Evaluation of use of wells criteria and D-dimer for screening of venous thromboembolism in a community hospital

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Abstract

Background: Venous thromboembolism is a leading cause of morbidity and mortality in the United States. Accurate and timely diagnosis may be confounded by its nonspecific clinical presentation. Wells Criteria is a reliable screening tool with high sensitivity; however, recent data suggest that D-dimer is increasingly used as a primary screening test. While the sensitivity of D-dimer is comparable to Wells Criteria, it lacks specificity and positive predictive value, thus resulting in increased hospitalizations, further testing and accompanied costs and complications.

Aims: Our goal was to assess utilization of D-dimer and Wells criteria in the emergency department as a screening tool for venous thromboembolism.

Materials and methods: This is a retrospective chart review conducted at a community hospital. All patients who had a D-dimer test at presentation to the emergency department were included in the study. Fisher’s Exact was used for statistical analysis.

Results: Contrary to current recommendations, 15 patients (9.3%) with low Wells Score and negative D-dimer had further imaging studies. Increased use of imaging resulted in an increased cost of care and possible exposure to the procedure related complications.

Conclusions: A review of the literature and our study both conclude that adherence to current guidelines for evaluation of venous thromboembolism is less than optimal. Adherence to the guidelines in evaluating these patients would have several beneficial outcomes including reducing the need for further imaging studies thus reducing healthcare costs and decreasing possible patient complications associated with such procedures.

Introduction

Venous thromboembolism (VTE) is a major cause of morbidity and mortality in United States [1]. Its prevalence is one patient per thousand people per year and out of 100,000 hospital admissions, 239 are from VTE [2-4]. Current recommendations, based on cumulative data, suggest using a two-step approach of utilizing Wells Criteria (Figure 1) for its high sensitivity and D-dimer for its high negative predictive value to triage patients quickly and effectively in the emergency department [5,6]. However, D-dimer use along with pulmonary computed tomographic angiography has increased for varying reasons [7-9]. D-dimer has very low positive predictive value and can be elevated in many other conditions, thus is not specific to VTE [7-12]. The purpose of this retrospective study was to evaluate the use of guidelines for diagnosis of VTE in a community based emergency department.

Methods

This is an institutional review board approved retrospective study. All patients who presented to our emergency department during January through June 2010 were considered for inclusion in the study. Patient charts were reviewed by a team of physicians for D-dimer level, documentation for Wells Score, lower extremity venous ultrasound, computed tomography chest pulmonary embolism protocol, ventilation quotient scan and demographic information. If there was no Wells Score documentation in the chart, a score was calculated using available information in the medical record. Zero was assigned for any missing data needed to calculate Wells Score. Statistical analysis was done using Fisher’s Exact Test.

Results

During the study period 10,651 patients presented to our emergency department, 346 (3.2%) had symptoms suggestive of VTE and were screened using D-dimer testing. Sixteen of the 346 screened (4.6%) had documented thromboembolic events. Average age of the patients was 55.4 years (range of 18-96), 63% of them being females. Table 1 shows detailed demographics of the study population. The average annual rate of venous thromboembolism was 0.23% in hospitalized patients. Wells Score was not documented in any of the patients screened for VTE.

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A significantly higher percentage of patients with a D-dimer result ≥0.50 mg/dl had a confirmed venous thromboembolism compared to patients with a D-dimer result < 0.50 mg/dl (9% vs. 1%, p-value=0.0004) (Table 2). Patients with a Wells Score that indicated a high or moderate probability of VTE had a much higher percentage of confirmed VTE than patients with a Wells score that indicated a low probability (20% vs. 18% vs. 2.6%, p=<0.0001) (Table 2). A vast majority, 87.5% (n=303), of the study population had low probability Wells Scores. Among them, 162 had negative D-dimers. Of the 162 patients, 15 (9.3%) had further imaging done (Figure 2).

### Table 1. Demographics of study population.

<table>
<thead>
<tr>
<th>Characteristics</th>
<th>Total number of patients (N)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age (years)</td>
<td>Range 18-96, average 55.4</td>
</tr>
<tr>
<td>Sex</td>
<td>Male · 37%, Female · 63%</td>
</tr>
<tr>
<td>D-dimer</td>
<td></td>
</tr>
<tr>
<td>&lt;0.51 %</td>
<td>170 (49%)</td>
</tr>
<tr>
<td>&gt;0.51 %</td>
<td>176 (51%)</td>
</tr>
<tr>
<td>Wells score</td>
<td></td>
</tr>
<tr>
<td>Low probability</td>
<td>303 (88%)</td>
</tr>
<tr>
<td>Medium probability</td>
<td>38 (11%)</td>
</tr>
<tr>
<td>High probability</td>
<td>5 (1%)</td>
</tr>
<tr>
<td>Venous thromboembolic events</td>
<td>16 (4.6%)</td>
</tr>
<tr>
<td>Deep venous thromboembolism (DVT)</td>
<td>3 (0.87%)</td>
</tr>
<tr>
<td>Pulmonary embolism (PE)</td>
<td>6 (1.7%)</td>
</tr>
<tr>
<td>DVT and PE</td>
<td>7 (2.0%)</td>
</tr>
</tbody>
</table>

### Table 2. Analysis of data based on D-dimer and Wells probability score.

<table>
<thead>
<tr>
<th>D-dimer</th>
<th>With VTE</th>
<th>Without VTE</th>
<th>p-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>&lt; 0.51</td>
<td>1 (1%)</td>
<td>169 (99%)</td>
<td>0.0004</td>
</tr>
<tr>
<td>&gt; 0.51</td>
<td>15 (9%)</td>
<td>161 (91%)</td>
<td></td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Wells Criteria</th>
<th>With VTE</th>
<th>Without VTE</th>
<th>p-value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Low probability</td>
<td>8 (2.6%)</td>
<td>295 (97.4%)</td>
<td>&lt;0.0001</td>
</tr>
<tr>
<td>Medium probability</td>
<td>7 (18%)</td>
<td>31 (82%)</td>
<td></td>
</tr>
<tr>
<td>High probability</td>
<td>1 (20%)</td>
<td>4 (80%)</td>
<td></td>
</tr>
</tbody>
</table>

### Discussion

VTE is a common condition (1:1,000 in general population) [2]. Pulmonary embolism is a common and potentially lethal disease accounting for more than 250,000 hospitalizations a year in the United States [1]. The incidence of deep venous thrombosis and pulmonary embolism is 20% or less among screened patients [13,14]. A missed diagnosis of VTE can lead to sudden death, chronic cardiopulmonary dysfunction, and impaired quality of life [4,15,16]. However, over testing increases costs and patients are exposed to undue risk with no added benefit. Diagnosing accurately avoids anticoagulation and its associated risk of bleeding in patients without disease [17].

There are multiple clinical prediction rules (in Table 3) available to predict likelihood of VTE [5, 18-20]. Patients are divided into high, moderate or low probability risk categories [4]. If likelihood is high or moderate, patients are scanned with computed tomography chest or lower extremity venous ultrasound directly. However, patients in the low risk group need further testing with D-dimer (Figure 1) to exclude the possibility of thromboembolic disease [6].

The most commonly used clinical prediction rule is Wells Criteria. Prevalence of deep venous thrombosis based on pretest probability...
of low, intermediate and high (as estimated by Wells Criteria) is 5%, 15% and 70% respectively [18]. For suspected pulmonary embolism, Wells Score and Revised Geneva Score are also widely accepted as standard measurements of pretest probability [20,21]. Revised Geneva Score is entirely based on clinical variables and thus independent from physician judgment [20]. In our study, the prevalence of venous thromboembolism in patients with low, intermediate and high clinical probability groups, using retrospectively calculated Wells Criteria, was 3%, 18% and 20% respectively, which is different from historical data [18]. Retrospective calculation of the Wells Score most likely underestimated the score, which would explain this difference.

D-dimer testing is done for the exclusion of VTE in low risk patients. The sensitivity of D-dimer in low to medium probability groups (as per Wells Criteria) varies from 94-96% (95 % CI 0.88-1.00) in multiple meta-analyses [10,11]. D-dimer levels below 0.5 mg/dl in low and intermediate clinical probability have a high negative predictive value, essentially ruling out VTE and no further studies are needed [12,22-24]. In our study only one patient (~0.5%) had confirmed VTE with D-dimer less than 0.50 mg/dl and 99% of patients (n=169) with D-dimer <0.50 had no VTE upon further investigations, in line with the previous data.

Testing D-dimer in patients with high clinical probability should not alter further diagnostic pathways [18]. In our study each of the five patients with high probability had D-dimer above 0.50 mg/dl. The post-test probability of pulmonary embolism remains above 3% despite a normal D-dimer, rendering the test futile in this subgroup [13]. D-dimer level above 2.0 mg/dl was predictive of the presence of pulmonary embolism, independently of the clinical score, with an odds ratio of 6.9 [25].

The fact that none of our patients had documented Wells Scores is not unusual. In a recent survey on clinical prediction rules, 68% of respondents (physicians) reported being familiar with at least one clinical prediction rule for pulmonary embolism [26]. Surveyed physicians also identified reasons for not using the prediction rules [26]. The clinical prediction rules for VTE are grossly underused or under documented and thus diagnostic decisions made were in discordance with the established recommendations [27-29].

In our study fifteen patients with negative D-dimer and low probability Wells Score had further work up done. This is not in line with established guidelines. This resulted in additional imaging studies and associated cost to the healthcare. Imaging cost on average was $1,289 per image based on reported Centers for Medicare and Medicaid Services charges (lower extremity venous doppler: $727, computed tomography chest with contrast: $1,586).

Limitations: Our single center study is limited by its retrospective nature and calculation of Wells Criteria based on chart review, which could have underestimated the actual Wells Score.
Conclusions

Every effort should be made to apply the clinical probability rules for VTE in emergency department patients before testing for D-dimer. This will result in substantial cost savings to the healthcare system. Furthermore, emergency departments will benefit from establishing guidelines and monitoring the use of clinical prediction rules for consistency, similar to other commonly utilized protocols including acute coronary syndrome and stroke.

References