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Using four different clinical tools as predictors for pain after total hip arthroplasty: a prospective cohort study

Anja Geisler1,2*, Josephine Zachodnik1, Jens Laigaard1, Laura S. Kruuse3, Charlotte V. Sørensen3, Magnus Sandberg2, Eva I. Persson2 and Ole Mathiesen1,4

Abstract

Background: Treatment of postoperative pain remains a significant clinical problem, and prediction of patients with a risk of higher postoperative pain levels is an important focus. We aimed to identify patients undergoing total hip arthroplasty (THA) with risk of higher pain levels at 24 h postoperatively by using four simple and easily available clinical tools.

Methods: This prospective observational cohort study included 102 patients having THA at Zealand University Hospital in Denmark. The following predictive tools were investigated for identifying patients with higher postoperative pain levels at 24 h postoperatively, both at rest and during mobilization: preoperative pain by peripheral venous cannulation (PVC) (dichotomized according to numerical rating scale pain ≤ 2/> 2 (PVC-Low/PVC-High) (primary outcome); the post anesthesia care unit (PACU) nurses’ expectations of patients pain levels; patients early pain levels at the PACU; and patients own forecast of postoperative pain levels. The Mann-Whitney U test was used to analyze comparisons between prediction groups. For the primary outcome we considered a \( p \)-value < 0.01 as statistically significant and for other outcomes a \( p \)-value of 0.05.

Results: We found no significant differences between the PVC groups for pain during mobilization at 24-h postoperatively: PVC-Low: 6 (4–8) (median, (IQR)) versus PVC-High: 7 (5–8) (median, (IQR)), \( p = 0.10 \); and for pain at rest: PVC-Low 2 (0–3) (median, (IQR)) versus PVC-High 3 (2–5) (median, (IQR)), \( p = 0.12 \). Other comparisons performed between predictive groups did not differ significantly.

Conclusions: In this prospective cohort study of 102 THA patients, we did not find that preoperative pain by PVC, when using a cut-off point of NRS ≤ 2, were able to predict postoperative pain at 24 h postoperatively. Neither did PACU nurses’ prediction of pain, patients forecast of pain, nor did maximum pain levels at the PACU.

Trial registration: Retrospectively registered 20th February 2018 at ClinicalTrials.gov (NCT03439566).

Keywords: Postoperative pain, Prediction, Total hip arthroplasty

Background

Despite considerable efforts in optimizing postoperative pain, this clinically important symptom remains a major challenge [1]. It is therefore important to identify individuals at risk of developing high postoperative pain levels, but clinically useful predictive tools are virtually absent [2]. A newer study indicated that pain intensity by preoperative peripheral venous cannulation (PVC), using a grouping according to numerical rating scale pain (NRS) ≤ 2 and > 2, was associated with pain levels at the post anesthesia care unit (PACU) [3]. This study, however, did not investigate the prediction of pain later at the surgical ward, which is particularly relevant, since sufficient pain treatment is a cornerstone for optimal rehabilitation [4].
Nurse’s prediction of patient outcomes has been investigated in different settings but with varying results [5, 6]. Therefore, it is relevant to investigate if experienced nurses at the PACU can predict which patients will suffer from higher levels of pain after PACU discharge.

As patients may be predisposed to certain levels of postoperative pain due to e.g. sex, preoperative pain, genetic variations [7], anxiety, or type of surgery [8, 9], it could be relevant to investigate if patient’s pain levels at the PACU, using moderate to severe pain (NRS > 3) as an indicator, can predict pain levels after PACU discharge.

A recent study reported that 47% of patients correctly predicted their pain levels 2 weeks after hand surgery, but they did not investigate the prediction of acute postoperative pain [10]. A subsequent commentary pointed out, that further prospective studies are needed regarding patient’s ability to forecast their disability and pain [11].

Total hip arthroplasty (THA) is a surgical procedure of which patients’ have reported moderate to severe postoperative pain [12]. Therefore, this population is relevant for investigating which patients are high pain responders including a particular focus on pain during recovery at the surgical ward. With this study, we hypothesized that different clinical parameters, nursing staff impressions and patients forecast could be used to predict postoperative pain. This study aimed to investigate if preoperative pain by PVC could be used to identify groups of THA patients with higher levels of pain during mobilization at 24 h postoperatively (primary outcome). Additionally, that PACU nurses’ capability of predicting patients with higher pain levels at the ward, pain levels at the PACU, and patients’ forecast preoperatively, could be used to identify patients with higher postoperative pain levels at 24 h after THA (secondary outcomes).

Outcomes
Primary outcome was: numeric rating scale (NRS) pain (0–10) during mobilization at 24 h postoperatively.
Secondary outcomes were: NRS-pain at rest at 24 h postoperatively, and 24 h intravenous morphine equivalent consumption, mg.

A correctly placed PVC was defined as a cannula placed in a vein on the dorsum part of the dominant hand. The NRS score was performed during the first attempt otherwise the patients were excluded. The allowed cannula sizes were 20G or 22G.

Anesthesia and analgesic treatment
All patients received spinal anesthesia (10–15 mg bupivacaine). The standard analgesic treatment that was provided for patients at the hospital was: perioperative methylprednisolone IV 125 mg (after induction of anesthesia). At the POTAs patients were supplied with opioids as needed, according to usual practice. At the ward, postoperatively, paracetamol 1000 mg OR every 6 h, and slow-release oxycodone 10 mg OR administrated twice a day, supplemented with oxycodone IV as needed.

Psychological profile
Patients’ psychological profile and relation to pain were tested using the validated self-administered questionnaire, Pain Catastrophizing Scale (PCS). The scale includes 13 items and assesses the extent of the patient’s catastrophizing thoughts and emotions associated with pain. Such thoughts or feelings are rated from zero (not at all) to five (all the time). The PCS has a maximum score of 52. A clinically relevant cut off for being a pain catastrophizer was considered as numbers above 30 [14].
**Supplemental data regarding PACU nurses’ prediction**

When PACU nurses were asked to state if they predicted a patient to be a high pain responder or not, they were also asked to tick a box with the following statements underpinning their choice: patients’ appearance, patients’ pain intensity, my own intuition, patients’ need of opioids, the patient’s expression of concern and anxiety, or optional additional information – described in their own words.

**Collection of data**

For evaluation of pain, the NRS-scale was used, 0 to 10 (0 = no pain and 10 = worst imaginable pain). All patients were instructed preoperatively in how to use the NRS tool [15]. The patients stated their own pain. All data were entered directly into the patient Case Report Form (CRF). All opioids were converted to IV morphine equivalents (eqv) (Additional file 1).

At inclusion, patients completed the PCS questionnaire and provided information about years of education after high school, civil status, employment, as well as their forecast of pain levels. The anesthesia nurse at the operation theater, who performed the peripheral cannulation, asked patients’ about the levels of NRS pain after placement of the PVC. The nurses on the PACU collected data on NRS pain after the spinal anesthesia had ceased, as well as performed a prediction of which patients they believed would experience moderate or severe pain at 24 h during mobilization. The primary investigator, a project nurse or research assistants performed the data collection at the ward at 24 h +/− 2 h postoperatively for pain and opioid consumption.

The following information was registered from the electronic patient records. Preoperative data: height, weight, sex, American Society of Anesthesiologists physical score (ASA), usual preoperative analgesic consumption and premedication. Perioperative data: analgesic and antiemetic treatment, duration of surgery. Postoperative data: analgesics used from 0 h to 24 h postoperatively, and length of stay (LOS). All data were registered in the CRF, imported to the statistical package IBM SPSS version 25, and the final data set was double-checked for errors.

The patients also filled out a diary from postoperative day one to five at home regarding pain, side effects, use of analgesics and quality of sleep. These data will be reported elsewhere.

**Sample size and statistics**

A sample size estimation was performed for NRS pain during mobilization based on data from a prior study that investigated a similar patient population [12]. To reduce the risk of spurious significant findings, we choose an alpha = 0.01 and a power of 0.9. Furthermore, we used a standard deviation (SD) of 2.5. We found 93 patients were needed to detect a minimal clinically important difference (MCID) of NRS-pain at 1.0. To compensate for the uncertainty of SD we aimed to include 100 consecutive patients undergoing THA.

For comparisons we defined groups based on the following:

1) Pain by preoperative PVC dichotomized to NRS ≤ 2 (PVC-Low) or NRS > 2 (PVC-High)
2) PACU nurses’ prediction of patients being a high pain responder or not (Nurse-Low, or Nurse-High)
3) Maximum NRS pain at the PACU dichotomized to NRS ≤ 3 / > 3 (when spinal anesthesia has ceased, Bromage score 0—1) (PACU-NRS ≤ 3, or PACU-NRS > 3)
4) Patients reporting of being a high pain responder or not (Forecast-Low, or Forecast-High)

Normal distribution was tested visually in histograms and Q-Q plots and quantitatively with Kolmogorov-Smirnov test. Data are presented with either numbers or percentages, median (IQR), mean (SD), and 95% CI, where relevant. Mann-Whitney U test (for non-parametric data) was used to analyze all comparisons between groups: PVC-High versus PVC-Low, Nurse-Low versus Nurse-High, Forecast-Low versus Forecast-High, and PACU-NRS ≤3 versus PACU-NRS > 3. For the primary outcome we considered a p-value < 0.01, and for other outcomes, a p-value < 0.05, as statistically significant. Bonferroni correction was used to counteract for mass-significance where relevant.

We performed an exploratory multiple linear regression, both adjusted an unadjusted analyses, using a dependent variable, NRS pain at 24 h during rest or mobilization, and adjusting for the following pre-defined covariates: sex, age, patients pain threshold, marital status, education, daily analgesic consumption, PCS, and employment status. To test for the possibility of multicollinearity, Pearson r for parametric data and Spearman rho for non-parametric data was used.

For evaluating and comparing predictive models we calculated Receiver Operating Characteristic curves (ROC). The true positive rate in the model (sensitivity) was plotted against the false positive rate (1 – specificity) for a given cut-off value of the predictive variable, thus aiming to determine the optimal cut-off value. Areas in the interval 0.9–1 represented excellent prediction, 0.8–0.9 good prediction, 0.7–0.8 fair prediction and 0.6–0.7 poor prediction [16, 17]. Statistical analyzes were expressed using IBM SPSS software version 25 for Windows (SPSS Inc. Chicago, IL).

**Results**

One hundred and fifty patients scheduled for THA were assessed for eligibility. After exclusions, 102 patients were included in the study for evaluation of the primary outcome. For further information see Fig. 1.
The study included 35 males and 67 females, mean age was 69 (19) (mean, (SD)) years, the surgery lasted in 53 (46–63) (median, (IQR)) min, and median LOS was 1 (1, 2) day. For further information, see Table 1.

**PVC related pain**

For the primary outcome, NRS pain during mobilization at 24 h, we found no significant difference between groups PVC-Low 6 (4–8) (median (IQR)) and PVC-High 7 (5–8) (median (IQR)) \((P = 0.10)\) (Table 2). For NRS pain at 24 h at rest, we found no significant difference between groups PVC-Low 2 (0–3) (median (IQR)) and PVC-High 3 (2–5) (median (IQR), \(p = 0.12\). For total 24 h IV morphine eqv. Consumption, we found no significant difference between groups PVC-Low 20 (15–24) (median (IQR)) mg and Group PVC-High 23 (15–28) (median (IQR) mg (median (IQR), \(p = 0.20\) (Table 2).

Explorative regression analyses were performed regarding the association between PVC-Low and PVC-High, and postoperative 24 h NRS pain during rest and mobilization, in both unadjusted and adjusted analyses. For 24 h NRS pain during mobilization, the unadjusted analyses demonstrated no significant difference in standardized Beta 0.88 (−0.18; 1.94) NRS (95% CI) \((p = 0.10)\) between groups defined by PVC pain. When adjusted for sex, age, patient’s forecast of own pain, marital status, education, daily analgesic consumption, PCS, and employment status, we found a significant difference in standardized Beta NRS (95% CI) 0.24 (0.14; 2.43) \((p = 0.03)\) between the groups defined by pain during PVC (Table 3). For NRS pain at rest at 24 h the unadjusted Beta showed a significant difference in NRS (95% CI), 1.13 (0.14; 2.12) \((p = 0.03)\) between groups defined by PVC pain. In the adjusted analyses, however, this difference became non-significant with a standardized Beta NRS (95% CI) 0.18 (−0.22; 2.06) \((p = 0.11)\) (Table 3). We did not find any multicollinearity of parameters in the adjusted analyses.

**PACU nurse prediction**

We found no significant differences between groups Nurse-Low and Nurse-High for NRS pain during mobilization at 24 h postoperatively Nurse-Low: 5 (4–8)
and Nurse-high: 6 (4–7) (median (IQR)), p = 0.78. No significant differences between groups were found for pain at rest at 24 h postoperatively, and for 24 h morphine consumption. See Table 2 for further details.

**Pain at the PACU**

For groups based on NRS pain ≤ 3 and NRS pain > 3 at the PACU, we found no significant differences between groups, for pain during mobilization at 24 h postoperatively, PACU-NRS ≤ 3: 5 (4–8) and PACU-NRS > 3: 7 (6–8) (median (IQR)), p = 0.74. No significant differences between groups were found for pain at rest. For 24 h morphine consumption, we found a significant difference between group PACU-NRS ≤ 3: 20 (15–25) mg and PACU-NRS > 3: 26 (18–33) mg (median (IQR)), p = 0.03, Bonferroni adj. See Table 2 for details.

**Patients forecast of pain**

We found no significant differences between groups Forecast-Low and Forecast-High regarding NRS pain during mobilization at 24 h postoperatively; Forecast-Low: 6 (4–8) and Forecast-High 6 (4–8) (median (IQR)), p = 0.79. No significant differences between groups for NRS pain 24 h during rest and opioid consumption was detected. For further information see Table 2.

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**Table 1** Demographics and baseline data

| Demographic/Baseline variable                        | Total population (n = 102) | Missing (n) | PVC-Low (n = 67) | Missing (n) | PVC-High (n = 35) | Missing (n) | PVC-Low vs High p-value |
|------------------------------------------------------|-----------------------------|-------------|------------------|-------------|-------------------|-------------|-------------------------|
| Sex m/f, (n)                                         | 35/67                       | 0           | 26/41            | 0           | 9/26              | 0           | 0.18                    |
| Age, yr, mean (SD)                                   | 69 (19)                     | 0           | 71 (8)           | 0           | 66 (10)           | 0           | 0.02                    |
| Height, cm, mean (SD)                                | 169 (8)                     | 15          | 168 (8)          | 11          | 169 (8)           | 4           | 0.58                    |
| Weight, kg, median (IQR)                             | 75 (65–85)                  | 15          | 75 (64–83)       | 11          | 75 (66–98)        | 4           | 0.41                    |
| ASA 1/2/3 (n)                                        | 21/62/16                    | 3           | 15/40/10         | 2           | 6/22/6            | 1           | 0.57                    |
| Education after high school                          | 24/71                       | 7           | 16/46            | 5           | 8/25              | 2           | 0.98                    |

**Table 2** All patients and comparisons between predictive groups

| Variable                                              | All patients (n = 102) | PVC-Low (n = 67) | PVC-High (n = 35) | p-value | PVC-Low (n = 49) | PVC-High (n = 32) | p-value | PVC-Low vs High p-value | PVC-NRS ≤ 3 (n = 90) | p-value | PVC-NRS > 3 (n = 12) | p-value | Forecast-Low (n = 79) | Forecast-High (n = 21) | p-value |
|-------------------------------------------------------|-------------------------|------------------|------------------|---------|-----------------|------------------|---------|-------------------------|----------------------|---------|-----------------------|---------|----------------------|-----------------------|---------|
| Pain (mobilization) 24 h postop.                      | 6 (4–8)                 | 6 (4–8)          | 7 (5–8)          | 0.10    | 5 (4–8)         | 6 (4–7)          | 0.78    | 5 (4–8)                 | 7 (6–8)              | 0.74    | 6 (4–8)               | 6 (4–8) | 0.79                 |                       |         |
| Pain (at rest) 24 h postop.                           | 2 (0–4)                 | 2 (0–3)          | 3 (2–5)          | 0.12    | 2 (0–4)         | 2 (0–4)          | 0.65    | 2 (0–4)                 | 3 (2–5)              | 0.22    | 2 (1–4)               | 2 (0–3) | 0.19                 |                       |         |
| Morphine consumption (eqv.), IV, mg, (0–24 h)         | 20 (15–25)              | 20 (15–24)       | 23 (15–28)       | 0.20    | 19 (15–23)      | 22 (15–29)       | 0.16    | 20 (15–25)              | 26 (18–33)           | 0.12    | 20 (15–28)            | 20 (15–23) | 0.35                |                       |         |

*Bonferroni correction. PVC Peripheral Venous Cannulation. PACU Post Anesthesia Care Unit. NRS Numerical Rating Scale. Data are median and interquartile range (IQR), pain are numerical rank scale (NRS). Nurse-Low means patients that the PACU nurse evaluates to be an ordinary pain responder and Nurse-High was evaluated to be a high pain responder. Forecast-Low means ordinary pain responder and Forecast-High means high pain responder, according to evaluation by patients themselves before surgery.
ROC curve analyses
We performed ROC curves for groups PVC-Low and PVC-High, and its capability of predicting pain during rest and mobilization 24 h, as well as 24 h opioid consumption. Since all ROC had AUC values less than 0.60, we did not consider these to be reliable predictors (Additional file 2).

Discussion
With this prospective observational cohort study of 102 patients undergoing THA, we investigated four simple and easily available clinical tools to predict patients with higher levels of postoperative pain 24 h after surgery. We did, however, not find any significant difference for postoperative pain during mobilization, at rest, between groups of patients defined by pain by PVC, PACU nurses’ prediction, pain levels at the PACU, and patients forecast of pain.

Pain from preoperative peripheral venous cannulation was previously investigated (3) for prediction of postoperative pain in a study with 180 patients undergoing laparoscopic cholecystectomy. Here, Persson and colleagues (3) reported that patients with NRS pain > 2 by PVC reported higher pain scores at rest and received more opioids within the first 90 min at the PACU, compared to those with NRS ≤ 2. Furthermore, in a newer and larger study, Persson and co-workers [18] repeated these findings, but this time in a population of 600 patients undergoing different surgical procedures, receiving different types of anesthesia, and using different places for venous cannulation. Still, they reported that NRS pain > 2 during PVC was associated with moderate to severe postoperative pain at rest at the PACU. We could not confirm these findings, as we found no significant differences for pain at rest at the PACU between PVC-Low and PVC-High. Our study differed from those of Persson and colleagues’ in several ways, including different patient populations, differences in age, and the type of anesthesia. Also, we measured pain both at rest and during mobilization, and not only at the PACU but also 24 h postoperatively. We did not find any association between PACU nurses’ prediction and patients’ pain levels after 24 h, neither at rest nor during mobilization. Interestingly, in the follow-up on background for their choice, we found that the PACU nurses stated that the forecasts of patients being low or a high pain responder, to be primarily based upon the appearance of the patient and on their intuition. These findings were supported by other studies demonstrating that when nurses assess if patients are in pain or not, they primarily base their decision on patients’ appearances and their non-verbal behavior [19, 20]. However, this failed to show applicability in this study.

In a recent study in 563 women having breast cancer surgery, Sipilä and colleagues [21] found that patient’s expectations of severe postoperative pain were associated with higher clinical pain intensity and increased initial oxycodone use at the PACU. In contrast, in another and smaller prospective study [10] investigating patients’ ability to forecast their disability and pain two weeks after hand surgery, only weak correlations between forecasted and realized pain was discovered [10]. In contrast, we did not find any differences in patients pain scores nor opioid usage for the first 24 h postoperatively, based on patients’ forecast on being a high pain responder or not. It is possible that differences in type of surgeries and patient populations between these and the present study influences the differences in outcomes, and further studies are needed to elaborate on this clinically relevant topic.

Strengths and limitations
In this prospective study it was a strength that all patients had spinal anesthesia and underwent the same

| Table 3 Multiple linear regression model for NRS pain by PVC. Adjusted and unadjusted |
|---------------------------------|-----------------|-----------------|-----------------|-----------------|-----------------|
| Dependent variable              | Independent variable | Beta Estimate NRS (95% CI) | P-value | R² |
| Pain at 24 h (mobilization)     | NRS by PVC ≤/>2    | 0.88 (-0.18;1.94) | 0.10 | 0.27 |
| Unadjusted                      |                  |                  |                |                |
| Pain at 24 h (mobilization)     | NRS by PVC ≤/>2    | 0.24* (0.14;2.43) | 0.03 | 0.20 |
| Adjusted                        |                  |                  |                |                |
| Pain at 24 h (at rest)          | NRS by PVC ≤/>2    | 1.13 (0.14;2.12) | 0.03 | 0.05 |
| Unadjusted                      |                  |                  |                |                |
| Pain at 24 h (at rest)          | NRS by PVC ≤/>2    | 0.18* (-0.22;2.06) | 0.11 | 0.13 |
| Adjusted                        |                  |                  |                |                |

Adjusted for; sex, age, patients pain threshold, marital status, education, daily analgesic consumption, PCS and employment
NRS Numerical Rating Scale. PVC Peripheral Venous Cannulation. CI Confidence Interval *Standardized Beta value
surgical procedure, which was performed by the same two orthopaedic surgeons, and with all patients receiving a standardized perioperative pain treatment. This minimizes variations and bias. Furthermore, PVC was performed by experienced anesthesia nurses, and all were placed in the patients’ dominant hand. Also, data at the ward were collected by a limited group of four investigators, reducing observer bias.

We used a cut-off point of NRS 2, for the division of groups, based on the study by Persson et al. [3]. The ROC curve analyses indicated that a cut-off point of NRS 2.5 could have been more appropriate for dividing patients into groups (Additional file 2). However, the AUC-value of the ROC was low and of limited reliability. Missing data was a limitation, especially the relatively large proportion of missing data on PACU nurses’ prediction could have influenced our results. Preoperative pain may serve as a predictor for postoperative pain levels [8]. It could therefore be considered as a limitation, that we did not register patients’ preoperative pain, but instead, as a proxy here for, registered preoperative analgesic consumption. Another limitation is the sample size calculation, as such calculations typically are based on an equal number of patients in the investigated groups. This was unfortunately not the case, as number of patients differed between the four investigated groups of predictors. As this was a clinical prospective cohort, and not a randomized trial, it can be argued that sample sizes are of limited value, especially when we did not have influence on the distribution of number of patients in the compared groups. Our study can, however, serve as base for sample size calculations of future studies, preferably including larger numbers of participants.

In perspective, the prediction of postoperative pain levels continues to be an important focus for future research, as individualized pain treatment has the potential to improve patient courses. Such research may well be based on the collection of big data including new types of analyses hereof, biomarkers, neuroimaging, physiological and psychological variables, and clinical data as well. It is possible, that results of studies using simple clinical tools, in the future, might be included in such big data in the guidance of pain treatment. Until then, the focus must be on effective pain treatment, regular assessment of pain with frequently monitoring of patients and their needs.

Conclusions
In this prospective cohort study of 102 THA patients, we did not find that preoperative pain by PVC, when using a cut-off point of NRS ≤ 2, were able to predict postoperative pain at 24 h after THA postoperatively. Neither did PACU nurses’ prediction of pain, patients forecast of pain, nor did maximum pain levels at the PACU.

Supplementary information
Supplementary information accompanies this paper at https://doi.org/10.1186/s12871-020-00959-2.

Additional file 1. Supplemental Digital Content 1: Opioid conversion.
Additional file 2. Supplementary information

Abbreviations
ASA: American Society of Anesthesiologists physical score; CRF: Case Report Form; Eqv: Equivalents; IQR: Inter Quartile Range; IV: Intravenous; LOS: Length of stay; NRS: Numeric Rating Scale; PACU: Post Anesthesia Care Unit; PVC: Peripheral Venous Cannulation; RCT: Randomized controlled trial; THA: Total Hip Arthroplasty; VAS: Visual Analogue Scale; VAS: Verbal Rating Scale; ZUHK: Zealand University Hospital, Køge

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Authors’ contributions
The study was designed by: AG, EP, OM. Study conduct: AG, JL, LK, CVS. Data analysis: AG, MS, OM. Critical revision and final approval of the manuscript: All authors.

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Availability of data and materials
The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

Ethics approval and consent to participate
Approved by the Danish Data Protection Agency (REG-158-2017) and The Danish Research Ethics Committee in the Region of Zealand (Reg. nr. J.nr. 17–000048). All participants were enrolled after giving verbal and written informed consent.

Consent for publication
Not applicable.

Competing interests
The authors declare that they have no competing interests.

Author details
1Department of Anesthesiology, Zealand University Hospital, Lykkebækvæj 1, 4600 Køge, Denmark. 2Department of Orthopaedics, Zealand University Hospital Hospital, Køge, Denmark. 3Department of Clinical Medicine, Faculty of Health Sciences, Copenhagen University, Copenhagen, Denmark. 4Department of Orthopaedics, Zealand University Hospital, Lykkebækvæj 1, 4600 Køge, Denmark.

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