



Increased Age, Weight, and Body Mass Index Increase the Likelihood of Total Knee Replacement Following Anterior Cruciate Ligament Reconstruction

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Disclosures

- David Flanigan, MD is a consultant for and receives research support from Vericel, Zimmer, and Smith & Nephew; is a consultant for ConMed-MTF and DePuy Mitek; and receives research support from MTF, Histogenics, Aesculap, Cartiheal, Anika Therapeutics, and Moximed
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Current Literature

- Incidence of end-stage knee OA that requires TKA is substantially greater in patients with prior ACLR¹
- Aging impairs the structural integrity of type II collagen and chondrocyte repairing capacity, resulting in greater susceptibility to developing knee OA²
- High BMI increases knee joint loading and inflammatory responses that accelerate cartilage degeneration and development of knee OA²
- Elevated systemic indices of inflammation have been linked to the presence and severity of OA^{3,4}



Purpose

- To investigate the association of age, BMI, and systemic indices of inflammation with TKA following ACLR.
- We hypothesized that an older age, higher BMI, and elevated systemic indices of inflammation associate with an increased likelihood of TKA following ACLR.



Materials and Methods

- Retrospective chart review identified 45 ACLR patients between 2009 and 2022 that met inclusion criteria
- Control groups were matched (1:1:1) based on sex and date of ACLR (± 1.0 y)

| Cases N=15 | Control 1 N=15 | Control 2 N=15 |
|---|---|---|
| <ul style="list-style-type: none">• Patients with CBC data and a subsequent TKA procedure performed for knee OA after ACLR. | <ul style="list-style-type: none">• Patients that underwent ACLR without a later documented knee OA diagnosis | <ul style="list-style-type: none">• Patients that underwent ACLR with later documented knee OA diagnosis but no TKA procedure |



| <div>TABLE 1</div> <div>Demographics</div> | | | | |
|--|--------------|------------------------|-------------------------|--------------|
| | ACLR and TKA | ACLR without OA or TKA | ACLR with OA but no TKA | Significance |
| N (m/f) | 15 (8/7) | 15 (8/7) | 15 (8/7) | N/A |
| Age at ACLR, y | 46.8 (5.8) | 40.3 (18.3) | 37.7 (15.8) | p = 0.05 |
| Height, m | 1.75 (0.17) | 1.68 (0.10) | 1.70 (0.17) | p = 0.15 |
| Body mass, kg | 108 (33) | 75.0 (23.7)* | 90.7 (29.7)* | p < 0.01 |
| BMI, kg/m² | 35.3 (10.8) | 25.0 (4.8)* | 26.2 (9.6)* | p < 0.01 |
| Additional procedures at ACLR, n/15 | 12/15 | 10/15 | 11/15 | p = 0.71 |
| ACLR to CBC, mos | 39.4 (39.1) | 12.6 (31.2)* | 17.0 (22.6) | p = 0.02 |
| ACLR to OA, mos | 32.2 (48.5) | NA | 22.2 (79.7) | p = 0.98 |
| ACLR to TKA or final follow-up, mos | 54.8 (49.5) | 88.7 (84.9) | 90.8 (83.1) | p = 0.08 |
| Data presented as median (interquartile range [IQR]) or counts (n). *p < 0.05 vs Cases. BMI, body mass index (kg/m²). NA, not applicable | | | | |

| <div>TABLE 2</div> <div>Cell Count Data</div> | | | | |
|---|--------------|------------------------|-------------------------|--------------|
| | ACLR and TKA | ACLR without OA or TKA | ACLR with OA but no TKA | Significance |
| WBC (K/μL) | 6.33 (2.16) | 7.10 (3.64) | 6.10 (3.77) | p = 0.24 |
| RBC (K/μL) | 4.72 (0.35) | 4.49 (0.71) | 5.03 (0.90) | p = 0.15 |
| Platelets (K/μL) | 266 (83) | 242 (36) | 230 (100) | p = 0.62 |
| Neutrophils (K/μL) | 3.74 (1.85) | 4.30 (3.65) | 3.43 (2.68) | p = 0.29 |
| Lymphocytes (K/μL) | 1.79 (0.48) | 1.90 (0.47) | 1.80 (0.95) | p = 0.37 |
| Monocytes (K/μL) | 0.59 (0.34) | 0.40 (0.31) | 0.50 (0.16) | p = 0.79 |
| PLR | 140 (62) | 122 (77) | 137 (50) | p = 0.86 |
| MLR | 0.26 (0.16) | 0.24 (0.18) | 0.27 (0.19) | p = 0.19 |
| NLR | 1.96 (.69) | 2.29 (1.70) | 2.34 (1.24) | p = 0.99 |
| SIRI | 1.20 (0.80) | 1.01 (0.10) | 1.24 (0.89) | p = 0.79 |
| SII | 495 (158) | 693 (580) | 482 (458) | p = 0.52 |
| Data presented as median (interquartile range [IQR]) or counts (n). WBC, white blood cell count. RBC, red blood cell count. PLR, platelet-to-lymphocyte ratio. MLR, monocyte-to-lymphocyte ratios. NLR, neutrophil-to-lymphocyte ratio. SIRI, systemic inflammation response index. SII, systemic immune- | | | | |

TABLE 3

Pearson Product Moment Linear Correlation Coefficients

| | Age at ACLR | | BMI | | ACLR to CBC | |
|--------------------|-------------|----------|-------|----------|-------------|----------|
| | r | Sig | r | Sig | r | Sig |
| WBC (K/μL) | 0.04 | p = 0.78 | -0.01 | p = 0.97 | -0.23 | p = 0.14 |
| RBC (K/μL) | 0.02 | p = 0.90 | 0.00 | p = 0.99 | -0.02 | p = 0.92 |
| Platelets (K/μL) | 0.24 | p = 0.12 | 0.26 | p = 0.09 | 0.05 | p = 0.74 |
| Neutrophils (K/μL) | 0.03 | p = 0.84 | -0.16 | p = 0.31 | -0.25 | p = 0.10 |
| Lymphocytes (K/μL) | 0.01 | p = 0.94 | 0.40 | p = 0.01 | -0.01 | p = 0.97 |
| Monocytes (K/μL) | 0.06 | p = 0.68 | 0.15 | p = 0.31 | -0.05 | p = 0.75 |
| PLR | 0.10 | p = 0.53 | -0.11 | p = 0.48 | 0.04 | p = 0.82 |
| MLR | 0.03 | p = 0.82 | -0.19 | p = 0.22 | 0.01 | p = 0.96 |
| NLR | -0.03 | p = 0.87 | -0.31 | p = 0.04 | -0.07 | p = 0.65 |
| SIRI | 0.01 | p = 0.96 | -0.14 | p = 0.35 | -0.15 | p = 0.34 |
| SII | 0.12 | p = 0.45 | -0.16 | p = 0.31 | -0.12 | p = 0.44 |

N=45. WBC, white blood cell count. RBC, red blood cell count. PLR, platelet-to-lymphocyte ratio. MLR, monocyte-to-lymphocyte ratios. NLR, neutrophil-to-lymphocyte ratio. SIRI, systemic inflammation response index. SII, systemic immune-inflammatory index.

| <div>TABLE 4</div> <div>Logistic Regression Odds Ratios</div> | | |
|--|---------------------------------------|--------------|
| | Odds Ratio (95% Confidence Intervals) | Significance |
| | | |
| Age at ACLR | 1.42 (1.08, 1.86) | p = 0.01 |
| BMI | 1.48 (1.12, 1.96) | p < 0.01 |
| PLR | 1.31 (0.96, 1.79) | p = 0.09 |
| MLR | 1.23 (0.89, 1.70) | p = 0.22 |
| NLR | 1.14 (0.92, 1.35) | p = 0.27 |
| SIRI | 0.99 (0.62, 1.57) | p = 0.96 |
| SII | 0.67 (0.43, 1.06) | p = 0.08 |
| Cases vs CON1 and CON2, response variable: TKA. WBC, white blood cell count. RBC, red blood cell count. PLR, platelet-to-lymphocyte ratio. MLR, monocyte-to-lymphocyte ratios. NLR, neutrophil-to-lymphocyte ratio. SIRI, systemic inflammation response index. SII, systemic immune-inflammatory index. | | |

Discussion and Conclusions

- Increasing age and BMI were risk factors for undergoing TKA following ACLR.
- Systemic indices of the immune system and inflammation were not significant predictors of undergoing TKA after ACLR



Limitations

- The age of the Control 1 group is higher than the general age for the population undergoing ACLR
 - this is probably a result of older patients being more likely to undergo a CBC with differentials due to co-morbidities, other conditions, or routine physicals
- It is feasible that some of the patients in the Control 2 group were knee arthroplasty candidates but did not undergo knee arthroplasty for multiple potential reasons



Future Directions

- Larger sample size to:
 1. Control for potential confounding variables
 2. Confirm the association between increased BMI and age with TKA following ACLR
- Further research is needed to develop targeted interventions to potentially delay or prevent the progression to TKA following ACLR



References

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