Maintenance of the paraspinal muscles may protect against radiographic knee osteoarthritis

Background: Knee osteoarthritis (OA) is an increasing health problem worldwide. So far, only obesity and quadriceps weakness are identified as modifiable risk factors for knee OA. Core muscle strengthening is becoming increasingly popular among older adults because of its ability to enhance the activities of daily living during old age. This study investigated the associations of the size and quality of the abdominal trunk muscles with radiographic knee osteoarthritis (RKOA).

Methods: From 2012 to 2016, data were collected from 146 males and 135 females (age 63.9±13.4 years, BMI 23.2±3.8 kg/m²) at annual musculoskeletal examinations, including knee radiographs and body composition analyses, by dual-energy X-ray absorptiometry. Cross-sectional areas of abdominal trunk muscles were measured using a single-slice computed tomography scan image obtained at the level of the umbilicus.

Results: The prevalence of RKOA was 21.2% in males and 28.1% in females. Compared to subjects without RKOA, subjects with RKOA were ~6 years older and had smaller paraspinal muscle (38.4±10.1 cm², ±8.7 vs 33.1±6 years older and had smaller paraspinal muscle cross sectional area, paraspinal muscle attenuation, lean body mass, fat mass (21.5±3.5 vs 24.5±4.4 kg/m², 16.7±7.0 vs 20.5±7.7 kg, respectively, both p<0.01). After adjusting for age and sex, smaller cross-sectional area/lower attenuation value of the paraspinal muscles was associated with RKOA (both p<0.05), while greater appendicular or total lean mass as well as greater FM was associated with RKOA. The size and quality of the paraspinal muscles were not associated with knee pain or habitual exercise.

Conclusion: Small, poor-quality paraspinal muscles may be linked to a higher risk of RKOA, but appendicular or total lean mass was not a good predictor of RKOA.

Keywords: radiographic knee osteoarthritis, paraspinal muscle cross sectional area, paraspinal muscle attenuation, lean body mass, fat mass

Introduction

The prevalence of knee osteoarthritis (OA), which is characterized by narrowing of the joint space and osteophyte formation, is an increasing problem worldwide. In Japan, radiographic knee OA is estimated to affect 25,300,000 people over 40 years of age, and a recent national survey ranked OA fourth among disabling diseases that cause affected people to require support with their activities of daily living.

The etiology of knee OA is complex, and several risk factors, including obesity, previous injury, female sex, advancing age, and quadriceps weakness, have been identified. Of these risk factors, only obesity and muscular strength are modifiable.
Therefore, weight loss and quadriceps resistance training are frequently recommended to alleviate knee pain and slow the progression of knee OA. However, for individuals with knee OA, weight loss through aerobic exercise (such as walking) and quadriceps training are both made difficult by pain during movement.

Core muscle strengthening exercises are becoming increasingly popular among older adults because these exercises enhance their ability to successfully perform the activities of daily living during old age. In fact, it has been demonstrated that trunk muscle activity occurs before the activity of the prime mover of the lower limb, and diminished core stability due to impaired abdominal trunk muscle is associated with lower extremity injury, including the knee. Improved core strength may also correct dynamic malalignment of the lower extremity by reducing the knee abduction moment. Thus, the objective of this study was to examine the association of abdominal trunk muscle size and quality with radiographic knee OA, independent of BMI or whole body/appendicular lean mass.

Methods
Study subjects
A total of 281 older adults who had undergone knee radiographs and dual-energy X-ray absorptiometry (DEXA) in the Center for Preventive Medicine of Keio University Hospital between September 2012 and March 2016 as part of screening for early signs of musculoskeletal degeneration were enrolled in this study. We obtained information on age, sex, body mass index (BMI; kg/m²), knee pain, and exercise habits from questionnaires. Habitual exercise was defined as exercising for at least 30 minutes per session, at least twice per week, for at least 1 year, according to the National Health and Nutrition Survey conducted by the Ministry of Health, Labour and Welfare in Japan.

Because this was a noninvasive retrospective study, informed consent was not obtained from the study subjects. All records were anonymized and de-identified prior to analysis. This study was approved by the local ethics committee of the Keio University School of Medicine (No. 20140323).

Imaging and measurements
Knee radiographs
Anteroposterior and lateral radiographs of bilateral knees were taken in a supine position at Keio University Hospital. The radiographs were read by a single, experienced knee surgeon, and Kellgren/Lawrence (K/L) grades were assigned as described in the K/L radiographic atlas. In the K/L grading system, radiographs are scored from 0 to 4, with higher grades being associated with more severe OA. The definition of K/L grade itself is, however, rather qualitative, especially for joint space narrowing. Unfortunately, our radiographs were taken in the non-weight-bearing position, but the K/L grade ≥ 2 under this condition is at least ≥ 2 if taken in a weight-bearing position. It is also obvious for experienced knee surgeons to determine the difference between K/L grade 1 (doubtful narrowing of joint space and possible osteophytic lipping) and grade 2 (definite osteophytes and possible narrowing of joint space). Therefore, in the current study, subjects with K/L grades ≥ 2 for at least one knee were considered to have radiographic knee osteoarthritis (RKOA).

DEXA scan
Whole-body DEXA scans (Lunar Prodigy® Advance; GE Healthcare Japan Corporation, Tokyo, Japan) were performed using enCORE software, version 9.2 (GE Healthcare). Standard scan mode was used for whole-body scans to measure total and regional (upper body, trunk, and legs) fat mass (FM) and lean body mass (LBM). Appendicular skeletal muscle mass (ASM) was defined as the LBM of both the upper and lower extremities. Skeletal muscle index (SMI) was defined as the ASM divided by height squared.

Computed tomography (CT) scan
Abdominal trunk muscle and visceral adipose tissue (VAT) were measured using a single-slice CT image obtained at the level of the umbilicus with 5 mm slice thickness.

The images were exported to a workstation in a standard DICOM (Digital Imaging and Communications in Medicine) format and were analyzed using sliceOmatic software (Tomovision, Magog, QC, Canada). The cross-sectional areas of psoas major, the paraspinous muscles (multifidus, longissimus, and iliocostalis lumbarum), and the anterolateral abdominal muscles (Ant Abd; the internal and external oblique muscles and transversus abdominis), as well as VAT, were measured by defining a range of CT attenuation values and a region of interest for each tissue. The attenuation values were –30 to –190 HU for VAT and 0–100 HU for muscle, as previously described. Low muscle attenuation (a low CT attenuation value for muscle) indicates increased muscle fat content and has been reported to be a marker of poor muscle quality and muscle weakness independent of age-associated loss in muscle mass; therefore, muscle attenuation was assessed simultaneously with cross-sectional area.
Statistical analysis
The significance of between-group differences (with and without RKOA) was assessed using analysis of variance (ANOVA). Homogeneity of variances was tested by Levene's test, and if the assumption was violated, Welch's test was used. Multivariable linear regression models were used to adjust for age, sex, and BMI or LBM/FM. Variance inflation factor (VIF) was used to check for multiple collinearity. Two-way ANOVA was also used to assess the interaction between RKOA and sex. Separate stepwise-regression analyses for RKOA were performed for males and females. IBM SPSS for Windows, Version 23 (IBM Corporation, Armonk, NY, USA) was used for all statistical analyses. The significance level was set at \( p<0.05 \), and all values are presented as mean±standard deviation (SD).

Results
Characteristics of the subjects
As shown in Table 1, subjects with radiographic knee osteoarthritis (RKOA) were older than those without RKOA; this was true for both males and females. In both sexes, BMI data showed a significant interaction between RKOA and sex \((p<0.05)\). In female subjects only, RKOA was associated with a higher BMI and greater FM. Although there was no other interaction between RKOA and sex, except for BMI, we have shown the data by sex, due to its large difference in body composition.

Analysis of the CT images showed that paraspinal muscle cross-sectional area was significantly lower among both males and females with RKOA, while paraspinal muscle CT attenuation was lower only in females with RKOA. In addition, female subjects with RKOA had more visceral adiposity and greater total adiposity than those without RKOA.

Multivariable analyses
Table 2 shows the results of multivariable linear regression models of body composition in subjects with and without RKOA, in which the subjects without RKOA serve as the referent group. After adjusting for sex and age (model 2), those with RKOA were heavier and had more LBM and FM than those without RKOA. In contrast, subjects with RKOA had smaller paraspinal muscle area and lower paraspinal muscle attenuation on CT, with concomitantly larger psoas muscle area, than those without RKOA. After adding BMI to the model (model 3), only the between-group difference in paraspinal muscle area remained significant. Surprisingly, after adjusting for age, sex, and BMI, lower FM was positively associated with RKOA. After adjusting for age, sex, height, and FM (model 4), greater total and appendicular LBM were more prominently associated with RKOA. After adjusting for age, sex, height, and LBM (model 5), the associations of smaller Ant abd and paraspinal muscle area with RKOA became more robust, while the association between greater FM and RKOA became insignificant.

Table 1 Body composition characteristics of the study subjects stratified using the K/L grading system for RKOA

| Characteristics | Male | Female |
|-----------------|------|--------|
| RKO A \(0–1\) (n=115) | \(>2\) (n=31) | \(0–1\) (n=97) | \(>2\) (n=38) |
| Age \(60.8\pm15.4\) | \(67.3\pm12.3\)** | \(63.8\pm11.6\) | \(71.0\pm8.6\)** |
| Height (cm) \(168.8\pm6.6\) | \(167.0\pm7.2\) | \(156.0\pm6.7\) | \(151.8\pm6.3\)** |
| BW (kg) \(67.7\pm11.2\) | \(69.1\pm11.5\) | \(52.3\pm9.8\) | \(56.5\pm10.9\)** |
| BMI (kg/m\(^2\)) \(23.7\pm3.3\) | \(24.7\pm3.4\) | \(21.5\pm3.5\) | \(24.5\pm4.4\)** |
| DEXA | | | |
| % body fat | \(24.4\pm7.2\) | \(25.3\pm6.6\) | \(30.9\pm7.8\) | \(35.2\pm7.2\)** |
| FM (kg) | \(17.1\pm7.5\) | \(17.9\pm6.9\) | \(16.7\pm7.0\) | \(20.5\pm7.7\)** |
| LBM (kg) | \(50.6\pm5.7\) | \(51.2\pm6.7\) | \(35.6\pm4.5\) | \(36.0\pm4.4\) |
| SMI | \(7.3\pm0.8\) | \(7.5\pm0.9\) | \(5.7\pm0.6\) | \(6.1\pm0.7\)** |
| CT at umbilical level | | | |
| Psoas (cm\(^2\)) | \(23.0\pm5.8\) | \(23.7\pm4.3\) | \(13.2\pm3.6\) | \(12.9\pm3.5\) |
| Psoas HU | \(49.2\pm4.2\) | \(48.8\pm4.2\) | \(45.7\pm4.2\) | \(44.3\pm4.1\) |
| Paraspinal (cm\(^2\)) | \(38.4\pm8.7\) | \(33.1\pm10.1**\) | \(24.1\pm7.1\) | \(20.7\pm7.5**\) |
| Paraspinal HU | \(47.3\pm6.5\) | \(44.5\pm7.1**\) | \(39.4\pm7.0\) | \(34.6\pm5.6**\) |
| Ant abd (cm\(^2\)) | \(49.8\pm10.5\) | \(47.2\pm13.9\) | \(27.8\pm8.1\) | \(25.8\pm8.6\) |
| Ant abd HU | \(44.0\pm6.8\) | \(42.3\pm6.2\) | \(38.9\pm7.6\) | \(34.7\pm4.9**\) |
| VAT (cm\(^2\)) | \(100.8\pm53.1\) | \(114.1\pm49.9\) | \(64.4\pm39.1\) | \(82.3\pm43.6**\) |

Notes: *\(p<0.05\) and **\(p<0.01\) vs subjects without RKOA, by ANOVA. Welch’s test was used for Ant abd HU in females, since the assumption of homogeneity of variances was violated. Data presented as mean±SD unless otherwise indicated.

Abbreviations: ANOVA, analysis of variance; Ant abd, anterolateral abdominal muscle; BMI, body mass index; BW, body weight; CT, computed tomography; DEXA, dual-energy X-ray absorptiometry; FM, fat mass; HU, Hounsfield unit; K/L, Kelgren/Lawrence; LBM, lean body mass; Paraspinal, paraspinal muscle; Psoas, psoas major muscle; RKO A, radiographic knee osteoarthritis; SMI, skeletal muscle index; VAT, visceral adipose tissue.
When data from males and females were analyzed separately, greater BMI and greater FM were associated with RKOA only in females, while greater LBM was associated with RKOA in both males and females after adjusting for age, height, and FM. In multivariate analyses, smaller paraspinal muscle area remained significantly associated with RKOA only in males, whereas in females, less paraspinal muscle attenuation was observed after adjusting for age. In the stepwise-regression analyses, when data from males and females were analyzed separately, paraspinal muscle attenuation, FM, and height were included in the final model for females, whereas paraspinal muscle area, LBM, and height were included in the final regression model for males.

### Association of body composition with knee pain and habitual exercise

Subjective knee pain was reported by 39.8% of the males and 62.1% of the females. For both sexes, there were no significant differences in body composition between subjects with and without knee pain, although the prevalence of RKOA was two times greater in females with knee pain than in females without knee pain.

Habitual exercise was associated with RKOA in males. Females who habitually exercised had a lower % body fat, larger psoas muscle and anterolateral muscle areas, greater muscle attenuation values, and greater ASM (appendicular LBM) than females who did not exercise regularly; in males,

#### Table 2 Multivariable analysis of body composition and trunk muscle characteristics in subjects with and without RKOA

| Model | 1 | 2 | 3 | 4 | 5 |
|-------|---|---|---|---|---|
| **All (n=281)** | | | | | |
| BMI (kg/m²) | 0.22*** | 0.27*** | – | 0.58*** | 0.16 |
| FM (kg) | 0.14* | 0.16** | –0.35** | – | 0.09 |
| LBM (kg) | –0.04 | 0.27* | 0.01 | 0.47** | – |
| ASM (kg) | –0.05 | 0.29* | 0.08 | 0.48*** | – |
| Psoas (cm²) | –0.05 | 0.27** | 0.11 | 0.24* | 0.08 |
| Psoas HU | –0.12* | 0.04 | 0.07 | 0.08 | 0.02 |
| Paraspinal (cm²) | –0.22*** | –0.19* | –0.27** | –0.18 | –0.35** |
| Paraspinal HU | –0.25*** | –0.21* | –0.13 | –0.15 | –0.17* |
| Ant abd (cm²) | –0.12* | 0 | –0.20* | –0.03 | –0.38*** |
| Ant abd HU | –0.20*** | –0.1 | 0 | –0.02 | –0.09 |
| **Male (n=146)** | | | | | |
| BMI (kg/m²) | 0.12 | 0.16 | – | 0.43* | 0.04 |
| FM (kg) | 0.05 | 0.07 | –0.25 | – | 0.02 |
| LBM (kg) | 0.04 | 0.15 | 0.07 | 0.25* | – |
| ASM (kg) | 0.02 | 0.15 | 0.08 | 0.26* | – |
| Psoas (cm²) | 0.05 | 0.22* | 0.17 | 0.22* | 0.14 |
| Psoas HU | –0.04 | 0.09 | 0.12 | 0.12 | 0.07 |
| Paraspinal (cm²) | –0.24** | –0.20* | –0.23* | –0.20 | –0.35** |
| Paraspinal HU | –0.17 | –0.11 | –0.05 | –0.08 | –0.09 |
| Ant abd (cm²) | –0.09 | –0.03 | –0.11 | –0.03 | –0.26* |
| Ant abd HU | –0.11 | –0.02 | 0.05 | 0.03 | –0.02 |
| **Female (n=135)** | | | | | |
| BMI (kg/m²) | 0.35*** | 0.33*** | – | – | 0.22 |
| FM (kg) | 0.23** | 0.25** | –0.48* | – | 0.14 |
| LBM (kg) | 0.04 | 0.16 | –0.04 | 0.29* | – |
| ASM (kg) | 0.05 | 0.18* | 0.02 | 0.30** | – |
| Psoas (cm²) | –0.04 | 0.12 | –0.05 | 0.07 | –0.09 |
| Psoas HU | –0.15 | 0 | 0.02 | 0.04 | –0.03 |
| Paraspinal (cm²) | –0.21* | –0.05 | –0.17 | –0.11 | –0.15 |
| Paraspinal HU | –0.31*** | –0.23* | –0.16 | –0.18 | –0.20 |
| Ant abd (cm²) | –0.11 | 0.05 | –0.17 | –0.02 | –0.24 |
| Ant abd HU | –0.27*** | –0.14 | –0.04 | –0.04 | –0.13 |

**Notes:** Values in columns represent standardized beta coefficients from linear regression model. 1: unadjusted; 2: adjusted for age and sex; 3: adjusted for age, sex, and BMI; 4: adjusted for age, sex, height, and FM; and 5: adjusted for age, sex, height, and LBM. *p<0.05, **p<0.01, and ***p<0.001, using multivariable linear regression models; subjects without radiographic knee OA are the referent group.

**Abbreviations:** Ant abd, anterolateral abdominal muscle; ASM, appendicular skeletal muscle mass; BMI, body mass index; FM, fat mass; HU, Hounsfield unit; LBM, lean body mass; Paraspinal, paraspinal muscle; Psoas, psoas major muscle; RKOA, radiographic knee osteoarthritis.
these differences were less clear. In contrast, paraspinal muscle area and attenuation did not differ between male or female subjects who did or did not habitually exercise (Table 3).

**Discussion**

The results of this study showed an association between lower spinal muscle area/attenuation and RKOA, in contrast to the lack of associations between lower appendicular or total lean mass and RKOA. Rather, greater LBM was associated with RKOA after adjusting for age, height, and total adiposity, while total adiposity was associated with RKOA only in females and was no longer associated with RKOA after adjusting for age and BMI or lean mass.

High BMI is a well-known risk factor for both the onset and progression of knee OA.

In a European cohort study (the Rotterdam study), a high BMI at baseline was associated with the incidence and progression of knee OA. Recently, a systematic review and meta-analysis showed an association between knee extensor muscle weakness and increased risk of knee OA; this suggests that fat rather than lean mass underlies the association between greater BMI and knee OA. However, several reports have shown that both total/lean leg mass and FM were associated with RKOA. Moreover, it has been reported that greater leg lean mass was a more powerful predictor of RKOA than greater BMI.

Segal et al reported that more thigh muscle mass was not associated with a lower risk of RKOA but that quadriceps strength was negatively associated with symptomatic knee OA and narrowing of the knee joint space. Although there is a strong positive association between muscle mass and strength, greater muscular mass does not necessarily mean greater muscular strength; the so-called low-quality muscle, common in elderly or sedentary individuals, may not be strong enough to alleviate the burden on the knee joints and may (similar to FM) even impose a higher mechanical burden on the knee.

The novel finding in our study is that, unlike total or lean leg mass, greater abdominal muscle area, especially paraspinal muscle area, is associated with less risk of RKOA. We do not have any mechanistic data; however, the paraspinal muscles are the only extensors of the trunk, and their stabilizing function could be of particular importance for balance and mobility in older adults, especially because people in this age group often suffer from a flexed (i.e., kyphotic) posture. Age-related kyphosis due to small and weak paraspinal muscles may increase the mechanical burden on the knee joint. Since this is a cross-sectional study, it is important to consider that a kyphotic posture could be the result of knee pain due to...
RKOA rather than the cause of RKOA. However, we found no associations between knee pain and abdominal muscle parameters, suggesting that the smaller paraspinal muscles in subjects with RKOA were not merely a consequence of knee pain.

A recent 15-year longitudinal observational study showed that the area and fatty infiltration of the paraspinal muscles were dependent on age and were not associated with physical activity level or degree or history of lumber back pain among typical Finnish males. In fact, familial aggregation (the combined effects of genes and a shared early environment) has been reported to be the strongest predictor of paraspinal muscle parameters. The lumbar paraspinal muscles help to stabilize the spine and maintain proper posture; thus, these muscles are used in any movement against gravity in daily life. Other abdominal muscles, including the anterolateral abdominal and psoas muscles, are more often used during the events such as heavy exercise or intentional brisk walking. In the current study, the paraspinal muscles were the trunk muscles that were least associated with habitual exercise. However, although no relationships were observed between habitual exercise and paraspinal muscle morphology, most of our study subjects performed aerobic exercise, which may place less stress on the lumbar extensor muscles than resistance exercise. An earlier case–control study showed that 70-year-old males with a history of aerobic exercise had similar arm and leg muscle areas as age-matched sedentary controls, whereas males who had resistance training had muscle area comparable with a group of sedentary 28-year-old subjects; this suggests that resistance training is key to maintaining muscle mass. Therefore, resistance exercises designed to stimulate the lumbar extensor muscles may be a promising treatment option for knee OA.

Muscle CT attenuation is a marker of fatty infiltration of the muscle, and lower muscle attenuation is associated with less muscular strength, independent of muscle area, in older people. Using magnetic resonance imaging (MRI), Kumar et al also reported more quadriceps intramuscular fat and less quadriceps strength in people with knee OA, without concomitant changes in quadriceps area. In the current study, lower paraspinal muscle attenuation values were also associated with RKOA, especially in females.

The effect of FM on knee OA was more apparent in females than in males, and this is consistent with previous observations. A recent study of Asian adults aged between 20 and 90 years found that advancing age was associated with knee OA in both males and females, while greater FM was associated with knee OA only in females. The reason for the observed sex difference is unclear; however, females have more fat and less muscle and are more prone to gaining fat in both adipose tissue and muscle, which worsens muscle quality. In a large-scale population-based cohort study of older Japanese adults (ROAD study), sitting on a chair was found to be a significant protective factor against RKOA, while standing, walking, climbing, and heavy lifting were associated with RKOA. In the current study, habitual exercise negatively affected RKOA only in males. It can be hypothesized that habitual exercise (which was almost exclusively walking in our study population) was not associated with RKOA in females because it prevents fat gain and fatty infiltration of the muscles.

The results of this study should be interpreted within the context of its limitations. Since this was a cross-sectional study, we cannot draw any conclusions about the causal or directional nature of the associations of muscle mass and quality with RKOA; although considering the lack of association between abdominal muscles and knee pain, it is unlikely that the observed association between the paraspinal muscles and RKOA was merely a consequence of knee pain. Moreover, although thigh/quadriceps muscle mass/strength is a well-known risk/protective factor for RKOA, we did not assess this directly. However, we did use DEXA to assess body composition and ASM, which is almost reflective of leg lean mass. We also did not assess abdominal muscle strength; however, muscle CT attenuation is a good index of muscular quality and strength. Our subjects were limited to medical checkup examinees and were more likely to be motivated for healthier lifestyle, and therefore, the selection bias also needs to be considered to generalize the findings. Finally, our definition of habitual exercise was not following the actual physical activity recommendations on a health perspective worldwide.

Despite these limitations, our current study adds to the current body of knowledge on the role of the size/quality of the paraspinal muscles in knee OA. The lack of a protective effect of total/appendicular lean mass also strongly suggests the importance of muscular size/quality rather than muscle quantity. So far, clinical recommendations to alleviate knee pain and slow the progression of knee OA consist of weight loss and resistance training of the quadriceps. Based on our observations, resistance exercise of the trunk extensor muscles or posture training may be other promising options for the treatment of knee OA. These exercises also have the advantage of placing no direct mechanical stress on the knee joints.
Conclusion
We observed that smaller paraspinal muscles were associated with RKOA and that total/appendicular lean mass did not protect against knee OA. This suggests the potential promising option of resistance training for the paraspinal muscles to alleviate knee OA.

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Disclosure
The authors report no conflicts of interest in this work.

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