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Less periprosthetic bone loss following the anterolateral approach to the hip compared with the direct lateral approach
A subgroup analysis from a randomized trial in patients with a femoral neck fracture

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Background and purpose — The loss of bone mineral in the proximal femur following hip arthroplasty may increase the fracture risk around uncemented stems. We hypothesized that the surgical approach to the hip might influence bone mineral changes around the femoral stem in patients with a femoral neck fracture (FNF).

Patients and methods — This was a pre-specified subgroup analysis (n = 51) of an ongoing randomized trial (n = 120) in patients with FNF. Participants were allocated to an uncemented hemiarthroplasty inserted through a direct lateral (Hardinge) approach or an anterolateral (modified Watson-Jones) approach. The 51 patients (mean age 83 (70–90) years, 33 women) were measured by dual-energy X-ray absorptiometry (DXA) to assess changes in periprosthetic bone mineral density (BMD).

Results — The mean change in total BMD differed between groups at 12 months in favor of the anterolateral group (4.8%, 95% CI 0.0–9.6; p = 0.05). DXA at 3 months displayed BMD loss in the proximal Gruen zones in the lateral group compared with the anterolateral group. Zone 1 (–5.0% vs. 2.7%), zone 2 (–4.3% vs. 4.1%), zone 6 (–6.5% vs. 0.0%) and zone 7 (–11% vs. –2.4%, all p < 0.05).

Interpretation — DXA measurements in this study indicate that surgical approach to the hip influences periprosthetic BMD. Clinical implications remain uncertain. Our conclusions should be interpreted with caution as we did not perform adjustments for multiple tests, possibly leading to inflation of false-positive findings.

Progressive periprosthetic bone loss around the femoral component is believed to contribute to aseptic loosening (Malchau et al. 1993) and late-occurring fractures around the implant (Lindahl 2007, Langslet et al. 2014). Periprosthetic fractures have emerged as a major reason for revision, especially in the elderly (Thien et al. 2014). Numerous reasons are believed to be responsible for changes in bone mass. Stem design seems to affect bone loss (Karrholm et al. 2002, Grant et al. 2005, Salenmyr et al. 2015) as well as stem sizes. It may be that daily activity, sex, and BMI influences BMD changes around the stem (Hayashi et al. 2012). Bone loss seems to be an inevitable event after stem insertion as part of the induced bone remodeling (Boe et al. 2011b). However, there is not much knowledge on the influence of the surgical approach to the hip joint and how this may affect bone remodeling around a femoral stem.

Dual-energy X-ray absorptiometry studies in patients receiving a total hip arthroplasty (THA) for osteoarthritis showed increased bone resorption in the direct lateral approach compared with the anterolateral approach (Perka et al. 2005, Merle et al. 2012). This may be due to compromised vascularization or possibly the alteration of the hip abductors and the musculoskeletal load to the proximal femur. Patients with femoral neck fracture are especially prone to periprosthetic fractures (Langslet et al. 2014, Skoldenberg et al. 2014) and bone remodeling and approach has not been studied in this patient group.

In this trial we hypothesized that the anterolateral (modified Watson-Jones) approach would give less bone loss around the femoral stem than the direct lateral (Hardinge) approach in patients with FNF.

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The first 56 patients included in the DXA sub-study and underwent DXA between February 2014 and March 2016 (Table 1). The endpoint was change in BMD as measured by DXA at 3 and 12 months. Patients between 70 and 90 years of age with adequate cognitive function and the ability to walk with or without a walking aid prior to fall were asked for their agreement to be enrolled and participate occurred after informed consent. Exclusion criteria were dementia, fractures in pathologic bone or patients not belonging to the hospital community. Those who displayed sepsis or local infection and not eligible to be treated with a hip prosthesis were excluded. 51 patients were enrolled in the DXA sub-study and underwent DXA between February 2014 and March 2016 (Table 1). The first 56 patients included in the main study (n = 120) were assessed for eligibility. 5 patients were not included.

The physician on duty evaluated whether the inclusion criteria were fulfilled and gave both verbal and written information about the trial. Randomization for surgery with either a direct lateral approach or an anterolateral approach was done by the same physician drawing a sealed envelope. Blinded study personnel recorded and monitored primary and secondary outcome measures.

We used the Corail stem (Depuy Orthopaedics Inc., Warsaw, IN, USA) intended for uncemented fixation, collared with standard offset and 135° neck angle. This is a titanium alloy straight stem with a grit-blasted surface and 155 µm of hydroxyapatite coating. The implant has a trapezoidal-like proximal cross section to provide rotational stability and self-locking, whereas the distal part is tapered. The SELF-CENTERING Bi-Polar Head was combined with an ARTICULEZE 28 mm femoral head, both from Depuy Synthes (West Chester, PA, USA) (Figure 1). Preoperative planning was performed using Sectra Medical Systems, Orthopaedic Package v5.5 (Sectra AB, Linköping, Sweden).

Patients were operated within 48 hours after sustaining their fracture. Operation was performed by 3 consultants in orthopedic hip surgery familiar with both approaches. The procedure was carried out under spinal anesthesia and all received the same standard analgesic protocol. Preoperatively 2 grams of cefalotin was given intravenously and a further 3 doses of 2 grams given over the next 24 hours. Low-dose heparin 40 mg (enoxaparin) was prescribed for 10 days.

The standard lateral decubitus position was selected for the direct lateral approach and the supine position for the anterolateral approach. For both procedures the femoral neck was resected and the femur reamed according to the preoperative planning or until rotational stability was achieved. The gluteal muscles were reinserted through osteosutures. There was no use of drainage and immediate full weight bearing was encouraged. Patients were examined with DXA within 3 days after surgery. Femoral BMD was measured on a GE Lunar Prodigy (GE Medical Systems, Madison, WI, USA).

BMD was measured postoperatively, at 3 and 12 months. The findings obtained at the postoperative scan were defined as baseline data.

The DXA measurement was performed by the technicians at the osteoporosis clinic, who were blinded to the allocated treatment.

Patients were positioned in the supine position with a triangle between the feet to obtain a standard rotation of the hip. Both hips were included. Readings started in the area about 2 centimeters proximal to the greater trochanter and distally to just below the femoral stem. The baseline scan was performed twice, and the patient moved between each scan. This was to estimate the precision expressed as coefficient of variation for the measurement procedure (Wilkinson et al. 2001). Changes in BMD related to the Gruen zones were then recorded and expressed as change in percentage using software from Orthopedic Hip for GE Lunar Prodigy (GE Healthcare, Chicago, IL, USA).

**Statistics**

Power calculations were based on previous studies on bone remodeling around the femoral stem (Boe et al. 2011b, Merle et al. 2012, Salemyr et al. 2015). We estimated a clinically

<table>
<thead>
<tr>
<th>Table 1. Baseline characteristics of included patients according to allocated surgical approach (figures are numbers unless stated otherwise)</th>
<th>Anterolateral (n = 26)</th>
<th>Lateral (n = 25)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mean (SD) age</td>
<td>82 (5.5)</td>
<td>84 (6.2)</td>
</tr>
<tr>
<td>Female / Male</td>
<td>17 / 9</td>
<td>16 / 9</td>
</tr>
<tr>
<td>ASA group I, II / III, IV</td>
<td>10 / 16</td>
<td>6 / 19</td>
</tr>
<tr>
<td>BMI (SD)</td>
<td>23 (4.0)</td>
<td>23 (3.2)</td>
</tr>
<tr>
<td>Dorr type a, B / C</td>
<td>22 / 4</td>
<td>20 / 5</td>
</tr>
<tr>
<td>Median stem size (range)</td>
<td>12 (9–18)</td>
<td>12 (10–16)</td>
</tr>
<tr>
<td>Mean (SD) HH5 b</td>
<td>84 (19)</td>
<td>84 (18)</td>
</tr>
</tbody>
</table>

*a* Dorr classification.

*b* Harris hip score estimated prior to fall.
important difference in BMD would be 10% (SD 10) between groups. To obtain a statistical power of 80% at the 0.05 level of significance 34 patients would be required, i.e. 17 in each treatment arm. We planned to include 50 patients to allow for loss to follow-up.

We used the double scans at baseline with repositioning between each scan to calculate the in-vivo precision error for the BMD procedures. Based upon the difference between these 2 scans the coefficient of variation (CV) was calculated for each ROI according to the formula: 
\[
CV\% = 100 \times \frac{\bar{\delta}}{\mu}
\]
where \( \overline{\delta} \) represents the standard deviation of the differences between the paired BMD measurements, and \( \mu \) is the overall mean of all the measurements for that ROI.

The change in BMD was calculated and the results expressed as percentage change with 95% confidence interval (CI) of postoperative values at 3 and 12 months for all regions of interest (ROI). The mean bone mineral density (g/cm²) postoperatively served as baseline. Bone density data were analyzed for normal distribution using histograms, Q-Q plots and the Shapiro–Wilk test. The groups were compared with Student’s t-test. A paired samples t-test was conducted to compare changes in BMD from baseline to follow-ups. The results were also reassessed with linear mixed models for repeated measurements, but we reported only results from t-tests since that statistical approach was pre-specified in the protocol. We did not perform adjustments for multiple tests.

A p-value of \( \leq 0.05 \) was considered statistically significant. SPSS Statistics 21 for Windows (IBM Corp, Armonk, NY, USA) was used for statistical analysis.

**Ethics, registration, funding, and potential conflicts of interest**

The trial was approved by the regional ethics committee (2013/1853/REK) and registered at ClinicalTrials.gov (ClinicalTrials.gov Identifier NCT02028468). The trial was reported based on the guidelines of the CONSORT Statement (Schulz et al. 2010) and designed in compliance with the Helsinki Declaration. All patients provided informed consent. The trial protocol was awarded an independent research grant of 25,000 Norwegian kroner from Smith and Nephew at the Norwegian Orthopedic meeting in 2013. Smith and Nephew did not perform adjustments for multiple tests. The classification of proximal femoral types according to Dorr et al. (1993) and the stem size inserted were similar in the 2 groups. The groups were comparable regarding stem alignment (Aldinger et al. 2009). Timed Up and Go test (TUG) performed at all 3 points of follow-up was similar between groups. A review of eligible medical records confirmed that the 2 groups were comparable regarding the prescription of bisphosphonates. At inclusion 4 patients were on osteoporosis medication. 7 patients died during the follow-up period and 5 patients did not attend the scheduled follow-up because of their health status. 1 patient with a periprosthetic fracture after a fall on the second postoperative day was excluded (Figure 2).

**BMD measurements**

The precision of the DXA measurements differed from 1.2% in Gruen zone 4 to 5.5% in Gruen zone 6 (Table 2). The 2 groups had similar BMD at the immediate postoperative measurement, both in the affected hip and in the contralateral hip. We found a continuous reduction in total periprosthetic BMD from baseline to 12 months. At 3 months there was a mean reduction in total periprosthetic BMD (4.2%, CI 2.4–6.1; \( p < 0.001 \)). Likewise there was a mean reduction in total periprosthetic bone at 12 months (5.8%, CI 3.3–8.3; \( p < 0.001 \)). At 3 months there was a mean reduction in total periprosthetic bone of 1.6% in the anterolateral group compared with

**Results**

**Patient characteristics**

51 patients were included in this sub-study. 26 were randomized to the anterolateral group and 25 to the direct lateral group. Mean age was 83 (70–90) years and 33 were female (Table 1). They were mainly ASA II (29%) and ASA III (58%) patients with a mean BMI of 23 (15–33). Time from admission to surgery and duration of surgery were similar in the 2 groups. The 2 groups had similar BMD at the immediate postoperative measurement, both in the affected hip and in the contralateral hip. We found a continuous reduction in total periprosthetic BMD from baseline to 12 months. At 3 months there was a mean reduction in total periprosthetic BMD (4.2%, CI 2.4–6.1; \( p < 0.001 \)). Likewise there was a mean reduction in total periprosthetic bone at 12 months (5.8%, CI 3.3–8.3; \( p < 0.001 \)). At 3 months there was a mean reduction in total periprosthetic bone of 1.6% in the anterolateral group compared with
In the direct lateral approach we found an early loss of BMD in all Gruen zones at 3 and 12 months. It was most pronounced between the baseline and the 3-month examination, with a loss of bone mineral in the remaining regions except zone 6, where BMD remained unchanged (Table 3). Statistical analysis confirmed a significant reduction of BMD in Gruen zones 1, 2, 6, and 7 (p < 0.05) in the lateral group compared with the anterolateral group: Gruen zone 1 (–5.0% vs. 2.7%), zone 2 (–4.3% vs. 4.1%), zone 6 (–6.5% vs. 0.0%), and zone 7 (–11% vs. –2.4%, all p < 0.05). There was a mean difference in Gruen zone 1 (7.7%, CI 0.0–15; p = 0.04), Gruen zone 2 (8.4%, CI 1.1–16; p = 0.02), Gruen zone 6 (6.5%, CI 0.2–13; p = 0.04), and Gruen zone 7 (8.8%, CI 0.1–18; p = 0.04), all in favor of the anterolateral group. The results were confirmed by a linear mixed model for repeated measurements analysis. At 12 months the tendency remained although now only significant in zone 6 (p < 0.05).

Discussion

We are not aware of any randomized trials examining the influence of the surgical approach on bone mineral changes in patients with femoral neck fractures. In this subgroup analysis of a randomized clinical trial reduction in BMD was higher in all Gruen zones in the direct lateral approach compared with the anterolateral approach after 3 months, which was statistically significant in the most proximal zones.

Changes in BMD around the femoral implant are a result of surgery-induced bone remodeling (Yamaguchi et al. 2000, Digas et al. 2009, Boe et al. 2011b, Tice et al. 2015). The complexity of this process is not fully understood as the etiology is thought to be multifactorial. Different implant designs, coatings (Flato et al. 2016) and stem sizes seem to influence periprosthetic bone remodeling (Karrholm et al. 2002, Nishino et al. 2013, Inaba et al. 2016). In our study, stem size was similar between groups and a single design only was used to minimize confounding effects. Activity may be a contributing factor (Hayashi et al. 2012) as well as loading of the proximal femur and local vascular status after the surgical trauma. The surgical technique itself is prone to influence BMD around the stem. Compaction of bone or excessive rasping prior to insertion of the stem is such a factor. An increase in bone density was measured by Kold et al. (2005)

Table 2. Precision of DXA measurements. Coefficients of variation (CV%)

<table>
<thead>
<tr>
<th>Gruen zone:</th>
<th>1</th>
<th>2</th>
<th>3</th>
<th>4</th>
<th>5</th>
<th>6</th>
<th>7</th>
</tr>
</thead>
<tbody>
<tr>
<td>CV%</td>
<td>3.2</td>
<td>2.4</td>
<td>2.1</td>
<td>1.2</td>
<td>3.8</td>
<td>5.5</td>
<td>4.7</td>
</tr>
</tbody>
</table>

Table 3. Periprosthetic changes in bone mineral density (BMD) around the hydroxyapatite-coated Corail stem measured by dual-energy X-ray absorptiometry (DXA)

<table>
<thead>
<tr>
<th>Gruen zone</th>
<th>Surgical approach</th>
<th>n</th>
<th>Postoperative mean BMD (g/cm²)</th>
<th>3-months change</th>
<th>12-months change</th>
</tr>
</thead>
<tbody>
<tr>
<td>1–7</td>
<td>AL</td>
<td>51</td>
<td>1.61 (1.55–1.68)</td>
<td>43 4.2 (2.4–6.1)</td>
<td>38 5.8 (3.3–8.3)</td>
</tr>
<tr>
<td>1</td>
<td>AL</td>
<td>26</td>
<td>0.81 (0.74–0.88)</td>
<td>21 2.7 (–2.9–8.4)</td>
<td>18 −3.8 (–13.6)</td>
</tr>
<tr>
<td></td>
<td>L</td>
<td>25</td>
<td>0.82 (0.76–0.89)</td>
<td>22 −5.0 (–10.0–0.5)</td>
<td>20 −7.4 (–15.0–3)</td>
</tr>
<tr>
<td>2</td>
<td>AL</td>
<td>26</td>
<td>1.79 (1.68–1.90)</td>
<td>21 4.1 (–2.8–11)</td>
<td>18 1.6 (–7.5–11)</td>
</tr>
<tr>
<td></td>
<td>L</td>
<td>25</td>
<td>1.91 (1.80–2.03)</td>
<td>22 −4.3 (–7.6–0.9)</td>
<td>20 −6.6 (–9.9–3.4)</td>
</tr>
<tr>
<td>3</td>
<td>AL</td>
<td>26</td>
<td>2.08 (1.95–2.21)</td>
<td>21 −1.5 (–6.2–3.2)</td>
<td>18 −1.3 (–7.6–4.9)</td>
</tr>
<tr>
<td></td>
<td>L</td>
<td>25</td>
<td>2.18 (2.05–2.31)</td>
<td>22 −4.9 (–8.0–0.1)</td>
<td>20 −6.1 (–8.5–3.8)</td>
</tr>
<tr>
<td>4</td>
<td>AL</td>
<td>26</td>
<td>1.84 (1.71–1.98)</td>
<td>21 −3.6 (–6.1–1.2)</td>
<td>18 −5.1 (–7.3–2.9)</td>
</tr>
<tr>
<td></td>
<td>L</td>
<td>25</td>
<td>1.87 (1.73–2.00)</td>
<td>22 −5.4 (–8.9–3.9)</td>
<td>20 −7.0 (–8.8–5.3)</td>
</tr>
<tr>
<td>5</td>
<td>AL</td>
<td>26</td>
<td>2.18 (2.04–2.31)</td>
<td>21 −5.7 (–8.0–3.4)</td>
<td>18 −5.9 (–9.5–2.3)</td>
</tr>
<tr>
<td></td>
<td>L</td>
<td>25</td>
<td>2.19 (2.05–2.34)</td>
<td>22 −8.0 (–11.0–4.6)</td>
<td>20 −6.1 (–9.1–3.2)</td>
</tr>
<tr>
<td>6</td>
<td>AL</td>
<td>26</td>
<td>1.50 (1.43–1.58)</td>
<td>21 0.0 (–4.4–4.6)</td>
<td>18 −1.9 (–8.9–5.1)</td>
</tr>
<tr>
<td></td>
<td>L</td>
<td>25</td>
<td>1.50 (1.41–1.60)</td>
<td>22 −6.5 (–11–2.0)</td>
<td>20 −11 (–16–5.6)</td>
</tr>
<tr>
<td>7</td>
<td>AL</td>
<td>26</td>
<td>0.98 (0.88–1.08)</td>
<td>20 −2.4 (–7.6–2.8)</td>
<td>18 −7.8 (–17.0–9)</td>
</tr>
<tr>
<td></td>
<td>L</td>
<td>25</td>
<td>0.97 (0.85–1.10)</td>
<td>22 −11 (–18–4.1)</td>
<td>20 −12 (–29–5.4)</td>
</tr>
</tbody>
</table>

The mean bone mineral density (g/cm²) postoperatively serves as baseline. Mean percentage change in BMD at 3 and 12 months (95% CI) for each Gruen zone and for the total periprosthetic region.

a AL = anterolateral; L = lateral.

b p < 0.05.

c One excluded due to the formation of heterotopic ossification.
blood supply may have influenced bone loss in the present study. Decreased blood flow rate to the greater trochanter in adult rabbits after vascularization of the greater trochanter and different loads on the proximal femur with functional load-bearing favoring bone remodeling (Rubin and Lanyon 1984). Thus, it seems reasonable that the surgical approach influences the process of periprosthetic bone adaptation, possibly contributing to the reported increased fracture risk around uncemented stems. Numbers from the Nordic Arthroplasty Register Association (NARA) show that nearly all of the periprosthetic fractures with uncemented stems occurred within the first 6 months after surgery (Gjertsen et al. 2012, Langslet et al. 2014, Thien et al. 2014, Inngul et al. 2015).

As literature on this issue is sparse we can only compare our results with studies on THR in osteoarthritis patients. Statistical power was based on sample size calculation from these studies and may not reflect the true nature of the problem. The limitations include some loss to follow-up and possibly the short period of observation.

Furthermore we allowed immediate full weight bearing in both groups including the TUG test but did not quantify the amount of mobilization possibly affecting proximal bone resorption. The analyses of the subgroup had a confirmatory statistical strategy with a single variable and a pre-specified hypothesis. We report on confidence intervals to emphasize clinical significance. No interim analysis was performed.

In summary, at 3 months we found a statistically significant reduction of bone mineral in the proximal Gruen zones in the direct lateral approach compared with the anterolateral approach. We did not perform adjustments for multiple tests, possibly leading to inflation of false-positive findings. Therefore, the results should be interpreted with caution.

The authors would like to express their gratitude and thanks to Isabel Priscilla Nunez for acquisition of data, database management, and organizing the follow-ups at the outpatient clinic. They thank the staff at the Osteoporosis Clinic, Soelandet Hospital, for performing the DXA examinations. The authors would like to thank physiotherapists Linda Hansen and Arild Ege.

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