ORIGINAL RESEARCH

Factors and Economic Outcomes Associated with Documented Difficult Intubation in the United States

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Healthcare Economics Outcomes Research, Medtronic, Mansfield, MA, USA **Purpose:** Establishing good mechanical ventilation is a critical component and prerequisite to a wide range of surgical and medical interventions. Yet difficulties in intubating patients, and a variety of associated complications, are well documented. The economic burden resulting from difficult intubation (DI), however, is not well understood. The current study examines the economic burden of documented DI during inpatient surgical admissions and explores factors that are associated with DI.

Patients and Methods: Using data from the Premier Healthcare Database, adult patients with inpatient surgical admissions between January 1, 2016 and December 31, 2018 were selected. International Classification of Diseases, Tenth Edition, Clinical Modification (ICD-10-CM) diagnosis codes were used to classify the patients into matched cohorts of DI and non-DI patients. **Results:** Patients in the DI group have mean inpatient costs and intensive care unit (ICU) costs that are substantially higher than patients without difficult intubations (\$14,468 and \$4,029 higher, respectively). Mean hospital length of stay and ICU length of stay were 3.8 days and 2.0 days longer, respectively (all p<0.0001, except ICU cost p=0.0001) in the DI group. Obesity, other chronic conditions, and larger hospital size were significantly associated with DI. **Conclusion:** DI is associated with higher average cost and longer average length of stay.

Keywords: cost burden, difficult intubation, airway management, health economics

Plain Language Summary

In this study, we compared the difference in hospital cost and length of hospital stay between two groups of patients. Both groups of patients were admitted to the hospital because of the need for a surgical procedure. When an anesthesiologist inserts a tube into the airway to ventilate a patient before administering anesthesia, the anesthesiologist might encounter some patients for whom the tube is difficult to insert. Patients with this characteristic were designated to the "difficult intubation" (DI) group. Patients without this characteristic were designated to the "non-difficult intubation" (non-DI) group. We made sure these two groups of patients were comparable in terms of similar age, gender, and disease conditions, and stayed in similar types of hospitals and had similar procedures. We found that when we compared the DI group to the non-DI group, on average, the hospital cost was \$14,468 more and length of hospital stay was 3.8 days longer for the DI group. This study highlights the significant cost burden associated with difficult intubation and suggests that further research to enhance clinicians' ability to predict DI would be beneficial.

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Introduction

During tracheal intubation, an anesthetist inserts a tube through the mouth or nose and down the trachea into the lungs. This is a common procedure for patients who

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undergo surgical procedures. Difficult intubation (DI), which increases the number of laryngoscopic attempts during tracheal intubation, has been associated with a variety of complications including oxygen desaturation, hypertension, dental damage, admission to the intensive care unit (ICU), and complications at extubation, as well as arrhythmias, bronchospasm, airway trauma, CICV ("can't intubate, can't ventilate"), and sequela of hypoxia (cardiac arrest, brain damage, and death).¹

Many factors influence the success rate of intubation. Studies have shown that the first-pass success rate of intubation in the operating room ranges from only 63% to 85%.² Up to 93% of difficult intubations are unanticipated.³ Patient risk factors, such as restrictions on neck movement, mouth opening, mallampati class⁴ and thyromental distance and neck extension are common factors.⁵ Many studies have tried to establish the best and most feasible method to predict difficult intubation, including Cormack-Lehane classification system⁶ and the Intubation Difficulty Scale (IDS).⁷ One study of experienced anesthesiologists, utilizing the widely used Cormack-Lehane classification system, found they did not score the same patients consistently over time, and that only 25% could correctly define all four grades of the Cormack-Lehane classification system.⁸ Under certain emergency circumstances (e.g., severe head trauma or suspected cervical spine injury), it may be impossible to fully utilize these various classification systems during physical examination to predict the difficulty of tracheal intubation.⁹

Before the advent of the ICD-10-CM coding system (International Classification of Diseases, Tenth Revision, Clinical Modification), there was no mechanism to quantify the cost burden of failed or difficult intubation from medical claims data. The implementation of ICD-10-CM introduced codes specific to DI, which enables researchers to identify cases with documented DI during surgical procedures. To our knowledge, there is no study currently available that examines the cost and healthcare utilization outcomes associated with difficult intubation in the hospital operating room setting. We utilized the ICD-10-CM coding system to classify patients into two groups, those with and without DI, to conduct a retrospective matched cohort study to fill this knowledge gap. This study will shed light from a hospital provider economic perspective on the consequence of DI in the inpatient hospital setting. The objective of this study was two-fold: First, to examine the healthcare utilization and cost associated with DI during an inpatient surgical procedure; and second, to explore the major patient characteristics and hospital characteristics of cases that were reported as DI.

Patients and Methods Construction and Content

A retrospective observational cohort study was conducted using three years of data (2016–2018) from the Premier Healthcare Database[®] (Premier, Inc., Washington, D.C.).

Data Source

The Premier[®] Database contains data from more than 600 United States (US) hospitals and includes de-identified Universal Billing (UB-04) data for approximately 1 out of every 5 discharges in the U.S.¹⁰ It also includes information for each discharge describing patient demographics, hospital characteristics, payer information, Medicare Severity-Diagnosis Related Group (MS-DRG), ICD-10-CM primary and secondary diagnosis and procedure codes, admission and discharge calendar quarter and year, and detailed hospital department-level billing and cost data for each discharge. As an administrative database containing de-identified data, institutional review board or ethics committee approval was not required for this study. Cost data represent the actual total costs to treat the patient and include both fixed and variable costs.¹⁰ Approximately 60% of hospitals in the Premier database report costs. For the remaining hospitals, costs are calculated by Premier using charges reported to Premier and department-specific cost-to-charge ratios reported by hospitals to the Centers for Medicare & Medicaid Services.

Cohort Selection

All adult patients who underwent inpatient surgical procedures between January 1, 2016 and December 31, 2018 were included in the initial cohort selection. Patients with difficult intubation (DI group) were identified using three ICD-10 codes on the discharge record, which correspond to "failed or difficult intubation" on an initial encounter (T88.4XXA), subsequent encounter (T88.4XXD), or sequela (T88.4XXS). The following patients were excluded from the study: 1) Pregnant women; 2) Cases where general anesthesia was not indicated in the master charge file; 3) Patients with a primary procedure that indicates the Major Diagnostic Category (MDC) was a) not documented, b) "Unrelated Operating Room Procedures" or c) "Invalid or Ungroupable MS-DRG"; and 4) "Pre-MDC" which included all organ transplants or tracheotomies. These exclusion criteria applied to both the DI group and the non-DI group. In addition, the DI group excluded patients with diagnosis code indicating failed or difficult intubation (ICD-10 codes: T88.4XXA, T88.4XXD, T88.4XXS) was present on admission and the non-DI group excluded patients with primary procedures that were not present in the DI group.

Pregnant women were excluded from the study because the level of difficulty of intubation and procedure type likely differ for pregnant women compared to nonpregnant adults. Admissions with primary procedures not in the DI group were excluded because the risk adjustment process includes an exact match on primary procedure as well as a match on propensity score between the DI and non-DI groups to ensure that cost difference is based on comparison of cases with the same primary procedure. The reasons for implementing the remaining exclusion criteria were to make the cohort groups comparable by ensuring all patients in the study underwent general anesthesia during a surgical procedure, all cases were inpatient admissions, and all patients in the DI group experienced difficult intubation during rather than prior to the admission.

Multiple steps were applied to identify the comparison group, non-DI, and implement risk adjustment to confirm comparison of demographically and clinically similar patients. The risk adjustment process involved two steps:

- Exact match between DI and non-DI patients on primary surgical procedures: to qualify for the non-DI group, patients must have the same primary procedure, identified using the ICD-10-PCS (International Classification of Diseases, Tenth Revision, Procedure Coding System) code, as in the DI group (Supplementary Table 1).
- 2. One to one DI vs non-DI propensity score matching within each primary procedure. The matching process between the DI and non-DI groups matched cases within each primary procedure based on propensity scores using a nearest neighbor matching method. The propensity score was calculated using a multivariable logistic regression model which included as predictor variables both patient characteristics (age group, gender, race, admission type (Urgent/Emergent or Elective), insurance type, comorbid conditions), and hospital characteristics (region of the country, teaching status, number of beds, urban/rural designation). The 12 comorbid conditions included in the propensity

score model were obesity, cancer, congestive heart failure (CHF), chronic obstructive pulmonary disease (COPD), cardiovascular disease, diabetes, diabetes with sequela, hypertension, peripheral vascular disease, mild liver disease, moderate/ severe liver disease, and paralysis. These 12 comorbid conditions were selected from the 17 comorbid conditions that make up the Charlson Comorbidity Index (CCI) using a "purposeful selection" process.

The "purposeful selection" process of determining which predictor variables to include in a logistic regression model was developed by researchers at the University of Arkansas for Medical Sciences the and University of Massachusetts, Amherst¹¹ and implemented for this research using an automated SAS macro developed by Bursac et al.¹² The "purposeful selection" process begins with a univariate analysis of the candidate predictive variables, in this case the 17 comorbid conditions. The initial model includes variables with p-values that fall below the pre-defined threshold for statistical significance (p < 0.25). An iterative process adds the variables that were not initially found to be statistically significant into the model one at a time to evaluate each variable for statistical significance in the new model, using a p-value threshold of p < 0.1, and confounding, which is defined as a 15% change in a parameter estimate in the new model compared to the original model. The variables that are statistically significant or determined to be confounders are retained in the new model. This iterative process results in a final variable selection that includes significant variables and important confounding variables.¹¹ The threshold values recommended by the authors of the "purposeful selection" process¹¹ were used to determine which variables to include-p < 0.25 for inclusion in the initial model, p < 0.1 to remain in the model after being added back in, and 15% change in a predictor variable.

Outcome Measures

Primary outcome measures were average cost of admission, hospital length of stay (LOS), post-operative Intensive Care Unit (ICU) admission rate, cost of ICU stay, and length of ICU stay. Post-operative ICU was defined as ICU admissions that occurred on the same day of surgery or within two days post-surgery. Costs for admissions in 2016 and 2017 were adjusted to 2018 dollars using the US Bureau of Labor Statistics Consumer Price Index for Medical Care.

Statistical Analysis

Descriptive univariate analysis was used to compare baseline patient demographics, clinical characteristics, and hospital characteristics between the DI and non-DI groups. Chi-squared test was used for categorical variables and the two sample *t*-test was used for continuous variables such as LOS or cost. A multivariable logistic regression model was used to calculate adjusted odds ratios (OR) to examine patient and hospital characteristics that are associated with documented DI. Variables modeled in the logistic regression included (1) patient characteristics: age group, gender, race, admission type (Urgent/Emergent or Elective), insurance type, comorbid conditions and (2) hospital characteristics: region, teaching status, number of beds, urban/rural designation. Statistical significance was defined as p-value <0.05. All statistical analyses were conducted using SAS[®] 9.4 for UNIX (SAS Institute Inc Cary, NC, USA).

This study did not directly involve human subjects. All patient data were de-identified and were compliant with the Health Insurance Portability and Accountability Act (HIPAA). Therefore, this study was exempt from Institutional Review Board (IRB) approval (45 CFR § 46.0001(b) (4)).

Results

Characteristics of the DI and Non-DI Groups Before Propensity Score Matching

A total of 2,233,751 cases met the eligibility criteria for the analysis. Of these, 609 cases were in the DI group and 2,233,142 cases were in the non-DI group. Patient and provider characteristics, and economic outcomes associated with each cohort, are described in Figure 1 and Table 1.

Prior to matching, patients in the DI group were more likely be male (59% vs. 45%, p<0.0001), black (21% vs. 10%, p<0.0001), less than 65 years old (57% vs. 52%, p=0.019) and have urgent or emergent admissions (54% vs. 39% p=<.0001) (Table 1). Notably, patients in the DI group were more likely to have comorbidities such as obesity (44% vs. 26%), cancer (14% vs. 9%), CHF (22% vs.12%), COPD (29% vs. 18%) and renal disease (20% vs.11%), compared to patients in the non-DI group. All p-values for these comorbidities were <0.0001 (Table 2). Compared to patients in the non-DI group, patients who were treated in a teaching hospital or a large hospital (500 beds or more) were more likely to



Figure I Patient selection criteria for inclusion in the study. Abbreviations: DI, difficult intubation; MDC, major diagnostic category; OR, operating room.

be documented as DI patients, 54% vs. 47% (p=0.0001) and 50% vs. 35% (p<0.0001) respectively (Table 1).

Patients in the DI group had significantly higher mean cost of hospital inpatient stay ($$45,251 \pm $43,613$ vs. $$22,056 \pm $27,008$); longer LOS (10.7 ± 12.1 days vs. 4.5 ± 5.8 days); higher post-operative ICU admission rate (60% vs. 16%); higher ICU costs ($$11,657 \pm $15,653$ vs. $$7,049 \pm $9,386$) and longer length of stay in the ICU (5.4 ± 5.8 days vs. 3.4 ± 3.9 days). All p-values were <0.0001 (Table 3).

Results Based on Matched Cohorts of Patients

Propensity score matching reduced the number of patients in each cohort to 606 (Table 3). The matching process
 Table I Patient and Hospital Demographic Characteristics and Logistic Regression Outputs Prior to Risk Adjustment Matching

Patient	Without	Difficult	With D	Difficult	p-value ^a	Comparing Difficult Intubation to No			
Characteristics	Intub	ation	Intub	ation	-		Difficult	Intubation	1
	2,233	8,142	60)9					
	Number of Patients	Percent of Patients	Number of Patients	Percent of Patients		Odds Ratio	95% Cor Lin	nfidence nits	p-value ^a
Age Category									
< 65 years	1,166,440	52%	347	57%	0.019	N/A			N/A
≥ 65 years	1,066,702	48%	262	43%		0.841	0.666	1.062	0.1454
						F	Reference g	roup=Age<	65
Gender									
Male	1,009,182	45%	358	59%	<0.0001	1.652	1.396	1.953	<0.0001
Female	1,223,960	55%	251	41%		N/A	N/A	N/A	N/A
						I	Reference g	roup=Fema	le
Race									
Black	226,988	10%	125	21%	<0.0001	2.069	1.679	2.551	<0.0001
White	1,775,839	80%	422	69%		N/A	N/A	N/A	N/A
Other	192,258	9%	52	9%		1.062	0.792	1.424	0.6894
Unspecified	38,057	2%	10	2%		1.01	0.538	1.895	0.9748
							Reference §	group=Whi	te
Insurance Type									
Medicaid	213,795	10%	74	12%	0.004	1.103	0.808	1.505	0.538
Managed Care	746,579	33%	183	30%		1.045	0.815	1.339	0.7294
Uninsured	71,561	3%	32	5%		1.562	1.032	2.363	0.0349
Other ins	90,998	4%	29	5%		1.162	0.775	1.742	0.4684
Medicare	1,110,209	50%	291	48%		N/A	N/A	N/A	N/A
						Reference group=Managed care			
Admission Type							_		
Elective	1.361.309	61%	279	46%	<0.0001	N/A	N/A	N/A	N/A
Urgent/Emergent	871,833	39%	330	54%		1.587	1.335	1.886	<0.0001
						Reference group=Elective			
Hospital						Hospital Characteristics			
Characteristics									
Hospital Region									
South	1,060,938	48%	303	50%	0.006	N/A	N/A	N/A	N/A
Northeast	280,004	13%	63	10%		0.862	0.647	1.147	0.3072
Midwest	516,277	23%	117	19%		0.868	0.696	1.083	0.2089
West	375,923	17%	126	21%		1.52	1.221	1.893	0.0002
							Reference	group=Sou	th
Teaching Hospital									
Teaching Hospital	1,039,632	47%	331	54%	0.0001	1.107	0.914	1.34	0.2981
Non-Teaching Hospital	1,193,510	53%	278	46%		N/A	N/A	N/A	N/A
						Refer	ence group [:]	=Teaching H	Hospital

(Continued)

Table I (Continued).

Patient Characteristics	Without Intub	Difficult ation	With D Intub	Difficult ation	p-value ^a	Compar	Comparing Difficult Intubation to N Difficult Intubation		
	2,233	3,142	60)9					
	Number of Patients	Percent of Patients	Number of Patients	Percent of Patients		Odds Ratio	95% Cor Lin	nfidence nits	p-value ^a
Bed Size									
500+ beds	777,012	35%	305	50%	<0.0001	1.567	1.25	1.965	<0.0001
300–500	671,676	30%	147	24%		0.94	0.746	1.185	0.6014
<300	784,454	35%	157	26%		N/A	N/A	N/A	N/A
						Reference group=<300			
Urban vs Rural									
Urban	2,004,270	90%	558	92%	0.127	1.038	0.774	1.393	0.8038
Rural	228,872	10%	51	8%		N/A	N/A	N/A	N/A
							Reference ;	group=Urba	an

Note: ^aChi-square test was used to calculate p-value.

Abbreviation: N/A, not applicable.

Table 2 Distribution of Comorbidules for rations with and without Difficult intubation rifer to hisk Adjustment ration

Patient Comorbidities	Number of Patients without Difficult Intubation	Percent of Patients without Difficult Intubation	Number of Patients with Difficult Intubation	Percent of Patients with Difficult Intubation	p-value ^a	Odds Ratio ^b	Lower 95% Confidence Limits	Upper 95% Confidence Limits	p-value ^a
Obesity	582,344	26%	265	44%	<0.0001	2.202	1.857	2.612	<0.0001
Cancer	190,379	9%	86	14%	<0.0001	1.886	1.458	2.44	<0.0001
Acquired Immune	2,484	0%	2	0%	0.108	1.654	0.41	6.67	0.4796
Deficiency Disease									
Congestive Heart Failure	265,728	12%	135	22%	<0.0001	1.236	0.989	1.545	0.0621
Chronic Obstructive	408,685	18%	176	29%	<0.0001	1.553	1.295	1.861	<0.0001
Pulmonary Disease									
Cardiovascular Disease	107,398	5%	55	9%	<0.0001	1.419	1.054	1.911	0.0211
Diabetes	447,307	20%	177	29%	<0.0001	1.146	0.954	1.378	0.145
Diabetes with Sequela	178,372	8%	107	18%	<0.0001	1.432	1.109	1.849	0.006
Hypertension	1,389,328	62%	452	74%	<0.0001	1.3	1.061	1.592	0.0114
Metastatic Solid Tumor	49,571	2%	24	4%	0.0039	1.123	0.712	1.772	0.6183
Myocardial Infarction	223,036	10%	94	15%	<0.0001	0.988	0.777	1.255	0.9201
Peripheral Vascular	173,699	8%	83	14%	<0.0001	1.289	1.007	1.649	0.0439
Disease									
Renal Disease	250,842	11%	120	20%	<0.0001	1.048	0.815	1.348	0.7135
Mild Liver Disease	79,086	4%	38	6%	0.0003	1.157	0.811	1.651	0.422
Moderate/Severe Liver	9,014	0%	10	2%	<0.0001	2.344	1.193	4.606	0.0134
Disease									
Paralysis	30,879	1%	20	3%	<0.0001	1.435	0.896	2.298	0.1332
Dementia	61,454	3%	16	3%	0.8509	0.919	0.553	1.527	0.7431

Notes: "Chi-square test was used to calculate p-value." Reference group for comorbidity odds ratio is patients without the comorbidity for each comorbid condition.

accounted for most differences in the distribution of demographic and hospital characteristics (Table 4) and comorbid conditions (Table 5). The difference between DI and non-DI matched cohorts for all outcome measures remained significant. Compared to the non-DI patient group, patients in the DI group had \$14,468 higher mean

						Before Ris	sk Adjustme	ent						
			Difficult	ntubation					2	Io Difficult I	ntubation			
	z	Mean	ICU Admission Rate	95% Lower CI	95% Upper CI	STD	z	Mean	ICU Admission Rate	95% Lower CI	95% Upper CI	STD	p-value	Mean Difference
Admission Cost (US Dollare)	609	\$45,251	I	\$41,780	\$48,722	\$43,613	2,233,142	\$22,056	I	\$22,020	\$22,091	\$27,008	<0.0001	\$23,196
Admission Length of	609	10.7	%09	9.7	11.7	12.1	2,233,142	4.5	16%	4.5	4.5	5.8	<0.0001	6.2
icu Cost (US Dollars)	367	\$11,657	I	\$10,051	\$13,264	\$15,653	358,359	\$7,049	I	\$7,018	\$7,079	\$9,386	<0.0001	\$4,609
ICU Length of Stay (Days)	367	5.4	I	4.8	6.0	5.8	358,359	3.4	I	3.3	3.4	3.9	000.0>	2.0
						After Ris	k Adjustme	int						
			Difficult I	ntubation					2	Io Difficult I	ntubation			
	z	Mean	ICU Admission	95% Lower	95% Upper	STD	z	Mean	ICU Admission	95% Lower	95% Upper	STD	p-value	Mean Difference
			Rate	ū	σ				Rate	Ū	CI			
Admission Cost (US Dollars)	909	\$45,233	I	\$41,748	\$48,717	\$43,679	606	\$30,765	I	\$28,242	\$33,288	\$31,624	000.0>	\$14,468
Admission Length of Stay (Days)	606	10.7	60%	9.8	11.7	12.1	606	6.9	33%	6.3	7.5	7.7	<0.0001	3.8
ICU Cost (US Dollars)	364	\$11,662	I	\$10,043	\$13,280	\$15,702	198	\$7,633	I	\$6,393	\$8,873	\$8,849	0.0001	\$4,029
ICU Length of Stay	364	5.4		4.8	6.0	5.8	198	3.4		2.9	4.0	3.8	<0.0001	2.0
(Uays)														
Notes: ICU admission rate, co Abbreviations: Cl, confidence	st (US c interva	dollars), and II; ICU, inten	length of stay (days) isive care unit; N, nu	are reported i imber of patien	for patients wh its; STD, stand	no had an IC ard deviation	U admission 0)–2 days aftei	r the primary surger	×				

Table 3 Cost and Utilization Differences Between Patients with and without Difficult Intubation Before and After Risk Adjustment

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Table 4 Demographic	Characteristics of	Patients with	n Difficult	Intubation	Hospital	Inpatient	Admissions	for	Patients	After	Risk
Adjustment Matching											

Patient Characteristics	Without Intub	Difficult ation	With Difficult Intubation		p-value ^a	Comparing Difficult Intubation to No Difficult Intubation			
	60)6	60	6			95% Cor Lin	nfidence nits	
	Number of Patients	Percent of Patients	Number of Patients	Percent of Patients		Odds Ratio	Lower Limit	Upper Limit	p-value ^a
Age Category									
< 65 years	324	53%	346	57%	0.204	N/A			N/A
≥ 65 years	282	47%	260	43%		0.948	0.676	1.331	0.7591
						l	Reference g	roup=Age<	:65
Gender									
Male	359	59%	356	59%	0.861	0.994	0.783	1.262	0.9595
Female	247	41%	250	41%		N/A	N/A	N/A	N/A
							Reference a	group=Fema	ale
Race									
Black	114	19%	125	21%	0.691	1.162	0.854	1.582	0.3384
White	437	72%	420	69%		N/A	N/A	N/A	N/A
Other	48	8%	51	8%		1.145	0.738	1.775	0.5456
Unspecified	7	1%	10	2%		1.453	0.532	3.971	0.4661
							Reference	group=Whi	te
Medicaid	87	14%	74	12%	0.293	0 822	0 528	1 279	0 3856
Managed Care	149	25%	182	30%	0.275	1 192	0.832	1.277	0.3392
Uninsured	34	<u>6%</u>	32	5%		0.958	0.534	1.707	0.8863
Other ins	29	5%	29	5%		1.036	0.578	1.855	0.9065
Medicare	307	51%	289	48%		N/A	N/A	N/A	N/A
		• • • •				Pof		-Managor	d care
Admission Type									
Elective	272	45%	277	46%	0.773	N/A	N/A	N/A	N/A
Urgent/Emergent	334	55%	329	54%		0.951	0.743	1.217	0.688
							Reference g	roup=Elect	ive
Hospital Characteristic	s								
Hospital Region									
South	325	54%	301	50%	0.495	N/A	N/A	N/A	N/A
Northeast	59	10%	62	10%		1.178	0.771	1.802	0.4486
Midwest	100	17%	117	19%		1.287	0.926	1.788	0.1338
West	122	20%	126	21%		1.126	0.821	1.545	0.462
							Reference	group=Sou	th
Teaching Hospital									
Teaching Hospital	347	57%	330	54%	0.325	0.884	0.667	1.17	0.3869
Non-Teaching Hospital	259	43%	276	46%		N/A	N/A	N/A	N/A
						F	। Reference व	roup=Teach	ing
[8		0

(Continued)

Patient Characteristics	Without Intub	Difficult ation	With D Intub	Difficult ation	p-value ^a	Comp	Comparing Difficult Intuba No Difficult Intubatio			
	60	06	60)6			95% Co Lin	nfidence nits		
	Number of Patients	Percent of Patients	Number of Patients	Percent of Patients		Odds Ratio	Lower Limit	Upper Limit	p-value ^a	
Bed Size										
500+ beds	319	53%	303	50%	0.638	1.027	0.735	1.435	0.8758	
300–500	141	23%	146	24%		0.99	0.705	1.39	0.9526	
<300	146	24%	157	26%		N/A	N/A	N/A	N/A	
							Reference group=<300			
Urban vs Rural										
Urban	558	92%	555	92%	0.753	0.939	0.609	1.447	0.7753	
Rural	48	8%	51	8%		N/A	N/A	N/A	N/A	
							Reference	group=Urb	an	

Table 4 (Continued).

Note: ^aChi-square test was used to calculate p-value. **Abbreviation:** N/A, not applicable.

cost of hospital inpatient stay ($$45,233 \pm $43,679$ vs. \$30,765 \pm \$31,624); 3.8 days longer LOS (10.7 \pm 12.1 days vs. 6.9 \pm 7.7 days); higher post-operative ICU admission rate (60% vs. 33%); \$4029 higher ICU costs (\$11,662 \pm \$15,702 vs. \$7,633 \pm \$8,849); and 2.0 days longer length of stay in the ICU (5.4 \pm 5.8 days vs. 3.4 \pm 3.8 days). All p-values were <0.0001 (Table 3).

Factors Associated with Documented DI

Both patient characteristics and hospital characteristics were independently associated with the likelihood of documented DI. Compared to patients who were white, female, or having elective surgical procedure, the odds of being in the DI group were 2.1 times higher for black patients (OR: 2.1, 95% CI: 1.7-2.6; p<0.0001); 1.7 times higher for male patients (OR: 1.7, 95% CI: 1.4-2.0, p<0.0001); and 1.6 times higher for patients with urgent or emergent admission (OR: 1.6; 95% CI: 1.3-1.9, p<0.0001) (Table 1). Additionally, when evaluating the risk associated with specific chronic conditions, the risk was higher for patients with the chronic condition than for patients without the specific condition. The risk was 2.2 times higher for patients with obesity (OR: 2.2; 95% CI: 1.9-2.6; p < 0.0001; 1.9 times higher for patients with cancer (OR: 1.9; 95% CI: 1.5-2.4; p<0.0001); 1.6 times higher for patients with COPD (OR: 1.6; 95% CI: 1.3-1.9; p<0.0001); and 1.4 times higher for patients with cardiovascular disease (OR: 1.4; 95% CI: 1.1–1.9; p=0.02) (Table 2). Hospital factors included hospital size and US Census Region. Compared to patients in hospitals with fewer than 300 beds, patients at hospitals with more than 500 beds were 1.6 times more likely to have documented DI cases (OR: 1.6, 95% CI: 1.3–2.0; p<0.0001) and patients at hospitals in the West region were 1.5 times more likely to have documented DI cases compared to hospitals in the South region (OR: 1.5, 95% CI: 1.2–1.9, p<0.0001) (Table 1).

Discussion

Airway management is fundamental to safe anesthetic practice. Difficult intubation, failed intubation, and wrongly placed tracheal tubes account for 2.3% of all anesthesia-related deaths.¹³ Problems with tracheal intubation were the most frequently recorded primary airway problem, which accounted for 39% of all events during anesthesia.⁹ To the best of our knowledge, this is the first study to use ICD-10-CM diagnosis codes to identify documented DI and explore factors associated with DI using Real World Data (RWD), and to examine the difference in healthcare utilization between DI and non-DI patient groups. This new information highlights the significant cost burden associated with DI. The variance in both cost and length of stay measures was high in the DI and non-DI groups, suggesting that while the average patient in the DI

Patient Comorbidities	Number of Patients without Difficult Intubation	Percent of Patients without Difficult Intubation	Number of Patients with Difficult Intubation	Percent of Patients with Difficult Intubation	p-value ^a	Odds Ratio ^b	Lower 95% Confidence Limits	Upper 95% Confidence Limits	p-value ^a
Obesity	217	36%	264	44%	0.0058	1.251	0.975	1.606	0.0781
Cancer	97	16%	86	14%	0.3775	0.96	0.665	1.386	0.8268
Acquired Immune	-	0%	2	0%	0.157	N/A	N/A	N/A	0.9794
Deficiency Disease									
Congestive Heart Failure	119	20%	133	22%	0.3217	1.16	0.845	1.594	0.3579
Chronic Obstructive	180	30%	175	29%	0.7523	0.948	0.73	1.232	0.6907
Pulmonary Disease									
Cardiovascular Disease	59	10%	54	9%	0.6213	0.877	0.571	1.35	0.5518
Diabetes	147	24%	176	29%	0.0596	1.201	0.909	1.587	0.1979
Diabetes with Sequela	85	14%	106	17%	0.0978	1.242	0.843	1.828	0.2727
Hypertension	447	74%	450	74%	0.8442	0.947	0.707	1.268	0.7141
Metastatic Solid Tumor	32	5%	24	4%	0.2737	0.788	0.424	1.465	0.4517
Myocardial Infarction	85	14%	94	16%	0.4662	1.109	0.788	1.562	0.553
Peripheral Vascular	92	15%	83	14%	0.462	0.87	0.614	1.232	0.4315
Disease									
Renal Disease	116	19%	120	20%	0.7717	0.915	0.642	1.304	0.6234
Mild Liver Disease	35	6%	38	6%	0.7172	1.244	0.724	2.14	0.429
Moderate/Severe Liver	16	3%	10	2%	0.2342	0.556	0.226	1.365	0.2003
Disease									
Paralysis	19	3%	20	3%	0.8707	1.188	0.583	2.418	0.6358
Dementia	19	3%	16	3%	0.6068	0.918	0.449	1.878	0.8145

Table 5 Patient and Hospital Demographic Characteristics and Logistic Regression Outputs After Risk Adjustment Matching

Notes: ^aChi-square test was used to calculate p-value. ^bReference group for comorbidity odds ratio is patients without the comorbidity for each comorbid condition. **Abbreviation:** N/A, not applicable.

group experiences higher costs and length of stay compared to the average patient in the non-DI group, at the individual patient level, these differences may be more difficult to detect.

Our finding of obesity as a risk factor for DI is consistent with other studies which have indicated that patients with obesity were at increased risk of an adverse airway event.^{9,14–16} Reasons for this include mechanical difficulty in securing the airway, increased risk of aspiration, increased risk of airway obstruction during intubation, and accelerated speed and extent of oxygen desaturation during airway obstruction.^{17–19} The current study also suggests that patient gender, race, and other chronic conditions are associated with risk of DI. This is consistent with previous reports that increasing age, American Society of Anesthesiologists physical status, and CCI are associated with failed and difficult intubation.^{20,21} Importantly, many of the factors associated with DI in our study are linked to obesity, including diabetes, cardiovascular disease, cancer, and liver disease.²²⁻²⁵ Further prospective research is required to determine whether some of the chronic conditions are directly related to DI, or if they are confounding factors associated with a major predictor of DI, such as obesity. Other conditions, including the presence of head and neck cancer, arthritis, ankylosing spondylitis, and diabetes with limited joint mobility, which can alter the patient's ability to lie flat are also associated with failed and difficult intubation, due to narrowing of the airway and challenges in positioning during intubation.^{26–28} In addition, large hospitals with >500 beds were more likely to have documented DI cases, which may reflect more consistent reporting standards often instituted at large hospitals. Although risk factors associated with DI have been identified, prediction of DI is a challenging task, and additional research to understand the predictors for DI would be beneficial, allowing clinicians to choose the appropriate method and device for patient airway management in challenging cases.^{16,29–31}

Implications and Future Actions

DI may occur in any type of surgical procedure and it can be costly when it happens. Our research highlights a need for clinicians who perform tracheal intubations to carefully consider options that could improve the chances of

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successful intubation on the first attempt. The advances in new technology in tracheal intubation in a hospital setting may provide alternative options. For example, several studies demonstrated the benefits of video laryngoscopy, including a shorter time required for tracheal intubation, a higher rate of successful intubations, and a reduced need for external manipulation of the larynx or other maneuvers in the case of an expected DI.^{32–35} The benefits and costs associated with alternative technologies should be considered through future research.

The use of administrative data for retrospective analysis has both strengths and weaknesses. Due to the large number of hospitals, we have a good representation of hospitals in all regions of the United States, as well as good representation of patients with different types of insurance coverage. However, we expect the administrative data to suffer significant under-documentation of DI in the medical claims data. Only 0.03% of admissions in this study were found to have documented DI. Elsewhere, the reported incidence of DI has been estimated to be 1.5-8.5%, while failed intubation occurred in 0.13–0.3% of the general population.³⁶ This discrepancy between expected and documented DI suggests that in our analysis, some admissions with DI may be assigned to the cohort assumed to have no DI. It is also possible that admissions that have documented DI have more serious complications resulting from DI than admissions that may have experienced undocumented DI. Therefore, the cost differential identified in the current study may be a conservative estimate overall, but not generalizable beyond cases of documented DI. Finally, our analysis did not include access to patient electronic medical records, limiting our ability to evaluate patient clinical characteristics to the data available in the administrative dataset.

Conclusion

In conclusion, the current study demonstrates the significant cost burden averaging \$14,468 and increased postoperative LOS averaging 3.8 days for patients with documented difficult tracheal intubation.

Abbreviations

DI, difficult intubation; CFR, Code Of Federal Regulations; CHF, congestive heart failure; CI, confidence interval; CICV, can't intubate can't ventilate; COPD, chronic obstructive pulmonary disease; HIPAA, Health Insurance Portability and Accountability Act; ICD-10-CM, International Classification Of Diseases, Tenth Edition, Clinical Modification; ICU, intensive care unit; IRB, Institutional Review Board; LOS, length of stay; MDC, Major Diagnostic Category; MS-DRG, Medicare Severity-Diagnosis Related Group; RWD, real world data; US, United States.

Data Sharing Statement

The data that support the findings of this study are available from Premier Inc. but restrictions apply to the availability of these data, which were used under license for the current study, and so are not publicly available. Data may be available from the authors upon reasonable request and with permission of Premier Inc.

Ethics Approval and Informed Consent

This is a retrospective cohort study using medical billing/ encounter data. No registry was used in current study. The data we used are de-identified and complied with Health Insurance Portability and Accountability Act (HIPAA) confidentiality requirements. Per US Title 45 CFR (Code of Federal Regulations) Part 46.101, institutional review board (IRB) approval for this study was not required under the exemption that this research involved the study of existing data and that the information was recorded in such a manner that the subjects could not be identified, directly or through identifiers linked to the subjects.

Author Contributions

All authors made substantial contributions to conception and design, acquisition of data, or analysis and interpretation of data; took part in drafting the article or revising it critically for important intellectual content; agreed to submit to the current journal; gave final approval of the version to be published; and agree to be accountable for all aspects of the work.

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Disclosure

All 3 authors are paid employees of Medtronic. The authors report no other conflicts of interest in this work.

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