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In vivo Measurement of Localized Tibiofemoral Cartilage Strains in Response to Dynamic Activity

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Abstract

Introduction—Altered local mechanical loading may disrupt normal cartilage homeostasis and play a role in the progression of osteoarthritis. Currently, there is limited data quantifying local cartilage strains in response to dynamic activity in normal or injured knees.

Purpose—The purpose of this study was to directly measure local tibiofemoral cartilage strains in response to a dynamic hopping activity in normal healthy knees. We hypothesize that local regions of cartilage will exhibit significant compressive strains in response to hopping, while overall compartmental averages may not.

Study Design—Controlled laboratory study.

Methods—Both knees of eight healthy subjects were MR imaged before and immediately after a dynamic hopping activity. Images were segmented and then used to create 3D surface models of bone and cartilage. These pre- and post-activity models were then registered using an iterative closest point technique to enable site-specific measurements of cartilage strain (defined as the normalized change in cartilage thickness before and after activity) on the femur and tibia.

Results—Significant strains were observed in both the medial and lateral tibial cartilage, with each compartment averaging a decrease of 5%. However, these strains varied with location within each compartment, reaching a maximum compressive strain of 8% on the medial plateau and 7% on the lateral plateau. No significant averaged compartmental strains were observed in the medial or lateral femoral cartilage. However, local regions of the medial and lateral femoral cartilage experienced significant compressive strains, reaching maximums of 6% and 3% respectively.

Conclusion—Local regions of both the femur and tibia experienced significant cartilage strains as a result of dynamic activity. An understanding of changes in cartilage strain distributions may help to elucidate the biomechanical factors contributing to cartilage degeneration after joint injury.

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Key Terms

knee; cartilage; osteoarthritis; strain; tibiofemoral; hopping; jumping; magnetic resonance imaging; stress test; biomechanics

Introduction

Normal physiologic loading of chondrocytes is important for the development and maintenance of healthy cartilage². However, altered mechanical stresses and strains, which may occur after ligament or meniscus injury^{4, 44}, may disrupt normal cartilage homeostasis and play a role in the progression of osteoarthritis (OA)^{1, 14}. Therefore, a number of studies have sought to understand how cartilage is loaded during activities of daily living. For example, some studies have used skin markers and in-ground force plates to estimate loading distributions between the medial and lateral compartments of the tibiofemoral joint^{7, 36}. Other studies have used biplanar radiographic images to measure joint kinematics and changes in joint space during activities such as walking and running^{18, 40}. To provide additional information regarding cartilage function, some recent studies have incorporated measurements of cartilage geometry with biplanar radiographic imaging to approximate acute cartilage strains during quasi-static loading and gait^{5, 6, 25, 44}. However, there is limited data directly measuring cartilage strains in response to dynamic activities such as running and jumping.

During activities of daily living, the knee joint experiences loads of several times body weight^{23, 29, 42}. As the cartilage deforms under these loads, water is extruded²⁸. After the load is removed, the low-permeability matrix of cartilage results in a time dependent recovery of deformation as fluid returns to the tissue. This sustained deformation can be measured using magnetic resonance (MR) imaging immediately before and after loading^{11, 13}. Previous studies have used this principle to measure changes in knee cartilage volume as a result of different dynamic activities, including running^{8, 20, 21, 30}, drop landings³⁰, and knee bends¹². While these studies contribute important information to the literature, compartmental volumetric changes may not be sensitive to variations in cartilage strain at different locations within the joint. Currently, there are limited site-specific measurements of cartilage strains in response to dynamic activities of daily living. Since mechanical loading can influence chondrocyte metabolism^{16, 22, 32, 37}, local measurements of cartilage strain are important to understanding normal cartilage physiology and may provide critical insights into the mechanisms leading to cartilage degeneration. Furthermore, such data may also provide baseline data for future studies evaluating cartilage loading in populations at high risk for the development of OA. Therefore, the objective of this study was to directly measure site-specific cartilage strains *in vivo* as a result of dynamic activity in normal knees. We hypothesized that local regions of cartilage will exhibit significant compressive strains in response to hopping, while compartmental averages of strain may not.

Methods

Institutional Review Board (IRB) approval was obtained before initiation of this study. Eight male subjects (mean age: 26.3 years, age range: 24–30 years) with no history of injury or surgery to either knee participated in this study. The mean body mass index (BMI) of these subjects was 22.8 kg/m² (range 21.1 to 25.1 kg/m²). Both knees of each subject were studied, for a total of 16 knees.

All subjects participated in this study on the morning of their testing day to reduce the potential for diurnal differences in cartilage thickness^{11, 48}. Subjects were asked to not perform any strenuous lower body activity the night prior to or the morning of their participation. Subjects first lay supine on a stretcher immediately outside of the MR suite for 45 minutes to minimize cartilage deformation prior to scanning^{6, 33, 39}. Subjects were then moved to a wheelchair without any weightbearing and transported to the MR suite where they underwent pre-activity imaging. Imaging was performed using a 3T scanner (Trio Tim, Siemens) with an eight-channel knee coil and the patient in a supine position with the knee relaxed. Sagittal plane images (field of view 16×16 cm, 512×512 pixels) of 1 mm thickness were generated using a double-echo steady state sequence (DESS, flip angle: 25°, TR: 17 ms, TE: 6 ms)^{11, 48}. Total scan time was approximately 9 minutes for each knee.

Subjects were transported via wheelchair to the hall adjacent to the MR suite and performed 60 single-legged hops of 0.6m on the tested leg. For each subject, the order of testing was alternated between left and right knees. During this hopping activity, the contralateral knee was kept in a flexed position off the ground. After completing the hopping activity, subjects were seated in a wheelchair and immediately transported into the MR suite to undergo post-activity MR imaging. The time from completion of the activity to the initiation of the post-activity MRI averaged 3.5 minutes (range: 3–4 minutes). After post-activity imaging was completed, testing was repeated on the contralateral knee.

The sagittal MR images were imported into solid modeling software (Rhinoceros, Robert McNeel and Associates, Seattle, Washington) and the outer bony cortex and cartilage surfaces of the femur and tibia were manually traced using non-uniform rational B-spline (NURBS) curves on each MR image by a single investigator (Figure 1a). These curves (Figure 1b) were then used to create a 3D surface mesh model of both bone and cartilage (Geomagic Studio, Morrisville, NC) (Figure 1c). This methodology has been previously validated for measuring cartilage thickness⁴⁴. Furthermore, a recent study indicated that the coefficient of repeatability for measuring cartilage thickness using this methodology is 0.03mm, which corresponds to a difference in cartilage thickness of 1%¹¹.

Cartilage thickness was calculated as the distance between each point on the cartilage surface and the nearest point on the bony surface. These calculations were used to generate cartilage thickness maps for each knee (Figure 2). The femoral and tibial bony surfaces of the pre-activity and post-activity models were aligned using an iterative closest point technique^{11, 33, 48}. This registration process enabled the site-specific measurement of cartilage strain using the pre-activity and post-activity models. Next, a grid system was created to sample strain at points spanning the femoral and tibial cartilage surfaces (Figure

3)^{11, 33, 48}. Nine evenly spaced grid points were created on each medial and lateral tibial plateau and 18 points were created on each medial and lateral femoral condyle. Strain was defined as the normalized change in thickness before and after activity and was calculated as the average across all of the points on model of the cartilage surface within a 2.5 mm radius of the respective grid point^{11, 48}. Overall compartmental strains were calculated as the average of the sampled points in a given tibial or femoral compartment.

Statistical analysis was performed using Statistica (StatSoft, Tulsa, OK). Single sample t-tests were used to determine whether averaged compartmental and local strains were different from zero. Differences were considered statistically significant where $p < 0.05$.

Results

Statistically significant compressive strains were exhibited in both the medial and lateral tibial plateaus as a result of single-legged hopping. In the medial compartment, there was an overall compressive strain of $5 \pm 1\%$ (mean \pm standard error of the mean) ($p < 0.001$). However, some regions exhibited significant local strains, while others did not (Figure 4a). For example, local compressive strain reached a maximum of $8 \pm 1\%$ ($p < 0.001$) in the lateral portion of the medial compartment. In the lateral compartment, an overall compressive strain of $5 \pm 1\%$ ($p < 0.001$) was exhibited. However, as was observed in the medial compartment, some regions exhibited significant strains while others did not. For example, local compressive strain reached a maximum of $7 \pm 1\%$ ($p < 0.001$) in the medial portion of the lateral compartment of the tibia (Figure 4b).

Overall, statistically significant averaged compartmental strains were not observed in either the medial or lateral femoral condyles as a result of single-legged hopping. In the medial compartment, an average compressive strain of $2 \pm 1\%$ ($p = 0.11$) was measured. However, local compressive strain reached a maximum of $6 \pm 2\%$ ($p < 0.01$) (Figure 5a) on the most anterior and medial portion of the medial femoral condyle. In the lateral compartment, an average compressive strain of $1 \pm 1\%$ ($p = 0.29$) was measured. However, there were significant compressive strains of $3 \pm 1\%$ ($p < 0.001$) (Figure 5b) in two local regions of the lateral femoral condyle.

Discussion

Mechanical loading plays an important role in normal cartilage homeostasis. Disruptions to the normal cartilage stress and strain distributions, which can occur after ligament or meniscus injury^{3, 4, 44}, can alter chondrocyte metabolism^{16, 22, 32, 37}, and potentially predispose the knee to degenerative changes^{1, 15}. Thus, baseline data characterizing the local mechanical environment of cartilage in response to in vivo loading conditions may provide valuable insights into the mechanisms contributing to the development and progression of OA. The present study measured variations in cartilage strains with location in the joint in response to a dynamic hopping activity. Specifically, we found significant averaged strains in both tibial compartments. However, some local regions of the tibia experienced significant strains, while other regions did not. Additionally, while no average compartmental changes were observed on either femoral condyle, some local regions of

femoral cartilage experienced significant strains. These results suggest that site-specific measurements of strain may provide important information regarding the local mechanical environment of cartilage that volumetric measurements of cartilage deformation may not be able to detect.

In comparison, our results regarding the average compartmental strains are consistent with previous studies measuring volumetric changes in cartilage in response to activity. For example, a previous study by Eckstein *et al*¹² measured compartmental volumetric changes as a result of a high impact single-legged jump landing activity. They found significant changes in the tibial cartilage volume in both compartments, with greater volume change in the lateral tibial plateau. They found no significant volumetric changes in the femoral cartilage in either compartment, which was similar to our finding of no significant averaged compartmental strains in the present study. Another study by Niehoff *et al*³⁰ investigating the effects of a high-impact double-legged jump landing also found significant average compartmental changes in cartilage thickness in both the medial (−2.2%) and lateral (−1.8%) tibial compartments. Again, no changes in the overall compartmental thickness were observed in either femoral condyle, which was also consistent with the averaged compartmental results of the present study. In contrast, the present study also observed local regions where significant cartilage strains were experienced on both the femur and the tibia. Together, these findings suggest that the site-specific measurements described in the present study may be more sensitive to variations in cartilage strain at different locations within the joint than volumetric deformations.

Our finding that there were fewer areas of significant local strains and lower compartmental strains exhibited in the femoral cartilage compared to the tibial cartilage may be explained by differences in loading and mechanical properties within the joint. For example, this may be a result of the larger area of femoral cartilage through which load is transmitted during dynamic flexion-extension movements^{25, 47}. In contrast, smaller areas of cartilage may be more consistently loaded throughout knee motion in the tibial cartilage. These differences in strain between the femur and tibia may also be due to the heterogeneity of mechanical properties of articular cartilage in the knee. Treppo *et al*⁴³ reported that femoral cartilage had a significantly greater equilibrium modulus and dynamic stiffness (at 0.1Hz) than tibial cartilage. As a result, the femoral cartilage may exhibit less strain than tibial cartilage when experiencing dynamic loads through the knee joint.

Interestingly, there were higher local compressive strains near the tibial spine compared to the peripheral regions in both tibial compartments. This pattern was most apparent in the middle region of both compartments. We believe that this may be due, in part, to the presence of the meniscus, which distributes load in the peripheral region of the tibial plateau^{19, 47}. Thus, regions of cartilage covered by the meniscus may experience less strain than regions where cartilage to cartilage contact occurs.

The single-legged hopping activity is one of many activities with which this method could be used to assess cartilage strains. For the purposes of this study, single-legged hops were chosen for multiple reasons. Single-legged hops are believed to place higher demands on the knee than walking or jogging and may represent dynamic movements such as jumping and

cutting during athletic activities³⁵. In addition, single-legged hops are used as a clinical tool to assess knee injury rehabilitation and as a return to sport evaluation^{31, 35}. Consideration was also given to the time sensitive nature in which the post-activity MR scan needed to be performed and the ability to ensure the safety of the subjects. This required that the activity could be performed in close proximity to the MR suite and in a controlled environment. We concluded that a hopping activity fulfilled these requirements. However, future studies might investigate the local strain patterns as a result of other dynamic activities, such as walking or running.

The method used in this study requires two measurements of cartilage: before the activity is performed and immediately after the activity is completed. Consequently, the measured deformation and calculated strain is the cumulative result of the entire activity. Therefore, though these are direct measurements of cartilage deformation, they may differ from the instantaneous cartilage strain induced throughout dynamic activity^{5, 6, 25}. Additionally, due to time required for subject transport, positioning and initiation of the imaging scan, some cartilage deformation may be recovered and our measurements may underestimate the true strains induced by the activity. However, we were able to begin imaging all knees within four minutes after the hopping activity was completed, which is consistent with prior studies³⁰. Furthermore, this amount of time appears to be relatively small compared to the time scale required for cartilage to completely recover from activity^{13, 45}. For example, a previous study indicated that 90 minutes was required for full volumetric recovery of the patellar cartilage after performing 100 deep knee bends¹³. Future studies might also use this methodology to evaluate the recovery of cartilage strain at various time points after exercise.

The current study investigated cartilage strains in young healthy male subjects only during a hopping activity. However, the strain patterns seen in normal healthy knees may be different between males and females due to anatomical differences between sexes, such as Q angle¹⁷. Future studies might evaluate differences in strain patterns between males and females. Additionally, future studies might use marker-based motion capture techniques^{38, 41} to investigate the influence of different motion patterns on cartilage deformation.

The ability to directly measure site-specific cartilage strains also has applications in the study of abnormal cartilage loading following ligament or meniscus injury. For example, this protocol could be used as a “stress test” to evaluate changes in the mechanical response of cartilage in these patient populations. Understanding how joint injury alters the mechanical environment of cartilage is important because these changes could directly affect chondrocyte metabolism^{16, 22, 32, 37} and disrupt normal cartilage homeostasis^{1, 14}. Thus, this methodology could be used to investigate cartilage strains in patients with ligament or meniscus injuries to provide insight into the mechanisms contributing to the high prevalence and early onset of post-traumatic OA in these patient populations^{26, 27, 34, 46}. Future studies may also seek to combine this method with other MR imaging modalities, such as T1 ρ or T2 mapping²⁴, which may allow for the measurement of zone-specific changes in cartilage strain due to variations in cartilage composition, structure, and mechanical properties with depth^{9, 10, 49}. The fusion of these modalities may also allow for the correlation of mechanical deformation and changes in water content as a result of dynamic activity³⁹.

Additionally, such data may also help to explain the interplay between altered strains and biochemical changes in cartilage and their role in the progression of OA²⁴.

In conclusion, our study demonstrated that local regions of tibiofemoral cartilage experienced significant compressive strains in response to a dynamic hopping activity. These local tissue strains varied from the compartmental averages, suggesting that site-specific measurements of cartilage strain may provide additional information regarding changes in the local mechanical environment that volumetric measures may not be sensitive enough to detect. In the future, this methodology and data could be used to evaluate the effects of soft tissue injuries (such as ligament or meniscus injuries) on cartilage strain distributions in response to dynamic activities of daily living. An understanding of these cartilage strain distributions may help to elucidate the biomechanical factors contributing to cartilage degeneration.

Acknowledgments

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Clinical relevance

Site-specific measurements of in vivo cartilage strains are important because altered loading is believed to be a factor contributing to the development and progression of osteoarthritis. Specifically, this methodology and data could be used to evaluate the effects of soft tissue injuries (such as ligament or meniscus tears) on cartilage strains in response to dynamic activities of daily living.

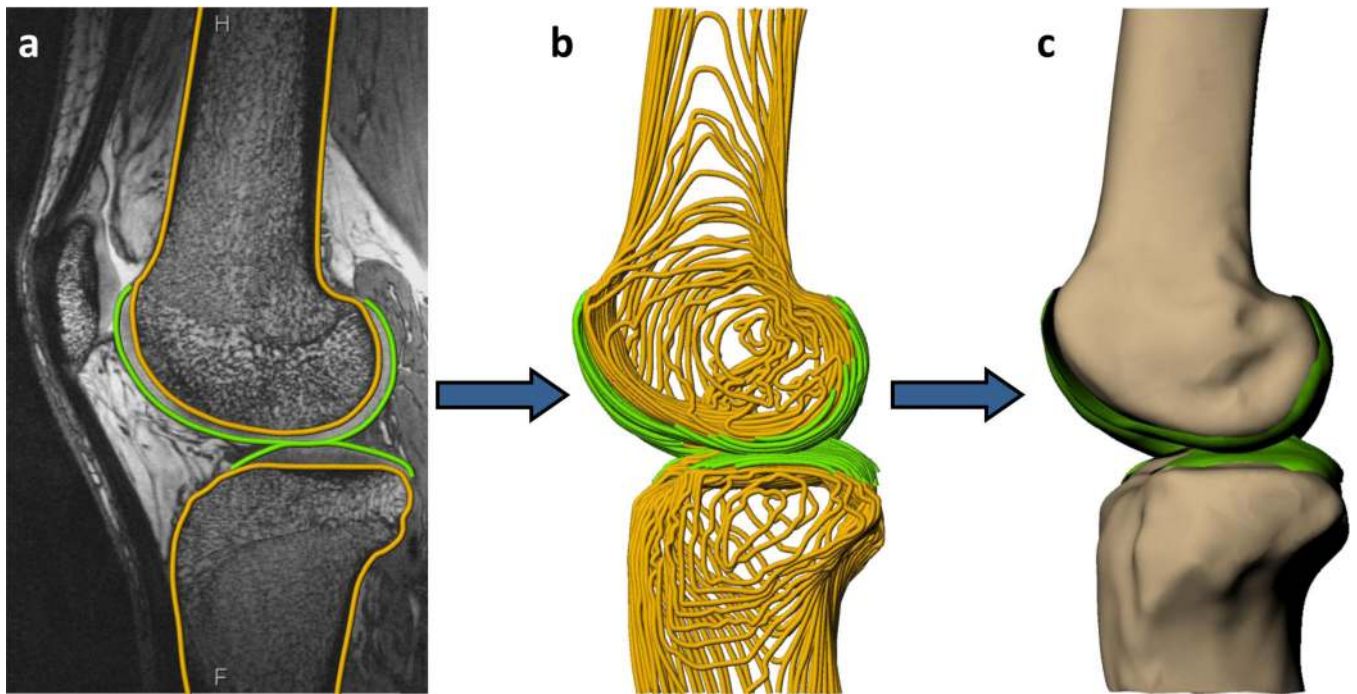


Figure 1.

a) The surfaces of the tibia, femur, and articular cartilage were segmented for each sagittal 3T MR slice, b) stacked to form a wireframe model, and c) converted to 3D surface mesh models.

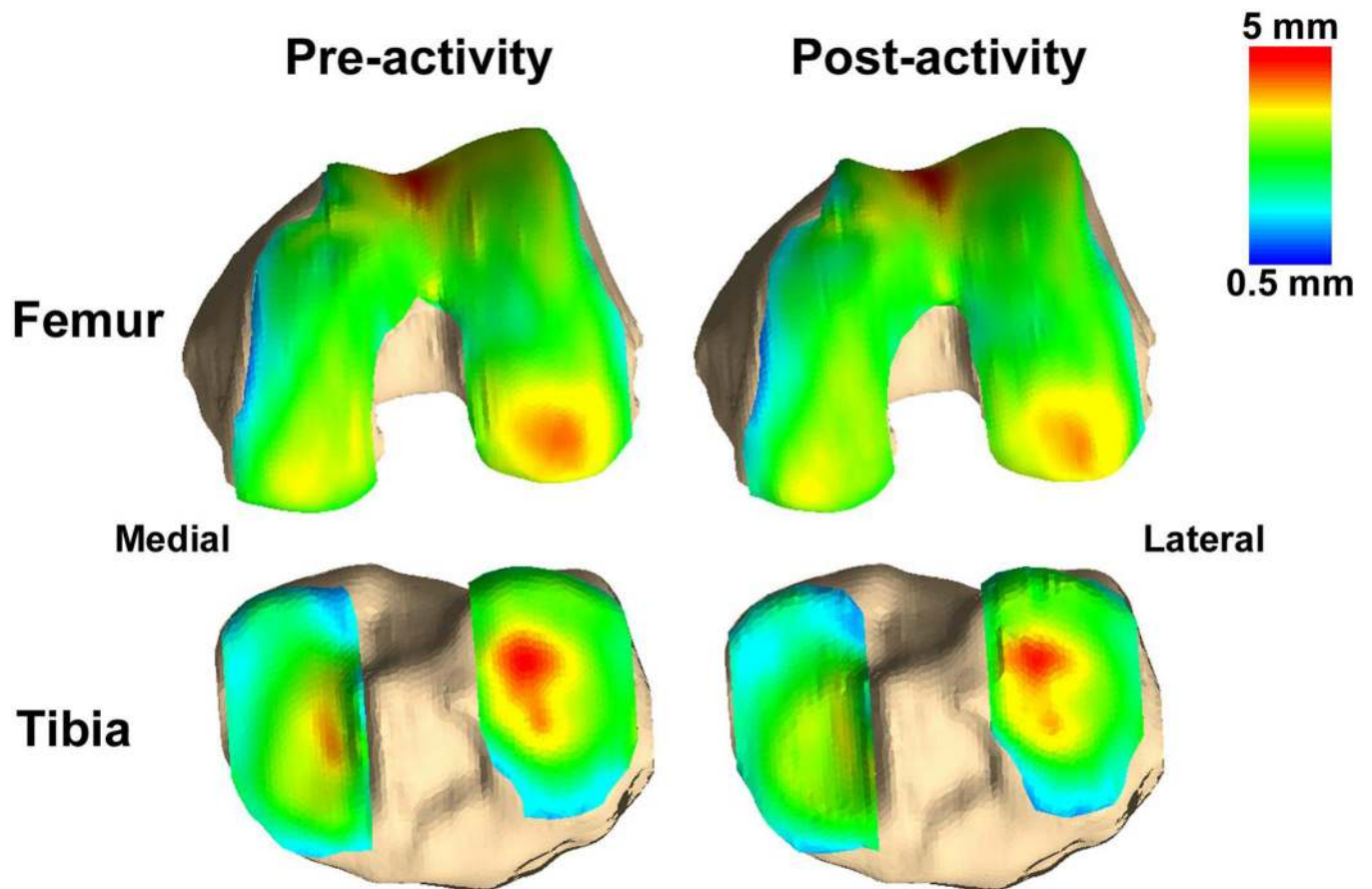


Figure 2. Representative cartilage thickness maps of pre-activity and post-activity femoral and tibial models. Thickness is represented in color, with thicker cartilage in red, and thinner cartilage in blue.

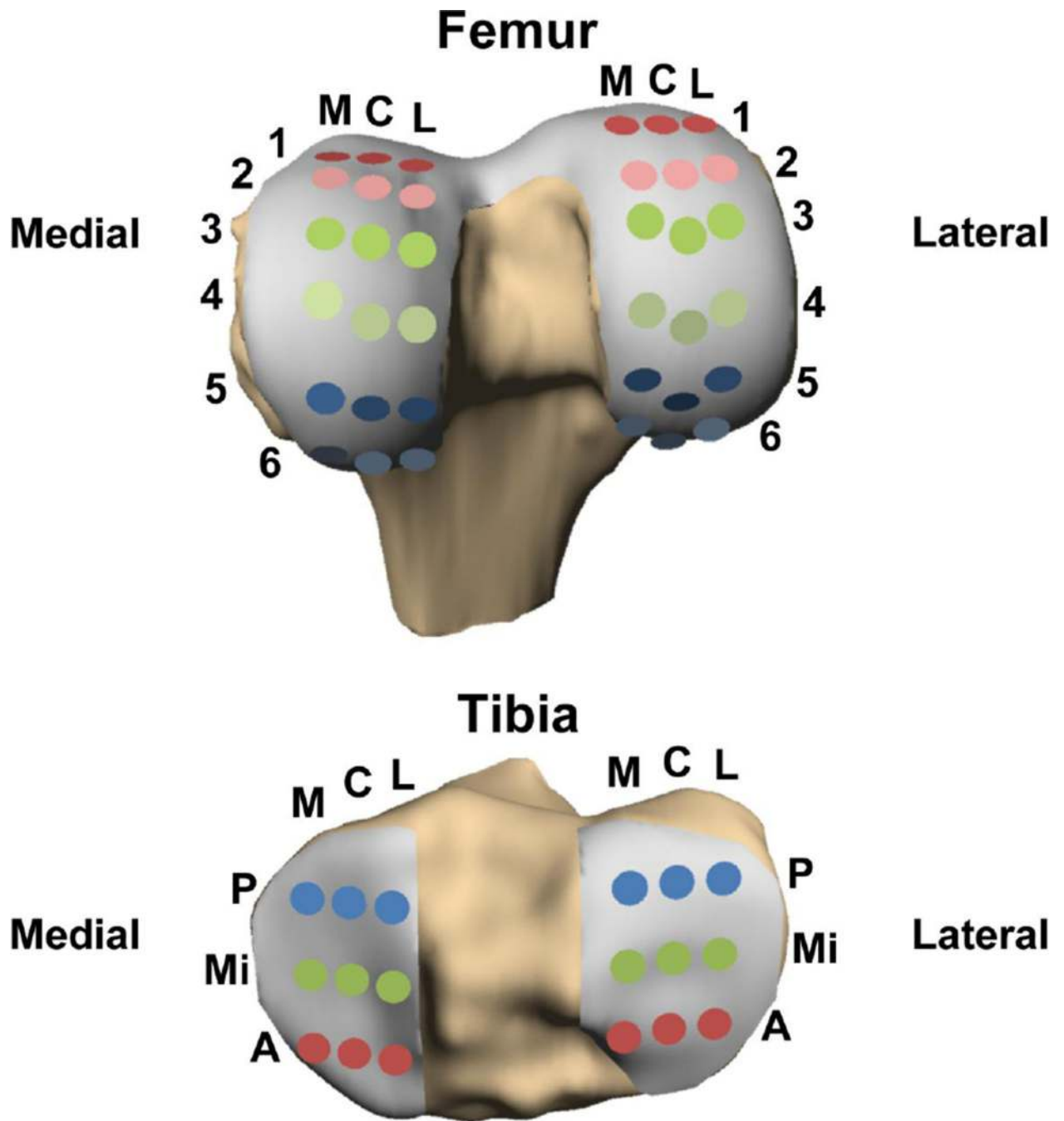


Figure 3. Femoral and tibial strain grids: 18 points on each femoral condyle and 9 points on each tibial plateau. L=lateral, C=center, M=medial, A=anterior, Mi=middle, P=posterior.

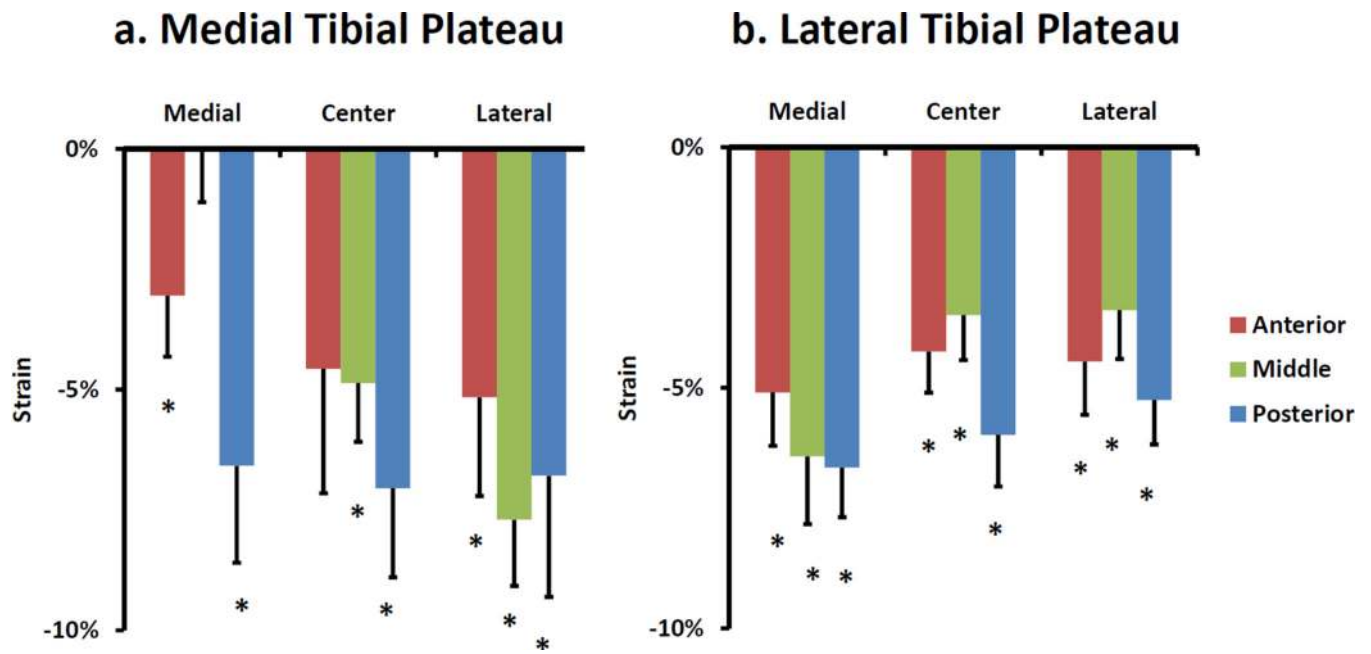


Figure 4.

Local strain results for a) medial and b) lateral tibial plateaus. Please see Figure 3 for point location legend. Error bars represent standard error of the mean. * $p < 0.05$ different from 0.

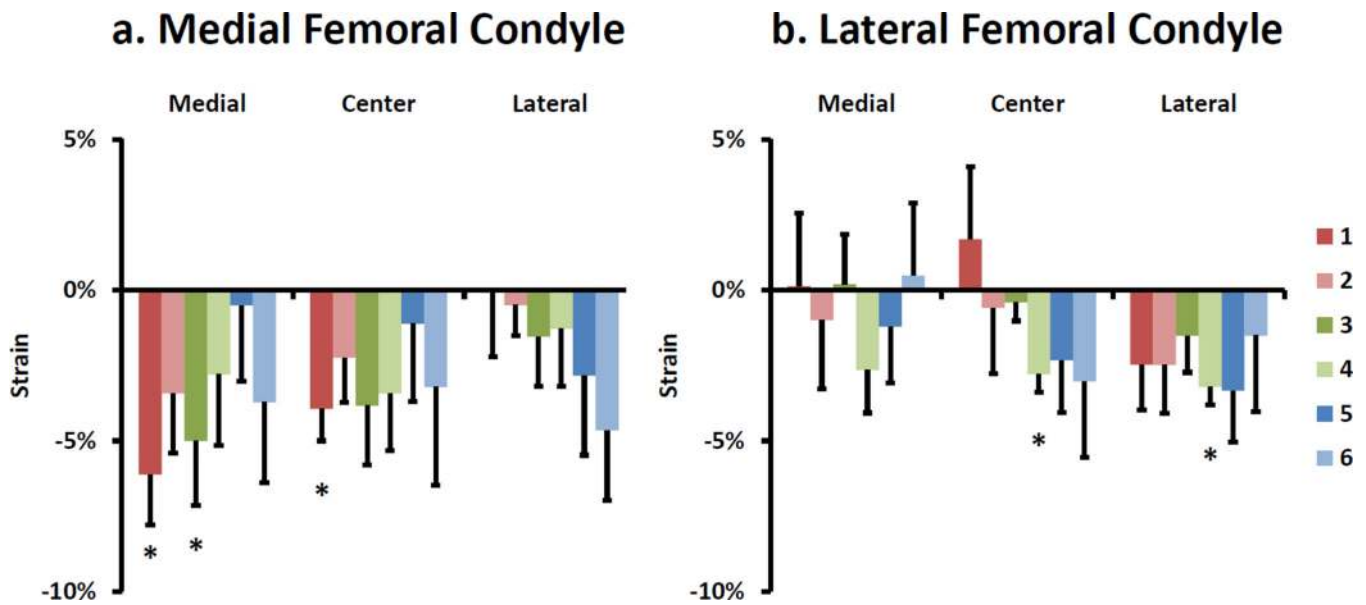


Figure 5. Local strain results for a) medial and b) lateral femoral condyles. Please see Figure 3 for point location legend. Error bars represent standard error of the mean. * $p < 0.05$ different from 0.