

Femoroacetabular impingement syndrome in middle-aged individuals is strongly associated with the development of hip osteoarthritis within 10-year follow-up a prospective cohort study (CHECK)

Agricola, Rintje; Van Buuren, Michiel M.A.; Kemp, Joanne L.; Weinans, Harrie; Runhaar, Jos; Bierma-Zeinstra, Sita M.A.

DOI

10.1136/bjsports-2024-108222

Publication date 2024

**Document Version**Final published version

**Published in**British Journal of Sports Medicine

Citation (APA)

Agricola, R., Van Buuren, M. M. A., Kemp, J. L., Weinans, H., Runhaar, J., & Bierma-Zeinstra, S. M. A. (2024). Femoroacetabular impingement syndrome in middle-aged individuals is strongly associated with the development of hip osteoarthritis within 10-year follow-up: a prospective cohort study (CHECK). *British Journal of Sports Medicine*, *58*(18), 1061-1067. Article 108222. https://doi.org/10.1136/bjsports-2024-108222

# Important note

To cite this publication, please use the final published version (if applicable). Please check the document version above.

Copyright

Other than for strictly personal use, it is not permitted to download, forward or distribute the text or part of it, without the consent of the author(s) and/or copyright holder(s), unless the work is under an open content license such as Creative Commons.

Takedown policy

Please contact us and provide details if you believe this document breaches copyrights. We will remove access to the work immediately and investigate your claim.



# Femoroacetabular impingement syndrome in middleaged individuals is strongly associated with the development of hip osteoarthritis within 10-year follow-up: a prospective cohort study (CHECK)

Rintje Agricola , <sup>1</sup> Michiel M A van Buuren, <sup>1</sup> Joanne L Kemp , <sup>2</sup> Harrie Weinans, <sup>3,4</sup> Jos Runhaar,<sup>5</sup> Sita M A Bierma-Zeinstra<sup>5</sup>

#### ► Additional supplemental material is published online only. To view, please visit the journal online (https://doi. org/10.1136/bjsports-2024-108222)

<sup>1</sup>Department of Orthopaedics and Sports Medicine, Erasmus Medical Center, Rotterdam, The Netherlands

<sup>2</sup>Latrobe Sports Exercise Medicine Research Centre. School of Allied Health, Human Services and Sport, La Trobe University, Bundoora, Victoria, Australia

<sup>3</sup>Department of Orthopaedics, University Medical Centre Utrecht, Útrecht, The Netherlands

<sup>4</sup>Biomechanical Engineering, Delft University of Technology, Delft, The Netherlands <sup>5</sup>Department of General Practice, Erasmus Medical Center, Rotterdam, The Netherlands

#### Correspondence to Rintje Agricola; r.agricola@erasmusmc.nl

Accepted 26 June 2024

# **ABSTRACT**

**Objective** The objective is to determine the association and absolute risk of femoroacetabular impingement syndrome (FAIS) for the development of radiographic hip osteoarthritis (RHOA).

**Methods** This is a nationwide, multicentre prospective cohort study (Cohort Hip and Cohort Knee) with 1002 individuals aged between 45 and 65 years. Hips without definitive RHOA (Kellgren-Lawrence (KL) grade≤1) at baseline and with anteroposterior pelvic radiographs at baseline and 10-year follow-up available (n=1386 hips) were included. FAIS was defined by the baseline presence of a painful hip, limited internal hip rotation≤25° and cam morphology defined by an alpha angle>60°. The outcomes were incident RHOA (KL grade≥2 or total hip replacement (THR)) and incident end-stage RHOA (KL≥3 or THR) within 10 years.

Results Of the 1386 included hips (80% women; mean age 55.7±5.2 years), 21 hips fulfilled criteria for FAIS and 563 hips did not fulfil any of the FAIS criteria (reference group; no symptoms, no signs, no cam morphology). Within 10-year follow-up, 221 hips (38%) developed incident RHOA and 15 hips (3%) developed end-stage RHOA (including 9 hips with THR). Adjusted for sex, age and body mass index, FAIS with cam morphology resulted in an OR of 6.85 (95% CI 2.10 to 22.35) for incident RHOA and 47.82 (95% CI 12.51 to 182,76) for incident end-stage RHOA, compared with hips not having any FAIS criteria. The absolute risk of FAIS was 81% for incident RHOA and 33% for incident end-stage RHOA.

**Conclusion** FAIS was strongly associated with the development of RHOA within 10 years. Although the baseline prevalence of FAIS was low, the high absolute risk of FAIS for RHOA warrants further studies to determine preventive strategies.

# Check for updates

@ Author(s) (or their employer(s)) 2024. Re-use permitted under CC BY-NC. No commercial re-use. See rights and permissions. Published by BMJ.

To cite: Agricola R, van Buuren MMA, Kemp JL, et al. Br J Sports Med Epub ahead of print: [please include Day Month Year]. doi:10.1136/ bjsports-2024-108222

**BMJ** Group

## INTRODUCTION

Osteoarthritis (OA) is a common and disabling disease with a large socioeconomic impact on individuals and society. 1-3 Hip OA is more prevalent among athletes who practised high-impact sports at an elite level, although the underlying mechanism is unknown, making preventive measures challenging. A potential risk factor that might account for this effect is cam morphology, which is both highly prevalent in athletes and associated with the development of radiographic hip OA (RHOA) in

# WHAT IS ALREADY KNOWN ON THIS TOPIC

- ⇒ Prospective cohort studies have shown an association between the radiographic presence of cam morphology and development of hip osteoarthritis (OA), although absolute risks were generally low.
- ⇒ Femoroacetabular impingement syndrome (FAIS) is a clinical condition which consists of a triad including symptoms, signs and radiographic findings (cam morphology).
- ⇒ Only two small cross-sectional studies have investigated the association between FAIS and cartilage defects in young athletic populations.

#### WHAT THIS STUDY ADDS

- ⇒ FAIS was strongly associated with a nearly sevenfold increased odds of hip OA within 10
- ⇒ The absolute risk of FAIS for development of hip OA was high (81%), with 33% developing endstage hip OA within 10 years.
- ⇒ By using simple and accessible measures (symptoms, a clinical hip examination and an anteroposterior pelvic radiograph), it is possible to distinguish a subgroup of people presenting with first onset hip pain at high risk for developing future hip OA.

# HOW THIS STUDY MIGHT AFFECT RESEARCH. **PRACTICE OR POLICY**

⇒ FAIS is an important risk factor for developing hip OA and warrants additional research to define preventive strategies.

several prospective cohort studies.<sup>5–7</sup> On the other hand, cam morphology is also highly prevalent in the asymptomatic population and does not necessarily lead to RHOA in all individuals.8 The absolute risk of cam morphology for the subsequent development of RHOA has been reported to be between 6% and 25% only.

The mechanism by which cam morphology might lead to OA is femoroacetabular impingement syndrome (FAIS), a motion-related clinical disorder which represents a premature contact between the proximal femur and acetabulum. 10-12 In a 2016 consensus meeting, it was agreed on that the diagnosis of FAIS cannot be made by the radiological presence of cam morphology alone; symptoms and



# Original research

 Table 1
 Baseline characteristics of the included and excluded participants

	Included participants, with at least one hip	Excluded participants, with both hips excluded	
	included (n=744)	(n=258)	P value
Age, years	55.7 (5.2)	56.6 (5.2)	0.01
BMI, kg/m <sup>2</sup>	26.2 (4.0)	26.0 (4.0)	0.43
Height, cm*	169.8 (8.4)	170.1 (8.0)	0.49
Weight, kg*	75.8 (13.2)	75.2 (14.2)	0.79
Sex			
Male	146 (20%)	64 (25%)	0.08
Female	598 (80%)	194 (75%)	

Values are mean (SD) or number (percentage).

Included participants had either one or both hips included in the analysis. In excluded participants, both hips were excluded.

Bold value represent statistical significant difference (p<0.05).

\*For height and weight, total n=848 persons (154 missing).

BMI, body mass index.

clinical signs consistent with FAIS should also be present. Symptoms include motion-related or position-related hip or groin pain, and clinical signs include a limited range of internal hip rotation or a painful sensation during the flexion–adduction–internal rotation (FADIR) test. <sup>10</sup> <sup>13</sup> <sup>14</sup> This triad (symptoms, signs and radiographic findings) should all be present to diagnose FAIS. <sup>10</sup> <sup>12</sup>

The presence of FAIS might better identify people at risk for RHOA, rather than the presence of a radiographic cam morphology alone. If this holds true, it might potentially enable preventive measures, as both surgical and non-surgical treatment options for FAIS are available. <sup>15</sup> <sup>16</sup> To date, there are only two small cross-sectional studies available which investigated the association between FAIS and cartilage defects in the athletic population aged < 50 years. <sup>17</sup> <sup>18</sup> To the best of our knowledge, no prospective studies on the association between FAIS and development of RHOA are available and has recently been identified as a research priority. <sup>19</sup>

The aim of this study was to investigate the association between FAIS at baseline and the development of RHOA within 10-year follow-up and to report corresponding absolute risks.

#### **METHODS**

# Study design and participants

The Cohort Hip and Cohort Knee (CHECK) study is a nation-wide multicentre prospective cohort study of 1002 Dutch individuals aiming to study the cause and course of complaints of OA as well as to identify markers for diagnosis and prognosis. Participants were eligible to enter the cohort if they had pain or stiffness in hip and/or knee and were aged 45–65 years. To be eligible, they should not yet have consulted their general practitioner for these symptoms, or the first consultation was within 6 months before entry. Participants with a pathological condition that could explain the symptoms were excluded (for hip: trauma, rheumatoid arthritis, known developmental dysplasia of the hip, Perthes disease, subluxation, osteochondritis dissecans, fracture, septic arthritis, Kellgren and Lawrence (K&L) grade 4 or total hip replacement (THR), previous hip surgery and individuals having only symptoms of bursitis or tendinopathy).

Data were obtained from 11 (general and university) hospitals. General practitioners were invited to refer eligible persons to one of those centres; advertisements in local newspapers were also used. For the hip, questionnaires and clinical hip

examination were obtained annually until 10-year follow-up. Radiographs of the hip were obtained at baseline (from October 2002 to December 2005), 2-year, 5-year, 8-year and 10-year follow-up. For the current study, baseline data were used for the exposure variables and the 10-year follow-up was used to define the outcome of RHOA. At both time points, weight-bearing anteroposterior (AP) pelvic views were obtained according to a standardised protocol which has been described previously. For the first 112 participants who entered the cohort, AP hip views instead of AP pelvic views were obtained. Of the 1002 participants at baseline, only hips free of definite RHOA (K&L≤1) were included for the current study. We followed the STrengthening the Reporting of OBservational studies in Epidemiology guidelines for observational studies.

#### **Exposure assessment**

The exposure was FAIS at baseline, defined as the presence of three criteria: symptoms, clinical signs and radiographic cam morphology, according to the Warwick agreement.<sup>10</sup>

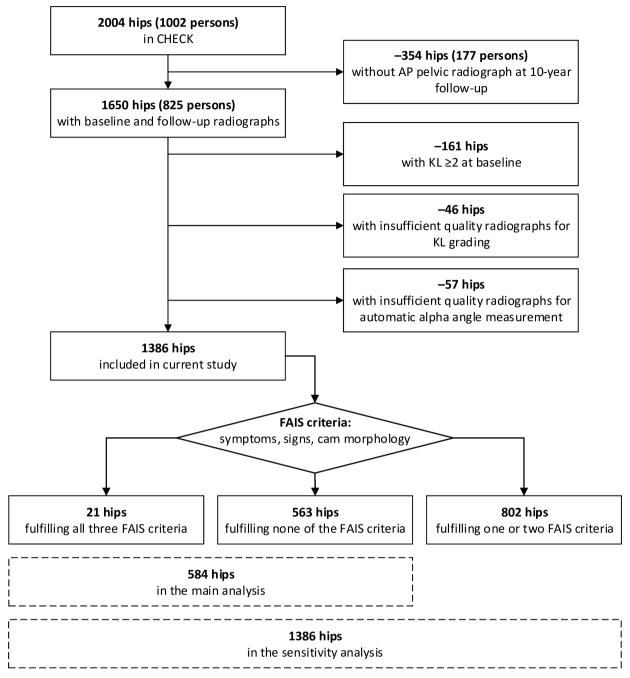
Symptoms were defined as the presence of self-reported hip pain. This was a dichotomous question and the same variable that determined at baseline whether they were eligible to participate in this cohort. For sensitivity analysis, we also used another variable of the presence/absence of self-reported hip pain or stiffness which was posed by a questionnaire ('do you have pain or stiffness in your hip, groin or upper thigh?'). Participants also had to indicate whether the symptoms were present in the right hip, left hip or both hips.

Clinical signs of FAIS were defined by a limited internal hip rotation of  $\leq 25^{\circ}$ . <sup>14</sup> Hip internal rotation was measured according to a standardised protocol by a goniometer in sitting position with the hip in 90° of flexion, which previously showed satisfactory reliability. <sup>22</sup> Due to a lack of consensus on the internal hip rotation threshold value to define FAIS, we performed a sensitivity analysis using a threshold of  $\leq 20^{\circ}$ .

Cam morphology was defined by the alpha angle on the baseline radiographs. This method has been described before.<sup>21</sup> In short, the shape of the proximal femur was manually outlined with a set of points using statistical shape modelling (SSM) software (ASM toolkit, Manchester University, Manchester, the UK). From this set of points, the alpha angle was automatically calculated using a custom Matlab script (V.7.1.0). The alpha angle was calculated by drawing a best-fitted circle around the femoral head. Then lines were drawn from the centre of the femoral head through the axis of the femoral neck and from the centre of the femoral head through the point where the bone leaves the fitted circle. We classified the presence of cam morphology by a validated alpha angle threshold value of >60°. <sup>23</sup> <sup>24</sup> As higher alpha angles increase the risk of developing hip OA, we also present results for an alpha angle threshold of>78°, which has previously been shown to best discriminate between hips that did and did not develop hip OA.<sup>23</sup> We previously reported an interobserver reliability of 0.73 and intraobserver reliability ranging from 0.85 to 0.99 for the alpha angle in this cohort.<sup>21</sup>

#### **Outcome assessment**

The primary outcome was incident RHOA as defined by a K&L grade≥2 or a THR at 10-year follow-up. The secondary outcome was incident end-stage RHOA as defined by a K&L grade≥3 or a THR at 10-year follow-up. All radiographs were scored for RHOA features using the Osteoarthritis Research Society International (OARSI) atlas, and RHOA was graded according to the K&L classification (grade 0–4) by experienced



**Figure 1** Flowchart of hips from cohort entry to hips included for the current study. AP, anteroposterior; CHECK, Cohort Hip and Cohort Knee; FAIS, femoroacetabular impingement syndrome; KL, Kellgren-Lawrence.

and well-trained readers. <sup>25-27</sup> The radiographs of all available time points (baseline, 2-year, 5-year, 8-year and 10-year follow-up) of each participant were scored simultaneously and previously showed substantial to almost perfect interobserver reliability with average prevalence adjusted bias adjusted kappa values ranging from 0.71 to 0.91 for the different radiographic OA features. <sup>28</sup> At baseline, we only included hips with a K&L grade≤1, indicating hips without definite signs of RHOA. This way, we minimalised the risk that cam morphology was misclassified due to osteoarthritic changes (eg, osteophyte formation, femoral head deformation).

#### Equity, diversity and inclusion

The majority of participants were women (80%) and participants were recruited from all socioeconomic levels. No particular

effort was made to include or exclude minorities. The authors are from varying career stages and disciplines, with two (33%) women.

#### Patient and public involvement

Two OA patients were part of the CHECK steering committee in the set-up of the study. Throughout the study period, regular patient and public meetings were held. Patients were involved in the design, interpretation of results and dissemination strategies.

#### Statistical analyses

The Shapiro-Wilk test was used to test for normality. Differences in baseline characteristics between included and excluded hips were evaluated by the Mann-Whitney U test for continuous

# Original research

**Table 2** Associations between femoroacetabular impingement syndrome and the development of incident radiographic hip osteoarthritis and incident end-stage radiographic hip osteoarthritis within 10-year follow-up

	Total n=584		Incident hip OA (KL 2-4 or THR) n=221		Incident end-stage hip OA (KL 3-4 or THR) n=15	
Exposure	N with condition	N without condition*+	OR (95% CI)	aOR† (95% CI)	OR (95% CI)	aOR† (95% CI)
FAIS (hip pain, internal hip rotation≤25°,cam morphology with alpha angle >60°)	21 (3.6%)	563 (96.4%)	7.5 (2.4 to 23.4)	6.9 (2.1 to 22.4)	34.6 (10.8 to 110.8)	47.8 (12.5 to 182.8)
FAIS with large cam morphology (hip pain, internal hip rotation≤25°,large cam morphology with alpha angle>78°)	14 (2.4%)	563 (96.4%)	6.3 (2.8 to 22.4)	5.6 (1.5 to 21.5)	51.9 (14.6 to 184.9)	88.4 (17.7 to 441.4)

<sup>\*</sup>The reference group for the predictor categories in this table consists of hips that did not have any of the stated conditions (eg, the reference group for FAIS are hips without cam, without hip pain, and without decreased internal rotation).

variables and by the  $\chi^2$  test for sex. The association between baseline hips with FAIS (with all three criteria present: symptoms, signs and cam morphology) as compared with hips not having any of these criteria and the development of hip OA within 10-year follow-up were calculated using logistic regression with generalised estimating equations (GEE). The strength of association was expressed in terms of OR with 95% CIs. The use of GEE allowed for modelling the correlation between the left and the right hip in the same person. To adjust for baseline confounders, sex was entered as a factor and body mass index (BMI) and age as a covariate in the GEE model. For sensitivity purposes, we repeated this analysis using a reference group of hips that could have one or two out of three criteria of FAIS instead of a reference group with complete absence of any FAIS feature; therefore, this reference group could also contain hips with for example pain and cam morphology but an IR>25°. The absolute risk of FAIS for RHOA was calculated and expressed as percentage. If available, baseline characteristics (age, sex, BMI) of follow-up visits were used if these were missing at baseline. Missing values from questionnaire data were excluded for analysis. All statistical analyses were performed in SPSS V.25.

#### **RESULTS**

#### Study population

Of the 1002 individuals (2004 hips) in the CHECK cohort, 825 (1650 hips) had AP pelvic radiographs available at 10-year follow-up (82%). Of these 1650 hips, 161 hips were excluded because a K&L grade of ≥2 at baseline, 46 hips were excluded because of missing or insufficient quality baseline radiographs for reliable K&L grading and 57 hips were excluded due to insufficient quality radiographs for outlining the bone with SSM and measuring the alpha angle; leaving 1386 hips. Participants with hips excluded were slightly older and taller and more likely to be man than included participants (table 1).

Of the included 1386 hips, 21 hips fulfilled the criteria of FAIS (symptoms, signs and cam morphology), 563 hips did not fulfil any of the FAIS criteria (reference group; no symptoms, no signs, no cam morphology) and 802 hips met one or two criteria of FAIS but did not meet all three criteria (figure 1). The number of hips which meet the separate criteria of FAIS (symptomatic vs asymptomatic, signs vs no signs) can be found in online supplemental table 1. Of the 1386 hips, 16 hips had missing data for baseline BMI. In six of those, BMI was available at 1 year and in eight hips at 2-year follow-up and these values were used. Two hips (one participant) had missing data throughout the follow-up and were excluded for the adjusted analysis. For the question 'Do you have pain or stiffness in your hip, groin or upper thigh?', used for the sensitivity analysis, 17 hips had missing values and were excluded.

#### FAI syndrome and risk of RHOA

From the 584 hips at baseline for the primary analysis, 453 hips (78%) had K&L grade 0 and 131 hips (22%) grade 1. Within 10-year follow-up, 221 hips (38%) developed incident RHOA and 15 hips (3%) developed incident end-stage RHOA (including 9 hips with THR due to hip OA). None of the participants with a THR had hip surgery prior to the THR. FAIS was present in 21 hips at baseline, of which 15 also had pain on internal hip rotation. Of the 21 hips with FAIS, 9 were women and 12 were men. This resulted in an overall hip FAIS prevalence of 1.5% and a sex-specific hip prevalence of 4.6% in men and 0.8% in women. Adjusted for confounders, FAIS was associated with both incident RHOA with an OR of 6.85 (95% CI 2.1 to 22.4) and end-stage RHOA (OR=47.82, 95% CI 12.5 to 182.8). Results of unadjusted analyses and results for FAIS with cam morphology defined by an alpha angle>78° are presented in table 2. The absolute risk of FAIS was 81.0% for incident RHOA and 33.3% for incident end-stage RHOA (table 3). The

		te risks			
Incident hip OA (K	Incident end-stage hip OA (KL≥3 or THR)				
Present (n=221)	Absent (n=363)	Absolute risk	Present (n=15)	Absent (n=569)	Absolute risk
morphology with alpha angle>	60°)				
17	4	81.0%	7	14	33.3%
204	359		8	555	
ernal hip rotation≤25°°, cam m	orphology with alpha	angle>78°)			
11	3	78.6%	6	8	42.9%
204	359		8	555	
	Incident hip OA (K Present (n=221) morphology with alpha angle> 17 204 ernal hip rotation≤25°°, cam months	Incident hip OA (KL≥2 or THR)  Present (n=221) Absent (n=363)  morphology with alpha angle>60°)  17 4  204 359  ernal hip rotation≤25°°, cam morphology with alpha  11 3	Present (n=221) Absent (n=363) Absolute risk morphology with alpha angle>60°)  17 4 81.0% 204 359 ernal hip rotation≤25°°, cam morphology with alpha angle>78°) 11 3 78.6%	Incident hip OA (KL≥2 or THR)  Present (n=221)  Absent (n=363)  Absolute risk  Present (n=15)  morphology with alpha angle>60°)  17  4  81.0%  7  204  359  8  ernal hip rotation≤25°°, cam morphology with alpha angle>78°)  11  3  78.6%  6	Incident hip OA (KL≥2 or THR)

<sup>†</sup>Adjusted ORs are adjusted for age, sex and body mass index.

aOR, adjusted OR; FAIS, femoroacetabular impingement syndrome; KL, Kellgren-Lawrence grade; THR, total hip replacement.

sensitivity analysis using the question 'do you have pain or stiffness in your hip, groin or upper thigh' from the questionnaire to define hip/groin pain instead of the self-reported hip pain variable showed similar statistically significant results (online supplemental table 2). Similar results were also found for the sensitivity analysis using an internal hip rotation threshold of  $\leq 20^{\circ}$  instead of  $\leq 25^{\circ}$  (online supplemental table 3). Results from the sensitivity analysis using all hips that did not fulfil all three criteria of FAIS (hip pain, decreased internal rotation and cam morphology) are presented in online supplemental table 4 and showed significant associations between FAIS and both incident RHOA and incident end-stage RHOA.

#### **DISCUSSION**

In this first prospective cohort study on the relationship between FAIS and the development of RHOA, we showed a strong relationship between FAI syndrome and development of RHOA within 10 years and corresponding high positive predictive values. Although the prevalence of FAIS was low in this cohort of people aged between 45 and 65 years, this subgroup of people at high risk for developing RHOA could be identified using simple and accessible measures (clinical hip examination and an AP pelvic radiograph) from people that present with first onset of either hip or knee complaints to the general practitioner.

Previous prospective cohort studies have only investigated the relationship between the radiographic presence of cam morphology and development of RHOA and consistently showed a positive association.<sup>5</sup> 6 29 The strength of association between cam morphology and development of hip OA in prospective cohort studies ranged between ORs of 2.1 (95% CI 1.6 to 2.9) and 9.7 (95% CI 4.7 to 19.8).  $^{21\,30-33}$  Two small studies investigated the cross-sectional relationship between FAIS and MRI detected cartilage defects in younger (18-50 years) individuals. 17 18 In one study, individuals with FAIS showed cartilage defects more frequently than asymptomatic controls. 18 In the other study, cam morphology was associated with cartilage defects and labral tears in athletes although the presence or absence of symptoms did not influence this association, meaning that the association found was similar between those with FAIS and asymptomatic controls. 17 Interestingly, in the same cohort, symptoms were associated with cartilage loss severity in men indicating that the relation between hip and groin pain, FAIS and (early) hip OA is still poorly understood.<sup>34</sup> The cartilage lesions found in these studies of people aged < 50 years might be a precursor of definite OA later in life, as found in this study of people aged>45 years.

The high absolute risks found in this study support the necessity for increased awareness for FAIS and justify more research into possible preventive options to halt or delay the progression from FAIS towards hip OA. Previous epidemiological studies investigating only the radiographic presence of cam morphology reported absolute risks for hip OA between 6% and 25%. This suggests that although the radiographic presence of cam morphology is strongly associated with hip OA, the majority of hips with cam morphology will still not develop OA. In contrast, the absolute risk for FAIS ranged between 33% and 81% dependent on the size of cam morphology and definition of OA used. Adding symptoms and limited internal rotation to the presence of cam morphology enhances the likelihood of the prediction of hip OA. Hips with cam morphology with a larger range of internal hip rotation might not cause impingement and could therefore be less likely to result in hip OA.

There are opportunities for both primary and secondary prevention. Primary prevention would include some sort of activity modification during growth, as cam morphology develops during adolescence when the proximal femoral growth plate is still open.35 The formation of cam morphology is triggered by the loads applied to a growing hip, resulting from athletic activities, as cam morphology is rare in non-athletes. 36 37 Although the exact mechanism in terms of loading pattern, frequency and duration of loading in cam morphology development is still unknown, and such prevention programmes might be challenging to implement, there is a theoretical opportunity for primary prevention. Challenges that come with primary prevention are the conflict between load reduction during a given timeframe and the recommendations for adolescents to engage in sports, particularly at an age where skill development and talent identification is important.<sup>38</sup> Secondary prevention might be more feasible to implement and can include strategies to prevent the cam morphology from causing intraarticular damage. One could think that activity modification to prevent impingement between cam morphology and the acetabulum, <sup>39</sup> strength training, <sup>40</sup> improving balance, <sup>41</sup> and functional movement, 42 education about the importance of exercise and physical activity,<sup>39</sup> and other forms of physiotherapist-led rehabilitation<sup>43</sup> that target impairments could all be useful. Another secondary preventive option could be to surgically remove cam morphology. Although recent randomised controlled trials show a clinical benefit of both approaches in the short term, there is no long-term evidence available on secondary prevention of hip OA. 15 16

There are several strengths and weaknesses in this study that need to be acknowledged. Strengths are the prospective design and large sample size. Despite the large sample size, only 21 hips fulfilled the criteria of FAIS. One of the reasons is the higher proportion of women in this cohort, who have a lower prevalence of cam morphology and FAIS than men. Given the low prevalence of FAIS, the strength of association needs to be interpreted with caution, as the CIs around the ORs are wide. Larger prospective studies are needed to refine the strength of association. In the definition of FAIS, we used limited internal hip rotation as the clinical sign, which has recently been described to be best for ruling in FAIS, although the quality of evidence of available studies was low.14 The Flexion Adduction Internal Rotation (FADDIR) test has also been described as a clinical sign of FAIS, and its use has rather been suggested to rule out FAIS, 13 14 but the FADIR test was not available in this cohort. We only used AP pelvic radiographs to quantify cam morphology, which might have led to an underestimation of cam morphology prevalence. We only examined FAIS with cam morphology because FAI with pincer morphology follows another mechanism, and the relation between pincer morphology with OA has previously shown to be inconsistent.<sup>29</sup> <sup>44</sup> Therefore, we cannot draw any conclusions on FAIS with pincer morphology, which will need further study. Finally, the control group of non-FAIS hips at baseline also included people with first onset of knee pain and might not represent a control group completely free of pain.

#### CONCLUSION

FAIS was strongly associated with development of hip OA within 10 years. Although the prevalence of FAIS was low, the majority of people with FAIS developed OA within 10 years. The high

# Original research

absolute risk of FAIS for developing hip OA warrants further studies on preventive strategies.

X Rintje Agricola @RintjeAgricola, Michiel M A van Buuren @mmavanbuuren and Joanne L Kemp @JoanneLKemp

Acknowledgements The authors would like to thank all the participants of the CHECK cohort. CHECK cohort study is initiated by the Dutch Arthritis Association and performed within Erasmus Medical Center Rotterdam, Kennemer Gasthuis Haarlem, Leiden University Medical Center, Maastricht University Medical Center, Martini Hospital Groningen/Allied Health Care Center for Rheumatism and Rehabilitation Groningen, Medical Spectrum Twente Enschede/Ziekenhuisgroep Twente Almelo, Reade, formerly Jan van Breemen Institute/VU Medical Center Amsterdam, St Maartens kliniek Nijmegen, University Medical Center Utrecht and Wilhelmina Hospital Assen.

**Contributors** All authors substantially contributed to the conception or design of the work; the acquisition, analysis or interpretation of data for the work and drafting the work (RA) or revising it critically (MMAvB, JLK, HW, JR, SMAB-Z) for important intellectual content and final approval of the version to be published and agreement to be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved. RA is the guarantor.

**Funding** The CHECK study was funded by the Dutch Arthritis Society. The current study was part of projects funded by the Dutch Arthritis Society (21-1-205) and ZonMw (VENI 09150161910071).

Competing interests None declared.

**Patient and public involvement** Patients and/or the public were involved in the design, or conduct, or reporting, or dissemination plans of this research. Refer to the Methods section for further details.

Patient consent for publication Not applicable.

**Ethics approval** This study involves human participants and was approved by IRB of Utrecht University Medical Centre. Participants gave informed consent to participate in the study before taking part.

**Provenance and peer review** Not commissioned; externally peer reviewed.

**Data availability statement** Data are available on reasonable request. Data are available on reasonable request. The data underlying this article cannot be shared publicly due to legal reasons as well as the privacy of individuals that participated in the study.

**Supplemental material** This content has been supplied by the author(s). It has not been vetted by BMJ Publishing Group Limited (BMJ) and may not have been peer-reviewed. Any opinions or recommendations discussed are solely those of the author(s) and are not endorsed by BMJ. BMJ disclaims all liability and responsibility arising from any reliance placed on the content. Where the content includes any translated material, BMJ does not warrant the accuracy and reliability of the translations (including but not limited to local regulations, clinical guidelines, terminology, drug names and drug dosages), and is not responsible for any error and/or omissions arising from translation and adaptation or otherwise.

**Open access** This is an open access article distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited, appropriate credit is given, any changes made indicated, and the use is non-commercial. See: http://creativecommons.org/licenses/by-nc/4.0/.

#### **ORCID** iDs

Rintje Agricola http://orcid.org/0000-0002-0645-093X Joanne L Kemp http://orcid.org/0000-0002-9234-1923

## **REFERENCES**

- 1 Glyn-Jones S, Palmer AJR, Agricola R, et al. Osteoarthritis. Lancet 2015;386:376–87.
- 2 Prieto-Alhambra D, Judge A, Javaid MK, et al. Incidence and risk factors for clinically diagnosed knee, hip and hand osteoarthritis: influences of age, gender and osteoarthritis affecting other joints. Ann Rheum Dis 2014;73:1659–64.
- 3 Hunter DJ, Schofield D, Callander E. The individual and socioeconomic impact of osteoarthritis. *Nat Rev Rheumatol* 2014;10:437–41.
- 4 Vigdorchik JM, Nepple JJ, Eftekhary N, et al. What is the association of elite sporting activities with the development of hip osteoarthritis? Am J Sports Med 2017:45:961–4
- 5 van Klij P, Heerey J, Waarsing JH, et al. The prevalence of cam and pincer morphology and its association with development of hip osteoarthritis. J Orthop Sports Phys Ther 2018;48:230–8.

- 6 van Buuren MMA, Arden NK, Bierma-Zeinstra SMA, et al. Statistical shape modeling of the hip and the association with hip osteoarthritis: a systematic review. Osteoarthr Cartil 2021;29:607–18.
- 7 Tang J, van Buuren MMA, Riedstra NS, et al. Cam morphology is strongly and consistently associated with development of radiographic hip osteoarthritis throughout 4 follow-up visits within 10 years. Osteoarthr Cartil 2023;31:1650–6.
- 8 Heerey J, Agricola R, Smith A, et al. The size and prevalence of bony hip morphology do not differ between football players with and without hip and/or groin pain: findings from the force cohort. J Orthop Sports Phys Ther 2021;51:115–25.
- 9 Agricola R, Waarsing JH, Arden NK, et al. Cam impingement of the hip: a risk factor for hip osteoarthritis. Nat Rev Rheumatol 2013;9:630–4.
- 10 Griffin DR, Dickenson EJ, O'Donnell J, et al. The warwick agreement on femoroacetabular impingement syndrome (FAI syndrome): an international consensus statement. Br J Sports Med 2016;50:1169–76.
- 11 Ganz R, Parvizi J, Beck M, et al. Femoroacetabular impingement: a cause for osteoarthritis of the hip. Clin Orthop Relat Res 2003;112–20.
- 12 Dijkstra HP, Mc Auliffe S, Ardern CL, et al. Oxford consensus on primary cam morphology and femoroacetabular impingement syndrome: part 1-definitions, terminology, taxonomy and imaging outcomes. Br J Sports Med 2022;57:325–41.
- 13 Reiman MP, Agricola R, Kemp JL, et al. Consensus recommendations on the classification, definition and diagnostic criteria of hip-related pain in young and middle-aged active adults from the international hip-related pain research network, Zurich 2018. Br J Sports Med 2020;54:631–41.
- 14 Ishøi L, Nielsen MF, Krommes K, et al. Femoroacetabular impingement syndrome and labral injuries: grading the evidence on diagnosis and non-operative treatment-a statement paper commissioned by the Danish Society of Sports Physical Therapy (DSSF). Br J Sports Med 2021;55:1301–10.
- 15 Palmer AJR, Ayyar Gupta V, Fernquest S, et al. Arthroscopic hip surgery compared with physiotherapy and activity modification for the treatment of symptomatic femoroacetabular impingement: multicentre randomised controlled trial. BMJ 2019;364:1185.
- 16 Griffin DR, Dickenson EJ, Wall PDH, et al. Hip arthroscopy versus best conservative care for the treatment of femoroacetabular impingement syndrome (UK fashion): a multicentre randomised controlled trial. Lancet 2018;391:2225–35.
- 17 Heerey J, Kemp J, Agricola R, et al. Cam morphology is associated with MRIdefined cartilage defects and labral tears: a case-control study of 237 young adult football players with and without hip and groin pain. BMJ Open Sport Exerc Med 2021;7:e001199.
- 18 Tresch F, Dietrich TJ, Pfirrmann CWA, et al. Hip MRI: prevalence of articular cartilage defects and labral tears in asymptomatic volunteers. A comparison with a matched population of patients with femoroacetabular impingement. J Magn Reson Imaging 2017:46:440–51.
- 19 Dijkstra HP, Mc Auliffe S, Ardern CL, et al. Oxford consensus on primary cam morphology and femoroacetabular impingement syndrome: part 2-research priorities on conditions affecting the young person's hip. Br J Sports Med 2022;57:342–58.
- 20 Wesseling J, Dekker J, van den Berg WB, et al. CHECK (cohort hip and cohort knee): similarities and differences with the osteoarthritis initiative. Ann Rheum Dis 2009;68:1413–9.
- 21 Agricola R, Heijboer MP, Bierma-Zeinstra SMA, et al. Cam impingement causes osteoarthritis of the hip: a nationwide prospective cohort study (CHECK). Ann Rheum Dis 2013;72:918–23.
- 22 Bierma-Zeinstra SM, Bohnen AM, Ramlal R, et al. Comparison between two devices for measuring hip joint motions. Clin Rehabil 1998;12:497–505.
- 23 Agricola R, Waarsing JH, Thomas GE, et al. Cam impingement: defining the presence of a cam deformity by the alpha angle: data from the CHECK cohort and Chingford cohort. Osteoarthr Cartil 2014;22:218–25.
- 24 van Klij P, Reiman MP, Waarsing JH, et al. Classifying cam morphology by the alpha angle: a systematic review on thresholdvalues. Orthop J Sports Med 2020;8.
- 25 KELLGREN JH, LAWRENCE JS. Radiological assessment of osteo-arthrosis. Ann Rheum Dis 1957:16:494–502.
- 26 Altman RD, Gold GE. Atlas of individual radiographic features in osteoarthritis, revised. Osteoarthr Cartil 2007;15:A1–56.
- 27 Macri EM, Runhaar J, Damen J, et al. Kellgren/lawrence grading in cohort studies: methodological update and implications illustrated using data from a Dutch hip and knee cohort. Arthritis Care Res (Hoboken) 2022;74:1179–87.
- 28 Damen J, Schiphof D, Wolde ST, et al. Inter-observer reliability for radiographic assessment of early osteoarthritis features: the CHECK (cohort hip and cohort knee) study. Osteoarthr Cartil 2014;22:969–74.
- 29 Casartelli NC, Maffiuletti NA, Valenzuela PL, et al. Is hip morphology a risk factor for developing hip osteoarthritis? A systematic review with meta-analysis. Osteoarthr Cartil 2021;29:1252–64.
- 30 Nelson AE, Stiller JL, Shi XA, et al. Measures of hip morphology are related to development of worsening radiographic hip osteoarthritis over 6 to 13 year follow-up: the iohnston county osteoarthritis project. Osteoarthritis Cartil 2016:24:443–50.
- 31 Nicholls AS, Kiran A, Pollard TCB, et al. The association between hip morphology parameters and nineteen-year risk of end-stage osteoarthritis of the hip: a nested case-control study. Arthritis Rheum 2011;63:3392–400.

- 32 Saberi Hosnijeh F, Zuiderwijk ME, Versteeg M, et al. Cam deformity and acetabular dysplasia as risk factors for hip osteoarthritis. Arthritis Rheumatol 2017;69:86–93.
- 33 Thomas GER, Palmer AJR, Batra RN, et al. Subclinical deformities of the hip are significant predictors of radiographic osteoarthritis and joint replacement in women. A 20 year longitudinal cohort study. Osteoarthr Cartil 2014;22:1504–10.
- 34 Heerey JJ, Srinivasan R, Agricola R, *et al.* Prevalence of early hip OA features on MRI in high-impact athletes. the femoroacetabular impingement and hip osteoarthritis cohort (force) study. *Osteoarthritis Cartil* 2021;29:323–34.
- 35 van Klij P, Heijboer MP, Ginai AZ, *et al.* Cam morphology in young male football players mostly develops before proximal femoral growth plate closure: a prospective study with 5-yearfollow-up. *Br J Sports Med* 2019;53:532–8.
- 36 van Klij P, Heijboer MP, Ginai AZ, et al. Clinical and radiological hip parameters do not precede, but develop simultaneously with cam morphology: a 5-year follow-up study. Knee Surg Sports Traumatol Arthrosc 2021;29:1401–10.
- 37 Pettit M, Doran C, Singh Y, et al. How does the cam morphology develop in athletes? A systematic review and meta-analysis. Osteoarthr Cartil 2021;29:1117–29.
- 38 Heerey JJ, van Klij P, Agricola R, et al. Preventing hip osteoarthritis in athletes: is it really a mission impossible? Br J Sports Med 2024;58:465–7.

- 39 Kemp JL, Risberg MA, Mosler A, et al. Physiotherapist-led treatment for young to middle-aged active adults with hip-related pain: consensus recommendations from the international hip-related pain research network, Zurich 2018. Br J Sports Med 2020;54:504–11.
- 40 Freke MD, Kemp J, Svege I, et al. Physical impairments in symptomatic femoroacetabular impingement: a systematic review of the evidence. Br J Sports Med 2016;50:1180.
- 41 Hatton AL, Kemp JL, Brauer SG, et al. Impairment of dynamic single-leg balance performance in individuals with hip chondropathy. *Arthritis Care Res (Hoboken)* 2014;66:709–16.
- 42 Kemp JL, Risberg MA, Schache AG, et al. Patients with chondrolabral pathology have bilateral functional impairments 12 to 24 months after unilateral hip arthroscopy: a cross-sectional study. J Orthop Sports Phys Ther 2016;46:947–56.
- 43 Kemp JL, Coburn SL, Jones DM, et al. The physiotherapy for femoroacetabular impingement rehabilitation study (physiofirst): a pilot randomized controlled trial. J Orthop Sports Phys Ther 2018;48:307–15.
- 44 Riedstra NS, Boel F, van Buuren M, et al. Pincer morphology is not associated with hip osteoarthritis unless hip pain is present: follow-up data from a prospective cohort study. Arthritis Care Res (Hoboken) 2024;76:644–51.

6 7

8

9

10

Supplemental material

	Incident hi (KL ≥2 or T		Incident end-stage hip OA (KL ≥3 or THR)			
Exposure	Present (n=221)	Absent (n=363)	Absolute risk	Present (n=15)	Absent (n=569)	Absolute risk
Hip pain						
Present (n=547)	264	283	48.3%	46	501	8.4%
Absent (n=839)	350	489		25	814	
Internal hip rotation ≤25°						
Present (n=18)	16	2	88.9%	6	12	33.3%
Absent (n=1367)	597	770		64	1303	33.370
Cam morphology alpha angle >60°						
Present (n=124)	87	37	70.2%	20	104	16.1%
Absent (n=1262)	527	735		51	1211	
Cam morphology alpha angle >78°						
Present (n=67)	47	20	70.1%	15	52	22.4%
Absent (n=1319)	567	752		56	1263	

3 KL = Kellgren-Lawrence grade; THR = total hip replacement

13

19

# 11 Supplementary Table 2: Sensitivity analysis using an alternative definition for hip pain.

	Total N = 567**		Incident hip O	A (KL 2-4 or THR)	Incident end-stage hip OA (KL 3-4 or THR)	
Exposure	N with condition	N without condition <sup>+</sup>	OR (95% CI)	aOR* (95% CI)	OR (95% CI)	aOR* (95% CI)
FAIS (hip pain, internal hip rotation ≤25°, cam morphology with alpha angle >60°)	18 (3.2%)	549 (96.8%)	12.5 (3 – 53)	11.5 (3 – 52)	30.0 (9 – 98)	33.6 (9 – 126)
FAIS (hip pain, internal hip rotation ≤25°, large cam morphology with alpha angle >78°)	13 (2.3%)	549 (96.8%)	8.6 (2 – 38)	7.8 (2 – 37)	37.4 (10 – 137)	52.9 (11 – 265)

- \* Adjusted odds ratios are adjusted for age, sex, and BMI
- \*\* values on hip pain were missing in 17 hips
- 16 \*: The reference group for the predictor categories in this table consists of hips that did not have any of the stated conditions (e.g. the reference group for
- 17 FAI syndrome are hips without cam, without hip pain, and without decreased internal rotation).
- aOR = adjusted odds ratio; KL = Kellgren-Lawrence grade; THR = total hip replacement; FAIS = femoroacetabular impingement syndrome

22

24

2627

28

# Supplementary Table 3: Sensitivity analysis using a threshold of ≤20° instead of ≤25° internal hip rotation to define FAIS

	Total N = 694		Incident hip O	A (KL 2-4 or THR)	Incident end-stage hip OA (KL 3-4 or THR)	
Exposure	N with condition	N without condition <sup>+</sup>	OR (95% CI)	aOR* (95% CI)	OR (95% CI)	aOR* (95% CI)
FAIS (hip pain, internal hip rotation ≤20°, cam morphology with alpha angle >60°)	13 (1.9%)	681 (98.1%)	4.7 (1 – 16)	4.3 (1 – 16)	24.7 (6 – 96)	36.8 (9 – 153)
FAIS (hip pain, internal hip rotation ≤20°, large cam morphology with alpha angle >78°)	9 (1.3%)	681 (98.1%)	5.0 (1 – 20)	4.4 (1 – 20)	44.5 (11 – 187)	133.1 (13 – 1322)

\* Adjusted odds ratios are adjusted for age, sex, and BMI

†: The reference group for the predictor categories in this table consists of hips that did not have any of the stated conditions (e.g. the reference group for

25 FAI syndrome are hips without cam, without hip pain, and with internal hip rotation >20°).

aOR = adjusted odds ratio; KL = Kellgren-Lawrence grade; THR = total hip replacement; FAIS = femoroacetabular impingement syndrome

- Supplementary Table 4: Associations between femoroacetabular impingement syndrome and the development of incident radiographic hip osteoarthritis and incident end-stage radiographic hip osteoarthritis within 10 years follow-up with all hips other than those fulfilling the FAIS criteria as
- 31 the reference group.

	Total n = 1386		Incident hip OA (KL 2-4 or THA) n = 614		Incident end-stage hip OA (KL 3-4 or THR) n=71	
Exposure	N with condition	N without condition <sup>+</sup>	OR (95% CI)	aOR* (95% CI)	OR (95% CI)	aOR* (95% CI)
FAIS (hip pain, internal hip rotation ≤25°, cam morphology with alpha angle >60°)	21 (1.5%)	1365 (98.5%)	3.9 (2 – 10)	3.5 (1 – 10)	8.5 (3 – 25)	8.1 (3 – 25)
FAIS with large cam morphology (hip pain, internal hip rotation ≤25°, large cam morphology with alpha angle >78°)	14 (1.0%)	563 (99.0%)	3.3 (1 – 8)	2.9 (1 – 8)	13.8 (5 – 40)	12.9 (4 – 41)

<sup>\*</sup> Adjusted odds ratios are adjusted for age, sex, and BMI

- <sup>+</sup>: The reference group for the predictor categories in this table consists of hips that did not have all three criteria of FAIS (e.g. the reference group for FAI syndrome could be hips wit cam morphology and hip pain, but without decreased internal rotation).
- aOR = adjusted odds ratio; KL = Kellgren-Lawrence grade; THR = total hip replacement; FAIS = femoroacetabular impingement syndrom