Introduction

Postoperative venous thromboembolism (VTE), including deep vein thrombosis (DVT) and pulmonary embolism (PE), is a life-threatening complication in patients undergoing surgery. Total knee arthroplasty (TKA) has a high risk of DVT and PE, and because a thromboembolic event is fatal, routine thromboprophylaxis is recommended after TKA. Different modalities including the use of pharmacological agents and mechanical compression devices have been introduced with the aim of decreasing the prevalence of DVT and PE after TKA. Among those, pharmacological agents are usually used to prevent DVT, but they can increase the risk of bleeding complications. Mechanical compression devices have no risk of bleeding complications, but they are relatively less effective.

Some reports have described a low prevalence of DVT and PE after TKA in Asian patients compared with western populations. It is generally believed that the rate of VTE is low in Asia, so prophylaxis is seldom used. The differences in VTE prevalence have been attributed to differences in genetic background and dietary habits. In contrast, some reports have shown that dietary habits of Asians have changed similar to those of westerners and that the incidence of thromboembolic events has increased.

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If the occurrence of a thromboembolic event is low, prophy-
laxis will result in unnecessary costs and complications related to bleeding. Therefore, whether pharmacological thromboprophylaxis is necessary to decrease the prevalence of DVT and PE after TKA in Asian patients is a legitimate question to raise. Mechanical compression thromboprophylaxis method may be sufficient for Asian patients.

The purpose of this study was to investigate the incidence of thromboembolic events (specifically, DVT and PE) and complications related to bleeding after TKA in patients treated with a mechanical compression device only and those treated with the same device in combination with a pharmacological agent.

We hypothesized that 1) there would be no statistically significant difference in the incidence of thromboembolic events including of DVT and PE between patients treated after TKA with mechanical compression only and those treated with compression and low-molecular-weight heparin (LMWH) and 2) bleeding-related complications would be less prevalent in those treated using mechanical compression only.

Materials and Methods

1. Patients

A total of 1,156 primary cemented TKAs were done in 745 patients from March 2009 to March 2015. Inclusion criteria were primary TKA using a mechanical compressive device with or without pharmacologic agents for prevention of DVT and PE. Exclusion criteria were the use of warfarin or aspirin before surgery for the treatment of other vascular, cardiac, or cerebral diseases and patients with any history of previous PE or DVT. We excluded 256 patients (380 knees) because they used warfarin or aspirin preoperatively and also for up to 48 hours following TKA for the aforementioned reasons. The remaining 489 patients (776 knees) were reviewed retrospectively. Of these, 233 patients (354 knees) used only a mechanical compression device without LMWH and 256 patients (422 knees) used the same mechanical compressive device along with LMWH. Of the 489 patients, 287 patients (574 knees) were reviewed retrospectively. Of these, 233 patients (354 knees) used only a mechanical compression device without LMWH and 256 patients (422 knees) used the same mechanical compressive device along with LMWH. The 489 patients, 287 patients (574 knees) underwent bilateral TKA with a one-week interval between procedures and 202 patients underwent unilateral TKA. The 489 patients comprised 86 males and 403 females, with a mean age of 68.6±7.6 years (range, 52 to 81 years) and mean body mass index (BMI) of 27.4 kg/m$^2$ (range, 23.0 to 30.5 kg/m$^2$). The average follow-up was 3.6 years (range, 2 to 5 years). All patients had osteoarthritis of the knee. The demographic data including age, sex, and BMI were compared between two groups. Histories related to thrombotic risk factors including smoking habits, congestive heart failure, and cancer were assessed and recorded. Preoperative coagulation assays (platelet count, prothrombin time, partial thromboplastin time, fibrinogen), full blood count, blood typing, and serum chemical profiling were performed for all patients.

2. Surgical Procedure

Surgeries were all performed by the same surgeon (SJY) using a medial parapatellar approach. The press-fit condylar TKA (Johnson & Johnson, Raynham, MA, USA) or Scorpio NRG (Stryker Inc., Mahwah, NJ, USA) was used with tourniquet inflation to 350 mmHg. After a skin incision, a capsular incision was performed to expose the medial and lateral condyles, with subsequent tibial plateau and medial soft tissue release. A measured resection technique was used for the proximal tibial and distal femoral bone cuts. The tibial cut was made perpendicular to the mechanical axis using an extra-medullary alignment guide, with the posterior tibial slope set at 0°. After thoroughly removing posterior femoral osteophytes, a trial implant was inserted to assess the joint space, varus/valgus stability, patellar tracking, and condylar lift-off. Cement was used for tibial and femoral fixation. Patellar resurfacing was not performed in all knees. Tibial osteophyte removal and chondral shaving were performed. If there was synovial thickening observed in a given patient, a synovectomy was performed as well.

Straight leg raising exercises were initiated from the day of surgery. The drain was removed from the second postoperative day and continuous passive motion was started. From the first postoperative week, weight bearing ambulation was allowed.

3. Thromboembolic Event Prophylaxis

The DVT-3000 impulse system (DS Maref, Gunpo, Korea) was used on both legs postoperatively for mechanical compression prophylaxis of all patients. Mechanical compression was started on the day of the operation and continued for 10 to 14 days after the operation until the patients were discharged home. In the bilateral TKA patients, the mechanical compressive device was applied from the first operation day to one day before the second operation. We used sleeves with three circumferential air chambers running the distance from the ankle to the knee. Pharmacological thromboprophylaxis involved LMWH in all patients. These patients received a subcutaneous injection of LMWH in the form of enoxaparin 4000 IU (Sanofi-Aventis, Gentilly, France) beginning 24 hours after surgery $^{18}$. 

4. Assessment of Thromboembolic Events

Color Doppler ultrasonography was performed for bilateral
common femoral veins, superficial veins, popliteal veins, and calf veins by skilled physicians seven days after surgery in all patients.

D-dimer levels were measured and PE and low extremity venography dynamic computed tomography were performed for patients suspected of DVT or PE. Symptoms of DVT and PE including chest pain, dyspnea, and painful swelling on lower extremities and positive findings on color Doppler ultrasonography were considered grounds for exclusion. For evaluation of bleeding tendency, the postoperative hemovac drainage amount and bleeding-related complications were checked and compared between the two groups. If hemoglobin was lower than 8 g/dL or the patient showed hemodynamic instability, transfusion was performed.

5. Statistical Analysis

Differences of parameters including postoperative hemovac drainage, age, BMI, preoperative coagulation assay, full blood count, and serum chemical profile tests between both groups were assessed using Student t-test or Wilcoxon nonparametric test. The incidence of thromboembolic events and bleeding-related complications between patients with and without PE were compared using chi-square and Fisher exact tests. Significance was set at p<0.05. The statistical analyses were performed using SPSS ver. 12.0 (SPSS Inc., Chicago, IL, USA).

Results

There were no significant differences between the two groups in demographic characteristics and preoperative coagulation assay results (Table 1). The incidence of symptomatic DVT was 15 of 375 knees (4.0%) in patients treated only using mechanical compression and 14 of 401 knees (3.4%) in patients treated with the combination of mechanical compression and LMWH. The prevalence of symptomatic PE was 5 of 375 knees (1.3%) in patients treated with mechanical compression alone and 5 of 401 knees (1.2%) in those treated with compression and LMWH.

Table 1. Demographics and Preoperative Coagulation Assay of Each Group

| Parameter                        | Mechanical compression only group (n=233) | Mechanical compression and pharmacological agent group (n=256) | p-value |
|----------------------------------|----------------------------------------|---------------------------------------------------------------|---------|
| Age (yr)                         | 64.3±10.7                              | 68.4±11.4                                                    | 0.174*  |
| Sex (M:F)                        | 35:49                                  | 50:74                                                        | 0.164*  |
| Body mass index (kg/m²)          | 26.2±3.0                               | 27.9±2.7                                                    | 0.153*  |
| Preoperative coagulation assay    |                                        |                                                               |         |
| Prothrombin time international normalized ratio | 1.14±0.4                              | 1.10±0.3                                                    | 0.156*  |
| Activated partial prothrombin time (sec) | 24.4±7.3                              | 27.0±7.1                                                    | 0.135*  |
| Platelet count (10³)             | 215±87                                 | 232±79                                                      | 0.154*  |

Values are presented as mean±standard deviation.

*Student t-test. Chi-squared test.

Table 2. Incidence of Thromboembolic Events in Each Group

| Complication              | Mechanical compression only group (n=233) | Mechanical compression and pharmacological agent group (n=256) | p-value |
|---------------------------|----------------------------------------|---------------------------------------------------------------|---------|
| Deep vein thrombosis (%)  | 15 (4)                                 | 14 (3)                                                       | 0.125   |
| Pulmonary embolism (%)    | 5 (1.3)                                | 5 (1.2)                                                      | 0.146   |

Table 3. Complications Related to Bleeding in Each Group

| Complication                  | Mechanical compression only group (n=233) | Mechanical compression and pharmacological agent group (n=256) | p-value |
|-------------------------------|----------------------------------------|---------------------------------------------------------------|---------|
| Hematoma (%)                  | 0 (0)                                  | 4 (2)                                                        | 0.024*  |
| Postoperative hemovac drainage (mL), mean±SD | 635±57                               | 813±84                                                      | 0.013*  |

SD: standard deviation.

*Chi-squared test. Student t-test.
The incidences of DVT and PE were not significantly different between the groups (Table 2). Postoperative hemovac drainage amount was 635±57 mL and 813±84 mL in the respective group, indicating a significant difference (Table 3). Thrombi that developed in 39 knees resolved completely regardless of their site or size without pharmacological treatment as assessed by sonogram and venogram performed 6 months postoperatively. No patient had symptomatic PE as indicated by the lack of abnormal findings on perfusion lung scan and the absence of symptoms.

Discussion

One of the principle findings of this study is the lack of statistical differences in the incidence of thromboembolic events including DVT and PE between treatment with a mechanical compressive device alone and compression delivered along with LMWH after TKA. Another principle finding is the greater bleeding tendency in terms of postoperative hemovac drainage in patients treated with the combination approach (i.e., greater bleeding after the use of LMWH).

There are LMWH, rivaroxaban, fondaparinux, warfarin, and aspirin in pharmacological methods for prophylaxis of DVT and PE. LMWH inhibits factor Xa. The major advantage of LMWH is that no monitoring is required, but it needs to be administered via a subcutaneous injection. Rivaroxaban is a direct Xa inhibitor that is taken orally and requires no monitoring. But rivaroxaban was associated with a higher rate of reoperation than was LMWH after TKA. Fondaparinux is a synthetic pentasaccharide and an indirect factor Xa inhibitor. The drug has had limited use in North America because of concerns related to bleeding.

Nagase et al. reported that the prevalence of postoperative PE after the use of mechanical prophylaxis alone was 0.66%, which was similar to 0.40% after the combined use of fondaparinux and mechanical prophylaxis in patients with total hip arthroplasty or TKA. Kim et al. reported that the prevalence of DVT was 94 of 1434 knees (6.6%) in patients with a mechanical compression device after TKA. There was a low prevalence of DVT and PE in Asian patients, and all thrombi resolved completely and spontaneously without treatment using anticoagulants in a study by Kim et al. Kim et al. performed a systematic review and meta-analysis that addressed the rate of VTE after orthopedic surgery in Asian patients without thromboprophylaxis. The pooled rates of proximal and symptomatic DVT were lower in Asians than those in western reports and none of these Asian patients with orthopedic surgery died from VTE. Especially, the prevalence of DVT in Koreans was lower than that in other Asians (Korean, 7.5%; South-East Asian, 11.8%; and Japanese, 11.0%).

The present study compared the incidence of DVT and PE between patients treated using only a mechanical compressive device and those treated using the device along with LMWH after TKA. This comparison has not been done before, except in a systematic review. In our study, the incidences of symptomatic DVT and PE were 15 of 375 knees (4.0%) and 5 of 375 knees (1.3%) in patients treated with mechanical compression alone. There was no significant difference compared to the group with the combined use of a mechanical compressive device and LMWH where the incidences were 14 of 401 knees (3.4%) and 5 of 401 knees (1.2%). These incidences are similar to those described in the previous systematic review study.

There were some limitations in our study. It was a retrospective study, not a prospective randomized study, subject to selection bias. However, to the best of our knowledge, there has been no randomized controlled trial conducted on this subject; therefore, a randomized prospective study needs to be performed. Second, the number of patients was relatively small considering the low incidences of DVT and PE. Because the incidences of DVT and PE were low, more patients will be needed to achieve statistical significance. Third, the study involved only Korean patients. Therefore, the results may not be applicable to other ethnic groups. Fourth, we used only LMWH as the pharmacological agent. Therefore, the group treated with a mechanical compressive device and LMWH in our study does not represent groups that use other pharmacological agents. Fifth, we did not include other groups for comparison such as a group that used only pharmacological agent or a group that used neither a pharmacological agent nor a mechanical compressive device. However, we aimed to investigate if the use of a mechanical compression device alone would be sufficient to prevent thromboembolic events after TKA in Asians. In our patients, the additional use of a pharmacological agent did not make a difference in prevention of thromboembolic events, and rather the bleeding-related complication rate was lower in the group with compression treatment only. Therefore, we think the comparison was appropriate for the study purpose.

Conclusions

Mechanical compression after TKA resulted in a low prevalence of DVT and PE, similar to the combined use of mechanical compression and LMWH as a pharmacological agent. Complications related to bleeding occurred less often in patients treated only using mechanical compression. Therefore, the use of mechanical
compression only for prophylaxis against DVT and PE can be an attractive option for Korean patients.

Conflict of Interest

No potential conflict of interest relevant to this article was reported.

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