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Prophylactic Antibiotic Choice and Risk of Surgical Site Infection After Hysterectomy

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Abstract

OBJECTIVE—To evaluate associations between prophylactic preoperative antibiotic choice and surgical site infection rates after hysterectomy.

METHODS—A retrospective cohort study was performed of patients in the Michigan Surgical Quality Collaborative undergoing hysterectomy from July 2012 to February 2015. The primary outcome was a composite outcome of any surgical site infection (superficial surgical site infections or combined deep–organ space surgical site infections). Preoperative antibiotics were categorized based on the recommendations set forth by the American College of Obstetricians and Gynecologists and the Surgical Care Improvement Project. Patients receiving a recommended antibiotic regimen were categorized into those receiving beta-lactam antibiotics and those receiving alternatives to beta-lactam antibiotics. Patients receiving non-recommended antibiotics were categorized into those receiving overtreatment (excluded from further analysis) and those receiving non-standard antibiotics. Multivariable logistic regression models were developed to estimate the independent effect of antibiotic choice. Propensity score matching analysis was performed to validate the results.

RESULTS—The study included 21,358 hysterectomies. The overall rate of any surgical site infection was 2.06% (N=441). Unadjusted rates of ‘any surgical site infection’ were 1.8%, 3.1% and 3.7% for beta-lactam, beta-lactam alternatives and non-standard groups, respectively. After adjusting for patient and operative factors within clusters of hospitals, compared to the beta-lactam antibiotics (reference group), the risk of ‘any surgical site infection’ was higher for the group receiving beta-lactam alternatives (OR 1.7, CI 1.27–2.07) or the non-standard antibiotics (OR 2.0, CI 1.31–3.1).

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CONCLUSION—Compared to women receiving beta lactam antibiotic regimens, there is a higher risk of surgical site infection after hysterectomy among those receiving a recommended beta lactam alternative or non-standard regimen.

Surgical site infections are associated with increased hospital length of stay and increased episode cost after surgery.^{1,2} In addition, surgical site infections are the most common reason for readmission after a wide variety of operations.³ The rate of overall surgical site infections (superficial, deep, and organ space) in hysterectomy has been reported to range between 1% and 4%.^{4,5} Hysterectomy is among the most common major operations in the United States (600,000 performed annually) and may result in 6,000 to 24,000 surgical site infections each year. Consequently, beginning October 1, 2015, inpatient post-hysterectomy surgical site infections were included in the Centers for Medicare and Medicaid Services calculations for Hospital-Acquired Condition Reduction Program metrics.⁶

Preoperative prophylactic antibiotic administration has been shown to consistently reduce the rate of postoperative surgical site infections.⁷ The American College of Obstetricians and Gynecologists (the College) has issued guidelines for choosing appropriate preoperative prophylactic antibiotics⁸; in addition, the Joint Commission's Surgical Care Improvement Project has issued a list of procedure-specific prophylactic antibiotics.⁹ Studies have shown that compliance with these guidelines varies across institutions and procedures,¹⁰ and regimens not in compliance have involved both undertreated and overtreated cohorts.

The objective of the current study is to quantify the effects of preoperative antibiotic choice on surgical site infection rates after hysterectomy using data from a statewide surgical collaborative.

MATERIALS AND METHODS

A retrospective cohort study was performed of patients in the Michigan Surgical Quality Collaborative (herein also referred to as the collaborative) undergoing hysterectomy from July 2012 to February 2015. The collaborative is funded by the Blue Cross Blue Shield of Michigan/Blue Care Network, and it includes patients from all insurance payers (public and private). At each participating hospital, a trained, dedicated nurse abstractor collects patient characteristics, intraoperative processes of care (including the details of preoperative antibiotics administered), and 30-day postoperative outcomes from general and vascular surgery and hysterectomy cases. To ensure complete capture of the data, nurse abstractors make phone calls to the patients to determine whether they were admitted to a hospital other than the one in which the index surgery was performed. To reduce sampling error, a standardized data collection methodology is used whereby data abstraction is performed on only the first 25 cases of an 8-day cycle (alternating on different days of the week for each cycle). The standardized data collection methodology is routinely validated through scheduled site visits, conference calls, and internal audits.^{11,12}

Patients were included in the study if they were older than 18 years of age and were undergoing abdominal, vaginal, laparoscopic, or robotic hysterectomy. Patients with gynecologic malignancy and those undergoing hysterectomy for benign indications were included in the study. Patients with no recorded antibiotic information and those with

missing surgical site infection information were excluded from the analysis. Michigan Surgical Quality Collaborative datasets provided to the researchers contain no patient, hospital or healthcare provider identifiers. Therefore, this study met the criteria for “exempt” status by the University of Michigan Institutional Review Board–Medical (HUM00073978).

The following information was available for analysis: age at the time of surgery; body mass index (BMI), calculated as weight/height (kg/m^2); covariates associated with performance status, including American Society of Anesthesiologists (ASA) classification score (defined as a dichotomous variable an ASA class <3 or 3)¹³; and preoperative medical history, including diabetes mellitus (defined as requiring oral hypoglycemic agents and/or insulin), hypertension (defined as documentation in preoperative evaluation or of receiving antihypertension medications), and smoking status (defined as having smoked cigarettes, cigars, or pipe, chewed tobacco, or used marijuana within the past year). Preoperative transfusion was defined as having receiving a minimum of one unit of whole blood or packed red blood cells during the 72 hours before surgery. Patients with a final diagnosis coded as 179 to 184 based on the primary International Classification of Diseases, Ninth Revision (ICD-9) were defined as having the diagnosis of gynecologic cancer. All other ICD-9 diagnoses were defined as benign final pathology.

Approach to hysterectomy was categorized as open (all abdominal hysterectomy cases and all cases converted from laparoscopic or robotic cases) or minimally invasive, which encompassed laparoscopic (including robotic-assisted cases) and vaginal (including laparoscopic-assisted cases). Surgical complexity was calculated by adding the relative value units for each surgical procedure recorded for the patient. Operative times were reported in hours, from the start of the surgery (incision) to the closing of the skin incision.

Surgical site infections within 30 days of surgery were defined by the Centers for Disease Control and Prevention criteria. A superficial surgical site infection involved only skin and the subcutaneous tissue of the incision. In this study, deep and organ space surgical site infections were both considered “deep surgical site infections” because the fascia and muscle layers of the vaginal cuff are contiguous with the organ space. The primary outcome of the study was a composite outcome of any surgical site infection. The term *any surgical site infection* indicates when there was either a superficial or deep surgical site infection.

Preoperative antibiotics were categorized based on the criteria set forth by the the American College of Obstetricians and Gynecologists⁸ and the Surgical Care Improvement Project.⁹ Patients receiving an antibiotic regimen recommended by ACOG or the Surgical Care Improvement Project were further categorized into those receiving beta-lactam antibiotics (e.g. cephalosporin, ampicillin-sulbactam, ertapenem) and those receiving alternatives to beta-lactam antibiotics (e.g. combination of clindamycin with gentamicin or quinolone). Patients receiving antibiotic regimens not recommended by the ACOG or the Surgical Care Improvement Project were categorized as those receiving overtreatment (e.g. recommended antibiotic with additional antibiotic) and those receiving non-standard antibiotics (e.g. Clindamycin alone). Patients who received overtreatment were excluded from the analysis because documented antibiotic resistance could account for such a decision. Figure 1 illustrates the development of the antibiotic categories.

For all included patients, descriptive and comparative statistics of demographics, comorbidities, operative details, and postoperative surgical site infections were analyzed. For bivariate analyses, chi-square analysis or Fisher's exact test was used. For continuous variables, parametric one-way analysis of variance or nonparametric Wilcoxon Mann-Whitney tests were used to assess significance in the bivariate relationship. To ascertain the independent effect of antibiotic categories included in the analysis, we constructed multivariate logistic regression models. Variables were excluded from model selection if they were not significant at a level of 0.1 in the bivariate analysis or if they were not related to the outcome in a clinically plausible manner.

For all logistic regression models, to account for violations in model assumptions due to non-independence of observations within clusters of data (hospital level), we used Huber-Eicker-White robust standard errors. These robust standard errors and the hospital-level clustering allowed the model to better reflect the collected data characteristics.^{13–15} We used STATA 14.0 SE for Macintosh (StataCorp LP, College Station, TX) for all analyses. Results of the logistic regression models were confirmed using propensity score matching (Appendix 1, available online at <http://links.lww.com/xxx>).

RESULTS

A total of 22,992 patients undergoing hysterectomy were available in the collaborative database. Excluded from the analysis were cases with no recorded antibiotic information (n=418; 1.8%) and with missing surgical site infection information (n=29; 0.1%). Patients who received overtreatment were excluded from the analysis (n=1,187; 5.1%). There were 21,358 (93%) who were included in the analysis (Fig 1).

Most of these patients received beta-lactam antibiotics (N=17,827; 79.1%), followed by the beta-lactam alternatives (N=2,878; 12.8%). The non-standard regimens were administered in 2.8% (N=653) of cases (Fig. 1). The majority of patients in the non-standard group received single agent antibiotics (Clindamycin alone – 67%; Gentamicin only – 8%) Details of the 15 different regimens included in this group are provided in Appendix 2, available online at <http://links.lww.com/xxx>.

The overall rate of any surgical site infection was 2.06% (N=441). Patients with any surgical site infection were older, had higher BMI, were more likely to have diabetes, were more likely to report tobacco use, received a preoperative transfusion, and had gynecologic cancer as a surgical indication. In addition, patients with any surgical site infection had higher use of open abdominal approach, higher median blood loss, higher complexity of surgery (measured by mean relative value units), and longer operative times (Table 1). Baseline comparison between the three groups of antibiotic categories is provided in Table 2. Patients receiving a beta-lactam antibiotics had lower incidence of tobacco use, ASA class 3, history of hypertension, and history of diabetes. The three groups did not differ in the operative time, blood loss, surgical complexity, and proportion of patients with malignancy. The beta-lactam antibiotics group had a higher proportion of patients undergoing open surgery than the other two groups. Unadjusted surgical site infection rates were 1.8% for beta-lactam antibiotics, 3.1% for beta-lactam alternatives, and 3.75% for non-standard antibiotics.

Details of the unadjusted rates of any surgical site infection, superficial surgical site infections, and deep surgical site infections are provided in Table 2.

Multivariate logistic regression models were constructed for any surgical site infection, superficial surgical site infections, and deep surgical site infections. Table 3 summarizes the independent effect of factors included in the regression models. Compared to the beta-lactam antibiotics (reference group), patients receiving the beta-lactam alternatives had increased risk of any surgical site infection (odds ratio [OR] 1.62, 95% confidence interval [CI] 1.27–2.07, $P < .001$), superficial surgical site infections (OR 1.5, 95% CI 1.04–2.09, $P = .03$), and deep–organ space surgical site infections (OR 1.7, 95% CI 1.27–2.4, $P < .001$). Similarly, compared to the beta-lactam antibiotics (reference group), patients receiving any non-standard regimen had at least twice the risk of any surgical site infection (OR 2.0 95% CI 1.31–3.1, $P < .001$), superficial surgical site infections (OR 2.5, 95% CI 1.46–4.34, $P < .001$) but did not differ significantly in the rate of deep–organ space (Table 3). The adjusted rate of any surgical site infection with respect to the antibiotic categories is shown in Figure 2. Results of the logistic regression were validated using propensity score matching (Appendix 1, <http://links.lww.com/xxx>).

The overall rate of non-standard antibiotics uses in the collaborative dropped from 5.2% to 2.5% over the study time period. (Fig. 3)

DISCUSSION

In this retrospective analysis of patients undergoing hysterectomy in the Michigan Surgical Quality Collaborative, we found that the choice of antibiotic regimen given before hysterectomy independently predicts the rate of any surgical site infection. Beta-lactam antibiotics (cephalosporins, ampicillin-sulbactam, ertapenem) are associated with the lowest rates of surgical site infections. Recommended beta-lactam alternatives (e.g. Clindamycin plus Gentamicin or Quinolone or Aztreonam) and patients receiving non-standard regimens (e.g. Gentamicin only, Clindamycin only) have a significantly higher risk of surgical site infections. One possible explanation is that beta-lactam antibiotics are highly effective against skin flora (*Streptococcus* species, *Staphylococcus aureus*, and coagulase-negative staphylococci), which are the predominant organisms that cause surgical site infections.^{16–18} Regimens that do not contain a beta-lactam antibiotic are inferior in controlling these organisms.⁷

Given this increased risk, patient-reported allergy to penicillin (PCN) should be thoroughly investigated to ascertain its validity and severity. Previous studies have shown that because of the fear of PCN anaphylaxis, clinicians frequently accept a diagnosis of PCN allergy without obtaining a detailed history of the reaction.¹⁹ In our study, approximately 12% of the patients received a beta-lactam alternative antibiotic regimen, a prevalence consistent with the self-reported PCN allergy described in the literature.²⁰ It is important to remember that cephalosporin cross-reactivity shown in skin testing is present in only 10% of patients with a true PCN allergy.^{19,21} Patients with negative results on PCN skin testing and those without a history of anaphylactic reaction to PCN can safely receive cephalosporin.^{19,22,23}

Routine use of PCN skin testing could potentially increase the use of cephalosporins and therefore reduce the use of alternative antibiotics in perioperative settings.^{24,25}

The current analysis quantifies the association of administering antibiotics not recommended by the College or by Surgical Care Improvement Project guidelines before hysterectomy. Wright et al¹⁰ reported that 2.3% of patients undergoing gynecologic surgery received antibiotics not recommended by the guidelines. However, the authors did not report the effect of non-adherence to guidelines on surgical site infection rates. In our study, the majority of patients who received a non-standard regimen received a single-agent antibiotic (clindamycin, gentamicin, or metronidazole). Previous studies have shown these single agents are inferior to cephalosporins.²⁶ Studies have also shown that adherence to Surgical Care Improvement Project's surgical site infection reduction bundle into surgical safety checklists can significantly improve antibiotic infusion timing and antibiotic selection.²⁷ In our study, for each quarter starting in July 2012, the percentage of patients in the collaborative who receive non-standard antibiotics has consistently decreased (Fig. 3). Although precise reasons of this improvement are likely multifactorial, the participation of hospitals in a functional collaborative encouraging evidence-based practices seems to improve the quality of surgical care across the hospitals.²⁸

This study has several strengths. The Michigan Surgical Quality Collaborative is a statewide collaborative that uses standardized data collection methods and dedicated nurse abstractors who are regularly audited for interrater reliability. Although the collaborative is limited to a single state, it includes a mix of academic and community hospitals, making the data more generalizable. In addition, our logistic regression modeling accounted for the clustering effect from physician/facility preferences. Studies have shown that the quality of data from collaboratives such as the Michigan Surgical Quality Collaborative and the National Surgical Quality Improvement Project is similar to that of chart review and much better than that of administrative claims-based databases.²⁹

Limitations of our study include reported heterogeneity in surgical site infection reporting in the literature; however, collaborative abstractors are trained to reduce variations in reporting. Moreover, the Centers for Disease Control and Prevention criteria for surgical site infection diagnosis may underestimate the true incidence of surgical site infections by excluding cases of cellulitis by as much as threefold.³⁰ Even though the nurse abstractors follow up with patients by phone within the 30-day period to avoid missing capturing complications if patients seek care in another hospital, potential for underreporting surgical site infections remains. Lastly, data on the appropriate timing and dosage of antibiotics were not available, and variations in these could have affected the conclusions of this study.

In summary, efforts to decrease surgical site infections could focus on adherence to recommended preoperative antibiotic guidelines and thorough evaluation of patient reported penicillin allergies in order to increase the number of patients receiving beta lactam antibiotics.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

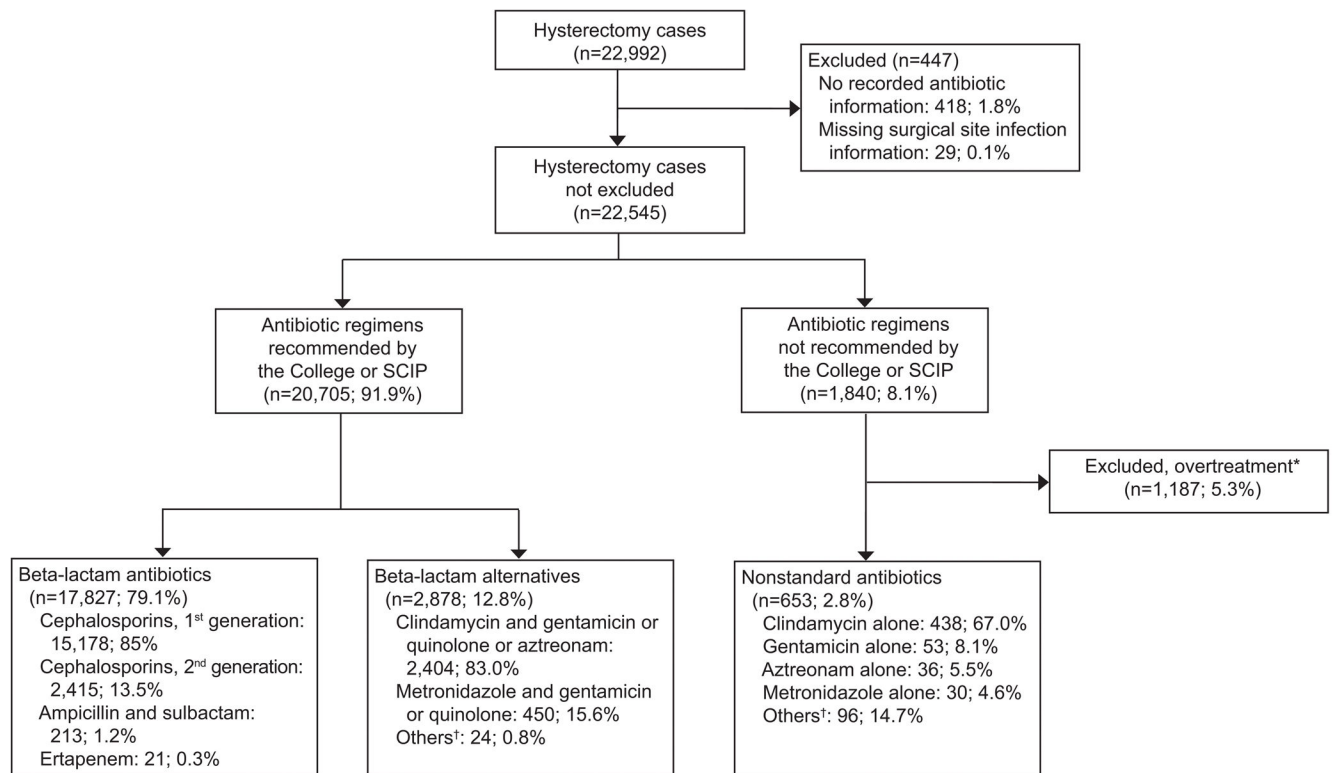
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**Fig. 1.**

Breakdown of the antibiotic categories based on the American College of Obstetricians and Gynecologists (the College) usage guidelines and the Surgical Care Improvement Project (SCIP) usage guidelines. Antibiotic regimen details available in Appendix 2, available online at <http://links.lww.com/xxx>. *Patients receiving additional antibiotics to those recommended by the College and SCIP guidelines were categorized as overtreatment.

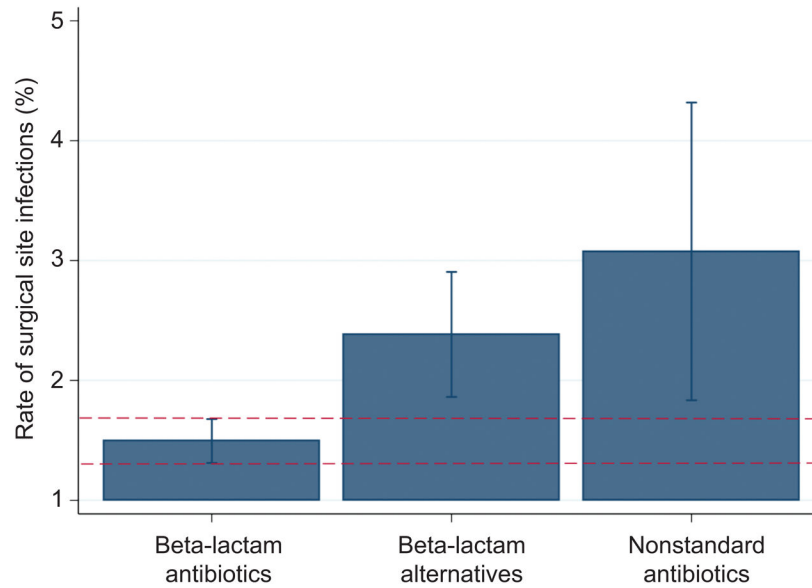


Fig. 2.

Adjusted rates of overall surgical site infection by antibiotic category. Rates adjusted for patient factors (age, body mass index, American Society of Anesthesiologists category, history of diabetes, gynecologic malignancy, and tobacco use) and operative factors (surgical time, blood loss, and surgical complexity). *Red dashed lines* indicate 95% confidence interval bounds for referent category.

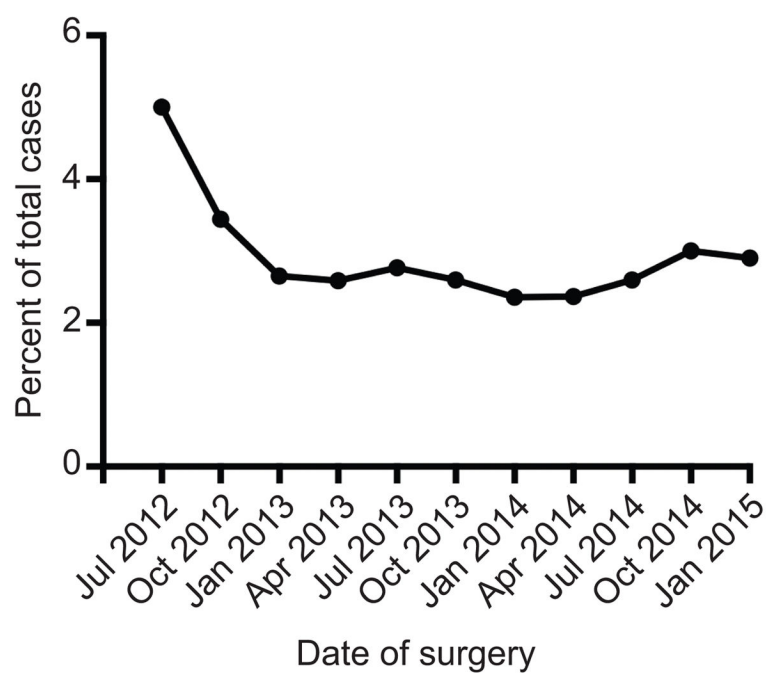


Fig. 3.

Proportion of patients over time enrolled in the Michigan Surgical Quality Collaborative receiving nonstandard antibiotic regimens per the American College of Obstetricians and Gynecologists usage guidelines and the Surgical Care Improvement Project usage guidelines.

Table 1

Predictors of Surgical Site Infection (unadjusted)

Variable	Overall N=22,358	Surgical Site Infection (any)		P
		Absent N=22,917	Present N=441	
Demographics and Comorbidities				
Age, y, median (SD)	48.1 (11.7)	48.1 (11.7)	47.9 (11.8)	.8
BMI, N (%)				
30 kg/m² obese	10,150 (47.5)	9,879 (97.3)	271 (2.7)	<.001
<30 kg/m² nonobese	11,208 (52.5)	11,038 (98.5)	170 (1.5)	
Diabetes, N (%)				
Present	1,984 (8.8)	1,911 (96.3)	73 (3.7)	<.001
Absent	20,561 (91.2)	20,169 (98.1)	392 (1.9)	
Smoker, N (%)				
Yes	4,991 (23.4)	4,864 (97.5)	127 (2.5)	.006
No	16,367 (76.6)	16,053 (98.1)	314 (1.9)	
ASA class, N (%)				
2	16,812 (78.7)	16,514 (98.2)	298 (1.8)	<.001
3	4,546 (21.2)	4,403 (96.8)	143 (3.2)	
History of hypertension, N (%)				
Present	6,358 (30)	6,209 (97.2)	176 (2.8)	<.001
Absent	14,973 (70)	14,708 (98.2)	265 (1.8)	
Preoperative transfusion, N (%)				
Yes	146 (0.7)	139 (95.2)	7 (4.8)	.02
No	21,212 (99.3)	20,778 (98)	434 (2)	
Final pathology, N (%)				
Cancer	1,997 (9.3)	1,911 (95.7)	86 (4.3)	<.001
Benign	19,261 (90.7)	19,006 (98.2)	355 (1.8)	
Surgical Factors				
Surgical approach, N (%)				
Open	5,797 (27.1)	5,569 (96.1)	228 (3.9)	<.001
Minimally invasive *	15,561 (72.9)	15,348 (98.6)	213 (1.4)	
Estimated blood loss, mL, median (IQR)	100 (50–200)	100 (50–200)	200 (100–350)	<.001
Surgical complexity, total RVU, mean (SD)	26.7 (14.1)	26.5 (13.9)	31.8 (21.7)	<.001
Operative time, h, median (SD)	2.2 (1.3)	2.1 (1)	2.5 (1.3)	<.001

Variable	Overall N=22,358	Surgical Site Infection (any)		<i>P</i>
		Absent N=22,917	Present N=441	
Antibiotic Type				
Beta-lactam antibiotics, N (%)	17,827 (83.5)	17,498 (98.2)	329 (1.8)	<.001
Beta-lactam alternatives, N (%)	2,878 (13.5)	2,790 (96.9)	88 (3.1)	
Non-standard	653 (3)	629 (96.3)	24 (3.7)	

ASA, American Society of Anesthesiologists; BMI, body mass index; PCN, penicillin; RVU, relative value units.

* Laparoscopic, vaginal, and robotic hysterectomy.

Table 2

Baseline Comparison of Characteristics Between the Antibiotic Groups

Variable	Beta-lactam antibiotics N=17,827	Beta-lactam alternatives N=2,878	Non-standard N=653	P
Demographics and Comorbidities				
Age, y, median (SD)	48 (11.4)	48.5 (12.3)	48 (12.2)	.09
BMI, kg/m ² , mean (SD)	30.8 (8)	31.6 (8)	31.7 (9)	.001
Diabetes present, N (%)	1,461 (8.2)	305 (10.6)	68 (10.4)	.001
Tobacco user, N (%)	4,088 (22.9)	730 (25.4)	173 (26.5)	.003
ASA class 3, N (%)	3,632 (20.4)	744 (25.9)	170 (26)	<.001
History of hypertension, N (%)	5,245 (29.4)	931 (32.3)	209 (32)	.03
Preoperative transfusion, N (%)	122 (0.7)	23 (0.8)	1 (0.2)	.1
Gynecologic cancer, N (%)	1,624 (9.1)	310 (10.8)	63 (9.6)	<.001
Perioperative Factors				
Surgical approach, open, N (%)	4,880 (27.4)	757 (26.3)	160 (24.5)	<.001
Estimated blood loss, mL, median (IQR)	100 (50–200)	100 (50–200)	100 (50–199)	.6
Surgical complexity, total RVU, mean (SD)	25.9 (12.8)	26.8 (13.4)	25.4 (13.4)	<.001
Operative time, h, mean (SD)	2.2 (1)	2.2 (1)	2 (1)	.2
Surgical Site Infection (unadjusted)				
Any, N (%)	329 (1.8)	88 (3.1)	24 (3.7)	<.001
Superficial, N (%)	165 (0.9)	41 (1.4)	15 (2.3)	<.001
Deep–organ, N (%)	167 (0.9)	47 (1.6)	10 (1.5)	.003

ASA, American Society of Anesthesiologists; BMI, body mass index; IQR, interquartile range; PCN, penicillin; RVU, relative value units.

Table 3

Logistic Regression Model: Independent Predictors of Surgical Site Infection)

Variable adjusted for in logistic regression model	Any SSI				Superficial SSI				Deep SSI			
	Unadjusted OR	Adjusted OR	95% CI	P	Unadjusted OR	Adjusted OR	95% CI	P	Unadjusted OR	Adjusted OR	95% CI	P
Antibiotic category												
Beta-lactam antibiotics	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref
Beta-lactam alternatives	1.7	1.62	1.27–2.07	<.001	1.6	1.5	1.04–2.09	.03	1.8	1.7	1.2–2.4	<.001
Non-Standard	2.1	2.02	1.31–3.1	<.001	2.5	2.5	1.46–4.34	.001	1.7	1.6	0.8–3.1	.1
Surgical time (per hour)	1.3	1.23	1.14–1.33	<.001	1.3	1.14	0.98–1.3	.08	1.3	1.26	1.14–1.39	<.001
BMI												
<30 kg/m ² nonobese	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref
30 kg/m ² obese	1.8	1.5	1.2–1.9	<.001	2.3	1.8	1.2–2.7	<.001	1.4	1.2	0.8–1.6	.2
Smoking status												
Nonsmoker	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref
Smoker		1.46	1.18–1.8	<.001	1.5	1.7	1.2–2.2	<.001	1.2	1.27	0.9–1.7	.14
ASA category												
<3	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref
3	1.8	1.12	0.9–1.4	.3	2.2	1.2	0.8–1.7	.5	1.4	1.1	0.8–1.5	.5
Surgical complexity (per RVU)	1.02	1.01	0.9–1.01	.13	1.02	1.01	0.9–1.02	.1	1.01	1.002	0.9–1.01	.9
Diabetes												
Absent	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref
Present	1.8	1.36	1.05–1.76	.02	2.2	1.5	1.04–2.2	.03	1.4	1.3	0.8–1.8	.24
Final pathology												
Benign	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref
Cancer	2.4	1.7	1.3–2.2	<.001	3	1.8	1.3–2.6	.01	1.8	1.5	0.9–2.5	.08
Surgical route												

Variable adjusted for in logistic regression model	Any SSI						Superficial SSI						Deep SSI					
	Unadjusted OR	Adjusted OR	95% CI	P	Unadjusted OR	Adjusted OR	95% CI	P	Unadjusted OR	Adjusted OR	95% CI	P	Unadjusted OR	Adjusted OR	95% CI	P	Unadjusted OR	Adjusted OR
	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref	Ref
MIS	3	2.6	2.1–3.1	<.001	4.2	3.5	2.6–5	<.001	2	1.9	1.37–2.6	<.001	2	1.9	1.37–2.6	<.001	2	1.9
Open																		

ASA, American Society of Anesthesiologists; BMI, body mass index; CI, confidence interval; MIS, minimally invasive surgery (vaginal, laparoscopic, and robotic hysterectomies); OR, odds ratio; PCN, penicillin; Ref, referent category; RVU, relative value units.