Breaking into a Sweat... and Risk of Osteoarthritis!



Osteoarthritis (OA) remains a highly prevalent disorder, both in North America and globally ^{1,2,3}. According to the US National Arthritis Data Workgroup, approximately 27 million Americans were estimated, based on the 2005 population size, to have clinical evidence of OA¹. Given that the prevalence of OA rises with increasing age, the national burden of OA continues to expand⁴.

When moderate to severe OA affects the lower extremity, particularly at the knee and hip joints, the related pain and stiffness has a tremendous influence upon virtually all ambulatory activities involved in daily living⁵. Further, in the circumstance when symptoms resulting from joint space narrowing, cartilage degeneration, and osteophyte formation become inadequately controlled by conventional nonmedicinal and pharmacologic interventions, the patient and physician, working together, broach the subject of surgical intervention to correct the underlying joint disorder. They seek resolution to the pain and functional impairment, and an optimal restoration to a full, vital functional capacity⁷.

Over the last 50 years, several notable demographic variables and modifiable host factors have been examined as potential risk factors for incident and prevalent OA. For example, there is clear evidence from population-based studies that age is related to OA risk. Not only are older adults at greater risk than their younger counterparts to develop OA, but the risk of incident OA, specifically at the knee, continues to rise into the septuagenarian and octogenarian decades of the life span⁴. In addition, gender is clearly related to risk of disease, with women experiencing higher rates of OA than men, at the hand, knee, and hip joints, after age 40 and beyond¹. Moreover, in cross-sectional data generated among 5193 black and white study participants aged 35-74 years in the First National Health and Nutrition Examination Survey, African American women appeared to have twice the rate of prevalent OA as the Caucasian women⁸.

Other host factors and health behaviors, in contrast to the above unalterable demographic variables, lend themselves

to modification, whereby the host can choose to augment, maintain, or diminish his/her exposure level. A prominent example is that of body weight, with a large number of published reports indicating that higher levels of body weight, and persons in the obese category (exceeding a body mass index $> 30 \text{ kg/m}^2$) face a heightened risk to develop knee OA^{9,10,11}.

In this context, the potential association of physical activity with the development of OA is of much interest. Notably, leisure-time physical activity (or exercise) is in principle, if perhaps not always in practice, a modifiable human behavior. Yet, a priori hypotheses as to how this exposure (physical activity) and outcome (OA) might be related span the gamut, including consideration for a beneficial, neutral, or even harmful relationship. Given that physical activity is frequently recommended for greater integration into the lives of the general population due to its touted benefits regarding cardiovascular morbidity and all-cause mortality¹², it is important to know how physical activity does or does not relate to the health status of a human lower extremity joint. Some postulate that exercise may be beneficial by strengthening periarticular tissues, including the quadriceps muscle, and thereby reduce the risk of incident OA. Others claim, to the contrary, that exercise may be detrimental to a healthy, fully functional lower extremity joint, by exposing the knee or hip joint to deleterious forces that compromise the structural integrity of articular components (including cartilage, ligament, and bone), and in turn accelerate the rate of OA development.

A number of prior reports have addressed this issue by focusing on former top-level male athletes and ex-elite female athletes, among whom a heightened risk of OA at both the knee and hip joints has been demonstrated ^{13,14}. But, what are we to understand of the relationship of physical activity to incident OA in the general population?

In this issue of *The Journal*, we read, with interest, prospective, population-based data from Australia that directly address this issue. The authors examine whether

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physical activity predicts a greater risk to develop endstage OA of the knee or hip joint that ultimately necessitates pursuit of joint replacement surgery¹⁵. An appealing aspect of the study was its focus on the regular man and woman. The report examined OA risk in the Melbourne Collaborative Cohort Study, initially consisting of 41,528 persons assembled between 1990 and 1994, the vast majority of whom were age 40-69 years at cohort entry. Assessment of exposure status was first ascertained at baseline, then reevaluated during the followup period, from 1995 to 1998 (first followup), and during a second followup assessment, conducted from 2003 to 2007. Physical activity was assessed at baseline by querying the participants regarding the frequency of "vigorous" activity, defined as exercise that "makes you sweat or feel out of breath!" The quantity of "less vigorous" activity, which "did not make you sweat or feel out of breath," was similarly ascertained, as was the frequency of walking as a form of exercise. The investigators further utilized a composite index of the vigorous, less vigorous, and walking frequencies to quantify the aggregate level of physical activity per week for each study participant.

Importantly, the outcome of arthroplasty was ascertained in the 5 year period of January 1, 2001, to December 31, 2005, by linkage of the demographic data in the Melbourne Cohort with the National Joint Replacement Registry, which monitors the occurrence and outcomes associated with knee and hip replacement in Australia. The underlying indication for which the surgery was performed was similarly recorded. Notably, during a mean 4.8 years of followup, 1.4% of the study participants (n = 541) underwent knee replacement for OA whereas 1.2% (n = 468) received a hip replacement. Not surprisingly, the cohort members who underwent these surgical procedures were on average older than the demographic composition of the cohort as a whole. They also appear to have been of higher average body mass index (implying a greater frequency of obesity in those who underwent joint replacement) and more female in composition than the overall cohort. Most noteworthy is that in regression models that took age, gender, and body mass index into account, higher levels of total physical activity were related to risk of undergoing knee replacement surgery for OA. Moreover, the association of physical activity with incident knee replacement remained evident when activity was examined according to level of vigorous exercise alone. In contrast, the frequency of less vigorous activity, as well as the frequency of walking, was not related to risk of undergoing knee joint replacement for OA. At the same time, none of these various measures of physical activity, in aggregate, or by vigorous activity alone, were related to conduct of hip replacement surgery.

These findings impart a number of important messages. First, that in the general population of Melbourne, Australia, engaging in greater amounts of total physical activity appears to increase the risk of subsequent severe OA, neces-

sitating knee replacement surgery. However, when the component parts of aggregate physical activity are subdivided and analyzed separately, only the most vigorous category of activity, that which results in breaking into a sweat, was related to OA risk. At the same time, it would be important to further identify what contribution to the observed harm incurred by vigorous exercise might be the result of overt or occult intraarticular damage, including ligamentous and meniscal derangements, which are known risk factors for OA¹⁶. One may also be eager to know what bearing a greater interval in time than that observed in the present report, even a decade or more, between the assessment of the exposure and the development of the outcome, might have on the exercise-OA relationship.

Second, less vigorous physical activity, as well as walking as a form of exercise, were not risk factors for undergoing knee arthroplasty, and therefore, need not worry the health-conscious mindset about a potential deleterious effect of less vigorous exercise, including walking, upon OA susceptibility. As such, the cardiovascular and all-cause mortality benefits of walking are not seemingly offset by an increase in risk of future advanced OA later in life. These findings complement prior population-based reports from the United States^{17,18} and furnish reassuring good news. They also do not implicate harm from aggregate or vigorous exercise to OA risk at the hip joint.

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