Revision and Conversion to Arthroplasty are Low Among Adolescents Undergoing Meniscal Allograft Transplantation Using the Bridge-In-Slot Technique at Midterm Follow-Up

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Running Title: Meniscal Allograft Transplants in Adolescents

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Title: Revision and Conversion to Arthroplasty are Low Among Adolescents Undergoing
 Meniscal Allograft Transplantation Using the Bridge-In-Slot Technique at Midterm Follow-Up

3 Abstract:

<u>Purpose</u>: To report midterm outcomes following primary medial and lateral meniscal allograft
transplantation (MAT) with fresh-frozen allografts implanted with the bridge-in-slot technique in
the adolescent patient population.

7 Methods: Adolescent patients less than 18 years old at the time of primary MAT from 1999-2016 8 were retrospectively identified. International Knee Documentation Committee (IKDC) subjective form, Lysholm, and Knee Injury and Osteoarthritis Outcome Score (KOOS) subscales scores were 9 10 collected preoperatively and at 1-year, 2-years, and a minimum of 5-years follow-up. Thresholds for achieving clinically significant outcomes were calculated and the proportion of patients 11 achieving minimal clinically important difference (MCID), patient acceptable symptomatic state 12 (PASS), and substantial clinical benefit (SCB) was determined. Meniscus reoperation (partial, 13 subtotal, or total meniscectomy, repair, or failure) and failure (revision MAT or conversion to 14 arthroplasty) rates were determined. 15

<u>Results</u>: Forty-four (female n=33; male n=11) of 62 identified patients met inclusion criteria and
were followed for a mean of 9.5 ± 3.8 years (range: 5.0-17.7). Lateral MAT was performed in most
patients (n=35/44; 80%). Isolated MAT was performed in 27 (61%) patients. Common
concomitant procedures included osteochondral allograft transplantation (OCA) (32%),
autologous chondrocyte implantation (ACI) (18%), and anterior cruciate ligament reconstruction
(ACLR) (14%). MCID, PASS, and SCB were achieved by patients at a minimum 5-year followup for IKDC (62%;76%;31%), Lysholm (62%;79%;23%), and KOOS questionnaires (Pain

[65%;81%;41%], Symptoms [58%;81%;47%], Activities of Daily Living [53%;77%;35%], Sport [86%;75%;50%], and Quality of Life [59%;81%;59%]), respectively. Eleven patients (25%) underwent reoperation at an average of 5.9 ± 4.5 years (range: 0.8 - 14.0) following MAT. Three (7%) patients met criteria for failure, requiring revision MAT an average of 3.8 ± 1.1 years (range: 2.8-4.9) post-transplant. No patients underwent arthroplasty. Overall survival free from failure at 1, 2, 5, and 10 years was 100%, 100%, 93%, and 93%, respectively. At the time of final followup, 80% of patients reported satisfaction with their current physical status.

<u>Conclusion:</u> Primary MAT in adolescent patients resulted in significant and durable functional
 improvements at mid- to long-term follow-up. At an average of 9.5 years postoperatively,
 meniscal reoperation rate was 25% while graft survival free of revision MAT was 93%.
 Adolescents undergoing MAT demonstrated similar functional outcomes and graft survivability
 when compared to available adult MAT literature.

35 <u>Level of Evidence</u>: Level IV, Retrospective Case Series

36 Introduction:

Youth meniscus injuries are increasingly common and may be attributable to increases in
organized athletic competition, sport specialization, and injury awareness.^{1–5} Meniscus injuries can
occur in isolation or in conjunction with other injuries such as anterior cruciate ligament (ACL)
tears, focal chondral injuries, and tibial fractures.^{1,5} Discoid menisci have also been identified as
risk factors for meniscus injury.^{6,7}

Current treatment guidelines for meniscus injuries recommend that meniscal tissue be 42 preserved whenever feasible, as subtotal or total meniscectomy has been shown to increase contact 43 forces and negatively impact long-term joint health.^{8,9} Meniscus preservation is especially relevant 44 in adolescents because this population will place greater demand on the knee joint over a longer 45 period of time than older adults.^{4,10} Although the meniscus is thought of as a relatively avascular 46 structure, meniscus tissue is richly vascular at birth and maintains some degree of vascularity to 47 allow for growth prior to regressing into the periphery by adulthood.^{11,12} Therefore, adolescent 48 menisci are generally thought to be more amenable to surgical repair given better blood flow and 49 healing potential.^{13,14} Longitudinal vertical and bucket-handle tears are fairly common meniscal 50 injury patterns and can routinely be managed with arthroscopic repair. However, radial, oblique, 51 horizontal, and complex tears are historically thought to fare poorly following repair and often go 52 on to partial meniscectomy.^{1,14,15} 53

Even following conservative partial meniscectomy, outcomes in youth populations are suboptimal. Long-term outcome studies have found that up to 80% of children and adolescents have evidence of degenerative changes as early as five years post-meniscectomy, while 75% and 60% report persistent pain and dissatisfaction with clinical outcomes, respectively.^{9,16} Previously, there was hesitation to indicate MAT in children and adolescents due to concerns about allograft

Iongevity and the unknown effects of transplantation on skeletal development.^{4,17} Initial short-term outcome reports have demonstrated consistent improvement in pain, function, and activity levels, suggesting that MAT is a viable treatment option for children and adolescents.^{6,10,18} However, the majority of available outcomes are limited by short-term follow-up. Studies with increased follow-up are needed in this patient population to evaluate for durability of clinical benefit as well as allograft survivorship.

The purpose of this investigation was to report midterm outcomes following primary medial and lateral meniscal allograft transplantation (MAT) with fresh-frozen allografts implanted with the bridge-in-slot technique in the adolescent patient population. The authors hypothesized that at a minimum five-year follow-up, MAT in the adolescent population would demonstrate significant improvements in patient-reported outcomes as well as allograft survivorship similar to reported rates of previous studies for adult populations.

71

72 Methods:

73 Patient Population:

Local Institutional Review Board approval was obtained prior to study initiation. A prospectively collected database from a single institution was retrospectively reviewed for patients who underwent primary MAT with a single surgeon between 1999-2016. Patients who underwent any concomitant procedures at the time of MAT were included. Inclusion criteria included the following: (1) primary MAT for symptomatic meniscal deficiency for which conservative treatment had failed, (2) age 18 years or younger at the time of primary surgery, and (3) minimum 5-year follow-up. Patient-reported outcomes and an updated surgical history were collected to

calculate reoperation and failure rates. Subsequent meniscal reoperation was defined as partial,
subtotal, or total meniscectomy, or meniscal repair. MAT failure was defined as revision MAT or
conversion to unicompartmental or total knee arthroplasty.¹⁹

84 <u>Surgical Technique:</u>

All MATs were performed by the senior author, a fellowship-trained orthopedic surgeon 85 who operates a high-volume joint preservation clinical practice. For both medial and lateral MAT, 86 the senior author utilized fresh-frozen, non-irradiated meniscal grafts fixated in bone troughs using 87 the bridge-in-slot technique.²⁰⁻²² As previously described, the Pollard method was used for 88 allograft matching.²³ If present, the senior author will also treat associated pathology or 89 abnormalities at the time of MAT, such as lower extremity malalignment, focal chondral defects, 90 or ligamentous instability. Realigning osteotomy was typically performed concomitantly with 91 MAT if patients had ≥ 5 degrees of mechanical axis malalignment on coronal radiographs. At the 92 time of transplantation, the meniscus is then debrided to a bleeding peripheral rim of 1- to 2-mm. 93 The anterior and posterior horns are subsequently resected. To prepare the meniscal slot 94 (approximately 8 mm wide and 10 mm deep), an initial slot guide is first made using a 4.5-mm 95 burr. In the presence of an open physis, intraoperative imaging can confirm the protection of the 96 physis in relation to the bone trough. A guide pin is then placed using an over-the-top style slot 97 guide. A 7-mm reamer is used to over-ream the guide pin and the slot is refined using a box cutter, 98 99 dilating rasp, and a bone-cutting shaver utilizing a dovetail technique. When possible, the back wall of the tibia is preserved to help prevent posterior displacement, but at times it may be 100 101 necessary to remove in order to improve the overall seating of the allograft. While the tibial slot is 102 being prepared, the allograft is thawed in normal saline on the sterile back surgical table.

The meniscal allograft bone bridge is created using reciprocating saw between the anterior 103 and posterior horn of the donor allograft. A 0-polydioxanone (PDS) suture is then placed through 104 105 the posterior third of the meniscus to assist with allograft insertion into the joint and positioning within the tibial slot. We do not generally employ transosseous sutures within the slot. With the 106 knee in flexion, a knotless suture anchor (SwiveLock, Arthrex, Inc., Naples, FL) is placed 107 108 anteriorly on the far side of the meniscal allograft to secure the allograft bone bridge to the recipient tibial tunnel. For circumferential fixation, approximately 8 to 10 vertical mattress sutures are 109 110 placed using an inside-out technique (FiberWire, Arthrex, Inc., Naples, FL). In one case where medial MAT and concomitant ACL reconstruction was performed, a modified bone block 111 technique was used. In this patient, the bone block was split and the posterior block was shuttled 112 into the joint after femoral fixation of the ACL graft. The posterior block was then fixated with 113 transosseous sutures, inside out repair performed, and finally transosseous sutures were used to 114 fixate the anterior horn. After confirming satisfactory meniscus placement and stability, incisions 115 are closed in standard fashion. 116

117

118 <u>Rehabilitation Protocol</u>:

Between postoperative weeks 0-2, patients begin heel touch weight bearing with crutches hinged knee brace locked in full extension for all activities. Exercise during this time period is limited to heel slides, quad sets, patellar mobs, and straight leg raises. Knee flexion is also limited from 0 to 90 degrees when non-weight bearing. During weeks 2-8 postoperatively, patients maintain heel touch weight bearing with crutches, with progression to full weight bearing by 6-8 weeks. When non-weight bearing during this timeframe, the knee immobilizer should continue to be worn and locked at 0-90 degrees but can be removed for sleep. Bracing can be discontinued at

126 8 weeks. Additionally, patients should continue range of motion exercises as tolerated between 127 weeks 2-8, with no restriction thereafter. Closed chain exercises are slowly introduced, with 128 gradual return to functioning activities at 12 to 20 weeks. Patients are allowed to return to full 129 activity by 5 months post-operatively once cleared by the attending physician.

130

131 <u>Data Collection</u>:

International Knee Documentation Committee (IKDC) subjective form, and Knee injury 132 133 and Osteoarthritis Outcome Score (KOOS subscales: Pain, Symptoms (SXS), Activities of Daily Living (ADL), Sport, and Quality of Life (QOL), and Lysholm scores were collected 134 preoperatively and at 1-year, 2-year, and at a minimum 5-year follow up. All procedures prior to 135 index surgery related to the operative knee, concomitant procedures, as well as the incidence of 136 reoperations and failures were recorded. Postoperative satisfaction was assessed with a "yes" or 137 "no" answer to the question, "Considering all the activities you have during your daily life, your 138 level of pain, and your functional impairment, do you consider that your current state is 139 satisfactory?" 140

141

142 <u>Statistical Analysis</u>

Descriptive statistics for binomial variables were reported as frequencies (proportions) and continuous variables were reported as means with standard deviations. Mann Whitney U, Wilcoxon signed rank, paired or unpaired t-tests were utilized when comparing preoperative and postoperative patient-reported outcome measures. Shapiro-Wilk testing determined normality of the data. Fisher's Exact test was used if frequencies for variables were less than 5 and Chi-square

analysis was used for comparing categorical variables. Thresholds for achieving clinically 148 significant outcomes (CSOs) were established using a distribution method for minimal clinically 149 important difference (MCID).²⁴ Patient acceptable symptomatic state (PASS) and substantial 150 clinical benefit (SCB) were derived using an anchor-based method.²⁴ A receiver operating 151 characteristic (ROC) curve with an area under the curve (AUC) of less than 0.7 and 0.8 was 152 153 considered an acceptable and excellent predictor, respectively. Kaplan-Meier survival analysis was utilized to determine survival probabilities. Cox proportional hazards modelling was utilized to 154 evaluate associations between preoperative and intraoperative variables with failure. P-values less 155 than 0.05 were considered statistically significant. Statistical analyses were performed using 156 RStudio Version 4.1.1 (RStudio, Boston, MA). 157

- 158
- 159 **Results**

160 Demographics

Sixty-two patients met study inclusion criteria during initial database review. Forty-four of 161 the 62 identified patients (71%) agreed to participate in the study and were consented prior to data 162 analysis. The remaining 18 patients who underwent MAT during the timeframe were unable to be 163 reached for follow-up (n=16) or refused study participation (n=2). The mean age of included 164 patients was 16.1 ± 1.9 years and 33 patients were female (75%) (Table 1). The mean BMI was 165 23.4 ± 3.4 kg/m² and patients were followed for a mean of 9.5 ± 3.8 years (range: 5.0-17.7). Medial 166 meniscus transplantation was performed in 9 patients (20%) and lateral meniscus transplantation 167 was performed in 35 patients (80%). Eleven patients (25%) were diagnosed with a discoid lateral 168 meniscal tear and were referred to the senior author for management for symptoms refractory to 169

170	meniscectomy. All included patients had undergone a prior meniscectomy at a mean time 1.2 \pm
171	0.9 years (range: $0.1 - 3.9$) prior to MAT. Nine (20%) and two (4.5%) patients were previously
172	treated with anterior cruciate ligament reconstruction (ACLR) or osteochondritis dissecans
173	fixation prior to primary MAT, respectively. In 27 (61%) patients, a concomitant procedure was
174	performed, the most common being osteochondral allograft transplantation (OCA) ($n = 14/44$,
175	32% [lateral formal condyle (LFC), $N = 13$; medial femoral condyle (MFC), $N = 1$), autologous
176	chondrocyte implantation (ACI) (n = 8/44, 18% [MFC, N = 1; LFC, N = 6; patellar and LFC, N =
177	1]) and ACLR (n = 6/44, 14%) (Table 1).

178

Patient-Reported Outcomes 179

Significant improvements in all patient-reported outcome measures for each postoperative 180 timepoint analyzed (1-year, 2-year, and most-recent follow-up) were appreciated when compared 181 to preoperative scores (p < .05, Figure 1, Supplemental Table 1). Of note, while Lysolm scores at 182 final follow-up were significantly improved when compared to baseline (p = .018), a significant 183 decrease was observed in Lysholm scores when comparing final follow-up outcome scores to 184 scores at 2 years postoperatively (p = .048). This was an isolated finding as no other PROM 185 demonstrated a significant decline in scores between the short-term and most recent follow-up 186 timepoints. 187

188

Clinically Significant Outcomes 189

Three patients met criteria for failure prior to 5-year follow-up and were excluded from CSO 190 analysis. Thresholds for achieving CSOs were calculated with all PROMs achieving an area under 191

the curve (AUC) \geq .700 for PASS (Supplemental Table 1). Conversely, only the Lysholm form had an acceptable AUC for SCB (AUC = .750). The proportions of patients achieving CSOs were calculated (Table 3). A majority of patients achieved MCID and PASS for each PROM analyzed. SCB for the subjective IKDC form was achieved by a minority of patients (N = 5/16, 31%). At the time of final follow-up, however, 80% of patients stated they were satisfied with their overall postoperative condition.

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200 <u>Reoperations and failures</u>

Fourteen patients (32%) underwent a reoperation on the ipsilateral knee at an average 5.0 201 202 \pm 4.3 years (range: 0.8 – 14.0) following MAT. Eleven patients underwent a reoperation and did not subsequently meet criteria for failure (Table 1). The most common reoperation was partial 203 meniscectomy (n=9), followed by subtotal (n=2) and total meniscectomy (n=2), and meniscal 204 repair (n=1). Two patients who had a subsequent total meniscectomy underwent RMAT at a mean 205 2.68 years following meniscectomy. The remaining two patients who received a subtotal 206 meniscectomy have not undergone subsequent surgeries at a mean 6.2 years of follow-up. An 207 additional three patients underwent non-meniscal reoperations including synovectomy and lysis of 208 adhesions (n=1), articular cartilage debridement (n=1), and OCA (n=1) at a mean 1.3 years 209 210 following MAT.

211

212 Female patients were significantly more likely to undergo a meniscal reoperation (p =
213 0.039). Kaplan-Meier estimations were performed to assess probability of requiring a reoperation.

Overall probability of being reoperation-free at 1, 2, 5, and 10 years was 96%, 91%, 80%, 71%, 214 respectively (Figure 2A). Three (7%) patients met criteria for failure at an average of 3.8 ± 1.1 215 years (range: 2.8-4.9) post-transplant. All three patients were treated with revision MAT. No 216 patients were converted to arthroplasty or received subsequent realignment osteotomy. MAT 217 survival free from failure at 1, 2, 5, and 10 years was 100%, 100%, 93%, 93%, respectively (Figure 218 219 2B). Log-rank testing found no difference in survival free from failure based on sex (p = .710), meniscus laterality (p = .485), or concomitant surgery of any kind (p = .234), including corrective 220 alignment procedures (p = .696), ACLR (p = .525), or cartilage procedures (p = .718). Log-rank 221 222 testing also found no difference in reoperation rates based on sex (p = .110), meniscus laterality (p= .910), or concomitant surgery of any kind (p = .118), including corrective alignment procedures 223 (p = .324), ACLR (p = .483), or cartilage procedures (p = .765). Reoperations and failure data 224 stratified by meniscal laterality is present in Supplemental Table 2. 225

226

227 **Discussion**

The primary findings of this study were Lysholm, IKDC subjective form, and all KOOS 228 subscale outcome measures were significantly improved in the short-term postoperative period 229 and were sustained at final, midterm follow-up. Additionally, a majority of patients achieved 230 MCID and PASS for all questionnaires examined at midterm follow-up. The survivorship free 231 232 from reoperation and failure at 5 years post-operatively was 71% and 93%, respectively. The results demonstrate that MAT is a safe, effective treatment option for indicated patients in this 233 select population even when performed at the time of other major concomitant procedures. This 234 235 study is of important clinical relevance given the overall lack of substantial literature documenting 236 MAT outcomes in the adolescent patients at midterm follow-up.

Previous data regarding meniscal allograft transplantation is largely based on outcomes in adult populations.^{25,26} Recent meta-analyses of adult MAT outcomes literature indicates MAT 238 survivability ranges between 85.8% and 89.2% depending on compartment laterality at five-years 239 and 74% at 10-years postoperatively.^{25,26} Despite the bevy of literature on MAT outcomes in 240 adults, only a handful of case series studying MAT within adolescent populations exist, and those 241 242 available are limited by short-term follow-up. Nevertheless, reported outcomes demonstrate a level of functional success and survivability consistent with the findings in adult populations and the 243 results of this case-series. 244

A previous clinical report of adolescent patients who underwent MAT performed by the 245 senior author was conducted in 2016. This study included 32 patients (mean age 15.4 years) at a 246 minimum 2-year follow-up.⁶ Similar to the present study, a significant proportion of patients (48%) 247 underwent a concomitant procedure at the time of MAT. Substantially fewer patients underwent 248 meniscal reoperation (6%) or any reoperation (22%) when compared to the present study (32%) 249 250 and 39%, respectively). Additionally, no revision MATs were performed in the 2016 paper by Riboh et al., while three were performed at a mean 3.8 years following primary MAT in the present 251 study. 252

More recently in 2021, Smith et al. completed a review of a public database of 49 children's 253 hospitals to evaluate the epidemiology of MAT procedures in patients 25 years old or younger.⁴ 254 255 The final cohort included 67 patients with a mean age of 16.6 years, though final follow-up timeframes were not reported. The most common concomitant procedures performed at time of 256 257 MAT were ACLR (n = 11; 16.4%) and either osteochondral graft or ACI (n = 6; 9%). The authors 258 reported a reoperation rate of 18% (n = 12) among the included patients, with reoperations occurring a median of 224 days (range: 47-949 days) after transplantation. However, because the 259

data was sourced from a public registry, direct patient follow-up could not be obtained, which
prevented analysis of both clinical outcomes and additional treatment that may have occurred at a
facility outside of the database.

263 In a study of 23 patients undergoing MAT at a median age of 17, Middleton et al. reported that all patient-reported outcomes measured had significantly improved at most-recent follow-up 264 265 (mean, 3.8 years) when compared to preoperative baseline. None of the 23 patients in the cohort met criteria for failure, defined as graft removal or meniscectomy resulting in functional 266 deficiency.¹⁰ Additionally, Kimura et al. reported successful outcomes in a small cohort of 6 267 pediatric patients at a mean follow-up of 5.2 years using meniscal allografts donated from adult 268 patients undergoing total knee arthroplasty. No failures were reported in this cohort, defined more 269 conservatively as graft removal, complications, or subsequent surgery.²⁷ With failure definitions 270 varying widely between studies, it is hard to compare failure rates. The present study reported a 271 7% failure rate due to revision MAT which is comparable to adult studies at similar time points.²² 272

Chondral concomitant procedures were performed with notable frequency in the present 273 study as well as in the studies published by both Riboh et al.⁶ and Smith et al.⁴ It is worth noting 274 that previous studies have demonstrated that concomitant chondral procedures did not affect 275 outcomes in MAT.²³ Similar results were redemonstrated in the present investigation as log-rank 276 testing found no difference in survivability or reoperation based on concomitant surgery of any 277 kind (Figure 2). The variable closest to significance for association with reoperations was sex (p =278 .110). There was a significantly greater frequency of female patients who underwent reoperation 279 and did not subsequently fail (p = .039). At a minimum 2-year follow-up, Frank et al., using the 280 281 same database as the present study, found no increase in reoperations between male and female patients under 40 years (p = .902). Female patients were, however, more likely to undergo 282

subsequent revision MAT (p = .003). The association between sex on reoperation rates has not been seen in other, large database studies.^{28,29}

Additionally, a majority of cases in the present study were of lateral MAT (80%). In a 2018 285 286 meta-analysis of midterm and long-term studies, Bin et al. found no differences in failure between lateral and medial MAT.²⁵ However, superior Lysholm and pain scores were observed for lateral 287 288 MAT. The study was limited, however, by a small sample size as only three studies reported PROMs (42 medial and 51 lateral MATs). Further, various allograft fixation and processing 289 techniques were included. More recent studies with fresh-frozen allografts using bone fixation 290 291 have not appreciated similar differences in PROMs.^{30,31} Lastly, the present study used the bone bridge using the trough or bridge-in-slot technique, which is one option among many others (such 292 as soft tissue fixation, bone plugs, keyhole bone bridge).^{32,33} Optimal fixation techniques remain 293 debated as comparisons are difficult due to frequency of concomitant procedures, different patient 294 populations, and surgeon experience. 295

The present study reported that 11 patients (25%) were diagnosed with a discoid lateral 296 meniscal tear as the primary indication for MAT. These tears have been shown to increase the risk 297 of early onset osteoarthritis and subsequent total knee arthroplasty (TKA).⁷ In 2016, Ramme et al. 298 demonstrated that in young women aged between 25-30 with torn discoid lateral menisci, MAT 299 reduced pain, improved function, and was more effective at delaying TKA when compared to 300 301 partial meniscectomy. Partial meniscectomy required TKA at an average of 12.5 years, compared to 17.3 years in the subset of patients that underwent MAT. However, MAT was more resource 302 intensive, requiring \$14,470 of care-related expenditure compared to \$10,430 in partial 303 meniscectomy patients.³⁴ The findings of Ramme et al., in conjunction with the outcomes of the 304

present study, suggests that MAT can serve as a viable treatment modality in the management ofdiscoid lateral meniscal tears.

307 Lastly, patients undergoing MAT may return to competitive and recreational sports, as tolerated and as has been previously reported by our group.^{35,36} Although the present study did not 308 examine return-to-sport data, it is crucial to have comprehensive discussions about these data to 309 310 establish expectations with adolescent patients. In the senior author's practice, patients are informed about the high risk of retear if they return to pivoting sports, which may necessitate 311 312 subsequent surgeries. Ultimately, thorough preoperative discussions that focus on patient-specific goals help patients assess the risk associated with a traumatic meniscal retear and the need to limit 313 their involvement in sports. 314

Overall, this study illustrates that midterm outcomes of MAT in this cohort of adolescent patients significantly increases in functional outcomes, with low probability of graft failure. However, further research of MAT outcomes in this demographic remains warranted. A primary area of future study should evaluate the chondroprotective effects of MAT in adolescents through imaging studies, especially in light of inconclusive results in studies of adult populations.³⁷ Longterm reporting of MAT outcomes remains necessary to better elucidate overall treatment prognosis and consequences of the procedure.

322

323 Limitations

This study is not without limitations. Performance bias may be evident as the cohort of patients in this study were treated by a single surgeon who operates a high-volume MAT practice. Therefore, the outcomes in this cohort may not be generalizable for adolescent patients undergoing

primary MAT at other institutions. Given the overall follow-up rate of 71%, a possible transfer 327 bias is present, as the eighteen patients who were not included may differ in outcomes from patients 328 in the present study, thus potentially confounding the results. Additionally, 61% of patients in this 329 cohort experienced concomitant procedures, which may impact functional outcomes, reoperation 330 rates, and graft survivability compared to patients who are treated with isolated MAT. The present 331 332 study was also confined to a retrospective analysis of functional scores and re-operations. Advanced imaging of each patient at final follow up to evaluate chondral or meniscal status of the 333 knee was not routinely obtained. Therefore, conclusions regarding about physeal safety following 334 MAT are limited.⁶ 335

336

337 Conclusion

Primary MAT in adolescent patients resulted in significant and durable functional improvements
at mid- to long-term follow-up. At an average of 9.5 years postoperatively, meniscal reoperation
rate was 25% while graft survival free of revision MAT was 93%. Adolescents undergoing
MAT demonstrated similar functional outcomes and graft survivability when compared to
available adult MAT literature.

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454 Figure Legends

Figure 1: Preoperative and postoperative patient-reported outcomes following primary meniscal 455 allograft transplantation were collected. Mean scores for Lysholm (Lys), International Knee 456 Documentation Committee (IKDC) subjective form, and Knee Injury and Osteoarthritis Outcome 457 458 Score (KOOS) subscale questionnaires increased at 1-year, 2-year, and most-recent follow-up timepoints when compared to baseline scores. ADL, activities of daily living; QOL, quality of life; 459 SXS, symptoms. Error bars represent standard deviation. * indicates a mean significantly greater 460 compared to preoperative scores (p < 0.05). ** indicates a significant increase compared to most-461 recent scores (p = 0.048). 462

463

Figure 2: Kaplan-Meier survival analysis for (A) meniscal reoperation and (B) graft failure
werestratified by sex. Overall probability of being reoperation-free at 1, 2, 5, and 10 years were
96%, 91%, 80%, 71%, respectively. Survival free from failure at 1, 2, 5, and 10 years were 100%,
100%, 93%, 93%, respectively. Log-rank testing demonstrated no significant difference in survival
distributions between male and female sex for reoperation and failure (p = 0.110 and 0.710,
respectively).

	F	Post-operative Stat	us	p-va	alues		
Characteristic	None, N = 30 ¹	Reop, N = 11 ¹	Fail, N = 3 ¹	None vs. Reop ²	None vs. Failure ²		
Sex (female)	20 (67%)	11 (100%)	2 (67%)	0.039	>0.999		
Age (years)	16.29 ± 1.11	15.90 ± 1.16	15.53 ± 2.16	0.303	0.594		
вмі	24.1 ± 2.9	21.6 ± 4.3	23.4 ± 3.6	0.072	0.810		
Lateral meniscus transplanted	24 (80%)	8 (73%)	3 (100%)	0.660	>0.999		
Concomitant procedure	21 (81%)	5 (45%)	1 (33%)	0.051	0.136		
OCA	12 (40%)	1 (9%)	1 (33%)	0.127	>0.999		
ACI	5 (17%)	3 (27%)	0 (0%)	0.660	>0.999		
нто	1 (3%)	0 (0%)	0 (0%)	>0.999	>0.999		
DFO	4 (13%)	0 (0%)	0 (0%)	0.559	>0.999		
MFX	3 (10%)	0 (0%)	0 (0%)	0.551	>0.999		
ACLR	5 (17%)	1 (9%)	0 (0%)	>0.999	>0.999		

Table 1: Demographics and Intraoperative Variable

¹categorical variables listed as n (% of respective cohort); continuous variables listed as mean \pm

SD

²Fisher's exact test; Wilcoxon rank sum test

OCA, osteochondral allograft transplantation; ACI, autologous chondrocyte implantation; HTO, high tibial osteotomy; DFO, distal femoral osteotomy; MFX, microfracture; ACLR, anterior cruciate ligament reconstruction.

Reoperations consist of any surgery related to the transplanted meniscus (arthroscopic evaluation due to meniscal symptoms, debridement, meniscectomy) but excludes revision meniscal allograft transplantation. Failure defined as revision meniscal allograft transplantation or conversion to unicompartmental or total knee arthroplasty.

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Table 2: Patient-Reported Outcomes in Patients Free of Graft Failure at 5-Year Minimum Follow-Up

Characteristic	Baseline	1-Year	P-value	2-Year	P-value	Most-Recent	P-value
Lysholm	49 ± 24	77 ± 18	.005	84 ± 19 ↓	.001	74 ± 16	.018
IKDC	38 ± 19	57 ± 27	.042	80 ± 19	<.001	71 ± 20	.005
ADL	72 ± 21	92 ± 10	.014	95 ± 10	.001	94 ± 8	.006
Pain	59 ± 19	84 ± 11	.016	87 ± 10	<.001	85 ± 11	.001
QOL	29 ± 20	51 ± 20	.001	65 ± 26	.007	60 ± 24	.006
Sport	31 ± 20	69 ± 13	.003	74 ± 15	.021	64 ± 25	.002
SXS	57 ± 16	77 ± 14	.001	82 ± 13	<.001	73 ± 17	.004

473 Patient-reported outcome scores at preoperative, 1-year, 2-year, and a minimum 5-year follow-

subjective form, and Knee osteoarthritis and outcome score (KOOS) subscales (ADL, activities

of daily living; Pain; QOL, quality of life; Sport; SXS, symptoms). Significance determined by

477 Wilcoxon Signed-Rank testing. Continuous variables listed as mean \pm SD.

478 \downarrow - significantly greater (p = .048), compared to most-recent follow-up

Table 3: Clinically Significant Outcomes at Five-Year Minimum						
Characteristic	$N = 41^{1}$					
MCID						
IKDC	10 / 16 (62%)					
Lysholm	8 / 13 (62%)					
Pain	11 / 17 (65%)					
Symptoms	11 / 19 (58%)					
Sport	12 / 14 (86%)					
ADL	9 / 17 (53%)					
QOL	10 / 17 (59%)					

⁴⁷⁴ up. Questionnaires included are Lysholm, International Knee Documentation Committee (IKDC)

Table 3: Clinically Significant Outcomes at Five-Year Minimum						
Characteristic $N = 41^{1}$						
PASS						
IKDC	25 / 33 (76%)					
Lysholm	23 / 29 (79%)					
Pain	26 / 32 (81%)					
Symptoms	26 / 32 (81%)					
Sport	24 / 32 (75%)					
ADL	24 / 31 (77%)					
QOL	25 / 31 (81%)					
SC	CB					
IKDC	5 / 16 (31%)					
Lysholm	3 / 13 (23%)					
Pain	7 / 17 (41%)					
Symptoms	9 / 19 (47%)					
Sport	7 / 14 (50%)					
ADL	6 / 17 (35%)					
QOL	10 / 17 (59%)					
Detions more included if they did not feil union	to fine montallow we					

¹Patients were included if they did not fail prior to five-year follow-up.

ADL, activities of daily living; IKDC, International Knee Documentation Committee subjective form; KOOS, Knee Injury and Osteoarthritis Outcome Score; MCID, minimal clinically important difference; PASS, patient acceptable symptomatic state; QOL, quality of life; SCB, substantial clinical benefit.

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Supplemental Table 1: Thresholds for Achieving Clinically Significant							
Outcomes							
	Threshold	SN	SP	AUC			
	MCID						
IKDC	11.5						
Lysholm	11.9						
KOOS							
Pain	12.0						
Symptoms	11.1						
Sport	13.2						
QOL	18.4						
ADL	13.9						
PASS							

IKDC	54.2	95.0	100.0	0.995
Lysholm	61.0	89.4	100.0	0.964
KOOS				
Pain	72.2	100.0	100.0	0.999
Symptoms	57.1	89.4	75.0	0.822
Sport	42.5	94.7	100.0	0.999
QOL	46.8	94.7	100.0	0.986
ADL	88.1	89.5	100.0	0.973
	<u>SCB</u>			
IKDC	<u>SCB</u> 30.6	60.0	85.7	0.686
IKDC Lysholm	<u>SCB</u> 30.6 24.5	60.0 75.0	85.7 71.4	0.686 0.750
IKDC Lysholm KOOS	<u>SCB</u> 30.6 24.5	60.0 75.0	85.7 71.4	0.686 0.750
IKDC Lysholm KOOS Pain	<u>SCB</u> 30.6 24.5 16.5	60.0 75.0 83.3	85.7 71.4 50.0	0.686 0.750 0.583
IKDC Lysholm KOOS Pain Symptoms	<u>SCB</u> 30.6 24.5 16.5 17.6	60.0 75.0 83.3 66.7	85.7 71.4 50.0 62.5	0.686 0.750 0.583 0.562
IKDC Lysholm KOOS Pain Symptoms Sport	<u>SCB</u> 30.6 24.5 16.5 17.6 40.0	60.0 75.0 83.3 66.7 40.0	85.7 71.4 50.0 62.5 100.0	0.686 0.750 0.583 0.562 0.667
IKDC Lysholm KOOS Pain Symptoms Sport QOL	<u>SCB</u> 30.6 24.5 16.5 17.6 40.0 9.4	60.0 75.0 83.3 66.7 40.0 50.0	85.7 71.4 50.0 62.5 100.0 71.4	0.686 0.750 0.583 0.562 0.667 0.464

Legend: Thresholds for minimal clinically important difference (MCID), patient acceptable symptomatic state (PASS), and substantial clinic benefit (SCB) were calculated using both revision and primary meniscal allograft transplantation cohorts. Patient reported outcome measures analyzed included Lysholm, subjective International Knee Documentation Committee (IKDC), and Knee Injury and Osteoarthritis Outcome Score (KOOS) subscales. ADL, activities of daily living; AUC, area under the curve; QOL, quality of life; SN, sensitivity; SP, specificity.

Supplemental Table 2: Reoperation and Failure Data Stratified by Meniscal							
Laterality							
Variable	Lateral ¹	Medial ¹	p-value				
Meniscal reoperation type			>0.999 ²				
None	24 (69%)	6 (67%)					
Partial meniscectomy	7 (20%)	2 (22%)					
Total Meniscectomy	3 (9%)	1 (11%)					
Repair	13%)	0 (0%)					
Time to reoperation	4.3 ± 4.1	8.0 ± 4.6	0.291 ³				

Supplemental Table 2: Reoperation and Failure Data Stratified by Meniscal							
Laterality							
Variable	Lateral ¹	Medial ¹	p-value				
Failure	3 (9%)	0 (0%)	>0.999 ²				
¹ categorical variables listed as n (% of transplantation group); continuous var ² Fisher's exact test ³ Wilcoxon rank-sum test	respective med iables listed as 1	ial or lateral menis mean ± standard de	cal allograft eviation.				



Preoperative and Postoperative Patient Reported Outcomes



