Ethnic Differences in the Relationship Between Obesity and Joint Pain and Function in a Joint Arthroplasty Population

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ABSTRACT. Objective. We investigated the influence of obesity on joint pain and function in Asians as compared to Caucasians with degenerative hip and knee arthritis.

Methods. We surveyed 1983 patients (1876 Caucasians and 107 Asians) undergoing primary hip or knee replacement surgery. Relevant covariates including demographic data, body mass index (BMI), sex, comorbidities, education, and ethnicity were recorded. Pain and joint functional status were assessed at baseline and at 1-year followup with the Western Ontario and McMaster University Osteoarthritis Index (WOMAC) pain and function scores.

Results. Asian patients presented for surgery at a significantly younger age and lower mean BMI, and reported greater pain and dysfunction than Caucasian patients. Multivariate linear regression modeling showed that for every level of BMI, Asian patients reported greater levels of joint pain and dysfunction. At a BMI of 30 kg/m², this translated to a 16.6% higher WOMAC score (p < 0.001).

Conclusion. Among patients with endstage osteoarthritis, at every level of BMI, joint pain and dysfunction are greater in Asians than in Caucasians. This difference is likely mediated through both mechanical and inflammatory effects. (First Release Aug 1 2008; J Rheumatol 2008;35:1874–7)

| Key Indexing Terms: | | | |
|---------------------|-----------|---------|----------|
| ARTHROPLASTY | ETHNICITY | OBESITY | OUTCOMES |

Obesity has been clearly defined as an independent risk factor for the development and progression of hip and knee osteoarthritis $(OA)^{1-4}$. The World Health Organization (WHO) has used body mass index (BMI) as a method of grading the level of adiposity, with a BMI $\geq 25 \text{ kg/m}^2$ defined as overweight and $\geq 30 \text{ kg/m}^2$ as obese. These cutpoints were derived from a primarily Caucasian (individuals whose ancestry is White/European) population as predictors of chronic disease and mortality⁵. It is presently unclear if these BMI cutpoints can be extrapolated to other ethnic groups in terms of degree of risk for chronic disease⁶.

It is well established that Asians (South Asians and East Asians) are at higher risk for type 2 diabetes, hypertension, and dyslipidemia at a lower BMI threshold compared to Caucasians⁷⁻⁹. Further, evidence suggests that for every level of BMI, Asians have a greater relative excess of adi-

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pose tissue and less lean muscle mass (LMM)^{10,11}. This excess adipose tissue may lead to a heightened inflammatory state through greater systemic levels of C-reactive protein (CRP) and interleukin levels^{12,13}.

Emerging literature demonstrates that the effect of obesity on OA is mediated by both mechanical and chemical effects, and that dyslipidemia can affect joint homeostasis¹¹⁻¹⁴. Fat cells release proteins called adipokines, of which leptin, adiponectin, and resistin are the most studied. Leptin has been shown to promote inflammation in arthritic joints, while adiponectin works to control the level of inflammation. Both proteins are found in synovial fluid and elevated levels of leptin have been found to be a key regulator of chondrocyte metabolism¹¹⁻¹⁶. South Asians, Chinese, Japanese, and Taiwanese have all been shown to have high circulating levels of leptin¹⁷⁻²¹. We hypothesized *a priori* that among a population awaiting knee and hip replacement surgery, for any given BMI, Asian patients would report greater pain and physical dysfunction than Caucasians.

MATERIALS AND METHODS

Study sample. Study patients were recruited from a single Canadian academic institution, the Toronto Western Hospital, prior to undergoing primary hip or knee replacement surgery. Our inclusion criteria for the study were ages 18–85, a diagnosis of primary or secondary OA, and ethnicity self-report as being either Asian or Europid (Caucasian).

All patients gave informed consent to participate in our study. All data were collected by an independent assessor not involved in the medical care

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of the patients. The study protocol was approved by the local ethics committee.

Collection of data. Baseline demographic data of age, sex, BMI, level of education, and ethnicity were recorded. Height and weight were collected through patient self-report and BMI was then calculated. Highest level of education was recorded as either higher education level (university or above) or low education level (high school or below). Ethnicity was recorded by patient self-report. "Asian" refers to individuals who classified themselves as South Asian (India, Pakistan, Bangladesh, and Sri Lanka) or East Asian (China, Japan, Taiwan, Korea). Baseline medical health was scored on the Charlson Comorbidity Illness Index²². A lower score on the Charlson Index represents a better health state. Functional status and pain level were assessed preoperatively and at 1-year followup with the Western Ontario and McMaster University Osteoarthritis Index (WOMAC) function and pain scores, respectively.²³ A higher WOMAC pain score indicates greater pain and a higher WOMAC function score indicates greater dysfunction.

Statistical analysis. Continuous data such as age, Charlson Index, and WOMAC pain and function scores were compared between groups using t-tests. Means and standard deviations are reported for all continuous variables. Binary data such as sex and level of education are reported with frequencies, and groups were compared with the corrected chi-squared test. Multivariate linear regression modeling was used to determine the influence of ethnicity and BMI on preoperative and 1-year WOMAC scores. The 6 dependent variables analyzed in separate models were preoperative and 1-year WOMAC pain, function, and overall scores. The independent variables entered into the models were age, sex, BMI, education, ethnicity, and Charlson Index.

All statistical analysis was done with SPSS version 13.0 (SPSS, Chicago, IL, USA). All reported p values are 2 tailed with an alpha of 0.05.

RESULTS

In our registry, we had complete data on 1983 out of 2825 (70.2%) total patients who composed our study cohort. Participants and nonparticipants were not significantly different in age, BMI, sex, or Charlson Index. There were 1876 (94.6%) Caucasian patients and 107(5.4%) Asian patients.

The mean age of the Caucasian patients was 70.5 years as compared to 66.9 years for the Asian patients (p = 0.003). There were significantly more men in the Caucasian group compared to the Asian group (p = 0.008) and the mean BMI was significantly greater in the Caucasian group at the time of surgery, 30.1 kg/m² versus 28.7 kg/m², respectively (p = 0.023).

Preoperatively, the WOMAC total score was 59.3 in the Asian patients and 50.8 in the Caucasian patients (p < 0.001). At 1-year followup postsurgery, the mean WOMAC total score was 25.1 in the Asian patients and 21.9 in the Caucasian patients (p = 0.23).

The overall data for our 2 cohorts is shown in Table 1.

In multivariate linear regression models, Asian ethnicity independently predicted a greater preoperative WOMAC pain, function, and total score after adjusting for age, sex, BMI, Charlson Index, and education (p = 0.001) At 1-year followup, Asian ethnicity was not predictive of WOMAC scores for any of 3 domains (p > 0.05). Table 2 shows the beta coefficients with 95% confidence intervals for the ethnicity variable from the adjusted analysis.

Table 1. Overall data comparing Caucasians and Asians.

| Characteristics | Caucasians, n=1876 | Asians, n=107 | р |
|----------------------------------|-----------------------|------------------|---------|
| Mean age (SD), yrs | 70.5 (12.0) | 66.9 (12.1) | 0.003 |
| Male, % | 42.7 | 29.6 | 0.008 |
| Mean BMI, kg/m ² (SD) | 30.1 (6.6) | 28.7 (4.6) | 0.023 |
| % Higher education | 48.9 | 51.4 | 0.625 |
| Mean Charlson index score (SD) | 0.71 (0.97) | 0.79 (0.93) | 0.384 |
| Baseline WOMAC | | | |
| Mean pain | 10.3 (4.0) | 11.8 (3.8) | < 0.001 |
| Mean function | 40.6 (14.5) | 47.5 (14.6) | < 0.001 |
| Mean overall | 50.8 (17.9) | 59.3 (17.7) | < 0.001 |
| 1 yr WOMAC | | | |
| Mean pain | 3.7 (3.7) | 4.2 (3.6) | 0.353 |
| Mean function | 18.3 (14.6) | 20.9 (14.4) | 0.223 |
| Mean overall | 21.9 (17.9) | 25.1 (17.4) | 0.233 |

BMI: body mass index; WOMAC: Western Ontario and McMaster University Osteoarthritis Index.

At the conventional threshold for obesity (BMI ≥ 30.0 kg/m²), the Asian cohort showed 16.6% greater overall level of pain and dysfunction as measured by the WOMAC score (Figure 1).

DISCUSSION

Our results demonstrate that Asians (South Asians and East Asians) report greater pain and dysfunction for every level of BMI as compared to Caucasian patients before joint replacement surgery. After the degenerative cartilage has been removed and replaced with a metallic implant, Asians and Caucasians show no significant differences in pain and level of function at 1 year. Other authors have shown that preoperative functional status predicts postoperative functional outcome in joint replacement surgery²⁴⁻²⁶; however, our data showed that the Asian population had a greater relative benefit from surgery than the Caucasian population.

We believe there are many potential contributing reasons for Asians to report greater pain and dysfunction despite a lower BMI compared to the Caucasian patients. For every level of BMI, Asians have a greater proportion of adipose tissue and less LMM^{10,11}, and LMM is known to be protective against joint pain and dysfunction². The greater level of adipose tissue in Asians also contributes to pain via a biochemical effect. Obesity has been shown to be associated with greater systemic levels of CRP, interleukins, and matrix metalloproteinases (MMP), factors that can all contribute to increased joint pain and inflammation¹²⁻¹⁴. Moreover, visceral adipocytes secrete leptin, a hormone shown to be involved in the degenerative process of chondrocytes^{15,16}, and large epidemiologic studies have shown that Asian populations have greater levels of circulating leptin than White populations¹⁷⁻²¹.

It is interesting to note that the Asian patients present for joint replacement surgery at a significantly younger age than

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Table 2. Linear regression modeling reporting beta coefficient for Asians compared to Caucasians adjusted for age, sex, BMI, Charlson Index, and education.

| | Preoperative Beta Coefficient (95% CI) Asian vs Caucasian | | p 1-year Postoperative Beta Coefficient (95% CI) Asian vs Caucasian | |
|---------------|---|---------|---|-------|
| WOMAC scores | | | | |
| Mean pain | 1.3 (0.5, 2.1) | 0.001 | 0.4 (-0.77, 1.5) | 0.456 |
| Mean function | 6.9 (4.0, 9.9) | < 0.001 | 3.2 (-1.0, 7.4) | 0.135 |
| Mean overall | 8.2 (4.6, 11.8) | < 0.001 | 3.6 (-1.5, 8.7) | 0.166 |

WOMAC: Western Ontario and McMaster University Osteoarthritis Index.

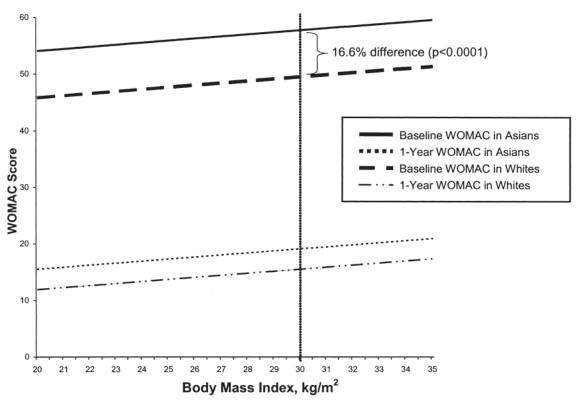


Figure 1. Total WOMAC scores at baseline and 1-year followup in Caucasians and Asians adjusted for age, sex, education, and Charlson Index.

the Caucasian patients. This may demonstrate that Asians develop degenerative arthritis at a younger age or that they have a more rapid course of progression and joint degeneration compared to Caucasian patients. These hypotheses warrant further study.

Our study has important clinical implications for physicians in that it suggests that although an Asian patient may present with a non-obese BMI (< 30.0 as currently defined by the WHO), he or she would still benefit from a weight loss and exercise program designed to reduce abdominal and general adipose levels and improve LMM. Further, we believe that the obesity cutpoints as defined by the WHO should be amended for the various ethnicities to more accurately reflect body habitus and risk of disease.

One potential limitation of our study is the 70% response rate among both the Caucasian and Asian patients. However, we demonstrated no difference in participants and nonparticipants in terms of age, sex, BMI, or baseline medical comorbidity, and we believe our conclusions remain valid and generalizable. Moreover, our population of 5% Asians is consistent with that of the downtown neighborhood of our hospital. Future large studies should be done separating the South and East Asian populations to confirm the same findings across groups.

We demonstrate that Asian patients present for surgery at a younger age and at a lower mean BMI, and complain of

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more joint pain and dysfunction prior to surgery compared to Caucasian patients. However, Asians and Caucasians both obtain significant benefit from joint replacement and have similar levels of function and pain 1-year postsurgery.

REFERENCES

- Bourne R, Mukhi S, Zhu N, Keresteci M, Marin M. Role of obesity on the risk for total hip or knee arthroplasty. Clin Orthop Rel Res 2007;465:185-8.
- Sowers MF, Yosef M, Jamadar D, Jacobson J, Karvonen-Gutierrez C, Jaffe M. BMI vs body composition and radiographically defined osteoarthritis of the knee in women: a 4-year follow-up study. Osteoarthritis Cartilage 2008;16:367-72.
- Abbate LM, Stevens J, Schwartz TA, Renner JB, Helmick CG, Jordan JM. Anthropometric measures, body composition, body fat distribution, and knee osteoarthritis in women. Obesity 2006;14:1274-81.
- Hochberg MC, Lethbridge-Cejku M, Scott WW Jr, Reichle R, Plato CC, Tobin JD. The association of body weight, body fatness and body fat distribution with osteoarthritis of the knee: data from the Baltimore Longitudinal Study of Aging. J Rheumatol 1995; 22:488-93.
- World Health Organization. Obesity: preventing and managing the global epidemic. Report on a WHO consultation on obesity; June 3–5, 1997; Geneva, Switzerland. WHO/NUT/NCD/98.1; Geneva; 1998.
- WHO/IASO/IOTF. The Asia-Pacific perspective: Redefining obesity and its treatment. Melbourne: Health Communications Australia; 2000.
- 7. Razak F, Anand SS, Shannon H, et al. Defining obesity cut points in a multiethnic population. Circulation 2007;115:2111-8.
- Lear SA, Toma M, Birmingham CL, Frohlich JJ. Modification of the relationship between simple anthropometric indices and risk factors by ethnic background. Metabolism 2003;52:1295–301.
- Forouhi NG, Jenkinson G, Thomas EL, et al. Relation of triglyceride stores in skeletal muscle cells to central obesity and insulin sensitivity in European and South Asian men. Diabetologia 1999;42:932–5.
- Chang CJ, Wu CH, Chang CS, et al. Low body mass index but high percent body fat in Taiwanese subjects: implications of obesity cutoffs. Int J Obes Relat Metab Disord 2003;27:253–9.
- Chowdhury B, Lantz H, Sjostrom L. Computed tomographydetermined body composition in relation to cardiovascular risk factors in Indian and matched Swedish males. Metabolism 1996;45:634–44.
- 12. Anand S, Razak F, Yi Q, et al. C-reactive protein as a screening test for cardiovascular risk in a multiethnic population. Arterioscler

Thromb Vasc Biol 2004;24:1509-15.

- Rexrode KM, Pradhan A, Manson JE, Buring JE, Ridker PM. Relationship of total and abdominal adiposity with CRP and IL-6 in women. Ann Epidemiol 2003;13:674-82.
- Toussirot E, Streit G, Wendling D. The contribution of adipose tissue and adipokines to inflammation in joint diseases. Curr Med Chem 2007;14:1095-100.
- Terlain B, Presle N, Pottie P, Mainard D, Netter P. Leptin: a link between obesity and osteoarthritis? Bull Acad Natl Med 2006;190:1421-35.
- 16. Dumond H, Presle N, Terlain B, et al. Evidence for a key role of leptin in osteoarthritis. Arthritis Rheum 2003;48:3118-29.
- Patel JV, Sosin M, Lin HS, et al. Raised leptin concentrations among South Asian patients with chronic heart failure. Int J Cardiol 2007;122:34-40.
- Wasim H, Al-Daghri NM, Chetty R, McTernan PG, Barnett AH, Kumar S. Relationship of serum adiponectin and resistin to glucose intolerance and fat topography in South-Asians. Cardiovasc Diabetol 2006;5:10.
- Wang TN, Huang MC, Chang WT, et al. G-2548A polymorphism of the leptin gene is correlated with extreme obesity in Taiwanese aborigines. Obesity 2006;14:183-7.
- Tong J, Fujimoto WY, Kahn SE, et al. Insulin, C-peptide and leptin concentrations predict increased visceral adiposity at 5- and 10 follow-ups in nondiabetic Japanese Americans. Diabetes 2005;54:985-90.
- Peng XD, Xie H, Zhao Q, Wu XP, Sun ZQ, Liao EY. Relationships between serum adiponectin, leptin, resistin, visfatin levels and bone mineral density, and bone biochemical markers in Chinese men. Clin Chim Acta 2008;387:31-5.
- Charlson M, Pompei P, Ales KL, Mackenzie CR. A new method of classifying prognostic comorbidity in longitudinal studies: development and validation. J Chron Dis 1987;40:373-83.
- 23. Bellamy N, Buchanan WW, Goldsmith CH, Campbell J, Stitt LW. Validation study of WOMAC: a health status instrument for measuring clinically important patient relevant outcomes to antirheumatic drug therapy in patients with osteoarthritis of the hip or knee. J Rheumatol 1988;5:1833-40.
- 24. Fortin PR, Clarke AE, Joseph L, et al. Outcomes of total hip and knee replacement: preoperative functional status predicts outcomes at six months after surgery. Arthritis Rheum 1999;42:1722-8.
- Fortin PR, Penrod JR, Clarke AE, et al. Timing of total joint replacement affects clinical outcomes among patients with osteoarthritis of the hip or knee. Arthritis Rheum 2002;46:3327-30.
- Lingard EA, Katz JN, Wright EA, Sledge CB. Predicting the outcome of total knee arthroplasty. J Bone Joint Surg Am 2004;86:2179-86.

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