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### Long-Term Mortality Trends Following Total Hip and Knee Replacement: A Population-Based Study

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#### Abstract

**BACKGROUND**—Mortality following total hip and knee replacement is lower than the general population, but it is unknown whether there are differences by surgery type, demographics and calendar year. Our objective was to evaluate trends and determinants of long-term mortality among patients with total hip and knee replacement.

**METHODS**—Using a historical cohort study design, we passively followed up population-based cohorts of total hip and total knee replacement patients with degenerative arthritis who underwent surgery between 1/1/1969 and 12/31/2008. Patients were followed up until death or 8/31/2014. Observed and expected survival was compared using standardized mortality ratios (SMR). Poisson regression models were used to examine relative mortality patterns by surgery type, age, sex, calendar year and time since surgery.

**RESULTS**—The overall age- and sex-adjusted mortality was significantly lower than the general population following both total hip (SMR: 0.82, 95% CI: 0.76, 0.88) and total knee (SMR = 0.80, 95% CI: 0.75, 0.86) replacement. Despite the low relative mortality within the first 8 years of surgery, we observed a worsening of relative mortality beyond 15 years after total knee replacement surgery. Both short and long term mortality improved over calendar time, and the improvement occurred about a decade earlier in total knee replacement than in total knee replacement.

**CONCLUSION**—Survival following total hip and total knee replacement is better than the general population for about 8 years following surgery. Secular trends are encouraging and suggest that survival following both procedures has been improving even further in recent years.

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#### Introduction

Total hip and knee replacement are safe and effective procedures for management of morbidity attributable to end-stage osteoarthritis. Despite the high prevalence and significant public health burden[1], life expectancy among individuals with osteoarthritis and the modifying role of total hip and knee replacement are poorly understood [2–4]. Although individuals with osteoarthritis are at a higher risk of death compared to individuals of the same age and sex in the general population [4], mortality following total hip and knee replacement is better than the general population[5-12], in part due to selection of healthier individuals to undergo surgery and, potentially in part due to benefits of increased mobility and activity levels following surgery[3]. So far, studies with follow-up beyond 5-years are limited. Some studies, mainly from the Nordic countries, show a bi-modal mortality pattern [5–7]. For example, within the Medicare population, investigators observed a convergence of mortality after 5 years following total hip replacement [5]. In a nationwide study from Sweden [7], mortality was lower than the general population for 12 years following knee replacement, but after that, it increased and became significantly worse than the general population. Although these studies suggest disappearance of the mortality difference over time, it is unclear when this occurs, and whether it is true in both total hip and knee replacement.

Regarding demographics, there is strong evidence for an age effect with high standardized mortality ratios among younger patients [7, 8, 11, 12]. Recent studies from England and Wales also suggest that early postoperative mortality fell substantially in recent years[13, 14], but it is unclear whether this improvement in mortality extends beyond the postoperative period which is not influenced by surgery itself but general population mortality trends. Indeed, considering the dramatic improvements in overall mortality in the general population[15], it would be reasonable to expect that the long-term mortality among total hip and knee replacement patients has followed the same pattern, but this has never been investigated.

With this background, our objective was to evaluate the trends and determinants of longterm mortality among patients with total hip and knee replacement in the United States. We sought to describe mortality differences by surgery type, demographics, calendar year and time since surgery.

#### Methods

We performed a historical cohort study in a population-based setting in the United States[16, 17]. Population-based epidemiologic studies in Olmsted County, Minnesota are feasible because of its relative geographic isolation from other urban centers and because nearly all medical care is delivered to local residents by a small number of providers. Data from all healthcare providers are contained in a single comprehensive medical records linkage system. Consequently, the unique population-based data resources in Olmsted County ensure virtually complete ascertainment and follow-up of total hip and knee replacement patients in a geographically defined community.

The study population comprised Olmsted County residents who underwent primary total hip (n=1645) and total knee (n=1980) replacement surgery for degenerative arthritis between 1/1/1969 and 12/31/2008[18]. Patients who underwent surgery for other indications and those who denied research authorization for use of their medical records for research were excluded. All patients were followed up longitudinally until death or 8/31/2014.

Statistical analyses were performed separately for the total hip and knee replacement cohorts. Death rates following total hip or knee replacement were evaluated using a personyears approach, in which the observed number of deaths in each cohort was compared to the expected number of deaths over the follow-up period. Expected survival was generated using the annual life tables. The results were reported as standardized mortality ratios (SMRs) with corresponding 95% confidence intervals (CIs). The confidence intervals were calculated assuming that the expected rates are fixed and the observed deaths follow a Poisson distribution. The confidence intervals are shown as dashed lines in the figures. Poisson regression models were used to model relative mortality by surgery type, age, calendar year of surgery and time since surgery. Although the relative mortality was slightly lower in males, the relative mortality patterns were similar in males and females; therefore, analyses were performed by combining data for male and female patients. Additionally, separate analyses were performed to evaluate relative mortality within the first year following surgery, and in the period following one year after surgery. For the former, all patients were included, but only deaths observed in the first year were analyzed, and followup greater than 365 days was censored at one year. For the latter, only patients with at least one year of follow-up were included. All analyses were performed using R version 3.0 (http://www.R-project.org/).

#### Results

Over a mean follow-up of 11.9 years (19573.7 person-years) of 1645 total hip replacement patients, 814 patients died. Similarly, over a mean follow-up of 10.8 years (21465.4 person-years) of 1980 total knee replacement patients, 842 patients died. Survival in both the total hip replacement and the knee replacement cohorts was significantly better than that expected in the general population with an SMR of 0.82 (95% CI 0.76, 0.88) following total hip replacement (Table 1).

Reduction in mortality was of similar magnitude in men and women for both procedures with an age-adjusted SMR of 0.82 (95% CI 0.75, 0.90) for women and 0.81 (95% CI 0.73, 0.90) for men in total hip replacement, and 0.82 (95% CI 0.75, 0.90) for women and 0.78 (95% CI 0.70, 0.86) for men in total knee replacement. Furthermore, the SMR decreased with increasing age at the time of surgery (Table 1).

Figures 1A and 1B illustrate relative death rates according to age at surgery adjusting for calendar year of surgery. Dashed lines indicate the 95% confidence intervals. In both total hip (Figure 1A) and total knee replacement (Figure 1B), the relative death rate was highest in younger ages and declined as age at surgery increased. The wide confidence intervals at younger ages are due to the low number of patients less than 50 years of age. The confidence

intervals for the relative death rate were below 1 after 60 years of age in total hip replacement and after 55 years of age in total knee replacement, indicating that their mortality was better than their peers of the same age in the general population. Moreover, consistent with the SMR estimates, the relative death rate curve for total knee replacement patients was lower than that for total hip replacement patients.

Figures 2A and 2B illustrate the relative death rates over time according to calendar year of surgery derived from age-adjusted Poisson regression models. We observed notable differences by surgery type. In total hip replacement (Figure 2A), the relative death rate was relatively flat around 1 (indicating mortality similar to the general population) until early 2000's after which it declined and became significantly better than the general population. In total knee replacement (Figure 2B), the relative death rate showed almost a linear decline over time, indicating steady improvements in survival of total knee replacement patients over time. In fact, as early as late1980's, the death rate following total knee replacement was significantly better than the general population (as indicated by the non-overlap of the confidence intervals with the unity line at 1) and continued to improve over time. These trends indicate that first, improvement in relative mortality occurred about a decade earlier in total knee replacement than in total hip replacement, and second, mortality rates following total knee replacement declined faster than the general population mortality.

We then examined relative death rates according to time since surgery. The relative death rate was significantly lower than the general population for about 8 years following both procedures (Figures 3A and 3B). The relative death rate was as low as 50% within the first year of total knee replacement surgery but there was a shift in relative mortality beyond 15 years (Figure 3B). The relative death rate following total knee replacement increased steadily during follow-up and became significantly worse than the general population beyond 18 years after surgery. Indeed, the relative death rates at 5 and 18 years following total knee replacement were 0.67 (95 % CI, 0.61, 0.74) and 1.19 (95 % CI, 1.01, 1.36), respectively. We did not observe such a shift in mortality in total hip replacement (Figure 3A).

We further examined to what extent these secular trends were driven by improvements in short-term (1-year) mortality in recent years, as shown in recent studies (Figure 4). Figures 4A and 4B illustrate relative death rates within the first year of surgery, and Figures 4C and 4D illustrate relative death rates beyond the first year following surgery. Overall, the downward trends in these figures indicate that improvements in mortality occurred both for short- and long-term mortality. In total hip replacement, improvement in 1<sup>st</sup> year mortality seems to have started in early 1990's but the pattern was not significant as indicated by the wide confidence intervals overlapping the unity line at 1 (Figure 4A). In total knee replacement, improvement in 1<sup>st</sup> year mortality started earlier and continued to improve thereafter (Figure 4B). Figures of relative death rates beyond the first year following surgery (Figure 4C and 4D) were similar to overall Figures 2A and 2B. Among total hip replacement patients who survived the first year (4C), improvement in mortality occurred only during the last decade. Among total knee replacement patients who survived the first year (Figure 4D), the relative death rate declined gradually and was significantly better than the general population by early 1990's.

#### Discussion

In this population-based study that spanned four decades, we evaluated mortality trends following total hip and knee replacement. Consistent with previous studies, mortality following both procedures is lower than the general population, and declines with increasing age at surgery. Despite the low relative mortality for about 8 years following surgery, mortality becomes worse than the general population beyond 15 years after total knee replacement surgery. Secular trends are encouraging and suggest that both short and long term mortality continues to improve in recent years. Finally, improvement in mortality seems to have occurred about a decade earlier in total knee replacement than in total hip replacement. Although the reasons for some trends, in particular the worsening of mortality beyond the first decade, remain to be determined with certainty, our encouraging findings reflect the success of the orthopedic community in improving the outcomes of their arthroplasty patients.

Our findings update and extend previous studies with findings broadly similar to those reported from population-based cohorts of similar age and sex distribution and length of follow-up [5–12]. For example, total hip replacement patients in Norway had 20% reduction in mortality (HR = 0.81) compared to the general population at eight years after surgery, despite having a higher risk of death within the first 2 months[8]. The reduced mortality following total hip and knee replacement is generally attributed to patient selection. The inverse relationship between age and mortality has been previously described [7, 8, 12], and similar to our findings, more pronounced in total hip replacement than in knee replacement.

In contrast to the overall and age-specific SMR estimates, there are only a handful of longterm mortality studies describing the bi-modal mortality pattern with the worsening of mortality beyond the first decade of surgery. Our findings are strikingly consistent with these studies [6, 7, 19]. Among almost 25,000 total hip replacement patients from Finland, Paavolainen et al [6] found that mortality was low during the first 5 years after surgery, but increased to unity (i.e., similar to general population) thereafter. In a cohort of more than 65,000 total knee replacement patients from Sweden[7], Robertsson et al. described reduced mortality during the first 12 postoperative years, but beyond 12 years, mortality increased and became significantly worse than the general population. Robertsson et al. attributed this finding to increased mortality in younger patients with longer follow-up, rather than an increased mortality in all patients. The reasons for the bimodal mortality pattern are unknown, and may potentially include underlying comorbidities and/or revision surgeries in these patients. Individuals who undergo surgery at a younger age may have more comorbidities and higher risk of death. With the ever growing number of individuals living with hip and knee replacement[20], there is a need to identify the drivers of the long-term morbidity and mortality in total hip and knee replacement.

Our analyses of secular trends indicate temporal improvements in mortality in both total hip and knee replacement patients. Although this was previously demonstrated for short-term mortality[13, 14, 21, 22], only one prior study from Norway reported declines in long-term mortality[8]. In a nationwide total hip replacement cohort, investigators observed a statistically significant difference in the SMR between 1987–90 and 1995–98 but a

significance test for trend was not significant. This is not surprising because our observations suggest that the mortality decline following total hip replacement may have occurred after 2000, and possibly about a decade later than in total knee replacement. With a mean follow-up of 11 years, we were able to show that the temporal improvements in in mortality is not just confined to the first year of surgery, and, at least in the case of total knee replacement, may have occurred even faster than the general population. Combined with the results of the short-term mortality studies[13, 14, 21], we can conclude that total hip and knee replacement patients have much lower mortality today than their counterparts about a decade ago.

Our study has a number of potential limitations. First and foremost, findings may not be generalizable to all joint replacement patients in the United States, in part due to racial and ethnic composition of the population during the calendar years under investigation, and in part due to variation in patient selection for joint replacement [23]. The generalizability of our mortality findings to demographically more diverse populations is unknown. Second, although our follow-up extended to August 2014, the cohort of joint replacement patients was limited to those who underwent surgery prior to 2008. Therefore, we cannot extrapolate our findings to patients who underwent surgery after 2008, who may have a different underlying mortality risk[21]. Third, due to the small cohort size, we had limited power for some of the analyses, such as mortality patterns among younger arthroplasty patients. Fourth, the reasons for the observed relative mortality trends, i.e. comorbidities, revisions are not addressed in the study. Our population-based design and inclusion of successive joint replacement cohorts since these procedures first started are unique strengths of the present study. In addition, our long and complete follow-up of all hip and knee replacement patients and ready access to general population mortality (expected) rates throughout the entire time period are advantages that made the current study feasible. In conclusion, there are notable differences in relative mortality trends following total hip and total knee replacement. Both short-term and long-term mortality are improving in recent years. These encouraging findings underscore the safety and effectiveness of total hip and knee replacement.

#### **Supplementary Material**

Refer to Web version on PubMed Central for supplementary material.

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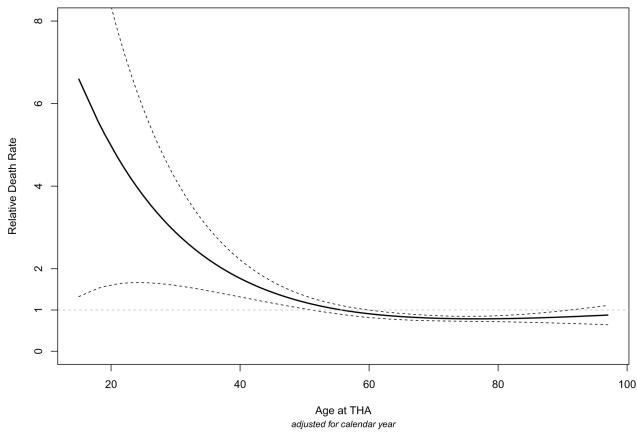
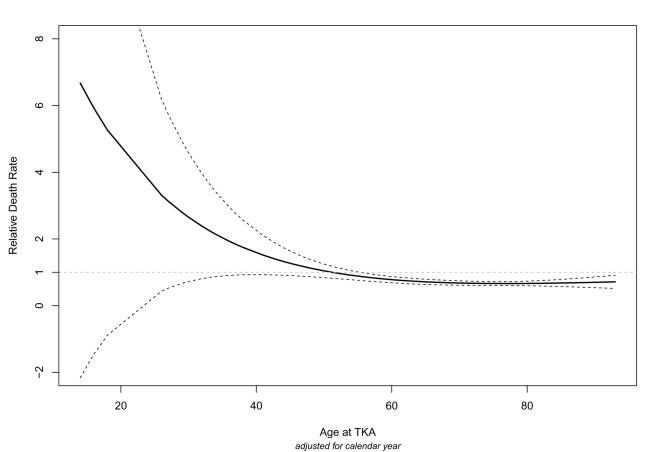




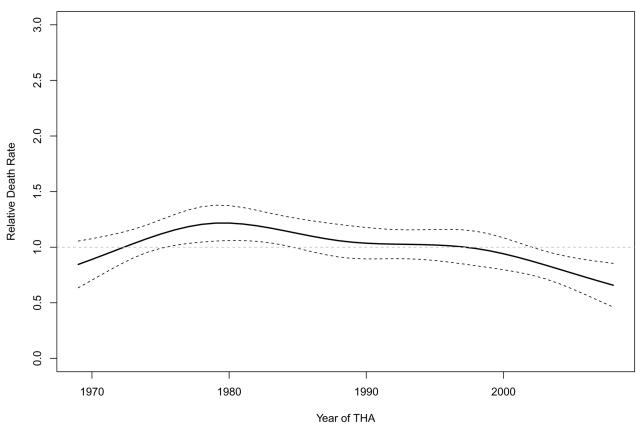
Figure 1B



#### Figure 1.

Figure 1A. Relative Death Rates (observed/expected) by Age: Total Hip Replacement Figure 1B. Relative Death Rates (observed/expected) by Age: Total Knee Replacement



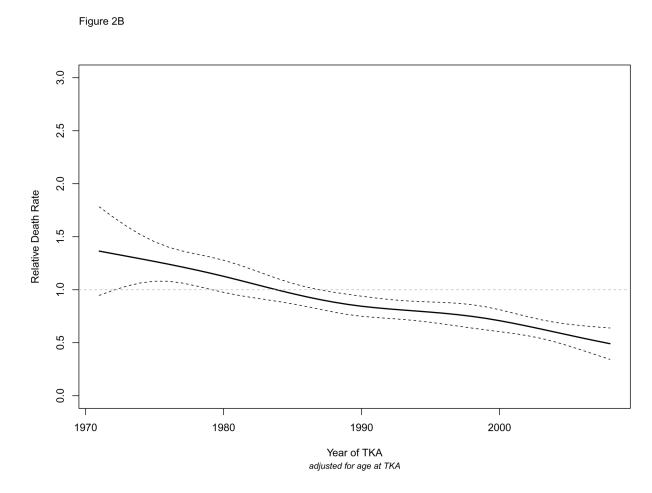


Year of THA adjusted for age at THA



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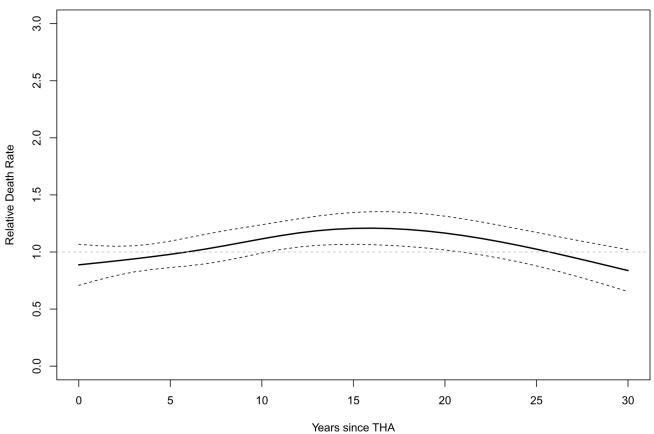


#### Figure 2.

**Figure 2 A.** Relative Death Rates (observed/expected) by Calendar Year of Surgery: Total Hip Replacement

**Figure 2 B.** Relative Death Rates (observed/expected) by Calendar Year of Surgery: Total Knee Replacement





adjusted for age at THA



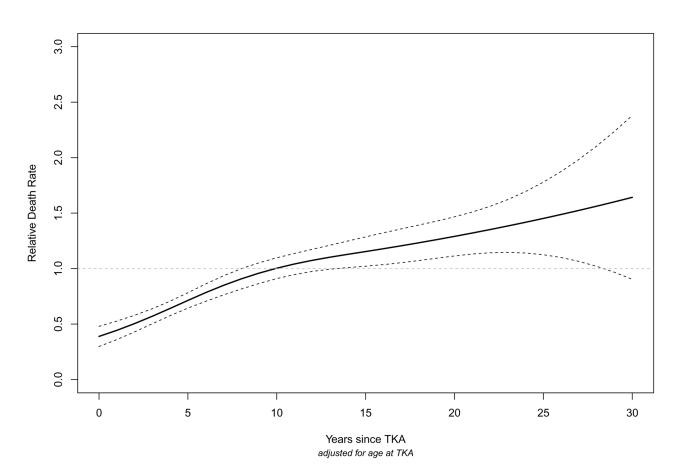


Figure 3.

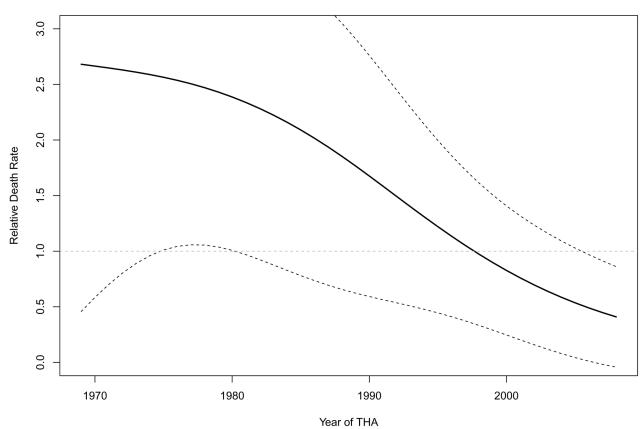
**Figure 3A.** Relative Death Rates (observed/expected) by Time since Surgery: Total Hip Replacement

**Figure 3B.** Relative Death Rates (observed/expected) by Time since Surgery: Total Knee Replacement

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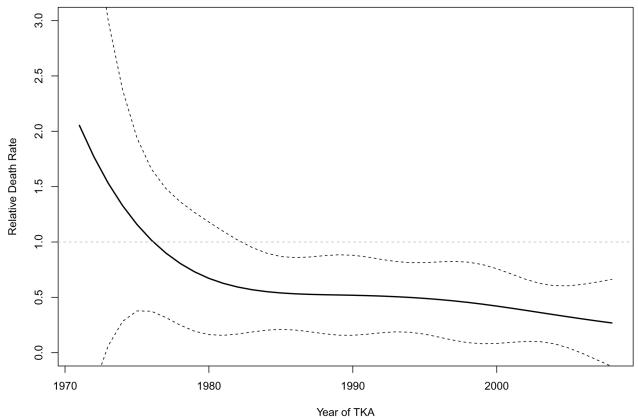
Figure 4A

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Year of THA adjusted for age at THA

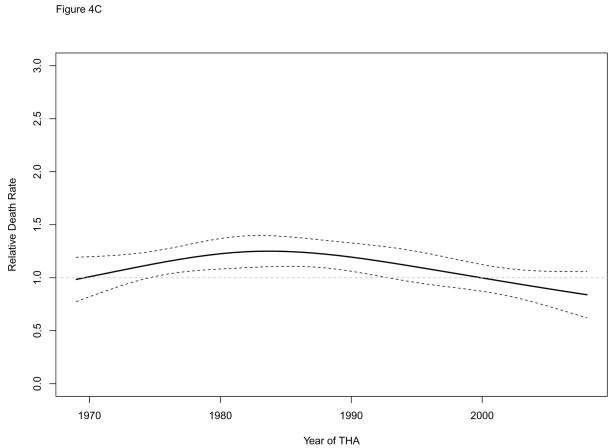




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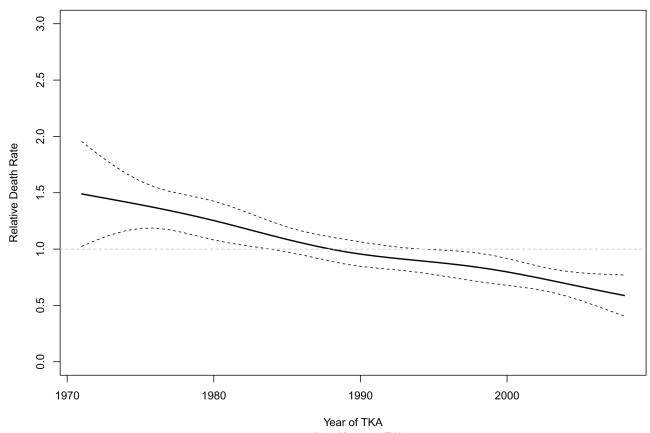


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adjusted for age at THA





adjusted for age at TKA

#### Figure 4.

Figure 4A. Relative Death Rates within the First Year of Surgery: Total Hip ReplacementFigure 4B. Relative Death Rates within the First Year of Surgery: Total Knee ReplacementFigure 4C. Relative Death Rates beyond the First Year: Total Hip ReplacementFigure 4D. Relative Death Rates beyond the First Year: Total Knee Replacement

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# Table 1

Observed and Expected Number of Deaths and Standardized Mortality Ratios (95% Confidence Intervals) in Total Hip and Knee Replacement

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	To	Total Hip Replacement	acement	Tot	Total Knee Replacement	lacement
	Observed	Expected	SMR (95% CI)	Observed	Expected	SMR (95% CI)
Overall	1001	1184	0.92 (0.87, 0.98)	1010	1200	0.84 (0.79, 0.90)
Osteoarthritis	814	995	0.82 (0.76, 0.88)	842	1049	0.80 (0.75, 0.86)
Sex						
Men	412	472	0.87 (0.79, 0.96)	401	504	0.80 (0.72, 0.88)
Women	619	712	0.95 (0.88, 1.03)	609	969	0.88 (0.81, 0.95)
Age groups, years						
<60	147	108	1.36 (1.15, 1.60)	86	69	1.25 (0.99, 1.54)
61–70	297	338	0.88 (0.78, 0.98)	260	293	0.89 (0.78, 1.00)
71–80	439	473	0.93 (0.84, 1.02)	482	589	0.82 (0.75, 0.90)
>80	208	265	0.79 (0.68, 0.90)	182	250	0.73 (0.63, 0.84)