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Hypothyroidism Increases 90-Day Complications and Costs Following Primary Total Knee Arthroplasty

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ABSTRACT

Background: Thyroid disease is common and often remains undetected in the US population. Thyroid hormone has an array of metabolic, immunologic, and musculoskeletal functions crucial to well-being. The influence of thyroid disease on perioperative outcomes following primary total knee arthroplasty (TKA) is poorly understood. We hypothesized that hypothyroidism was associated with a higher risk of postoperative complications and 90-day costs following primary TKA.

Methods: The Medicare standard analytical files were queried using International Classification of Disease codes between 2005 and 2014 to identify patients undergoing primary TKA. Patients with a diagnosis of hypothyroidism were matched by age and gender on a 1:1 ratio. Ninety-day postoperative complication rates, day of surgery, and 90-day global period charges and reimbursements were compared between matched cohorts.

Results: A total of 2,369,594 primary TKAs were identified between 2005 and 2014. After age and gender matching, each cohort consisted of 98,555 patients. Hypothyroidism was associated with greater odds of postoperative complications compared to matched controls (odds ratio 1.367, 95% confidence interval 1.322–1.413). The 90-day incidence of multiple postoperative medical and surgical complications, including periprosthetic joint infection, was higher among patients with hypothyroidism. Day of surgery and 90-day episode of care costs were significantly higher in the hypothyroidism cohort.

Conclusion: This study demonstrated an increased risk of multiple postoperative complications and higher costs among patients with hypothyroidism following primary TKA. Surgeons should counsel patients on these findings and seek preoperative optimization strategies to reduce these risks and lower costs in this patient population.

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Total knee arthroplasty (TKA) is a cost-effective [1,2] option for reducing pain and improving function among patients with end-stage knee arthritis [3]. Not surprisingly, it is one of the most frequently performed procedures in the United States [4–6],

imposing an enormous economic burden on the healthcare system [7]. Although complications following TKA are rare [8], the personal and fiscal costs can be overwhelming [9–12]. Attention has recently focused on identifying risk factors associated with adverse events [13–20], and worse clinical outcomes [21–24], following TKA, presuming the identification of modifiable variables may allow for preoperative intervention, potentially improving outcomes and decreasing cost.

Between 4.6% and 12% of the US population are affected by thyroid disease [25,26]. Thyroid hormone has a number of systemic and musculoskeletal functions including bone remodeling and maintaining articular cartilage health [27,28]. Research suggests an increased risk of osteoarthritis among people with thyroid disease

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[29,30]. The reported prevalence of hypothyroidism in the total joints population is up to 18%, which is significantly higher than in the general population [31]. Hypothyroidism has previously been associated with an increased risk of periprosthetic joint infection (PJI) in a retrospective, single center study [32]. However, no prior research has evaluated the influence of thyroid disease on postoperative outcomes or cost following primary TKA. Therefore, we sought to evaluate the influence of hypothyroidism on complication rates and cost of primary TKA using a large national database. We hypothesized that, following primary TKA, hypothyroidism was associated with a higher risk of postoperative complication as well as an increase in the 90-day episode of care cost compared to patients without thyroid disease.

Materials and Methods

A retrospective, case-control study was performed to investigate the effects of hypothyroidism on 90-day complications and costs following TKA. The PearlDiver Supercomputer (Warsaw, IN) was utilized to query the entire Medicare sample from 2005 to 2014 by way of the standard analytical files. Medicare releases 100% of their inpatient and outpatient hospital billing data across all service locations nationwide. These data are deidentified and cohorts less than 11 are removed and these data are sent to PearlDiver for distribution subscribers. The dataset used for this study is composed of 100% of the records of Medicare patients from 2005 to 2014 without any patient identifiers. This publically and commercially available HIPAA compliant server provides the ability to identify and track patient information based on International Classification of Disease (ICD) and Current Procedural Terminology (CPT) codes. The ICD and Current Procedural Terminology codes utilized in this search have been previously used in the literature [33,34]. Patients with ICD-9 codes for hypothyroidism were identified based on codes 2440 (postsurgical hypothyroidism), 2441 (other/postablative hypothyroidism), 2442 (iodine hypothyroidism), 2443 (iatrogenic hypothyroidism), and 2448 (acquired hypothyroidism not elsewhere classifiable/other specified acquired hypothyroidism). Patients with a diagnosis of hyperthyroidism (ICD-9 codes 242.0-242.3 and 242.9) were excluded. No other exclusion criteria were utilized. All diagnosis slots were used to identify a diagnosis of hyperthyroidism or hypothyroidism. A comparison of the Charlson Comorbidity Index (CCI), a measurement of morbidity and mortality risk shown to be associated with worse outcomes following orthopedic surgery [33], was performed to determine the similarity between cohorts. Following the creation of the cohorts, the server performs a random selection of patients to match 2 cohorts with the same age and gender distribution with the only differentiating factor being the presence or absence of hypothyroidism. This also provides a method to randomize each cohort to match the same criteria, which allows confounders to be randomly present in both cohorts. Patients were matched based on age, gender, and CCI at a 1:1 ratio prior to cost and outcome analysis (Table 1).

Ninety-day complication rates were tracked through ICD-9 coding. Day of surgery and 90-day global period charges and reimbursements were analyzed as markers of costs, using previously described techniques [33,35]. We selected a 90-day global period to represent the most common time period for bundle payment initiatives as depicted in the Comprehensive Care for Joint Reconstruction model.

Given that the data were extracted from a HIPAA compliant server, the current study did not require Institutional Review Board approval. Odds ratios (ORs) and 95% confidence intervals were calculated. Independent samples 2-tailed t-tests were performed for the comparison of means. A *P*-value of <.05 was used to define

Table 1

Matched Medicare Beneficiaries Undergoing TKA During the Study Period With and Without a Diagnosis of Hypothyroidism.

| Patient Demographics | Number of Patients (%) |
|----------------------|------------------------|
| Age group | |
| 64 and under | 11,513 (11.7) |
| 65-69 | 28,730 (29.2) |
| 70-74 | 24,142 (24.5) |
| 75-79 | 19,543 (19.8) |
| 80-84 | 11,160 (11.3) |
| 85 and over | 3467 (3.5) |
| Gender | |
| Female | 81,306 (82.5) |
| Male | 17,249 (17.5) |

Patient age and gender demographics for the control (n = 98,555) and hypothyroid (n = 98,555) cohorts.

statistical significance [36]. All data were analyzed using the software Statistical Package for Social Sciences [SPSS] version 23 (Chicago, IL). No external funding source was used for the conduct of this study.

Results

A total of 2,369,594 primary TKAs were identified as having been performed between 2005 and 2014. During the same time period, within the database, a total of 1,277,014 patients had a diagnosis of hypothyroidism. After age and gender matching and randomization, each cohort comprised 98,555 patients (Table 1). Due to the matching process, the gender and age distributions among both cohorts were the same. Comparison of CCI scores between cohorts revealed no statistically significant difference (*P* = .892). Analysis of the cumulative 90-day outcomes demonstrated that following primary TKA, patients with a diagnosis of hypothyroidism were at increased odds of medical and surgical complications (9.7% vs 7.09%, OR 1.367, 95% CI 1.322-1.413) compared to the matched cohort. Additionally, patients with a diagnosis of hypothyroidism experienced a significantly higher rate of many individual medical complications compared to the matched cohort (Table 2). Similarly, postoperative infections were significantly more frequent among patients with hypothyroidism (Table 3). Compared to the matched cohort, patients with hypothyroidism incurred significantly greater day of surgery (\$54,459.39 standard deviation (SD): \$36,026.99 vs \$52,181.17 SD: \$33,476.67) and 90-day episode of care (\$61,622.39 SD: \$47,900.80 vs \$57,871.69 SD: \$42,333.69) charges and reimbursements (*P* < .001) (Table 4).

Discussion

Although the prevalence of thyroid disease among patients undergoing total joint arthroplasty is reportedly high [32], previous evaluation of its influence on outcomes following primary TKA is limited. This study found hypothyroidism to be a significant risk factor for postoperative medical and surgical complications and that it is associated with increased episode of care costs following primary TKA.

Patients with hypothyroidism were found to have significantly greater odds of thromboembolic disease including deep venous thrombosis (OR 1.252) and pulmonary embolism (OR 1.206) compared to matched controls. Additionally, hypothyroidism was associated with greater odds of acute postoperative anemia (OR 1.326) requiring a blood transfusion (OR 1.428). These findings may be explained by the direct and indirect effects of thyroid hormone on platelet maturation and function, the synthesis and action of coagulation factors, and the maintenance of blood viscosity [37].

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