

Operative timing is associated with increased morbidity and mortality in patients undergoing emergency general surgery: a multisite study of emergency general services in a single academic network

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Background: Despite the widespread implementation of the acute care surgery (ACS) model, limited access to operating room time represents a barrier to the optimal delivery of emergency general surgery (EGS) care. The objective of this study was to describe the effect of operative timing on outcomes in EGS in a network of teaching hospitals.

Methods: We conducted a retrospective review of EGS operations performed at 3 teaching hospitals in a single academic network. Time of operation was categorized as daytime (8 am to 5 pm), after hours (5 pm to 11 pm) or overnight (11 pm to 8 am). Time to operation was calculated as the interval from admission to operative start time and categorized as less than 24 hours, 24–72 hours and greater than 72 hours.

Results: After we excluded nonindex cases, trauma cases and cases occurring more than 5 days after admission, 1505 EGS cases were included. We found that 39.0% of operations were performed in the daytime, 46.3% after hours and 14.8% overnight. In terms of time to operation, 52.3% of operations were performed within 24 hours of admission, 33.4% in 24–72 hours and 14.3% in more than 72 hours. The overall complication rate was 20.6% (310 patients) and the overall mortality rate was 3.8% (57 patients). After multivariable analysis, time to operation more than 72 hours after admission was independently associated with increased odds of morbidity (odds ratio [OR] 1.64, 95% confidence interval [CI] 1.09–2.47), while overnight operating was associated with increased odds of death (OR 3.15, 95% CI 1.29–7.70).

Conclusion: Increasing time from admission to operation and overnight operating were associated with greater morbidity and mortality, respectively, for EGS patients. Strategies to provide timely access to the operating room should be considered to optimize care in an ACS model.

Contexte : Même si le modèle de chirurgie en soins actifs (CSA) est largement répandu, l'accès limité aux blocs opératoires représente un obstacle à la chirurgie générale chez les patients des services d'urgence (CGSU). L'objectif de cette étude était de décrire l'effet du moment de l'intervention sur l'issue des CGSU dans un réseau d'hôpitaux universitaires.

Méthodes : Nous avons procédé à une revue des CGSU effectuées dans 3 hôpitaux d'enseignement d'un réseau universitaire. Le moment opératoire était catégorisé selon que les interventions étaient effectuées le jour (8 h 00 à 17 h 00), le soir (17 h 00 à 23 h 00) ou la nuit (23 h 00 à 8 h 00). Le délai opératoire représentait l'intervalle entre l'admission et le début de l'intervention et était réparti selon les catégories suivantes : moins de 24 heures, de 24 à 72 heures et plus de 72 heures.

Résultats : Après exclusion des cas non index, des cas de traumatologie et des cas survenus plus de 5 jours après l'admission, 1505 CGSU ont été incluses. Nous avons constaté que 39,0 % des interventions avaient été effectuées le jour, 46,3 % le soir et 14,8 % la nuit. Pour ce qui est du délai opératoire, 52,3 % des interventions ont été effectuées dans les 24 heures suivant l'admission, 33,4 % dans les 24 à 72 heures et 14,3 % plus de 72 heures après l'admission. Le taux global de complications a été de 20,6 % (310 patients) et le taux de mortalité global a été de 3,8 % (57 patients). Après analyse multivariée, le délai opératoire de plus de 72 heures suivant l'admission a été associé de manière indépendante à un risque accru de morbidité (rapport ces cotes [RC] 1,64, intervalle de confiance [IC] de 95 % 1,09 à 2,47), tandis que les interventions effectuées la nuit ont été associées à un risque de décès plus élevé (RC 3,15, IC de 95 % 1,29 à 7,70).

Conclusion : L'augmentation du délai entre l'admission et l'intervention et les interventions de nuit ont été associées à une morbidité et une mortalité plus élevées, respectivement, chez les patients soumis à des CGSU. Des stratégies visant à offrir un accès rapide aux blocs opératoires sont à envisager pour optimiser le modèle de CSA.

The advent of the acute care surgery (ACS) model brought the promise of improved outcomes for the emergency general surgery (EGS) patient population. Over the last decade, extensive research has highlighted the benefits of the ACS model for patient- and hospital-specific outcomes.¹⁻⁷

Despite the widespread implementation of the ACS model in Canada, lack of access to the operating room is a common issue facing many EGS services across the country. A recent national review of EGS services in Canada found that less than half of hospitals that employ the ACS model have dedicated access to the OR for EGS patients during the day. Not surprisingly, nearly 50% of all EGS procedures in Canada continue to take place after hours or overnight, between 5 pm and 8 am.⁸

The impacts of after-hours operating and delay to operative intervention have been described in the literature. For hip fractures, for example, studies have shown that after-hours operating and operative delay lead to increased morbidity and mortality.^{9,10} Furthermore, implementation of dedicated daytime operating room resources to address hip fractures has resulted in significant improvements in disease-specific outcomes, in part by limiting operative delay and overnight operating.^{11,12}

Similar research in EGS is less conclusive. Multiple studies have examined the impact of after-hours operating on appendectomy and cholecystectomy outcomes, with mixed results.¹³⁻¹⁸ Delay to operative intervention has been difficult to study because of the diversity of disease processes, severity of presentation and variable comorbidities in EGS patients.

Despite the implementation of ACS models in our academic network, access to the OR represents a substantial barrier to timeliness of care for our EGS patients. The purpose of this study was to describe the impact of operative timing on EGS outcomes in our network. Our primary objectives were to determine the association, if any, of time elapsed from admission to operation, with morbidity and mortality of EGS patients and to determine the association, if any, of timing of operation (day, after hours, overnight) with morbidity and mortality of EGS patients. Secondarily, we sought to characterize the timing of operation of EGS patients in our centre and to obtain detailed information on disease distribution.

METHODS

Study design and data collection

This study was a retrospective review of all EGS operative cases performed throughout the 3 adult teaching hospitals in our academic health sciences network (Hamilton Health Sciences and St. Joseph's Healthcare Hamilton). Data were collected on all EGS cases performed from Jan. 1 to Dec. 31, 2014. Cases were identified through each hospital's operating room administrative data set. Data were

derived through this data set and electronic chart review. Ethics approval was obtained from the Hamilton Integrated Research Ethics Board.

Context

Each of the 3 sites used a different EGS model of care: an ACS model with allocated daytime operating room access, an ACS model with no allocated daytime operating room access, or general surgeon on call (24 h) with no ACS team and no allocated daytime operating room access. In addition, each hospital serviced unique patient populations (Table 1).

Inclusion and exclusion criteria

All nonelective EGS operations performed on adult patients by the ACS service or on-call surgeon were reviewed. Nonemergent and elective semiurgent operations were excluded. In addition, we excluded all secondary cases (e.g., repeat laparotomies) and cases that were not categorized as general surgery cases. Trauma cases were excluded because the physiology and management of patients with traumatic injuries treated operatively were not comparable to those of EGS patients.

Cases in which the patient underwent an operation more than 5 days after admission were excluded. In choosing a timing cut-off, we considered that most EGS patients will require surgery within hours to days, but several situations would alter this assumption. These would include conditions requiring preoperative investigation or procedures (choledocholithiasis) or initial nonoperative management (small bowel obstruction), or situations in which the patient requires preoperative medical optimization. Most of these situations could occur within 5 days. Expanding our cut-off further would probably capture procedures performed on patients admitted for other reasons who subsequently developed an emergency surgical condition. Such patients would have different baseline characteristics than other patients, and the time from admission to operation would not be as representative as a potential marker of delayed operative management.

Sensitivity analyses

We tested the above assumptions using 3 sensitivity analyses. The first 2 altered time to operation cut-offs. First, we included all patients to test whether limiting our analysis to patients having surgery in the few days after admission produced results that differed dramatically from those produced when we included patients with prolonged admission. Second, we restricted the analysis to patients who had surgery within 10 days of admission. A third sensitivity analysis maintained the 5 day cut-off but excluded disease categories that we anticipated would be associated with a preoperative delay (choledocholithiasis, small bowel obstruction).

Table 1. Summary of patient characteristics and operative timing

Characteristic	No. (%) of patients*			
	Overall n = 1505	Site 1† (ACS, day OR) n = 548	Site 2‡ (ACS, no day OR) n = 593	Site 3§ (No ACS, no day OR) n = 364
Age, yr, mean ± SD	51.9 ± 19.35	54.1 ± 19.28	50.3 ± 19.70	51.3 ± 18.68
Female sex	819 (54.4)	303 (55.3)	332 (56.0)	184 (50.6)
Disease type				
Cholecystitis	364 (24.4)	132 (24.5)	148 (25.0)	84 (23.3)
Choledocholithiasis	63 (4.2)	19 (3.5)	34 (5.8)	10 (2.8)
Appendiceal disease	355 (23.6)	107 (19.5)	152 (25.6)	96 (26.4)
Large bowel disease	189 (12.6)	80 (14.6)	68 (11.5)	41 (11.3)
Small bowel disease	164 (10.9)	79 (14.4)	54 (9.1)	31 (8.5)
Hernia	129 (8.6)	38 (6.9)	60 (10.1)	31 (8.5)
Perianal disease	87 (5.8)	42 (7.7)	15 (2.5)	30 (8.2)
Gastric disease	46 (3.1)	12 (2.2)	25 (4.2)	9 (2.5)
Breast disease	12 (0.8)	2 (0.4)	3 (0.5)	7 (1.9)
Hepatopancreatobiliary disease	1 (0.1)	1 (0.2)	0 (0.0)	0 (0.0)
Other	80 (5.3)	27 (4.9)	32 (5.4)	21 (5.8)
Time to operation				
< 24 h	761 (52.3)	297 (54.3)	248 (44.4)	216 (61.5)
24–72 h	487 (33.4)	174 (31.8)	225 (40.3)	88 (25.1)
72–120 h	208 (14.3)	76 (13.9)	85 (15.2)	47 (13.4)
Time of operation				
Daytime	586 (39.0)	254 (46.5)	168 (28.3)	164 (45.2)
After hours	695 (46.3)	209 (38.3)	328 (55.3)	158 (43.5)
Overnight	221 (14.7)	83 (15.2)	97 (16.4)	41 (11.3)
Adverse outcomes				
Complications	310 (20.6)	109 (19.9)	118 (19.9)	83 (22.8)
In-hospital mortality	57 (3.8)	25 (4.6)	13 (2.2)	19 (5.2)

ACS = acute care surgery; OR = operating room; SD = standard deviation.
 *Unless indicated otherwise.
 †ACS initiated in 2011. Surgical specialties: oncology, hepatobiliary, colorectal, urology, arthroplasty, plastic.
 ‡ACS initiated in 2013. Surgical specialties: bariatric, thoracic, otolaryngology, obstetrics, urology.
 §No ACS. Surgical specialties: trauma, cardiac, neurosurgery, spine, vascular, orthopedics, plastics/burns.

Variables

Outcome variables included complications, defined as any postoperative adverse event occurring during hospital admission, and in-hospital mortality. Explanatory variables included the demographic characteristics age and sex. Time to operation was calculated by subtracting the date and time of the start of the operation from the date and time of admission. This was further categorized as less than 24 hours, 24–72 hours and more than 72 hours. Time of operation was categorized as daytime (Monday to Friday, 8 am to 5 pm), after hours (Monday to Friday, 5 pm to 11 pm; Saturday and Sunday, 8 am to 11 pm) or overnight (11 pm to 7 am). Other variables collected were pre- and postoperative diagnosis and operative intervention performed.

Statistical analysis

Summary and descriptive statistics were tabulated and reported as means with standard deviations for normally

distributed data. Univariable analysis was conducted comparing categorical variables using the χ^2 test or Fisher exact test. Generalized linear mixed-effects logistic regression was used to determine independent effects of preoperative factors on complications and mortality. The dependent variables were postoperative complications and mortality. Independent variables were age (by deciles), sex, disease type, time to operation (by category) and timing of operation (daytime, after hours, overnight). Certain variables were excluded from regression analysis because there were too few cases (i.e., hepatopancreatobiliary operations) or no events (i.e., no deaths among patients who underwent breast operations) or because of heterogeneity (i.e., the “other” operation category). Hospital site was used as a random effects variable to account for the potential clustering effect by hospital. The variables were selected for forced inclusion; no stepwise selection process was used. Adjusted odds ratios (ORs) with 95% confidence intervals (CIs) are reported. Analysis was performed using R version 3.5.3 (R Foundation).

RESULTS

Patient characteristics

In total, 1874 nonelective EGS operative cases were identified. After we excluded 139 nonindex cases (e.g., repeat laparotomy, abdominal wall closure), 14 trauma cases, and 216 cases delayed more than 5 days after admission, 1505 cases were included in the analysis (Fig. 1). The mean age of the patient population was 53 years. Fifty-four percent of patients were women ($n = 819$). Appendectomy and cholecystectomy represented just over 50% of cases. Overall, the in-hospital complication and mortality rates were

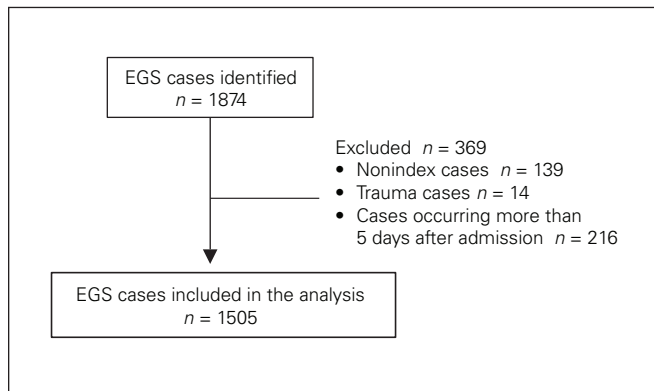


Fig. 1. Patient flowchart. EGS = emergency general surgery.

20.6% ($n = 310$) and 3.8% ($n = 57$), respectively (Table 1). Surgical site infection (SSI) was the most common complication (5.1%, $n = 77$), followed by postoperative ileus (3.6%, $n = 54$) and pneumonia (2.8%, $n = 42$) (Fig. 2).

Time to operation

In the majority of cases (52.3%), operations occurred within 24 hours of admission; operations occurred within 24–72 hours of admission and more than 72 hours after admission in 33.4% and 14.3% of cases, respectively (Table 1). The rate of in-hospital complications was greatest among cases with time to operation greater than 72 hours (17.1%, 22.2% and 32.7% for cases performed in less than 24 h, within 24–72 h and more than 72 h after admission, respectively; $p < 0.001$; Table 2). There was no difference in mortality. After multivariable analysis, time to operation greater than 72 hours was independently associated with an increased risk of morbidity (OR 1.64, 95% CI 1.09–2.47, $p = 0.02$) (Table 3, Fig. 3).

Time of operation

More operations were performed after hours (46.3% of cases, $n = 695$) than during the daytime (39.0%, $n = 586$) and overnight (14.8%, $n = 221$). Overall, on univariate

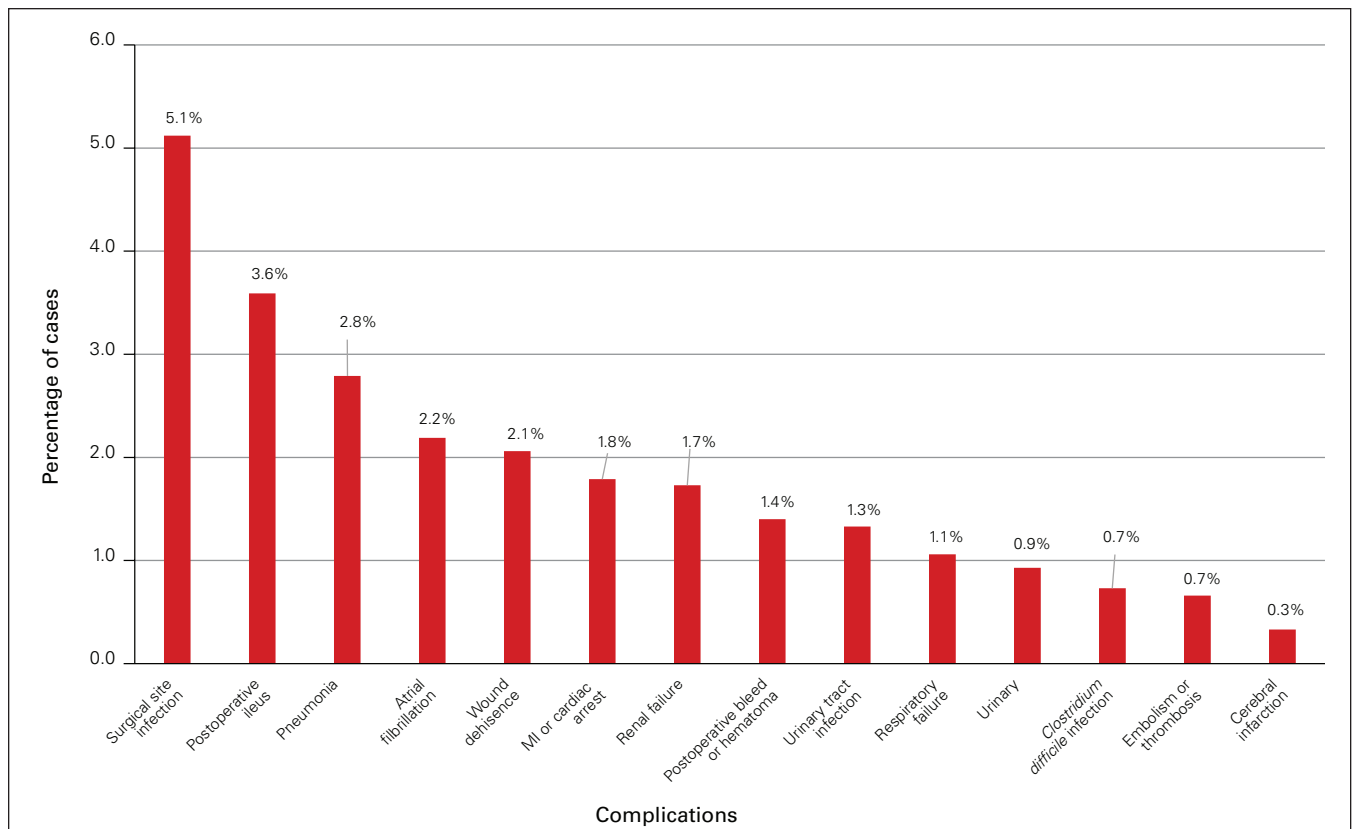


Fig. 2. In-hospital complications. MI = myocardial infarction.

analysis there was no difference in the rate of complications or mortality (Table 2). However, regression analysis revealed that overnight operations were independently associated with an increased risk of death compared with daytime operations (OR 3.15, 95% CI 1.29–7.70, $p = 0.01$) (Table 3, Fig. 3).

Multivariable analysis

In addition to the above findings, morbidity was independently associated with pathology type. With ref-

erence to cholecystitis, the cases relating to small bowel, large bowel, gastric disease and hernias were associated with an increased risk of complications (Table 3). With the exception of hernia cases, the same trend held true for mortality.

Sensitivity analyses

Details of the sensitivity analysis can be found in Appendix 1, Supplemental Tables S1 and S2 (available at canjsurg.ca/012919-a1). When we expanded the analysis to include patients who had surgery within 10 days of admission, and when we subsequently included all patients, there was a marginally increased risk of complications if the operation was performed after 72 hours (OR 1.83, 95% CI 1.28–2.62; OR 2.52, 1.83–3.46). Otherwise there was no major difference from our final model.

With regard to mortality, expansion of the inclusion criteria did not alter the main finding that overnight operations are associated with an increased risk of mortality compared with daytime operations. There was an association with timing greater than 72 hours after admission that occurred only in the model including all patients.

When we restricted the analysis to patients expected to be booked for an operation without preoperative delay, no substantive changes were seen in either the complication or mortality models (Appendix 1, Supplemental Tables S1 and S2).

Cholecystitis

In 364 cases (24.4%), the operation was performed to treat cholecystitis; 98.0% of these operations were completed laparoscopically. Time to operation was as follows: 45.5% within 24 hours of admission, 39.8% within 24–72 hours of admission and 14.8% more than 72 hours after admission (Table 4). A majority of cases occurred during the daytime (42.3%), while 44.2% took place after hours and 13.4% overnight.

The incidence of complications rose with increasing time to operation: patients experienced complications in 6.2% cases in which the operation was performed within 24 hours, in 14.3% of cases in which the operation was performed between 24 and 72 hours and in 19.2% of cases in which the operation was performed more than 72 hours after admission ($p = 0.01$). There was no difference in complication rate based on time of day ($p = 0.89$). The mortality rate was low for biliary disease, and no significant difference was found for time to operation or operative timing.

Table 2. Complications and mortality

	Complications		Mortality	
	No. (%) of patients	<i>p</i> value	No. (%) of patients	<i>p</i> value
Time to operation				
< 24 h	130 (17.1)	< 0.001	24 (3.2)	0.12
24–72 h	108 (22.2)		19 (3.9)	
> 72 h	68 (33.0)		13 (6.2)	
Time of operation				
Daytime	115 (19.6)	0.39	15 (2.6)	0.07
After hours	142 (20.4)		27 (3.9)	
Overnight	53 (24.0)		13 (5.9)	

Table 3. Regression analyses for complications and mortality

Factor	Complications		Mortality	
	OR (95% CI)	<i>p</i> value	OR (95% CI)	<i>p</i> value
Age, deciles	1.38 (1.27–1.50)	< 0.001	1.59 (1.30–1.95)	< 0.001
Male sex (Ref female)	1.26 (0.94–1.69)	0.12	0.95 (0.50–1.81)	0.88
Disease type				
Cholecystitis	Ref	—	Ref	—
Choledocholithiasis	1.06 (0.46–2.45)	0.89	1.93 (0.19–19.43)	0.58
Appendicitis	1.24 (0.75–2.02)	0.40	1.24 (0.24–6.32)	0.80
Large bowel disease	4.10 (2.59–6.49)	< 0.001	6.70 (1.87–24.10)	< 0.001
Small bowel disease	4.93 (3.05–7.95)	< 0.001	7.47 (2.03–27.50)	< 0.001
Hernia	1.87 (1.08–3.23)	0.03	2.00 (0.39–10.25)	0.41
Perianal disease	0.65 (0.26–1.62)	0.35	3.37 (0.53–21.57)	0.20
Gastric disease	5.22 (2.56–10.65)	< 0.001	18.50 (4.25–80.45)	< 0.001
Breast disease*	1.23 (0.14–10.76)	0.85	Ref	—
Time to operation				
< 24 h	Ref	—	Ref	—
24–48 h	1.24 (0.88–1.73)	0.22	0.83 (0.38–1.80)	0.64
> 72 h	1.64 (1.09–2.47)	0.02	1.21 (0.53–2.76)	0.66
Time of operation				
Daytime	Ref	—	Ref	—
After hours	1.03 (0.74–1.43)	0.86	1.53 (0.71–3.29)	0.27
Overnight	1.32 (0.86–2.05)	0.21	3.15 (1.29–7.70)	0.01

CI = Confidence interval; OR = odds ratio; Ref = reference.
*Breast disease was excluded from mortality regression because there were no postoperative mortality events.

Appendicitis

Appendectomies accounted for 355 cases (23.6%), of which 96.9% were completed laparoscopically. Time to operation was as follows: 69.9% of operations were performed within 24 hours, 22.6% within 24–72 hours and 7.4% more than 72 hours after admission (Table 4). In almost half of cases (49.3%) the operation was performed after hours (49.3%), and in the remainder of the cases the operation was performed during the daytime (33.8%) or overnight (16.9%). There was no difference in complication or mortality rates based on operative timing or time to operation.

DISCUSSION

The ACS model has been seen as a solution to many challenges in EGS.¹⁻⁷ Patients may anticipate more coordinated care, a shorter wait time for an operation and a shorter

hospital stay.¹⁻³ Hospitals hope to be able to streamline services, improve bed flow and improve cost efficiency.⁴⁻⁶ Surgeons look to the many potential benefits of ACS services, such as reduced disruption to elective practice, increased job satisfaction and perhaps even the prospect of equalization of disparities among surgeons.^{7,19} Even learners stand to benefit, given the potential for high-intensity exposure to a large volume of critically ill patients requiring operative and nonoperative management.

Although many of these improvements are worth pursuing in their own right, the core of the ACS concept has always been service to the EGS patient. A recent meta-analysis confirmed that, for biliary and appendiceal disease, this model reduces complications and length of admission.²⁰ Unfortunately, these improvements disappear without daytime access to the operating room.

Timeliness of care is paramount for the emergency surgical patient. In our patient population, operations occurring

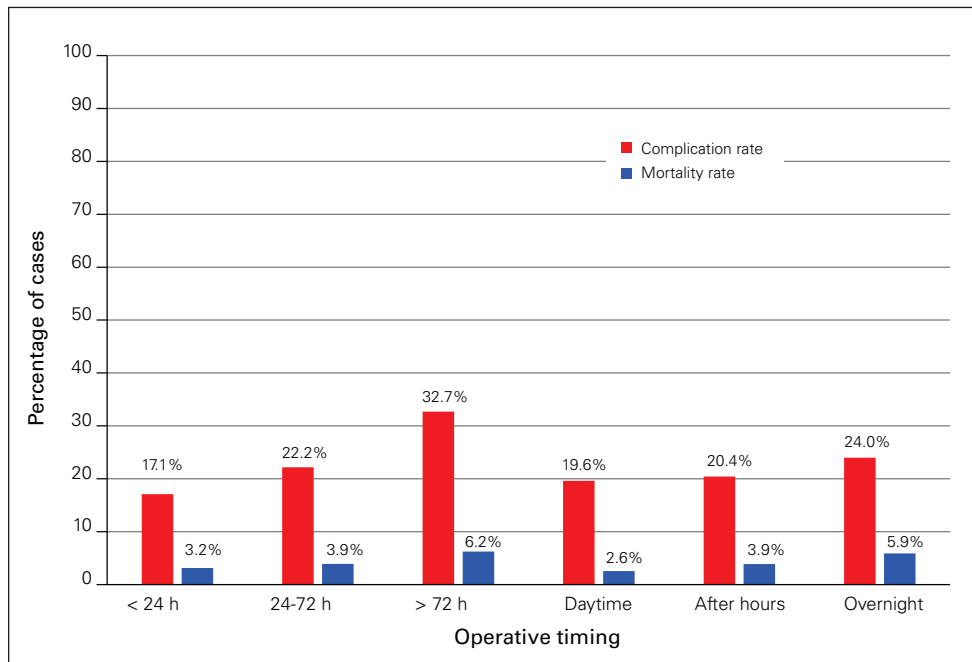


Fig. 3. Complication and mortality rates by time to operation (< 24 h, 24–72 h, > 72 h) and by time of operation (daytime, after hours, overnight).

more than 72 hours after admission were associated with an increased risk of complications, irrespective of disease process. Likewise, overnight operations were associated with a 3 times higher odds of death than daytime operations. Similar results have been previously published. In a detailed analysis, McIsaac and colleagues effectively found that delay to the OR was associated with increased hospital costs, length of stay and in-hospital mortality.²⁴ Although their study accounted for a broad range of emergency surgical conditions, from orthopedic to urologic, we were able to demonstrate that this finding holds true for the specific subset of diseases that general surgeons encounter.

Table 4. Complications and mortality in cholecystitis and appendicitis

	Time to operation				Time of operation			
	No. (%) of patients			p value	No. (%) of patients			p value
	< 24 h	24–72 h	> 72 h		Daytime	After hours	Overnight	
Cholecystitis	160 (45.4)	140 (39.8)	52 (14.8)	—	154 (42.3)	161 (44.2)	49 (13.5)	—
Complications	10 (6.2)	20 (14.3)	10 (19.2)	0.01	16 (10.4)	20 (12.4)	4 (8.2)	0.72
Mortality	1 (0.6)	0 (0)	2 (3.8)	0.06	0 (0)	3 (1.9)	0 (0)	0.36
Appendicitis	235 (69.9)	76 (22.6)	25 (7.4)	—	120 (33.8)	175 (49.3)	60 (16.9)	—
Complications	22 (9.4)	13 (17.1)	3 (12.0)	0.19	15 (12.5)	17 (9.7)	8 (13.3)	0.62
Mortality	2 (0.8)	1 (1.3)	0 (0)	0.66	1 (0.8)	1 (0.6)	1 (1.7)	0.75

In spite of the risks associated with increased time to the operating room, a substantial proportion of cases in our study occurred during higher risk time frames. Our centre is not alone; a prior study on ACS services across Canada established that competition for access to the operating room is a common problem.⁸ Monitoring is also a challenge. As outlined by a recent systematic review on the impact of the ACS model on timeliness of care and patient outcomes, a lack of standard metrics and the absence of a dedicated comprehensive EGS database limit the ability to monitor performance and identify areas for quality improvement.²¹

Although the ACS model has been a beacon of hope for addressing these issues, attention must be paid to timely access to the operating room. Other groups have found that even after-hours operating confers increased risk to patients undergoing general and vascular surgery.^{22,23} Improved daytime operating room resources would be a potential solution to both time and timeliness, and if dovetailed with the ACS model it could be advantageous for patients, providers and trainees alike.

Strengths of the present study include the diverse set of EGS delivery models, including 1 non-ACS hospital. The EGS literature is heterogeneous, typically examining either emergency surgery as a whole (general, trauma, orthopedics, cardiothoracic)²²⁻²⁴ or a single EGS pathology (perforated peptic ulcer, appendectomy, diverticulitis),²⁵⁻³¹ making it difficult to draw clear conclusions on the effect of time to operation. We were able to capture the disease processes for which the patients required surgery with improved precision and to account for this in our regression models. These factors may improve the generalizability of our results to other patients who may be encountered by the general surgeon on call, allowing for better care and empowered advocacy.

Limitations

A number of important limitations must also be acknowledged. Similar to other studies, our time to operation calculation was made using the time of admission, with the assumption that this is when the decision to operate was made. We attempted to account for this using sensitivity analyses, but these are no substitute for documented decisions. Importantly, data on patient comorbidities and physiologic status were unavailable for analysis, and these play an important role in determining timing of surgery and postoperative outcomes in this patient population. Future studies should endeavour to account for all possible relevant inputs, keeping in mind that this will require even larger high-quality data sets to avoid overfitting in multivariable analysis. Finally, time elapsed before operation could be increased for a myriad reasons, often by medical necessity, and it is difficult to know on a patient level why this is the case. Regardless, clinicians ought to be attentive to this matter, and while tending to their patients' immediate needs they should advocate for timely surgery.

CONCLUSION

Within our academic network comprising 3 EGS services of various service-model types, increasing length of time between admission and operation and overnight operating were associated with greater morbidity and mortality for EGS patients. Although multiple factors are probably at play, a lack of access to daytime operating room resources across our institutions is a modifiable system-based factor contributing to these findings. Moving forward, optimization of care for EGS patients in an ACS model should consider strategies to provide timely access to the operating room.

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Contributors: M. Meschino, A. Giles, M. Saddik, R. Nenshi, K. Vogt and P. Engels designed the study. M. Meschino acquired the data, which M. Meschino, A. Giles, T. Rice, A. Doumouras, R. Nenshi, L. Allen, K. Vogt and P. Engels analyzed. M. Meschino, A. Giles and P. Engels wrote the article, which all authors critically reviewed. All authors approved the final version of the article to be published.

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