






ORIGINAL RESEARCH

Alert-Triggered Patient Education Versus Nurse Feedback for Nonadministered Venous Thromboembolism Prophylaxis Doses: A Cluster-Randomized Controlled Trial

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BACKGROUND: Many hospitalized patients are not administered prescribed doses of pharmacologic venous thromboembolism prophylaxis.

METHODS AND RESULTS: In this cluster-randomized controlled trial, all adult non-intensive care units (10 medical, 6 surgical) in 1 academic hospital were randomized to either a real-time, electronic alert-triggered, patient-centered education bundle intervention or nurse feedback intervention to evaluate their effectiveness for reducing nonadministration of venous thromboembolism prophylaxis. Primary outcome was the proportion of nonadministered doses of prescribed pharmacologic prophylaxis. Secondary outcomes were proportions of nonadministered doses stratified by nonadministration reasons (patient refusal, other). To test our primary hypothesis that both interventions would reduce nonadministration, we compared outcomes pre- versus postintervention within each cohort. Secondary hypotheses were tested comparing the effectiveness between cohorts. Of 11 098 patient visits, overall dose nonadministration declined significantly after the interventions (13.4% versus 9.2%; odds ratio [OR], 0.64 [95% CI, 0.57–0.71]). Nonadministration decreased significantly ($P<0.001$) in both arms: patient-centered education bundle, 12.2% versus 7.4% (OR, 0.56 [95% CI, 0.48–0.66]), and nurse feedback, 14.7% versus 11.2% (OR, 0.72 [95% CI, 0.62–0.84]). Patient refusal decreased significantly in both arms: patient-centered education bundle, 7.3% versus 3.7% (OR, 0.46 [95% CI, 0.37–0.58]), and nurse feedback, 9.5% versus 7.1% (OR, 0.71 [95% CI, 0.59–0.86]). No differential effect occurred on medical versus surgical units. The patient-centered education bundle was significantly more effective in reducing all nonadministered ($P=0.03$) and refused doses ($P=0.003$) compared with nurse feedback (OR, 1.28 [95% CI, 1.0–1.61]; $P=0.03$ for interaction).

CONCLUSIONS: Information technology strategies like the alert-triggered, targeted patient-centered education bundle, and nurse-focused audit and feedback can improve venous thromboembolism prophylaxis administration.

REGISTRATION: URL: <https://www.clinicaltrials.gov>; Unique identifier: NCT03367364.

Key Words: deep vein thrombosis ■ patient-centered care ■ prophylaxis ■ pulmonary embolism ■ randomized trial ■ venous thromboembolism

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CLINICAL PERSPECTIVE

What Is New?

- This trial shows that each intervention, (1) a real-time electronic, alert-triggered, patient-centered education bundle and (2) nurse performance feedback, independently and successfully reduced missed doses of prescribed venous thromboembolism (VTE) prophylaxis among hospitalized patients.

What Are the Clinical Implications?

- Hospital-associated VTE is a largely preventable condition when hospitalized patients are prescribed and receive appropriate VTE prophylaxis; however, many doses of prescribed prophylaxis are not administered to hospitalized patients.
- These interventions demonstrate that missed doses of VTE prophylaxis in hospitalized patients can be significantly reduced by engaging patients and nurses with timely, informative, actionable information about the importance of VTE prophylaxis.

Nonstandard Abbreviations and Acronyms

PCORI Patient-Centered Outcomes Research Institute

Hospitalization is a primary risk factor for venous thromboembolism (VTE),¹ and deep vein thrombosis and pulmonary embolism are leading causes of preventable harm among patients in the hospital.^{2,3} As a result, national organizations focused on health care quality improvement, including the Agency for Healthcare Research and Quality and The Joint Commission, measure and report VTE prevention practices.^{4,5} Efforts to improve prescription of VTE prophylaxis for hospitalized patients have achieved great success.^{6–9} However, evidence shows that many doses of prescribed pharmacologic VTE prophylaxis are not administered and most often attributed to patient refusal or poor patient–nurse communication.^{10–14} Missing even 1 dose of a VTE prophylaxis medication has been associated with developing VTE events.^{15,16} VTE prophylaxis medications are some of the most frequently nonadministered medications for hospitalized patients.^{17,18}

Reasons for nonadministration of prescribed pharmacologic VTE prophylaxis have been attributed to nurse attitudes, knowledge, and beliefs on the need for prophylaxis,¹⁴ and patients refusing doses. Moreover, recent American Society of Hematology guidelines do not address nonadministration of prescribed doses,^{19,20}

which is a missed opportunity to encourage best practices to prevent VTE. Previous interventions have targeted nonadministration practices and significantly improved administration of prescribed prophylaxis doses, including comprehensive VTE-specific nurse education²¹ and widespread patient education.^{11,22} Most recently, we showed that a real-time, patient-centered, education bundle (bundle) implemented on 4 adult non-intensive care units can reduce nonadministered doses by >40% in hospitalized patients.²³ This bundle intervention was part of an extramurally funded research project and had substantial human resources that are not sustainable long term or generalizable to a wide range of hospitals.

The current study is part of a dissemination effort. The goal was to scale use of the bundle to other adult non-intensive care units and transition implementation from the research team to the nurses for integration in their unit's routine medication administration process without additional support of a research nurse, as was done in the original trial. This approach has already been shown to be effective at a community hospital.²⁴ We aimed to test the effectiveness of unit staff nurses implementing the bundle triggered by a real-time electronic alert.²³ While planning our intervention, individualized nurse performance feedback and coaching (nurse feedback) was suggested as an alternative approach to reduce nonadministration of prescribed in-hospital pharmacologic VTE prophylaxis. We decided to implement both strategies simultaneously as part of a rigorously performed cluster-randomized trial. Our primary a priori hypothesis was that each intervention would improve medication administration. Because patient refusals account for the majority of nonadministered doses,¹⁰ and the bundle has significantly decreased refusals in prior studies,^{23,24} our secondary a priori hypotheses were that the alert-triggered patient education bundle would be more effective than nurse feedback in reducing overall nonadministered doses for any reason and in reducing patient refusal.

METHODS

Data are available from the Johns Hopkins Medicine Institutional Data Trust for researchers who meet the criteria for access to confidential data. The data set in question contains protected health information, which by nature of the study cannot be deidentified. For the purpose of this study, identifiable information included patient identifiers, nurse identifiers, dates of hospitalization, dates/times of medication administration, and hospital location, which are all required components for the analysis that would potentially enable individuals to identify specific patients and nurses. To request access to the data, please reach out to Dr. Daniel

E. Ford, MPH, Vice Dean for Clinical Investigation at dford@jhmi.edu.

Study Design and Setting

ENACT (Patient Education Bundle Versus Nurses Feedback and Coaching to Prevent Missed Doses of Venous Thromboembolism Prophylaxis) was a cluster-randomized controlled trial involving all 16 adult non-intensive care medical (n=10) and surgical (n=6) units at The Johns Hopkins Hospital, an urban academic tertiary care hospital. The Johns Hopkins Medicine Institutional Review Board approved the trial and provided a waiver of consent. The trial was registered on clinicaltrials.gov (<https://clinicaltrials.gov/ct2/show/NCT03367364>).

Randomization and Blinding

We performed a stratified block randomization to balance the distribution of medicine (n=10) and surgery (n=6) units and of baseline dose administration performance (high versus low performers) in each intervention arm. We leveraged unit-specific data on VTE prophylaxis nonadministration practices of nurses from the electronic health record (EHR) system, and in a research team meeting rank ordered units by performance. After group consensus on the block assignment, an actual physical coin toss (heads or tails) within each block assigned units to either the bundle or the nurse feedback intervention. Eight units (3 surgical, 5 medical) were assigned to each arm. Our biostatisticians (J.W., G.Y.) were blinded to unit intervention assignments.

Interventions

Patient-Centered Education Bundle

The bundle was developed and validated in the original study and the details published.^{23–26} Briefly, the bundle used the same educational approach, which included a 1-on-1 personalized discussion with the patient, supplemented by a 2-page paper handout (available in 13 languages), and a 10-minute video (bit.ly/bloodclots).²³ In the original study, a Patient-Centered Outcomes Research Institute (PCORI)-funded research nurse educator implemented the bundle, asking the patient their preferred mode of education, spending a median of 10 minutes on the patient intervention.²³ In the current study, the nursing teams providing routine clinical care on the units replaced that funded nurse educator and implemented the bundle. The bundle process following the alert involved engaging the bedside nurse who documented the nonadministered dose to determine the cause and intervening with education targeting the nurse, the patient, or both. To prepare units for implementation, we provided standardized educational

materials (posters, badge cards, and slides) and trained nurses to deliver the education in real time, offering them scripted talking points when discussing VTE prevention with patients. Because the intent of dissemination was independent implementation of the bundle during routine medication administration, fidelity to the intervention and patient preferences to which education they preferred were not tracked.

Real-Time Alert

At The Johns Hopkins Hospital, every patient is VTE risk assessed on admission using a computerized clinical decision support tool in the EHR, which has been in place for over a decade.^{12,27,28} Prescription of appropriate VTE prophylaxis has improved dramatically via numerous quality improvement interventions and is well over 90% in many patient populations.^{7,29,30} When the bedside nurse documented a nonadministered dose in the EHR, it automatically generated an alert in real-time to a pager and triggered the bundle intervention. Each unit could choose which team members would hold the pager (eg, nurse manager, charge nurse, nurse educator) and respond in a timely manner to the alert.

Nurse Feedback

The research team provided unit nurse managers with a monthly performance scorecard describing VTE prophylaxis administration practices for each individual nurse (number of doses prescribed, number of doses administered, and number of doses refused) (Figure S1). The unit nurse managers were exclusively responsible for conducting the nurse feedback intervention and used their preferred style. Although the research team recommended options to provide feedback (eg, 1-on-1 coaching, sharing data via email, posting blinded/unblinded results), all decisions and actions were left to the nurse manager for each participating unit. We did not collect information on approaches used. When performing our analysis, we used an intention-to-treat concept, aggregating missed doses for all patients in this cohort, regardless of what, if any, nurse feedback was provided.

Data Collection and Variables

Data on patient characteristics (age, sex, race), length of stay, and number of prescribed and administered prophylaxis doses were extracted directly from the EHR, achieving 100% automated data capture. The EHR system required documentation of all administered and nonadministered prescribed prophylaxis doses and the reason when not administered. We categorized reasons for nonadministration into patient refusal and other reasons, as documented by the nurse. We included patient visits (a hospitalization) with ≥ 1

dose of VTE prophylaxis prescribed. Baseline data included the period July 1, 2017 to December 31, 2017. We introduced the interventions to units in January 2018 and excluded these data from our analysis (washout period). We collected postintervention data from February 1, 2018 through April 30, 2018.

Outcomes

The primary outcome was the proportion of all prescribed pharmacologic VTE prophylaxis doses not administered. Secondary outcomes were the proportions of nonadministered doses from patient refusal or for other reasons.

Statistical Analysis

We performed a power calculation using historical data of prescribed VTE doses to determine whether we would have sufficient power for our planned analyses. Individual dose was the unit of analysis. Patient demographic and clinical characteristics for the baseline and postintervention periods are described in aggregate by intervention arm. The primary analysis was a pre- versus postintervention comparison to determine if both cohorts improved administration of prophylaxis, treating individual floors as their own historical control to account for differences by floor. We conducted 2-sample *t* tests with equal variance to compare mean (SD) for age, and χ^2 tests to compare proportions for sex and race. Nonparametric Wilcoxon rank sum tests were used to compare the number of prescribed doses per patient visit and length of stay; these were reported as median (interquartile range) and mean (SD). Our secondary analyses compared each intervention to determine which (if either) was more effective in reducing nonadministered doses, comparing overall reasons for missed doses, patient refusal, and other reasons not related with patient refusal.

For our primary and secondary hypotheses, mixed-effects logistic regression models and Poisson regression models with random intercepts for unit and nurse were used to account for the nurse-unit correlation when comparing VTE prophylaxis nonadministration by intervention arm and time. We included indicator variables for arm (nurse feedback versus patient bundle), time periods (pre- versus postintervention) and interaction term between arm and time periods as predictors. For the subgroup analysis by hospital unit, we included indicator variables for arm, pre- versus postintervention periods, and hospital unit (surgery versus medicine), as well as all the interaction terms among indicator variables for arm, time, and hospital unit as predictors. We used a multiple outputation approach to account for multiple VTE doses per patient across nurses and/or units.³¹ This approach randomly selects 1 VTE prophylaxis dose per patient and repeats the

procedure 1000 times to bootstrap the conditional odds ratio (OR) and proportions with corresponding 95% CI and *P* values for these comparisons, reducing hierarchical structure to the nurse-unit level.³¹ All comparisons were specified a priori and performed using a 0.05 α level for statistical significance. Statistical analyses were performed by a blinded team of biostatisticians using Stata version 14.1 MP–Parallel Edition (StataCorp, College Station, TX).

RESULTS

Of 12 958 patient visits prescribed pharmacologic VTE prophylaxis, 1860 were excluded (multiple floors *n*=495, doses in multiple periods *n*=1365), resulting in 11 098 visits from 9657 unique patients included in our analysis (Figure 1). Of 11 098 patient visits, 6158 were in the bundle arm and 4940 in the nurse feedback arm. A significant difference was observed in the racial distribution of patients between the bundle and nurse feedback arms in the preintervention period (White: 55.9% versus 49.3%, Black: 35.5% versus 42.3%, and other races: 8.6% versus 8.4%; *P*<0.001), but no significant differences were observed within unit type and between periods (Table 1). Median length of stay and median number of prescribed VTE doses per patient visit were significantly higher on nurse feedback units in the preintervention period. The proportion of nonadministered doses for all reasons for the patient education bundle was 12.2% (95% CI, 8.5–17.6) preintervention and 7.4% (95% CI, 5.1–10.8) postintervention, and for nurse feedback was 14.7% (95% CI, 10.3–21.2) preintervention and 11.2% (95% CI, 7.7–16.4) postintervention (Table 2).

VTE Prophylaxis Medication Administration

Overall, the odds of nonadministration of a pharmacologic VTE prophylaxis dose significantly decreased from the pre- to the postintervention period (OR, 0.64 [95% CI, 0.57–0.71]). Both the nurse feedback (OR, 0.72 [95% CI, 0.62–0.84]) and the bundle (OR, 0.56 [95% CI, 0.48–0.66]) interventions significantly decreased the odds of nonadministration after implementation. Our analysis of intervention effect for reducing nonadministration found the bundle to be more effective (exponentiated 2-way interaction term [ratio of the nurse feedback OR versus the bundle] 1.28 [95% CI, 1.02–1.61]; *P*=0.03; Table 2).

Nonadministered Doses Stratified by Reason

Patient refusal was the most common reason for dose nonadministration (Table 2). Refused doses for the

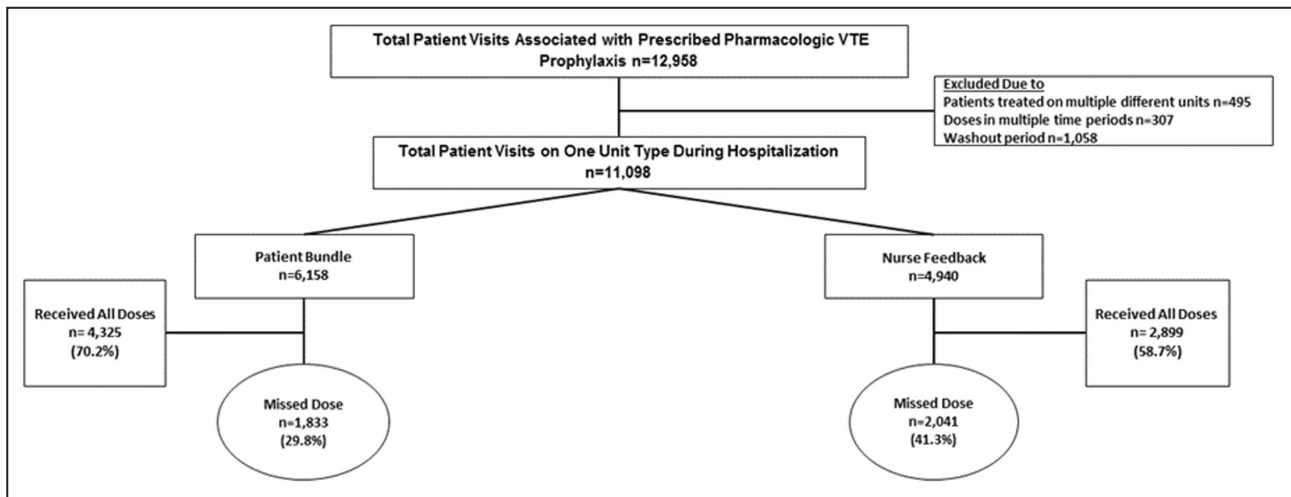


Figure 1. Consolidated Standards of Reporting Trials (CONSORT) flow diagram of hospital units receiving the patient-centered education bundle vs nurse feedback interventions.

Patient visits reflect patients with prescribed pharmacologic venous thromboembolism (VTE) prophylaxis doses during 1 hospital encounter. In the patient-centered education bundle arm, a nonadministered prophylaxis dose triggers an alert leading to the delivery of the patient-centered education bundle intervention. The patient-centered education bundle is only delivered once to patients. Nurse leaders in the nurse feedback arm received monthly scorecards detailing VTE prophylaxis administration practices by individual nurse.

patient education bundle were 7.3% (95% CI, 4.3–12.4) preintervention and 3.7% (95% CI, 2.1–6.4) postintervention, and for nurse feedback were 9.5% (95% CI, 5.6–16.1) preintervention and 7.1% (95% CI, 4.2–12.2) postintervention (Table 2). The overall odds of dose refusal significantly decreased after the intervention (OR, 0.59 [95% CI, 0.51–0.68]; $P < 0.001$). Both interventions significantly decreased the odds of dose refusal ($P < 0.001$). The bundle had a larger decrease in odds of refused doses compared with nurse feedback (OR, 0.46 versus OR, 0.71) and was significantly more effective (2-way interaction term [ratio of the nurse feedback OR versus the bundle OR] 1.53, [95% CI, 1.16–2.05]; $P = 0.003$; Table 2). There was a smaller, yet still statistically significant decrease in nonadministration for other reasons, and the bundle and nurse feedback were equally effective (OR, 0.80 and OR, 0.79), and the ratio between the 2 odds ratios is not statistically different from 1 (exponentiated 2-way interaction term 0.99 [95% CI, 0.72–1.35]; $P = 0.93$).

Stratified Analysis by Unit Type (Surgery Versus Medicine)

Overall, both unit types (surgery and medicine) had significantly lower odds of nonadministered doses for patient refusal and other reasons postintervention ($P < 0.001$; Table 3). On the medicine units, the bundle led to a statistically significant improvement in the odds of refusal compared with nurse feedback (OR 0.49 [95% CI, 0.38–0.64] versus OR, 0.79 [95% CI, 0.63–0.97]; $P = 0.008$). There were no significant differential effects for any other comparisons.

Time Trend Analysis

Figure 2 shows an overall graph of the conditional probability of nonadministered doses by month, including unit type strata for surgery (Figure 2B) and medicine (Figure 2C) units. These analyses demonstrate improvements in the postimplementation period. Similar improvements were observed for both unit types.

DISCUSSION

In this cluster-randomized controlled trial, we found significant reductions in nonadministered doses of pharmacologic VTE prophylaxis among hospitalized patients following implementation of both the patient-centered education bundle and the nurse feedback intervention. The trial implemented 2 unique strategies: a real-time EHR alert prompting delivery of patient-centered education materials, or a monthly scorecard reporting the dose administration percentages for unit nurses relative to their peers. Although both interventions improved administration practices, the bundle showed a superior effect, evident in a 44% reduction in the odds of nonadministration and a 54% reduction in patient refusal of pharmacologic prophylaxis. This finding is important, because we sought to determine if frontline nurses could implement the bundle and effectively reduce nonadministered doses as part of their routine work, even without the interventions being done by a research-funded nurse. These data replicate a similar, yet smaller, scale intervention at a community hospital.²⁴ When stratified by unit type, both

Table 1. Demographic and Clinical Characteristics for Pre- and Postintervention Visits by Patient Bundle and Nurse Feedback Arms

Characteristics	Patient education bundle		P value	Nurse feedback		P value	P value comparing pre patient bundle and pre nurse feedback
	Period			Period			
	Pre, n=4025	Post, n=2133		Pre, n=3342	Post, n=1598		
Unique patients	3497	1926		2811	1423		
Unique nurses	529	446		476	421		
Mean age, y (SD)*	56.1 (16.8)	56.1 (16.1)	0.97	56.1 (16.2)	56.7 (16.2)	0.17	0.99
Sex, n (%)†							
Men	2107 (52.3)	1126 (52.8)	0.74	1754 (52.5)	864 (54.1)	0.30	0.91
Women	1918 (47.7)	1007 (47.2)		1588 (47.5)	734 (45.9)		
Race, n (%)†							
Black	1428 (35.5)	755 (35.4)	0.79	1414 (42.3)	724 (45.3)	0.14	<0.001
White	2251 (55.9)	1212 (56.8)		1646 (49.3)	744 (46.6)		
Other	346 (8.6)	166 (7.8)		282 (8.4)	130 (8.1)		
Unit type, n (%)†							
Medicine units	1569 (39.0)	839 (39.3)	0.98	1849 (55.3)	882 (55.2)	0.93	<0.001
Surgery units	2456 (61.0)	1394 (60.7)		1493 (44.7)	716 (44.8)		
No. of prescribed doses per patient visit (Q1–Q3)							
Median (IQR)	6.0 (3.0–12.0)	6.0 (3.0–11.0)	0.45	7.0 (3.0–13.0)	6.0 (3.0–1.0)	0.001	<0.001
Mean (SD)‡	9.22 (11.3)	8.81 (10.1)		10.44 (12.4)	8.88 (9.7)		
Length of stay, d (Q1–Q3)							
Median (IQR)	4.0 (2.0–7.0)	4.0 (2.0–7.0)	0.68	5.0 (3.0–9.0)	5.0 (3.0–8.0)	0.64	<0.001
Mean (SD)‡	6.49 (9.6)	6.52 (8.94)		7.41 (8.4)	7.22 (7.9)		

Patient education bundle refers to the Patient-Centered Education Bundle. IQR indicates interquartile range; and Q, quartile.

*P values calculated using 2-sample *t* tests with equal variances.

†P values calculated using χ^2 tests.

‡P values calculated using Wilcoxon rank sum tests.

interventions showed a statistically significant improvement in the odds of both nonadministered and refused doses, confirming that the intervention was beneficial for both surgery and medicine patients, and the effect size was similar in this larger scale implementation, which grew the intervention from 4 to 16 floors at our hospital.

To our knowledge, this is the first study to implement multiple quality improvement interventions in a randomized fashion, targeting this key point of failure in VTE prevention, administration of VTE prophylaxis doses. In addition, we harnessed health information technology to automate the intervention and empowered nurses and nurse leaders to use evidence-based medicine in daily practice. The monthly scorecard in this trial was the only such clinical performance feedback provided to nurses on participating units during the study period. This trial expands on our previous evaluation of the impact of the bundle, in which we achieved significant improvements in nonadministered doses of VTE prophylaxis.^{23,24}

In the current trial, we demonstrated that the bundle delivered after an EHR real-time alert can be scaled for real-world use, independent of a dedicated research

team or extra financial resources. Our approach used an implementation science framework to translate evidence into routine practice.³² We developed and studied the bundle first and showed its effectiveness.^{23,25} Then, nursing staff took ownership and integrated it into routine, daily practice, and finally, we independently measured whether their efforts were making significant improvements in delivering prescribed prophylaxis doses. This approach has achieved sustained success in previous quality improvement research.^{33,34} Evidence from quality improvement research can contribute to effective and efficient patient-centered care.³⁵ Synthesis from such evidence has informed health care policy and formulated guidelines.³⁶ However, translating evidence for VTE prophylaxis into practice can be an unpredictable and slow process.^{32,37} Some areas of uncertainty may relate to structural and cultural influences or the health care professionals involved in crystallizing these findings into practice.¹⁴

Studies consistently show that audit and feedback changes behaviors and improves professional practice.^{30,38,39} Yet, there are different degrees to which feedback has affected sustained change.^{6,40} Shojania and colleagues reviewed a variety of quality

Table 2. Proportion of Doses Nonadministered for All Reasons, Refused, and Other Reasons: Comparisons Between Overall Pre- and Postintervention by Patient Bundle and Nurse Feedback Arms*†

Period	Overall	Patient education bundle	Nurse feedback	OR nurse feedback/patient education bundle (95% CI)	P value*
Any nonadministered dose					
Preintervention, % (95% CI)	13.4 (10.3–17.5)	12.2 (8.5–17.6)	14.7 (10.3–21.2)	1.24 (0.68–2.26)	0.49
Postintervention, % (95% CI)	9.2 (7.0–12.1)	7.4 (5.1–10.8)	11.2 (7.7–16.4)	1.59 (0.86–2.94)	0.14
OR, post/pre, (95% CI)	0.64 (0.57–0.71)	0.56 (0.48–0.66)	0.72 (0.62–0.84)		
P value*	<0.001	<0.001	<0.001		
Refused dose					
Preintervention, % (95% CI)	8.3 (5.6–12.2)	7.3 (4.3–12.4)	9.5 (5.6–16.1)	1.35 (0.59–3.06)	0.48
Postintervention, % (95% CI)	5.2 (3.5–7.8)	3.7 (2.1–6.4)	7.1 (4.2–12.2)	2.07 (0.89–4.83)	0.09
OR, post/pre (95% CI)	0.59 (0.51–0.68)	0.46 (0.37–0.58)	0.71 (0.59–0.86)		
P value*	<0.001	<0.001	<0.001		
Other reasons for nonadministered doses (not patient refused)					
Preintervention, % (95% CI)	3.6 (3.1–4.2)	3.4 (2.8–4.3)	3.8 (3.0–4.7)	1.09 (0.80–1.50)	0.57
Postintervention, % (95% CI)	2.9 (2.4–3.5)	2.8 (2.2–3.6)	3.0 (2.3–3.9)	1.08 (0.75–1.56)	0.68
OR, post/pre (95% CI)	0.79 (0.68–0.93)	0.80 (0.64–0.99)	0.79 (0.63–0.99)		
P value*	0.004	0.04	0.04		

Patient education bundle refers to the Patient-Centered Education Bundle. OR indicates odds ratio.

*P values for the ORs were calculated using multiple outputation of the mixed-effects logistic regression models.

†Exponentiated 2-way interactions were performed including pre vs post time period, and nurse feedback vs patient bundle: OR, 1.28 (95% CI, 1.02–1.61); $P=0.03$ for any nonadministered dose; OR, 1.53 (95% CI, 1.16–2.05); $P=0.003$ for patient refused dose; OR, 0.99 (95% CI, 0.72–1.35); $P=0.93$ for other reasons for nonadministered doses (excluding patient refused).

improvement strategies aimed at provider adherence to care practices for diabetes management and alluded to the greater chance of success when using multiple targeted strategies over 1 strategy alone.⁴¹ Strategies that address multiple layers of defect-free VTE prophylaxis have also been suggested.⁴² Our current study highlights the advantage of a multifaceted approach aimed at numerous critical steps to optimal VTE prevention, namely nurse administration and patient acceptance of prophylaxis. In addition, we implemented the interventions as part of an overarching strategy at our institution, which already has a computerized clinical decision support tool that stratifies patients by major risk categories and has improved prescription of risk-appropriate VTE prophylaxis.^{7,8,28,43} Our targeted alert approach is supported by previous research that recommends reducing unneeded electronic alert interventions to mitigate alert fatigue.^{44,45}

This dissemination study relied on nurses' endorsement and welcoming of the intervention into routine clinical practice. The easily adaptable nature of our intervention was an added benefit and required minimal deviation from nurses' workflow, which already includes dedicated time for patient education. The vast majority of nurses had already been exposed to targeted VTE prophylaxis knowledge through a prior study of required online VTE education module.²¹ The nurse education module improved dose administration, and therefore, our baseline may have been better than hospitals naive to any nurse education

intervention. The results of this study are predicated on first providing education to nurses about the importance of delivering all prescribed prophylaxis doses for VTE prevention, which is a critical first step before implementing the patient-centered education bundle or the nurse feedback intervention.

A key element highlighted by previous quality improvement efforts to optimize VTE prophylaxis administration is to target interventions to appropriate clinical groups.^{38,46–48} Although surgical attendings did not benefit from VTE prophylaxis prescription feedback,³⁸ surgical residents' significantly improved prophylaxis prescribing practices following performance feedback.³⁰ Although audit and feedback can be useful tools, implementation may be influenced by structural or institutional capacities and differential buy-in that can effect intervention success.^{46–48} In the current trial, our findings show that providing targeted feedback to nurses can be successful. Although feedback improved practice, it was not as effective as the bundle. Perhaps adding feedback in a multipronged approach alongside other interventions would show added benefit.

Our study has several limitations. First, our trial was in a single tertiary care center with robust health information technology and a modifiable EHR system, limiting the generalizability of our findings to other settings. However, delivery of the bundle can be adapted based on workflows in different settings. At hospitals without an EHR system, the approach potentially can

Table 3. Subgroup Analysis by Hospital Unit (Surgery Versus Medicine) on the Proportion of Nonadministered Venous Thromboembolism Prophylaxis Medication Doses by Patient Bundle and Nurse Feedback Arms^{a,†,‡}

Surgery		Medicine						
Period	Patient education bundle	Nurse feedback	OR nurse feedback/patient bundle (95% CI)	P value ^a	Patient education bundle	Nurse feedback	OR nurse feedback/patient education bundle (95% CI)	P value ^a
Any nonadministered dose								
Preintervention, % (95% CI)	6.3% (4.6%–8.6%)	10.0% (7.4%–13.7%)	1.68 (1.00–2.84)	0.05	18.6% (14.5%–23.7%)	18.9% (14.8%–24.0%)	1.03 (0.68–1.55)	0.89
Postintervention (95% CI)	3.3% (2.3%–4.7%)	6.7% (4.6%–9.5%)	2.11 (1.18–3.79)	0.01	12.2% (9.2%–16.1%)	15.2% (11.7%–19.9%)	1.31 (0.84–2.06)	0.24
OR post/pre (95% CI)	0.50 (0.39–0.65)	0.63 (0.49–0.81)			0.60 (0.49–0.74)	0.77 (0.63–0.93)		
P value ^a	<0.001	<0.001			<0.001	0.01		
Patient refused dose								
Preintervention (95% CI)	2.7% (1.8%–4.2%)	5.5% (3.6%–8.5%)	2.13 (1.07–4.22)	0.03	13.5% (9.7%–18.8%)	13.8% (10.0%–19.1%)	1.04 (0.62–1.75)	0.89
Postintervention (95% CI)	1.0% (0.6%–1.8%)	3.2% (1.9%–5.2%)	3.12 (1.39–6.99)	0.01	7.3% (5.0%–10.6%)	11.2% (7.9%–15.9%)	1.64 (0.93–2.89)	0.09
OR post/pre (95% CI)	0.37 (0.23–0.59)	0.55 (0.39–0.77)			0.49 (0.38–0.64)	0.79 (0.63–0.97)		
P value ^a	<0.001	<0.001			<0.001	0.02		
Other reason for nonadministered dose (excluding patient refused)								
Preintervention (95% CI)	3.0% (2.4%–3.9%)	3.5% (2.7%–4.7%)	1.18 (0.80–1.74)	0.41	3.8% (3.0%–4.9%)	4.0% (3.1%–5.1%)	1.04 (0.72–1.49)	0.84
Postintervention (95% CI)	1.9% (1.4%–2.6%)	2.8% (1.9%–4.2%)	1.50 (0.90–2.50)	0.12	4.0% (3.0%–5.4%)	3.2% (2.4%–4.2%)	0.77 (0.51–1.17)	0.22
OR post/pre (95% CI)	0.62 (0.46–0.83)	0.79 (0.55–1.13)			1.05 (0.78–1.43)	0.78 (0.59–1.05)		
P value ^a	0.002	0.19			0.74	0.10		

Patient education bundle refers to the Patient-Centered Education Bundle. OR indicates odds ratio.

^aP values for the odds ratios were calculated using multiple imputation of the mixed-effects logistic regression models.

[†]Exponentiated 2-way interactions were calculated using multiple imputation of the mixed-effects logistic regression models. [†]Exponentiated 2-way interactions were performed including pre vs post time period, and nurse feedback vs patient bundle units for surgery: OR, 1.26 (95% CI, 0.87–1.81); P=0.22 for any nonadministered dose; OR, 1.47 (95% CI, 0.83–2.60); P=0.19 for patient refused dose; OR, 1.27 (95% CI, 0.79–2.06); P=0.33 for other reasons for nonadministered doses (excluding patient refused). Medicine: OR, 1.28 (95% CI, 0.96–1.70); P=0.09 for any nonadministered dose; OR, 1.58 (95% CI, 1.13–2.20); P=0.01 for patient refused dose; OR, 0.74 (95% CI, 0.49–1.13); P=0.16 for other reasons for nonadministered doses (excluding patient refused).

[‡]Exponentiated 3-way interactions were performed to calculate the ratio of ratios between ORs that examine whether the differences in ORs are different, including pre vs post time period, and nurse feedback vs patient bundle units, and surgery vs medicine with no significant differences observed: OR, 0.94 for any nonadministered dose; OR, 0.80 for patient refused dose; OR, 0.09 for other reasons for nonadministered doses (excluding patient refused).

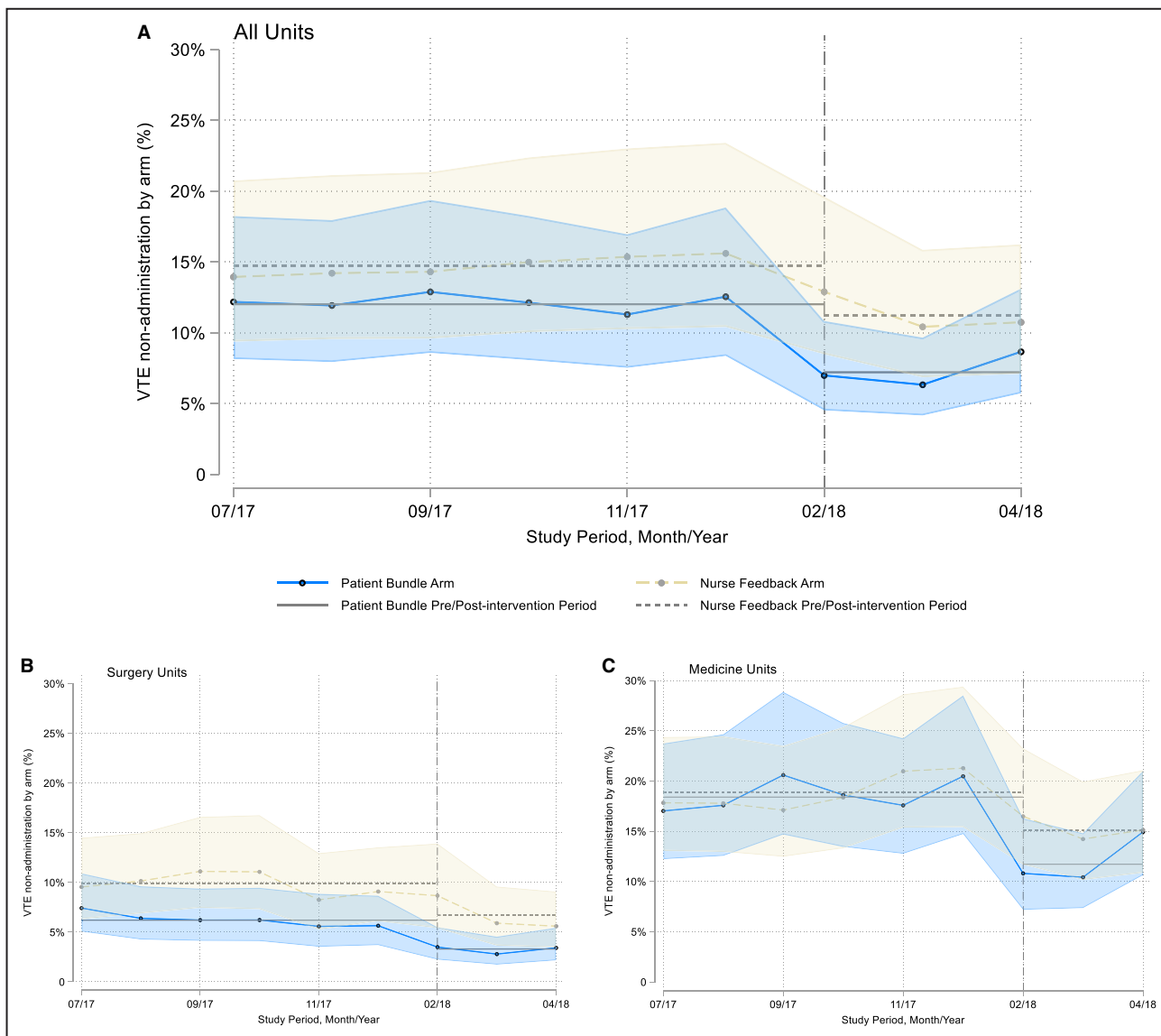


Figure 2. Time series analysis for all units (A) of the patient-centered education bundle and nurse feedback arms stratified by surgery (B) and medicine (C) hospital units.

This trend analysis reflects the monthly data for nonadministered doses of venous thromboembolism (VTE) prophylaxis in the preintervention (July 1, 2017 to December 31, 2017) and postintervention (February 1, 2018 to April 30, 2018) periods, comparing both intervention arms. Data for January 2018 were excluded (washout period). Our findings show that no changes are evident before the postimplementation period.

be implemented without the triggered alert if the bedside nurse uses the same educational bundle at the time of dose administration refusal. The materials are freely available for widespread use (bit.ly/bloodclots), and were created with patient stakeholder input.²³ Second, differences in patient demographics and length of stay between the arms in the preintervention period may have influenced our findings. We mitigated this through our randomization approach and analytic models, which provided a fair distribution of patients by arm and accounted for any residual bias. Moreover, each unit served as its own historical control

for the pre- and postintervention comparisons, also limiting bias in our analysis. Third, nurses and/or patients moving between units may have contaminated the data. However, at the time of this study, nurses at our hospital were predominantly assigned to specific units; thus, moving between units was likely not a problem. We also limited contamination by excluding patients who moved between units. Finally, we did not control intervention implementation, and consciously decided not to jeopardize buy-in by overburdening frontline nurses and nurse leaders with vast amounts of added data collection (ie, which patients got which

bundle elements, how exactly nurses received individual or group feedback). Although we would have liked to know exactly which patients were intervened upon and the frequency and methods that nurse leaders used to deliver feedback, these data are not available. A benefit of this intention to treat approach was the demonstrated real-world applicability for what originally began as funded clinical research.

CONCLUSIONS

Our study provides clear evidence that supports the use of information technology strategies combined with targeted patient-centered education to bolster best practices of VTE prophylaxis medication administration. Future research and quality improvement efforts should target strategies that leverage information technology solutions to scale and translate evidence into practice to improve a wider variety of clinical practices.

ARTICLE INFORMATION

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for the integrity of the data and the accuracy of the data analysis. J. Wang, and Dr Yenokyan from the Department of Biostatistics in the Johns Hopkins Bloomberg School of Public Health conducted the data analysis.

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Supplemental Material

Figure S1

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SUPPLEMENTAL MATERIAL

Figure S1. Sample of Monthly Scorecard Received by Nurse Leaders on Nurse Feedback Units

Rank	Unique Identifier	CURRENT MONTH				Previous Month
		Number of Doses Prescribed	Proportion Administered	Proportion missed	Proportion Refused	Proportion Administered
1	Nurse A	50	100%	0.0%	0.0%	100%
2	Nurse B	45	100%	0.0%	0.0%	100%
3	Nurse C	40	100%	0.0%	0.0%	100%
4	Nurse D	35	100%	0.0%	0.0%	94.0%
5	Nurse E	30	100%	0.0%	0.0%	91.0%
6	Nurse F	20	100%	0.0%	0.0%	93.0%
7	Nurse G	20	95.0%	0.0%	5.0%	92.0%
8	Nurse H	50	94.0%	2.0%	4.0%	81.0%
9	Nurse I	40	90.0%	5.0%	5.0%	80.0%
10	Nurse J	25	92.0%	0.0%	8.0%	82.0%
11	Nurse K	25	88.0%	8.0%	4.0%	80.0%
12	Nurse L	20	85.0%	0.0%	15.0%	79.0%
Overall Unit		400	95.3%	1.3%	3.4%	89.0%
Overall Department		3000	89.0%	3.7%	7.3%	88.0%

A monthly scorecard of venous thromboembolism (VTE) prophylaxis administration was given to the nurse leader on each unit participating in the nurse feedback arm. The scorecard reflected individual nurse and unit-level administration practices. Data included the number of doses prescribed and the proportions administered, missed, and refused. Nurses were ranked according to the percentage of doses not administered and performance color-coded based on reaching a 96% goal. We used the 96% goal because leaders of The Johns Hopkins Hospital and the health system established this as the common goal for quality improvement efforts aimed at externally reported core measures. Thus, it was a goal familiar to many frontline staff and unit leaders. Performance on administration is coded green when $\geq 96\%$, yellow when between 90.0%-95.9%, and red when $<90.0\%$. Nursing leaders provided peer coaching to nurses who performed below 96% administration, or in the yellow and red zones.