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1 **Comparison of intra subject repeatability of quantitative fluoroscopy and static**
2 **radiography in the measurement of lumbar intervertebral flexion translation**

3

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23 **Comparison of intra subject repeatability of quantitative fluoroscopy and static**
24 **radiography in the measurement of lumbar intervertebral flexion translation**

25

26 Low back pain patients are sometimes offered fusion surgery if intervertebral translation,
27 measured from static, end of range radiographs exceeds 3mm. However, it is essential to
28 know the measurement error of such methods, if selection for back surgery is going to be
29 informed by them. Fifty-five healthy male (34) and female (21) pain free participants aged
30 21-80 years received quantitative fluoroscopic (QF) imaging both actively during standing
31 and passively in the lateral decubitus position. The following five imaging protocols were
32 extracted from 2 motion examinations, which were repeated 6 weeks apart: 1. Static during
33 upright free bending. 2. Maximum during controlled upright bending, 3. At the end of
34 controlled upright bending, 4. Maximum during controlled recumbent bending, 5. At the
35 end of controlled recumbent bending. Intervertebral flexion translations from L2-S1 were
36 determined for each protocol and their measurement errors (intra subject repeatability)
37 calculated. Estimations using static, free bending radiographic images gave measurement
38 errors of up to 4mm, which was approximately twice that of the QF protocols. Significantly
39 higher ranges at L4-5 and L5-S1 were obtained from the static protocol compared with the
40 QF protocols. Weight bearing ranges at these levels were also significantly higher in males
41 regardless of the protocol. **Clinical decisions based on sagittal translations of less than 4mm**
42 **would therefore require QF imaging.**

43

44 **Keywords**

45 low back pain, spinal injuries, spinal surgery, quantitative fluoroscopy, spine, kinematics,
46 quantitative imaging biomarkers.

47

48

49 Introduction

50 Low back pain is responsible for the world's largest number of days lost to disability ¹ and
51 although its diagnosis is often problematical, it is agreed that mechanics generally, and
52 segmental stability in particular, plays a significant role ²⁻⁴. However, the measurement of
53 segmental stability in patients is problematical due to lack of a unified concept of the
54 condition. Yet while biomechanical measurements alone are not considered to be good
55 predictors of prognosis, patients with sufficiently severe symptoms may be offered fusion
56 surgery if intervertebral translation exceeds 4mm⁵. There are many imaging methods for
57 determining this, but practicality and economics dictates that it is generally performed using
58 standing end-range radiographs⁶.

59

60 For this measurement, a radiograph is taken in the neutral standing position and then with
61 the patient flexing forward as far as possible. This is repeated with the patient bending
62 backwards into extension. On the resulting images lines are drawn on adjacent vertebrae
63 from which to measure the translation or sliding movement between vertebrae. This is
64 generally preferred by clinicians to angular movement for the assessment of stability⁷.

65 However, it has long been recognised that inaccuracies and population variations using this
66 technique may limit its usefulness and make selection of a cut off for excessive translation
67 difficult⁸. Static views have also been found to underestimate intervertebral translation
68 compared to dynamic imaging and the lateral decubitus position to better detect excessive
69 motion in spondylolisthesis cases ^{9,10}. Furthermore, complexity increases if the patient also
70 has spinal stenosis ¹¹ or if revision surgery is being considered ¹².

71

72 Recently, advances in fluoroscopic imaging have made it possible to register and track multi
73 segmental vertebral image sequences throughout the entire motion. This method is called
74 quantitative fluoroscopy (QF) and has been able to identify motion patterns that
75 discriminate patients with chronic, nonspecific back pain from pain free controls¹³⁻¹⁷. It has
76 also been used to measure positional changes at individual levels, where for translation, it
77 has been reported to have an accuracy of 0.1mm and inter-observer repeatability of 1.1mm
78 (agreement) and ICC 0.533-0.988 (reliability)^{18 19}. Given the ubiquity of fluoroscopes in
79 general hospitals, these might be repurposed to provide an alternative method for
80 measuring inter vertebral translation in such patients.

81

82 Continuous standardised motion measurement has a number of potential advantages. First,
83 although the motion is not 'naturally performed', controlled motion enables standardisation
84 for trunk range, velocity, ramp up and ramp down speeds and is therefore potentially more
85 reproducible. Second, QF can be conducted either actively weight-bearing or passively in
86 recumbence, to avoid muscle contraction, or guarding, and to test the passive structures
87 with minimal uncontrolled movement variation²⁰. Third, the option of a passive recumbent
88 examination has the advantage of additional patient comfort, where upright bending may
89 be inhibited by pain. Fourth, the range of translation may be measured at the end of the
90 maximum range of the segment, which may not coincide with its range at the end of the
91 trunk bending motion (Fig 1).

92

93 As QF allows for a number of protocols for measuring intervertebral translation it was
94 thought useful to assess the measurement properties of these in terms of random and intra
95 subject variability for measuring maximum displacement. In addition, a direct comparison

96 of end range vs through range translation is lacking, as is measurement during free and
97 guided bending²¹. The main aim of this study was to compare the intra subject variability,
98 or measurement error, of 5 methods for measuring intervertebral flexion translation to
99 determine the level of difference that could be detected by each. The evaluation of
100 extension was not included as the standing range of lumbar spine extension is small (20°)²².

101

102 **Methods**

103 **Participants**

104 Fifty-five healthy control participants were recruited from staff, students and visitors of the
105 AECC University College (Bournemouth, UK). To be eligible, participants had to be aged 21-
106 80 years, BMI<30, with no history of previous back or abdominal surgery or
107 spondylolisthesis, no medical radiation exposure of >8mSV in the previous 2 years and no
108 current pregnancy. Participants also had to have been free of any back pain that limited
109 their normal activity for more than 1 day in the previous year. All imaging was carried out
110 in accordance with AECC UC Local Rules and ethical approval was obtained from the
111 National Research Ethics Service (South West 3, 10/H0106/65). Written Informed consent
112 was obtained from all individual participants included in the study. All images of models
113 were submitted with the express permission and signed informed consent of the model for
114 publication of identifying information/images in an online open-access publication.

115

116 **Data collection**

117 Participants (median age 30 years, range 21 to 69), received fluoroscopic imaging of their
118 lumbar spines during both lying (passive recumbent) and standing (weight-bearing guided)
119 flexion. In passive recumbent flexion. For passive imaging they lay unconstrained in the

120 lateral decubitus position on a motorised table that flexed their upper body to 40° flexion
121 and return during fluoroscopic screening (Atlas Clinical Ltd.) (Fig 2a). They were then
122 imaged whilst weight-bearing, standing with their right side against the motion frame using
123 the same controller apparatus as for the recumbent procedure (Fig 2b).

124

125 With their pelvises stabilised and during active voluntary motion, participants were guided
126 through a standardised range of 60° standing flexion and return by a moving arm. The
127 motion controllers accelerated at 6°s^{-2} for the first second followed by a uniform 6°s^{-1}
128 thereafter. The guiding arm was then removed, and the participants were asked to bend
129 forward freely to the end of their comfortable range (weight-bearing unguided flexion) (Fig
130 2c). Single fluoroscopic images were obtained at the beginning and end of the weight-
131 bearing unguided flexion motion. Fluoroscopic motion sequences were recorded at 15 Hz
132 using a Siemens Arcadis Avantic digital C-arm fluoroscope (Siemens GMBH) and stored in
133 DICOM format. They were then exported to a computer workstation and analysed using
134 manual first image registration (Fig 3) and thereafter using bespoke frame-to-frame
135 tracking using codes written in Matlab (V2011a, The Mathworks Inc). These measurements
136 were repeated 6 weeks later by the same operator using the same equipment at
137 approximately the same time of day for the determination of intra-subject measurement
138 error²¹.

139

140 Image Analysis

141 Sagittal plane translation was calculated using the method of Frobin et al in vertebral body
142 units (VBU) which were converted to millimetres for presentation by multiplying the result
143 by 35 - the standard chosen for vertebral body depth in millimetres²³. In order to address

144 the degree of translation that could be considered excessive, sagittal plane translation of
145 each intervertebral level from L2-S1 was determined and the levels pooled to provide
146 means and upper reference ranges of variation (+1.96SD) for the following five
147 measurement protocols:

- 148 1. Maximum IV translation during passive recumbent flexion
- 149 2. IV translation at maximum bend of passive recumbent flexion
- 150 3. Maximum IV translation during guided weight-bearing flexion
- 151 4. IV translation at maximum bend of guided weight-bearing flexion
- 152 5. IV translation at maximum bend of unguided weight-bearing flexion (reflective of
153 traditional static radiograph acquisition)

154

155 Statistical analysis

156 All data were tested for normality using the Shapiro-Wilk test. The significance of
157 differences was calculated using 2-way paired t-tests for normally distributed data and the
158 Wilcoxon test for non-normal data. Repeatability was calculated using the following
159 formula, where S_w is the within-subject standard deviation. The repeatability coefficient, or
160 measurement error, estimates the magnitude of the within-subject change that can be
161 expected 95% of the time and represents the Minimum Detectable Change (MDC₉₅)²¹.

162 Source data for this study are available by application to the corresponding author.

$$163 \text{ Repeatability coefficient (MDC}_{95}) = 2.77S_w$$

164 The association between test-retest differences and their means were assessed using
165 Kendall's tau. As no significant and/or substantial associations were found, the data were
166 not transformed for the calculation of MDC₉₅.

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Results

Fifty-five participants (21F, 34M) were recruited and all provided complete data. These data were mainly distributed non-normally, resulting in a nonparametric approach to statistical comparisons. Participants' characteristics were: height 1.75m (range 1.53-1.90), weight 74.9kg (range 47.6-112.4) and BMI 24.2 (range 16.9-31.8). The median effective dose received per participant was 0.27mSv for weight bearing motion (range 0.20-0.68), 0.18mSv for recumbent motion (range 0.11-0.31) and 0.04mSv for single frame maximum bend images (range 0.01-0.09).

The median translations for pooled L2-S1 levels were less than 2mm regardless of protocol while the static uncontrolled protocol gave significantly higher translation ranges than any of the controlled protocols ($p < 0.001$, (Wilcoxon) (Table 1.). Intra class correlations were moderate to substantial, showing acceptable reliability for all protocols, however, the measurement error was highest (3.36mm) for the static uncontrolled protocol, compared with the highest error of the controlled protocols (2.14mm). This reflects an error in excess of 200% of the baseline translation for the static protocol compared with a maximum of 163% for controlled weight bearing. Weight bearing measurements, both guided and unguided, gave slightly higher ranges than passive recumbent testing, but similar values when measured at the end of the motion and during it.

191

192 Table 1.

Translation ranges, reliability and measurement error for five measurement protocols (L2-S1 pooled)

Measurement	Protocol	n	Median translation (IQR)		Reliability ICC _{2,1} (95% CI)
			mm		
			Baseline	Follow up	
During motion 40 deg	Passive recumbent	219	0.74 (0.69)	0.86 (0.78)	0.639 (0.528 , 0.724)
End of motion 40 deg	Passive recumbent	219	0.74 (0.58)	0.86 (0.53)	0.611 (0.486 , 0.706)
During motion 60 deg	Active weight bearing	216	1.21 (1.26)	1.21 (1.37)	0.550 (0.413 , 0.655)
End of motion 60 deg	Active weight bearing	216	1.22 (1.05)	1.31 (1.08)	0.782 (0.715 , 0.833)
End of uncontrolled flexion	Active weight bearing	200	1.54 (1.42)	1.47 (1.67)	0.697 (0.605 , 0.768)

193

194 When taken level by level, the median baseline translation of L2-3 was significantly greater
 195 during guided weight bearing continuous measurement than unguided weight bearing static
 196 measurement ($p < 0.001$), whereas the converse was true for L4-5 and L5-S1 ($p < 0.001$)
 197 (Wilcoxon) (Fig4).

198

Figure 4 about here

199 The measurement errors at L4-5 and L5-S1 for static uncontrolled measurements at around
 200 4mm were approximately double those of controlled ones (around 2mm), however, for L2-3
 201 and L3-4 these differences were less marked (Fig 5).

202

203

Figure 5. about here

204 The baseline median translation range at L5-S1, when measured using the static,
 205 uncontrolled maximum bend protocol, was significantly higher for males than for females
 206 ($P < 0.001$) (Mann Whitney). In addition, for the weight bearing controlled bending
 207 protocols, L3-4 and L4-5 ranges were higher for males ($p < 0.01$), while recumbent sequences
 208 measured during the motion gave higher ranges at L2-3 for females ($p < 0.05$). Age above
 209 and below the inter-quartile ranges did not have any significant effect on translation range

210 for any level or protocol. It should also be noted that L5-S1 translation, measured using
211 controlled motion protocols, returned very small values when measured during motion as
212 opposed to at its end, while weight bearing measurements returned more variation and less
213 consistency than recumbent ones (Fig 6 a-e).

214

215 Figure 6a-e about here

216 Discussion

217 This research found that static radiographs gave twice the measurement error of QF and
218 higher L4-5 and L5-S1 ranges when used to measure flexion translation. In effect, this
219 means that it is not possible to detect translation of under 4mm using static radiographs,
220 2.5mm using weight bearing QF or 2.0mm using recumbent QF. Furthermore, the
221 normative ranges for each protocol are different for males and females, but not in older
222 people. A cut-off at 4mm for inferring instability is consistent with much of the literature as
223 reviewed by Leone et al, however, as recognised by Nizard et al, population variation and
224 lack of standardisation have made any such cut off somewhat tenuous^{5,24}. Nevertheless,
225 Posner et al's criterion for selecting patients with instability for fusion treatment, which
226 defines a cut off of 8% of vertebral body depth for anterior translation is generally accepted,
227 although this would amount to only 2.8mm using a standard intervertebral body depth of
228 35mm²⁵⁻²⁷.

229 In this study, measurement at the end of uncontrolled motion using static radiographs was
230 more variable than using QF. At L4-5 and L5-S1, this returned approximately twice the
231 measurement error of the QF protocols, while static, uncontrolled weight bearing
232 measurements were similar to guided weight bearing QF measurements at L2-3 and L3-4.

233 The least population variability and measurement error was found when participants were
234 imaged during passive recumbent motion, as has been recommended for the detection of
235 excessive translation in spondylolisthesis¹⁰.

236

237 The 4mm measurement error for weight bearing, static, unguided, end of range
238 measurements was especially applicable to L4-5 and L5-S1. These levels are frequently of
239 interest in terms of translatory slip, however, this may be uncommon in back pain
240 populations. A recent study of aberrant motion in chronic, nonspecific back pain did not
241 find translation to be greater in patients than healthy controls¹⁷. Even in patients with
242 spondylolisthesis, excessive translation is also not necessarily a feature, while in older
243 individuals with degenerative spinal stenosis, bone loss, arthritic outgrowth and vertebral
244 mal-alignment may make the measurement of translation using any current form of
245 radiographic imaging additionally problematical^{5,28}.

246

247 The tendency for static views, acquired at the end of trunk motion, to give different values
248 from QF may be thought to be because the range of trunk motion at the end of a weight-
249 bearing unguided flexion motion could be greater than 60°, which is the standard range of
250 flexion used for standing guided weight-bearing QF¹⁹. However, free bending resulted in
251 only approximately 0.5mm greater translation than controlled bending to 60°. Indeed, the
252 median ranges of translation found in this study, by all of the protocols, compare favourably
253 with those found in a separate study of healthy volunteers²⁹. However, although studies of
254 intervertebral translation in back pain patients have concluded that it is related to age and
255 disc height, it does not differentiate patients from controls^{17,30,31}. This may be partially a
256 result of the uncontrolled variation associated with current measurement methods.

257 However, composite disc degeneration throughout the lumbar spine has been associated
258 with disproportionate sharing of angular motion between the lumbar spine segments in
259 chronic, nonspecific back pain patients¹⁶. Thus, it may be that it is the distribution of
260 degenerated discs in the lumbar spine, rather than large changes in ranges of motion at
261 individual levels, that is most closely associated with symptoms in chronic, nonspecific low
262 back pain³².

263

264 Finally, the qualitative use of fluoroscopy tends to be associated with prolonged exposures,
265 raising the expectation of higher radiation dosage. However, the QF protocols are, by
266 definition, quantitative and in this study resulted in effective radiation dosages of less than
267 0.3mSv each. This is considerably less than the 1.3mSv quoted as the typical effective dose
268 expected for a series of X-rays of the lumbar spine for diagnostic purposes^{33,34}. This makes
269 continued use of plain radiographs difficult to justify for most cases where degrees of
270 increased translation that are not measurable might be acted upon.

271

272 Limitations

273 The present study did not include extension motion; however, its purpose was to compare
274 radiographic techniques for their measurement properties while minimising radiographic
275 exposure. The levels considered also did not include L1 because the intensifier diameter
276 was too small to permit it.

277

278 Further work

279 These methods, although tested on a healthy asymptomatic population here, have also
280 been utilised to evaluate back pain populations^{16,17}. Therefore, this study should be
281 repeated in symptomatic cohorts to establish repeatability and variability of translation.

282

283 **Conclusion**

284 Quantitative fluoroscopic measurement of lumbar intervertebral flexion translation in
285 healthy control participants during passive recumbent QF gave significantly lower values
286 than static, weight-bearing unguided imaging. It also resulted in lower population variation
287 and approximately half the measurement error, which for static images during uncontrolled
288 motion was in the region of 4mm. Thus, clinical decisions based on smaller amounts of
289 sagittal translation would require QF imaging.

290

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388

389 **Figure legends:**

390 Figure 1. Example of continuous translational motion from L2-S1 in a healthy control

391 participant showing the points of maximum translation (coloured arrows) compared to the
392 point of the patient's maximum trunk bend

393 Figure 2. Dynamic acquisition of fluoroscopy sequences: a) controlled passive recumbent
394 flexion, b) controlled active weight bearing flexion, c) uncontrolled weight bearing flexion

395 Figure 3. Sagittal lumbar spine fluoroscopic image showing computer reference templates

396 Figure 4. Median baseline translations (interquartile range) for each level from L2-S1 for five
397 measurement methods

398 Figure 5. Measurement error (MDC₉₅) for translations for each level from L2-S1 for five
399 measurement methods

400 Figure 6. Box plots showing median intervertebral translations from L2-5 at baseline
401 (hatched box) and follow-up (clear box) measured a) at end of uncontrolled weight bearing

402 flexion b) during controlled weight bearing flexion c) at end of controlled weight bearing

403 flexion d) during recumbent flexion e) at end of recumbent flexion

404