

# Return to Preoperative Function After Autologous Cartilage Implantation of the Knee in Active Military Servicemembers

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**Background:** Autologous chondrocyte implantation (ACI) has been shown to provide adequate durability, pain relief, and improved long-term functional outcomes in the average patient, but proof of its efficacy in individuals with greater than average physical demands is scarce. Further knowledge is required to understand which patients may benefit from ACI and to identify which risk factors are associated with failure to return to the preinjury activity level.

**Purpose:** To determine the occupational outcomes, rates of reoperation, and variables predictive of suboptimal outcomes after ACI.

**Study Design:** Case series; Level of evidence, 4.

**Methods:** All active-duty military servicemembers in the United States who underwent ACI of the knee between 2004 and 2014 were identified. Demographic information, injury characteristics, surgical variables, and clinical and surgical outcomes were extracted from the medical record. Univariate and multivariate analyses were used to determine significant independent predictors of clinical and surgical failures.

**Results:** A total of 90 patients (91 knees) met the inclusion criteria. The cohort was predominantly male (86%), with a mean age of  $34.5 \pm 6.3$  years (range, 20-50 years). The most common location of the articular cartilage lesion was the patellofemoral compartment (54 lesions, 59%), and the mean Outerbridge grade and size were  $3.8 \pm 0.4$  and  $4.00 \pm 2.77$  cm<sup>2</sup> (range, 1.2-15.0 cm<sup>2</sup>), respectively. A total of 72 patients (79%) had at least 1 previous knee procedure. Nearly three-quarters of patients (71%) underwent concomitant procedures. At a mean follow-up of  $59.9 \pm 27.1$  months (range, 24.0-140.1 months), 60% of our patients reported significant improvement in knee pain and did not require further surgical intervention. Multivariate analysis identified age <30 years as the only significant independent predictor of both clinical ( $P = .011$ ) and overall failure ( $P = .014$ ). Moderate-demand military occupational specialties ( $P = .036$ ), exclusive involvement of the patellofemoral compartment ( $P = .045$ ), and use of a periosteal patch ( $P = .0173$ ) were additionally found to be independent predictors of surgical failure.

**Conclusion:** Treatment of articular cartilage defects of the knee with ACI in physically active young individuals can return nearly two-thirds of individuals to daily activity with decreased pain and improved function. Risk factors for failure after ACI surgery were age younger than 30 years, lower demand occupation, exclusive involvement of the patellofemoral compartment, prior microfracture, and use of a periosteal patch.

**Keywords:** autologous chondrocyte implantation; ACI; athletic activity; knee; articular cartilage

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Focal articular cartilage defects of the knee can cause significant pain and disability and can ultimately lead to degenerative arthritis if left untreated. While there are a number of treatment options, each must be tailored to the demographic and functional demands of the patient. Management may also differ significantly depending on lesion size, location, knee alignment, meniscal integrity, and ligament status. There are various operative options that can be divided into 3 broad categories: palliative (eg, chondroplasty), reparative (eg, microfracture and subchondral drilling), and restorative (eg, osteochondral autograft transplantation [OATS], autologous chondrocyte

implantation [ACI], and osteochondral allograft transplantation [OCA]).<sup>12,13</sup>

ACI was first published as a restorative treatment option in 1994.<sup>2,8</sup> Compared with alternative techniques, ACI offers the ability to re-create hyaline-like cartilage with reproducible longevity and wear characteristics.<sup>22</sup> ACI has been shown to reduce pain in young active populations and improve knee function by returning athletes to their pre-injury level of activity at rates between 73% and 84%.<sup>3,10,20</sup>

The active-duty military population is similarly subject to regular daily lower extremity demands. However, ACI has not been evaluated on a large scale in the military population. When deciding to intervene surgically, the military orthopaedic surgeon must consider the techniques that will allow patients to return to their pre-injury level of function. Accordingly, the aim of the current study was to evaluate the short- to midterm surgical and functional outcomes of United States (US) military service members after ACI of the knee. Additionally, we sought to determine the rates of reoperation and variables predictive of suboptimal outcomes.

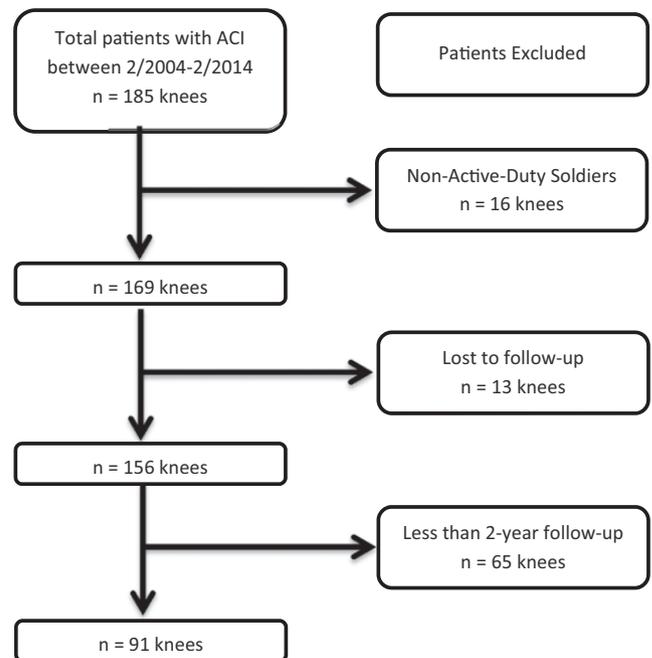
## METHODS

We conducted a retrospective review of the Management Analysis and Reporting Tool (M2) database to identify all tri-service US active-duty military servicemembers who underwent ACI (Current Process Terminology [CPT] code 27412) between the years 2004 and 2014 in the Military Health System. The M2 is an established health care management database that can be used to perform clinical outcome research related to a variety of knee<sup>6,24</sup> injuries and other musculoskeletal conditions.<sup>12</sup>

Inclusion criteria were patients with primary ACI for a high-grade, Outerbridge grade III or IV chondral defect of the knee, minimal 2-year clinical follow-up, and active-duty military status at the time of surgery. Exclusion criteria were applied to patients with nonmilitary or retired status at time of surgery, clinical follow-up of less than 2 years, hybrid chondral procedure other than ACI, previous ipsilateral ACI of the knee, and/or those with insufficient documentation (Figure 1).

Demographic data, including patient identifiers, age, sex, military rank, and branch of service, were initially extracted from the M2 database. These identifiers were cross-referenced with the military electronic medical record (Armed Forces Health Longitudinal Technology Application [AHLTA] and e-Profile [version 3.17; Medical Operational Data System]), and chart review was conducted to confirm the diagnosis and procedure. Further analysis yielded additional patient-based (laterality, race, body mass index [BMI], military rank, military occupational specialty [MOS], tobacco use), lesion-related (location, size, grade), and surgical variables (type of patch, prior and concomitant procedures [eg, tibial tubercle or proximal tibial osteotomy]).

Rank was stratified into junior enlisted (E1-5) versus senior enlisted (E6-9) and commissioned officers (O1-6). Junior enlisted soldiers are less able to modify their daily



**Figure 1.** Primary autologous chondrocyte implantation (ACI) patient selection diagram.

physical activities than their senior enlisted or officer counterparts. Junior enlisted rank has furthermore been correlated with lower socioeconomic and education status, as these servicemembers are typically younger, earn lower salaries, and more often have only secondary school education level at the time of enlistment.<sup>19</sup> MOS designations were classified collectively as combat arms (CA), combat support (CS), or combat service support (CSS) based on inherent occupational demands. CA and CS occupational specialties are actively engaged in conflict and generally require greater functional demands than those of the more ancillary CSS MOS.

The primary outcomes of interest were (1) clinical failure, defined as inability to return to active military duty or medical separation from the military secondary to persistent disability attributable to the operative extremity and (2) surgical failure, constituting revision chondral procedure or conversion to arthroplasty due to persistent symptomatology after index ACI. Overall failure was defined as meeting the criteria for clinical and/or surgical failure. Complications and reoperations other than revision procedures were also recorded.

## Statistical Analysis

Summary statistics including means and standard deviations were reported for continuous variables. Frequencies were calculated for categorical variables. Bivariate logistic regression analysis was performed to test the significance of demographic, injury-related, and surgical risk factors as predictors of clinical, surgical, and overall (either clinical or surgical, or both) failure. Multivariate regression analysis was subsequently performed for any

TABLE 1  
Patient Demographics and Clinical Profile<sup>a</sup>

Patients/knees, n	90/91
Follow-up, mo, mean (SD)	59.9 (27.1)
Age, y, mean (SD)	34.5 (6.3)
Sex, n (%)	
Men	78 (86)
Women	13 (14)
Body mass index (kg/m <sup>2</sup> ), mean (SD)	25.5 (3.6)
Service, n (%)	
Army	81 (89)
Navy/Air Force/Marines/Coast Guard	10 (11)
Rank, n (%)	
Junior enlisted	12 (13)
Senior enlisted	62 (68)
Officer	17 (19)
Military occupational specialty, n (%)	
Combat arms	25 (28)
Combat support	24 (26)
Combat service support	42 (46)
Tobacco, n (%)	
Yes	35 (38)
No	56 (62)

<sup>a</sup>Percentages were calculated per total number of knees (N = 91).

risk factor with  $P < .05$  on initial bivariate analysis. Significant independent predictors were defined as those with  $P < .05$  and 95% confidence interval (CI) for the odds ratio (OR) exclusive of 1.0 on multivariable logistic regression analysis. ORs with corresponding 95% CIs and  $P$  values are reported for each variable. All statistical calculations were performed using SAS version 9.3 (SAS Institute Inc).

## RESULTS

### Demographics

A total of 90 patients (91 knees) met the inclusion criteria (Table 1). The mean patient age was  $34.5 \pm 6.3$  years (range, 20-50 years), and 86% were male. The mean BMI was  $25.5 \pm 3.6$  kg/m<sup>2</sup>; 35% of patients had a BMI  $\geq 30$  kg/m<sup>2</sup>. More than one-third of patients (38%) reported regular use of tobacco products. MOS was divided nearly equally between CA/CS (54%) and CSS (46%).

### Surgical Variables

With regard to lesion characteristics, nearly three-quarters (77%) were isolated, and most (59%) commonly involved the patellofemoral compartment (Table 2). The mean overall lesion Outerbridge grade and size were  $3.8 \pm 0.4$  and  $4.00 \pm 2.77$  cm<sup>2</sup> (range, 1.2-15.0 cm<sup>2</sup>), respectively. The largest lesions were located in the patellofemoral compartment ( $5.61 \pm 4.13$  cm<sup>2</sup>). Most patients had undergone at least 1 previous knee procedure (79%), which was most often either chondroplasty (37%), meniscal debridement (21%), and/or microfracture (17%) (Table 3).

A type I/III collagen xenograft bilayer membrane was used in the majority (69%) of cases, while the remainder

TABLE 2  
Articular Cartilage Injury Characteristics<sup>a</sup>

Outerbridge grade, mean (SD)	3.8 (0.4)
Defects on index knee, n (%)	
Multiple ( $\geq 2$ locations)	21 (23)
Single	70 (77)
Defects by location, n (%)	
Patella	40 (44)
Trochlea	35 (38)
Medial femoral condyle	28 (31)
Lateral femoral condyle	10 (11)
Medial tibial plateau	1 (1)
Lateral tibial plateau	1 (1)
Defects by knee compartment, n (%)	
Patellofemoral compartment	54 (59)
Bipolar patellofemoral defects <sup>b</sup>	11 (12)
Tibiofemoral compartment	27 (30)
Bipolar tibiofemoral compartment <sup>c</sup>	1 (1)
Total defect surface area, cm <sup>2</sup> , mean (SD)	4.00 (2.77)
Total defect surface area by knee compartment, cm <sup>2</sup> , mean (SD)	
Patellofemoral	5.61 (4.13)
Tibiofemoral	5.36 (4.52)
Defect surface area, cm <sup>2</sup> , n (%)	
$\geq 6$	24 (26)
4-5.9	28 (31)
2-3.9	25 (28)
$< 2$	14 (15)

<sup>a</sup>Percentages were calculated per total number of knees (N = 91).

<sup>b</sup>Bipolar patellofemoral lesions represent lesions present on the articular surface of both the patella and the trochlea.

<sup>c</sup>Bipolar tibiofemoral lesions represent lesions present on the articular surface of both the femoral condyle and the adjacent tibial plateau.

used a periosteal patch harvested from the ipsilateral tibia (31%). Most patients (71%) underwent concomitant procedures, including tibial tubercle osteotomy (60%), high tibial osteotomy (7%), and meniscal allograft transplantation (4%) (Table 3). The indications for concomitant anteromedializing tibial tubercle osteotomy when performed included lateral subluxation of the patella, distal and lateral patellar chondral defects, and/or tibial tuberosity trochlear groove distance  $> 15$  mm. Any patient who underwent a high tibial osteotomy was found to have a concurrent varus mechanical alignment with a chondral defect located within the medial compartment of the knee.

### Outcomes

At a mean follow-up of  $59.9 \pm 27.1$  months (range, 24.0-140.1 months), 60% of patients had significant improvement of their knee pain and did not require any further surgical intervention. Furthermore, 82% of patients reported minimal to no pain at final follow-up. Nearly one-third (35%) of patients were considered clinical failures due to the inability to return to military duty or were medically separated due to persistent, rate-limiting symptoms in the operative extremity. At the time of final follow-up, 10 patients (11%) qualified as surgical failures. Of these, 6

TABLE 3  
Prior and Concomitant Knee Procedures<sup>a</sup>

Procedure	n (%) <sup>b</sup>
Any knee surgery prior to ACI procedure	72 (79)
Chondroplasty	34 (37)
Meniscal debridement	19 (21)
Microfracture	15 (17)
Anterior cruciate ligament reconstruction	12 (13)
Meniscal repair	5 (6)
Lateral release	5 (6)
High tibial osteotomy	4 (4)
Osteochondral autograft transplantation	2 (2)
Meniscal allograft transplantation	1 (1)
Medial collateral ligament reconstruction	1 (1)
Posterior cruciate ligament reconstruction	1 (1)
Any concomitant surgery during ACI procedure	65 (71)
Tibial tubercle osteotomy	55 (60)
High tibial osteotomy	6 (7)
Meniscal allograft transplantation	4 (4)
Lateral release	3 (3)
Microfracture	1 (1)
Anterior cruciate ligament reconstruction	1 (1)
Medial patellofemoral ligament reconstruction	1 (1)

<sup>a</sup>ACI, autologous chondrocyte implantation.

<sup>b</sup>Percentages were calculated per total number of knees in each group.

(7%) patients required revision chondral procedure (3 microfracture, 3 OATS), and an additional 4 (4%) individuals underwent subsequent knee arthroplasty (3 total knee arthroplasty, 1 patellofemoral arthroplasty).

A total of 16 complications (18%) occurred in 16 patients and included graft hypertrophy (8%, n = 7) with first-generation technique, arthrofibrosis (6%, n = 5), superficial infection requiring irrigation and debridement (2%, n = 2), as well as proximal tibial fracture (1%, n = 1) and nonunion at the site of concomitant tibial tubercle osteotomy (1%, n = 1). Thirty patients (33%) underwent a total of 36 reoperations, composed mostly of hardware removal (12%, n = 11), second-look diagnostic arthroscopy (8%, n = 7), periosteal graft debridement (8%, n = 7), and manipulation under anesthesia (6%, n = 5) (Table 4). Of the 7 patients with second-look arthroscopy, indications were persistent pain despite normal examination and diagnostic findings (n = 2), arthrofibrosis (n = 2), mechanical symptoms (n = 2), and loose body formation (n = 1). However, none of these second-look arthroscopies documented any graft hypertrophy at the previous ACI location. Both cases of infection required a single repeat irrigation and debridement and were resolved with oral antibiotic use. One patient was noncompliant with premature weightbearing and activity outside of his brace, and he sustained a displaced proximal tibial fracture at 8 weeks postoperatively after combined patellar ACI and tibial tubercle osteotomy that required open reduction and internal fixation (ORIF) with a lateral proximal tibia plate. One additional patient had a persistent symptomatic nonunion that underwent curettage and revision ORIF with successful union.

TABLE 4  
Outcomes and Complications After ACI<sup>a</sup>

Outcome/complication	n (%) <sup>b</sup>
Clinical outcomes	
Persistent pain	16 (18)
Return to duty	54 (59)
Deployment	23 (25)
Clinical failure	
Medical separation	32 (35)
Surgical failure	10 (11)
Arthroplasty	4 (4)
Chondral procedure	6 (7)
Microfracture	3 (3)
Osteochondral autograft transplant	3 (3)
Other reoperation besides surgical failures	
Removal of hardware	11 (12)
Manipulation under anesthesia	5 (6)
Periosteal graft hypertrophy debridement	7 (8)
Diagnostic arthroscopy	7 (8)
Irrigation and debridement	2 (2)
ORIF nonunion	1 (1)
ORIF proximal tibia fracture	1 (1)
Meniscal debridement	1 (1)
Lateral release	1 (1)
Complications	
Periosteal graft hypertrophy	7 (8)
Periosteal patch	6 (7)
Collagen xenograft bilayer membrane	1 (1)
Arthrofibrosis	5 (6)
Infection	2 (2)
Osteotomy nonunion	1 (1)
Proximal tibia fracture	1 (1)

<sup>a</sup>ACI, autologous chondrocyte implantation; ORIF, open reduction internal fixation.

<sup>b</sup>Percentages were calculated per total number of knees (N = 91).

## Risk Factors

Univariate logistic regression analysis yielded several risk factors for clinical, surgical, and overall failure (Table 5). Subsequent multivariate analysis identified age <30 years as the only significant independent predictor of both clinical (OR, 4.01; 95% CI, 1.37-11.73) and overall failure (OR, 3.84; 95% CI, 1.32-11.16) (Table 6). Moderate demand, CS MOS (OR, 19.59; 95% CI, 1.21-316.12), exclusive involvement of the patellofemoral compartment (OR, 6.98; 95% CI, 1.04-46.80), and use of a periosteal patch (OR, 10.75; 95% CI, 1.52-75.81) were additionally found to be independent predictors of surgical failure.

## DISCUSSION

Resolution of symptoms and return to activity after ACI for focal articular cartilage defects of the knee present a significant challenge. Our study presents a retrospective case series of physically active young servicemembers with symptomatic focal articular cartilage defects of the knee treated with ACI. Within this analysis, we demonstrate that age <30 years, combat support, exclusive involvement of the patellofemoral compartment, prior microfracture, and use of a periosteal patch are predictors of failure after ACI.

TABLE 5  
Univariate Logistic Regression Analyses of Clinical, Surgical, and Overall Failures<sup>a</sup>

Variable	Clinical Failure, OR (95% CI)	P Value	Surgical Failure, OR (95% CI)	P Value	Overall Failure, OR (95% CI)	P Value
Age <sup>b</sup>	0.92 (0.86-0.99)	.033 <sup>c</sup>	0.99 (0.89-1.10)	.841	0.94 (0.88-1.01)	.087
<30 y	2.99 (1.14-7.82)	.026 <sup>c</sup>	2.03 (0.52-7.94)	.307	2.86 (1.10-7.48)	.031 <sup>c</sup>
Female sex	1.71 (0.52-5.62)	.374	0.64 (0.07-5.51)	.684	1.97 (0.60-6.44)	.261
Tobacco use	2.11 (0.87-5.09)	.098	1.08 (0.28-4.12)	.916	1.84 (0.78-4.36)	.167
BMI <sup>b</sup>	1.02 (0.90-1.14)	.810	0.97 (0.81-1.17)	.748	1.00 (0.89-1.13)	.972
≥30 kg/m <sup>2</sup>	1.17 (0.48-2.87)	.731	0.77 (0.19-3.20)	.718	1.07 (0.45-2.58)	.878
Army branch of service	5.58 (0.67-46.21)	.111	1.13 (0.13-9.94)	.916	6.85 (0.83-56.61)	.074
Military occupational specialty						
Combat arms	0.85 (0.30-2.42)	.757	7.81 (0.82-74.34)	.074	1.08 (0.39-2.99)	.877
Combat support	1.08 (0.38-3.06)	.885	10.79 (1.18-98.80)	.035 <sup>c</sup>	1.16 (0.42-3.23)	.775
Combat service support	—	—	—	—	—	—
Exclusive patellofemoral lesion(s)	1.18 (0.45-3.09)	.741	4.86 (1.11-21.25)	.036 <sup>c</sup>	1.71 (0.67-4.38)	.263
Multiple (≥2) lesions	1.53 (0.57-4.16)	.402	0.82 (0.16-4.17)	.807	1.19 (0.44-3.21)	.725
Total size, cm <sup>2b</sup>	1.00 (0.99-1.01)	.331	1.00 (0.99-1.01)	.338	1.00 (0.99-1.01)	.766
≥6	5.08 (0.93-27.75)	.061	0.55 (0.07-4.28)	.568	2.12 (0.52-8.67)	.298
4-5.9	4.50 (0.84-23.99)	.078	0.46 (0.06-3.68)	.465	1.88 (0.47-7.45)	.372
2-3.9	2.33 (0.41-13.20)	.338	1.14 (0.18-7.19)	.887	1.41 (0.34-5.81)	.638
<2	—	—	—	—	—	—
Any previous procedures	2.39 (0.72-7.93)	.156	2.57 (0.31-21.66)	.385	3.00 (0.91-9.93)	.072
Previous microfracture procedure	0.62 (0.18-2.15)	.454	4.24 (1.03-17.48)	.046 <sup>c</sup>	1.02 (0.33-3.17)	.970
Concomitant procedure	0.82 (0.32-2.10)	.677	0.35 (0.09-1.33)	.123	0.55 (0.22-1.38)	.200
Periosteal patch	1.29 (0.52-3.25)	.584	6.67 (1.58-28.16)	.010 <sup>c</sup>	1.86 (0.76-4.60)	.177

<sup>a</sup>BMI, body mass index; OR, odds ratio.

<sup>b</sup>Age, BMI, and total size were analyzed as continuous variables.

<sup>c</sup>Statistically significant at *P* < .05.

TABLE 6  
Multivariate Analysis for Risk Factors for Clinical, Surgical, and Overall Failures<sup>a</sup>

Outcome	Odds Ratio	95% CI	P Value
Clinical failure			
Age <30 y <sup>a</sup>	4.01	1.37-11.73	.011
Surgical failure			
Military occupational specialty			
Combat arms	10.58	0.73-153.58	.084
Combat support	19.59	1.21-316.12	.036
Combat service support	—	—	—
Exclusive patellofemoral lesion(s)	6.98	1.04-46.80	.046
Previous microfracture procedure	2.06	0.33-12.80	.437
Periosteal patch	10.75	1.52-75.81	.017
Overall failure			
Age <30 y <sup>b</sup>	3.84	1.32-11.16	.014

<sup>a</sup>Selected multivariate regression analysis performed on variables identified to be *P* < .05 on univariate analysis.

<sup>b</sup>Age was analyzed as a continuous variable.

The military represents an ideal population for evaluating the outcomes of restorative chondral procedures of the lower extremities. On average, active-duty patients are younger, maintain regular lower extremity physical activity, and, ultimately, desire return to a high level of function. At a mean 59.9-month follow-up, the current

study demonstrated that 60% of patients who had ACI for a symptomatic focal chondral defect of the knee were able to return to military duty and their occupational demands. This rate is slightly lower than previously reported success rates of 73% to 84%<sup>3,9,11,17,21,22,25</sup> but may be commensurate with the intense physical profile of this demographic and the limited time frame allotted for rehabilitation within the military framework.

A systematic review by Campbell et al<sup>3</sup> reported on cartilage repair in the athlete's knee. The authors sought to determine which surgical techniques returned athletes to competition and correlated patient- and defect-specific factors that were associated with successful return to sport. The authors reported an 84% rate of returning athletes to sport participation after ACI for a focal symptomatic cartilage lesion. They concluded that younger patients with smaller chondral defects, shorter duration of symptoms, knee without history of microfracture, and those who participated in a dedicated rehabilitation protocol would have a better prognosis. Additionally, a retrospective case series assessing return to sports after ACI<sup>21</sup> showed that 73.1% of patients were able to resume some form of sports activity. The authors of this study further analyzed the different sports activities that patients were involved in prior to and after ACI. Their findings revealed that after ACI, patients transitioned from high-impact, start-stop sports to less demanding, low-impact and endurance activities.

When critically evaluating our clinical failure rate of 35%, analogous findings from the civilian literature reveal

an inability to return to previous sports activity in 16% to 27% of patients after ACI of the knee.<sup>3,11,21</sup> Within our cohort, we attribute this mainly to the increased activity-related demands placed on our population and the larger overall size of articular lesions ( $5.06 \pm 3.78 \text{ cm}^2$ ) treated. Additionally, it is exceedingly difficult to expeditiously restore military patients to active duty status due to a number of unique considerations, including the inherent strenuous occupational demands, physically active lifestyle, limited timeline allotted for full recovery (ie, 6-12 months depending on service branch), and variable capacity for activity modifications or work restrictions. An additional challenge in the military framework is defining realistic expectations for not only the patient but also the patient's superiors within the chain of command. A patient's specific occupation or function (ie, MOS) is often one of the most critical variables in determining return to unrestricted duty. While protective physical limitations can be imposed by the treating orthopaedic surgeon, a medical separation may still be initiated for a given patient if the anticipated recovery time exceeds certain established thresholds, despite a favorable prognosis for return to unrestricted duty and continued clinical improvement at up to 24 months postoperatively.<sup>21</sup> If patients within the military system were afforded greater latitudes for modified daily activity or occupational specialty postoperatively, we believe that this would translate to more favorable short- to midterm overall outcomes after ACI.

We also found that patients younger than 30 years were more prone to overall failure postoperatively. This is in stark contrast to previous studies that have largely found that younger patient age is associated with superior outcomes.<sup>3,17</sup> Nawaz et al<sup>17</sup> reported on the mid- to long-term results of skeletally mature individuals undergoing ACI, with 78% and 51% graft survival at 5 and 10 years, respectively, and younger patients serving as the ideal candidates for ACI due to limited degeneration of adjacent compartments. In the current population, these disparate findings may reflect disproportionately greater physical demands associated with junior military status as well as the importance of peak physical fitness in career progression. Furthermore, younger servicemembers often have broader occupational skill sets, lesser degrees of specialized training, and limited mission-critical leadership roles, making them easier to replace. While failure in the general population may be more attributable to age-related decline in cartilage quality and viability, younger military patient populations are more likely to fail due to more activity-related reasons and inability to modify their occupational activities.

In the current series, patellofemoral lesions accounted for 2 of 3 defects, and this subgroup was found to be at increased risk for surgical failure when compared with articular lesions exclusive to the tibiofemoral compartment. In addition, these defects were larger than lesions located elsewhere ( $5.61 \text{ cm}^2$  vs  $5.36 \text{ cm}^2$ ). Traditionally, success after ACI for lesions within the patellofemoral joint has been much lower when compared with other locations within the knee (60%-80% vs 80%-95%, respectively).<sup>2,9,15,18,22,23</sup> The general consensus is that

the treatment options are less reliable for restoration of patellofemoral lesions. This is due to the unique articular topography and the biomechanical forces that create a formidable challenge to articular cartilage restoration,<sup>15</sup> particularly in the presence of bipolar or so-called "kissing" lesions ( $n = 12$ ) that are typically contraindicated for cell-based treatment. Although not demonstrated in our results, other studies have shown that correction of rotational malalignment and/or offloading of symptomatic bipolar defects with a tibial tubercle osteotomy can improve the outcomes after ACI in the patellofemoral joint by correcting the abnormal shear and compressive forces.<sup>5,7,23</sup>

Prior marrow stimulation techniques (microfracture, abrasion chondroplasty) were associated with unsatisfactory surgical outcomes on univariate analysis but failed to achieve statistical significance on multivariate testing. Current trends in the surgical management of articular cartilage defects show comparatively lower rates of cell-based chondral restoration and osteochondral reconstruction when compared with other palliative techniques, such as microfracture or limited chondroplasty, as the initial surgery of choice.<sup>4,13,16</sup> Ostensibly, this practice of initially selecting a microfracture or limited chondroplasty first could complicate outcomes for revision cartilage surgery with subsequent ACI, in large due to reactive, osseous overgrowth with intralesional osteophyte formation and disruption of the normal subchondral plate architecture.<sup>14</sup> Minas et al<sup>14</sup> compared ACI after previous marrow stimulation techniques with individuals undergoing primary ACI for symptomatic chondral defects. Although objective functional knees scores were not obtained in the study, the authors demonstrated an increased failure rate of ACI after marrow stimulation (26%) versus a control group (8%). Zaslav et al,<sup>25</sup> in a prospective multicenter study, further demonstrated that although there is an increased failure rate for patients who had ACI after failed microfracture, these individuals can still have clinically significant improvements in both pain and function. Similar functional outcome scores and failure rates were found in studies with mid- to long-term follow-up.<sup>15,20</sup> Minas et al<sup>15</sup> showed a 62% survival rate of ACI grafts at 15 years among patients with failed marrow stimulation surgery (microfracture, subchondral drilling, or abrasion chondroplasty), and patients with prior microfracture (44%) had significantly worse survivorship than those with primary ACI (79%) for chondral defects. Pestka et al,<sup>20</sup> in a retrospective matched-pair study, analyzed ACI after failed microfracture treatment versus those who underwent ACI alone for articular cartilage defects of the knee. Their findings demonstrated significant differences between groups. The group with prior microfracture had lower survival rates (75% vs 96%, respectively), and lower Knee Injury and Osteoarthritis Outcome Score (KOOS) pain and KOOS activities of daily living subscores.

Graft hypertrophy requiring debridement occurred in 7 (8%) knees from the entire cohort, including 1 patient with a second-generation technique. The association between graft hypertrophy and first-generation ACI technique is well established in the literature, and patients treated

with a periosteal patch may require secondary debridement in up to 15% to 50% of cases.<sup>9,15,18</sup> Additionally, periosteal patch coverage with ACI may also portend an increased risk of surgical or clinical failure. Subgroup analysis of those patients in the current cohort with the first-generation ACI group revealed that 6 patients (21%) required arthroscopic debridement, with 4 of these 6 meeting the criteria for overall failure (microfracture, n = 1; patellofemoral arthroplasty, n = 1; medical discharge, n = 2). We theorize that repeated insult to the hyaline-like cartilage regenerated at the graft site may disrupt the architectural reorganization and structural integrity, increasing the risk for abnormal loading patterns and subsequent failure.

A large percentage of our cohort (71%) had a concomitant surgical procedure at the time of ACI, which may serve as a source of confounding. In particular, it may be difficult to ascertain whether outcomes were attributable to adjunctive procedures, such as offloading osteotomy, or chondral restoration alone. As with any retrospective series from the military setting, certain inherent limitations must be acknowledged. Foremost, the definition of clinical failure may be overly stringent and lack external validity to other less-active or sedentary patient populations. Similarly, knee-related medical discharge may reflect inadequate rehabilitation or compliance with commonly accepted postoperative limitations typically observed at up to 12 to 18 months after ACI surgery,<sup>1</sup> and this could artificially inflate rates of clinical failure at short-term follow-up. Additionally, there is the potential for nonresponder bias with patients who complete their military service obligation and exit the military or choose to follow up in the civilian health care network. Given its retrospective nature, the current study also lacks validated patient-reported outcomes and objective knee measures, which would increase the generalizability of our findings and allow for quantification of pain and functional outcomes.

## CONCLUSION

This study evaluated the short- to midterm clinical and occupational outcomes of ACI in a physically active military population and demonstrated that nearly two-thirds of individuals can return to daily activity with improved pain and function after ACI surgery. Specific risk factors for failure after ACI surgery were age younger than 30 years, lower demand occupation, exclusive involvement of the patellofemoral compartment, prior microfracture, and use of a periosteal patch. Future research is needed to provide long-term outcomes; however, the present findings allow us to appropriately counsel the expected outcomes for a young active patient who would undergo ACI for a symptomatic chondral defect of the knee.

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