Graft Selection in Anterior Cruciate Ligament Reconstruction

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Abstract: Surgical reconstruction of the anterior cruciate ligament (ACL) is often indicated to restore functional stability and prevent early degeneration of the knee joint, as there is little biological healing capacity of the native ACL. Although a reconstructed ACL does not fully restore the original structure or biomechanics properties of the native ACL, the graft used for reconstruction must not only have structural and mechanical properties that closely resemble those of the native ligament, it must also have minimal antigenicity and enough biological potential to incorporate into host bone. There are several considerations in graft selection: autograft versus allograft, and soft tissue grafts versus grafts with bone plugs. Commonly used grafts include bone-patella tendon-bone, hamstring, and quadriceps; among allografts, options further include tibias anterior and posterior, Achilles, an peroneal tendons. Optimal graft selection is not only dependent on graft properties, but perhaps more importantly on patient characteristics and expectations. The purpose of this review is to summarize the relevant biological, biomechancial, and clinical data regarding various graft types and to provide a basic framework for graft selection in ACL reconstruction.

Key Words: anterior cruciate ligament reconstruction, autograft, allograft, bone patellar tendon bone, hamstrings, quadriceps

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R upture of the anterior cruciate ligament (ACL) is a frequent injury in the general population, with an incidence up to 75 per 100,000 person-years, particularly in younger active individuals involved in cutting or contact sports. With increased early sports participation, incidence in the pediatric population is also rising. It is known that there is little biological healing potential in the native ACL. Thus, surgical reconstruction of the ACL is often indicated to restore functional stability during athletic activity and prevent early degeneration of the knee joint. Use a Surgical reconstruction is performed using a variety of graft types, each with its own benefits and disadvantages. Despite the large volume of research and improvements in surgical technique, there is still considerable debate regarding the ideal graft choice for any particular patient.

Although a reconstructed ACL does not fully restore the original structure or biomechanical properties of the native ACL,⁵ the graft used for reconstruction must not only

have structural and mechanical properties that closely resemble those of the native ligament, it must also have minimal antigenicity and enough innate biological healing potential to adequately incorporate into the host bone. In selecting graft types, there are several considerations: autograft versus allograft, and soft tissue only grafts versus grafts with bone plugs. The commonly used autograft types are bone-patella tendon-bone (BTB), hamstring, quadriceps (with or without a patellar bone plug); among allografts, additional options include tibialis anterior and posterior, peroneal, and Achilles tendon. Optimal graft selection depends not only on graft properties, but more importantly on patient characteristics and expectations. It is essential for the surgeon to have a thorough understanding of the surgical techniques, the basic biology of graft-bone healing, and patient expectations regarding donor site morbidity, postoperative recovery and return to activity, and potential longer term outcomes.

AUTOGRAFTS

Autografts, in particular BTB and quadrupled hamstring (Fig. 1), have been most used for ACL reconstruction. Theoretically, from a biological healing perspective, autografts are preferred because they consist of viable autogenous tissue and avoid the risk of disease transmission, maximizing the speed and likelihood of complete biological integration at the grafthost junction. Within the autografts, major considerations are donor site morbidity, and graft-tunnel healing. Clinically, the most commonly used autografts are BTB and hamstring, 6-8 followed by quadriceps tendon.

Bone-Patellar Tendon-Bone Autograft

BTB autograft was the first graft used for ACL reconstruction. The presence of bone plugs at both ends of the graft allow for retention of a native tendon-bone interface and thus graft-host integration via bone-to-bone healing. Creeping substitution at the bone-to-bone graft-tunnel interface creates a true bony junction, and is known to be stronger than soft tissue-bone healing that occurs through a fibrovascular scar. In laboratory studies, complete graft integration has been shown to occur more rapidly with a bone-bone interface compared with tendon-bone. Despite these findings, in the clinical setting, good graft incorporation can be achieved with all graft types (Fig. 2).

Biomechanically, BTB autograft has similar properties to the native ACL, with slightly higher ultimate tensile load, but a smaller cross-sectional area ^{10,12} (Table 1). In addition to the smaller cross-sectional area, the dimensions of the BTB graft are fixed and dependent on the patient's native anatomy; patella baja or alta will lead to a shorter or longer graft, respectively. Variations in this graft length can lead to technical challenges, specifically graft-tunnel mismatch. Another theoretical disadvantage to BTB grafts is aperture micromotion. Laboratory studies have shown that motion is greatest and healing is slowest at tunnel apertures. ¹⁴ Given

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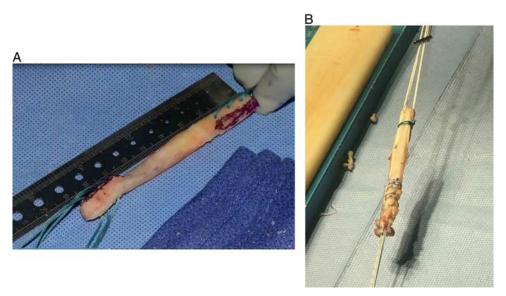


FIGURE 1. Intraoperative images of BTB (A) and hamstring (B) autografts after harvest and preparation. BTB indicates bone-patella tendon-bone.

the BTB graft's fixed tendon length and decreased crosssectional area, the mismatch in cross-sectional area at the tunnel aperture theoretically allows greater motion; intraoperatively this is often addressed by using aperture fixation. Nonetheless, despite the few theoretical biomechanical disadvantages, BTB grafts are known to lead to improved postoperative stability and decreased rates of early rerupture compared with other graft types. ^{15,16}

There are several clinical disadvantages of using BTB autograft. First is donor site morbidity, as BTB is harvested from the central third of the patellar tendon with bone blocks from the patella and tibia, through a larger incision than hamstring harvest. Anterior knee pain is more common after BTB harvest, with incidence reported up to 42%, ¹⁷ although this has not been shown to change functional

scores, overall activity level, or return to same level of play. ^{15,17,18} A larger incision carries a theoretical increased risk for infection although this has not been shown in clinical studies, with reported infection rates ranging from 0.1% to 0.3%. ¹⁹ Intraoperative and postoperative patella fracture due to the disruption of the extensor mechanism has been reported as a rare complication. ^{20,21} Fractures occur after direct or indirect trauma or as a result of physiological forces applied on a mechanically weaker patella, ²¹ tend to be transverse, and after proper treatment do not change overall outcomes at midterm follow-up. ²² Proper harvesting technique and the use of bone graft to fill the patellar void after BTB harvest, may theoretically decrease postoperative fracture risk. Another rare complication that has been reported is patellar tendon rupture. ²³ At long-term follow-

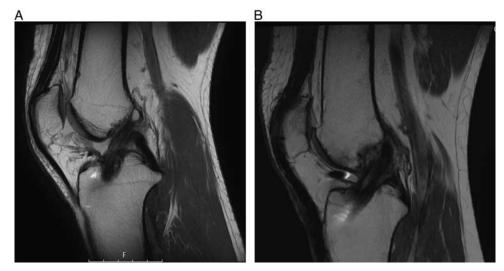


FIGURE 2. Postoperative sagittal proton density magnetic resonance images of healed ACL reconstruction after hamstring autograft (A), and quadriceps autograft (B). Grafts show intact fibers and good incorporation at the tunnel interface. ACL indicates anterior cruciate ligament.

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TABLE 1. Biomechanical Properties of ACL Graft Options*

	Tensile Load (N)	Stiffness (N/mm)	Cross-sectional Area (mm²)
Native ACL	2160	242	44
Autograft			
Bone-patellar tendon-bone	2977	620	35
Semitendinosus tendon	1216	186	14.0
Gracilis tendon	838	170	7.6
Quadruple hamstring	4090	776	53
Quadriceps tendon	2352	463	62
Allograft			
Bone-patellar tendon-bone	1403	224	
Achilles	1189	743	105
Tibialis anterior	3012	343	

^{*}Adopted from data by Noyes et al, 13 West and Harner, 10 and Mehran et al. 6

up, it has been suggested, based on nonrandomized trials, that patients who have undergone BTB autografts may be at a slightly increased risk of radiographic evidence of osteoarthritic change at 15 to 20 years postoperative; however, the clinical significance of these findings is still unclear as functional scores and activity level remain high. 17,24

Hamstring Autograft

Hamstring autograft is typically harvested through a small longitudinal incision over the medial proximal tibia. The insertions of the semitendinosus and gracilis are identified deep to the Sartorius fascia, and tendons are harvested by stripping off the muscle and detaching proximally. Because of the small incision size, no additional trauma to the extensor mechanism, and no bony disruption, donor site morbidity is much less with hamstring harvest than BTB harvest. Biologically, however, as a purely soft tissue graft, the absence of a bone plug necessitates tendon-bone healing and the generation of fibrovascular scar tissue rather than reconstitution of the native direct insertion with a fibrocartilage transition zone. In controlled laboratory studies, tendon-bone healing has been shown to be slower than bone-bone healing, with hamstring grafts showing inferior initial pullout strength to BTB, but no difference by 6 weeks in a canine model.²⁵ This longer period of initial graft integration may also be present in clinical practice, as large studies have shown greater rates of early re-rupture after ACL reconstruction with hamstring compared with BTB, ^{16,26,27} although many of these are midsubstance failures.

Biomechanically, a quadrupled hamstring graft achieves greater ultimate tensile load, stiffness, and cross-sectional area than both BTB autograft and the native ACL^{6,10} (Table 1). Compared with BTB grafts, the likelihood of graft tunnel mismatch is less likely; however, hamstring autograft size may be limited by variability in native hamstring anatomy. It has been shown that there is increased risk of failure after ACL reconstruction with hamstring grafts of 8 mm and smaller in diameter. ^{28,29} Several recent large retrospective studies have modeled the relationship between increase in hamstring graft size and reduced risk of failure, ^{30,31} with Snaebjornsson et al³⁰ reporting a 0.85× lower likelihood of revision surgery with every 0.5 mm increase in hamstring autograft diameter,

based on national registry data from Sweden. Graft size can be roughly predicted using the patient's height and preoperative MRI scan²⁸; however, in the event of a small graft or technical problem during graft harvest, augmentation using allograft tendon can be performed. Inaddition, smaller hamstring grafts have been associated with particularly high failure rates in patients below 20 years of age.²⁸

Aside from early failure and small graft diameter, there are several important postoperative considerations when using hamstring autograft. Hamstring weakness after tendon harvest is a concern, as it has been demonstrated that peak isokinetic torque is decreased after hamstring autograft compared with BTB autograft at 5 years after ACL reconstruction.³² In light of this reported postoperative loss of hamstring strength, some authors recommend avoiding hamstring autograft in high-level athletes.³³ However, there also exist studies suggesting that there is no difference in postoperative knee flexion strength after hamstring harvest compared with BTB.³⁴ Further high-quality studies are required to fully elucidate the likelihood of postoperative hamstring weakness and the role of rehabilitation programs in its prevention.

Another concern with hamstring grafts is tunnel widening and its possible association with increased postoperative laxity. Radiographic tunnel widening has been reported, at a higher rate following hamstring than BTB autograft, in both short and intermediate-term follow-up, and in a systematic review of randomized trials. 6,35,36 The hypothesized mechanism is graft micromotion, which in animal models has been shown to be greatest at tunnel apertures and least at tunnel exits. 14 Fixation technique likely affects graft micromotion, as suspensory fixation is thought to lead to greater graft micromotion at the apertures, although this is yet to be shown in large studies. Tunnel widening is a concern because it may be associated with increased laxity postoperatively following ACL reconstruction with hamstring autograft. Although some studies have shown no difference in laxity, 8 several large studies and systematic reviews have shown increased likelihood of positive Lachman, pivot shift, and laxity on KT-1000 measurement. 15,37,38 Despite the increased objective laxity, there is no difference in patient-reported outcome scores (PROs), and the long-term effects on joint degeneration are unclear, as the use of hamstring autograft has been associated with decreased risk of osteoarthritis at long-term follow-up in 2 retrospective series. 17,24

Comparison of Outcomes After BTB Versus Hamstring Autograft

Many studies have compared clinical outcomes and retear rates after BTB and hamstring autograft. Post-operative PROs are similar, with both grafts yielding high patient satisfaction and PRO scores. ^{15,17,24,39–42} In addition, the time to safely return to play has been shown to be similar between the 2 types of grafts. ^{42,43}

There are, however, several outcomes that differ after BTB and hamstring autograft. BTB is associated with greater stability on KT-1000 and pivot shift tests. ^{15,41} More importantly, BTB grafts have been shown in several separate cohorts to have lower retear rate than hamstring grafts, with up to a 2 to 4-fold higher retear risk with hamstring autograft. ^{8,16,26,27} This difference in retear rates is especially important for high-risk patients who are young and play cutting and pivoting sports and thus already have a high risk for retearing their graft. Nonetheless, despite the lower

ACL indicates anterior cruciate ligament.

retear rate, BTB grafts are associated with greater anterior knee pain than hamstring autograft.^{15,40} In addition, an increased risk of postoperative quadriceps weakness has been reported after BTB autograft harvest.⁴⁴

There is limited long-term follow-up data in the literature, but there are 2 recent studies with 15- and 20-year follow-up that suggest that this increased risk of re-rupture persists after the immediate and intermediate postoperative period. 17,24 Thompson et al²⁴ reported an 18% and 10% failure rate, respectively at 20 years after hamstring and BTB autograft use. Interestingly, both long-term follow-up studies suggest increased incidence of radiographic osteoarthritis in the BTB group. The clinical significance of this finding is unclear, as despite the radiographic degenerative changes, all functional and PROs scores were similar between the 2 groups. Further long-term studies need to be conducted to determine the effect of graft choice on long-term knee health as there are many confounding factors that affect long-term outcome studies.

Quadriceps Autograft

Another autograft option, which has gained interest recently, is the quadriceps tendon. 45 Graft harvest is typically performed using a separate anterior superior incision over the distal quadriceps tendon, and the central third of the tendon is harvested. Depending on surgeon preference, a bone block from the superior patella may be harvested as well to maintain a native tendon-bone interface. Benefits to quadriceps autograft are consistently greater graft cross-sectional area, reduced risk of anterior knee pain and kneeling pain, and decreased risk of patellar fracture compared with BTB. 45-48

There are several recent clinical comparative studies investigating outcomes after quadriceps tendon versus other autografts for ACL reconstruction. A prospective randomized study of 39 patients by Lund et al⁴⁶ found that compared with BTB autograft, at 1- and 2-year follow-up, there were no differences in outcome scores or laxity testing after quadriceps autograft, but there was a lower incidence of graft site pain and kneeling pain. A more recent, nonrandomized retrospective study of 90 patients at minimum 3 years postoperative by Cavaignac et al⁴⁹ compared outcomes after quadriceps versus hamstring autograft. Their results suggest similar to slightly improved outcome scores, improved stability on KT-1000 measurement, and greater likelihood of negative Lachman following quadriceps tendon autograft, with no difference in morbidity. A similar recent retrospective comparative study of quadriceps versus hamstring showed no difference in postoperative outcome scores, return to preinjury activity level, or morbidity.⁵⁰ Results from larger, ongoing clinical studies with more patients and longer follow-up will provide further insight into the benefits and risks of using quadriceps tendon in ACL reconstruction.

ALLOGRAFTS

Allografts are commonly used in ACL reconstruction, with reports of 22% to 42% of ACL reconstructions being performed with allografts in the United States.^{51,52} Options include allograft BTB, quadrupled hamstring, quadriceps, Achilles, tibialis anterior and posterior, and peroneals. Anatomic origin of the allograft affects biomechanical properties, with greatest load to failure seen in looped tibialis anterior or posterior (Table 1), greatest stiffness in

quadriceps, and lowest load to failure and stiffness in non-looped tibialis anteriorgrafts. 53,54

Graft processing is an important consideration in selecting an allograft. Irradiation has been associated with higher rates of graft failure in numerous clinical studies. A dose-response relationship has been established with higher levels of gamma irradiation being associated with decreased load to failure. 55-57 Electron beam sterilization is thought to be less detrimental to the structural and biomechanical properties of allografts.^{58,59} Aside from radiation, there are chemical sterilization techniques as well. Among these, peracetic acid, ethylene oxide, and supracritical CO₂ treatment have been shown to decrease stiffness and load to failure in various types of allograft.⁵³ In general, the less processed an allograft is, the more structurally and biomechanically stable it is; however, processing helps to decrease both disease transmission and the host's immuneinflammatory response to the graft. Transmission of viruses such as hepatitis B, hepatitis C, and HIV, is a rare but feared phenomenon, with an estimated risk of about 1 in 1,667,000.6 Although true graft immune rejection has not been reported, fresh frozen allografts have been shown to be associated with a postoperative inflammatory synovitis possibly reflecting a latent immunologic response.60

Several other notable considerations for allografts are storage method and donor characteristics. Freezing at -80° C for > 30 days, > 3 freeze-thaw cycles, and treatment with various preservatives have all been shown to have negative effects on biomechanical properties. Freezing donor characteristics, increased donor age has been shown to have a negative correlation with ultimate tensile strength and modulus of elasticity. Although these allograft factors are likely un-modifiable for each individual case, the surgeon should become familiar with the local tissue bank's methods for acquisition, processing, and storage.

Autograft Versus Allograft

The clinical literature generally shows superior outcomes following ACL reconstruction with autograft compared with allograft, although good results have been reported with allograft as well. The many permutations of the allograft versus autograft comparison, given the different anatomic types and processing methods, make interpreting individual studies in the greater context of clinical practice somewhat difficult. Furthermore, in clinical studies, the different types of allografts are often grouped together into a single cohort, causing confounding from various graft-related factors discussed above. Nonetheless, there has been an abundance of recent literature focusing on outcomes after allograft versus autograft ACL reconstruction.

Several notable studies of ACL reconstruction using autograft BTB versus allograft BTB have shown superior outcomes in the autograft group. Krych et al⁶² reported a 5-fold increase in risk of rerupture after BTB allograft compared with BTB autograft. When excluding irradiated and chemically processed grafts, they saw no difference in failure rate between BTB allograft and autograft; however, their meta-analysis/systematic review only included 6 studies. Kraeutler et al⁶³ reported similar results with an approximately 3-fold increase in risk of graft failure after BTB allograft compared with BTB autograft (12.7% vs. 4.3%). They also showed increased knee laxity, decreased single-leg hop test results, and lower subjective satisfaction after BTB allograft. Notably, however, patients in the

allograft group had significantly less knee pain, and processing and irradiation of allografts was not differentiated.

With regard to graft processing, it is generally believed that sterilization techniques alter the biomechanical properties of a graft, and that the more processed a graft is, the worse it performs. Park et al⁶⁴ performed a systematic review of irradiated versus nonirradiated allografts at minimum 2-year follow-up, showing lower outcome scores, decreased stability to Lachman, pivot-shift, and KT-1000 testing, and increased risk of revision compared with nonirradiated allografts. Several other recent studies, including by Yao and colleagues and Tian et al⁶⁵ have shown similar results.

With greater failure rates and worsened biomechanical properties after use of irradiated grafts, some authors have suggested that nonirradiated allografts may have similar outcomes to autografts. Mariscalo et al⁶⁶ performed a retrospective study comparing BTB autografts and nonirradiated allografts, reporting no significant difference between the 2 groups in postoperative laxity, outcome scores, or failure rates. Most recently, Maletis et al⁶⁷ performed a large registrybased retrospective cohort study of 14,015 ACL reconstructions with BTB autograft, hamstring autograft and soft tissue allograft subcategorized based on processing and irradiation. Using an adjusted model for age, sex, and race, they found that BTB autograft had the lowest revision risk, followed by hamstring autograft [hazard ratio (HR), 1.51], allografts irradiated with <1.8 Mrad with chemical processing (HR, 2.19) and without chemical processing (HR, 2.31), and allografts irradiated with > 1.8 Mrad with chemical processing (HR, 5.03) and without chemical processing (HR, 6.30). In their cohort, nonprocessed allografts and those irradiated with <1.8 Mrad had a similar risk of revision compared with hamstring autografts.

Caution must still be taken when applying these findings to patients in practice, as various patient factors must be considered in addition to graft type when discussing failure rate and long-term outcomes. Kaeding et al⁶⁸ reported, as part of the Multicenter Orthopaedic Outcomes Network trials, that although patients undergoing allograft ACL reconstruction had an overall 4× greater chance of graft rupture, they saw a notable impact from age, with a higher failure rate in all grafts in younger patients.

GRAFT SELECTION IN THE PEDIATRIC PATIENT

Given the increasing incidence of pediatric ACL tears,² ACL reconstruction in the skeletally immature population is becoming a growing area of interest. The main strategies for ACL reconstruction in the pediatric population, depending on the patient's skeletal maturity and surgeon preference, are transphyseal reconstruction, all-epiphyseal reconstruction using soft tissue graft and bone tunnels in the epiphysis, and physeal-sparing reconstruction using iliotibial (IT) band autograft without bone tunnels, also known as the modified Macintosh technique.⁶⁹ Trans-epiphyseal technique is similar to adult ACL reconstruction and is used in patients nearing skeletal maturity; however, oftentimes more vertical tunnels are preferred to leave a smaller footprint crossing the physis. In more immature patients, a completely physeal sparing approach can be used in which tunnels are entirely in the epiphysis; intraoperative 3D imaging or navigation can be used to aid in confirming tunnel placement. Only soft tissue grafts (not allografts) should be used for ACL reconstruction in pediatric patients with open physes. 70 The quadrupled hamstring graft is most

common, although quadriceps tendon graft may be used. The patella tendon should not be harvested in pediatric patients with open physes to avoid damage to the tibial tubercle apophysis. Allografts are not indicated in pediatric patients in most cases, as allografts are known to have a higher failure risk.

Alternatively, the modified Macintosh procedure uses the native IT band, with its distal insertion onto Gerdy tubercle left intact, threaded into the notch in an "over the top position" and brought over the anterior tibial plateau between the tibial spines and under the intermeniscal ligament as the ACL graft. Although the literature on ACL reconstruction in the skeletally immature patient is quite limited, 72 good outcomes have been reported. After IT band autograft physeal sparing ACL reconstruction, at 6-year postoperative, the failure rate has been reported to be 6.6% with excellent clinical outcomes and no limb length discrepancy or angular deformity. 73

GENERAL ALGORITHM FOR GRAFT SELECTION IN ADULT PATIENTS

Despite the abundance of studies on ACL graft types, there is still no ideal graft, and there is no generally accepted algorithm for graft selection for a given patient. In the clinical setting, all graft factors discussed above must be considered, such as biomechanical properties, presence of a bone plug, donor site morbidity and graft availability (in places where allografts are not routinely used). Applying the literature on failure rate and clinical outcomes, several key clinical factors to emphasize are age, desired activity type and level, occupation, barriers to rehabilitation, and general expectations, as these factors affect failure rate, outcomes, and satisfaction. In the setting of revision ACL reconstruction or multiligamentous reconstruction, previous tunnels, grafts, and implants, and availability of autograft donor sites must be considered.

In general, for primary ACL reconstruction, adult patients can be divided into 3 main categories: elite athletes and young highly active patients, recreational athletes and active individuals, and older or less active patients.

Overall, in the setting of a primary isolated ACL reconstruction, the preferred graft of choice for consideration in the youngest, most active group is BTB autograft, although with the expanding literature, quadriceps tendon autograft may become an increasingly viable option in this population as well. Given that hamstring autograft and allograft have both shown higher failure rates and increased postoperative laxity, these choices are less ideal although they are still frequently used, especially in the revision or multiligamentous setting or if the patient has pre-existing anterior knee pain, or is required to kneel frequently due to occupation.

For the patients in the middle category, who are moderately active but not necessarily elite athletes, and are still physiologically young, the graft of choice to consider is frequently hamstring autograft. Hamstring autograft has a lower failure and revision rate compared with allografts, and also avoids the donor site morbidity associated with BTB harvest. Although there may remain some residual laxity when compared with a BTB graft, it is unclear whether this difference is clinically relevant.

For older, less active patients, if nonoperative treatment fails, the graft of choice remains hamstring autograft; however, in patients willing to accept an increased risk of graft failure, allograft may be considered.

CONCLUSIONS

Graft options for ACL reconstruction fall into 2 general categories, autograft and allograft. Frequently used autografts are BTB, quadriceps, and hamstring, whereas frequently used allografts include BTB, hamstring, tibialis anterior and posterior, peroneals, and Achilles. Biologically and biomechanically, the presence of a bone plug allows for bone-to-bone healing and retention of a normal native direct insertion site at the donor bone plug. In laboratory and clinical studies, autograft outperforms irradiated and processed allografts. Allograft may be considered for less active patients who are willing to accept an increased risk of graft failure. Within the autograft category, BTB has demonstrated the lowest failure rate and superior objective stability measurements; however, it is associated with increased anterior knee pain; overall clinical outcome scores are similar among the various autografts. The growing body of literature will continue to identify and provide further insight into the strengths and weaknesses of various graft types as we continue to refine our understanding of ACL healing and rehabilitation.

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