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'What's my risk of sustaining an ACL injury while playing football (soccer)?' A systematic review with meta-analysis

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Abstract

Objective—To estimate the incidence proportion (IP) and incidence rate (IR) of ACL injury in football players.

Design—Systematic review with meta-analysis.

Data sources—PubMed, CINAHL and SPORTDiscus electronic databases were searched from inception to 20 January 2017.

Contributors All authors contributed significantly to the conception, design and writing of this manuscript.

Competing interests None declared.

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Eligibility criteria for selecting study—Studies that reported the total number of participants/population by sex, total number of ACL injuries by sex and total person-time by sex were included.

Results—Twenty-eight studies were included. The IP and IR of ACL injury in female football players were 2.0% (95% CI 1.2% to 3.1%) and 2.0/10 000 athlete exposures (AEs) (95% CI 1.6 to 2.6; I^2 =91%) over a period of one season to 4 years. The IP and IR of ACL injury in male players were 3.5% (95% CI 0.7% to 8.2%) and 0.9/10 000 AEs (95% CI 0.7 to 1.1; I^2 =94%). Studies that evaluated matched cohorts of female and male players showed no difference in IP (relative risk=1.2; 95% CI 0.9 to 1.6; P=0.47) over a period of one season to 4 years. Women were at greater risk than men (incidence rate ratio (IRR)=2.2; 95% CI 1.6 to 3.1; I^2 =83%; P<0.001). When accounting for participation level, the difference in IR between women and men was greatest for intermediate players (IRR=2.9; 95% CI 2.4 to 3.6) compared with amateur (IRR=2.6; 95% CI 1.4 to 4.8) and elite (IRR=2.0; 95% CI 1.1 to 3.4) players.

Summary/conclusion—Overall, more men sustained ACL injury in football. There was no difference in the relative risk of ACL injury between female and male football players in a window that spanned one season to 4 years. The IR of ACL injury among women was 2.2 times higher than the IR of ACL injury among men. The reported sex disparity in ACL injury was independent of participation level.

INTRODUCTION

ACL rupture is one of the most serious injuries associated with participation in football (soccer). ACL injury can have serious consequences for activity level and quality of life. 4 Many athletes with ACL injury do not return to full sport participation, gain weight, and develop degenerative knee conditions, such as osteoarthritis, later in life. 5-7

Female football players may have a higher risk of these problems than their male counterparts. ⁸⁹ This sex disparity requires attention from the sports medicine community because of the increasing popularity of football among women. The number of women playing football increased from the 26 million reported in the most recent Big Count survey in 2006 to over 30 million reported in the Women's Football Survey in 2014. ¹⁰¹¹ In addition, programmes with the goal of further increasing women's participation were announced in the last Women's Football Symposium. ¹² To safely achieve this goal requires actions to attract women to football and foster their successful, long-term participation. For the latter, addressing sex disparities in the risk for serious injuries, such as ACL rupture, is key.

Previous studies in collegiate athletes estimated that female football players were at approximately three times the risk of sustaining ACL injury relative to male soccer players from 1989 to 1993, from 1989 to 2002 and from 1989 to 2004. ^{13–15} A 2007 meta-analysis estimated that the incidence rate (IR) of ACL injury was more than 2.5 times higher among female football players relative to male football players. ⁸ Most studies included in this meta-analysis documented ACL injuries in college-level football players. It is not clear whether participation level influenced the documented sex differences in ACL injury IR. The sex differences in incidence proportion (IP; number of new injuries relative to total number of athletes exposed) seem to be less pronounced. ⁹ A possible reason is that the participation of

female football players has been historically characterised by fewer training sessions and games, but that difference is rapidly changing. ^{10–12} In the last 10 years, there have been no additional systematic reviews or meta-analyses aggregating available studies. The aims of this systematic review were to: (1) provide current, pooled estimates of IP and IR of ACL injury in female and male football and (2) evaluate whether sex differences in ACL injury were affected by participation level.

METHOD

A systematic review was conducted using the Preferred Reporting Items for Systematic Reviews and Meta-Analyses guidelines. We searched the PubMed and EBSCO host (CINAHL, SPORTDiscus) electronic databases from database inception to 20 January 2017. We used the search phrase: anterior cruciate ligament AND (injury OR tear OR rupture) AND (incidence OR prevalence OR epidemiology). The terms soccer and football were not used because this search was also used in a larger meta-analysis that included all sports. ¹⁶ The articles pertinent to football were selected for this meta-analysis. We limited the search to peer-reviewed articles published in English language. Experts in the field were contacted for further study suggestions, and references from review papers were examined to identify any further relevant articles for potential inclusion. Publication details from all studies identified in the literature search were exported to bibliographic software (EndNote X7, Thomson Reuters, USA).

Selection criteria

One assessor screened articles for inclusion. Any grey areas of inclusion were brought to discussion with the second assessor, and any disagreements were arbitrated by a third assessor. Articles were screened first by title, then by abstract and finally by full text according to the inclusion and exclusion criteria (box 1). Full texts were retrieved when the title or abstract met the selection criteria, or when the status (include or exclude) could not be determined from the title and/or abstract alone.

Data extraction and management

The primary variables extracted were the number of ACL injuries, exposure for men and exposure for women and participation level (amateur, intermediate or elite). The amateur category included recreational, amateur and high school football players. ^{17–25} The intermediate category included collegiate and upper division amateur football players. ^{172226–28} The elite category included professional and elite football players. ^{29–36} One author recorded all of the pertinent data from the included articles and another author independently reviewed those data for accuracy and completeness. The studies included in the IP and IR analyses are listed in online supplementary file 1.

The reported person-time unit was not uniform across studies. To establish a common metric, we calculated athlete exposure (AE). When the number of player-hours was reported, AEs were estimated by multiplying the number of player-hours by two. 2024253032–3537–40 This was to estimate the average length of training sessions and games with warm-ups included. Play-er-days were treated as AEs as athletes typically participate in one training

session or game per day. This unit was only used in studies that investigated high school and college athletes. 17 For studies that reported IR by sex, the number of AEs and the reported IRs were used to calculate the number of ACL injuries by sex (number of ACL injuries by sex=total AE by sex × rate numerator by sex/rate denominator by sex). 21

Whenever possible, the IP and IR were calculated. The IP was calculated by dividing the number of new ACL injuries by the total number of participants over the given time period specified in each study. The denominator for IP ranged from one season to 4 years. The IR was calculated by dividing the total number of new ACL injuries by the total number of exposures. We emailed the authors of studies where the number of ACL injuries by sex could not be estimated. If the study authors did not have access to the information or did not respond to our email, the study was excluded from the meta-analysis. ^{1941–43}

Risk of bias assessment

Risk of bias was assessed by two authors independently using the Quality Assessment Tool for Observational Cohort and Cross-Sectional Studies. 44 This tool dichotomously assesses criteria including participation rate, whether exposure data were collected prospectively, whether the timeframe was sufficient to allow for the outcome to occur and number of participants lost to follow-up after baseline. If the criterion was present, the item was given a 'Y'. If the criterion was absent or was not reported, the item was given an 'N'. Any discrepancies in assessment were discussed. For discrepancies that could not be resolved, a third author was consulted for arbitration. Studies where interventions were administered were treated as cohort studies in our analyses and assessed using the same tool. Studies that fulfilled fewer than 50% of criteria were deemed to be of low quality.

Statistical analysis

Injury data were analysed using R (V3.3.2, the R Foundation for Statistical Computing). We used the packages meta and metafor with the functions metarate for IR, metaprop for IP single ratio, metainc for incidence rate ratio (IRR) and metabin for IP binary data ratio (weighted for individual study size). When the number of ACL injuries was zero, 0.1 was substituted so the measure could be used in meta-analysis. For calculating IP and IR for the total population, female football players only and male football players only, all studies that presented data were included in the respective analyses. When calculating the relative risks (RRs) and IRRs, only studies that included men and women in the same study were used. For example, studies that included only women were used to calculate female IP and IR, but were not used to calculate RR or IRR.

Injury risk proportions for individual studies and pooled estimates were summarised in forest plots for the following subgroups: woman, man and combined. A pooled estimate for the RR of ACL injury in women compared with men was calculated and summarised in a forest plot. Raw injury IRs for individual studies and pooled estimates were summarised in forest plots for the following groups and subgroups: woman, man and combined. Pooled IRRs for women compared with men were calculated and summarised in forest plots. Heterogeneity was assessed using the I² statistic, 45 which estimates the variation resulting from heterogeneity and not chance. 45 I² values over 75% were considered high. 45

Publication bias was assessed using funnel plots with SE as the measure of the study size on the y axis and the ratio on the x axis. A P value of 0.05 was considered significant for all statistical analyses.

Sensitivity analyses

To account for heterogeneity among studies, we used sensitivity analysis to assess the effect of each individual study on the overall rate of ACL injury. This was done by removing one study, performing the analysis, investigating the effect of the removal on heterogeneity, returning the study to the analysis and repeating the procedure. This was repeated for each main analysis.

To account for bias, we attempted to perform separate analyses on studies that met the criteria for item 11 (reliability and validity of measurement of the dependent variable) and item 14 (statistical adjustment for potential confounding variables). However, there were too few studies that met the criteria. Therefore, we included all studies in the final analysis (regardless of heterogeneity), and we expected that combining sports with varying risks of ACL injury would result in high heterogeneity.

Subgroup analyses

To account for differences in participation levels, we compared IRs from amateur, intermediate and elite athletes. To identify changes in the rates of ACL injury since the rates established in 2007,⁸ we calculated rates in studies published after 2007.

RESULTS

The final search yielded 3774 abstracts. After removal of duplicates, there were 1300 records. After title and abstract screening, 1155 articles were excluded. The remaining 145 articles were manually cross-referenced, and experts were consulted to identify additional relevant articles, resulting in the inclusion of a further 17 articles. We screened the full-text articles of 162 articles, and 28 studies were included (figure 1 and online supplementary file 1).

Risk of bias assessment

Three studies fulfilled 75% or more of the criteria, and 23 studies fulfilled 50% or more of the criteria (online supplementary file 2). Five studies fulfilled fewer than 50% of the criteria and were deemed to be of low quality. It is possible that some studies may have been judged as being at high or moderate risk of bias simply because they did not report particular information, including the number of eligible individuals, how outcomes were measured and attrition.

The pooled ACL injury IP for all football players was 2.7% (95% CI 1.6% to 4.1%; I^2 =98%; figure 2). The pooled ACL injury IP for female football players was 2.0% (95% CI 1.2% to 3.1%; I^2 =97%; figure 3). The pooled ACL injury IP for male football players was 3.5% (95% CI 0.7% to 8.2%; I^2 =99%; figure 4). In studies that included only women or only men, the IP was greater in men than in women. However, we found that female and male football

players were at equal risk for injury (RR=1.2; 95% CI 0.9 to 1.6; $I^2 = 0\%$; P = 0.47; figure 5). A list of studies used for each analysis are located in online supplementary file 1.

ACL injury incidence among all football players was 1.7/1000 AEs (95% CI 1.4 to 2.1; $I^2 = 93\%$; figure 6). Injury incidence among women was 2.0 ACL injuries per 10 000 AEs (95% CI 1.6 to 2.6; $I^2 = 91\%$; figure 7), and injury incidence among men was 0.9 ACL per 10 000 AEs (95% CI 0.7 to 1.1; $I^2 = 94\%$; figure 8). When controlling for exposure, women had a 2.2-fold increase in IR of ACL injury relative to men (IRR=2.2; 95% CI 1.6 to 3.1; $I^2 = 83\%$; P<0.001; figure 9).

The IRR was highest for intermediate players (IRR=2.9; 95% CI 2.4 to 3.6). The IRR was lower for amateur (IRR=2.6; 95% CI 1.4 to 4.8) and elite (IRR=2.0; 95% CI 1.1 to 3.4) players (Figure 10).

The subgroup analysis of studies published after 2007 indicates that the IRR has remained stable over time (IRR=2.5; 95% CI 2.1 to 3.02; figure 11).

Sensitivity analyses

Excluding single studies from each analysis did not alter the statistical heterogeneity (I² statistic). Heterogeneity remained high (greater than 75%) throughout the procedure. Excluding studies that did not fulfil item 11 in the risk of bias assessment did not statistically alter the pooled estimates for ACL injury incidence, and heterogeneity remained high. Excluding studies that did not fulfil item 14 in the risk of bias assessment did not statistically alter the pooled estimates for total ACL IR and female ACL injury IR. There were insufficient studies to conduct sensitivity analyses (excluding studies that did not fulfil item 14) for the other ACL injury outcomes.

Publication bias

The funnel plot for RR indicated that almost all studies fell within the expected parameters, most with low SE indicating that most studies were large (online supplementary file 3). A majority of studies reported that women had greater IP than men. The funnel plot for IRR indicated that most studies fell within the expected parameters (online supplementary file 4). SE was relatively low, indicating that studies were large, and a majority of studies reported that women were at increased risk of ACL injuries relative to men. The studies are not evenly distributed in the funnel, with studies missing from the lower left quadrant. Studies in the lower left quadrant would represent smaller studies that report a greater IP or IR of ACL injuries in men compared with women.

DISCUSSION

The IP and IR of ACL injury in female football players were 2.0% (95% CI 1.2% to 3.1%) and 2.0/10 000 AEs (95% CI 1.6 to 2.6; I^2 =91%) over a period of one season to 4 years, respectively. The IP and IR of ACL injury in male football players were 3.5% (95% CI 0.7% to 8.2%) and 0.9/10 000 AEs (95% CI 0.7 to 1.1; I^2 = 94%), respectively. Female football players had a similar ACL injury IP to male football players. Female football players had a 2.2-fold increase in the IR of ACL injury compared with men. Women remained at greater

risk of ACL injury regardless of participation level, suggesting that injury risk may be independent of participation level. The IRR identified in our review (2.50) is similar to previous research (2.67).⁸

We aimed to answer the following questions: (1) what are the IP and the IR of ACL injuries for all football players, female football players and male football players? and (2) are there sex differences in IP and IR of ACL injuries? When pooling data to answer our review questions, there was a trade-off between including more studies and producing spurious estimates when pooling statistically and clinically heterogeneous data. We did not exclude studies based on either risk of bias or level of evidence, and we judged the risk of bias in included studies as moderate because most included studies (23/28) met 50% or more of the criteria. Many criteria that were not met were not relevant to epidemiological study quality, but the large number of included studies may enhance the generalisability of our results. Based on the Centre for Evidence Based Medicine, these estimates could be judged as level 4 evidence (Grade of Recommendation, C). With regard to publication bias, there were few studies published indicating that men were at greater risk of sustaining ACL than women. While it is possible that this dearth is due to publication bias, it seems more likely that most studies tend to find that women are at greater risk of sustaining ACL injury than men. Therefore, we deem that publication bias is low.

While we found a difference in the IR of ACL injury between men and women, we did not find a difference in IP This contrasts with previous research. IR is the rate of new injuries, and IP is the number of newly injured individuals. If the rate at which ACL injury occurs is more than twice as high in women, it seems reasonable to expect that a higher proportion of women would sustain this injury than men. However, this assumes that exposure is similar between men and women. The higher IR of ACL injury among female athletes associated with the absence of sex effects on IP suggests that men have greater exposure than women (ie, men participate in more training sessions and games). Given the number of female sports teams is increasing, the difference in participation rates between men and women might be expected to reduce. An increase in the number of championships, games per championship and training sessions played by women is also likely. Therefore, an increase in IP among female football players may follow.

Among football players, women had a 2.2 times greater incidence of ACL injury compared with men. Our estimate of the ACL IR for female football players was 2.5, which is similar to previous research. The fact that the IR among female football players has remained steady, despite increasing evidence of the effectiveness of injury prevention interventions, is disappointing. Even with increased knowledge about injury prevention, we did not find evidence that the incidence of ACL injury in female football players has reduced. The disparity between the effects of intervention studies and injury outcomes in female athletes highlights the need to bridge the gap between scientific advances about ACL injury prevention and clinical practice. If primary ACL injury prevention programmes are effective, then the challenge becomes to identify and overcome factors that may be limiting the clinical application of scientific evidence.

Effective prevention programmes require substantial resources and time commitments. These programmes require intervention providers to supervise athletes to ensure correct technique and can be perceived as time-consuming. ²¹⁵⁰⁵¹ Care must be taken in designing, implementing and encouraging adherence to prevention programmes to ensure prophylactic effects. ⁴⁰⁵²⁵³ Even where these factors are taken into consideration, the most effective prevention programmes are associated with a high number needed to treat. ⁵⁴ Optimistic estimates suggest that 108 athletes have to be treated to prevent one ACL injury. ⁵⁴ Therefore, these programmes may have greatest value when a large population of football players is considered.

Methods issues

There are three issues related to systematic review methods that must be considered when reading and using the results of our systematic review.¹⁶

Risk of bias assessment—Poor internal validity (bias within studies) may have contributed to statistical heterogeneity. The tools available to assess risk of bias in observational (epidemiological) studies evaluate criteria that do not necessarily contribute to bias. For example, existing tools assess for randomisation of subjects, but this criterion is not relevant for prognosis studies. Items like this negatively impact risk of bias even though they are irrelevant to the study design or research question. Despite the limitations regarding the assessment of risk of bias, there was bias in included studies. Bias was not further reduced when investigating football alone relative to investigating all sports combined. ¹⁶ Therefore, caution should be exercised in generalising rates and risks.

Sources of variability—The high I^2 statistics in our analyses indicate that the variability in injury rates was due to heterogeneity rather than to chance. Heterogeneity remained high in studies investigating football alone relative to studies investigating a range of sports. ¹⁶ There are at least three possible explanations for high heterogeneity in our review: (1) including a variety of study designs, including randomised controlled trials and cohort studies, (2) variable exposure data and estimating missing exposure for some studies—conversion of play- er-days and player-hours to AEs may have resulted in an underestimate of the true exposure which would contribute to an overestimation of the IRs and (3) differences in participation level.

Sensitivity and subgroup analyses—We used sensitivity and subgroup analyses based on participation level to explore potential sources of heterogeneity. However, statistical heterogeneity remained high in all sensitivity and subgroup analyses. We were unable to identify the main sources of statistical heterogeneity, and this reduces the robustness of the estimates in our systematic review.

Limitations

IPs and rates may have been overestimated by the inclusion of reinjuries. However, the inclusion of reinjuries is likely a small source of bias. Some studies only included non-contact mechanisms of injury. Therefore, the estimates provided here may be underestimates

of the true rates, although it is difficult to be certain. IRs were based on estimated exposure for 18 of 38 studies included for rate analyses.

We did not assess the influence of age on the injury estimates because many studies did not specify participant age or age range. Controlling for age would have substantially reduced the number of studies eligible for inclusion. While the search was systematic and comprehensive, we acknowledge the risk for publication and language bias.

Future research—Whenever possible, future research should investigate injury epidemiology in male and female athletes simultaneously to allow for sex comparisons in future meta-analyses. This research should use the most specific exposure units possible, such as player-hours, and should detail exact length of follow-up in months or years. Because sport season lengths can vary by location, age and participation level, detailing follow-up length this way is particularly important for standardising the period of time over which exposures are occurring.

CONCLUSION

Female football players had 2.2-fold increase in the IR of ACL injury relative to male football players over 4 years. The sex difference in ACL injury rates was independent of participation level.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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Inclusion and exclusion criteria

Inclusion criteria

- Research conducted in football
- Total number of ACL injuries and total number of individuals in the population by sex reported
- ▶ Reported data such that the number of ACL injuries by sex could be calculated.

Exclusion criteria

- Further analyses on previously reported studies
- Studies written in languages other than English
- Review articles.

What is known already known?

▶ In 2007, the incidence rate ratio (IRR) of ACL injuries in women compared with men in football was estimated to be 2.7.

What are the new findings?

- Female and male football players had similar ACL injury IP in a window that spanned one season and 4 years.
- ► The incidence rate of ACL injury among female football players over 4 years was 2.2 times higher than the incidence of ACL injury among male football players.
- The sex differences in ACL injury rates were independent of participation level.
- The IRR in football players has remained unchanged since 2007 (2.7 vs 2.5).

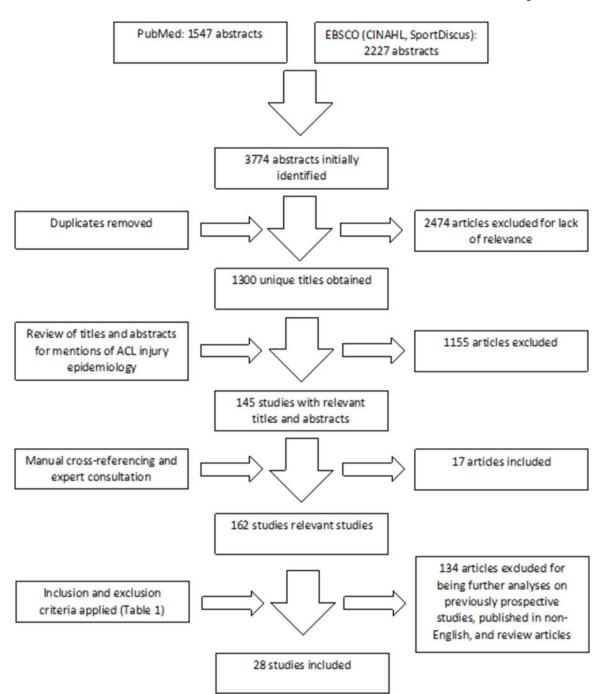


Figure 1. Flow chart summary of the literature search process.

Author	Total Number of ACL Injuries	Total Population			Proportion	95%-CI	Weight
Caraffa et al,1996	80	600		-	0.133	[0.107; 0.163]	4.9%
Faude et al,2005	11	165		10.0	0.067	[0.034; 0.116]	4.3%
Gilchrist et al,2008	25	1435	-		0.017	[0.011; 0.026]	5.1%
Giza et al,2005	8	202			0.040	[0.017; 0.077]	4.4%
Hagglund et al,2009	16	467			0.034	[0.020; 0.055]	4.9%
Heidt et al,2000	9	300			0.030	[0.014; 0.056]	4.7%
Kiani et al,2010	5	1506	+		0.003	[0.001; 0.008]	5.1%
Krutsch et al,2016	16	408	-		0.039	[0.023; 0.063]	4.8%
LeGall et al,2008	12	119			0.101	[0.053; 0.170]	4.0%
Mandelbaum et al,2005	73	5703	4		0.013	[0.010; 0.016]	5.2%
Ostenburg and Roos,2000	5	123	-	-	0.041	[0.013; 0.092]	4.0%
Padua et al,2015	7	829	-		0.008	[0.003; 0.017]	5.0%
Pfeiffer et al,2006	1	433	+-		0.002	[0.000; 0.013]	4.8%
Quisquater et al,2013	608	833396			0.001	[0.001; 0.001]	5.2%
Rahnama et al,2009	33	390	_	_	0.085	[0.059; 0.117]	4.8%
Soderman et al,2000	5	140	-		0.036	[0.012; 0.081]	4.2%
Steffen et al,2008	9	2020	+		0.004	[0.002; 0.008]	5.1%
Steffen et al,2016	29	447	-	-	0.065	[0.044; 0.092]	4.8%
Tegnander et al,2008	2	181	-		0.011	[0.001; 0.039]	4.4%
Walden et al,2011	21	4564	+		0.005	[0.003; 0.007]	5.2%
Walden et al,2012	35	962	-		0.036	[0.025; 0.050]	5.0%
Random effects model		854390	<u></u>		0.027	[0.016; 0.041]	100.0%
Heterogeneity: $I^2 = 98\%$, $\tau^2 =$: 0.0069, <i>p</i> < 0.01			1			
			0.05	0.1 0.15			
		Male	at Greater Risk	Female at Gre	ater Risk		

Figure 2. Forest plot for the incidence proportion and 95% CI of ACL injury in football players for men and women combined.

Author	Number of ACL Injuries F	emale Population	Proportion	95%-CI	Weight
Faude et al,2005	11	165	0.067	[0.034; 0.116]	4.8%
Gilchrist et al,2008	25	1435 —	0.017	[0.011; 0.026]	6.3%
Giza et al,2005	8	202	0.040	[0.017; 0.077]	5.0%
Hagglund et al,2009	8	228	0.035	[0.015; 0.068]	5.2%
Heidt et al,2000	9	300	0.030	[0.014; 0.056]	5.4%
Kiani et al,2010	5	1506 🖃	0.003	[0.001; 0.008]	6.3%
LeGall et al,2008	12	119	0.101	[0.053; 0.170]	4.3%
Mandelbaum et al,2005	73	5703	0.013	[0.010; 0.016]	6.5%
Ostenburg and Roos,2000	5	123	0.041	[0.013; 0.092]	4.4%
Padua et al,2015	6	481	0.012	[0.005; 0.027]	5.8%
Pfeiffer et al,2006	1	433 +	0.002	[0.000; 0.013]	5.8%
Quisquater et al,2013	29	37170	0.001	[0.001; 0.001]	6.6%
Soderman et al,2000	5	140	0.036	[0.012; 0.081]	4.6%
Steffen et al,2008	9	2020 🔄	0.004	[0.002; 0.008]	6.4%
Steffen et al,2016	29	447	0.065	[0.044; 0.092]	5.8%
Tegnander et al,2008	2	181	0.011	[0.001; 0.039]	4.9%
Walden et al,2011	21	4564	0.005	[0.003; 0.007]	6.5%
Walden et al,2012	15	310	0.048	[0.027; 0.079]	5.5%
Random effects model Heterogeneity: $I^2 = 97\%$, $\tau^2 =$	0.004, <i>p</i> < 0.01	55527 🔷	0.020	[0.012; 0.031]	100.0%

Figure 3. Forest plot for incidence proportion and 95% CI of ACL injury for female football players.

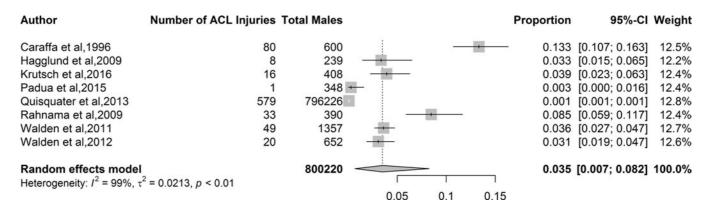


Figure 4. Forest plot for incidence proportion and 95% CI of ACL injury for male football players.

Author	Number of ACL Injuries	Famale	Number of ACL Injuries	Male Population	Risk Ratio	RR	95%-CI Weight
Author	Number of ACL injuries	Population	Number of ACL injuries	Population	RISK RATIO	KK	95%-Ci weight
Hagglund et al,2009	8	228	8	239		1.05	[0.40; 2.75] 10.0%
Padua et al,2015	6	481	1	348	 . 	-4.34	[0.52; 35.90] 2.1%
Quisquater et al,2013	29	37170	579	796226	-	1.07	[0.74; 1.56] 66.5%
Walden et al,2012	15	310	20	652	-	1.58	[0.82; 3.04] 21.5%
Random effects model Heterogeneity: $I^2 = 0\%$, τ^2		38189		797465	—	1.20	[0.88; 1.62] 100.0%
				Male at	0.1 0.5 1 2 10 Greater Risk Female at Gr	eater Ris	sk

Figure 5.Forest plot for RR and 95% CI of ACL injury for football players—woman versus man. <1=reduced RR in women; >1=reduced RR in men. RR, relative risk.

Author	Total Number of ACL Injuries	Total Athlete Exposure/10000	Incidence Rate	Rate	95%-CI	Weight
Agel et al,2016	97	136.0		0.71	[0.58; 0.87]	5.8%
Beynnon et al,2014	35	29.0		1.21	[0.87; 1.68]	5.3%
Faude et al,2005	11	1.8		6.11	[3.38; 11.03]	4.2%
Gilchrist et al,2008	25	8.8	-	2.84	[1.92; 4.20]	5.1%
Giza et al,2005	8	17.8		0.45	[0.22; 0.90]	3.7%
Gwinn et al,2000	53	21.5	-	2.47	[1.88; 3.23]	5.5%
Hagglund et al,2009	16	6.3	-	2.54	[1.56; 4.15]	4.6%
Hewett et al,1999	3	2.2	- 1	1.36	[0.44; 4.23]	2.3%
Hootman et al,2007	579	333.5		1.74	[1.60; 1.88]	6.0%
Joseph et al,2013	140	170.1		0.82	[0.70; 0.97]	5.8%
Kiani et al,2010	5	6.7	■	0.75	[0.31; 1.79]	3.1%
Krutsch et al,2016	16	7.5	 	2.13	[1.31; 3.48]	4.6%
LeGall et al,2008	12	4.8		2.50	[1.42; 4.40]	4.3%
Mandelbaum et al,2005	73	20.5	-	3.56	[2.83; 4.48]	5.7%
Mountcastle et al,2007	353	260.6		1.35	[1.22; 1.50]	6.0%
Ostenburg and Roos,2000	3	0.5		6.00	[1.94; 18.60]	2.3%
Pfeiffer et al,2006	1	1.5	*		[0.09; 4.73]	1.0%
Soderman et al,2000	5	0.7	- 8	7.14	[2.97; 17.16]	3.1%
Stanley et al,2016	157	165.3	•	0.95	[0.81; 1.11]	5.9%
Steffen et al,2008	9	6.7		1.34	[0.70; 2.58]	3.9%
Tegnander et al,2008	2	1.5		1.33	[0.33; 5.33]	1.7%
Walden et al,2011	21	13.9	*	1.51	[0.99; 2.32]	4.9%
Walden et al,2012	35	21.7	#	1.61	[1.16; 2.25]	5.3%
Random effects model				1.69	[1.36; 2.10]	100.0%
Heterogeneity: $I^2 = 93\%$, $\tau^2 =$	= 0.2037, p < 0.01				a 6 T	
			5 10 15			
		Male	at Greater Risk Female at Gre	ater Ris	sk	

Figure 6. Forest plot for total incidence rate and 95% CI of ACL injury in football players for men and women combined.

Author	Number of Female ACL Injuries	Athlete Exposure/10000	Incidence Rate	Rate	95%-CI	Weight
Agel et al,2016	71	71.0	+	1.00	[0.79; 1.26]	5.9%
Beynnon et al,2014	26	14.2	*	1.83	[1.25; 2.69]	5.4%
Faude et al,2005	11	1.8		6.11	[3.38; 11.03]	4.6%
Gilchrist et al,2008	25	8.8	-	2.84	[1.92; 4.20]	5.4%
Giza et al,2005	8	17.8		0.45	[0.22; 0.90]	4.3%
Gwinn et al,2000	22	4.0		5.50	[3.62; 8.35]	5.3%
Hagglund et al,2009	8 2	2.7	-	2.96	[1.48; 5.92]	4.3%
Hewett et al,1999	2	1.4	-10	1.43	[0.36; 5.71]	2.2%
Hootman et al,2007	411	146.8	•	2.80	[2.54; 3.08]	6.1%
Joseph et al,2013	96	78.6	+	1.22	[1.00; 1.49]	6.0%
Kiani et al,2010	5	6.7	*	0.75	[0.31; 1.79]	3.6%
LeGall et al,2008	12	4.8		2.50	[1.42; 4.40]	4.7%
Mandelbaum et al,2005	73	20.5	-	3.56	[2.83; 4.48]	5.9%
Mountcastle et al,2007	51	40.9	-	1.25	[0.95; 1.64]	5.8%
Ostenburg and Roos,2000	3	0.5		— 6.00	[1.94; 18.60]	2.8%
Pfeiffer et al,2006	1	1.5	*	0.67	[0.09; 4.73]	1.3%
Soderman et al,2000	5	0.7		7.14	[2.97; 17.16]	3.6%
Stanley et al,2016	99	70.2		1.41	[1.16; 1.72]	6.0%
Steffen et al,2008	9	6.7	-	1.34	[0.70; 2.58]	4.4%
Tegnander et al,2008	2	1.5		1.33	[0.33; 5.33]	2.2%
Walden et al,2011	21	13.9	-	1.51	[0.99; 2.32]	5.3%
Walden et al,2012	15	5.2	-	2.88	[1.74; 4.78]	5.0%
Random effects model			♦	2.03	[1.57; 2.62]	100.0%
Heterogeneity: $I^2 = 91\%$, $\tau^2 =$: 0.2736, <i>p</i> < 0.01			•	,	
			5 10 15			

Figure 7. Forest plot for incidence rate and 95% CI of ACL injury for female football players.

Author	Number of Male ACL Injuries	Athlete Exposure/10000	Incidence Rate	Rate	95%-CI	Weight
Agel et al,2016	26	65.0	=	0.40	[0.27; 0.59]	9.9%
Beynnon et al,2014	9	14.7		0.61	[0.32; 1.18]	7.7%
Gwinn et al,2000	31	17.6	-	1.76	[1.24; 2.50]	10.2%
Hagglund et al,2009	8	3.6	- 10	2.22	[1.11; 4.44]	7.4%
Hewett et al, 1999	1	0.9		- 1.11	[0.16; 7.89]	2.1%
Hootman et al,2007	168	186.7		0.90	[0.77; 1.05]	11.4%
Joseph et al,2013	44	91.5	-	0.48	[0.36; 0.65]	10.6%
Krutsch et al,2016	16	7.5	-	2.13	[1.31; 3.48]	9.1%
Mountcastle et al,2007	302	219.7	+	1.37	[1.23; 1.54]	11.5%
Stanley et al,2016	58	95.1		0.61	[0.47; 0.79]	10.8%
Walden et al,2012	20	16.5	-	1.21	[0.78; 1.88]	9.5%
Random effects model Heterogeneity: I ² = 92%,	T-2		\(\rightarrow\)	0.97	[0.71; 1.33]	100.0%
			1 2 3 4 5 6 7			

Figure 8. Forest plot for incidence rate and 95% CI of ACL injury for male football players.

Author	Number of ACL Injuries	Famale Athlete Exposure/10,000		Male Athlete Exposure/10,000	Incidence Rate Ratio	IRR	95%-CI	Weight
Agel et al,2016	71	71.0		65.0	-101		[1.60; 3.92]	11.5%
Beynnon et al,2014	26	14.2	9	14.7		2.99	[1.40; 6.38]	8.3%
Gwinn et al,2000	22	4.0	31	17.6		3.12	[1.81; 5.39]	10.5%
Hagglund et al, 2009	8	2.7	8	3.6		1.33	[0.50; 3.55]	6.5%
Hewett et al, 1999	2	1.4	1	0.9		1.29	[0.12; 14.18]	1.7%
Hootman et al,2007	411	146.8	168	186.7		3.11	[2.60; 3.72]	13.9%
Joseph et al,2013	96	78.6	44	91.5		2.54	[1.78; 3.63]	12.5%
Mountcastle et al,2007	51	40.9	302	219.7	*	0.91	[0.67; 1.22]	13.0%
Stanley et al,2016	99	70.2	58	95.1	-	2.31	[1.67; 3.20]	12.8%
Walden et al,2012	15	5.2	20	16.5		2.38	[1.22; 4.65]	9.2%
Random effects model Heterogeneity: $I^2 = 83\%$, τ					<u> </u>	2.20	[1.57; 3.08]	100.0%
					0.1 0.5 1 2 10			
				Male a	at Greater Risk Female at Grea	ater Ris	sk	

Figure 9. Forest plot for incidence rate and 95% CI of ACL injury for football players—woman versus man. <1=reduced relative risk in women;>1=reduced relative risk in men. IRR, incidence rate ratio.

Author I	Number of ACL Injuries	Famale Athlete Exposure/10,000	Number of ACL Injuries A	Male Athlete Exposure/10,000	Incidence Rate Ratio	IRR	95%-CI Weight
Level = Amateur Beynnon et al,2014 Gwinn et al,2000 Hewett et al,1999 Joseph et al,2013 Mountcastle et al,2007 Random effects model Heterogeneity: I ² = 27%, 2 ²	15 2 2 2 96 1	0.1 1.4		11.7 2.5 0.9 91.5 9.0		5.00 [1.29 [2.54 [0.40 [1.49; 17.73] 2.4% 1.10; 22.82] 1.6% 0.12; 14.18] 0.7% 1.78; 3.63] 20.5% 0.05; 3.06] 0.9% 1.41; 4.81] 26.2%
Level = Intermediate Agel et al,2016 Beynnon et al,2014 Gwinn et al,2000 Hootman et al,2007 Mountcastle et al,2007 Random effects model Heterogeneity: I² = 7%, x² =	71 11 5 411 4	71.0 2.8 0.7 146.8 2.3	6 1 168	65.0 3.0 1.2 186.7 3.4	-	1.96 [[1.60; 3.92] 14.7% (0.73; 5.31] 3.7% (1.00; 73.37] 0.8% (2.60; 3.72] 40.9% (0.32; 4.40] 2.2% (2.35; 3.55] 62.4%
Level = Elite Hagglund et al,2009 Walden et al,2012 Random effects model Heterogeneity: $l^2 = 0\%$, $\tau^2 =$ Random effects model Heterogeneity: $l^2 = 14\%$, τ^2			8 20	3.6 16.5	•	2.38 [1.98 [0.50; 3.55] 3.8% (1.22; 4.65] 7.6% 1.14; 3.44] 11.5% 2.17; 3.23] 100.0%
				Male a	0.1 0.5 1 2 10 t Greater Risk Female at G	Greater Ris	k

Figure 10. Forest plot for incidence rate and 95% CI of ACL injury in football players—woman versus man by participation level. <1=reduced relative risk in women; >1=reduced relative risk in men. IRR, incidence rate ratio.

Study	Experiment Events Tin	al ne Events	Control Time	Ir	cidence Rate Ratio	IRR	95%-CI	Weight (fixed)	Weight (random)
After2007 = 2 Agel et al Gwinn et al Joseph et al Stanley et al Walden et al Fixed effect model Random effects model Heterogeneity: I ² = 0%, τ ²		50 31 80 44 77 58	65.00000 17.56280 91.45510 95.08197 16.49230			3.16 2.54 2.31 2.36 2.49	[1.60; 3.92] [1.83; 5.45] [1.78; 3.63] [1.67; 3.20] [1.21; 4.61] [2.06; 3.01] [2.07; 3.02]	6.8% 2.9% 10.2% 12.4% 2.4% 34.7%	11.5% 10.5% 12.5% 12.8% 9.2% 56.5%
After2007 = 1 Beynnon et al Hagglund et al Hewett et al Hootman et al Mountcastle et al Fixed effect model Random effects model Heterogeneity: 1 ² = 92%, x		80 8 40 1 71 168 1 80 302 2	14.73810 3.56805 0.85130 186.66667 219.68120	_		1.32 1.26 3.11 0.91 2.25	[1.40; 6.39] [0.49; 3.51] [0.11; 13.87] [2.60; 3.72] [0.67; 1.22] [1.95; 2.59] [0.83; 3.85]	2.2% 1.7% 0.3% 37.2% 23.8% 65.3%	8.3% 6.5% 1.7% 13.9% 13.0% — 43.5%
Fixed effect model Random effects model Heterogeneity: I ² = 83%, τ				0.1	0.5 1 2		[2.08; 2.61] [1.57; 3.07]	100.0%	100.0%

Figure 11. Forest plot for incidence rate and 95% CI of ACL injury in football players—woman versus man by year of publication. <1=reduced relative risk in women; >1=reduced relative risk in men. IRR, incidence rate ratio.