## ORIGINAL RESEARCH

## Implementation of a Mortality Prediction Rule for Real-Time Decision Making: Feasibility and Validity

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**BACKGROUND:** A previously published, retrospectively derived prediction rule for death within 30 days of hospital admission has the potential to launch parallel interdisciplinary team activities. Whether or not patient care improves will depend on the validity of prospectively generated predictions, and the feasibility of generating them on demand for a critical proportion of inpatients.

**OBJECTIVE:** To determine the feasibility of generating mortality predictions on admission and to validate their accuracy using the scoring weights of the retrospective rule.

DESIGN: Prospective, sequential cohort.

SETTING: Large, tertiary care, community hospital in the Midwestern United States

**PATIENTS:** Adult patients admitted from the emergency department or scheduled for elective surgery

**RESULTS:** Mortality predictions were generated on demand at the beginning of the hospitalization for 9312

The systematic deployment of prediction rules within health systems remains a challenge, although such decision aids have been available for decades.<sup>1,2</sup> We previously developed and validated a prediction rule for 30-day mortality in a retrospective cohort, noting that the mortality risk is associated with a number of other clinical events.<sup>3</sup> These relationships suggest risk strata, defined by the predicted probability of 30-day mortality, and could trigger a number of coordinated care processes proportional to the level of risk.<sup>4</sup> For example, patients within the higher-risk strata could be considered for placement into an intermediate or intensive care unit (ICU), be monitored more closely by physician and nurse team members for clinical deterioration, be seen by a physician within a few days of hospital discharge, and be considered for

Received: April 30, 2014; Revised: July 22, 2014; Accepted: July 27, 2014

2014 Society of Hospital Medicine DOI 10.1002/jhm.2250 Published online in Wiley Online Library (Wileyonlinelibrary.com). (92.9%) out of a possible 10,027 cases. The area under the receiver operating curve for 30-day mortality was 0.850 (95% confidence interval: 0.833-0.866), indicating very good to excellent discrimination. The prospectively generated 30-day mortality risk had a strong association with the receipt of palliative care by hospital discharge, in-hospital mortality, and 180-day mortality, a fair association with the risk for 30-day readmissions and unplanned transfers to intensive care, and weak associations with receipt of intensive unit care ever within the hospitalization or the development of a new diagnosis that was not present on admission (ie, complication).

**CONCLUSIONS:** Important prognostic information is feasible to obtain in a real-time, single-assessment process for a sizeable proportion of hospitalized patients. *Journal of Hospital Medicine* 2014;9:720–726. © 2014 Society of Hospital Medicine

advance care planning discussions.<sup>3–7</sup> Patients within the lower-risk strata might not need the same intensity of these processes routinely unless some other indication were present.

However attractive this conceptual framework may be, its realization is dependent on the willingness of clinical staff to generate predictions consistently on a substantial portion of the patient population, and on the accuracy of the predictions when the risk factors are determined with some level of uncertainty at the beginning of the hospitalization.<sup>2,8</sup> Skepticism is justified, because the work involved in completing the prediction rule might be incompatible with existing workflow. A patient might not be scored if the emergency physician lacks time or if technical issues arise with the information system and computation process.<sup>9</sup> There is also a generic concern that the predictions will prove to be less accurate outside of the original study population.<sup>8-10</sup> A more specific concern for our rule is how well "present on admission" diagnoses can be determined during the relatively short emergency department or presurgery evaluation period. For example, a final diagnosis of heart failure might not be established until later in the hospitalization, after the results of diagnostic testing and clinical response to treatment are known. Moreover, our

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Additional Supporting Information may be found in the online version of this article.

retrospective prediction rule requires an assessment of the presence or absence of sepsis and respiratory failure. These diagnoses appear to be susceptible to secular trends in medical record coding practices, suggesting the rule's accuracy might not be stable over time.<sup>11</sup>

We report the feasibility of having emergency physicians and the surgical preparation center team generate mortality predictions before an inpatient bed is assigned. We evaluate and report the accuracy of these prospective predictions.

## METHODS

The study population consisted of all patients 18 years of age or less than 100 years who were admitted from the emergency department or assigned an inpatient bed following elective surgery at a tertiary, community teaching hospital in the Midwestern United States from September 1, 2012 through February 15, 2013. Although patients entering the hospital from these 2 pathways would be expected to have different levels of mortality risk, we used the original prediction rule for both because such distinctions were not made in its derivation and validation. Patients were not considered if they were admitted for childbirth or other obstetrical reasons, admitted directly from physician offices, the cardiac catheterization laboratory, hemodialysis unit, or from another hospital. The site institutional review board approved this study.

The implementation process began with presentations to the administrative and medical staff leadership on the accuracy of the retrospectively generated mortality predictions and risk of other adverse events.<sup>3</sup> The chief medical and nursing officers became project champions, secured internal funding for the technical components, and arranged to have 2 project comanagers available. A multidisciplinary task force endorsed the implementation details at biweekly meetings throughout the planning year. The leadership of the emergency department and surgical preparation center committed their colleagues to generate the predictions. The support of the emergency leadership was contingent on the completion of the entire prediction generating process in a very short time ("within the time a physician could hold his/her breath"). The chief medical officer, with the support of the leadership of the hospitalists and emergency physicians, made the administrative decision that a prediction must be generated prior to the assignment of a hospital room.

During the consensus-building phase, a Web-based application was developed to generate the predictions. Emergency physicians and surgical preparation staff were trained on the definitions of the risk factors (see Supporting Information, Appendix, in the online version of this article) and how to use the Web application. Three supporting databases were created. Each midnight, a past medical history database was updated, identifying those who had been discharged from the study hospital in the previous 365 days, and whether or not their diagnoses included atrial fibrillation, leukemia/lymphoma, metastatic cancer, cancer other than leukemia, lymphoma, cognitive disorder, or other neurological conditions (eg, Parkinson's, multiple sclerosis, epilepsy, coma, and stupor). Similarly, a clinical laboratory results database was created and updated real time through an HL7 (Health Level Seven, a standard data exchange format<sup>12</sup>) interface with the laboratory information system for the following tests performed in the preceding 30 days at a hospital-affiliated facility: hemoglobin, platelet count, white blood count, serum troponin, blood urea nitrogen, serum albumin, serum lactate, arterial pH, arterial partial pressure of oxygen values. The third database, admission-discharge-transfer, was created and updated every 15 minutes to identify patients currently in the emergency room or scheduled for surgery. When a patient registration event was added to this database, the Web application created a record, retrieved all relevant data, and displayed the patient name for scoring. When the decision for hospitalization was made, the clinician selected the patient's name and reviewed the pre-populated medical diagnoses of interest, which could be overwritten based on his/her own assessment (Figure 1A,B). The clinician then indicated ("yes," "no," or "unknown") if the patient currently had or was being treated for each of the following: injury, heart failure, sepsis, respiratory failure, and whether or not the admitting service would be medicine (ie, nonsurgical, nonobstetrical). We considered "unknown" status to indicate the patient did not have the condition. When laboratory values were not available, a normal value was imputed using a previously developed algorithm.<sup>3</sup> Two additional questions, not used in the current prediction process, were answered to provide data for a future analysis: 1 concerning the change in the patient's condition while in the emergency department and the other concerning the presence of abnormal vital signs. The probability of 30-day mortality was calculated via the Web application using the risk information supplied and the scoring weights (ie, parameter estimates) provided in the Appendices of our original publication.<sup>3</sup> Predictions were updated every minute as new laboratory values became available, and flagged with an alert if a more severe score resulted.

For the analyses of this study, the last prospective prediction viewed by emergency department personnel, a hospital bed manager, or surgical suite staff prior to arrival on the nursing unit is the one referenced as "prospective." Once the patient had been discharged from the hospital, we generated a second mortality prediction based on previously published parameter estimates applied to risk factor data ascertained retrospectively as was done in the original

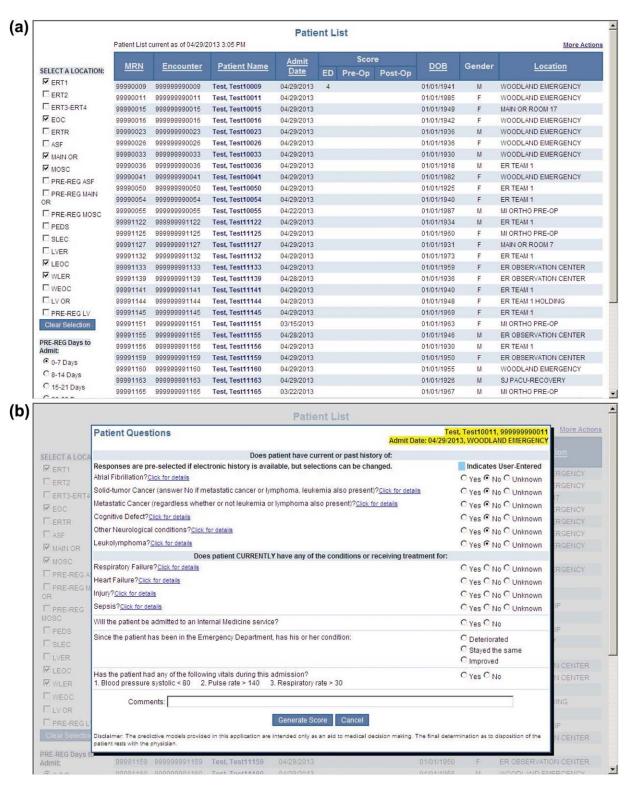


FIG. 1. Screen shots of the Web application used to generate predictions (A) Patient list. The clinician in the emergency department or surgical preparation center selects the patient to be scored. (B) Diagnosis-based risk factors to be entered. The clinician provides an answer to each question and/or reviews information that has been prepopulated from the past medical history database. Clinical laboratory values and demographic information are electronically provided. After the diagnosis information has been supplied, the clinician presses the "Generate Score" button to obtain the predicted 30-day mortality.

article<sup>3</sup>; we subsequently refer to this prediction as "retrospective." We will report on the group of patients who had both prospective and retrospective scores (1 patient had a prospective but not retrospective score available).

The prediction scores were made available to the clinical teams gradually during the study period. All scores were viewable by the midpoint of the study for emergency department admissions and near the end of the study for elective-surgery patients. Only 2 changes in care processes based on level of risk were introduced during the study period. The first required initial placement of patients having a probability of dying of 0.3 or greater into an intensive or intermediate care unit unless the patient or family requested a less aggressive approach. The second occurred in the final 2 months of the study when a large multispecialty practice began routinely arranging for high-risk patients to be seen within 3 or 7 days of hospital discharge.

#### **Statistical Analyses**

SAS version 9.3 (SAS Institute Inc., Cary, NC) was used to build the datasets and perform the analyses. Feasibility was evaluated by the number of patients who were candidates for prospective scoring with a score available at the time of admission. The validity was assessed with the primary outcome of death within 30 days from the date of hospital admission, as determined from hospital administrative data and the Social Security Death Index. The primary statistical metric is the area under the receiver operating characteristic curve (AROC) and the corresponding 95% Wald confidence limits. We needed some context for understanding the performance of the prospective predictions, assuming the accuracy could deteriorate due to the instability of the prediction rule over time and/ or due to imperfect clinical information at the time the risk factors were determined. Accordingly, we also calculated an AROC based on retrospectively derived covariates (but using the same set of parameter estimates) as done in our original publication so we could gauge the stability of the original prediction rule. However, the motivation was not to determine whether retrospective versus prospective predictions were more accurate, given that only prospective predictions are useful in the context of developing realtime care processes. Rather, we wanted to know if the prospective predictions would be sufficiently accurate for use in clinical practice. A priori, we assumed the prospective predictions should have an AROC of approximately 0.80. Therefore, a target sample size of 8660 hospitalizations was determined to be adequate to assess validity, assuming a 30-day mortality rate of 5%, a desired lower 95% confidence boundary for the area under the prospective curve at or above 0.80, with a total confidence interval width of 0.07.13 Calibration was assessed by comparing the actual proportion of patients dying (with 95% binomial confidence intervals) with the mean predicted mortality level within 5 percentile increments of predicted risk.

### **Risk Strata**

We categorize the probability of 30-day mortality into strata, with the understanding that the thresholds for defining these are a work in progress. Our hospital currently has 5 strata ranging from level 1 (highest mortality risk) to level 5 (lowest risk). The corresponding thresholds (at probabilities of death of 0.005, 0.02, 0.07, 0.20) were determined by visual inspection of the event rates and slope of curves displayed in Figure 1 of the original publication.<sup>3</sup>

# Relationship to Secondary Clinical Outcomes of Interest

The choice of clinical care processes triggered per level of risk may be informed by understanding the frequency of events that increase with the mortality risk. We therefore examined the AROC from logistic regression models for the following outcomes using the prospectively generated probability as an explanatory variable: unplanned transfer to an ICU within the first 24 hours for patients not admitted to an ICU initially, ICU use at some point during the hospitalization, the development of a condition not present on admission ("complication"), receipt of palliative care by the end of the hospitalization, death during the hospitalization, 30-day readmission, and death within 180 days. The definition of these outcomes and statistical approach used has been previously reported.<sup>3</sup>

## RESULTS

Mortality predictions were generated on demand for 7291 out of 7777 (93.8%) eligible patients admitted from the emergency department, and for 2021 out of 2250 (89.8%) eligible elective surgical cases, for a total of 9312 predictions generated out of a possible 10,027 hospitalizations (92.9%). Table 1 displays the characteristics of the study population. The mean age was 65.2 years and 53.8% were women. The most common risk factors were atrial fibrillation (16.4%) and cancer (14.6%). Orders for a comfort care approach (rather than curative) were entered within 4 hours of admission for 32/9312 patients (0.34%), and 9/9312 (0.1%) were hospice patients on admission.

## Evaluation of Prediction Accuracy

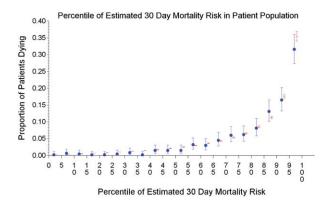
The AROC for 30-day mortality was 0.850 (95% confidence interval [CI]: 0.833-0.866) for prospectively collected covariates, and 0.870 (95% CI: 0.855-0.885) for retrospectively determined risk factors. These AROCs are not substantively different from each other, demonstrating comparable prediction performance. Calibration was excellent, as indicated in Figure 2, in which the predicted level of risk lay within the 95% confidence limits of the actual 30-day mortality for 19 out of 20 intervals of 5 percentile increments.

# Relationship to Secondary Clinical Outcomes of Interest

The relationship between the prospectively generated probability of dying within 30 days and other events is quantified by the AROC displayed in Table 2. The 30-day mortality risk has a strong association with the receipt of palliative care by hospital discharge, inhospital mortality, and 180-day mortality, a fair

Risk Factors	No.	Without Imputation	No.	With Imputation
Clinical laboratory values within preceding 30 days				
Maximum serum blood urea nitrogen (mg/dL)	8,484	22.7 (17.7)	9,312	22.3 (16.9)
Minimum hemoglobin, g/dL,	8,750	12.5 (2.4)	9,312	12.4 (2.4)
Minimum platelet count, 1,000/UL	8,737	224.1 (87.4)	9,312	225.2 (84.7)
Maximum white blood count, 1,000/UL	8,750	10.3 (5.8)	9,312	10.3 (5.6)
Maximum serum lactate, mEq/L	1,749	2.2 (1.8)	9,312	0.7 (1.1)
Minimum serum albumin, g/dL	4,057	3.4 (0.7)	9,312	3.2 (0.5)
Minimum arterial pH	509	7.36 (0.10)	9,312	7.36 (0.02)
Minimum arterial pO <sub>2</sub> , mm Hg	509	73.6 (25.2)	9,312	98.6 (8.4)
Maximum serum troponin, ng/mL	3,217	0.5 (9.3)	9,312	0.2 (5.4)
Demographics and diagnoses			,	( )
Age, y			9,312	65.2 (17.0)
Female sex			9,312	5,006 (53.8%)
Previous hospitalization within past 365 days			9,312	3,995 (42.9%)
Emergent admission			9,312	7,288 (78.3%)
Admitted to a medicine service	9,312	5,840 (62.7%)		
Current or past atrial fibrillation	9,312	1,526 (16.4%)		
Current or past cancer without metastases, excluding leukemia or ly	9,312	1,356 (14.6%)		
Current or past history of leukemia or lymphoma	9,312	145 (1.6%)		
Current or past metastatic cancer	9,312	363 (3.9%)		
Current or past cognitive deficiency	9,312	844 (9.1%)		
Current or past history of other neurological conditions (eg, Parkinso	9,312	952 (10.2%)		
Injury such as fractures or trauma at the time of admission	9,312	656 (7.0%)		
Sepsis at the time of admission			9,312	406 (4.4%)
Heart failure at the time of admission	9,312	776 (8.3%)		
Respiratory failure on admission	9,312	557 (6.0%)		
Outcomes of interest			,	· · /
Unplanned transfer to an ICU (for those not admitted to an ICU) within	8,377	86 (1.0%)		
Ever in an ICU during the hospitalization	9,312	1,267 (13.6%)		
Development of a condition not present on admission (complication)	9,312	834 (9.0%)		
Within hospital mortality	9,312	188 (2.0%)		
Mortality within 30 days of admission			9,312	466 (5.0%)
Mortality within 180 days of admission			9,312	1,070 (11.5%)
Receipt of palliative care by the end of the hospitalization			9,312	314 (3.4%)
Readmitted to the hospital within 30 days of discharge (patients alive	9,124	1,302 (14.3%)		
Readmitted to the hospital within 30 days of discharge (patients alive	9,312	1,302 (14.0%)		

NOTE: Data are presented as mean (standard deviation) or number (%). Abbreviations: ICU, intensive care unit.



**FIG. 2.** Calibration of the prediction rule. The horizontal axis displays intervals of 5 percentile increments of the predicted risk of dying within 30 days of admission (prospectively collected covariates). The vertical axis indicates the proportion of patients who actually died. The red dash marks represent the mean predicted mortality risk (and corresponding 95% confidence limits) for patients within the interval. The blue solid dot represents the actual proportion of patients within the interval who died, with the blue vertical hash marks indicating the 95% confidence limits for the proportion dying.

association with the risk for 30-day readmissions and unplanned transfers to intensive care, and weak associations with receipt of intensive unit care ever within the hospitalization or the development of a new diagnosis that was not present on admission (complication). The frequency of these events per mortality risk strata is shown in Table 3. The level 1 stratum contains a higher frequency of these events, whereas the level 5 stratum contains relatively few, reflecting the Pareto principle by which a relatively small proportion of patients contribute a disproportionate frequency of the events of interest.

#### DISCUSSION

Emergency physicians and surgical preparation center nurses generated predictions by the time of hospital admission for over 90% of the target population during usual workflow, without the addition of staff or resources. The discrimination of the prospectively

<b>TABLE 2.</b> Area Under the Receiver Operating
Characteristic Curve Secondary Outcomes
of Interest Associated With 30-Day Mortality Risk

In-hospital mortality	0.841 (0.814-0.869)
180–day mortality	0.836 (0.825-0.848)
Receipt of palliative care by discharge	0.875 (0.858-0.891)
30-day readmission (patients alive at discharge)	0.649 (0.634-0.664)
Unplanned transfer to an ICU (for those not	0.643 (0.590-0.696)
admitted to an ICU) within 24 hours	
Ever in an ICU during the hospitalization	0.605 (0.588-0.621)
Development of a condition not present	0.555 (0.535-0.575)
on admission (complication)	

NOTE: Data are presented as Mann–Whitney (95% Wald confidence limits) using the calculated probability of dying within 30 days and its logarithm as the explanatory variable. Abbreviations: ICU, intensive care unit.

generated predictions was very good to excellent, with an AROC of 0.850 (95% CI: 0.833-0.866), similar to that obtained from the retrospective version. Calibration was excellent. The prospectively calculated mortality risk was associated with a number of other events. As shown in Table 3, the differing frequency of events within the risk strata support the development of differing intensities of multidisciplinary strategies according to the level of risk.<sup>5</sup> Our study provides useful experience for others who anticipate generating real-time predictions. We consider the key reasons for success to be the considerable time spent achieving consensus, the technical development of the Web application, the brief clinician time required for the scoring process, the leadership of the chief medical and nursing officers, and the requirement that a prediction be generated before assignment of a hospital room.

Our study has a number of limitations, some of which were noted in our original publication, and although still relevant, will not be repeated here for space considerations. This is a single-site study that used a prediction rule developed by the same site, albeit on a patient population 4 to 5 years earlier. It is not known how well the specific rule might perform in other hospital populations; any such use should therefore be accompanied by independent validation studies prior to implementation. Our successful experience should motivate future validation studies. Second, because the prognoses of patients scored from the emergency department are likely to be worse than those of elective surgery patients, our rule should be recalibrated for each subgroup separately. We plan to do this in the near future, as well as consider additional risk factors. Third, the other events of interest might be predicted more accurately if rules specifically developed for each were deployed. The mortality risk by itself is unlikely to be a sufficiently accurate predictor, particularly for complications and intensive care use, for reasons outlined in our original publication.<sup>3</sup> However, the varying levels of events within the higher versus lower strata should be noted by the clinical team as they design their team-based processes. A follow-up visit with a physician within a few days of discharge could address the concurrent risk of dying as well as readmission, for example. Finally, it is too early to determine if the availability of mortality predictions from this rule will benepatients.<sup>2,8,10</sup> During the study period, we fit implemented only 2 new care processes based on the level of risk. This lack of interventions allowed us to evaluate the prediction accuracy with minimal additional confounding, but at the expense of not yet

Risk Strata	30-Day Mortality, Count/Cases (%)	Unplanned Transfers to ICU Within 24 Hours, Count/Cases (%)	Diagnosis Not Present on Admission, Complication, Count/Cases (%)	Palliative Status at Discharge, Count/Cases (%)	Death in Hospital Count/Cases (%)
1	155/501 (30.9%)	6/358 (1.7%)	58/501 (11.6%)	110/501 (22.0%)	72/501 (14.4%)
2	166/1,316 (12.6%)	22/1,166 (1.9%)	148/1,316 (11.3%)	121/1,316 (9.2%)	58/1,316 (4.4%)
3	117/2,977 (3.9%)	35/2,701 (1.3%)	271/2,977 (9.1%)	75/2,977 (2.5%)	43/2,977 (1.4%)
4	24/3,350 (0.7%)	20/3,042 (0.7%)	293/3,350 (8.8%)	6/3,350 (0.2%)	13/3,350 (0.4%)
5	4/1,168 (0.3%)	3/1,110 (0.3%)	64/1,168 (5.5%)	2/1,168 (0.2%)	2/1,168 (0.2%)
Total	466/9,312 (5.0%)	86/8,377 (1.0%)	834/9,312 (9.0%)	314/9,312 (3.4%)	188/9,312 (2.0%)
				Death or Readmission	
	Ever in ICU,	30-Day F	Readmission,	Within 30 Days,	180-Day Mortality
Risk Strata	Count/Cases (%)	Count/Cases (%)		Count/Cases (%)	Count/Cases (%)
1	165/501 (32.9%)	106/429 (24.7%)		243/501 (48.5%)	240/501 (47.9%)
2	213/1,316 (16.2%)	275/1,258 (21.9%)		418/1,316 (31.8%)	403/1,316 (30.6%)
3	412/2,977 (13.8%)	521/2,934 (17.8%)		612/2,977 (20.6%)	344/2,977 (11.6%)
4	406/3,350 (12.1%)	348/3,337 (10.4%)		368/3,350 (11.0%)	77/3,350 (2.3%)
5	71/1,168 (6.1%)	52/1,166 (4.5%)		56/1,168 (4.8%)	6/1,168 (0.5%)
Total	1,267/9,312 (13.6%)	1,302/9,124 (14.3%)		1,697/9,312 (18.2%)	1,070/9,312 (11.5%)

NOTE: Abbreviations: ICU, intensive care unit.

#### Cowen et al | Generating Mortality Predictions

knowing the clinical impact of this work. After the study period, we implemented a number of other interventions and plan on evaluating their effectiveness in the future. We are also considering an evaluation of the potential information gained by updating the predictions throughout the course of the hospitalization.<sup>14</sup>

In conclusion, it is feasible to have a reasonably accurate prediction of mortality risk for most adult patients at the beginning of their hospitalizations. The availability of this prognostic information provides an opportunity to develop proactive care plans for highand low-risk subsets of patients.

#### Acknowledgements

The authors acknowledge the technical assistance of Nehal Sanghvi and Ben Sutton in the development of the Web application and related databases, and the support of the Chief Nursing Officer, Joyce Young, RN, PhD, the emergency department medical staff, Mohammad Salameh, MD, David Vandenberg, MD, and the surgical preparation center staff.

Disclosure: Nothing to report.

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