

Predictors of total hip replacement in community based older adults: a cohort study

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Abstract

Background

Hip osteoarthritis (OA) commonly affects older adults and leads to high morbidity. There is no preventative treatment available and total hip replacement (THR) is offered for end stage disease. Known predictors of THR include pain and radiographic OA. Hip structure has also been shown to worsen hip OA and predict THR. A better understanding of predictors of THR can aid in triaging patients and researching preventative strategies. The purpose of this study is to describe predictors of THR in community dwelling older adults.

Methods

At baseline, participants had assessment of radiographic OA and cam impingement (from pelvic radiographs), shape mode scores (from dual energy X-ray absorptiometry (DXA)) and hip bone mineral density (BMD) (from DXA). After 2.6 and 5 years, participants reported hip pain using WOMAC (Western Ontario and McMaster Universities Osteoarthritis Index), and had hip structural changes assessed using magnetic resonance imaging (MRI). Risk of THR was analysed using mixed-effect Poisson regression.

Results

Incidence of THR for OA over 14 years was 5.0% (40 / 802). As expected, WOMAC hip pain and hip radiographic OA both predicted risk of THR. Additionally, shape mode 2 score (decreasing acetabular coverage) (RR 1.57 per SD; 95% CI 1.01-2.46), shape mode 4 score (non-spherical femoral head) (RR 0.65/SD; 95% CI 0.44-0.97), cam impingement ($\alpha > 60^\circ$) (RR 2.66/SD; 95% CI 1.38-5.13), neck of femur BMD (RR 1.85/SD, 95% CI 1.4-2.44) and bone marrow lesions (BMLs) increased risk of THR (RR 5.62/unit; 95% CI 1.1 – 28.81). There was a trend for hip effusions to increase the risk of THR (RR 1.88/SD; 95% CI 0.24 to 14.78).

Conclusion

In addition to hip pain and radiographic hip OA, measures of hip shape, cam impingement, BMD and BMLs independently predict risk of THR. This supports the role of hip bone geometry and structure in the pathogenesis of end stage hip OA and has identified factors that can be used to improve prediction models for THR.

Introduction

Hip osteoarthritis (OA) is a common musculoskeletal condition that is a major contributor to disability globally.(1) There are currently no treatments available that prevent hip OA, or slow the disease trajectory. Once disease is advanced, total joint replacement surgery is offered. Whilst these surgeries are successful and have high levels of patient satisfaction(2) they are expensive and have a finite life. Better

understanding of predictors of hip replacement provides some scope for prevention of hip replacement and may aid treatment decisions.

There is ongoing debate as to whether associations exist between radiographic and clinically defined hip OA. The inconsistent literature might be due to different definitions of hip OA, different radiographic protocols and scoring methods.(3) However, both predict risk of total hip replacement.(4) Recently, hip morphology has been identified as having an important role in the progression of hip OA.(5-8) Particular patterns of hip shape such as reduced acetabular coverage(8), non-spherical femoral heads(8) and cam impingement (abnormally shaped head of femur leading to abnormal contact between femoral head and acetabulum)(9) predict progression of hip OA and risk of THR.(8-11) Hip bone marrow lesions (BMLs), hip cartilage defects(12-16) and higher BMD of the proximal femur(17) are independent risk factors for progression of hip OA. Greater BMD also increases risk of THR(18, 19); hip BMLs and cartilage defects may do likewise but these associations have not been studied. No studies have reported on all these risk factors in the same population or community based populations and few have adjusted for pain and/or radiographic osteoarthritis. When they have the result for hip shape became negative making it uncertain if these risk factors are independent.(6) Thus, the aim of this study was to examine the effect of hip structural factors as risk factors for THR independent of hip pain and radiographic measures of hip OA in community dwelling older adults.

Patients And Methods

Study design and setting

The Tasmanian Older Adult Cohort (TASOAC) study is a prospective, population-based cohort study, which aimed to identify factors associated with development and progression of OA and osteoporosis in older adults. Men and women aged 50–80 years in 2002 were selected from the electoral roll, which is the most complete population listing for adult Australians, in Southern Tasmania (population 229,000) using sex-stratified random sampling (response rate 57%). Participants were excluded if they lived in an aged care facility, or had contraindications to magnetic resonance imaging. The Southern Tasmanian Health and Medical Human Research Ethics Committee approved the study, and we obtained written informed consent from all participants.

Baseline data (Phase 1) were collected from February 2002 to September 2004 in 1099 participants. Follow up data (Phases 2, 3 and 4) were collected on average 2.6 (n = 875), and 5 years (n = 769) later. Participants who had a hip replacement prior to Phase 1 were excluded from analyses in this manuscript (n = 13).

Outcome: Total Hip Replacement

Incidence of primary THR was determined by data linkage to the Australian Orthopaedic Association National Joint Replacement Registry (AOANJRR), and includes data from both public and private

hospitals. Data validation against State and Territory Health Department data is done using a sequential multi-level matching process.(20) Matched data were then obtained; this included the date, side of joint replacement, primary or revision joint replacement and the reason for the procedure (e.g., OA, fracture of neck of femur, osteonecrosis, inflammatory arthritis, tumour). In this study, we only considered primary THRs that were due to OA. We include data from the AOANJRR between 1 March 2002 and 21 September 2016. These data excluded participants who died, collected from the Tasmanian Death Registry and who left Australia, which was collected from TASOAC questionnaires.

BMI

Body mass index (BMI) was calculated (weight (in kilograms)/height (in metres)²) using weight measured to the nearest 0.1 kg (with shoes, socks, bulky clothing and headwear removed) using a single pair of calibrated electronic scales (Seca Delta Model 707), and height measured to the nearest 0.1 cm (with shoes and socks removed) using a stadiometer.

Hip pain

Self-reported hip pain over the past 30 days was assessed by questionnaire at Phase 2 and 3 using the Western Ontario and McMaster Universities Osteoarthritis (WOMAC) index.(8, 21) Briefly, the WOMAC pain scale has five items, each rated on a 10-point numeric rating scale from 0 (no pain) to 9 (most severe pain). Each pain item was summed to create a total pain score (0–45).

Hip radiographs and assessment of hip radiographic OA (ROA) and cam impingement

Anteroposterior radiographs of the pelvis were obtained at Phase 1, with the individual standing with both feet internally rotated by 10 degrees. Radiographs were read by two trained readers using the OARSI (Osteoarthritis Research Society International) grading system.(22) Radiographic features of joint space narrowing (JSN) (axial and superior) and osteophytes (superior, acetabular and femoral) of both hips were graded separately on a 4-point scale (range 0–3 where 0 is no disease and 3 is severe disease. Data from these four features were summed (range 0–12).

The α angle measures the extent to which the femoral head deviates from spherical and is used to quantify cam impingement. It is measured by first drawing the best fitting circle around the femoral head, and then a line through the centre of the neck and the centre of the head. From the centre of the femoral head, a second line is drawn to the point where the superior surface of the head-neck junction first departs from the circle. The angle between these two lines is the α angle. We defined cam impingement by using a previously published standardised cut off point of 60° either in one or both hips.(23) The α angle was calculated by drawing a circle of best fit based on the statistical shape modeling (SSM) points

around the femoral head using custom code in MatLab (v 9.0). This method has good reliability as was shown previously with intraclass correlation coefficient (ICC) for inter-observer reliability of 0.73 and intra-observer reliability of 0.85–0.99.(9)

DXA Imaging and Statistical Shape Modelling (SSM)

Participants had dual-energy X-ray absorptiometry (DXA) images taken of the left hip, unless contra-indicated, using a Hologic Delphi densitometer (Hologic Inc., Waltham, MA, USA) as part of the Phase 1 assessment. Participants were excluded from DXA scanning if their weight exceeded 130 kg ($n = 3$). Left hip images were used to assess bone mass; examined as areal BMD at neck of femur (g/cm^2). This is calculated by dividing the bone mineral content (BMC) by the area measured. Precision was estimated to be 2% in vivo.

Statistical shape modelling (SSM) was used to describe hip shape variation within the study population. Briefly the proximal femur and acetabulum were modelled for each image using a template of 85 points placed on defined anatomical landmarks using the Active Shape Modelling toolkit (University of Manchester, UK).(24, 25) The images and points were transferred to the Shape software (University of Aberdeen, UK), where they were rotated and scaled using the Procrustes transform and then subjected to Principal Component Analysis to generate independent, orthogonal modes of variation. The modes of variation were then normalized to a mean of 0 and expressed as standard deviations from the mean. The modes of variation described decreasing amounts of variation within the model with the first 6 modes describing 68% of the total model variation.

Magnetic resonance imaging (MRI)

A subgroup ($n = 250$) had MRI. The right hip was imaged in the sagittal plane during visits at phases 2 and 3 using a 1.5 T GE Signa whole-body magnetic resonance scanner, as previously described.(8) Subchondral BMLs and effusion-synovitis were assessed on the short T1 inversion recovery (STIR)–weighted, fat saturation, 2-dimensional fast spin-echo sequence using OsiriX software (Mac version, University of Geneva, Geneva, Switzerland). BMLs were identified as areas of increased signal intensity adjacent to the subchondral bone on the femoral head and/or the acetabulum.(8) Hip effusion-synovitis was identified and assessed in STIR images from phases 2 and 3. The observer (HGA) manually selected the MRI slice with the largest effusion-synovitis and determined the maximum cross-sectional area (CSA) of the bright region by manually drawing contours around the outer edges, as previously described. Inter-rater reliability was excellent (0.84).(8) BMLs and effusions were dichotomised as present ($\text{CSA} > 0$) or absent ($\text{CSA} = 0$).

Statistical analysis

Differences between participants who did and did not have hip replacements were assessed using Students' t-tests and chi-squared tests.

Risk of THR in addition to the 'base model' (WOMAC hip pain score, and radiographic hip OA score) was assessed using mixed-effect Poisson regression, in which each potential risk factor was designated as a fixed effect and participant identification as a random effect. Models were run for each hip separately using the xt function, with side-specific WOMAC pain score and data from radiographs (ROA and alpha angle) used for risk of THR of each hip, while data from DXA (BMD and SSM) and MRI (BML, effusion) had data from one hip only (left hip for DXA, right hip for MRI) and was used to predict risk of THR in either hip. Standard errors were adjusted using the sandwich (robust) estimator of variance.(26) We used WOMAC hip pain as continuous data (range 0–35), but collapsed radiographic hip OA scores into categories as effect sizes were similar within groups. The relationship between each of the risk factors and the incidence of THR during follow-up was assessed using Cox proportional hazards regression models. Hazard ratios (HRs) and 95% confidence intervals (CIs) were calculated. Model assumption was checked and confirmed using the proportional hazards test.(27) We performed a sensitivity analysis, using a competing risk regression model to account for competing risks, which occurred within the study time frame (death, left Australia).

We used Stata 15.0 (StataCorp LP) for all statistical analyses. Statistical significance was defined as a p value ≤ 0.05 (two tailed).

Results

Eight hundred and two (802) participants had WOMAC hip pain data and data on radiographic hip grade. Of these, 40 individuals had at least one THR for OA, 15 participants had bilateral THR. Those who received a hip replacement were more likely to be smokers, have greater WOMAC hip pain scores, greater neck of femur BMD, more severe radiographic hip OA, more likely to have a BML, higher mode 2 and lower mode 4 shape scores, and were more likely to have a cam impingement in either left or right hip (Table 1). Study participants were followed for an average of 12.1 years (maximum 14 years).

Table 1
Summary of participant characteristics, by hip replacement status

	No hip replacement (mean \pm SD) (n = 762)	Hip replacement (mean \pm SD) (n = 40)	p
Left hip replacement only (n, %)	-	16 (40%)	
Right hip replacement only (n, %)	-	9 (23%)	
Bilateral hip replacement (n, %)	-	15 (38%)	
Age (years)	62.5 (7.3)	63.3 (7.1)	0.51
Sex (female: %)	51	45	0.44
Body mass index (kg/m ²)	27.8 (4.6)	28.1 (4)	0.65
Current Smoker (%)	11	23	0.03
WOMAC hip pain, P2 (range 0–45)	2.3 (5.2)	7.3 (8.7)	< 0.001
Neck of femur BMD (g/cm ²), left hip.	0.77 (0.12)	0.83 (0.14)	< 0.001
Radiographic hip OA score, mean of both hips, (score 0–4+)	0.68 (1.03)	1.91 (2.11)	0.001
Hip BML (P2 or P3) (%)	22	57	0.03
Hip effusion (P 2 or P3) (%)	83	83	1.00
Mode 2, left hip (SD from the mean)	-0.05 (0.97)	0.4 (1.3)	0.01
Mode 4, left hip (SD from the mean)	-0.01 (0.98)	-0.4 (1)	0.016
Cam impingement ($\alpha \geq 60^\circ$), mean of both hips	39	67	< 0.001
WOMAC: Western Ontario and McMaster Universities Osteoarthritis Index			
BMD: bone mineral density			
BML: bone marrow lesion			

As expected, WOMAC hip pain and radiographic hip OA predicted risk of THR. In addition, greater mode 2 scores (decreasing acetabular coverage) and lower shape mode 4 scores (non-spherical femoral head) predicted risk of THR. Cam impingement also increased risk of THR, as did higher BMD at the neck of femur. MRI detected BMLs and hip effusion, increased the risk of THR, with significant associations with

BMLs in the sub population with MRI available. The addition of age, sex and BMI to the model did not alter the risk of THR independent of WOMAC hip pain and radiographic hip OA (Table 2).

Table 2
Risk factors for THR in addition to WOMAC hip pain and radiographic hip OA

	Relative Risk (RR) (95% CI)
Base model: WOMAC hip pain and radiographic hip OA, n = 802	
WOMAC hip pain	1.08 (1.05 to 1.12)
Radiographic hip OA	
Scores 0	1 (reference)
Scores 1–3 (Grade 1)	2.36 (1.17 to 4.77)
Scores 4+ (Grade 2 or 3)	10.42 (4.49 to 24.18)
Base model plus	
Shape mode 2, n = 617	1.57 (1.01 to 2.46)
Shape mode 4, n = 617	0.65 (0.44 to 0.97)
Presence of cam impingement (α angle $\geq 60^\circ$), n = 786	2.66 (1.38 to 5.13)
Hip BMLs, n = 215	5.62 (1.1 to 28.81)
Hip effusions, n = 215	1.88 (0.24 to 14.78)
Neck of femur BMD (per SD), n = 802	1.85 (1.40 to 2.44)
Age, sex and BMI n = 802	1.01 (0.96 to 1.05)
	0.79 (0.4 to 1.55)
	0.98 (0.92 to 1.04)
WOMAC: Western Ontario and McMaster Universities Osteoarthritis Index	
BMD: bone mineral density	
BML: bone marrow lesion	
BMI: body mass index	

We also investigated associations between hip ROA score, hip pain score and cam impingement with incidence of THR over time. The highest cumulative hazard (30% after 14 years of observation) was observed in participants with greater ROA score and higher pain scores (WOMAC pain score ≥ 4) (Fig. 1). Similarly, cam impingement increased incidence of THR by approximately 10% over the study timeframe

(Fig. 2). Sensitivity analyses were performed to account for competing risks (predominantly competing risk of death) but these did not change the results (data not shown).

Discussion

This prospective population-based cohort study of older adults showed that abnormal hip shape (decreasing acetabular coverage and non-spherical femoral head), cam deformity, higher BMD and BMLs predicted the risk of THR independent of WOMAC hip pain score and radiographic hip OA. Hip effusions also increased risk of THR but this association did not reach statistical significance. Age, sex, and BMI did not predict THR independently of pain and radiographic hip OA. These results, if replicated, can be used to develop predictive models for THR.

This study extends the literature that hip shape and cam impingement increase risk of THR, independent of hip pain and radiographic hip OA. It is worth noting that whilst cam impingement and shape modes are calculated differently, they are capturing similar aspects of hip morphology and are, therefore, not completely independent measures.(28) Both measures, however, reflect changes in the bone, rather than the cartilage, and show that hip OA is driven strongly by bone shape. Of the 6 modes which accounted for 68% of the total variation in the population, mode 2 (decreasing acetabular coverage) and mode 4 (non-spherical femoral head) have been previously associated with THR in this sample(8). They were included in this manuscript for completeness and to compare with other structural measures. Hip shape; specifically, flattening of the femoral head (non-spherical femoral head)(6, 7) and decreasing acetabular coverage(5) were found to predict THR in different community based populations. One study adjusted for pain,(6) which negated the association. Cam impingement (structural malformation of the femoral head-neck junction) has also been found to be a risk factor for THR and accelerated hip OA in a community population study, but this study did not adjust for pain.(9)

A recent cohort study showed that the combination of radiographic hip OA and higher BMD as well as the BMD difference between the most affected hip and the contralateral hip predicted progression of hip OA, which included THR.(17) This study is the first to show that BMD, independent of radiographic hip OA, is a predictor for THR. A higher BMD may reflect the presence of osteophytes in hip OA(29) or bone hyperplasia reported previously(30) and may be enhanced by the surgical recommendation of patient selection of those with good bone quality for THR.(31) The former is unlikely as we adjusted for hip ROA including osteophytes. We do not have data on the latter.

We demonstrated no associations between advancing age, BMI or sex. A cross sectional study found that those with a higher BMI ($> 35 \text{ kg/m}^2$) had a THR at a younger age compared to those with BMI $< 25 \text{ kg/m}^2$ (32) and prospective cohort studies have identified increased risk in older, obese people(33–35) and an increased risk in men.(34) However, none of these studies assessed relationships independent of hip OA and pain, suggesting that the findings in these cohort studies is mediated by or confounded by hip pain and ROA.

Changes in hip structures seen on MRI (eg. BMLs, cartilage defects) have been previously demonstrated in patients with hip OA.(15, 16) Similarly, particular hip shapes correlated to MRI features of hip OA.(8) In this cohort, MRI changes were associated with a higher risk of THR, with BMLs reaching statistical significance. This is consistent with data for the knee where BMLs are a strong independent predictor of TKR.(36) We are not aware of any previous data for the hip. Hip effusion was associated with a higher risk of THR, however this did not reach statistical significance, possibly due to the smaller number of participants in this model (n = 215).

Limitations of this study include the difference in number of participants in some models based on the data from which predictors were collected. In particular, MRI data were only available for a subset of the cohort (215 participants), however the smaller sample size was unlikely to be the reason that why effusions did not predict THR (RR 1.88 (0.24 to 14.78), p = 0.50). However, the risk estimates for BMLs from the MRI data are consistent with the knee literature, suggesting that these associations are in the clinically important range. Whilst patient access to THR may be a potential confounder, data from this cohort demonstrates that socio-economic status does not predict time to hip replacement (unpublished in-house data), demonstrating that the publicly funded hospital system in Australia has enabled timely access to THR in TASOAC participants regardless of their socio-economic status. Study participants could be lost to follow up due to death, illness or leaving Australia. However, as we were able to perform sensitivity analyses for competing risks (due primarily to death), and results did not change, we conclude that data are not biased by loss to follow up. Strengths of this study are the large cohort of participants, the prospective design and long-term follow up, the completeness of the AOANJRR data, and the analysis of multiple variables in the same population cohort.

Conclusion

In this community-based study, hip structural changes as well as MRI changes predicted the risk of THR. These risk factors were independent of hip pain and radiographic hip OA, which has not been shown previously. Such factors can lead to better predictive models for THR and enhance our understanding of the pathogenesis of hip OA.

Abbreviations

Osteoarthritis

OA

Total hip replacement

THR

Dual energy X-ray absorptiometry

DXA

Bone mineral density

BMD

Western Ontario and McMaster Universities Osteoarthritis Index

WOMAC

Magnetic resonance imaging

MRI

Bone marrow lesion

BML

Tasmanian Older Adult Cohort

TASOAC

Australian Orthopaedic Association National Joint Replacement Registry

AOANJRR

Body mass index

BMI

Osteoarthritis Research Society International

OARSI

Statistical shape modeling

SSM

Declarations

Ethics approval and consent to participate:

The Southern Tasmanian Health and Medical Human Research Ethics Committee approved the study, and we obtained written informed consent from all participants.

Consent for publication:

Not applicable

Availability of data and materials:

The datasets used and analysed during the current study are available from author G Jones (graeme.jones@utas.edu.au) on reasonable request.

Competing Interests:

The authors declare that they have no competing interests

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Authors Contributions:

VM, LL, HA, RA, JG, FS, IM, GC, FC and GJ were major contributors in writing the manuscript. LL and CB analysed the dataset. GJ designed the study. All authors read and approved the final manuscript.

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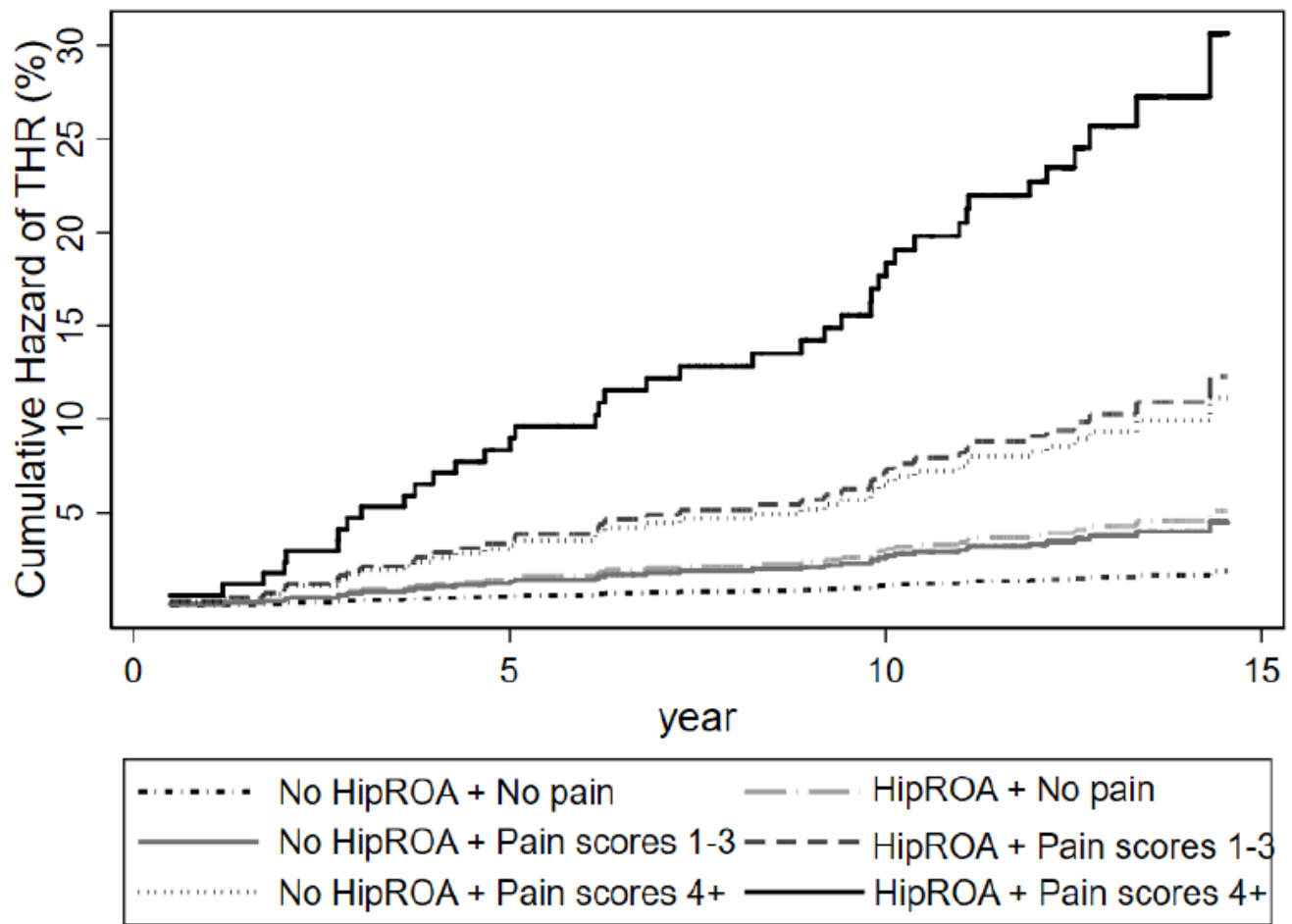
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Figures



Cox proportional hazards regression

Figure 1

Cumulative hazard of THR, by presence of radiographic hip OA and WOMAC hip pain intensity

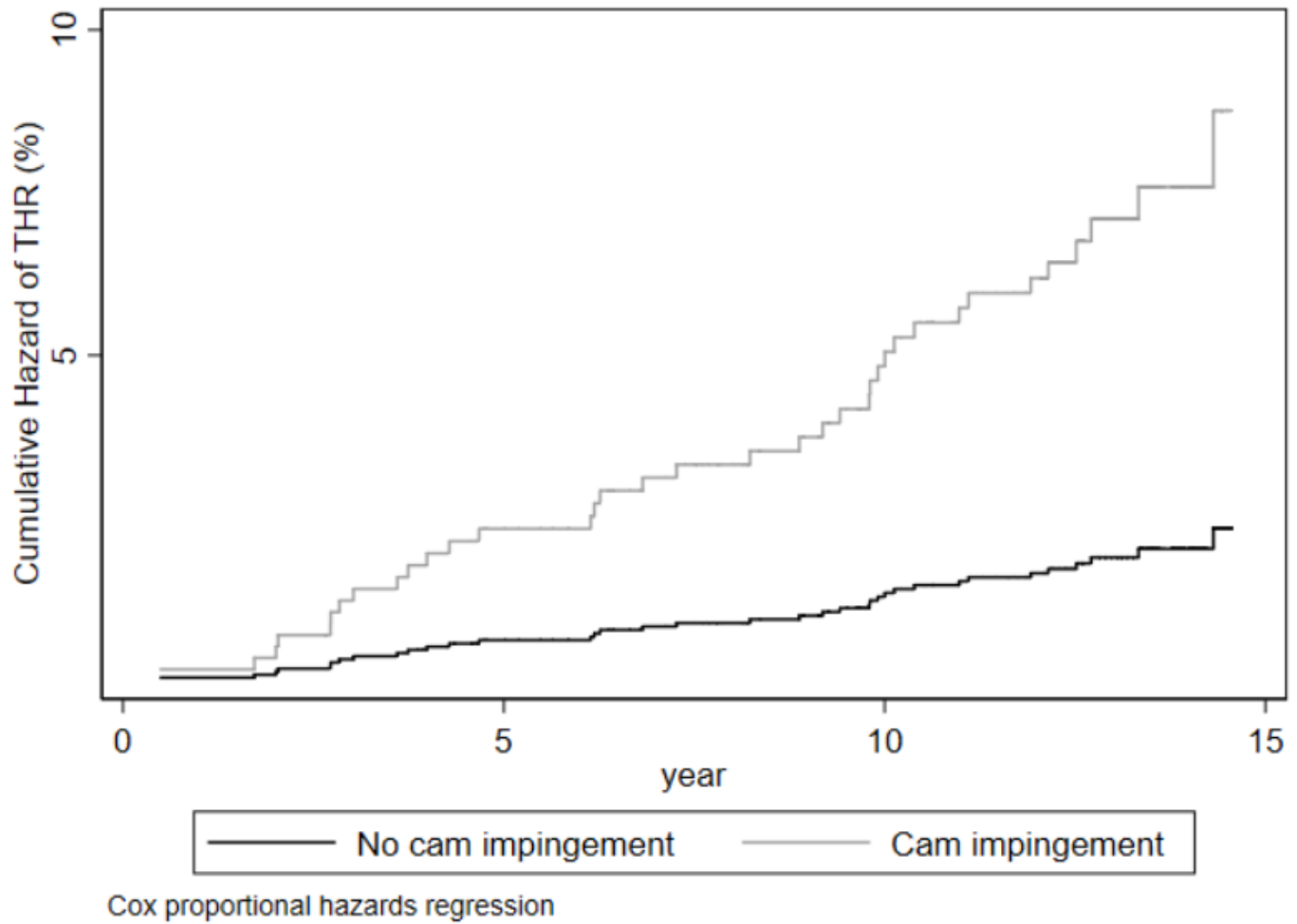


Figure 2

Cumulative hazard of THR, by presence of cam impingement