Subsidence following anterior lumbar interbody fusion (ALIF): a prospective study

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Background: Anterior lumbar interbody fusion (ALIF) is a widely used surgical technique for disorders of the lumbar spine. One potential complication is the subsidence of disc height in the post-operative period. Few studies have reported the rate of subsidence in ALIF surgery prospectively. We prospectively evaluated the rate of subsidence in adult patients undergoing ALIF.

Methods: Results were obtained by reviewing scans of 147 patients. Disc heights were measured on radiographic scans taken pre-operatively in addition to post-operatively immediately, at 6 weeks and at 18 months. The anterior and posterior intervertebral disc heights were measured. Subsidence was defined as greater than or equal to 2 mm loss of height.

Results: A total of 15 patients (10.2%) had subsidence, with 7 being male. Each case was of delayed cage subsidence (DCS) >6 weeks postoperatively. The mean subsidence was 4.7 mm (range, 2.4–7.8). Mean anterior disc height was 8.6±0.4 mm preoperatively, which improved to 15.1±0.5 mm at latest follow-up. Mean posterior disc height was 4.7±0.2 mm preoperatively, which improved to 8.7±0.4 mm at latest follow-up. The mean lumbar lordosis (LL) angle was 42.5°±10.8° and the mean local disc angle (LDA) was 6.7°±4.0°. The 91.2% (n=114/125) of patients with appropriate radiological follow-up demonstrated fusion by latest follow-up. There was no correlation between subsidence rate with patient reported outcomes [Visual Analog Scale (VAS), Oswestry Disability Index (ODI) and Short Form 12 Item survey (SF-12)] and fusion rates. There was a significant negative correlation between LL and extent of subsidence (Pearson correlation =−0.754, P=0.012).

Conclusions: In conclusion, we found that the subsidence rate at follow-up was generally low following standalone ALIF for this patient series. Patient clinical outcomes and bony fusion rates were not significantly influenced by subsidence.

Keywords: Subsidence; anterior lumbar interbody fusion (ALIF); disc height; prospective; cohort study; surgery; spine; radiographic; fusion

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Introductions

Vertebral interbody fusion is a surgical intervention commonly undertaken in selected patients with discogenic pain and/or mechanically unstable spinal levels (1,2). The most common location for interbody fusion is in the lumbar spine, as this region experiences the most mechanical strain and is most prone to pathology (1). Although multiple approaches can successfully achieve interbody fusion, an anterior surgical approach is widely accepted as having an acceptable success rate with few complications (1,3,4).

Anterior lumbar interbody fusion (ALIF) has become
a widely utilized surgical intervention across the world. Known complications have been studied in ALIF and correlated with potential clinical outcomes. One of these potential complications is the subsidence of disc height in the post-operative period (5-7). Previous research has implied that disc height is increased acutely following surgery as a factor of operative distraction, but decreases over the next weeks to months (6). Generally the disc height has not been shown to decrease to less than the preoperative disc height, however this has been documented in some cases (6,8,9). The time frame of subsidence has been reported as varying from 1 week to 8 months, presenting a large, non-specific time window (9).

Although it was previously concluded that implant subsidence was not correlated with clinical outcome, no trials have consequently shown that mechanical stability is unaffected by subsidence, which is empirically of concern. The primary hypothesis of this paper is that subsidence is more prevalent than previously believed, being seen in up to 100% of ALIF cases, and is an acceptable aspect of the progression to fusion with this technique.

Disc height has previously been estimated using a variety of techniques, all of which use substitute measurements to extrapolate an average disc height. Proxy measurements have included: average disc height using the most anterior and most posterior points of adjacent end-plates, mid-point height measured perpendicular to the inferior endplate, and measurement of the mid-point distance from the inferior endplate of the level above, to the inferior endplate of the level in question (10-12).

This study measured disc heights in 147 patients operated from 2011 to 2013 by the same surgical team, pre-operatively, immediately post-operatively, 6 weeks post-operatively and 6 months postoperatively and/or most recent scan. To the authors’ knowledge, this is one of the largest and most extensive assessment of subsidence in ALIF literature to date.

Methods

Results were obtained by reviewing scans of 147 patients, all of whom underwent surgery by the same senior neurosurgeon across two hospitals. Disc heights were measured on CT scans taken pre-operatively, immediately post-operatively, on erect X-rays at 6 weeks post-operatively, CT scans at 6-month-postoperatively when evaluating fusion and using the latest available CT scan performed as clinically indicated, such as suspicion of incomplete fusion. Clearance for the prospective study was obtained through the Human Research Ethics Committee of New South Wales Health (reference No. 11/183). Patients undergoing ALIF surgery were included in the study with indications: degenerative disc disease without radiculopathy, degenerative disc disease with radiculopathy, spondylolisthesis, failed posterior fusion, adjacent segment disease and scoliosis requiring correction. Exclusion criteria were patients with concurrent local or systemic infection, neoplasia, significant cardiac disease, fever (>38.5 °C), or metal allergy; as well as patients who were pregnant or breast-feeding, who were mentally incompetent, who had a history of alcohol or drug abuse, and who were at increased risk of vascular or bowel complications related to the anterior approach.

Patients received stand-alone PEEK integral cage devices. In particular, the vast majority (89.1%) received the SynFix-LR PEEK integral cage device (DePuy Synthes, West Chester, PA, USA) with four diverging intrinsic screws and anterior locking plate, without anterior tension band plating nor posterior instrumentation. The implant sizing varied across patients in accordance with the disc height of neighbouring healthy lumbar discs, ranging from 12–19 mm height with either 8° or 12° lordotic angle to ensure sufficient distraction. Bone graft substitute i-FACTOR (Cerapedics, Westminster, CO, USA) was used for 136 patients and is comprised of anorganic bone matrix bound to anorganic P-15 small peptide, together facilitating attachment of osteogenic cells. For the remaining 11 patients, recombinant human bone morphogenetic protein 2 (rhBMP2) (Medtronic Sofamor Danek, Memphis, TN, USA) was used. In 7 (4.8%) patients there had been a previous fusion performed and no patients required additional posterior pedicle screw fixation to augment ALIF.

The radiological parameters for subsidence and fusion were measured by a neurosurgeon (P.R.). Fusion rates were assessed using reconstructed axial and coronal fine-cut CT scans. Criteria for established fusion were bridging trabecular formation across the intervertebral disc space with the absence of radiolucency spanning more than half of the implant. The anterior and posterior intervertebral disc heights were measured and averaged. Endplate levels were taken as a straight-line average of the endplate as seen on the most central image in all planes, using the most anterior and posterior points excluding osteophytes. Osteophytes were identified as superficial extrusions of bone anteriorly or posteriorly beyond the main vertebral
body. This allows for reliable disc height estimation without being confounded by central disc erosion. However, it can be a difficult measurement in images with significant anterolisthesis, retrolisthesis, or osteophyte formation. The local disc angle (LDA) was determined by the angle formed by the intersection of the inferior endplate line and the superior endplate line of the index disc level. Lumbar lordosis (LL) was measured between the superior endplate of L-1 to the superior endplate of S-1 using the Cobb method.

Subsidence was defined as greater than or equal to 2 mm loss of height. Early subsidence was defined as occurring within the first 6 weeks postoperatively and delayed subsidence as after 6 weeks postoperatively. Cranial (inferior endplate) subsidence was classed as type 2 and caudal (superior endplate) was considered type 1 subsidence as per Malham et al.’s study (11).

Clinical outcome was measured preoperatively and postoperatively using the Oswestry Disability Index (ODI) and the Visual Analog Scale (VAS) back pain score. Questionnaire data from the Short Form 12 Item survey (SF-12) were compiled in a custom-designed database. Preoperative and postoperative study outcomes were examined with analysis of variance and repeated-measures general linear models adjusted for age and sex. Analyses were based on 2-sided tests with values of P<0.05 considered significant with Bonferroni correction when appropriate. Correlation studies were performed using Pearson’s coefficients to investigate relations between changes in radiologic parameters and improvements in VAS, ODI, and SF-12 scores. Data analysis and statistical evaluation was conducted using IBM SPSS Statistics 22 (IBM Corporation, Armonk, NY, USA).

### Results

#### Patient demographics

A total of 147 patients underwent ALIF in our study. The mean age of the study group was 57.3±13.6 years. Sixty-five of the patients were female (44.2%). From the total cohort, 17 patients (11.6%) were smokers and 7 patients (4.8%) had diabetes. Only one patient used corticosteroids, due to comorbid Crohn’s disease. 74 patients (50.3%) underwent ALIF at the L4/L5 level, 95 patients (64.6%) at the L5/S1 level and 29 patients (19.7%) underwent multilevel fusion (Table 1). Indications for ALIF are detailed in Table 2.

#### Operative outcomes

Fifteen patients (10.2%) demonstrated subsidence of mean 4.7 mm (range, 2.4–7.8). These were all delayed cage subsidence (DCS). The mean age of the patients with subsidence was 67 years and the male to female ratio was equivocal (7:8). Four of these patients were overweight and another had pseudoarthrosis.

There were 15 patients with type 1 (caudal endplate) subsidence; 3 with anterior, 2 with posterior and 10 with both anterior and posterior subsidence. No patients had type 2 (cranial endplate) subsidence alone, but 2 patients had both type 1 and type 2 subsidence.

The preoperative anterior disc height was 8.6±0.4 mm, which improved to 16.9±0.4 mm immediately postoperatively, 15.5±0.5 mm at 6-week follow-up, and 15.1±0.5 mm at latest follow-up. For posterior disc height, preoperatively this was 4.7±0.2 mm, which improved to 9.1±0.3 mm immediately postoperatively, 9.4±0.4 mm at 6-week follow-up, and 8.7±0.4 mm at latest follow-up (Figure 1).

The 91.2% (n=114/125) of patients with appropriate

### Table 1 Baseline demographics

<table>
<thead>
<tr>
<th>Demographic</th>
<th>Mean ± standard deviation; or proportion (%) (n=147)</th>
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<tbody>
<tr>
<td>Age (years)</td>
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<tr>
<td>Females</td>
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<tr>
<td>Smokers</td>
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</tr>
<tr>
<td>Diabetes</td>
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<tr>
<td>L4/L5 level</td>
<td>50.3</td>
</tr>
<tr>
<td>L5/S1 level</td>
<td>64.6</td>
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<tr>
<td>Multilevel</td>
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</tbody>
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### Table 2 Indications for surgery

<table>
<thead>
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<th>Indication</th>
<th>Proportion of patients (%)</th>
</tr>
</thead>
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<tr>
<td>DDD without radiculopathy</td>
<td>14.8</td>
</tr>
<tr>
<td>DDD with radiculopathy</td>
<td>54.8</td>
</tr>
<tr>
<td>DDD with radiculopathy &amp; stenosis</td>
<td>0.7</td>
</tr>
<tr>
<td>DDD with radiculopathy &amp; spondylolisthesis</td>
<td>1.5</td>
</tr>
<tr>
<td>Spondylolisthesis</td>
<td>28.2</td>
</tr>
</tbody>
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radiological follow-up demonstrated fusion by latest follow-up whilst 11 patients did not demonstrate fusion. Appropriate CT scans at 6 months were not available for 22 patients. In our study, only 19.7% of patients received multilevel ALIF, that is, ALIF at two or more adjacent levels. Of these, only 2 patients received three-level ALIF, for which all segments fused, and only 1 received four-level ALIF, for which only immediate postoperative follow-up was available.

The mean LL angle was 42.5°±10.8° and the mean LDA was 6.7°±4.0°. The mean cage height, length and width was 13.4±1.4, 37.8±1.6 and 30.0±0.4 mm respectively. The mean cage lordosis was 9.0°±1.8°.

**Functional outcomes**

All functional outcomes demonstrated improvement, with reduction in VAS and ODI scores and increase in SF-12 scores. Preoperative VAS pain score was mean 7.1±0.2, reducing post-operatively to 2.7±0.2 (P<0.0001) (Figure 2). The preoperative ODI was also significantly reduced post-operatively (P<0.0001) from mean 57.8±2.0 to 28.8±1.8 (Figure 3). Preoperative SF-12 physical component summary (PCS) was 33.2±1.7, which was increased to 41.7±0.9 post-operatively (P<0.0001). Similarly, preoperative SF-12 mental component summary (MCS) score was 38.0±1.2, and increased to 48.9±1.0 post-operatively (P<0.0001) (Figure 4).

**Correlation analysis**

No significantly association was found between subsidence rates and changes in VAS (P=0.36), ODI (P=0.55), SF-12 PCS (P=0.69) or SF-12 MCS (P=0.64). There was also no significant association between degree of subsidence with fusion rates (P=0.85). In our cohort, we did not find significant correlations between subsidence and demographic factors including age, sex, spondylolisthesis,
steroid use, diabetes, smoking, BMI or LDA. There was a significant negative correlation between LL with extent of subsidence (Pearson correlation = -0.754, P=0.012), i.e., the greater the LL correction, the lower the extent of subsidence on follow-up.

Discussion

An anterior approach for lumbar fusion is a procedure used to treat spine instability with the goal of providing a solid and lasting union of the affected vertebrae (1,10,13,14). An anterior approach to the spine, as opposed to a posterior approach, does not disturb the paravertebral muscles and ligaments, which are involved in spine mobility and stability (15). As a result there is improved spinal mobility and decreased muscle pain postoperatively in the anterior approach (16). Anterior exposure also provides greater surgical versatility and exposure intraoperatively (1). In addition, ALIF has been shown to be effective in decompressing the intervertebral foramen and restoring disc interspace height (9).

A complication that can arise from ALIF is subsidence, which is a decrease in the vertical height of the vertebral disc space prior to complete fusion. Subsidence is an important consideration as the reduction in disc space can detrimentally affect mechanical correction and clinical outcomes (10). In the present study, we found that the overall subsidence rate at follow up was generally low following ALIF for this patient series. In the patients where subsidence was observed it was mostly attributed to the caudal endplate. There was no significant correlation between degree of subsidence with changes in clinical outcomes including VAS back pain, ODI, SF-12 PCS and MCS scores or fusion rates. This result is broadly consistent with prior published studies focusing on posterior lumbar interbody fusion (17-20). Collectively, these results suggest that bony fusion as well as patient reported measures of pain and disability were not significantly impaired by subsidence of the ALIF cage into the vertebral body.

A number of factors have been proposed to influence subsidence rates. Inappropriate intraoperative endplate preparation of the vertebral bodies with extensive endplate resection has been suggested as one factor contributing to subsidence (21,22). However, endplate preparation can be controlled in ALIF as there is direct visualization of the disc space and removal of the ALL creates a non-rigid

![Figure 3 ODI scores preoperative and postoperative. ODI, Oswestry Disability Index.](image)

![Figure 4 Preoperative and postoperative SF-12 PCS and MCS. SF-12, Short Form 12 Item survey; PCS, physical component score; MCS, mental component score.](image)
space during disc preparation. Marchi et al. demonstrated that in stand-alone lateral interbody fusion, wide cages avoid subsidence and restore segmental lordosis to a greater extent (23). Furthermore, it has been found that ALIF conducted solely using bone grafts can result in high subsidence rates of up to 100% (6,8). Metal interbody cages have thus been used as an alternative as they can better maintain disc space height during the process of bone graft incorporation into the fusion mass (9). Sandhu et al. (24) found that subsidence in sheep was reduced when a threaded cage was used as opposed to an iliac crest bone graft. Further clinical studies have also found lower subsidence rates when using a metal interbody cage in ALIF (9,10,25). In the present study, we also found that improved correction of segmental LL was significantly associated with reduced subsidence extent on follow-up, which further supports the notion of adequate lordosis correction during stand-alone ALIF.

More recently, non-absorbable, biocompatible PEEK cages have been used as they are radiolucent and have similar mechanical properties to bone (16,26,27). The radiolucency of this PEEK cage material has allowed for improved ease and clarity during assessment of fusion (28,29). The PEEK cages have also been proposed to have load-bearing properties superior to metal cages, and thus can further reduce subsidence rates (16). This would support the results of our current study, which found a low 8.7% rate of subsidence in standalone ALIF using PEEK cages. In the current study, the use of cages with a modulus of elasticity close to bone in our cohort receiving stand alone ALIF resulted in significantly improved functional outcomes with reduced VAS and ODI scores along with higher SF-12 MCS and PCS scores post-operatively.

In terms of the site of implant subsidence, the predominated location was found to be in the caudal superior endplate. This is most likely due to variations in strength across endplate regions (9). In a study using human cadavers, Grant et al. (30) found that the posterior region of the endplate was stronger than the anterior region, the vertebral periphery was stronger than the center and the caudal inferior endplate was approximately 40% stronger than the cervical superior endplate. The results of this biomechanical study are consistent with not only our findings but also that of Choi et al. (9), who reported a subsidence rate of 39.1% at the superior endplate and 17.3% at the inferior endplate. Kumar et al. (31) also found greater subsidence rates at the posterior endplate, supporting the findings of Grant et al. (11,30). Variations in bone mineral density have been shown to play a role in subsidence and thus could potentially explain the differences in bone strength at various regions of the endplate (32).

All the patients in this study developed DCS, which was defined as occurring at greater than 6 weeks post-operatively. This finding was consistent with that of Choi et al. (9), who reported a median period for cage subsidence at 2.75 months. Cheung et al. (8) also reported onset of subsidence mostly within the first 3 months after ALIF using an iliac bone graft. However, Kumar et al. (31) found that subsidence occurred within 15 days when using a femoral bone graft. Thus, there are large variations in subsidence onset time reported in the literature when using both interbody cages and bone grafts (8,9,31). The large variations in subsidence onset as reported in the literature may be due to variations in patient BMI between studies. A higher BMI has been associated with greater subsidence as it can accelerate disc degeneration and increase spinal instability (33). Differences in cage design have also been shown to have an effect on the strength of the implant and thus its ability to resist subsidence (34). Additionally, surgical technique has also been implicated, with rigorous endplate preparation resulting in loss of bony endplate, leading to subsidence.

It is important to recognize the limitations of this study (35). Firstly, there was significant heterogeneity in the study due to usage of different graft materials along with innate variations in the included patient population. Although bone mineral density has been implicated as a potential contributor to subsidence, this was not measured nor controlled for in our study (32). The patient series was also from a single spine surgeon, limiting the external validity of the findings despite allowing consistency in surgical technique. Nevertheless our study is the largest prospective study investigating the subsidence in ALIF to date. Further, the measurement of subsidence was conducted using fine cut CT scans and the technique was standardized across all patients.

**Conclusions**

In conclusion, we found that the subsidence rate at follow-up was generally low following standalone ALIF for this patient series. In the patients where subsidence was observed, it was mostly attributed to the caudal endplate. There was no significant correlation found between degree of subsidence with improvements in clinical outcomes in terms of VAS, ODI and SF-12 scores or fusion rates.
Acknowledgements

None.

Footnote

Conflicts of Interest: The authors have no conflicts of interest to declare.

Ethical Statement: The study was approved by the Human Research Ethics Committee of New South Wales Health (No. 11/183) and written informed consent was obtained from all patients.

References


Contributions: (I) Conception and design: PJ Rao, RJ Mobbs; (II) Administrative support: RJ Mobbs; (III) Provision of study materials or patients: RJ Mobbs; (IV) Collection and assembly of data: PJ Rao, K Phan, MM Maharaj, G Gaing; (V) Data analysis and interpretation: PJ Rao, K Phan, MM Maharaj; (VI) Manuscript writing: All authors; (VII) Final approval of manuscript: All authors.