Risk-Stratified Thromboprophylaxis Effects of Aspirin versus Low-Molecular-Weight Heparin in Orthopaedic Trauma Patients

Nathan N O'Hara¹, Robert V O'Toole, Katherine Frey², Deborah M. Stein, Renan C Castillo³, Gregory T Altman, Julie Agel⁴, Michael J Bosse, Anthony R Carlini⁵, Eben A Carroll⁶, Yasmin Degani, Reza Firoozabadi⁷, Joshua Layne Gary⁸, Greg E Gaski, Elliott Haut⁹, COL (ret) Roman A Hayda, Robert A Hymes, Madhav A Karunakar, Conor P Kleweno¹⁰, Todd Owen McKinley¹¹, William T. Obremskey¹², Gerard Slobogean¹³, Kuladeep R Sudini, Tara J Taylor⁵, Heather A Vallier¹⁴, Stephen James Warner¹⁵, Michael John Weaver¹⁶, John C Weinlein ¹University of Maryland, ²Johns Hopkins University, ³John Hopkins Bloomberg School of Public Health, ⁴Harborview Med

¹University of Maryland, ²Johns Hopkins University, ³John Hopkins Bloomberg School of Public Health, ⁴Harborview Med Ctr, ⁵Johns Hopkins Bloomberg School of Public Health, ⁶Wake Forest Univ School Of Medicine, ⁷UW Medicine, ⁸Keck School of Medicine of USC Department of Ortho, ⁹The Johns Hopkins University School of Medicine, ¹⁰University of Washington Orthopaedics, ¹¹IU Health Physicians, ¹²Vanderbilt Ortho Inst, ¹³University of Maryland School of Medicine, ¹⁴Metrohealth Med Ctr, ¹⁵McGovern Medical School At Uthealth, ¹⁶Brigham and Women's Hospital

INTRODUCTION: The PREVENT CLOT trial concluded thromboprophylaxis with aspirin was noninferior to low-molecular-weight heparin (LMWH) in preventing death in orthopaedic trauma patients. This secondary analysis sought to determine if the effects of aspirin versus LMWH differed based on patients' baseline risk of venous thromboembolism (VTE).

METHODS: This PREVENT CLOT secondary analysis included 12,211 adult patients indicated for thromboprophylaxis due to a surgically treated extremity fracture or pelvis or acetabulum fracture. We used the Caprini Score to calculate patients' baseline risk of VTE, stratifying patients into risk quartiles ranging from low (<1%) to high risk (>10%). The primary outcome was a composite of thromboembolic outcomes within 90 days. We assessed treatment effects using the win ratio method. This approach paired all patients in each treatment group and compared the outcomes hierarchically, starting with death, followed by pulmonary embolism, deep vein thrombosis, then bleeding when the patients could not be differentiated on a more severe outcome. The win ratio is the number of wins in the aspirin arm divided by the number of wins in the LMWH arm.

RESULTS: In the high-risk quartile (n=3,052), 46% of patients had a femur fracture, 42% had a pelvis or acetabulum fracture, 48% had a thoracic injury, 39% had a spinal injury, and 35% had a head injury. Low-risk quartile (n=3,053) patients most commonly had a tibia fracture (67%), and few had thoracic (5%), head (1%), or spinal (1%) injuries. In the high-risk quartile, there was no statistical difference in the effect of aspirin compared to LMWH on the composite outcome (win ratio, 0.95; 95% CI, 0.83 - 1.09, p=0.48). This result was consistent in the low-risk quartile (win ratio, 1.09; 95% CI, 0.85 - 1.40, p=0.51), low-medium risk quartile (win ratio, 1.01; 95% CI, 0.82 - 1.26, p=0.90), and medium-high risk quartile (win ratio, 0.98; 95% CI, 0.83 - 1.15, p=0.78).

DISCUSSION AND CONCLUSION: Thromboembolic outcomes were similar when either aspirin or LMWH is used for prophylaxis. This result held true even when considering patients at highest risk of VTE.