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BMJ Open

Ankle osteoarthritis and its association with severe ankle injuries, ankle surgeries and health-related quality of life in recently retired professional male football and rugby players

Journal:	BMJ Open
Manuscript ID	bmjopen-2020-036775
Article Type:	Original research
Date Submitted by the Author:	02-Jan-2020
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Keywords:	EPIDEMIOLOGY, ORTHOPAEDIC & TRAUMA SURGERY, Foot & ankle < ORTHOPAEDIC & TRAUMA SURGERY, Orthopaedic sports trauma < ORTHOPAEDIC & TRAUMA SURGERY, SPORTS MEDICINE

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3 4	1	Ankle osteoarthritis and its association with severe ankle injuries, ankle surgeries and health-related
5 6	2	quality of life in recently retired professional male football and rugby players
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2			
3 4	24	Abstract	
5 6	25	Objectives	
7 8	26	To determine (i) the prevalence of ankle osteoarthritis (OA) among former professional football and	
9 10 11	27	rugby players, (ii) assess the association between ankle injuries or ankle surgeries with ankle OA, and (ii	i)
12 13	28	compare the mental and physical quality of life (QoL) between former professional football and rugby	
14 15	29	players with and without OA.	
16 17	30		
18 19 20	31	Methods O	
21 22	32	We conducted a questionnaire based observational study with a cross-sectional design. Former	
23 24	33	professional football and rugby players were recruited by the football players worldwide (FIFPRO) and	
25 26 27	34	the International Rugby Players (IRP). Information concerning ankle OA, sustained ankle injuries and	
27 28 29	35	ankle surgeries was gathered (medical record or team doctor). Health related QoL was assessed using	
30 31	36	the PROMIS physical and mental health scores.	
32 33	37		
34 35 26	38	Results	
37 38	39	Overall, 553 former professional football (n = 401) and rugby players (n = 152) were enrolled in the	
39 40	40	study (response rate of 56%). Ankle OA prevalence among former professional football and rugby	
41 42	41	players was 9.2% and 4.6%, respectively. Football players were more likely to suffer from ankle OA	
43 44 45	42	following every ankle injury and/or surgery. Football and rugby players with ankle OA had similar	
43 46 47	43	PROMIS physical and mental health scores to the norm for the general population.	
48 49	44		
50 51	45	Conclusion	
52 53	46	Former professional football and rugby players had a higher ankle OA prevalence than the general	
54 55 56 57	47	population (3.4%). Football players are more likely to suffer from ankle OA following every ankle injury	
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3 4	48	and/or surgery. No clinical relevant difference was seen for physical or mental health related QoL among
5 6	49	football and rugby players. Preventive measures for ankle injuries are recommended.
7 8 9	50 51	Strengths and limitations
10 11 12	52	- Overall good response rate (56%) and a large group of participating football players (401)
13 14 15	53	- This study had a cross-sectional design, therefore no causal-association can be determined
16 17 18 19	54	- There were only 7 rugby players with OA, implying a potential serious type II error.
20 21	55	- Research based (retrospective) questionnaires are dependent on the accuracy and
22 23	56	understanding of the participant. Professional athletes might generally remember the number
24 25	57	of severe ankle injuries or surgeries resulting in a training or match absence of \geq 4 weeks.
26 27 28	58	However, we cannot exclude that participants were not able to precisely recall all their
28 29 30	59	sustained severe ankle injuries and ankle surgeries or that they did not consult their last (team)
31 32 33	60	doctor
34 35 36	61	- Our OA definition did not include a clinical and radiological evaluation.
30 37 38 39 40 41 42 43 44 45 46 47 48 49 50 51 52 53 54 55 56 57	62	
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63	Background
64	Football and Rugby Union (hereafter 'rugby') are among the most popular team sports worldwide. In
65	both sports, ankle injuries represent a significant proportion of the total injuries sustained. ^{1–5} Previous
66	ankle injuries are an important risk factor for ankle osteoarthritis (OA). ^{6,7}
67	
68	Drawer et al. (2002) estimates the overall risk of injury in professional football to be 1,000 times higher
69	than in high-risk industrial occupations, with ankle injuries contributing 10-18% of all injuries. ^{2,8} The
70	Union of European Football Associations (UEFA)Champions League Injury Study reported an overall
71	ankle injury rate of 1 injury per 1000 hours. ² Lateral ankle sprains (51%) were the most prevalent sub-
72	type of ankle injuries. ² In professional rugby, epidemiological studies show that ankle injuries contribute
73	5-20% of all injuries. ^{1,3,9,10} Similar to professional football, lateral ankle sprains are the most common
74	ankle injuries with an injury rate of 3.8 - 4.5 injuries per 1,000 player hours depending on the position on
75	the field. ¹¹
76	
77	Two retrospective cohort studies from the general population (n = 390 and n = 639 patients
78	respectively), reported that 70-78% of ankle OA cases were associated with a previously sustained ankle
79	injury. These previous injuries included fractures, isolated osteochondral defects of the talus and ankle
80	ligament injuries. ^{6,7} Among these post-traumatic ankle OA cases, 28% were associated with a previously
81	sustained ligamentous injury. ⁷ To our knowledge, there are no prospective studies on the association of
82	ankle injuries and ankle OA.
83	
84	In the general population the prevalence of ankle OA is estimated to be 3.4% in people over 50 years of

age. ¹² The reported prevalence of ankle OA 16 to 22 years after a professional football career is 12-86 19%.^{13–15} No data are available for ankle OA among former professional rugby players. In comparison to

a general population control group, the prevalence of OA in any joint and the incidence of joint replacement for any joint was estimated to be 4 and 6 times higher, respectively, among former elite rugby players.¹⁶ This has been associated with a significantly poorer mental and physical health-related quality of life (QoL).¹⁶ Several other studies report the association of general OA with mental health symptoms and physical health-related QoL among former elite athletes across professional team sports including football and rugby.^{17,18} Therefore, the impact of OA on retired players must be considered. To date, the epidemiological evidence concerning the prevalence of ankle OA in professional football is either outdated or limited, and in the case of rugby non-existent, while data on the association between previous ankle injury and ankle OA is scarce in both sports. The objective of this study was threefold. Firstly, to evaluate the prevalence of ankle OA in recently retired professional football and rugby players. Secondly, to determine the association between ankle injury and/or ankle surgery and ankle OA. Thirdly, to compare the mental and physical QoL in players with and without ankle OA. Our hypotheses were that (i) recently retired professional football and rugby players have a higher ankle OA prevalence than the general population (as found in the literature), (ii) ankle injuries and/or ankle surgeries are associated with ankle OA, and (iii) the mental and physical QoL of recently retired professional football and rugby players with OA is lower compared to those players without OA.

2		
2 3 4	109	Methods
5 6	110	Design
7 8	111	We conducted a questionnaire based observational study with a cross-sectional design. Ethical approval
9 10 11	112	for the study was provided by the Ethical Committee of the Yokohama City Sports Medical Center (17.003;
11 12 13	113	Yokohama, Japan) and the Medical Ethics Review Committee of the Academic Medical Center
14 15	114	(W16_366#16.431; Amsterdam, The Netherlands). The study was reported according to the
16 17	115	'Strengthening the Reporting of Observational Studies in Epidemiology' statement and conducted in
18 19	116	accordance with the Declaration of Helsinki (2013). ¹⁹
20 21 22	117	
22 23 24	118	Participants
25 26	119	Retired professional football and rugby players were randomly recruited to reduce recruitment bias, by
27 28	120	the Football Players Worldwide (FIFPRO) and the International Rugby Players (IRP). Inclusion criteria were:
29 30 31	121	(a) being a retired male professional football or rugby player; (b) being younger than 50 years of age; (c)
31 32 33	122	being able to read and comprehend texts in English, French or Spanish.
34 35	123	
36 37	124	The definition of a retired professional football or rugby player was that he (i) has trained to improve
38 39 40	125	performances, (ii) has competed in the highest or second highest national league, and (iii), has had training
40 41 42	126	and competition as a major activity (way of living) or focus of personal interest, devoting several hours in
43 44	127	all or most of the days for these activities, and exceeding the time allocated to other types of professional
45 46	128	or leisure activities. If interested, players received information about the study, gave their informed
47 48 40	129	consent and completed an electronic questionnaire. Once completed, the electronic questionnaires were
49 50 51	130	saved automatically on a secured electronic server that only the principal investigator could access.
52 53	131	
54 55	132	Sample size calculation for ankle OA prevalence indicated that 138 participants per sport were needed
56 57		
58 59 60		6 For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml

(power of 80%, confidence interval of 95%; absolute precision of 5%) under the assumption of an
anticipated population proportion of 10%.²⁰ Expecting a response rate of approximately 50%, we intended
to reach at least 300 participants per sport.

37 Dependent variable: Ankle osteoarthritis

The clinical diagnosis of ankle OA by a medical professional was retrospectively determined using a single question ('Have you been diagnosed with ankle osteoarthritis by a medical professional?'). Participants were given the definition of ankle OA (based on the NICE criteria; adapted for age), determined as the damage of the ankle joint's cartilage that leads to activity-related joint pain with either no morning jointrelated stiffness or morning stiffness that lasts no longer than 30 minutes.²¹ For this question, participants were requested to consult either their medical record or their most recent medical professional.

28 144

145 Independent variable: Severe ankle injury and related surgery

History of a severe ankle injury during their professional football or rugby career was examined through a single question for the sequential recording of single or multiple injury events over time. Similarly, history of an ankle related surgery was recorded through a single question for the sequential recording of a single or multiple ankle surgeries over time. In our study, a severe ankle injury was defined as an injury that involved the ankle joint, occurred during team activities (training or match), and led to absence from either training or a match for more than 28 days ^{18,19}. For this question, participants were requested to consult either their medical records or their most recent medical doctor.

48 153

154 Health-related quality of life

155 The Patient-Reported Outcomes Measurement Information System Global Health short form (PROMIS-156 GH) assesses multiple domains related to health-related quality of life such as health, functioning, pain,

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social activities and fatigue.²⁰ The PROMIS-GH has been validated in several populations and languages among which are English, French and Spanish (for detailed information, see www.nihpromis.org).²⁰ The Global Mental Health and Global Physical Health scores were calculated based on 10 items each measured on a 5-point scale (from 1 to 5) and subsequently converted.²⁰ These subscale scores ranged from 0 to 100, with a higher score indicating better quality of life and a mean score of 50 indicating the norm for the general population.²²

- - Procedures

An anonymous questionnaire (electronic and/or paper) available in English, French and Spanish was compiled (LimeSurvey Professional). It included the following descriptive variables: age, body-height, body-weight, duration of professional football or rugby career, level of play, duration and nature of retirement, and current employment status (employed or not employed). Additionally, this questionnaire included questions for the outcomes previously elaborated on (ankle OA, ankle injury, ankle surgery, QoL). Information about the study was sent per email to potential participants by FIFPRO and IRP. Participants interested in the study, gave their informed consent and completed the electronic questionnaire. Participants were asked to complete the questionnaire within 2 weeks, reminders being sent after 2 and 4 weeks. For privacy reasons, the responses to the questionnaires were coded and anonymized. Once completed, the electronic questionnaires were automatically saved on a secured electronic server that only the principal investigator could access. Players participated voluntarily and did not receive any reward for their participation.

Statistical analyses

Data analysis was performed using the statistical software IBM SPSS 24.0 for Windows. Analyses were conducted separately for retired professional football and rugby players. Data were assessed (visually) for

normality and presented as mean \pm standard deviation (SD), median (interquartile range [IQR]), or frequency (proportion in %) as appropriate. Body mass index (BMI) was calculated using the provided weight and height of participants (kg/m^2) . Prevalence of ankle OA, overall and within the age categories of \leq 40 years and >40 years, was calculated as the proportion of the number of participants with ankle OA relative to the total number of participants in each category.²⁰ A logistic regression analysis was performed to determine the association of the number of severe ankle injuries and/or ankle surgeries (continuous independent variable) with ankle OA (dichotomous dependent variable). Severe ankle injuries and or ankle-related surgeries were also expressed using 3 pre-determined categories (0, 1 and >1). This was adjusted for age and BMI, both having been identified as risk factors for OA.^{23,24} Descriptive analyses of health-related quality of life (Global Physical Health and Global Mental Health) were conducted, while comparisons between groups (retired players with ankle OA vs. retired players without ankle OA) were made using a Mann-Whitney U test for independent samples.²⁰

1 2		
2 3 4	195	Results
5 6	196	Participants
7 8	197	From a total of 750 football and 326 rugby players contacted, respectively 401 and 152 gave their
9 10 11	198	written informed consent and completed the questionnaire (overall response rate of 56%).
12 13	199	Characteristics of the two groups including the pre-determined sub-categories for severe ankle injuries
14 15	200	and ankle surgeries (0, 1 and >1) are illustrated in Table 1.
16 17	201	
18 19 20	202	Prevalence of ankle osteoarthritis, severe injury and surgery
20 21 22	203	The prevalence of patient reported ankle OA (including the pre-determined categories of \leq 40 years
23 24	204	and > 40 years) among recently retired professional football and rugby players, is presented in Table 2.
25 26	205	The prevalence of ankle OA in former professional football players was 9.2% (n = 37). Overall, 94% of
27 28 20	206	players with ankle OA were documented to have sustained ≥1 ankle injury and 62% had undergone ≥1
30 31	207	ankle surgery. In the non-OA group (n = 364), 54% had sustained ≥1 ankle injury and 14% had
32 33	208	undergone ≥1 ankle surgery. Logistic regression analysis (Table 3) showed that football players who
34 35	209	had sustained a severe ankle injury were 1.3 (95%CI: 1.1 - 1.6) times more likely to report ankle OA (p <
36 37 20	210	0.001). Football players who had undergone ankle surgery were 2.1 (95%CI: 1.4 – 3.1) times more likely
39 40	211	to report ankle OA (p< 0.001).
41 42	212	
43 44	213	The prevalence of ankle OA in former professional rugby players was 4.6% (n = 7). Overall, six out of
45 46 47	214	seven (86%) rugby players with ankle OA reported to have sustained ≥1 ankle injury and four out of
47 48 49	215	seven (57.1%) had undergone ≥1 ankle surgery. In the non-OA group (n = 145), 58.2% had sustained ≥1
50 51	216	ankle injury and 18.4% had undergone ≥1 ankle surgery. Logistic regression analysis (Table 3) showed no
52 53	217	significant association between either severe ankle injuries (OR = 1.1 95%CI: 0.8 – 1.5) or ankle surgeries
54 55	218	(OR = 2.0 95%CI: 0.8 – 4.7) and ankle OA among former professional rugby players.
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Table 1: Descriptive characteristics of recently retired professional football players and rugby players

45 46 47

1			Football			Rughy	
2							
3 4 5		Total (n = 401)	Ankle OA (n = 37)	No ankle OA (n = 364)	Total (n = 152)	Ankle OA (n = 7)	No ankle OA (n = 145)
6 7 8	Age in years * (median; IQR; min-max / mean ± SD)	36 (32-40; 25-50)	39 (36-42; 27-49)*	36 (32-40; 25-50)*	40 ± 6	41 ± 7	40 ± 6
9 10 11	Height in cm (mean ± SD / median; IQR; min-max)	181 ± 7	180 ± 7	182 ± 7	186 (180-191; 166-203)	191 (173-192; 172-192)	186 (180-191; 166-203)
12 13 14	Weight in kg (in kg (mean ± SD))	82 ± 9	82 ± 13	82 ± 9	100 (91-114; 59-170)	108 (89-123; 82-130)	100 (91-113; 59-170)
15 16 17	BMI in kg/m² (median; IQR; min-max))	24.8 (23.5-26.2; 20.1-32.7)	24.8 (22.9-26.6; 20.3-29)	24.7 (23.5-26.1; 20.1-32.7)	29.0 (27.1-31.4; 20.4-52.5)	32.2 (28.5-33.7; 27.4-35.3)*	28.9 (27.0-31.2; 20.4-52.5)*
18 19 20	Employed in % (n)	89% (350)	92% (33)	89% (317)	91% (139)	86% (6)	92% (133)
21 22 23	Professional career duration in years (median; IQR; min- max)	13 (10-15; 2-19)	13 (12-16; 5-18)	13 (9-15; 2-19)	10 (7-15; 2-20)	10 (8-12.5; 8-13)	10 (7-15; 2-20)
24 25 26	Retirement duration in years (median; IQR; min-max)	4 (2-7; 1-17)	5 (3-9; 1-12)	4 (2-7; 1-17)	8 (3-12; 0-22)	12 (2-16; 1-19)	8 (4-11; 0-22)
27 28	Level of play top league; % (n)	81% (314)	92% (34)	80% (280)	80% (106)	86% (6)	80% (100)
29 30 31	Forced retirement % (n)	29% (113)	27% (10)	30% (103)	36% (55)	57% (4)	35% (51)
32 33 34	Severe ankle injuries (median; IQR; min-max)	1 (0-2; 0-10)	2 (1-3; 0-10)	1 (0-2; 0-10)	1 (0-2; 0-20)	1 (1-2; 0-3)	1 (0-2; 0-20)
35 36 37	Ankle surgeries (median; IQR; min-max)	0 (0-0; 0-5)	1 (0-2; 0-5)	0 (0-0; 0-5)	0 (0-0; 0-4)	1 (0-1; 0-2)	0 (0-0; 0-4)
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* Statistically significant d	interence between ankie OA and no ankie OA. Abbreviations: OA = Osteoarthritis; IQK = Interquartile range; n = number	
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		Football			Rugby	
	Total (n = 401)*	≤ 40 years (n = 271)	> 40 years (n = 83)	Total (n = 152)	≤ 40 years (n = 82)	> 40 years (n = 66)
Prevalence ankle OA in % (95%Cl)	9.2% (6.4 –12.1)	8.5% (5.2 – 11.8)	12.0% (5.0 – 19.1)	4.6% (1.2 – 8.0)	2.4% (0.9 – 5.8)	4.5% (0.5 – 9.6)
	0 ankle injuries	1 ankle injury	> 1 ankle injury	0 ankle injuries	1 ankle injury	> 1 ankle injury
Severe ankle injuries total (%, (n))	42.5% (156)	25.3% (93)	32.2% (118)	40.5% (60)	27.7% (41)	31.8% (47)
- With ankle OA (%, (n))	5.7% (2)	28.6% (10)	65.7% (23)	14.3% (1)	42.9% (3)	42.9% (3)
- Without ankle OA (%, (n))	46.4% (154)	25.0% (83)	28.6% (95)	41.8% (59)	27.0% (38)	31.2% (44)
	No ankle surgeries	1 ankle surgery	> 1 ankle surgery	No ankle surgeries	1 ankle surgery	> 1 ankle surgery
Ankle surgeries total in % (n)	81.5% (299)	10.9% (40)	7.6% (28)	79.7% (118)	15.5% (23)	4.7% (7)
- With ankle OA % (n)	38.2% (13)	29.4% (10)	32.4% (11)	42.9% (3)	42.9% (3)	14.3% (1)
- Without ankle OA % (n)	85.9% (286)	9% (30)	5.1% (17)	81.6% (115)	14.2% (20)	4.3% (6)

Table 2: Subcategories for prevalence of reported ankle OA, severe injuries and ankle surgeries in retired football and rugby players

 *The sum of patients in the subcategories \leq 40 years and > 40 years does not equal the "Total number of participants" due to missing data (regarding age). Abbreviations: OA = Osteoarthritis; n = number; CI = confidence interval;

1							
2 3		Table 2: Accessiation (a	dds ratio and OE% CI) a	of couere ankle iniuru a	nd ankla surgary with a	nkla ostooarthritis	
4		amona recently retired	professional football d	and ruaby players	na ankie surgery with a	TIKIE OSLEOUT LITTLIS	
5			Footbal	l (n = 401)	Rugby (by (n = 152)	
6			Unadjusted	Adjusted	Unadjusted	Adjusted	
7 8		Severe ankle injury (OR, 95%Cl)	1.34 (1.17-1.53)	1.33 (1.14-1.55)	1.01 (0.74-1.38)	1.05 (0.75-1.47)	
9 10		Ankle surgery (OR, 95%Cl)	2.20 (1.61-3.00)	2.09 (1.41-3.08)	1.99 (0.92-4.3)	1.96 (0.81-4.71)	
12 13 14 15 16 17 18 19 20 21 22 23 24 25 26 27 28 29 30 31 32 33 4 35 36 37 38 39 40 41 42	219	interval					
43 44 45 46			For p	eer review only - http:	//bmjopen.bmj.com/si	te/about/guidelines.	

Mental and physical QoL

For the PROMIS physical health score a statistically significant lower outcome was found for football (p = 0.01) and rugby players (p = 0.009) with

OA (48 and 45 points respectively) compared to players without OA (51 and 51 points respectively). The PROMIS mental health score was not

significantly different between players with OA vs players without OA. Data is presented in Table 4.

Table 4: Mental and physical quality of life (0-100) of recently retired football and rugby players with ankle OA compared to players without ankle OA

	Football			Rugby				
	Total (n = 401)	With ankle OA (n = 37)	Without ankle OA (n = 364)	p-value	Total (n = 152)	With ankle OA (n = 7)	Without ankle OA (n = 145)	p-value
PROMIS mental health (median; IQR; min-max)	51; 46-56; 31-68	51; 46-53; 31-63	53; 46-56; 31-68	0.251	52; 46-56; 28-68	48; 48-53; 46-56	53; 46-56; 28-68	0.567
PROMIS Physical health* (median; IQR; min-max)	51; 46-56; 27-68	48; 42-54; 35-62*	51; 48-58; 27-68*	0.010	51; 45-54; 30-68	45; 37-48; 37-51*	51; 37-56; 30-68*	0.009

*Statistically significant difference; Abbreviations: OA = Osteoarthritis; IQR = Interguartile range; n = number

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Table 5: Post-Hoc analysis regarding mental and physical quality of life of recently retired football and rugby players who reported to have been forced to retire, and quality of life of players who reported to have been forced to retire with ankle OA compared to players without ankle OA

	PROMIS mental health (median; IQR; min-max)	p-value	PROMIS physical health (median; IQR; min-max)	p-value
Total				
Voluntarily retired* (n = 369)	53; 48-56; 31-68*	n < 0.001	54; 48-58; 30-68*	n < 0.001
Forced to retire* (n = 168)	48; 44-53; 28-68*	ρ< 0.001	48; 42-51; 27-68*	μ< 0.001
Voluntarily retired with OA* (n = 30)	51; 46-53; 36-63		51; 45-54; 35-62*	
Voluntarily retired without OA* (n = 339)	53; 48-59; 31-68	p = 0.074	54; 48-58; 30-68*	p = 0.001
Forced to retire with OA (n = 14)	48; 46-55; 31-63	p = 0.841	42; 40-46; 37-58	p = 0.159

Post-hoc analysis

In the literature, forced retirement is a risk factor for post-career mental health symptoms.²⁵ Following data analysis, we were concerned by the seemingly high percentage of football and rugby players who stopped due to a career ending injury. As we were analysing the impact ankle OA had on QoL, we felt the effect of a career ending injury on QoL to be in line with our initial objectives. Consequently we performed a post-hoc analysis to determine any associations between reported forced retirement and mental and physical PROMIS health scores. In this study, 31% of players reported to have been forced to retire, specifically: 29% of football players and 36% of rugby players.

voluntary retirement and in conjunction with ankle OA is presented in Table 5. No post-hoc sub-analysis was done for football and rugby players separately, regarding voluntary retirement in conjunction with ankle OA, due to the low number of ankle OA cases among rugby players. However mental and physical health scores in football and rugby due to forced retirement is presented separately in Table 5.

The post-hoc sub-analysis of all players regarding mental and physical health scores associated with

2						
3		Forced to retire	48; 44-53; 28-68		48; 42-52; 27-68	
4		without OA (n = 154)				
5						
0 7		Football*	1			
, 8		Voluntarily retired*	53; 48-56; 31-68*		54; 48-58; 30-68*	
9		(n = 272)				
10		Forced to retire*	48; 44-53; 31-63*	p < 0.001	48; 42-51; 27-62*	p < 0.001
11		(n = 113)				
12		Du~bu*				
13						
14		voluntarily retirea*	53; 48-56; 31-68*		54; 48-58; 35-68*	
15		(11 = 97) Forced to retire*	18: 11-56: 28-68*	p = 0.018	18:37-51:30-68*	p < 0.001
10 17		(n = 55)	48, 44-30, 28-08		48, 37-34, 30-08	
17		*Statistically significant diff	erence: Abbreviations: IOR = In	terauartile ranae	: OA = Osteoarthritis: n = nun	nber
19	246	Statistically significant all		terquartie range,		
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2 3	248	Discussion
4 5	249	Our most important findings were that (i) the prevalence of ankle OA among former professional
6 7	250	football and rugby players was, 9.2% (95%CI 6.4 –12.1) and 4.6% (95%Cl (1.2 – 8.0), respectively; (ii)
8 9 10	251	previous severe ankle injuries and ankle surgeries are associated with ankle OA among former
10 11 12	252	professional football players; and (iii) ankle OA does not lead to reduced mental or physical QoL in
12 13 14	253	former professional football and rugby players.
15 16	254	
17 18	255	Ankle OA in football and rugby compared to other sports and the general population
19 20	256	Several studies have reported symptomatic ankle OA prevalence among a variety of former elite
21 22 22	257	athletes, ranging from 0% in volleyball (n = 22), long distance running (n = 30), elite high-jump (n = 30)
23 24 25	258	and ballet (n = 27) to 10% among former military parachutists (n = 40). $^{26-30}$ In our study, the prevalence
25 26 27	259	of ankle OA among former professional football (9.2%) and rugby players (4.6%) was higher than the
28 29	260	estimated prevalence among the general population (3.4 %). ¹² Murray et al. (2018) invited participants
30 31	261	with self-reported ankle pain (11.7%) from the general population for a radiograph (radiological atlas of
32 33	262	Foot OA). ^{12,31}
35 36	263	
37		
38 39	264	Our study reports a lower ankle OA prevalence compared to previous studies among former
40 41	265	professional football players (12-19%). ¹³⁻¹⁵ In these studies, data was similarly collected using cross-
42 43	266	sectional retrospective self-reported questionnaires. ^{13,14} No previous data is available concerning ankle
44 45 46	267	OA among former professional rugby players. The low ankle OA prevalence among rugby players in this
40 47 48	268	study is likely due to either an under reporting of ankle OA among rugby players or a type II error.
49 50	269	
51 52	270	Association of severe ankle injuries and ankle surgeries with ankle OA
53 54	271	In our study, football players were 1.3 (95%CI: 1.1 - 1.6) and 2.1 (95%CI: 1.4 – 3.1) times more likely to
55 56	272	develop ankle OA for every ankle injury and ankle surgery, respectively. This is in line with previous
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> studies that showed an association between ankle OA and ankle injuries, stating traumatic ankle injuries to be the main cause of ankle OA (70-78%).^{6,7} Consequences of ankle injuries are thought to be due to either direct damage the articular surfaces or change in ankle joint biomechanics (e.g. through chronic ankle instability due to ankle ligament damage).³² No association was seen for severe ankle injuries or ankle surgeries and ankle OA among rugby players. Although the best estimate for the association of ankle surgeries and OA is similar to football (OR = 2), the 95%CI includes 1 (0.8 – 4.7). This may be due to insufficient power among rugby players, as only 7 reported ankle OA.

281 Health-related quality of life

Filbay et al. (2019) reported factors such as age, type of sport (contact/collision sports), OA and involuntary retirement from sport to possibly negatively impact QoL.^{25,33–35} However the positive impacts associated with elite sport participation, such as pride of accomplishments, social network (through sport participation) and coping and adjustment to musculoskeletal pain (due to resilience and determination), may persist and even compensate reduced physical QOL beyond retirement of the player's athletic career.³³

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Our post-hoc sub-analysis revealed 29% of football players and 36% of rugby players were forced to retire. Although we reported several instances of statistically significant differences in our study for both physical and mental PROMIS health scores, all were similar to the estimated norm for the general population (50 points). Unfortunately, there is no robust minimally clinically important difference (MCID) threshold for the PROMIS-10 global short form applicable for this population. Consequently, it is difficult to interpret the results. Nevertheless, although statistically significant, the PROMIS mental and physical health scores observed, for both football and rugby players, are unlikely clinically significant.

1 2		
2 3 4	296	Based on our data, ankle OA does not lead to reduced mental or physical QoL in former professional
5 6	297	football and rugby players.
7 8	298	
9 10 11	299	Strengths and limitations
12 13	300	We had an overall good response rate (56%) and a large group of participating football players (401).
14 15	301	However, several limitations should be mentioned. Firstly, this study had a cross-sectional design,
16 17	302	therefore no causal-association can be determined. Secondly, there were only 7 rugby players with OA,
18 19 20	303	implying a potential serious type II error. Thirdly, recruitment procedures were blinded to the research
21 22	304	team, a non-response analysis could not be performed. Fourthly, research based (retrospective)
23 24	305	questionnaires are dependent on the accuracy and understanding of the participant. Professional
25 26	306	athletes might generally remember the number of severe ankle injuries or surgeries resulting in a
27 28 29	307	training or match absence of ≥4 weeks. However, we cannot exclude that participants were not able to
30 31	308	precisely recall all their sustained severe ankle injuries and ankle surgeries or that they did not consult
32 33	309	their last (team) doctor. Finally, our OA definition did not include a clinical and radiological evaluation.
34 35	310	
36 37 38	311	Implications for practice
39 40	312	The epidemiological data from this study regarding ankle OA prevalence among former professional
41 42	313	football and rugby players, and its association with both severe ankle injuries and ankle surgeries, is an
43 44	314	essential part of protecting the athlete's health. In both professional football and rugby, a higher
45 46 47	315	prevalence of ankle OA can be expected compared to the general population. Furthermore, As the
47 48 49	316	majority are post-traumatic, preventive measures and adequate rehabilitation are recommended.
50 51	317	Awareness should be raised, regarding the increased risk of developing ankle OA following each ankle
52 53	318	injury and surgery. Additionally primary and secondary prevention programmes should be implemented.
54 55	319	Exercise based prevention intervention programs, such as the FIFA11+ in football and the movement
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2 3 4	320	control injury prevention programme in rugby, have found reduced ankle injury incidence and									
5 6	321	burden. ^{36,37} Finally, management of ankle OA is key for retired players, focusing on healthy lifestyle									
7 8	322	including physical activity, pain alleviation, function and minimising disability. ^{38–40} With this intention,									
9 10	323	FIFPRO has developed an "After Career Consultation". ⁴¹ Future studies will evaluate its relevance.									
11 12 12	324										
15 14 15	325										
16 17		New findings and impact on clinical practice in the future									
18 19 20		• The prevalence of ankle OA in both professional football (9.2%) and rugby (4.6%), is higher									
21 22		than the estimated prevalence in the general population (3.4%).									
23 24 25		Awareness should be raised among football and rugby players, regarding the increased risk of									
26 27 28		developing ankle OA following each ankle injury and ankle surgery.									
29 30		Primary and secondary prevention programmes should be implemented, as these might help									
32 33		in reducing the risk of developing ankle OA.									
34 35 36		• Management of ankle OA is key for retired players. With this intention, FIFPRO has developed									
37 38 39		an "After Career Consultation". Future studies will evaluate its relevance.									
40 41 42	326 327										
43 44	328	Conclusion									
45 46 47	329	This cross-sectional study found an athlete reported ankle OA prevalence of 9.2% and 4.6% among									
48 49	330	recently retired professional football and rugby players, respectively. Severe ankle injuries and ankle									
50 51	331	surgeries were associated with a higher prevalence of ankle OA in former professional football players.									
52 53	332	No clinically relevant differences in QoL were reported in players with OA compared to players without									
54 55 56	333	OA. As a higher prevalence of ankle OA can be expected in both professional football and rugby									
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3 4	334	compared to the general population, and the majority are post-traumatic, primary and secondary
5 6	335	preventive measures for ankle injuries and an optimal diagnosis and treatment plan are recommended.
/ 8 9	336	
10 11	337	Acknowledgements
12 13	338	The Authors would like to thank all football and ruby players' associations involved. The authors are
14 15 16	339	grateful to all retired professional football and rugby players who participated in the study.
10 17 18	340	
19 20	341	Contributorship statement
21 22	342	Authorship was determined according to the ICMJE authorship recommendations. All authors were
23 24 25	343	involved in the design of the study, data analysis and data interpretation. VG and GMMJK were
25 26 27	344	responsible for data collection. LP drafted the manuscript, with critical review provided by all authors.
28 29	345	All authors approved the final version of the manuscript.
30 31	346	
32 33	347	Funding
34 35 36	348	This research received specific grant from any funding agency in the public, commercial or non-profit
37 38	349	sectors.
39 40	350	
41 42	351	Competing interests
43 44 45	352	None declared.
45 46 47	353	
48 49	354	Ethics approval
50 51	355	Ethical approval for the study was provided by the Ethical Committee of the Yokohama City Sports
52 53	356	Medical Center (17.003; Yokohama, Japan) and the Medical Ethics Review Committee of the Academic
54 55 56 57	357	Medical Center (W16_366#16.431; Amsterdam, The Netherlands)
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2 3 4	358	
5 6	359	Patient and public involvement
7 8	360	This research was done without patient involvement. Patients were not invited to comment on the
9 10 11	361	study design and were not consulted to develop patient relevant outcomes or interpret the results.
12 13	362	Patients were not invited to contribute to the writing or editing of this document for readability or
14 15	363	accuracy.
16 17	364	
18 19 20	365	
20 21 22	366	Data sharing statement
23 24	367	No additional data available
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STROBE Statement—Checklist of items that should be included in reports of cross-sectional studies

	Item No	Recommendation
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract $-$
		(b) Provide in the abstract an informative and balanced summary of what was done and what was found – $$
Introduction		
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported – $\sqrt{2}$ p. 4 line 64 – 96
Objectives	3	State specific objectives, including any prespecified hypotheses $-\sqrt{2}$ p. 4 line 98 $-$ 104
Methods		
Study design	4	Present key elements of study design early in the paper – $\sqrt{2}$ p. 6 line 108 – 114
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment, exposure, follow-up, and data collection $-\sqrt{2}$; p. 6 line 116 – 133
Participants	6	(a) Give the eligibility criteria, and the sources and methods of selection of participants $-\sqrt{2}$; p. 6 line 116 – 133
Variables	7	Clearly define all outcomes, exposures, predictors, potential confounders, and effect modifiers. Give diagnostic criteria, if applicable $-\sqrt{2}$ p. 6 line 135 – 159
Data sources/	8*	For each variable of interest, give sources of data and details of methods of
measurement		assessment (measurement). Describe comparability of assessment methods if there is more than one group $-\sqrt{2}$ p. 6 line $161 - 172$
Bias	9	Describe any efforts to address potential sources of bias $-\sqrt{2}$ p. 6 line 117
Study size	10	Explain how the study size was arrived at $-\sqrt{2}$; p. 6 – 7 line 130 – 133
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If applicable, describe which groupings were chosen and why $-\sqrt{2}$ p. 8-9 line 175 – 188
Statistical methods	12	(a) Describe all statistical methods, including those used to control for confounding
		(b) Describe any methods used to examine subgroups and interactions
		(c) Explain how missing data were addressed
		(d) If applicable, describe analytical methods taking account of sampling strategy
		(e) Describe any sensitivity analyses
		– √: p. 8-9 line 175 – 188
Results		
Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially
		eligible, examined for eligibility, confirmed eligible, included in the study,
		completing follow-up, and analysed – $\sqrt{:}$ p. 10 line 192 – 196
		(b) Give reasons for non-participation at each stage – $\sqrt{:}$ p. 10 line 192 – 196
		(c) Consider use of a flow diagram
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and
		information on exposures and potential confounders – $\sqrt{2}$ p 11 table 1
		(b) Indicate number of participants with missing data for each variable of interest
Outcome data	15*	Report numbers of outcome events or summary measures – $\sqrt{2}$ p 11 – 14, table 2 -4
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and
		their precision (eg, 95% confidence interval). Make clear which confounders were
		adjusted for and why they were included – $\sqrt{\mathbf{p} 11 - 14, \mathbf{table 2 - 4}}$
		(b) Report category boundaries when continuous variables were categorized

		(<i>c</i>) If relevant, consider translating estimates of relative risk into absolute risk for a meaningful time period
Other analyses	17	Report other analyses done—eg analyses of subgroups and interactions, and sensitivity analyses $-\sqrt{2}$ n 15 line 227 -237 + n 16 table 5
Discussion		Sensitivity analyses $= \langle \cdot, p \rangle$ is, fine $227 = 257 + p$ to table 5
Key results	18	Summarise key results with reference to study objectives – $\sqrt{: p \ 17, line \ 242 - 245}$
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or imprecision. Discuss both direction and magnitude of any potential bias – $\sqrt{:}$ p 3, line 53 - 62
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations, multiplicity of analyses, results from similar studies, and other relevant evidence – $\sqrt{:}$ p 19 - 20, line 293 - 317
Generalisability	21	Discuss the generalisability (external validity) of the study results p 19 - 20 , line 293 - 305
Other information		
Funding	22	Give the source of funding and the role of the funders for the present study and, if applicable, for the original study on which the present article is based p 21, line 339 - 340

*Give information separately for exposed and unexposed groups.

Note: An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.

BMJ Open

Ankle osteoarthritis and its association with severe ankle injuries, ankle surgeries and health-related quality of life in recently retired professional male football and rugby players: A Cross-sectional observational study

Journal:	BMJ Open
Manuscript ID	bmjopen-2020-036775.R1
Article Type:	Original research
Date Submitted by the Author:	26-Mar-2020
Complete List of Authors:	Paget, Liam; Amsterdam UMC - Locatie AMC, Orthopaedic surgery; Amsterdam Collaboration for Health and Safety in Sports, AMC/VUMC IOC Research Center (ACHSS) Aoki, Haruhito; St. Marianna University School of Medicine; Yokohama City Sports Medical Center Kemp, Simon; Rugby Football Union, Sports Medicine Lambert, Mike; University of Cape Town, UCT/MRC Research Unit for Exercise Science and Sports Medicine Readhead, Clint; South Africa Rugby Union, Medical; University of Cape Town Division of Exercise Science and Sports Medicine Stokes, Keith; University of Bath, Department for Health; Rugby Football Union, Viljoen, Wayne; South African Rugby Union, Medical; University of Cape Town Division of Exercise Science and Sports Medicine Reurink, G; Sports Medicine, OLVG, The Sport Physician Group; Academic Center for Evidence-based Sports medicine (ACES) Tol, J; Academic Center for Evidence-Based Sports medicine (ACES); Aspetar Orthopaedic and Sports Medicine Hospital Kerkhoffs, Gino; Amsterdam UMC - Locatie AMC, Orthopaedic surgery; Amsterdam Collaboration for Health and Safety in Sports (ACHSS) Gouttebarge, Vincent; Amsterdam UMC - Locatie AMC, Orthopaedic surgery; Football Players Worldwide (FIFPRO)
Primary Subject Heading :	Sports and exercise medicine
Secondary Subject Heading:	Rehabilitation medicine, Sports and exercise medicine
Keywords:	EPIDEMIOLOGY, ORTHOPAEDIC & TRAUMA SURGERY, Foot & ankle < ORTHOPAEDIC & TRAUMA SURGERY, Orthopaedic sports trauma < ORTHOPAEDIC & TRAUMA SURGERY, SPORTS MEDICINE

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review on

1		
2 3 4	1	Ankle osteoarthritis and its association with severe ankle injuries, ankle surgeries and health-related
5 6	2	quality of life in recently retired professional male football and rugby players: A Cross-sectional
7 8	3	observational study
9 10 11	4	Liam D. A. Paget ^{1,2,3} , Haruhito Aoki ^{4, 5} , Simon Kemp ⁶ , Mike Lambert ⁷ , Clint Readhead ^{7, 8} , Keith Stokes ^{6, 9, 10} ,
12 13	5	Wayne Viljoen ^{7, 8} , Gustaaf Reurink ^{1,2,3,11} , Johannes L. Tol ^{2,3,12} , Gino M. M. J. Kerkhoffs ^{1,2,3} , Vincent
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49 50	23	
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59 60		For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml

2 3	28	Abstract	
4 5	20		
6 7	29	Objectives	
, 8 9	30	To determine (i) the prevalence of ankle osteoarthritis (OA) among former professional football and	
10 11	31	rugby players, (ii) assess the association between ankle injuries or ankle surgeries with ankle OA, and (ii	i)
12 13	32	compare the mental and physical quality of life (QoL) between former professional football and rugby	
14 15	33	players with and without OA.	
16 17	34		
18 19 20	35	Methods	
20 21 22	36	We conducted a questionnaire based observational study with a cross-sectional design. Former	
23 24	37	professional football and rugby players were recruited by the football players worldwide (FIFPRO) and	
25 26	38	the International Rugby Players (IRP). Information concerning ankle OA, sustained ankle injuries and	
27 28 20	39	ankle surgeries was gathered (medical record or most recent medical professional). Health related	
29 30 31	40	QoL was assessed using the PROMIS physical and mental health scores.	
32 33	41		
34 35	42	Results	
36 37	43	Overall, 553 former professional football (n = 401) and rugby players (n = 152) were enrolled in the	
38 39 40	44	study (response rate of 56%). Ankle OA prevalence among former professional football and rugby	
41 42	45	players was 9.2% and 4.6%, respectively. Football players were more likely to suffer from ankle OA	
43 44	46	following every ankle injury and/or surgery. Football and rugby players with ankle OA had similar	
45 46	47	PROMIS physical and mental health scores to the norm for the general population.	
47 48 40	48		
50 51	49	Conclusion	
52 53	50	Former professional football and rugby players had a higher ankle OA prevalence than the general	
54 55	51	population (3.4%). Football players are more likely to suffer from ankle OA following every ankle injury	
56 57			
58 59 60		For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml	2

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2 3 4	52	and/or surgery. No clinical relevant difference was seen for physical or mental health related QoL among
5 6	53	football and rugby players. Preventive measures for ankle injuries are recommended.
7 8	54	
9 10 11	55	Strengths and limitations
12 13 14	56	- Overall good response rate (56%) and a large group of participating football players (401)
15 16 17	57	- This study had a cross-sectional design, therefore no causal-association can be determined
17 18 19	58	- This study contributes to the literature, where the current available data concerning ankle OA in
20 21 22	59	professional football and rugby players is outdated or non-existent.
23 24	60	- The data is based on (retrospective) questionnaires and is therefore dependent on the accuracy
25 26 27	61	and understanding of the participant. Our OA definition did not include a clinical and
27 28 29	62	radiological evaluation.
31 32 33 34 35 36 37 38 39 40 41 42 43 44 45 46 47 48 49 50 51 52 53 54 55	63	
56 57 58 59 60		3 For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml

Football and Rugby Union (hereafter 'rugby') are among the most popular team sports worldwide. In both sports, ankle injuries represent a significant proportion of the total injuries sustained.^{1–5} Previous ankle injuries are an important risk factor for ankle osteoarthritis (OA).^{6,7}

Drawer et al. (2002) estimates the overall risk of injury in professional football to be 1,000 times higher than in high-risk industrial occupations, with ankle injuries contributing 10-18% of all injuries.^{2,8} The Union of European Football Associations (UEFA)Champions League Injury Study reported an overall ankle injury rate of 1 injury per 1000 hours.² Lateral ankle sprains (51%) were the most prevalent sub-type of ankle injuries.² In professional rugby, epidemiological studies show that ankle injuries contribute 5-20% of all injuries.^{1,3,9,10} Similar to professional football, lateral ankle sprains are the most common ankle injuries with an injury rate of 3.8 - 4.5 injuries per 1,000 player hours depending on the position on the field.¹¹

Two retrospective cohort studies from the general population (n = 390 and n = 639 patients respectively), reported that 70-78% of ankle OA cases were associated with a previously sustained ankle injury. These previous injuries included fractures, isolated osteochondral defects of the talus and ankle ligament injuries.^{6,7} Among these post-traumatic ankle OA cases, 28% were associated with a previously sustained ligamentous injury.⁷ To our knowledge, there are no prospective studies on the association of

- ankle injuries and ankle OA.

In the general population the prevalence of ankle OA is estimated to be 3.4% in people over 50 years of age. ¹² The reported prevalence of ankle OA 16 to 22 years after a professional football career is 12-19%.^{13–15} No data are available for ankle OA among former professional rugby players. In comparison to

3 4	88	a general population control group, the prevalence of OA in any joint and the incidence of joint
5 6	89	replacement for any joint was estimated to be 4 and 6 times higher, respectively, among former elite
7 8	90	rugby players. ¹⁶ This has been associated with a significantly poorer mental and physical health-related
9 10	91	quality of life (QoL). ¹⁶
11 12 13	92	
14 15	93	Several other studies report the association of general OA with mental health symptoms and physical
16 17	94	health-related QoL among former elite athletes across professional team sports including football and
18 19 20	95	rugby. ^{17,18} Therefore, the impact of OA on retired players must be considered. To date, the
20 21 22	96	epidemiological evidence concerning the prevalence of ankle OA in professional football is either
23 24	97	outdated or limited, and in the case of rugby non-existent, while data on the association between
25 26	98	previous ankle injury and ankle OA is scarce in both sports.
27 28	99	
29 30 31	100	The objective of this study was threefold. Firstly, to evaluate the prevalence of ankle OA in recently
32 33	101	retired professional football and rugby players. Secondly, to determine the association between ankle
34 35	102	injury and/or ankle surgery and ankle OA. Thirdly, to compare the mental and physical QoL in players
36 37 20	103	with and without ankle OA. Our hypotheses were that (i) recently retired professional football and rugby
30 39 40	104	players have a higher ankle OA prevalence than the general population (as found in the literature), (ii)
41 42	105	ankle injuries and/or ankle surgeries are associated with ankle OA, and (iii) the mental and physical QoL
43 44	106	of recently retired professional football and rugby players with OA is lower compared to those players
45 46	107	without OA.
47 48	108	
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59 60		For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml

1 2		
- 3 4	110	Methods
5 6	111	Design
7 8	112	We conducted a questionnaire based observational study with a cross-sectional design. Ethical approval
9 10 11	113	for the study was provided by the Ethical Committee of the Yokohama City Sports Medical Center (17.003;
12 13	114	Yokohama, Japan) and the Medical Ethics Review Committee of the Academic Medical Center
14 15	115	(W16_366#16.431; Amsterdam, The Netherlands). The study was reported according to the
16 17	116	'Strengthening the Reporting of Observational Studies in Epidemiology' statement and conducted in
18 19 20	117	accordance with the Declaration of Helsinki (2013). ¹⁹
20 21 22	118	
23 24	119	Participants
25 26	120	Retired professional football and rugby players were randomly recruited to reduce recruitment bias, by
27 28 20	121	the Football Players Worldwide (FIFPRO) and the International Rugby Players (IRP). Inclusion criteria were:
29 30 31	122	(a) being a retired male professional football or rugby player; (b) being younger than 50 years of age; (c)
32 33	123	being able to read and comprehend texts in English, French or Spanish.
34 35	124	
36 37	125	The definition of a retired professional football or rugby player was that he (i) has trained to improve
38 39 40	126	performances, (ii) has competed in the highest or second highest national league, and (iii), has had training
40 41 42	127	and competition as a major activity (way of living) or focus of personal interest, devoting several hours in
43 44	128	all or most of the days for these activities, and exceeding the time allocated to other types of professional
45 46	129	or leisure activities. If interested, players received information about the study, gave their informed
47 48 49	130	consent and completed an electronic questionnaire. Once completed, the electronic questionnaires were
50 51	131	saved automatically on a secured electronic server that only the principal investigator could access.
52 53	132	
54 55	133	Sample size calculation for ankle OA prevalence indicated that 138 participants per sport were needed
56 57 58		
59 60		6 For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml

1 2		
- 3 4	134	(power of 80%, confidence interval of 95%; absolute precision of 5%) under the assumption of an
5 6	135	anticipated population proportion of 10%. ²⁰ Expecting a response rate of approximately 50%, we intended
7 8	136	to reach at least 300 participants per sport.
9 10 11	137	
12 13	138	Dependent variable: Ankle osteoarthritis
14 15	139	The clinical diagnosis of ankle OA by a medical professional was retrospectively determined using a single
16 17	140	question ('Have you been diagnosed with ankle osteoarthritis by a medical professional?'). Participants
18 19 20	141	were given the definition of ankle OA (based on the NICE criteria; adapted for age), determined as the
20 21 22	142	damage of the ankle joint's cartilage that leads to activity-related joint pain with either no morning joint-
23 24	143	related stiffness or morning stiffness that lasts no longer than 30 minutes. ²¹ For this question, participants
25 26	144	were requested to consult either their medical record or their most recent medical professional.
27 28 29 30 31 32 33	145	
	146	Independent variable: Severe ankle injury and related surgery
	147	History of a severe ankle injury during their professional football or rugby career was examined through
34 35	148	a single question for the sequential recording of single or multiple injury events over time. Similarly,
36 37	149	history of an ankle related surgery was recorded through a single question for the sequential recording of
38 39 40	150	a single or multiple ankle surgeries over time. In our study, a severe ankle injury was defined as an injury
40 41 42	151	that involved the ankle joint, occurred during team activities (training or match), and led to absence from
43 44	152	either training or a match for more than 28 days ^{18,19} . For this question, participants were requested to
45 46	153	consult either their medical records or their most recent medical professional.
47 48 40	154	
49 50 51	155	Health-related quality of life
52 53	156	The Patient-Reported Outcomes Measurement Information System Global Health short form (PROMIS-
54 55	157	GH) assesses multiple domains related to health-related quality of life such as health, functioning, pain,
56 57 50		
59 60		7 For peer review only - http://bmjopen.bmj.com/site/about/guidelines.xhtml

social activities and fatigue.²⁰ The PROMIS-GH has been validated in several populations and languages among which are English, French and Spanish (for detailed information, see www.nihpromis.org).²⁰ The Global Mental Health and Global Physical Health scores were calculated based on 10 items each measured on a 5-point scale (from 1 to 5) and subsequently converted.²⁰ These subscale scores ranged from 0 to 100, with a higher score indicating better quality of life and a mean score of 50 indicating the norm for the general population.²²

Procedures

An anonymous questionnaire (electronic and/or paper) available in English, French and Spanish was compiled (LimeSurvey Professional). It included the following descriptive variables: age, body-height, body-weight, duration of professional football or rugby career, level of play, duration and nature of retirement, and current employment status (employed or not employed). Additionally, this questionnaire included questions for the outcomes previously elaborated on (ankle OA, ankle injury, ankle surgery, QoL). Information about the study was sent per email to potential participants by FIFPRO and IRP. Participants interested in the study, gave their informed consent and completed the electronic questionnaire. Participants were asked to complete the questionnaire within 2 weeks, reminders being sent after 2 and 4 weeks. For privacy reasons, the responses to the questionnaires were coded and anonymized. Once completed, the electronic questionnaires were automatically saved on a secured electronic server that only the principal investigator could access. Players participated voluntarily and did not receive any reward for their participation.

8 178

179 Statistical analyses

Data analysis was performed using the statistical software IBM SPSS 24.0 for Windows. Analyses were
 conducted separately for retired professional football and rugby players. Data were assessed (visually) for

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normality and presented as mean \pm standard deviation (SD), median (interquartile range [IQR]), or frequency (proportion in %) as appropriate. Body mass index (BMI) was calculated using the provided weight and height of participants (kg/m^2) . Prevalence of ankle OA, overall and within the age categories of ≤40 years and >40 years, was calculated as the proportion of the number of participants with ankle OA relative to the total number of participants in each category.²⁰ We chose to dichotomize age at 40 years, as our population included former professional athletes. This is opposed to Murray et al. (2018) concerning the general population (dichotomized at 50 years of age), and in line with Song et al. (2019) studying former professional American Football players. We made the assumption that although no-longer professionals, these former professional athletes would no longer be competing at a decent level at the age of 40. We therefore found 40 years to be a more relevant age to dichotomise than 50 years.^{12,23} A logistic regression analysis was performed to determine the association of the number of severe ankle injuries and/or ankle surgeries (continuous independent variable) with ankle OA (dichotomous dependent variable). Severe ankle injuries and or ankle-related surgeries were also expressed using 3 pre-determined categories (0, 1 and >1). This was adjusted for age and BMI, both having been identified as risk factors for OA.^{24,25} Descriptive analyses of health-related quality of life (Global Physical Health and Global Mental Health) were conducted, while comparisons between groups (retired players with ankle OA vs. retired players without ankle OA) were made using a Mann-Whitney U test for independent samples.²⁰

43 200 Patient and public involvement

This research was done without patient involvement. Patients were not invited to comment on the study design and were not consulted to develop patient relevant outcomes or interpret the results. Patients were not invited to contribute to the writing or editing of this document for readability or accuracy.

1

2		
3 4	205	Results
5 6	206	Participants
7 8 0	207	From a total of 750 football and 326 rugby players contacted, respectively 401 and 152 gave their
9 10 11	208	written informed consent and completed the questionnaire (overall response rate of 56%).
12 13	209	Characteristics of the two groups including the pre-determined sub-categories for severe ankle injuries
14 15	210	and ankle surgeries (0, 1 and >1) are illustrated in Table 1.
16 17	211	
18 19 20	212	Prevalence of ankle osteoarthritis, severe injury and surgery
21 22	213	The prevalence of patient reported ankle OA (including the pre-determined categories of \leq 40 years
23 24	214	and > 40 years) among recently retired professional football and rugby players, is presented in Table 2.
25 26 27	215	The prevalence of ankle OA in former professional football players was 9.2% (n = 37). Overall, 94% of
27 28 29	216	players with ankle OA were documented to have sustained ≥1 ankle injury and 62% had undergone ≥1
30 31	217	ankle surgery. In the non-OA group (n = 364), 54% had sustained ≥1 ankle injury and 14% had
32 33	218	undergone ≥1 ankle surgery. Logistic regression analysis (Table 3) showed that football players who
34 35 26	219	had sustained a severe ankle injury were 1.3 (95%CI: 1.1 - 1.6) times more likely to report ankle OA (p <
30 37 38	220	0.001). Football players who had undergone ankle surgery were 2.1 (95%CI: 1.4 – 3.1) times more likely
39 40	221	to report ankle OA (p< 0.001).
41 42	222	
43 44 45	223	The prevalence of ankle OA in former professional rugby players was 4.6% (n = 7). Overall, six out of
45 46 47	224	seven (86%) rugby players with ankle OA reported to have sustained ≥1 ankle injury and four out of
48 49	225	seven (57.1%) had undergone \geq 1 ankle surgery. In the non-OA group (n = 145), 58.2% had sustained \geq 1
50 51	226	ankle injury and 18.4% had undergone ≥1 ankle surgery. Logistic regression analysis (Table 3) showed no
52 53	227	significant association between either severe ankle injuries (OR = 1.1 95%CI: 0.8 – 1.5) or ankle surgeries
54 55 56	228	(OR = 2.0 95%CI: 0.8 – 4.7) and ankle OA among former professional rugby players.
57 58 59		10

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Table 1: Descriptive characteristics of recently retired professional football players and rugby players

1 · 2			Football			Rugby	
∠ 3		Total	Ankle OA	No ankle OA	Total	Ankle OA	No ankle OA
4		(n = 401)	(n = 37)	(n = 364)	(n = 152)	(n = 7)	(n = 145)
5							
6 7	Age in years *	36	39	36	40 ± 6	41 ± 7	40 ± 6
8	(mean + SD)	(32-40, 23-30)	(30-42, 27-49)*	(52-40, 25-50)*			
9	Height in cm	181 ± 7	180 ± 7	182 ± 7	186	191	186
10	(mean ± SD / median; IQR;				(180-191; 166-203)	(173-192; 172-192)	(180-191; 166-203)
11	min-max)						
12 13	Weight in kg	82 ± 9	82 ± 13	82 ± 9	100	108	100
14	(III KY (IIIEUII ± SD))				(91-114; 59-170)	(89-123; 82-130)	(91-113; 59-170)
15	BMI in kg/m ²	24.8	24.8	24.7	29.0	32.2	28.9
16 17	(median; IQR; min-max))	(23.5-26.2; 20.1-32.7)	(22.9-26.6; 20.3-29)	(23.5-26.1; 20.1-32.7)	(27.1-31.4; 20.4-52.5)	(28.5-33.7; 27.4-35.3)*	(27.0-31.2; 20.4-52.5)*
18	Employed in %	89%	92%	89%	91%	86%	92%
19 20	(n)	(350)	(33)	(317)	(139)	(6)	(133)
20 21	Professional career duration	13	13	13	10	10	10
22	in years (median; IQR; min-	(10-15; 2-19)	(12-16; 5-18)	(9-15; 2-19)	(7-15; 2-20)	(8-12.5; 8-13)	(7-15; 2-20)
23	max)						
24	Retirement duration in years	4	5	4	8	12	8
25	(median; IQR; min-max)	(2-/; 1-1/)	(3-9; 1-12)	(2-/; 1-1/)	(3-12; 0-22)	(2-16; 1-19)	(4-11; 0-22)
20	Level of play top league; %	81%	92%	80%	80%	86%	80%
28	(n)	(314)	(34)	(280)	(106)	(6)	(100)
29	Forced retirement %	29%	27%	30%	36%	57%	35%
30 31	(n)	(113)	(10)	(103)	(55)	(4)	(51)
32	Severe ankle iniuries	1	2	1	1 🥒	1	1
33	(median; IQR; min-max)	(0-2; 0-10)	(1-3; 0-10)	(0-2; 0-10)	(0-2; 0-20)	(1-2; 0-3)	(0-2; 0-20)
34							
35	Ankle surgeries	0	1	0	0	1	0
30 37	(median; IQR; min-max)	(0-0; 0-5)	(0-2; 0-5)	(0-0; 0-5)	(0-0; 0-4)	(0-1; 0-2)	(0-0; 0-4)
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Joreviations: CA = Ost. * Statistically significant difference between ankle OA and no ankle OA. Abbreviations: OA = Osteoarthritis; IQR = interguartile range; n = number

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		Football			Rugby	
	Total (n = 401)*	≤ 40 years (n = 271)	> 40 years (n = 83)	Total (n = 152)	≤ 40 years (n = 82)	> 40 years (n = 66)
Prevalence ankle OA in % (95%Cl)	9.2% (6.4 –12.1)	8.5% (5.2 – 11.8)	12.0% (5.0 – 19.1)	4.6% (1.2 – 8.0)	2.4% (0.9 – 5.8)	4.5% (0.5 – 9.6)
	0 ankle injuries	1 ankle injury	> 1 ankle injury	0 ankle injuries	1 ankle injury	> 1 ankle injury
Severe ankle injuries total (%, (n))	42.5% (156)	25.3% (93)	32.2% (118)	40.5% (60)	27.7% (41)	31.8% (47)
- With ankle OA (%, (n))	5.7% (2)	28.6% (10)	65.7% (23)	14.3% (1)	42.9% (3)	42.9% (3)
- Without ankle OA (%, (n))	46.4% (154)	25.0% (83)	28.6% (95)	41.8% (59)	27.0% (38)	31.2% (44)
	No ankle surgeries	1 ankle surgery	> 1 ankle surgery	No ankle surgeries	1 ankle surgery	> 1 ankle surgery
Ankle surgeries total in % (n)	81.5% (299)	10.9% (40)	7.6% (28)	79.7% (118)	15.5% (23)	4.7% (7)
- With ankle OA % (n)	38.2% (13)	29.4% (10)	32.4% (11)	42.9% (3)	42.9% (3)	14.3% (1)
- Without ankle OA % (n)	85.9% (286)	9% (30)	5.1% (17)	81.6% (115)	14.2% (20)	4.3% (6)

*The sum of patients in the subcategories \leq 40 years and > 40 years does not equal the "Total number of participants" due to missing data (regarding age). Abbreviations: OA = Osteoarthritis; n = number; CI = confidence interval;

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Table 3: Association (odds ratio and 95% CI) of severe ankle injury and ankle surgery with ankle osteoarthritis					
among recently retired professional football and rugby players					
	Football (n = 401)	Rugby (n = 152)			

Unadjusted Adjusted ^v Unadjusted Adjusted ^v ievere ankle injury OR, 95%(CI) 1.34 (1.17-1.53) 1.33 (1.14-1.55) 1.01 (0.74-1.38) 1.05 (0.75-1.47) Ankle surgery OR, 95%(CI) 2.20 (1.61-3.00) 2.09 (1.41-3.08) 1.99 (0.92-4.3) 1.96 (0.81-4.71) Adjusted for age and body mass index (BMI); Abbreviations: OR = Odds Ratio; n = number; CI = confidence interval Image: Confidence interval		1000041	· (··· ······	1,0201	
Severe ankle injury 1.34 (1.17-1.53) 1.33 (1.14-1.55) 1.01 (0.74-1.38) 1.05 (0.75-1.47) OR, 95%Cl) 2.20 (1.61-3.00) 2.09 (1.41-3.08) 1.99 (0.92-4.3) 1.96 (0.81-4.71) Adjusted for age and body mass index (BMI); Abbreviations: OR = Odds Ratio; n = number; Cl = confidence nterval 1.01 (0.74-1.38) 1.96 (0.81-4.71)		Unadjusted	$Adjusted^\psi$	Unadjusted	$Adjusted^{\psi}$
OR, 95%Cl) 2.20 (1.61-3.00) 2.09 (1.41-3.08) 1.99 (0.92-4.3) 1.96 (0.81-4.71) OR, 95%Cl) 'Adjusted for age and body mass index (BMI); Abbreviations: OR = Odds Ratio; n = number; Cl = confidence 'Adjusted for age and body mass index (BMI); Abbreviations: OR = Odds Ratio; n = number; Cl = confidence	Severe ankle injury	1.34 (1.17-1.53)	1.33 (1.14-1.55)	1.01 (0.74-1.38)	1.05 (0.75-1.47)
Ankle surgery 2.20 (1.61-3.00) 2.09 (1.41-3.08) 1.99 (0.92-4.3) 1.96 (0.81-4.71) 'Adjusted for age and body mass index (BMI); Abbreviations: OR = Odds Ratio; n = number; CI = confidence nterval Image: Confidence nterval	(OR, 95%CI)				
OR, 95%CI) 'Adjusted for age and body mass index (BMI); Abbreviations: OR = Odds Ratio; n = number; CI = confidence nterval	Ankle surgery	2.20 (1.61-3.00)	2.09 (1.41-3.08)	1.99 (0.92-4.3)	1.96 (0.81-4.71)
'Adjusted for age and body mass index (BMI); Abbreviations: OR = Odds Ratio; n = number; Cl = confidence nterval	(OR, 95%CI)				
nterval	'Adjusted for age and l	body mass index (BMI)	; Abbreviations: OR = 0	Odds Ratio; n = number,	: CI = confidence
	nterval				

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Mental and physical QoL

For the PROMIS physical health score a statistically significant lower outcome was found for football (p = 0.01) and rugby players (p = 0.009) with

OA (48 and 45 points respectively) compared to players without OA (51 and 51 points respectively). The PROMIS mental health score was not

significantly different between players with OA vs players without OA. Data is presented in Table 4.

Table 4: Mental and physical quality of life (0-100) of recently retired football and rugby players with ankle OA compared to players without ankle OA

		Football				Rugby			
	Total (n = 401)	With ankle OA (n = 37)	Without ankle OA (n = 364)	p-value	Total (n = 152)	With ankle OA (n = 7)	Without ankle OA (n = 145)	p-value	
PROMIS mental health (median; IQR; min-max)	51; 46-56; 31-68	51; 46-53; 31-63	53; 46-56; 31-68	0.251	52; 46-56; 28-68	48; 48-53; 46-56	53; 46-56; 28-68	0.567	
PROMIS Physical health* (median; IQR; min-max)	51; 46-56; 27-68	48; 42-54; 35-62*	51; 48-58; 27-68*	0.010	51; 45-54; 30-68	45; 37-48; 37-51*	51; 37-56; 30-68*	0.009	

*Statistically significant difference; Abbreviations: OA = Osteoarthritis; IQR = Interquartile range; n = number

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Post-hoc analysis

242 In the literature, forced retirement is a risk factor for post-career mental health symptoms.²⁶ Following 243 data analysis, we were concerned by the seemingly high percentage of football and rugby players who 244 stopped due to a career ending injury. As we were analysing the impact ankle OA had on QoL, we felt 245 the effect of a career ending injury on QoL to be in line with our initial objectives. Consequently we 246 performed a post-hoc analysis to determine any associations between reported forced retirement and 247 mental and physical PROMIS health scores. In this study, 31% of players reported to have been forced to 248 retire, specifically: 29% of football players and 36% of rugby players. 249 250 The post-hoc sub-analysis of all players regarding mental and physical health scores associated with 251 voluntary retirement and in conjunction with ankle OA is presented in Table 5. No post-hoc sub-analysis 252 was done for football and rugby players separately, regarding voluntary retirement in conjunction with 253 ankle OA, due to the low number of ankle OA cases among rugby players. However mental and physical

health scores in football and rugby due to forced retirement is presented separately in Table 5.

Table 5: Post-Hoc analysis regarding mental and physical quality of life of recently retired football and rugby players who reported to have been forced to retire, and quality of life of players who reported to have been forced to retire with ankle OA compared to players without ankle OA

9 10			PROMIS mental health (median; IQR; min-max)	p-value	PROMIS physical health (median; IQR; min-max)	p-value
11 12		T / /				
12						
13 14 15		Voluntarily retired* (n = 369)	53; 48-56; 31-68*	n < 0.001	54; 48-58; 30-68*	n < 0.001
16		Forced to retire* (n = 168)	48; 44-53; 28-68*	ρ<0.001	48; 42-51; 27-68*	ρ< 0.001
17 18 19		Voluntarily retired with OA* (n = 30)	51; 46-53; 36-63		51; 45-54; 35-62*	
20 21 22		Voluntarily retired without OA* (n = 339)	53; 48-59; 31-68	p = 0.074	54; 48-58; 30-68*	p = 0.001
23 24		Forced to retire with OA (n = 14)	48; 46-55; 31-63		42; 40-46; 37-58	
25 26 27		Forced to retire without OA (n = 154)	48; 44-53; 28-68	p = 0.841	48; 42-52; 27-68	p = 0.159
28 29		Football*				
30 31		Voluntarily retired* (n = 272)	53; 48-56; 31-68*		54; 48-58; 30-68*	
32 33 34		Forced to retire* (n = 113)	48; 44-53; 31-63*	p < 0.001	48; 42-51; 27-62*	p < 0.001
35		Rugby*				
36 37		Voluntarily retired* (n = 97)	53; 48-56; 31-68*	n - 0.019	54; 48-58; 35-68*	n < 0.001
38 39		Forced to retire* (n = 55)	48; 44-56; 28-68*	ρ – 0.018	48; 37-54; 30-68*	ρ< 0.001
40		*Statistically significant differ	ence; Abbreviations: IQR = Int	erquartile rang	e; OA = Osteoarthritis; n = numb	per
41 42	258					
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3	260	Discussion
4 5	261	Our most important findings were that (i) the prevalence of ankle OA among former professional
6 7 0	262	football and rugby players was, 9.2% (95%Cl 6.4 –12.1) and 4.6% (95%Cl (1.2 – 8.0), respectively; (ii)
8 9 10	263	previous severe ankle injuries and ankle surgeries are associated with ankle OA among former
11 12	264	professional football players; and (iii) ankle OA does not lead to reduced mental or physical QoL in
13 14	265	former professional football and rugby players.
15 16 17	266	
17 18 19	267	Ankle OA in football and rugby compared to other sports and the general population
20 21	268	Several studies have reported symptomatic ankle OA prevalence among a variety of former elite
22 23	269	athletes, ranging from 0% in volleyball (n = 22), long distance running (n = 30), elite high-jump (n = 30)
24 25	270	and ballet (n = 27) to 10% among former military parachutists (n = 40). ^{$27-31$} In our study, the prevalence
26 27 28	271	of ankle OA among former professional football (9.2%) and rugby players (4.6%) was higher than the
20 29 30	272	estimated prevalence among the general population (3.4 %). ¹² Murray et al. (2018) invited participants
31 32	273	with self-reported ankle pain (11.7%) from the general population for a radiograph (radiological atlas of
33 34	274	Foot OA). ^{12,32}
35 36 27	275	
37 38 39	276	Our study reports a lower ankle OA prevalence compared to previous studies among former
40 41	277	professional football players (12-19%). ^{13–15} In these studies, data was similarly collected using cross-
42 43	278	sectional retrospective self-reported questionnaires. ^{13,14} No previous data is available concerning ankle
44 45	279	OA among former professional rugby players. The low ankle OA prevalence among rugby players in this
46 47 48	280	study is likely due to either an under reporting of ankle OA among rugby players or a type II error.
49 50	281	
51 52	282	Association of severe ankle injuries and ankle surgeries with ankle OA
53 54	283	In our study, football players were 1.3 (95%CI: 1.1 - 1.6) and 2.1 (95%CI: 1.4 – 3.1) times more likely to
55 56 57	284	develop ankle OA for every ankle injury and ankle surgery, respectively. This is in line with previous
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3 4	285	studies that showed an association between ankle OA and ankle injuries, stating traumatic ankle injuries	
5 6	286	to be the main cause of ankle OA (70-78%). ^{6,7} Consequences of ankle injuries are thought to be due to	
7 8	287	either direct damage the articular surfaces or change in ankle joint biomechanics (e.g. through chronic	
9 10 11	288	ankle instability due to ankle ligament damage). ³³ No association was seen for severe ankle injuries or	
12 13	289	ankle surgeries and ankle OA among rugby players. Although the best estimate for the association of	
14 15	290	ankle surgeries and OA is similar to football (OR = 2), the 95%CI includes 1 (0.8 – 4.7). This may be due	
16 17	291	to insufficient power among rugby players, as only 7 reported ankle OA.	
18 19 20	292		
20 21 22	293	Health-related quality of life	
22 23 24 25 26 27 28 29 30 31 32 33	294	Filbay et al. (2019) reported factors such as age, type of sport (contact/collision sports), OA and	
	295	involuntary retirement from sport to possibly negatively impact QoL. ^{26,34–36} However the positive	
	296	impacts associated with elite sport participation, such as pride of accomplishments, social network	
	297	(through sport participation) and coping and adjustment to musculoskeletal pain (due to resilience and	
	298	determination), may persist and even compensate reduced physical QOL beyond retirement of the	
34 35	299	player's athletic career. ³⁴	
36 37	300		
38 39 40	301	Our post-hoc sub-analysis revealed 29% of football players and 36% of rugby players were forced to	
40 41 42	302	retire. Although we reported several instances of statistically significant differences in our study for both	
43 44	303	physical and mental PROMIS health scores, all were similar to the estimated norm for the general	
45 46	304	population (50 points). Unfortunately, there is no robust minimally clinically important difference	
47 48 40	305	(MCID) threshold for the PROMIS-10 global short form applicable for this population. Consequently, it is	
50 51	306	difficult to interpret the results. Nevertheless, although statistically significant, the PROMIS mental and	
52 53	307	physical health scores observed, for both football and rugby players, are unlikely clinically significant.	
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3 4	308	Based on our data, ankle OA does not lead to reduced mental or physical QoL in former professional	
5 6	309	football and rugby players.	
7 8 9	310		
) 10 11	311	Strengths and limitations	
12 13	312	We had an overall good response rate (56%) and a large group of participating football players (401).	
14 15 16	313	However, several limitations should be mentioned. Firstly, this study had a cross-sectional design,	
10 17 18	314	therefore no causal-association can be determined. Secondly, there were only 7 rugby players with OA,	
19 20	315	implying a potential serious type II error. Thirdly, recruitment procedures were blinded to the research	
21 22	316	team, a non-response analysis could not be performed. Fourthly, research based (retrospective)	
23 24 25	317	questionnaires are dependent on the accuracy and understanding of the participant. Professional	
25 26 27	318	athletes might generally remember the number of severe ankle injuries or surgeries resulting in a	
28 29	319	training or match absence of ≥4 weeks. However, we cannot exclude that participants were not able to	
30 31	320	precisely recall all their sustained severe ankle injuries and ankle surgeries or that they did not consult	
32 33	321	their most recent medical professional. Finally, our OA definition did not include a clinical and	
34 35 36	322	radiological evaluation.	
37 38	323		
39 40	324	Implications for practice	
41 42	325	The epidemiological data from this study regarding ankle OA prevalence among former professional	
43 44 45	326	football and rugby players, and its association with both severe ankle injuries and ankle surgeries, is an	
46 47	327	essential part of protecting the athlete's health. In both professional football and rugby, a higher	
48 49	328	prevalence of ankle OA can be expected compared to the general population. Furthermore, As the	
50 51	329	majority are post-traumatic, preventive measures and adequate rehabilitation are recommended.	
52 53	330	Awareness should be raised, regarding the increased risk of developing ankle OA following each ankle	
54 55 56 57 58	331	injury and surgery. Additionally primary and secondary prevention programmes should be implemented	1. 20
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- 3 4	332	Exercise based prevention intervention programs, such as the FIFA11+ in football and the movement
5 6 7 8	333	control injury prevention programme in rugby, have found reduced ankle injury incidence and
	334	burden. ^{37,38} Finally, management of ankle OA is key for retired players, focusing on healthy lifestyle
9 10 11	335	including physical activity, pain alleviation, function and minimising disability. ^{39–41} With this intention,
12 13	336	FIFPRO has developed an "After Career Consultation". ⁴² Future studies will evaluate its relevance.
14 15	337	
16 17	338	Conclusion
18 19 20	339	This cross-sectional study found an athlete reported ankle OA prevalence of 9.2% and 4.6% among
20 21 22	340	recently retired professional football and rugby players, respectively. Severe ankle injuries and ankle
23 24	341	surgeries were associated with a higher prevalence of ankle OA in former professional football players.
25 26	342	No clinically relevant differences in QoL were reported in players with OA compared to players without
27 28 20	343	OA. As a higher prevalence of ankle OA can be expected in both professional football and rugby
29 30 31	344	compared to the general population, and the majority are post-traumatic, primary and secondary
32 33	345	preventive measures for ankle injuries and an optimal diagnosis and treatment plan are recommended.
34 35	346	
36 37 39	347	Acknowledgements
30 39 40	348	The Authors would like to thank all football and ruby players' associations involved. The authors are
41 42	349	grateful to all retired professional football and rugby players who participated in the study.
43 44	350	
45 46	351	Contributorship statement
47 48 49	352	Authorship was determined according to the ICMJE authorship recommendations. All authors (LP, $\mathrm{HA},$
50 51	353	SK,ML,CR,KS,WV,GR,JT,GK,VG) were involved in the design of the study, data analysis and
52 53	354	data interpretation. VG and GK were responsible for data collection. LP drafted the manuscript, with
54 55	355	critical review provided by all authors (LP, HA, SK, ML, CR, KS, WV, GR, JT, GK, VG). All
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3 4	356	authors approved the final version of the manuscript (LP, HA, SK, ML, CR, KS, WV, GR, JT, GK,
5 6	357	VG).
7 8	358	
9 10 11	359	Funding
12 13	360	This research received no specific grant from any funding agency in the public, commercial or non-profit
14 15	361	sectors.
16 17 19	362	
10 19 20	363	Competing interests
21 22	364	None declared.
23 24	365	
25 26 27	366	Ethics approval
27 28 29	367	Ethical approval for the study was provided by the Ethical Committee of the Yokohama City Sports
30 31	368	Medical Center (17.003; Yokohama, Japan) and the Medical Ethics Review Committee of the Academic
32 33	369	Medical Center (W16_366#16.431; Amsterdam, The Netherlands)
34 35	370	
36 37 38	371	Data sharing statement
39 40	372	No additional data available
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STROBE Statement—Checklist of items that should be included in reports of cross-sectional studies

	Item No	Recommendation
Title and abstract	1	(a) Indicate the study's design with a commonly used term in the title or the abstract
The and abstract	1	(a) indicate the study s design with a commonly used term in the title of the abstract $-$
		(b) Provide in the abstract an informative and balanced summary of what was done
		and what was found $-$
Introduction		
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported –
Dackground/rationale	2	$\sqrt{1}$ n 4 line 64 – 96
Objectives	3	State specific objectives, including any prespecified hypotheses $-\sqrt{2}$ n 4 line 98 –
objectives	5	104
Madha da		
Methods Study design	1	Present low alaments of study design early in the paper of n 6 line 109 114
Study design	4	Present key elements of study design early in the paper $-\sqrt{2}$, p. 6 line 108 -114
Setting	5	Describe the setting, locations, and relevant dates, including periods of recruitment,
Doutioinonto	6	exposure, follow-up, and data collection -3 : p. 6 line 116 -155
Participants	0	(a) Give the eligibility criteria, and the sources and methods of selection of $a_{1}^{(a)}$
Variablas	7	participants – V. p. 6 line 116 – 135
variables	/	medifiere Ciue diagnostie criterie if emplicable al in 6 line 125 150
Data anumana/	0*	For each surgical of interact, since courses of data and datails of matheds of
Data sources/	0.	rol each variable of interest, give sources of data and details of interiods of
measurement		more than one group with 6 line 161 - 172
Diag	0	$\frac{1}{12}$
Study size	9	Explain how the study size was arrived at a/a $p = 6$, 7 line 120, 122
Study Size	10	Explain how the study size was arrived at $- v$. p. $6 - 7$ line 130 - 135
Quantitative variables	11	describe which grownings were shown and why the 8 0 line 175 188
Statistical matheda	12	describe which groupings were chosen and why -3 : p. 8-9 line $1/3 - 188$
Statistical methods	12	(a) Describe an statistical methods, including those used to control for controllating
		(b) Describe any methods used to examine subgroups and interactions
		(c) Explain now missing data were addressed
		(a) If applicable, describe analytical methods taking account of sampling strategy
		(\underline{e}) Describe any sensitivity analyses
		- V: p. 8-9 line 1/5 - 188
Results		
Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially
		eligible, examined for eligibility, confirmed eligible, included in the study,
		completing follow-up, and analysed – \forall : p. 10 line 192 – 196
		(b) Give reasons for non-participation at each stage – $\sqrt{2}$: p. 10 line 192 – 196
		(c) Consider use of a flow diagram
Descriptive data	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and
		information on exposures and potential confounders – \forall : p 11 table 1
		(b) Indicate number of participants with missing data for each variable of interest
Outcome data	15*	Report numbers of outcome events or summary measures – $\sqrt{2}$ p 11 – 14, table 2 -4
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and
		their precision (eg, 95% confidence interval). Make clear which confounders were
		adjusted for and why they were included – $\sqrt{\mathbf{p} 11 - 14}$, table 2 -4
		(b) Report category boundaries when continuous variables were categorized

	(c) If relevant, consider translating estimates of relative risk into absolute risk for a
	meaningful time period
17	Report other analyses done—eg analyses of subgroups and interactions, and
	sensitivity analyses – $\sqrt{:}$ p 15, line 227 – 237 + p 16 table 5
18	Summarise key results with reference to study objectives – $\sqrt{2}$ p 17, line 242 - 245
19	Discuss limitations of the study, taking into account sources of potential bias or
	imprecision. Discuss both direction and magnitude of any potential bias – $\sqrt{:}$ p 3,
	line 53 - 62
20	Give a cautious overall interpretation of results considering objectives, limitations,
	multiplicity of analyses, results from similar studies, and other relevant evidence – $$:
	p 19 - 20, line 293 - 317
21	Discuss the generalisability (external validity) of the study results p 19 - 20 , line 293
	- 305
22	Give the source of funding and the role of the funders for the present study and, if
	applicable, for the original study on which the present article is based p 21, line 339
	- 340
	17 18 19 20 21 22

*Give information separately for exposed and unexposed groups.

Note: An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.

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