 for hybrid fusion (SMD –0.03; 95% CI, –0.22, 0.16; P=0.76) (*Figure 6A*). Three separate studies compared postoperative NDI in hybrid fusion and ACCF with similar mean NDI scores at follow-up ranged from 14.2 to 14.9 for hybrid fusion and 14 to 17.19 for ACCF (SMD –0.39; 95% CI, –0.90, 0.12; P=0.14) (*Figure 6B*).

Fusion

Five separate studies compared the percentage of patients who were fused at follow-up in ACDF and hybrid fusion. The percentage of patients fused at follow-up ranged from 97.7% to 100% in the ACDF group and 94.4% to 100% in the hybrid fusion group (OR 3.50; 95% CI, 0.87, 14.05; P=0.08) (*Figure 7A*). Five separate studies compared the percentage of patients who were fused at follow-up in hybrid fusion and ACCF. The percentage of patients fused at follow-up in the hybrid fusion group was significantly

А	A	CDF		Hybri	d Fus	ion		Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl
Guo 2011	13.7	1.9	43	13.1	2.3	53	20.1%	0.28 [-0.12, 0.68]	
Liu Hou 2012	14.1	1.6	69	13.8	1.9	72	20.9%	0.17 [-0.16, 0.50]	
Liu Qi 2012	14.8	2.2	103	13.9	2.8	96	21.4%	0.36 [0.08, 0.64]	
Wei 2019	13.6	0.7	38	14.9	0.8	24	17.8%	-1.73 [-2.34, -1.13]	
Xu 2019	14	1.8	40	13.6	2.1	42	19.8%	0.20 [-0.23, 0.64]	
Total (95% CI)			293			287	100.0%	-0.10 [-0.65, 0.45]	-
Heterogeneity: Tau² = Test for overall effect:				df=4 (P	< 0.0	0001);1	r≊= 90%		
В									

Hybrid Fusion		ACCF			Std. Mean Difference		Std. Mean Difference		
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% CI
Guo 2011	13.1	2.3	53	13	2	24	20.1%	0.04 [-0.44, 0.53]	_
Liu 2009	14.333	0.536	12	14.344	0.651	16	14.1%	-0.02 [-0.77, 0.73]	
Liu Hou 2012	13.8	1.9	72	14.5	1.8	39	22.3%	-0.37 [-0.77, 0.02]	
Liu Qi 2012	13.9	2.8	96	14.5	2.7	87	24.8%	-0.22 [-0.51, 0.07]	
Wei 2019	14.9	0.8	24	14.3	0.6	36	18.7%	0.86 [0.32, 1.40]	
Total (95% CI)			257			202	100.0%	0.03 [-0.36, 0.43]	
Heterogeneity: Tau² =			•	4 (P = 0.)	005); I ² =	= 73%		-	-1 -0.5 0 0.5 1
Test for overall effect	:Z=0.15 i	(P = 0.8	8)						

Figure 5 Forest plot comparing postoperative mJOA score for ACDF, hybrid fusion, and ACCF group. SD, standard deviation; IV, inverse variance; CI, confidence interval; df, degrees of freedom; ACDF, anterior cervical discectomy and fusion; mJOA, modified Japanese Orthopedic Association scale; ACCF, anterior cervical corpectomy fusion.

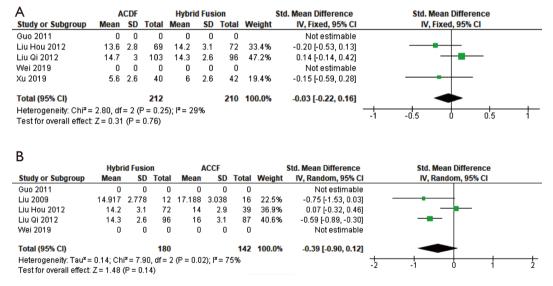


Figure 6 Forest plot comparing postoperative NDI score for ACDF, hybrid fusion, and ACCF group. SD, standard deviation; IV, inverse variance; CI, confidence interval; df, degrees of freedom; ACDF, anterior cervical discectomy and fusion; ACCF, anterior cervical corpectomy fusion; NDI, Neck Disability Index.

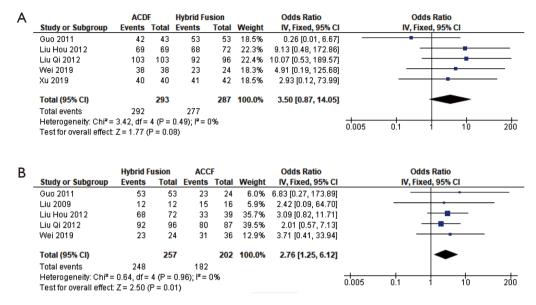


Figure 7 Forest plot comparing fusion rates for ACDF, hybrid fusion, and ACCF group. SD, standard deviation; IV, inverse variance; CI, confidence interval; df, degrees of freedom; ACDF, anterior cervical discectomy and fusion; ACCF, anterior cervical corpectomy fusion.

higher than in the ACCF group (94.4–100% vs. 84.6– 95.8%, respectively; OR 2.76; 95% CI, 1.25, 6.12; P=0.01) (*Figure 7B*).

Complications

Five studies compared the frequency of total complications in ACDF and hybrid fusion groups. The total complication rates were similar between the two groups (OR 0.70; 95% CI, 0.44, 1.13; P=0.14). Five studies comparing complication rates in hybrid fusion and ACCF groups showed significantly lower complication rates in the hybrid group compared to ACCF group (0–22.92% vs. 6.2–43.6%, respectively; OR 0.54; 95% CI, 0.34, 0.84; P=0.006). Four studies compared the frequency of implant failure or mesh subsidence in hybrid fusion and ACCF. The frequency of implant failure or mesh subsidence in the hybrid group ranged from 4.17% to 9.4% fusion which was significantly lower than in ACCF that ranged from 8.05% to 19.4% (OR 0.42; 95% CI, 0.20, 0.88; P=0.02).

Discussion

Degenerative cervical myelopathy, with an annual incidence of 41 per million in North America, is among the most common indications for cervical spine surgery (1,2). While cervical myelopathy has a variable natural history, patients often experience progressive symptoms that ultimately

require surgical intervention (2,3,46). Although there is a near consensus that patients with progressive degenerative cervical myelopathy require treatment, there is a wide range of surgical option. Retrospective studies have examined anterior versus posterior approaches without clear superiority in radiographic and patient outcomes (2,47). Regarding anterior only approaches, surgeons have utilized a variety of surgical approaches to address spinal cord compression. ACDF has been employed to treat cervical spinal cord compression without postvertebral involvement. When there is stenosis behind the vertebral body, an ACDF may not be sufficient and an ACCF is required. The ACCF's downsides include increased operative blood loss, increased hardware complications including device subsidence, and less ability to create and maintain cervical lordosis. ACDF also has shown to have increased rates of pseudoarthrosis with multi-level procedures (4,43,48,49). We have presented a case series of patients undergoing hybrid fusion and performed a meta-analysis to analyze the superiority of three-level ACDF, two-level corpectomy, and hybrid fusion in the treatment of three-level degenerative cervical myelopathy.

In terms of radiographic parameters, we analyzed cervical lordosis at follow-up and overall change in cervical lordosis. In our case series, we found an average postoperative C2–7 cervical lordosis of 11.8 degrees which was less than the range of average postoperative C2–7 cervical lordosis of

the hybrid group included in the meta-analysis. When comparing hybrid fusion to ACCF there was a statistically significant increase in postoperative cervical lordosis in the hybrid group. Regarding the change in cervical lordosis from preoperative to follow-up; ACDF showed a statistically significant increase when compared to hybrid fusion which showed a statistically significant increase compared to ACCF. Zhao et al. 2018, a meta-analysis that analyzed pre and postoperative C2-7 cervical lordosis between ACDF and hybrid groups were additionally unable to find a statistically significant difference in postoperative lordosis between the two procedures (50). Shamji et al. 2013 also showed a statistically significant increase in change in cervical lordosis with ACDF as compared to HS (4). Comparing hybrid fusion to ACCF, Liu et al. 2015 showed a trend towards increased postoperative C2-7 cervical lordosis in the hybrid group however their results, unlike ours, were not statistically significant (51). A notable difference between their meta-analysis and ours, however, is that they included Odate et al. 2016, a study that focused on fourlevel degenerative cervical myelopathy and included threelevel corpectomies in the ACCF group (31). We excluded that study as our meta-analysis was focused only on threelevel procedures.

Regarding PRO we found an average postoperative JOA of 11 (range, 6-14) in our case series of hybrid fusion patients which was less than the mean postoperative JOA included in the meta-analysis. We were unable to find a statistically significant difference between the three surgical approaches with meta-analysis in terms of JOA at follow up, JOA recovery rate, and NDI at follow up. However, when Wei et al. 2019 was removed from the meta-analysis, there was a statistically significant increase in postoperative JOA in the ACDF group over the hybrid group and a near statistically significant increase in postoperative JOA in the ACCF group over the hybrid group. The reason that results from Wei et al. 2019 favored hybrid fusion in terms of postoperative JOA when compared to ACCF and ACDF is unclear but represents a fairly significant outlier in data collected thus far. Wei et al. 2019 only included patients with intramedullary increased signal intensity (ISI) on preoperative MRI and showed that patients undergoing hybrid fusion had a statistically significant improvement in ISI over patients undergoing ACDF or ACCF, this factor may explain why postoperative JOA (a clinical measure of cervical myelopathy) was so favored in the hybrid group (43). Our outcomes agree with Zhao et al., however regarding postoperative JOA, only agree with the conclusions of Shamji et al. when Wei et al. is excluded (4,50).

We analyzed multiple operative factors including estimated blood loss, operative time, fusion, and complications. Regarding operative blood loss, our case series of 10 patients undergoing hybrid fusion for threelevel degenerative cervical myelopathy had an average estimated blood loss of 107.5 mL, significantly less than average estimated blood loss in the hybrid group used in meta-analysis. From our meta-analysis, we found that patients undergoing three-level ACDF had significantly less blood loss than patients undergoing hybrid fusion who in turn had significantly less blood loss than patients undergoing two-level ACCF. This relationship directly follows the number of corpectomies that occur in each procedure, which is where the majority of operative blood loss occurs and has been previously shown in other metaanalyses that involve hybrid fusion (50,51). Regarding operative time our meta-analysis failed to show a statistically significant difference between the three procedures. Regarding fusion, our case series had a postoperative fusion rate of 100%, however, one patient did require additional posterior fusion during the initial hospitalization, although this was reportedly for persistent stenosis. Regarding hybrid fusion and ACCF, our meta-analysis definitively showed that hybrid fusion had a statistically significant increased fusion rate. This is synonymous to the conclusion of meta-analysis Liu et al. 2015 (51). The overall conclusion that fusion rate is highest in ACDF followed by hybrid Fusion and lowest in ACCF may be supported by the decreased rigidity of a multilevel construct (48). The risk of instrumentation failure after a two-level corpectomy is well known and posterior instrumentation is often required especially in patients with significant loss of cervical lordosis. The total complication rate from our case series was 40% with postoperative dysphagia being most common. This is higher than the total complication rate from our meta-analysis which ranged from 0% to 22.9%. Regarding ACDF and hybrid fusion, our meta-analysis showed a trend towards decrease in frequency of total complications in the ACDF group however this was not statistically significant. Zhao et al. 2018 did show a statistically significant decrease in the frequency of complications with ACDF compared to hybrid fusion, however, used a publication in their meta-analysis which is not available to be read in English. Without that publication, they would have been unable to show significance (50). Regarding hybrid fusion and ACDF, our meta-analysis did show a statistically significant decrease in the frequency of complications with hybrid fusion. This

agrees with previous data shown by meta-analysis Liu *et al.* (51). When further analyzing complications for implant failure and corpectomy device subsidence we found a statistically significant decrease in the hybrid group when compared to the ACCF group.

Two biomechanical studies have been done that explain some of the relationships shown in our meta-analysis. Singh et al. compared the rigidity of three anterior cervical constructs (ACDF, ACCF, HS) using three-dimensional motion analysis (6). The authors noted that multiple ACDF and HS provided a higher degree of rigidity in flexion/extension and lateral bending compared to twolevel ACCF. Hussain et al. examined motion, disc stress, and facet loading at the adjacent level in three anterior cervical techniques using finite element modeling (52). They found that the ACCF model showed the greatest disc stress and facet loading followed by HS and ACDF. In addition, these biomechanical changes were seen more in the cephalad adjacent level. Anterior cervical plating after HS constructs has more points of fixation than ACCF allowing for more rigidity and decreased modes of failure. Together these biomechanical properties may explain why hybrid fusion is superior to ACCF in fusion percentage, frequency of complications, frequency of implant failure/ mesh subsidence, and C2-7 cervical lordosis at follow-up.

There are several limitations to our study. No randomized-controlled trials were included in our metaanalysis and the included data was all retrospective. Articles were limited to English only which did leave out previous articles included in the meta-analysis of this topic. Relatively few studies, although more than in any other meta-analysis on this topic, were included and increasing the number of studies may increase the power to identify more statistically significant relationships on this topic. Follow-up was relatively short in the included studies.

Conclusions

To our knowledge this study presents the largest published meta-analysis of patients undergoing three-level ACDF, two-level corpectomy, and hybrid fusion in the treatment of three-level degenerative cervical myelopathy. We have shown that in comparison to three-level ACDF, hybrid fusion has a similar outcome, complication, and fusion success rate. Hybrid fusion has increased postoperative C2–7 cervical lordosis, a higher fusion rate, lower total complication rate, lower implant failure/mesh subsidence rate, and lower average blood loss than two-level cervical

corpectomy. A hybrid cervical fusion for patients with postvertebral degenerative cervical myelopathy and three-level degenerative cervical myelopathy is preferable to a twolevel cervical corpectomy.

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Footnote

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Table S1 Summary of studies included in the meta-analysis

Study	Liu e <i>t al.</i> 2009	Guo <i>et al.</i> 2011	Liu Hou <i>et al.</i> 2012	Liu Qi et al. 2012	Xu <i>et al.</i> 2019	Wei <i>et al.</i> 2019
OR time (minutes)	NR	ACDF: 97.4±17.1; hybrid: 105.9±19.1; ACCF: 119.2±16.4	ACDF: 143.6±31.7; hybrid: 129.4±25.9; ACCF: 116.5±29.8	NR	ACDF: 112.4±23.7; hybrid: 126.5±28.4	ACDF: 142.2±13.8; hybrid: 125.6±5.4; ACCF: 112.2±12.3
EBL (mL)	NR	ACDF: 143.3±72.7; hybrid: 161.6±78.8; ACCF: 208.5±136.9	ACDF: 107.5±49.6; hybrid: 141.5±52.8; ACCF: 172.3±68.2	NR	ACDF: 172.1±32.2; hybrid: 193.2±33.3	ACDF: 102.3±8.3; hybrid: 141.3±10.1; ACCF: 161.5±25.5
Fusion rate (%)	HS: 100; ACCF: 94	ACDF: 97.7%; hybrid: 100%; ACCF: 95.8%	ACDF: 100; hybrid: 94.4; ACCF: 84.6	ACDF: 100; hybrid: 95.83; ACCF: 91.95	ACDF: 100; hybrid: 97.6	ACDF: 100%; hybrid: 95.8%; ACCF: 86.1%
Total complication rates	Hybrid: 0%; ACCF: 6.2%	ACDF: 2.3%; hybrid: 15.1%; ACCF: 25%	ACDF: 21.7%; hybrid: 22.2%; ACCF: 43.6%	ACDF: 15.53%; hybrid: 22.92%; ACCF: 26.44%	ACDF: 2.5%; hybrid: 4.8%	ACDF: 15.8%; hybrid: 16.7%; ACCF: 41.7%
Dysphagia/hoarseness	NR	NR	ACDF: 15.94%; hybrid: 12.5%; ACCF: 15.38%	ACDF: 9.7%; hybrid: 8.3%; ACCF: 5.7%	ACDF: 2.5%; hybrid: 4.8%	ACDF: 10.5%; hybrid: 8.3%; ACCF: 11.1%
Hematoma	0%	ACDF: 0%; hybrid: 1.9%; ACCF: 4.2%	ACDF: 1.4%; hybrid: 0%; ACCF: 2.6%	NR	0%	ACDF: 0%; hybrid: 0%; ACCF: 2.8%
C5 palsy	HS: 0%; ACCF: 6.2%	ACDF: 0%; hybrid:1.9%; ACCF: 0%	ACDF: 2.9%; hybrid: 2.8%; ACCF: 10.3%	ACDF: 3.88%; hybrid: 8.33%; ACCF: 11.49%	0%	ACDF: 2.6%; hybrid: 4.2%; ACCF: 2.8%
Implant failure	0%	ACDF: 0%; hybrid:0%; ACCF: 4.2%	See next column	See next column	0%	See next column
Titanium mesh subsidence	0%	ACDF: NA; hybrid: 9.4%; ACCF: 12.5%	see next column	see next column	0%	see next column
"Graft migration, collapse or displacement" = implant failure + mesh subsidence	0%	ACDF: 0%; hybrid: 9.4%; ACCF: 16.7%	ACDF: 0%; hybrid: 5.5%; ACCF: 12.82%	ACDF: 0%; hybrid: 4.17%; ACCF: 8.05%	0%	ACDF: 0%; hybrid: 4.2%; ACCF: 19.4%
CSF leak	0%	ACDF: 0%; hybrid: 1.9%; ACCF: 4.2%	ACDF: 1.4%; hybrid: 0%; ACCF: 0%	ACDF: 1.94%; hybrid: 1.04%; ACCF: 0%	0%	ACDF: 5.3%; hybrid: 0%; ACCF: 2.8%
CL preop (degrees)	NR	ACDF: -0.1±12.6; hybrid: 9.7±8.6; ACCF:8.7±9.5	ACDF:11.87±11.71; hybrid: 13.75±10.98; ACCF: 12.07±11.82	NR	ACDF: 11.8±9.4; hybrid: 12.5±9.9	ACDF: 11.9±5.0; hybrid: 12.8±2.0; ACCF: 12.4±5.2
CL postop (degrees)	NR	ACDF: 14.9±7.2; hybrid: 17.8±7.7; ACCF: 11.4±7.5	ACDF: 24.27±10.17; hybrid: 23.21±9.55; ACCF: 15.63±12.41	NR	ACDF: 18.2±8.0; hybrid: 14.3±7.4	ACDF: 16.8±5.4; hybrid: 17.3±1.9; ACCF: 14.1±5.1
CL change (degrees)	NR	ACDF: 15.1±9.9; hybrid: 8.1±5.2; ACCF: 2.7±4.5	NR	NR	ACDF: 6.5±7.7; hybrid: 1.8±8.0	ACDF: 4.9±3.7; hybrid: 4.5±0.4; ACCF: 1.7±3.2
JOA initial	HS: 11.167±0.807; ACCF: 10.875±0.646	ACDF: 8.3±1.7; hybrid: 8.1±2.2; ACCF: 7.7±1.6	ACDF: 10.8±1.8; hybrid: 11.2±1.9; ACCF: 10.6±1.4	ACDF: 10.2±2.7; hybrid: 11.3±2.5; ACCF 10.7±3.1	ACDF:10.6±2.0; hybrid: 10.3±1.9	ACDF: 10.1±0.5; hybrid: 10.3±0.7; ACCF: 10.2±1.0
JOA follow up	HS: 14.333±0.536; ACCF: 14.344±0.651	ACDF: 13.7±1.9; hybrid: 13.1±2.3 ACCF: 13.0±2.0	ACDF: 14.1±1.6; hybrid: 13.8±1.9; ACCF: 14.5±1.8	ACDF: 14.8±2.2; hybrid: 13.9±2.8; ACCF: 14.5±2.7	ACDF: 14.0±1.8; hybrid: 13.6±2.1	ACDF: 13.6±0.7; hybrid: 14.9±0.8; ACCF: 14.3±0.6
JOA recovery rate (percentage)	HS: 55.833±3.961; ACCF: 56.831±8.850	ACDF: 60±20; hybrid: 60±20 ACCF: 60±10	NR	NR	NR	ACDF: 50.8±9.0; hybrid: 69.1±10.8; ACCF: 59.4±5.7
NDI initial	HS: 34.250±2.832; ACCF: 34.625±3.364	NR	ACDF: 35.1±2.9; hybrid: 34.7±2.6; ACCF: 35.3±3.0	ACDF: 35.6±3.3; hybrid: 34.9±2.9; ACCF: 35.2±2.8	ACDF: 28.6±6.8; hybrid: 27.1±7.0	NR
NDI follow up	HS: 14.917±2.778; ACCF: 17.188±3.038	NR	ACDF: 13.6±2.8; hybrid: 14.2±3.1; ACCF: 14.0±2.9	ACDF: 14.7±3.0; hybrid: 14.3±2.6; ACCF: 16.0±3.1	ACDF: 5.6±2.6; hybrid: 6.0±2.6	NR

ACDF, anterior cervical discectomy and fusion; ACCF, anterior cervical corpectomy fusion.