Systematic Review

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Anterior cruciate ligament allografts and low-dose gamma irradiation: a systematic review

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ABSTRACT

Anterior cruciate ligament (ACL) rupture is a common injury requiring surgical reconstruction in active patients. Autografts are considered the gold standard due to superior early biomechanical performance, but allografts offer advantages such as reduced operative time and elimination of donor-site morbidity. Irradiation is the primary sterilisation method; however, high doses impair mechanical integrity. Low-dose irradiation has been proposed to preserve graft strength while ensuring sterility. A systematic review was performed using EMBASE, PubMed and Scopus, supplemented with backward snowballing. Cohort and randomised studies comparing low-dose irradiated allografts to autografts in vivo were included. Primary outcomes were graft failure/revision rates, knee laxity (KT-1000/2000, Lachman, pivot shift) and patient-reported measures (IKDC, Lysholm). Eight studies comprising over 10,000 patients were included. Autografts demonstrated lower graft failure rates in younger (<22 years) patients, with revision rates of 10.1% for allografts vs. 2.9% for autografts. In older cohorts, outcomes were largely equivalent. Arthrometric testing showed greater knee laxity with allografts, but this did not consistently translate into differences in patient-reported outcomes. Return-to-sport was faster with autografts, reflecting earlier graft incorporation, while long-term outcomes converged once allografts remodelled. Autografts remain the graft of choice for younger, highdemand patients due to superior early stability and lower revision rates. Low-dose irradiated allografts may represent a viable alternative in older or less active populations. High-quality, prospective studies with standardised protocols are required to clarify long-term efficacy and guide graft selection.

Keywords: ACL, Allograft, Gamma radiation, Low-dose

INTRODUCTION

Anatomy and micro-anatomy of anterior cruciate ligament

The ACL is a key stabilizer of the knee joint, resisting anterior tibial translation and rotational loads.^{1,2} Structurally, it comprises an anteromedial (AM) and a posterolateral (PL) bundle, with functional variation across the flexion-extension range.³ The ACL is predominantly made of type I collagen, with regional specialisation in collagen type and proteoglycan content

aiding in its viscoelastic and load-bearing properties as seen in Table 1.⁴ Its hierarchical microstructure, including crimped collagen fibrils and fascicles within vascularized connective tissue, enables it to deform under stress and protect the joint.⁵ Injury disrupts this architecture, with restoration of crimp structure being critical for functional recovery.⁶

Anterior cruciate ligament injury and grafts

ACL rupture is the most common knee ligament injury, with an incidence of 14 per 100,000 annually in the UK,

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rising sharply among athletes. 9,10 Surgical reconstruction is typically indicated for active patients with full ruptures or symptomatic instability. Graft choices include autografts, bone-patellar tendon-bone (BPTB), hamstring or quadriceps tendon and allografts from cadaveric tissue.11 Autografts are commonly used but may result in harvest-site morbidity.¹² The incidence of anterior knee pain is influenced by patient characteristics and rehabilitation as well as the type of graft.¹³ Surgically, grafts are fixed in bone tunnels drilled in the femur and tibia, with fixation achieved via interference screws, cortical suspensory devices or cross-pins depending on graft type and surgeon preference. Anatomical singlebundle and double-bundle reconstructions are both practiced, the latter aiming to replicate native AM and PL bundle functions more closely albeit double-bundle reconstructions are quoted in the literature to be less than 2% of all reconstructions. 14 Allografts are typically reserved for older, multi-revision patients to avoid extended surgical time and harvest complications or younger athletes whose performance may be impacted by harvest-site morbidity. 15,16

proceeds Graft healing through inflammation. revascularization and remodeling. In bone-including grafts like BPTB, healing occurs via bone ingrowth at the graftbone interface.¹⁷ In contrast, soft-tissue grafts require fibrovascular integration, a slower process influenced by graft type and mechanical loading. Despite surgical advances, there is a lack of consensus on evidence-based post-operative timelines for healing and return to sport. 18 Outcomes are evaluated via stability tests (Lachman, pivot shift, KT-1000), graft failure rates and validated patientreported outcome measures. According to these outcomes, autografts appear to have marginally better outcomes compared to allografts although the clinical significance of this is debatable.19

Anterior cruciate ligament allografts

Allografts are typically sterilized with irradiation and ethylene glycol in order to prevent disease transmission or an immune response by the graft-receiving patient. Although allografts offer shorter operative times, faster recovery and no risk of donor-site morbidity, there are questions over their functional suitability after this sterilization. Conrad et al found that irradiated tendons had lower load to failure, lower stiffness and lower Young's Modulus when compared to non-irradiated 'control' tendons in an in-vitro study.²⁰ A 2013 meta-analysis of prospective studies also found that allografts were associated with higher rates of graft failure than autografts although functional outcomes in non-failure cases were similar.²¹

This suggests a link between allograft sterilisation and their observed loss of mechanical capabilities. Haut and Powlison (1990) has shown that irradiation can reduce the mechanical strength of collagen by cleaving its peptide bonds along the polypeptide chain, although historically 3-

4 Mrad of radiation were typically used. The disruption of these peptide bonds will result in lower tensile strength and lower load to failure.²² The extent of disruption to the collagen peptide bonds is dose-dependent.²³

Nonirradiated ACL allografts have shown no significant differences to autografts in 'functional outcomes', including levels of activity, stiffness and pain.²⁴ However, they are not first line due to increased infection risk from the absence of irradiation sterilisation.²⁵ Although the rates of infection after ACL repair are reportedly low, severe complications such as septic arthritis support use of precautions. Preventing infection avoids delayed return to activity, graft removal and revision surgery, as well as inferior functional long-term outcomes.²⁶ Irradiated allografts have been associated with compromised mechanical stability and loss of tensile strength in ex-vivo studies involving both ovine and human models.^{27,28} Lowtemperature, low dose gamma irradiation can inactivate enveloped and nonenveloped DNA and RNA, allowing for effective sterilisation whilst partially preserving the mechanical properties of the allograft and as such have been selected as the focus for this review.²⁹

Tejwani et al, retrospectively reported graft failure rates of ACL reconstructions performed with different dosage irradiated allografts (Table 2).30 A dose-dependent relationship was established between revision rates of allografts and irradiation sterilisation. Furthermore, no differences were found between soft tissue grafts such as tibialis posterior or hamstring tendons and BPTB grafts. In the case of non-irradiated patellar tendon allografts, there is insufficient evidence to suggest that mechanically, they are inferior to BPTB autografts.^{21,31} Thus, leading to the consideration of low dose irradiation for sterilisation as a method of preserving the biomechanical properties of the graft; reducing the risk of graft failure in allografts. It is hypothesized that use of low-dose gamma irradiation for allografts can produce the same mechanical outcomes as non-irradiated allografts due to preservation of the grafts' biomechanical properties. However, infection risk and/or re-injury, surgery waiting time and quality of life must all be considered when selecting a graft.

Aims and objectives

This project aims to evaluate the clinical efficacy of low-dose irradiated allografts for ACL reconstruction in vivo. Low-dose irradiated allograft outcomes will be compared to those of the current gold standard—autografts.

METHODS

Search strategy

A comprehensive search of EMBASE, PUBMED and SCOPUS was performed using the terms 'allograft AND (ACL repair OR ACL reconstruction OR) AND (gamma radiation OR low dose radiation) AND autograft'. Due to the lack of MeSH terms surrounding low dose radiation or

gamma radiation only non-MeSH terms were used. Appendix 1 demonstrates the generation of search results through each search term in the included databases. Searches conducted from inception to January 2025. Appendix 1 demonstrates the generation of search results through each search term in the included databases.

Studies were included on the condition the full-text paper was available in the English language and data sets were available for review. Abstracts were then screened to ensure they were cohort or randomized studies on humans and were in-vivo. Abstracts were also reviewed to confirm comparison was between allograft and autograft controls. Backward snowballing from review papers was used to identify papers that were missed through the primary search. The eligible studies were then de-duplicated using reference software Endnote.

Study selection

Following the first-pass search, the papers deemed eligible then underwent a full-text review according to the inclusion and exclusion criteria (Table 3). In order to be included, the paper methodology had to be a cohort or randomized control trial comparing allografts to autografts. Outcome measures had to be quantitative and reproducible variables, hence any studies with patient reported outcomes only were excluded. The studies left were included in the report. In this review we followed PRISMA 2020 to ensure transparent reporting: we completed the 27-item checklist (covering rationale, search methods, eligibility criteria, selection, risk of bias, synthesis and certainty) and included the PRISMA 2020 flow diagram (Figure 2) to document records identified, screened, excluded and included.

Quality assessment

The critical appraisal skills programme (CASP) tool for assessing cohort and randomized studies was used to measure the quality of the included studies (Table 4). Variables from each study were compared for autografts vs. low-dose irradiated allografts including graft failure/revision surgery rates, arthrometer score of knee joint laxity and pivot shift measurements. International Knee Documentation Committee Subjective Knee Form (IKDC) scores were also reviewed. Included studies were further reviewed by two independent reviewers and if considered appropriate were selected for in-depth analysis in the literature analysis section of this report.

Evaluation of findings

The findings from each study were presented in table 4 in order to compare variables including graft failure/revision rates, arthrometer scores of knee joint laxity and pivot shift measurements. The significances of the findings were summarised in the context of autografts vs. low-dose irradiated allografts. International Knee Documentation Committee Subjective Knee Form (IKDC) scores were

also considered in included studies which also provided other objective outcomes. Five studies were selected for in-depth analysis in the literature analysis section of this report (Table 5).

Following a search of the three databases and accumulation of papers identified through systematic reviews, the inclusion and exclusion criteria were applied as summarised in Figure 2.

Table 4 presents papers included in this study following the literature search with the addition of Tejwani et al, which has been selected as a seminal paper in comparing the effect of gamma irradiation on allografts at different dosages over a follow-up period of 2-years.³⁰ No meta-analysis was performed due to heterogeneity in irradiation doses and follow-up of the included studies and instead, a qualitative synthesis was performed.

RESULTS

The results of the literature search, alongside a summary of each included study's findings is presented in table 4. The bias concerns for each paper have been identified and discussed.

Graft failure rate

Graft failure rates vary across studies due to differing definitions and low incidence in follow-up cohorts. While young, high-activity patients exhibit higher failure rates (~7%), general population risk remains low, limiting the utility of graft rupture as a sole outcome measure. ^{30,32} Mechanical assessments offer a more consistent comparison. Ghodadra et al used Lachman, pivot shift and KT-1000 arthrometry, finding no significant differences in knee laxity between autografts and low-dose irradiated allografts over a one-year follow-up. ³³ This was the case when 2.5 Mrad or lower radiation was used, suggesting that the dose-dependent collagen damage is perhaps not relevant in doses under 2.5 Mrad.

However, grouping processed and unprocessed allografts may obscure subtle differences due to omitted variable bias. Objective measures like the KT-1000 reduce examiner variability and outperform clinical tests such as the Lachman (which loses post-reconstruction sensitivity) and the pivot shift (which has high specificity but moderate sensitivity). Despite mixed evidence, low-dose irradiated allografts appear biomechanically comparable to autografts, supporting their broader use across age groups.

Age as a factor

While small studies lack power to assess graft failure rates meaningfully, Maletis et al analysed 5,586 ACL reconstructions and found higher revision rates in allografts (3.6%) compared to autografts (1.9%), especially in patients under 22 years old.⁴⁴ In contrast, outcomes in patients over 22 showed no significant

difference between low-dose irradiated allografts and autografts, aligning with Hulet et al, who reported superior allograft performance in patients over 25 (Figure 3).³⁴

Age bias complicates interpretation, only 18.4% of the allograft group were under 22, versus 49.9% in the autograft group, suggesting clinician selection influenced by known age-dependent outcomes. Despite its strengths, the Maletis study lacked control over variables such as surgical timing and rehab protocols. Additionally, cost and logistical challenges of allografts, may limit their feasibility, especially in resource-constrained settings. Nevertheless, the data support cautious use of allografts in older adults while reaffirming autografts as the preferred option in younger, more active patients. Clinician selection bias may overestimate allograft risks in young patients; propensity matching in future studies could address this.

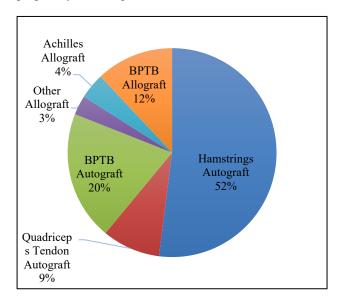


Figure 1: Proportion of different ACL graft types used currently in the UK.⁴⁶

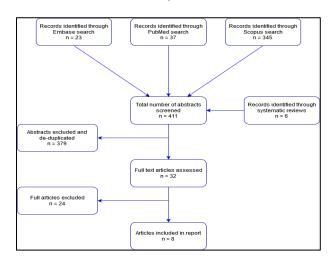
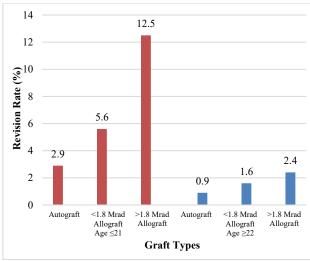


Figure 2: A flow-chart representing the study inclusion process based on preferred reporting items for systematic reviews and meta-analyses (PRISMA) guidelines.⁴⁷



*Statistically significant differences in values have been marked.

Figure 3: Reported revision rates (%) of different graft types by stratified by age. 44

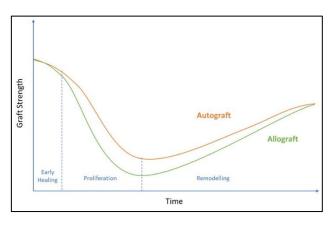


Figure 4: Different proposed timelines for graft healing in autografts and low-dose irradiated allografts.³¹

Are patient-reported outcome scores useful in measuring Anterior cruciate ligament graft success

In a five-year prospective RCT, Li et al found greater knee laxity in low-dose irradiated allografts (3.5 mm) compared to autografts (2.1 mm) via KT-1000 measurements, though Lachman and pivot shift tests showed no difference.³⁶ These findings were corroborated by Sun et al, Objective differences did not translate into statistically significant disparities in patient-reported outcome measures (PROMs).^{24,37}

In Li et al no significant differences were found in Lysholm scores, Tegner activity levels or IKDC scores between graft types over a five-year follow-up.³⁶ This suggests that while biomechanical superiority exists, the functional experiences of patients may not differ appreciably, a finding with important implications for graft selection in routine clinical practice. If subjective function

is preserved, the relevance of small differences in laxity may be less impactful for certain patient populations.

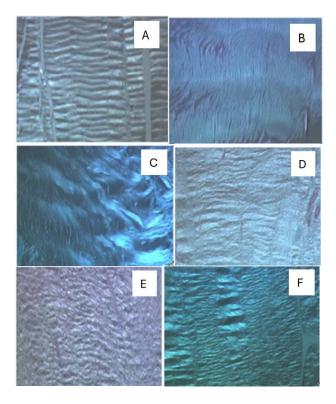


Figure 5: Light microscopy of ACL fibres and grafts during healing.

Graft choice after primary reconstruction failure

The MARS cohort, comprising 1205 patients and 87 surgeons, reported improvements in IKDC scores following revision ACL surgery in both autograft and low-dose irradiated allograft groups.³⁸ However, autograft recipients showed greater functional gains and there was a higher, though underpowered, re-rupture risk in allograft patients. No differences were noted between BPTB and

soft tissue grafts within each graft type. Selection bias likely influenced results, as patients with prior autograft failure were more often assigned to the allograft group, particularly athletes who inherently face higher re-rupture risk.³⁹ The 2016 MARS analysis further noted surgeon preference as the strongest predictor of revision outcomes, though patient perceptions and prior graft dissatisfaction often influenced graft choice. Despite limitations such as non-randomization and short follow-up, findings suggest autografts may offer superior outcomes in revision ACL surgery, especially in younger, high-demand individuals.

Does the post-operative recovery time differ between allografts and autografts

Rihn et al conducted a four-year retrospective study comparing ACL reconstructions using autografts and low-dose irradiated allografts.⁴⁰ While knee laxity and IKDC scores were statistically similar between groups, autograft patients returned to weight-bearing one week earlier and resumed running one month sooner than those with allografts. This supports a consistent pattern across the literature autografts facilitate faster return to physical activity, a factor not fully captured by traditional outcome scores.⁴¹ Biological differences in graft remodeling help explain this disparity. Autografts undergo remodeling more quickly (6-12 months), while allografts show delayed tissue integration (11-18 months).⁴²

Although allografts may regain strength over a longer period, their slower adaptation limits early functional gains (Figure 4).³¹ These findings suggest that short- to mid-term recovery is generally superior with autografts, even if long-term mechanical properties converge. However, disparity in rehabilitation protocols must be considered between the studies. While autografts enable faster early return a meta-analysis suggests higher long-term preinjury activity restoration with allografts, possibly due to reduced donor morbidity.^{40,43}

Table 1: Comparing ligament and tendon composition, adapted from Amis and Marieswaran et al.^{7,8}

Component	Ligament (%)	Tendon (%)
Cellular material	20% (fibroblasts)	20 (tenocytes–fibroblast-like cells)
Collagen	70%	80
Type I collagen	90%	99
Type III collagen	10%	1
Elastin	Abundant	Scarce
Proteoglycans and GAGs	5%	<2

Table 2: Summary of ACL allograft revision rates for each dosage group reported in Tejwani et al.³⁰

Irradiation dosage (Mrad)	Number of cases	Revision rate (%)
None	1185	2.28
<1.2	726	2.89
1.2-1.8	2911	2.23
>1.8	1146	3.75*

^{*}Statistical significance when compared to the non-irradiated control group.

Table 3: A summary of the inclusion and exclusion criteria, including their justification, for identifying suitable papers in this report.

Exclusion criteria	Inclusion criteria	Justification of criteria		
No controls	Autografts of the same graft type as control	This project aims to evaluate the efficacy of allografts that have been prepared using low dose gamma irradiation. Thus, only papers which include comparisons against the current gold standard for ACL repairs, autografts, were considered for this project.		
No full-text paper available	Full-text paper available in English			
No data available	Data sets available for review	For the purposes of evaluation of the findings of each included study, studies without		
Case-studies and letters to editors	Cohort or randomised studies	the full-text available and those without their data published were also excluded.		
Animal studies	Human studies only	Furthermore, only human studies performed in vivo were considered in order for the		
In vitro studies	In vivo studies	findings of this project to have inferences for clinical practice.		
Outcomes measured with only patient reported data	Outcomes measured with quantitative and reproducible variables	Furthermore, due to lack of reliability of self-reported patient data especially in studies without randomised blinding, this project excluded any studies without objective and quantitative data as their measurable outcome.		

Table 4: Characteristics of the 8 included papers from the literature search and backward snowballing, including study design, number of patients, outcomes measured and definitions for low-dose irradiated allografts. The key findings of each paper are summarised along with their significance in this report.

Reference	Study type (follow-up time)	No. of patients	Irradiation dosage & graft types	Outcome assessed	Key findings	Significance of findings	Bias concerns
Maletis et al ⁴⁴	Retro- spective cohort (2 years)	5586	Unprocessed, <1.8- Mrad and ≥1.8- Mrad BPTB	Aseptic revision	There were 37 (3.6%) revisions in BPTB allograft cases and 85 (1.9%) in BPTB autograft cases. The results varied significantly for age categories >22 and =21. In over 22-year olds there was no difference in revision rates for autografts and allografts which received <1.8 Mrad. In the 21-year-old or younger patients, there was a stark difference in revision rates from 2.9% to 10.1% in autografts and allografts with low dose gamma radiation respectively.</th <th>This study supports the idea that autografts are mechanically superior <i>in vivo</i> to allografts. However, it raises suspicion over the effects of age on the healing of allografts. This being a retrospective study, times between injury and surgery were not standardised and thus a prospective study of similar size would be beneficial to the literature.</th> <th>High risk of bias from age differences between each group's population. High risk of bias from disproportionate representation of allograft and autograft groups.</th>	This study supports the idea that autografts are mechanically superior <i>in vivo</i> to allografts. However, it raises suspicion over the effects of age on the healing of allografts. This being a retrospective study, times between injury and surgery were not standardised and thus a prospective study of similar size would be beneficial to the literature.	High risk of bias from age differences between each group's population. High risk of bias from disproportionate representation of allograft and autograft groups.
MARS cohort ³⁸	Prospective cohort (2 years)	1205	Unprocessed, <1.8Mrad Various Grafts	Re-rupture rates	This study analysed re-rupture rates in repeat ACL repairs. It was found that patients with an	This study suggests autograft use in repeat ACL reconstructions shows lower	Despite being a prospective study, there is no

Continued.

Refere	nce	Study type (follow-up time)	No. of patients	Irradiation dosage & graft types	Outcome assessed	Key findings	Significance of findings	Bias concerns
					IKDC scores	autograft revision were found to be 2.78 times less likely of sustaining a subsequent graft rupture compared with subjects who received an allograft. However, patient reported outcomes such as activity or pain were inconsistent with contrasting results using one scoring tool versus another.	risk of graft rupture. This study is unique in its investigation of repeat reconstructions. The data on individual graft types is not given however and low failure rates meant the power to detect a difference was low. As such, failure rates reported are unreliable.	randomisation. This may introduce some bias as the previous reconstruction's graft type may have influenced graft choice in this study.
Li et a	36 ∗	Prospective Randomised (5 years)	102	2.5 Mrad Various Grafts	Knee joint laxity Pivot shift IKDC scores	No significant difference was found in Lachman test and pivot shift tests between allograft and autograft groups. The same was not the case with arthrometer readings and autograft patients showed more stable knee joints. Patient reported scores were not statistically different. Graft failure rates were not reported due to the small N number of the study.	Despite the arthrometer reading suggesting autografts were superior, this did not translate to functional differences in the context of patient experience. This suggests autograft and allograft repairs functioned equally in the context of patient experience. A larger N number is needed with a similar follow-up time in order to comment on failure rates.	Moderate risk of bias due to this being a single surgeon study.
Ghoda al ³³ *	dra et	Retrospectiv e Cohort (1 year)	238	Unprocessed, 1- 1.3Mrad PT	Re-rupture rates Lachman's test Pivot shift Arthromete r measure	Laxity measured by arthrometer did not increase after the 6-week initial testing for either allograft and autograft group. There were no significant differences found between each of the groups in arthrometer, pivot shift and Lachman's test measurements. No differences between the unprocessed and low dose gamma irradiated grafts were found.	Found no differences between low dose irradiated allografts and autografts across 3 different mechanical testing modalities. However, they first compared the unprocessed allograft to the low dose allograft groups and found no significant difference, before then comparing a combined allograft group to an autograft group. This can	Moderate risk of bias due to single surgeon and single examiner being used for reconstruction and further testing.

Continued.

Reference	Study type (follow-up time)	No. of patients	Irradiation dosage & graft types	Outcome assessed	Key findings	Significance of findings	Bias concerns
						potentially skew the statistical tests.	
Tejwani et al ³⁰	Retrospectiv e Cohort (2 years)	5968	Unprocessed, <1.2, 1.2-1.8, >1.8 Mrad Various Grafts	Revision rates	This study investigated the effects of different dosages of allografts with each other, as well as the effects of other processing methods for allografts including chemical preparation. It found that within the first year of ACL repair, non-irradiated, low dose irradiated and high dose irradiated allografts had no significant differences in graft failure rates. However, after a year, low dose irradiated showed a lower risk for failure than the higher dose.	This study comments solely on non-irradiated vs. low dose vs. high dose irradiated allografts. Despite the longer follow-up time and large N number, there was no randomisation of groups and no standardisation of graft types. As only revision rates were considered, graft failures which did not lead to revision surgery were not considered in the results.	High risk of bias due to the non-randomised and non-standardised variables in each ACL repair recorded.
Rihn et al ⁴⁰ *	Retrospectiv e Cohort (4 years)	102	2.5 Mrad BPTB	Activity level Knee joint laxity IKDC scores Radiograph findings	This study found there to be no statistical significance between knee joint laxity of allograft and autograft treated patients when adjusted for age. There was statistically significant differences in the raw data before age adjusting. There were no significant differences for patient reported IKDC scores and return to activity scores between the two groups.	This study demonstrates the ability of low dose sterilised allografts to produce the same clinical outcomes as autografts in ACL repairs. The patient reported and objectively measured factors were statistically indifferent. Although it must be considered that this is with age adjusted results and raw data showed statistical differences between the groups.	Moderate risk of bias from age differences between each group's population and time from injury to reconstruction. Due to retrospective data, there are no comparisons to laxity before graft repair.
Sun et al ²⁴	Prospective Randomised (2.5 years)	99	2.5 Mrad BPTB	Re-rupture rates Arthromete r Pivot shift IKDC scores	This study found there to be no functional differences in the knees of allograft vs. autograft treated patients including range of motion, vertical jump and IKDC scores. However, they found statistical differences in	This study suggested that short term clinical outcomes are affected in low dose irradiated allografts compared to autografts. The higher rates of failure for the allograft group found in this	Moderate risk of bias due to this being a single surgeon study. These studies were carried out by the same group.

Continued.

Reference	Study type (follow-up time)	No. of patients	Irradiation dosage & graft types	Outcome assessed	Key findings	Significance of findings	Bias concerns
					knee laxity for side-to-side differences in anterior tibial displacement, pivot shift and Lachman test.	study supported the choice of autografts in ACL repairs.	
Sun et al ³⁷	Prospective Randomised (2.5 years)	67	2.5 Mrad Hamstring Tendon	Arthromete r Pivot shift IKDC scores	This study found the rate of laxity to be higher in allograft treated knees than in the autograft group. There was significantly higher rotational instability in the allograft group than in the autograft. However, there were no functional differences reported by patients in IKDC scores and activity testing.	This study suggested that short term clinical outcomes are affected in low dose irradiated allografts compared to autografts.	

^{*}Papers will be discussed in further detail. All irradiation dosages have been converted to Mrad units for ease of comparison. One rad is defined as 0.01 Joules of energy absorbed by 1kg of tissue, 1 Mrad is equivalent to 1 million of the rad unit. Bias concerns for each paper have been categorised as low (green), moderate (amber) and high (red).

Table 5: List of studies selected for further analysis and justifications for their selection.

Study	Justification for in-depth analysis
Maletis et al ⁴⁴	This study was chosen due to its large sample size allowing for reporting of graft failure rates with adequate statistical power. Furthermore, data provided in this study was stratified by age and allowed discussion of agerelated outcomes of ACL autografts compared to low-dose irradiated allografts.
MARS cohort ³⁸	This study was the only one included in this report which considered revision surgery following an initial graft failure. It included prospective data shared by 52 clinical centres which makes the findings of this study applicable for a wider population across ages and levels of activity.
Li et al ³⁶	This study was selected due to its long follow-up time and variety of measured outcomes. These included both objective mechanical testing and patient reported scores to evaluate the success of autografts compared to low-dose irradiated allografts.
Ghodadra et al ³³	This study was selected due to its incorporation of both graft failure and mechanical testing of a graft post- operatively as outcome measures. This allowed for an evaluation of graft rupture as a suitable measure of graft success.
Rihn et al ⁴⁰	This study was selected as it was the only study included in this report to look at rates of return to physical activity as an outcome measure. Its long follow-up period and use of both patient-reported factors and mechanical testing add context to its findings regarding return to activity rates in autografts compared to low-dose irradiated allografts.

Table 6: Summary of laxity measurements defined as side-to-side difference of anterior tibial displacement of the graft knee compared to the contralateral knee across 3 studies which used KT-1000 and KT-2000 arthrometry.

C4J	Laxity measured (mm)	Laxity measured (mm)				
Study	Autograft	Low-dose irradiated allograft				
Li et al ³⁶ *	2.1±1.6	3.5±1.2				
Sun et al ²⁴ *	2.4±0.6	5.5±3.6				
Sun et al ³⁷ *	2.3±1.1	3.6±2.8				
Rihn et al ⁴⁰ *	1.3±2.3	2.2±2.0				

^{*}All studies reported statistically significant laxity measurements between the autograft and low-dose irradiated allograft groups.

DISCUSSION

Principal findings and interpretation

Across eight studies (>10,000 patients), autografts showed lower failure/revision risk in younger, high-demand patients, whereas outcomes were broadly comparable in older cohorts. Specifically, revision risk was higher with allografts in patients ≤21 years (10.1% vs 2.9%), but not different in those >22 when low-dose irradiation was used, indicating that age/activity level modifies any irradiation-related decrement in performance. 30,44 Objective laxity was generally greater after low-dose γ-irradiated allografts-typically ≤2.5 Mrad in the RCTs-by KT-1000/2000 yet these differences rarely translated into worse PROMs (IKDC/Lysholm/Tegner) over 2–5 years. 24,36,37,40 Clinically, autografts supported earlier functional milestones (e.g., weight-bearing and running) consistent with faster ligamentization. 40,42

Dose effects and graft processing

The largest cohort to stratify dose reported no early (<1 year) difference among non-irradiated, low-dose (<1.8 Mrad) and higher-dose groups, but beyond one year failure risk rose with >1.8 Mrad, supporting a dose-response detriment to collagen integrity at moderate doses (30). These clinical observations align with bench data showing irradiation-induced collagen chain seission and reduced tensile properties in a dose-dependent fashion. ^{22,23,28}

Comparison with previous syntheses

Our pattern greater early laxity yet similar short- to midterm PROMs for irradiated allografts—accords with prior meta-analyses finding higher failure with allografts overall but small between-group differences in functional scores. ^{21,25,29} The age-contingent effect we observed echoes registry-based and multi-centre series in which autografts outperform allografts in young, cutting/pivoting athletes, while differences attenuate with age and lower activity levels. ^{38,44}

Revision settings

In revision ACLR, the MARS cohort found improved IKDC with both graft types but higher (under-powered) rerupture risk after allograft and greater functional gains

after autograft, suggesting autograft preference in young revisions where tissue quality and loading demands are greatest.³⁸

Clinical implications

For patients ≤21 years or returning to pivoting sport, autografts remain preferable given lower failure risk, earlier functional recovery and superior early stability. ^{38,40,42,44} Low-dose (≤~1.8–2.5 Mrad) irradiated allografts are reasonable in older or less active individuals where avoiding donor-site morbidity and operative time is prioritized, with the caveat that early laxity may be higher and ligamentization slower. ^{24,36,37,40,42} Doses >1.8 Mrad should be avoided when possible due to higher late failure risk. ³⁰

CONCLUSION

This review examined the comparative effectiveness of autografts and low-dose irradiated allografts in ACL repair, particularly focusing on mechanical integrity, patient-reported outcomes and long-term graft viability.

Autografts consistently show superior early biomechanical outcomes, particularly in younger patients. Several studies, including Li et al, report better knee laxity with autografts despite no difference in patient-reported outcomes such as IKDC scores.³⁶ This was the case when 2.5 Mrad or lower radiation was used, suggesting that the dose-dependent collagen damage is perhaps not relevant in doses under 2.5 Mrad. While subjective measures are important for assessing quality of life and functional return to sport, they may fail to capture clinically significant mechanical deficits. Notably, knee laxity is a better predictor of re-injury risk and potential osteoarthritis development, favoring autograft use for reducing long-term joint degeneration.

Revision rates further reinforce this preference. Maletis et al showed autografts to have significantly lower failure rates than allografts in patients under 22, with no difference in older cohorts. Harmonia This may be linked to the slower remodeling capacity of allografts and potential complications with bone incorporation, particularly when using donor bone from older individuals. Advances in soft tissue graft fixation now allow for effective use of hamstring or quadriceps autografts in skeletally immature patients, supporting a broader autograft strategy.

Although some evidence suggests that allografts can ultimately regain equivalent or greater strength through prolonged remodeling, short follow-up durations common in the literature may underestimate their long-term potential.³¹ Still, delayed return to sport and higher early laxity rates raise concerns about allograft suitability in younger or high-demand individuals. However, a meta-analysis of 17 studies comparing return to pre-injury activity levels between BPTB allograft and autograft recipients found a significantly higher proportion of successful returns in the allograft group (68.3%) compared to the autograft group (57.1%), with the odds ratio favoring allografts.⁴³

Interpretation of the literature is complicated by methodological inconsistencies, retrospective designs, non-randomized graft selection, lack of standardised rehabilitation protocols and variation in surgical techniques (e.g., single vs double-bundle reconstructions). These factors contribute to outcome heterogeneity and limit definitive conclusions. Moreover, the inconsistency in defining graft failure, particularly in athletic populations where contralateral injuries are common, adds further complexity.

In summary, autografts remain the graft of choice for younger, active patients due to their superior mechanical stability, faster incorporation and lower failure rates. Lowdose irradiated allografts may be suitable alternatives in older, less active individuals, but their long-term efficacy remains uncertain without higher-quality, standardised studies. Future research should priorities long-term, prospective trials that integrate mechanical outcomes with patient-reported data and clearly defined revision criteria. The cost and logistical challenges of low-dose irradiated allografts must be considered. Given the increased cost in sourcing an allograft and the irradiation sterilisation process, autografts may still be the preferred option by health services. However, in patients for whom avoiding donor-site morbidity is paramount, patients who are at risk of longer operative times and patients in whom immunologic reactions are unlikely, an alternative to autografts is viable.45

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REFERENCES

- 1. Butler DL, Noyes FR, Grood ES. Ligamentous restraints to anterior-posterior drawer in the human knee. A biomechanical study. JBJS. 1980;62(2):259-70.
- 2. Markolf KL, Mensch JS, Amstutz HC. Stiffness and laxity of the knee--the contributions of the supporting structures. A quantitative in vitro study. J Bone Joint Surg. 1976;58(5):583-94.
- 3. Petersen W, Zantop T. Anatomy of the anterior cruciate ligament with regard to its two bundles.

- Clinical orthopaedics and related research. 2007;454:35-47.
- Birch HL, Thorpe CT, Rumian AP. Specialisation of extracellular matrix for function in tendons and ligaments. Muscles, Ligaments Tendon J. 2013;3(1):12-22.
- Georgiev GP, Iliev A, Kotov G, Kinov P, Slavchev S, Landzhov B. Light and electron microscopic study of the medial collateral ligament epiligament tissue in human knees. World J Orthoped. 2017;8(5):372-8.
- Haslauer CM, Proffen BL, Johnson VM, Murray MM. Expression of modulators of extracellular matrix structure after anterior cruciate ligament injury. Wound repair and regeneration. Wound Heal Soc European Tissue Repair Soc. 2014;22(1):103-10.
- Amis AA. The Biomechanics of Ligaments. In: Poitout DG, editor. Biomechanics and Biomaterials in Orthopedics. London: Springer London; 2004: 550-63.
- 8. Marieswaran M, Jain I, Garg B, Sharma V, Kalyanasundaram D. A Review on Biomechanics of Anterior Cruciate Ligament and Materials for Reconstruction. Appl Bionics Biomech. 2018;2:45.
- 9. Jameson SS, Dowen D, James P, Serrano-Pedraza I, Reed MR, Deehan D. Complications following anterior cruciate ligament reconstruction in the English NHS. Knee. 2012;19(1):14-9.
- Grassi A, Macchiarola L, Filippini M, Lucidi GA, Della Villa F, Zaffagnini S. Epidemiology of Anterior Cruciate Ligament Injury in Italian First Division Soccer Players. Sports Health. 2019:1941738119885642.
- 11. Ish Kumar D, Rehan Ul H, Sudhir K. Graft choices for anterior cruciate ligament reconstruction. Indian Journal of Orthopaedics. 2015;49(2):127-8.
- 12. Mouarbes D, Menetrey J, Marot V, Courtot L, Berard E, Cavaignac E. Anterior Cruciate Ligament Reconstruction: A Systematic Review and Meta-analysis of Outcomes for Quadriceps Tendon Autograft Versus Bone-Patellar Tendon-Bone and Hamstring-Tendon Autografts. Am J Sports Med. 2019;47(14):3531-40.
- 13. Peebles KC, Jacobs C, Makaroff L, Pacey V. The use and effectiveness of exercise for managing postural orthostatic tachycardia syndrome in young adults with joint hypermobility and related conditions: A scoping review. Auton Neurosci. 2024;252:103156.
- 14. Aga C, Kartus J-T, Lind M, Lygre SHL, Granan L-P, Engebretsen L. Risk of Revision Was Not Reduced by a Double-bundle ACL Reconstruction Technique: Results From the Scandinavian Registers. Clinical Orthopaedics and Related Research®. 2017;475(10):2503-12.
- 15. Vyas D, Rabuck SJ, Harner CD. Allograft anterior cruciate ligament reconstruction: indications, techniques and outcomes. J Orthopaed Sports Phys Ther. 2012;42(3):196-207.
- Paul BR, Robaina J, Parmar R, Carter T, Shah A. Myths and facts about allograft use in anterior

- cruciate ligament reconstruction: a detailed review of the literature. JBJS Rev. 2025;13(4):67.
- Van Dyck P, Zazulia K, Smekens C, Heusdens CHW, Janssens T, Sijbers J. Assessment of anterior cruciate ligament graft maturity with conventional magnetic resonance imaging: a systematic literature review. Orthop J Sports Med. 2019;7(6):2325967119849012.
- 18. Cavanaugh JT, Powers M. ACL Rehabilitation Progression: Where Are We Now? Current reviews in musculoskeletal medicine. 2017;10(3):289-96.
- Zhu S, Ma R, Qiao R, Zhang J, Hu Y. The results of anterior cruciate ligament reconstruction with allograft versus autograft: a systematic review and meta-analysis. Eur J Orthop Surg Traumatol. 2025;35(1):337.
- Conrad B, Rappé M, Horodyski M, Farmer K, Indelicato P. The effect of sterilization on mechanical properties of soft tissue allografts. International J Bank, Eng Transplan Cells Tissues Incorp Adv Tissue Bank. 2013;14(3):359-66.
- 21. Hu J, Qu J, Xu D, Zhou J, Lu H. Allograft versus autograft for anterior cruciate ligament reconstruction: An up-to-date meta-analysis of prospective studies. International Orthopaed. 2013;37(2):311-20.
- 22. Gibbons MJ, Butler DL, Grood ES, Bylski-Austrow DI, Levy MS, Noyes FR. Effects of gamma irradiation on the initial mechanical and material properties of goat bone-patellar tendon-bone allografts. J Orthopaed Res. 1991;9(2):209-18.
- 23. Cheung DT, Perelman N, Tong D, Nimni ME. The effect of γ-irradiation on collagen molecules, isolated α-chains and crosslinked native fibers. J Biomed Mat Res. 1990;24(5):581-9.
- 24. Sun K, Tian S, Zhang J, Xia C, Zhang C, Yu T. Anterior cruciate ligament reconstruction with BPTB autograft, irradiated versus non-irradiated allograft: a prospective randomized clinical study. Knee Surgery, Sports Traumatology, Arthroscopy. 2009;17(5):464-74.
- 25. Kan SL, Yuan ZF, Ning GZ, Yang B, Li HL, Sun JC, et al. Autograft versus allograft in anterior cruciate ligament reconstruction: A meta-analysis with trial sequential analysis. Medicine. 2016;95(38):185.
- Waterman BR, Arroyo W, Cotter EJ, Zacchilli MA, Garcia ESJ, Owens BD. Septic Arthritis After Anterior Cruciate Ligament Reconstruction: Clinical and Functional Outcomes Based on Graft Retention or Removal. Orthopaed J Sports Med. 2018;6(3):2325967118758626-.
- 27. McGilvray KC, Santoni BG, Turner AS, Bogdansky S, Wheeler DL, Puttlitz CM. Effects of (60)Co gamma radiation dose on initial structural biomechanical properties of ovine bone--patellar tendon--bone allografts. Cell Tissue Bank. 2011;12(2):89-98.
- Balsly CR, Cotter AT, Williams LA, Gaskins BD, Moore MA, Wolfinbarger L. Effect of low dose and moderate dose gamma irradiation on the mechanical

- properties of bone and soft tissue allografts. Cell and Tissue Banking. 2008;9(4):289-98.
- Dashe J, Parisien RL, Cusano A, Curry EJ, Bedi A, Li X. Allograft tissue irradiation and failure rate after anterior cruciate ligament reconstruction: A systematic review. World J Orthop. 2016;7(6):392-400.
- Tejwani SG, Chen J, Funahashi TT, Love R, Maletis GB. Revision risk after allograft anterior cruciate ligament reconstruction: association with graft processing techniques, patient characteristics, and graft type. The American J Sports Med. 2015;43(11):2696-705.
- 31. McDermott I, Thomas NP. Tendon allografts in the knee. The Knee. 2005;12(6):401-4.
- 32. Wiggins AJ, Grandhi RK, Schneider DK, Stanfield D, Webster KE, Myer GD. Risk of Secondary Injury in Younger Athletes After Anterior Cruciate Ligament Reconstruction: A Systematic Review and Meta-analysis. The American J Sports Med. 2016;44(7):1861-76.
- 33. Ghodadra NS, Mall NA, Grumet R, Sherman SL, Kirk S, Provencher MT, Bach Jr BR. Interval arthrometric comparison of anterior cruciate ligament reconstruction using bone–patellar tendon–bone autograft versus allograft: Do grafts attenuate within the first year postoperatively. The American J Sports Med. 2012;40(6):1347-54.
- 34. Hulet C, Sonnery-Cottet B, Stevenson C, Samuelsson K, Laver L, Zdanowicz U, Stufkens S, Curado J, Verdonk P, Spalding T. The use of allograft tendons in primary ACL reconstruction. Knee Surgery, Sports Traumatol, Arthros. 2019;27(6):1754-70.
- 35. Dashe J, Parisien RL, Cusano A, Curry EJ, Bedi A, Li X. Allograft tissue irradiation and failure rate after anterior cruciate ligament reconstruction: A systematic review. World J Orthoped. 2016;7(6):392-400.
- 36. Li J, Wang J, Li Y, Shao D, You X, Shen Y. A Prospective Randomized Study of Anterior Cruciate Ligament Reconstruction With Autograft, γ-Irradiated Allograft and Hybrid Graft. Arthroscopy: The J Arthrosc Related Surg. 2015;31(7):1296-302.
- 37. Sun K, Zhang J, Wang Y, Xia C, Zhang C, Yu T, et al. Arthroscopic anterior cruciate ligament reconstruction with at least 2.5 years' follow-up comparing hamstring tendon autograft and irradiated allograft. Arthroscopy. 27. United States: 2011 Arthroscopy Association of North America. Published by Elsevier Inc. 2011: 1195-202.
- 38. MARS. Effect of graft choice on the outcome of revision anterior cruciate ligament reconstruction in the Multicenter ACL Revision Study (MARS) Cohort. Am J Sports Med. 2014;42(10):2301-10.
- 39. Griffith TB, Allen BJ, Levy BA, Stuart MJ, Dahm DL. Outcomes of repeat revision anterior cruciate ligament reconstruction. The American J Sports Med. 2013;41(6):1296-301.
- Rihn JA, Irrgang JJ, Chhabra A, Fu FH, Harner CD.
 Does irradiation affect the clinical outcome of

- patellar tendon allograft ACL reconstruction. Knee Surgery, Sports Traumatol, Arthrosc. 2006;14(9):885-96.
- 41. Sepúlveda F, Sánchez L, Amy E, Micheo W. Anterior Cruciate Ligament Injury: Return to Play, Function and Long-Term Considerations. Current Sports Med Rep. 2017;16(3):67.
- 42. Moretti L, Bizzoca D, Cassano GD, Caringella N, Delmedico M, Moretti B. Graft Intra-Articular Remodeling and Bone Incorporation in ACL Reconstruction: The State of the Art and Clinical Implications. J Clin Med. 2022;11(22):87.
- 43. Bottoni CR, Smith EL, Shaha J, Shaha SS, Raybin SG, Tokish JM, et al. Autograft Versus Allograft Anterior Cruciate Ligament Reconstruction: A Prospective, Randomized Clinical Study With a Minimum 10-Year Follow-up. Am J Sports Med. 2015;43(10):2501-9.
- 44. Maletis GB, Chen J, Inacio MC, Love RM, Funahashi TT. Increased risk of revision after anterior cruciate ligament reconstruction with bone–patellar tendon–bone allografts compared with autografts. Am J Sports Med. 2017;45(6):1333-40.
- 45. Caldwell PE, Shelton WR. Indications for allografts. Orthopedic Clinics. 2005;36(4):459-67.

- 46. Grassi A, Zaffagnini S, Marcheggiani Muccioli GM, Roberti Di Sarsina T, Urrizola Barrientos F, Marcacci M. Revision anterior cruciate ligament reconstruction does not prevent progression in one out of five patients of osteoarthritis: a meta-analysis of prevalence and progression of osteoarthritis. J Joint Disord Amp Orthop Sports Med. 2016;1(1):16.
- 47. Liberati A, Altman DG, Tetzlaff J, Mulrow C, Gøtzsche PC, Ioannidis JPA, et al. The PRISMA statement for reporting systematic reviews and meta-analyses of studies that evaluate healthcare interventions: explanation and elaboration. BMJ. 2009;339:2700.
- 48. Scheffler S, Unterhauser F, Weiler A. Graft remodeling and ligamentization after cruciate ligament reconstruction. Knee surgery, sports traumatology, arthroscopy. Off J ESSKA. 2008;16:834-42.

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