

**ORIGINAL CONTRIBUTION**

**Textbook Outcome After Ileal Pouch–Anal Anastomosis for Ulcerative Colitis: A Nationwide Multicenter Study and Its Implications for Long-Term Quality of Life**

**Running head:** Textbook Outcomes in Surgery for Ulcerative colitis

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**Funding/Support:** None reported.

**Financial Disclosures:** None reported.

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**ACKNOWLEDGMENTS:** The authors express their sincere gratitude to all members of the Japan Society of Laparoscopic Colorectal Surgery Collaborative Group for their invaluable contributions to this study.

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ACCEPTED

## **ABSTRACT**

**BACKGROUND:** Textbook outcome is a composite quality measure increasingly used in oncologic surgery but has not been applied to ulcerative colitis. Restorative proctocolectomy with ileal pouch–anal anastomosis is the standard surgical treatment for ulcerative colitis, yet a comprehensive assessment of surgical quality remains lacking. Pouchitis, the most common long-term complication after ileal pouch–anal anastomosis, significantly impairs quality of life and may lead to pouch failure. Linking perioperative success to long-term pouch function is therefore clinically relevant.

**OBJECTIVE:** To evaluate the feasibility and clinical relevance of textbook outcome as a perioperative quality measure and its association with long-term pouchitis after ileal pouch–anal anastomosis for ulcerative colitis.

**DESIGN:** Nationwide multicenter retrospective cohort study.

**SETTING:** Japan’s collaborative ulcerative colitis surgery registry including high-volume academic and affiliated hospitals.

**PATIENTS:** A total of 1,109 patients undergoing ileal pouch–anal anastomosis for ulcerative colitis between 2005 and 2019.

**MAIN OUTCOME MEASURES:** Textbook outcome was defined as absence of 90-day mortality, major complications (Clavien–Dindo  $\geq$  III), reoperation, readmission, and prolonged hospital stay. Logistic regression identified predictors of textbook outcome; Cox regression evaluated its association with pouchitis.

**RESULTS:** Textbook outcome was achieved in 57% of patients. In multivariable logistic regression, male sex (OR 0.74, 95% CI: 0.57–0.95), higher preoperative albumin (OR 1.22, 95% CI: 1.01–1.47), and the presence of UC-associated neoplasia (OR 1.48, 95% CI: 1.03–2.14) were

independently associated with achieving textbook outcome. Textbook outcome achievement increased significantly over time (OR per year 1.04, 95% CI: 1.01–1.07), coinciding with greater adoption of minimally invasive surgery. Failure to achieve textbook outcome was independently associated with higher pouchitis risk (HR 1.67, 95% CI: 1.12–2.49).

**LIMITATIONS:** Retrospective design and registry-based data may involve unmeasured confounders.

**CONCLUSIONS:** Textbook outcome is a feasible and clinically meaningful quality metric in ulcerative colitis ileal pouch–anal anastomosis surgery. Its achievement, improving over time with minimally invasive techniques, correlates with reduced pouchitis risk and may serve as a benchmark for surgical quality and outcomes in ulcerative colitis. (See **Video Abstract**)

**KEY WORDS:** Ileal pouch anal anastomosis; Pouchitis; Textbook outcome; Ulcerative colitis.

**RESULTADO TÍPICO TRAS UNA ANASTOMOSIS ILEOANAL CON BOLSA ILEAL PARA LA COLITIS ULCEROSA: UN ESTUDIO MULTICÉNTRICO A NIVEL NACIONAL Y SUS IMPLICACIONES PARA LA CALIDAD DE VIDA A LARGO PLAZO.**

**ANTECEDENTES:** El resultado del libro de texto es una medida de calidad compuesta que se utiliza cada vez más en la cirugía oncológica, pero que no se ha aplicado a la colitis ulcerosa. La proctocolectomía restauradora con anastomosis ileoanal es el tratamiento quirúrgico estándar para la colitis ulcerosa, pero aún no se ha realizado una evaluación exhaustiva de la calidad quirúrgica. La pouchitis, la complicación a largo plazo más común tras la anastomosis ileoanal, deteriora significativamente la calidad de vida y puede provocar el fallo de la bolsa. Por lo tanto, relacionar el éxito perioperatorio con la función a largo plazo de la bolsa es clínicamente relevante.

**OBJETIVO:** Evaluar la viabilidad y la relevancia clínica del resultado teórico como medida de calidad perioperatoria y su asociación con la pouchitis a largo plazo tras la anastomosis ileoanal para la colitis ulcerosa.

**DISEÑO:** Estudio de cohorte retrospectivo multicéntrico a nivel nacional.

**ENTORNO:** Registro colaborativo japonés de cirugía de colitis ulcerosa, que incluye hospitales académicos y afiliados de gran volumen.

**PACIENTES:** Un total de 1109 pacientes sometidos a anastomosis ileoanal por colitis ulcerosa entre 2005 y 2019.

**PRINCIPALES MEDIDAS DE RESULTADO:** El resultado teórico se definió como la ausencia de mortalidad a los 90 días, complicaciones graves (Clavien-Dindo  $\geq$  III),

reintervención, reingreso y estancia hospitalaria prolongada. La regresión logística identificó los predictores del resultado ideal; la regresión de Cox evaluó su asociación con la pouchitis.

**RESULTADOS:** Se logró el resultado ideal en el 57 % de los pacientes. En la regresión logística multivariable, el sexo masculino (OR 0,74, IC del 95 %: 0,57-0,95), un nivel más alto de albúmina preoperatoria (OR 1,22, IC del 95 %: 1,01-1,47) y la presencia de neoplasia asociada a la CU (OR 1,48, IC del 95 %: 1,03-2,14) se asociaron de forma independiente con el logro del resultado ideal. El logro de resultados óptimos aumentó significativamente con el tiempo (OR por año 1,04, IC del 95 %: 1,01-1,07), coincidiendo con una mayor adopción de la cirugía mínimamente invasiva. El hecho de no lograr resultados óptimos se asoció de forma independiente con un mayor riesgo de pouchitis (HR 1,67, IC del 95 %: 1,12-2,49).

**LIMITACIONES:** El diseño retrospectivo y los datos basados en registros pueden implicar factores de confusión no medidos.

**CONCLUSIONES:** El resultado ideal es una medida de calidad viable y clínicamente significativa en la cirugía de anastomosis ileoanal con bolsa ileal para la colitis ulcerosa. Su consecución, que mejora con el tiempo gracias a las técnicas mínimamente invasivas, se correlaciona con un menor riesgo de pouchitis y puede servir como referencia para la calidad quirúrgica y los resultados en la colitis ulcerosa. (*AI-generated translation*)

## INTRODUCTION

Ulcerative colitis (UC) is a chronic inflammatory condition of the colon with a rising global incidence, including in US, Europe and Asia.<sup>1-3</sup> Although most patients can be managed with medical therapy, surgery remains indicated for those with refractory disease, dysplasia, or neoplasia. Restorative proctocolectomy with ileal pouch–anal anastomosis (IPAA) is the standard surgical treatment, offering both disease control and continence preservation.<sup>4,5</sup> However, the procedure is technically complex and carries risks of short-term morbidity as well as long-term pouch-related complications.<sup>4,5</sup>

In oncologic surgery, textbook outcome (TO) has emerged as a composite quality indicator that consolidates endpoints such as mortality, complications, and readmission into a single benchmark.<sup>6-8</sup> In contrast, outcomes following surgery for UC are usually reported as isolated parameters, limiting assessment of overall care quality. Although minimally invasive surgery (MIS) is increasingly adopted in UC and has demonstrated favorable perioperative outcomes,<sup>9,10</sup> its broader impact—together with other perioperative factors—on comprehensive surgical quality has yet to be examined.

Long-term pouch-related complications remain a significant concern following IPAA. Among these, pouchitis is the most common late complication, capable of markedly reducing quality of life and, in some cases, leading to pouch failure.<sup>11,12</sup> Although several large Western and Japanese series have described short-term outcomes after IPAA,<sup>13-15</sup> the relationship between comprehensive perioperative success and long-term pouch health has not been clarified.

Therefore, the objectives of this study were to (i) determine the rate of TO achievement following IPAA for UC in a large nationwide multicenter cohort from Japan, (ii) identify factors

associated with TO, and (iii) examine the clinical implications of TO achievement, with particular emphasis on pouchitis.

## **MATERIAL AND METHODS**

### **Study Design and Data Source**

This retrospective cohort study utilized a nationwide multicenter registry of UC surgery in Japan, which gathers perioperative and long-term outcomes from tertiary referral hospitals. Ethical approval was obtained at all participating institutions, and the need for individual consent was waived owing to the study's retrospective design.

### **Patients**

All consecutive patients who underwent restorative proctocolectomy with IPAA for UC between 2005 and 2019 were included. Both single-stage and staged procedures were eligible. Patients who did not undergo pouch construction or had incomplete data were excluded. This cross-sectional study has been reported in line with the STROBE guidelines.

### **Definition of Textbook Outcome**

TO was defined as the absence of all of the following adverse events: 90-day mortality, major postoperative complications (Clavien–Dindo grade  $\geq$ III), reoperation within 90 days, readmission within 90 days, and prolonged hospital stay (above the cohort's 75th percentile).

This definition was adapted from oncologic surgery literature and tailored for UC.

### **Outcomes and Variables**

The primary endpoint was TO achievement. In addition to evaluating TO as the primary endpoint, our primary analysis examined predictors of achieving TO. The secondary endpoint was the incidence of pouchitis during follow-up. Covariates included age, sex, BMI, disease duration, preoperative albumin, ASA class, surgical approach (open vs MIS), emergency status,

and UC-associated neoplasia. Pouchitis was defined as episodes requiring hospitalization, as captured in the registry.

### **Statistical Analysis**

Continuous variables were expressed as medians (IQR) and compared with the Wilcoxon rank-sum test; categorical variables were compared with chi-squared or Fisher's exact tests.

Univariable and multivariable logistic regression identified predictors of TO. Pouchitis incidence was assessed using Kaplan–Meier and Cox regression analyses. Trends in TO over time were examined with logistic regression using surgical year as a continuous variable. A  $p$  value  $< 0.05$  was considered significant. Analyses were conducted in R (version 4.2.0).

## **RESULTS**

### **Baseline Characteristics**

A total of 1,109 patients (97.4%) who underwent IPAA for ulcerative colitis between 2005 and 2019 were analyzed (Fig. 1). Median age was 42 years (IQR, 29–56), and median BMI was 19.8 kg/m<sup>2</sup> (IQR, 17.7–22.4). Of these, 450 (40.6%) were female and 250 (22.5%) had UC-associated neoplasia (UCAN). Preoperative albumin, ASA-PS classification, urgency, and surgical approach are summarized in Table 1. Overall, 636 patients (57.3%) achieved TO. Stratified analyses (Table 2) showed that non-TO patients were more often male, had lower albumin, and underwent emergency surgery more frequently, whereas UCAN was more common among TO achievers. In the entire cohort, prolonged hospital stay occurred in 24.0%, major complications (Clavien–Dindo  $\geq$ III) in 19.9%, readmission in 14.2%, reoperation in 6.5%, and 90-day mortality in 0.3%. Figure 2 illustrates TO determinants, with prolonged stay, complications, and mortality contributing most to TO failure. TO achievement increased significantly over time (OR, 1.04 per year; 95% CI, 1.01–1.07;  $p = 0.02$ ; Fig. 3). MIS was associated with higher TO rates than open

surgery, and its use rose steadily across the study period (Supplemental Figure 1A and 1B at <http://links.lww.com/DCR/C600>).

### **Predictors of Textbook Outcome**

In univariate analyses, male sex, open surgery, lower preoperative albumin, emergency presentation, higher ASA-PS, and absence of UC-associated neoplasia were significantly associated with non-achievement of TO. In multivariable logistic regression, female sex (OR 0.74, 95% CI: 0.57–0.95), higher preoperative albumin (OR 1.22, 95% CI: 1.01–1.47), and the presence of UC-associated neoplasia (OR 1.48, 95% CI: 1.03–2.14) were independently associated with achieving a Textbook Outcome. Conversely, male sex, lower albumin levels, and the absence of UC-associated neoplasia were related to TO non-achievement. Surgical approach (MIS vs open) demonstrated borderline significance in multivariable analysis ( $p = 0.06$ ; Table 3).

### **Association Between TO and Pouchitis**

TO achievement was significantly associated with a lower incidence of pouchitis, defined as episodes requiring hospitalization (Fig. 4A). In univariate analysis, TO non-achievement ( $p = 0.024$ ) and younger age ( $p < 0.01$ ) emerged as significant risk factors; both remained independent predictors in multivariable analysis (Table 4). The cumulative incidence of pouchitis was higher among non-TO patients, with the greatest divergence observed in the early postoperative period (Fig. 4B; Supplemental Fig. 2 at <http://links.lww.com/DCR/C601>).

## **DISCUSSION**

This nationwide multicenter study offers three key insights into surgical quality after IPAA for UC. First, it establishes the feasibility of applying TO to UC surgery—a benign but technically demanding setting where conventional oncologic benchmarks are less applicable. Second, TO achievement increased significantly over time, paralleling the adoption of MIS and reflecting

improvements in perioperative care. Third, TO non-achievement was independently associated with a higher risk of pouchitis, linking perioperative success to long-term pouch health. Although TO is well established in oncologic surgery,<sup>8,16,17</sup> its application in benign but technically complex procedures has been limited, with only a few recent reports in transplantation.<sup>18,19</sup> A recent NSQIP-based abstract reported TO rates for several IBD procedures using a 30-day composite definition.<sup>20</sup> While this provides an important early benchmark for short-term perioperative quality, it did not include postoperative mortality or long-term functional outcomes such as pouchitis. In contrast, our nationwide study applied a comprehensive 90-day TO definition and demonstrated an association between perioperative quality and long-term pouch outcomes after IPAA.

Restorative proctocolectomy with IPAA, however, is a demanding operation associated with considerable perioperative morbidity, making comprehensive assessment of surgical quality equally important. With the incidence of UC continuing to rise in both Western countries and Asia, including Japan,<sup>1,2</sup> the surgical burden is expected to grow, reinforcing the need for robust quality metrics such as TO. This study provides the first nationwide multicenter evaluation of TO in UC IPAA, revealing that only 57% of patients achieved TO, thereby highlighting substantial opportunities for quality improvement in this high-risk procedure.

The adoption of MIS for UC has grown rapidly over the past two decades.<sup>18,19</sup> In our nationwide cohort, TO rates rose in parallel with MIS uptake, reflecting temporal improvements in perioperative quality. Although MIS was significantly associated with TO in univariable analysis, its effect diminished in multivariable analysis, reaching only marginal significance. This may be explained by the steep learning curve of laparoscopic IPAA,<sup>21</sup> institutional differences in expertise, and case selection bias, as more complex or urgent cases were often

managed with open surgery. Nonetheless, the temporal trend strongly suggests that MIS has contributed to improved outcomes. Looking ahead, early data<sup>22</sup> indicate that robotic-assisted IPAA may further improve perioperative quality by reducing conversion rates and blood loss, though cost and training challenges remain.

Pouchitis is the most common long-term complication after IPAA, with significant effects on quality of life and pouch survival.<sup>12,23</sup> Large cohorts from Europe, North America, and Japan have reported functional outcomes and risk factors for pouch failure.<sup>24–26</sup> In our study, pouchitis requiring hospitalization occurred in ~6% of patients and was independently linked to TO non-achievement. Although hospitalization for pouchitis may be less frequent in some healthcare systems, this stringent definition ensured consistent identification of clinically significant episodes in our cohort. Perioperative morbidity, such as pelvic sepsis or anastomotic leakage, may initiate chronic inflammation or structural damage, increasing susceptibility to later pouchitis. Prior studies have identified pelvic sepsis and anastomotic complications as strong predictors of pouch dysfunction and failure,<sup>27,28</sup> and large series have confirmed their detrimental impact on pouch survival.<sup>29,30</sup> Although this mechanism remains theoretical, our findings support the view that early comprehensive perioperative success influences long-term pouch health. Notably, the divergence in pouchitis risk between TO and non-TO patients was most pronounced in the early postoperative period, with hazards converging over time, suggesting that perioperative recovery quality may primarily influence early susceptibility rather than the long-term phenotype of chronic pouchitis. These results suggest that beyond isolated complications, the cumulative burden of perioperative events—captured by TO—affects long-term outcomes, underscoring TO's value as a holistic quality metric. Pouchitis encompasses heterogeneous clinical patterns, including early-onset and chronic forms, each of which may arise through

distinct biological pathways. While perioperative factors are likely to exert greater influence during the early postoperative period, later or chronic pouchitis may involve mechanisms that extend beyond perioperative recovery alone. This distinction provides additional context for understanding how perioperative quality relates to the long-term risk of clinically meaningful pouchitis.

This study has limitations. Its retrospective design introduces potential unmeasured confounders, and institutional differences in surgical practice and perioperative management may have affected outcomes. The TO definition, adapted from oncologic surgery, may not fully capture quality in UC. Finally, the observed effect of MIS should be interpreted with caution, as early adoption, learning curves, and selection bias could have influenced the results.

## **CONCLUSIONS**

TO was achieved in over half of UC IPAA patients and correlated with improved long-term pouch outcomes. Its rising rates alongside the adoption of MIS highlight TO's value as a benchmark for surgical quality. These results support TO as a comprehensive metric in UC surgery, bridging perioperative success with long-term, patient-centered outcomes.

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## LEGEND

1. Study flow diagram. From 2005 to 2019, 1,109 patients who underwent ileal pouch–anal anastomosis for ulcerative colitis were included after applying exclusion criteria.
2. Distribution of textbook outcome (TO) parameters. Bar graphs show the individual achievement rates for each element (90-day mortality, major complications, reoperation, readmission, and prolonged length of stay), with the overlaid line indicating cumulative achievement rates as conditions are added sequentially. The accompanying table summarizes both the *Population (%)*, defined as the proportion of patients fulfilling each element individually, and the *Conditional (%)*, defined as the cumulative proportion of patients fulfilling all elements sequentially up to that point.
3. Annual trends in textbook outcome (TO) achievement. Scatter plot with regression line demonstrating the rise in TO rates over time. Logistic regression confirmed a significant upward trend (OR, 1.04 per year (95% CI, 1.01–1.07),  $p = 0.02$ ).
4. Association between textbook outcome (TO) and pouchitis. (A) Bar plot showing the ratio for pouchitis. (B) Kaplan–Meier curves of pouchitis incidence stratified by TO status, showing significantly higher pouchitis in non-TO patients.

Table 1. Baseline characteristics of the study cohort (n = 1,109).

Variable	Level	Overall (n=1109)
Age, years		42 [29–56]
BMI, kg/m <sup>2</sup>		19.8 [17.7–22.4]
Albumin, g/dL		3.4 [2.6–4.0]
Disease duration, years		5 [1–12]
Gender	Female	450 (40.6%)
	Male	659 (59.4%)
Steroid use	No	484 (43.6%)
	Yes	625 (56.4%)
Immunosuppressive therapy	No	482 (43.6%)
	Yes	624 (56.4%)
Emergency surgery	No	805 (72.6%)
	Yes	304 (27.4%)
Surgical approach	MIS	780 (70.3%)
	Open	329 (29.7%)
ASA class	Low (0–1)	243 (21.9%)
	High (≥2)	866 (78.1%)
UC-associated neoplasia	No	859 (77.5%)
	Yes	250 (22.5%)
Surgery stage	1	79 (7.1%)
	2	806 (72.7%)
	3	224 (20.2%)
Pouchitis requiring admission	No	1041 (93.9%)
	Yes	68 (6.1%)
Ileus requiring admission	No	896 (80.8%)
	Yes	213 (19.2%)
Ileus requiring reoperation	No	158 (74.2%)
	Yes	55 (25.8%)

Table 2. Comparison of patient characteristics stratified by TO achievement.

Variable	Level	Overall	non-TO	TO (n=636)	p value
Age, years		42.0 [29.0–56.0]	42.0 [29.0–	42.0 [30.0–	0.57
BMI, kg/m <sup>2</sup>		19.8 [17.7–22.4]	19.5 [17.4–	20.1 [17.9–	0.11
Gender	Female	450 (40.6%)	173	277	<b>0.02</b>
	Male	659 (59.4%)	300	359	
Disease duration, years		5.0 [1.0–	5.0 [1.0–	5.0 [2.0–	0.22
Albumin, g/dL		3.4 [2.6–	3.3 [2.5–	3.5 [2.7–	<b>&lt;0.001</b>
Steroid use	No	484 (43.6%)	192	292	0.09
	Yes	625 (56.4%)	281	344	
Immunosuppressive therapy	No	482 (43.6%)	206	276	1
	Yes	624 (56.4%)	266	358	
Emergency surgery	No	805 (72.6%)	328	477	<b>0.04</b>
	Yes	304 (27.4%)	145	159	
Surgical approach	MIS	780 (70.3%)	312	468	<b>0.01</b>
	Open	329 (29.7%)	161	168	
ASA class	Low(0-1)	243 (21.9%)	86	157	<b>0.01</b>
	High(2≤)	866 (78.1%)	387	479	
UC-associated neoplasia	No	859 (77.5%)	391	468	<b>&lt;0.001</b>
	Yes	250 (22.5%)	82	168	
Pouchitis requiring admission	No	1041	435	606	<b>0.03</b>
	Yes	68	38	30	
Ileus requiring admission	No	896 (80.8%)	373	523	0.18
	Yes	213 (19.2%)	100	113	
Surgery stage	1	79	34	45	0.06
	2	806 (72.7%)	328	478	
	3	224 (20.2%)	111	113	
Ileus requiring reoperation	No	158 (74.2%)	75	83	0.92
	Yes	55	25	30	

Values are presented as median [IQR] or frequency (proportion).

Table 3. Univariate and multivariate logistic regression for TO achievement. (OR > 1 = protective, OR < 1 = risk)

Variable	Level /	Univariate		Multivariate		Interpretation
		OR [95% CI]	p-	OR [95% CI]	p-	
Age (per year)	–	1.00	0.53	1.00 [0.99–1.00]	0.57	—
Male sex	Female	0.75	0.02	0.74 [0.57–0.96]	<b>0.02</b>	Risk
BMI (per kg/m <sup>2</sup> )	–	1.01	0.54	1.00 [0.96–1.03]	0.83	—
Disease duration	–	1.01	0.12	0.99 [0.97–1.01]	0.26	—
Open surgery	MIS	0.70	0.01	0.79 [0.60–1.03]	0.1	—
Albumin (per g/dL)	–	1.30	<0.001	1.22 [1.01–1.47]	<b>0.04</b>	Protective
Emergency surgery	Elective	0.75	0.04	1.13 [0.82–1.57]	0.47	—
ASA High (≥2)	Low (0–1)	0.61	0.01	0.74 [0.51–1.07]	0.11	—
UC-associated	No tumor	1.71	<0.001	1.48 [1.03–2.11]	<b>0.04</b>	Protective

Logistic Regression Analysis, Forced entry

Table 4. Risk factors for pouchitis after IPAA by univariate and multivariate analysis.

Variable	Reference	Univariate		Multivariate	
		OR [95%	p-	OR [95%	p-
Age (per year)	—	0.97 [0.96–0.99]	<0.01	0.97 [0.96–	<b>&lt;0.01</b>
Gender Female	Male	1.11 [0.67–1.86]	0.69	1.13 [0.68–	0.64
BMI (per kg/m <sup>2</sup> )	—	0.94 [0.88–1.01]	0.1	0.98 [0.91–	0.63
Disease duration (per	—	0.98 [0.94–1.01]	0.15	1.00 [0.96–	0.93
Surgical approach Open	MIS	1.32 [0.78–2.18]	0.3	1.09 [0.62–	0.76
Albumin (per g/dL)	—	0.89 [0.67–1.18]	0.42	0.93 [0.64–	0.71
Emergency surgery	Elective	1.29 [0.75–2.15]	0.35	1.05 [0.54–	0.88
ASA (High $\geq$ 2)	Low (0–1)	1.30 [0.63–2.44]	0.45	1.09 [0.50–	0.82
UC-associated	No tumor	0.58 [0.27–1.09]	0.11	0.92 [0.39–	0.85
TO achieved	No achieved	0.57 [0.34–0.93]	0.02	0.60 [0.36–	<b>0.049</b>

Figure 1

Figure 1. A flow chart of the patients included

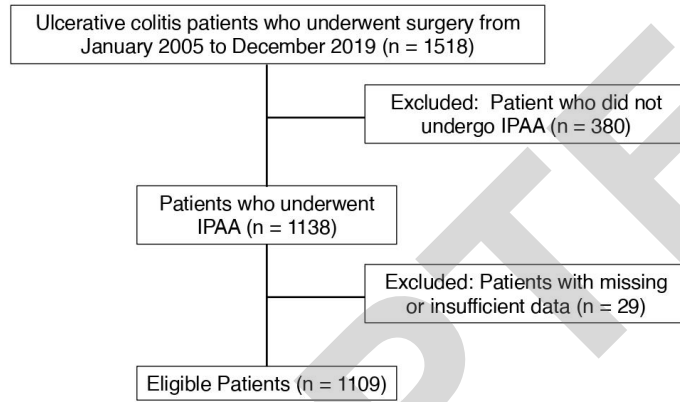
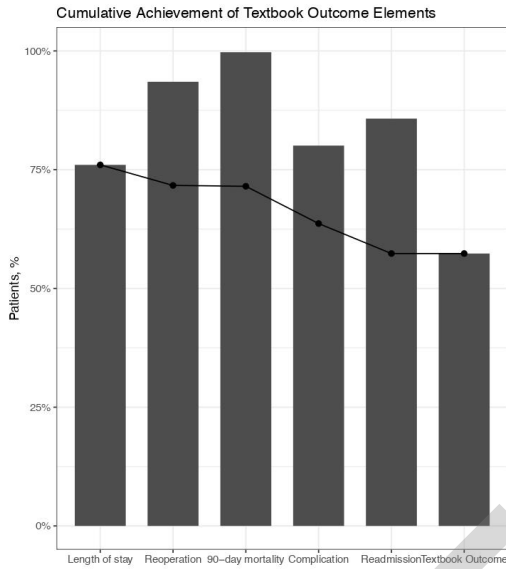


Figure 2

Figure 2. Cumulative bar plot



	Population (%)	Conditional (%)
Length of stay	76.0	76.0
Reoperation	93.5	71.7
90-day mortality	99.7	71.5
Complication	80.1	63.7
Readmission	85.8	57.3
Textbook outcome	57.3	57.3

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Figure 3

Figure 3. TO rate across years

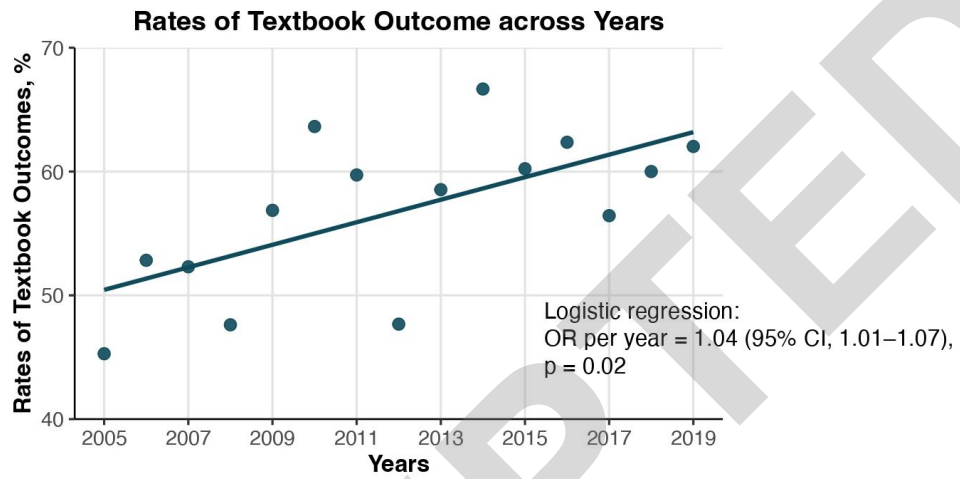


Figure 4

Figure 4.

