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the ESTES SnapAppy Group

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the ESTES SnapAppy Group , Bass , G A , Mohseni , S , Ryan , E J , Tolonen , M , Aaltonen , G , Sillman , I S , Haapanen , S , Lampela , H , Sammalkorpi , H , Eskola , S , Laakso , A , Back , J , Kettunen , U , Nummi , A M , Nykänen , T , Rantala , R , Mäkäräinen-Uhlbäck , E J , Meriläinen , S A & Rintala , J M J 2023 , 'Clinical practice selectively follows acute appendicitis guidelines ' , European Journal of Trauma and Emergency Surgery , vol. 49 , bÿ p p . 45 56 . https://doi.org/10.1007/s00068-022-02208-2

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ORIGINAL ARTICLE



Clinical practice selectively follows acute appendicitis guidelines

Gary Alan Bass^{1,2,3,4} · Shahin Mohseni² · Éanna J. Ryan⁵ · Maximilian Peter Forssten² · Matti Tolonen⁶ · Yang Cao⁷ · Lewis J. Kaplan^{1,8} on behalf of ESTES SnapAppy Group

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Abstract

Introduction Acute appendicitis is a common surgical emergency, and the standard approach to diagnosis and management has been codified in several practice guidelines. Adherence to these guidelines provides insight into independent surgical practice patterns and institutional resource constraints as impediments to best practice. We explored data from the recent ESTES SnapAppy observational cohort study to determine guideline compliance in contemporary practice to identify opportunities to close evidence-to-practice gaps.

Methods We undertook a preplanned analysis of the ESTES SnapAppy observational cohort study, identifying, at a patient level, congruence with, or deviation from WSES Jerusalem guidelines for the diagnosis and management of acute appendicitis and the Surviving Sepsis Campaign in our cohort. Compliance was then correlated with the incidence of postoperative complications.

Results Four thousand six hundred and thirteen (4613) consecutive adult and adolescent patients with acute appendicitis were followed from date of admission (November 1, 2020, and May 28, 2021) for 90 days. Patient-level compliance with guideline elements allowed patients to be grouped into those with full compliance (all 5 elements: 13%), partial compliance (1–4 elements: 87%) or noncompliance (0 elements: 0.2%). We identified an excess postoperative complication rate in patients who received noncompliant and partially compliant care, compared with those who received fully guideline-compliant care (36% and 16%, versus 7.3%, p < 0.001).

Conclusions The observed diagnostic and treatment practices of the participating institutions displayed variability in compliance with key recommendations from existing guidelines. In general, practice was congruent with recommendations for preoperative antibiotic surgical site infection prophylaxis administration, time to surgery, and operative approach. However, there remains opportunities for improvement in the choice of diagnostic imaging modality, postoperative antibiotic stewardship to timely discontinue prophylactic antibiotics, and the implementation of ambulatory treatment pathways for uncomplicated appendicitis in the healthy young adult.

Keywords Acute appendicitis · Observational cohort · Appendectomy · Guidelines

Introduction

Acute appendicitis is one of the most prevalent general surgical emergencies worldwide [1–4]. The combination of resectional source control (appendectomy) and

The members of the ESTES SnapAppy Group are mentioned in the Acknowledgements section.

- ☐ Gary Alan Bass gary.bass@pennmedicine.upenn.edu
- Shahin Mohseni mohsenishahin@yahoo.com

Extended author information available on the last page of the article

perioperative antimicrobial pharmacotherapy have served as the 'gold standard' treatment strategy for most with acute appendicitis [5]. However, decision-making for those with acute appendicitis has recently become more nuanced around treatment approach and timing, especially related to acute SARS-CoV-2 infection. Acknowledging this, the World Society of Emergency Surgery (WSES) identifies acute appendicitis as a global research priority in acute care surgery [6, 7].

Discouragement of the use of laparoscopy by guidelines from learned societies early in the COVID-19 pandemic accentuated a drift in 'usual practice' away from operative management for a variety of surgical conditions,



including appendicitis [8, 9]. Unrelatedly, the observation that early appendicitis may be treated with antibiotics alone in selected patients had already prompted randomized controlled trial investigation [10–12]. However, the maturation of these trial results suggested poorer outcomes in nonoperative versus operative treatment of appendicitis [12–17]. Furthermore, as the pandemic progressed, and the virus mutated, it became apparent that the perioperative morbidity and mortality associated with acute SARS-CoV-2 infection appeared overstated [18–21]. Nonetheless, the influence of rapidly articulated professional organization guidance—ahead of evidence—and the rise of nonoperative management supported the growth of heterogenous patterns of practice for patients with acute appendicitis, arose that diverged from [22–26]

Consensus guidelines are often developed use the modified Delphi method, an iterative approach which originated in business management decision-making theory [22]. This approach leverages aggregated expert opinion anchored in data from previous scientific inquiry to harmonize clinical practice across healthcare settings [23]. The resulting guidelines, which exist for the diagnosis and treatment of many surgical diseases, are often sponsored by medical professional organizations, such as the European Society of Trauma and Emergency Surgery (ESTES), the Eastern Association for the Surgery of Trauma (EAST), the Society for Critical Care Medicine (SCCM), and the World Society of Emergency Surgery (WSES). Prompt initiation of empiric antimicrobial therapy and early definitive source control reduce the risk of infection-related complications including organ failure and mortality [24–26]. Antimicrobial stewardship, informed by the STOP-IT trial [27, 28], advocates for the discontinuation of antibiotics in a truncated time frame after achieving adequate source control [29]. These findings have been codified in several medical professional organization guidelines, such as the joint SCCM/European Society for Intensive Care Medicine (ESICM)-sponsored Surviving Sepsis Campaign (revised 2021) [30] and the WSES Jerusalem Consensus Guidelines for the Diagnosis and Management of Acute Appendicitis (revised 2020) [31].

In the SnapAppy study of usual care in appendicitis, we accrued a simultaneous multi-national observational sample of adult patients undergoing operation for acute appendicitis to determine current practice patterns. This non-interventional approach provides granular insights into current practice and how it embraces or deviates from published guidance. Since guideline adoption is generally nonuniform and may require decades to demonstrate widespread adoption, deviation is reasonably anticipatable [32, 33]. Furthermore, recognizing that individual clinicians direct bedside care within the context of institutional expertise and resources, we hypothesized that there would be broad divergence from

established guidelines regarding antibiotic agent selection, antibiotic cessation timing, diagnostic modality selection, and operative approach in managing acute appendicitis.

Methods

Protocol

Using the 'snapshot audit methodology,' a validated prospective observational approach to studying epidemiology, treatment effectiveness and inter-institutional variations in practice patterns and guideline adherence [34], we constructed a defined dataset, in line with a pre-specified protocol registered with ClinicalTrials.gov (Trial # NCT04365491), which captures the contemporary epidemiology of appendicitis across 71 centers in 14 European, Middle Eastern, and North American countries (Bahrain, Estonia, Finland, Iran, Ireland, Israel, Italy, Portugal, Romania, Spain, Sweden, Switzerland, UK, and USA. We enrolled all consecutive adult patients (over 15 years of age) admitted with acute appendicitis in a 90-day window between November 1, 2020, and May 28, 2021, and followed those patients for 90 days post-admission (up to August 31, 2021). The study complied with the Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) guidelines [35] and the Declaration of Helsinki [36].

Center eligibility

To capture as broad a practice base as possible, any unit undertaking adult emergency general surgery was eligible to register patients in the study, unrestricted by minimum case volume, or center-specific limitations. The study protocol, and an invitation to participate, was disseminated to registered members of the European Society of Trauma and Emergency Surgery (ESTES), and through national surgical societies using societal email lists, social media announcements, announcement on the ESTES website, and through peer-to-peer word of mouth.

Patient eligibility

All adult and adolescent patients (over 15 years of age) admitted for acute appendicitis were included in the current study. Acknowledging that there are several appendicitis grading systems, for simplicity and convenience, we used the AAST Anatomic Disease Severity grading system, which uses clinical, radiographic, operative, and pathologic criteria to assign an incrementing ordinal severity score of 1 (mild disease limited to the organ) to 5 (widespread severe disease) [37–40]. The American Society of Anesthesiologists (ASA) risk-stratification classification was reported. Patients



who demonstrated mesenteric adenitis, or ovarian or colonic pathology were specifically excluded.

Data capture

Data were recorded contemporaneously and stored on a secure, user-encrypted online platform (SMARTTrial®) without patient-identifiable information. Centers were asked to validate that all eligible patients during the study period had been entered and to attain > 95% completeness of data field entry prior to final submission. The database was closed for analysis on November 1, 2021. The SnapAppy protocol was designed so that usual patient follow-up pathways could be utilized to obtain outcomes data. No additional visits or changes to routine follow-up were made. However, local investigators were encouraged to be proactive in identifying post-diagnosis events (or lack thereof), within the limits of usual follow-up. These included reviewing the patient notes (paper and electronic) during admission and before discharge to note in-hospital complications, reviewing hospital systems to check for re-attendances or re-admissions, and reviewing postoperative radiology reports.

Outcome measures

The primary outcome measures were documented adherence to practices recommended in the Surviving Sepsis Campaign 2021, and the WSES Jerusalem Guidelines 2020 for the diagnosis and management of acute appendicitis [30, 31]. Specifically, we focused on the following diagnostic and treatment guidelines that embrace antimicrobial stewardship: diagnosis by ultrasound or clinical examination in patients under the age of 40 years, and by CT in those over 40, the administration of first dose of preoperative antibiotics within 3 h of diagnosis, operative source control within 24 h of diagnosis, the omission of postoperative antibiotics in uncomplicated appendicitis, and the discontinuation of antibiotics within 3-5 days following adequate source control (Table 1). We tallied the five guideline elements described (Table 1), allowing patients to be grouped into those with full compliance (all 5 elements), partial compliance (1-4 elements), or noncompliance (0 elements). Secondary outcomes were inpatient length of stay and the incidence of postoperative complications (overall, and specific complications including surgical site infection [superficial, deep, organ space], superficial wound dehiscence, and postoperative ileus).

Statistical analysis

All descriptive analyses were conducted with the statistical software the jamovi project (2022). *jamovi*. (version 2.3), [Computer Software, Retrieved from https://www.jamovi.

org.] running the *gtsummary*, *ClinicoPath* and *ggstatsplot* packages. Normally distributed variables were presented as means and standard deviations (SDs), while the median and interquartile range (IQR) were used for non-normally distributed variables. Categorical variables were summarized as counts and percentages.

Ethical considerations.

All participating centers secured IRB approval or equivalent. No patient consent was sought since the current study was purely observational. All data were de-identified when uploaded to the secure study database (SMARTTrial®), in compliance with HIPAA and EU GDPR legislation.

Results

Patient demographics

Four thousand six hundred and thirteen (4613) consecutive adult patients with acute appendicitis were followed from date of admission (November 1, 2020, and May 28, 2021) for 90 days. Their median age was 36 years (IQR 25–51), and there was a slight male preponderance (55.2%). Patients had a mean body mass index of 26.5 (\pm 12.1). Most patients were risk stratified as low risk to undergo general anesthesia by the American Society of Anesthesiologists (ASA) classification (ASA \leq 2: 90.4%). Approximately 2% of patients had an active COVID-19 infection on admission, with a further 2.6% reporting prior infection.

Diagnostic modality

The diagnosis of acute appendicitis was made on clinical findings in 512 (11.1%) patients, combining clinical findings and transabdominal ultrasound (either at point of care or in the radiology department) in 1436 (31.3%) and with axial imaging (CT) in 2644 (57.6%). Ninety percent of patients over the age of 40 underwent diagnostic CT.

Source control

Of the 4,613 patients diagnosed with acute appendicitis who underwent a source control intervention, 4,391 (95.2%) underwent appendectomy, with the vast majority being operated upon within 24 h (87.4%). Interventional radiological drainage without operation was undertaken in 1.2% of cases. Most appendectomies were performed laparoscopically (85.8%). The conversion rate was approximately 3%, and the incidence of primary open surgery was 9.9%. Frank pus was observed in 16.8% of cases at operation. Most appendix specimens were submitted for histopathological evaluation (97.4%); 1.4% of patients had



Table 1 Guideline compliance elements from the WSES Jerusalem Consensus Guidelines (2020) for the Diagnosis and Treatment of Acute Appendicitis, and the Surviving Sepsis Campaign Guidelines (2021) [30, 31]

Guideline domain	Recommendation/suggestion	Quality of evidence	Strength of recommendation
Diagnosis			
Clinical and Ultrasound	WSES 1.7 We recommend the routine use of a combination of clinical parameters and US to improve diagnostic sensitivity and specificity and reduce the need for CT scan in the diagnosis of acute appendicitis	Moderate	Strong; 1B
Point-of-care Ultrasound	WSES 1.10 We recommend POCUS as the most appropriate first- line diagnostic tool in both adults and children, if an imaging investigation is indicated based on clinical assessment	Moderate	Strong; 1B
СТ	WSES 1.9 We suggest that in high-risk patients younger than 40 years old (with AIR score 9–12 and Alvarado score 9–10 and AAS≥16), CT may be avoided before proceeding to diagnostic+/−therapeutic laparoscopy	Moderate	Weak; 2B
СТ	WSES 1.12 We recommend cross-sectional imaging before surgery for patients with normal investigations but non-resolving right iliac fossa pain, and those over the age of 40 years. After negative imaging, initial nonoperative treatment is appropriate. However, in patients with progressive or persistent pain, explorative laparoscopy is recommended to establish/exclude the diagnosis of acute appendicitis or alternative diagnoses	High	Strong; 1A
Surgical treatment			
Source control time to OR	WSES 3.1 We recommend planning laparoscopic appendectomy for the next available operating list within 24 h in case of uncomplicated acute appendicitis, minimizing the delay wherever possible	Moderate	Strong; 1B
Source control time to OR	WSES 3.2 We recommend against delaying appendectomy for acute appendicitis of any grade needing surgery beyond 24 h from the admission	Moderate	Strong; 1B
Antimicrobial stewardship			
Time to first-dose antibiotics	SSC 14 For adults with possible sepsis without shock, we suggest a time-limited course of rapid investigation and if concern for infection persists, the administration of antimicrobials within 3 h from the time when sepsis was first recognized	Weak	Very low
Preoperative antibiotics	WSES 7.1 We recommend a single preoperative dose of broad- spectrum antibiotics in patients with acute appendicitis undergo- ing appendectomy	High	Strong; 1A
Duration of postoperative antibiotics	WSES 7.1 We recommend against postoperative antibiotics for patients with uncomplicated appendicitis	High	Strong; 1A
Duration of postoperative antibiotics	WSES 7.2 We recommend against prolonging antibiotics longer than 3–5 days postoperatively in case of complicated appendicitis with adequate source-control	High	Strong; 1A

some form of appendiceal neoplasm upon histopathological evaluation.

Perioperative antimicrobial stewardship

While almost all patients appropriately received preoperative antibiotic therapy (96.6%), the SSC target of a first dose within 3 h for those with confirmed or highly suspected sepsis was achieved in just 42.3% of patients. The most commonly prescribed antimicrobials were amoxicillin/clavulanic

acid, or a cephalosporin with or without metronidazole (77.5%); other local variations were noted.

While data on postoperative antibiotic prescription are missing for 366 (8.3%) patients who underwent appendectomy, 62.5% of patients received postoperative antibiotics. Postoperative antibiotics were continued in 1,589 (49.7%) patients with Grade I (uncomplicated) appendicitis and 578 (71.4%) of patients with Grade II (gangrenous, nonperforated) appendicitis (see Table 2). The mean (SD; range) duration of postoperative antibiotics in Grade I and Grade



II appendicitis was found to be 3.7 (4.1; 0–50) days and 4.3 (3.7; 0–21) days, respectively.

Composite measure of compliance and postoperative complications

We tallied patient-level compliance with the five guideline elements described above (Table 1) allowing patients to be grouped into those with full compliance (all 5 elements: 13%), partial compliance (1–4 elements: 87%) or noncompliance (0 elements: 0.2%). We identified an excess postoperative complication rate in patients who received noncompliant and partially compliant care, compared with those who received fully guideline-compliant care (36% and 16%, versus 7.3%, p < 0.001). While the Clavien–Dindo 30-day complication classification and the absolute numbers for individual infective complications (wound infection, postoperative ileus, etc.) are tabulated in Table 3, the small event rate means that statistically significant differences are unlikely to have clinical significance. Country-level comparisons in compliance rates are tabulated in Table 4, with full compliance ranging from 0 to 45% (mean = 13%, SD = 14%). Greatest rates of full compliance were seen in patients treated in Romania, USA, and Estonia. Partial compliance was more common, ranging between 55 and 100%.

Table 2 Select guideline compliance metrics—grouped by AAST disease severity grade and overall patients

Characteristic	Grade I $(N=3203)$	Grade II $(N=809)$	Grade III $(N=277)$	Grade IV $(N=288)$	Grade V $(N=36)$	All Patients $(N=4613)$	p value
Method of diag- nosis							< 0.001 ^b
CT	1929 (60.5%)	157 (19.4%)	253 (92%)	275 (95.8%)	30 (83.3%)	2644 (57.6%)	
Over 40	929 (88.3%)	105 (84.0%)	186 (94.4%)	204 (97.6%)	20 (90.9%)	1444 (90.0%)	
Ultrasound	951 (29.8%)	453 (56.1%)	19 (6.9%)	9 (3.1%)	4 (11.1%)	1436 (31.3%)	
Clinical	306 (9.6%)	198 (24.5%)	3 (1.1%)	3 (1%)	2 (5.6%)	512 (11.1%)	
Missing	17	1	2	1	0	21	
Time to first-dose antibiotics ^a							
<1 h	359 (11.8%)	102 (13%)	33 (12%)	43 (15.9%)	8 (22.9%)	545 (12.4%)	
1–3 h	817 (26.9%)	229 (29.2%)	91 (33.1%)	83 (30.6%)	13 (37.1%)	1233 (28%)	
3–6 h	796 (26.2%)	248 (31.7%)	80 (29.1%)	77 (28.4%)	8 (22.9%)	1209 (27.5%)	
>6 h	1061 (35%)	204 (26.1%)	71 (25.8%)	68 (25.1%)	6 (17.1%)	1410 (32.1%)	
Missing	170	26	2	17	1	216	
First-dose antibiotics within 3 h	1346 (42%)	357 (44.1%)	126 (45.5%)	143 (49.7%)	22 (61.1%)	1994 (43.2%)	0.015 ^b
Time to OR ^a							
<6 h	555 (17.9%)	132 (16.8%)	58 (21.7%)	39 (18.5%)	12 (34.3%)	796 (18.1%)	
6–12 h	1109 (35.8%)	285 (36.3%)	102 (38.2%)	49 (23.2%)	11 (31.4%)	1556 (35.4%)	
12-24 h	1062 (34.3%)	279 (35.5%)	80 (30%)	72 (34.1%)	7 (20%)	1500 (34.1%)	
>24 h	369 (11.9%)	89 (11.3%)	27 (10.1%)	51 (24.2%)	5 (14.3%)	541 (12.3%)	
Missing	108	24	10	77	1	220	
Preoperative antibiotics	3092 (96.6%)	787 (97.3%)	271 (97.8%)	268 (93.4%)	36 (100%)	4454 (96.6%)	0.013 ^b
Missing	1	0	0	1	0	2	
Postoperative antibiotics	1589 (49.7%)	578 (71.4%)	263 (94.9%)	279 (97.6%)	36 (100%)	2745 (59.6%)	< 0.001 ^b
Missing	4	0	0	2	0	6	
Postoperative antibiotic duration~	3.7 (4.1; 0–50)	4.3 (3.7; 0—21)	8.5 (4.2; 0–21)	10.5 (5.3; 0–33)	11 (5; 1–21)	4.6 (4.6; 0–50)	<0.001°
Missing	474	83	13	15	3	588 ^d	

^aTime from diagnosis (hours); [~]mean days (SD; range)



^bPearson's Chi-squared test

^cLinear model ANOVA

^dIncludes 222 patients who did not undergo surgery; thus the true missing figure is 366

Outcomes for appendicitis patients

Appendicitis patients had a median length of hospital stay of 2 days. Fewer than 1% of patients required ICU care. A post-operative complication was suffered by 17.3% of patients within 30 days of surgery, with the most common being the development of a pelvic abscess (3.2%), ileus (2.6%), or a surgical site infection (1.9%). However, only 2.8% of all patients suffered a severe complication (Clavien–Dindo classification \geq 3a), while 1.6% required reoperation. A total of 7 patients (0.2%) died within the first month after surgery. Median (IQR) length of hospital stay in days was significantly shorter in patients who received fully compliant care (1.2, 0.8–1.7 days), compared with those who received partially compliant (2.0, 1.4–3.6 days) or noncompliant care (4.3, 3.2–8.1 days) [p < 0.001].

Discussion

The SnapAppy analysis of the current state of acute appendicitis management offers valuable insights regarding practice patterns and outcomes across widely disparate health systems and settings. Besides affording comparisons between different countries, these data allow us to explore how current practice interfaces with existing evidence regarding that care. To that end, we have compared observational data with

two major guidelines—that of the World Society of Emergency Surgery for the management of patients with acute appendicitis and that of the Surviving Sepsis Campaign for the management of patients with sepsis or septic shock (Table 1) [30, 31]. While the SSC guidelines may be viewed as primarily appropriate for medical disease, or postoperative infection such as pneumonia, recent iterations of the guidelines also highlight the importance of source control for patients with a source controllable lesion [30]. Therefore, appendectomy for acute appendicitis provides an ideal opportunity to achieve rapid and effective source control in the absence of perforation.

Guidelines provide an evidence base upon which clinicians may rely to inform care decisions [41]. Often, guidelines are generated using a modified Delphi consensus approach that yields recommendations, suggestions, best practices, and the recognition that data are sufficiently lacking for some topics to preclude guidance [22, 23]. Some guidelines, but not all, are accompanied by a 'bundle' that provides an implementation strategy that translates to the bedside [41]. When there is no offered implementation strategy, the individual clinician, group, or facility must devise how best to incorporate the evidence base into their practice. Unsurprisingly, the completeness with which evidence is embraced within practice is often less than uniform, creating what has been termed an 'evidence-to-practice' gap [42]. The abrogation of such gaps across all disciplines is a key

Table 3 Crude complication rate, grouped by full (5/5), partial (1–4/5) or noncompliance (0/5) with selected guidelines

Characteristic N		Fully compliant N=588 (13%)	Partially compliant N=4014 (87%)	Noncompliant $N=11 (0.2\%)$	<i>p</i> -value	
Any complication	4,610	43 (7.3%)	624 (16%)	4 (36%)	< 0.001	
Wound infection	4,604	4 (0.7%)	81 (2%)	2 (18%)	0.002	
Wound dehiscence	4,604	4 (0.7%)	26 (0.6%)	0 (0%)	0.80	
Pelvic abscess	4,604	3 (0.5%)	143 (3.6%)	0 (0%)	< 0.001	
Subphrenic abscess	4,604	0 (0%)	8 (0.2%)	0 (0%)	0.61	
Postoperative sepsis	4,604	0 (0%)	33 (0.8%)	0 (0%)	0.050	
Postoperative ileus	4,604	3 (0.5%)	117 (2.9%)	0 (0%)	< 0.001	
Clavien–Dindo 30-day complications	4,415				< 0.001	
1		11 (1.9%)	156 (4.1%)	1 (10%)		
2		8 (1.4%)	170 (4.4%)	2 (20%)		
3a		1 (0.2%)	68 (1.8%)	0 (0%)		
3b		1 (0.2%)	48 (1.3%)	0 (0%)		
4a		0 (0%)	2 (< 0.1%)	0 (0%)		
4b		0 (0%)	1 (<0.1%)	0 (0%)		
5		0 (0%)	7 (0.2%)	0 (0%)		
None		545 (96%)	3,387 (88%)	7 (70%)		
Missing		22	175	1		

Selected guidelines were: preoperative antibiotic administration, first-dose antibiotics within 3 h of diagnosis (Surviving Sepsis Campaign 2021), time to OR less than 24 h, omission of postoperative antibiotics in uncomplicated appendicitis, and diagnosis using clinical examination or ultrasound in patients under 40 years of age, or using CT in patients over 40 years



Table 4 Full (5/5), partial (1–4/5) or noncompliance (0/5) with selected guidelines, divided by country

Country	Fully compliant		Partially compliant		Noncompliant		Total
	\overline{n}	%	\overline{n}	%	\overline{n}	%	n
Bahrain	10	5%	211	95%	1	0%	222
Estonia	54	27%	147	73%	0	0%	201
Finland	47	7%	594	93%	0	0%	641
Iran	0	0%	88	100%	0	0%	88
Ireland	27	6%	418	94%	1	0%	446
Israel	0	0%	28	100%	0	0%	28
Italy	7	6%	113	94%	0	0%	120
Portugal	8	5%	161	95%	0	0%	169
Romania	9	45%	11	55%	0	0%	20
Spain	197	17%	962	83%	0	0%	1159
Sweden	112	11%	947	89%	0	0%	1059
Switzerland	2	7%	26	93%	0	0%	28
UK	10	6%	139	88%	9	6%	158
USA	105	38%	169	62%	0	0%	274
Overall	588	13%	4014	87%	11	0%	4613

Selected guidelines were: preoperative antibiotic administration, first-dose antibiotics within 3 h of diagnosis (Surviving Sepsis Campaign 2021), time to OR less than 24 h, omission of postoperative antibiotics in uncomplicated appendicitis, and diagnosis using clinical examination or ultrasound in patients under 40 years of age, or using CT in patients over 40 years

priority of the expanding field of implementation science. The evidence-to-practice gap bears important implications for care quality, safety, and cost, rendering it imperative to understand what supports the creation of the gap.

Established drivers of the evidence-to-practice gap include individual, environment, and system factors. Established practice is generally comfortable for the clinician, and inertia may impede change. The notion that trial patients from whom the data are derived are different from those the clinician treats has some validity, especially when the data flow from Randomized Clinical Trials with restrictive entry criteria. However, a host of other assessments, like this study, is not restricted and provides evidence that may be broadly applied. Other clinician factors include a lack of awareness of newly published evidence, especially if the data are published behind a pay wall. Guidelines, on the other hand, are often freely accessible, even if only as an executive summary. Additional impediments are those related to the environment of care and the healthcare system. The lack of an electronic health record (EHR), as is the case in many of our contributing centers, may confound obtaining longitudinal data for assessment. Practice within a resource limited space, including those within low- and middle-income countries (LMIC), may constrain diagnostic modality availability, as well as the rapeutic agent selection. Patients operated after 24 h may reflect lack of OR availability in resource constrained settings, a constraint that would not be remedied by an Emergency General Surgery service—an initiative that durably decreased time to OR

for time sensitive conditions [43, 44]. Clinician availability may reduce the oversight required for timely termination of perioperative antibiotics, in the absence of an established pathway, or advanced practice provider (APP) to help guide care. These aspects likely contributed to the low frequency of an enhanced recovery after surgery (ERAS) approach that includes same day discharge for uncomplicated appendicitis in otherwise healthy adults [45, 46].

Nonetheless, our data demonstrate that practice can conform to guideline-based recommendations-which suggests the lack of an evidence-to-practice gap—across very different healthcare systems (USA, Estonia, Romania), at least in participating centers. Despite the ability to embrace guideline recommendations in select sites, most centers demonstrated failures in guideline adoption. These failures were most notable in antibiotic cessation timing and inpatient care (time to operation, and the lack of outpatient care for uncomplicated appendicitis). While not readily apparent on an individual patient basis, collectively, these choices exert major impacts on cost and serve as a driving force for the genesis of multidrug-resistant organisms [47-49]. Individual clinician management approaches influence microbial ecology within the hospital and the community, including chronic care facilities. Accordingly, antimicrobial stewardship programs (ASPs) have arisen to help guide appropriate antibiotic selection and cessation. ASPs bring additional clinicians onto the team to support embracing new evidence and



incorporating it into practice but are insufficient in isolation. Other approaches are warranted to systematically close the gap.

A variety of methods may help close the evidence-topractice gap and are presented below in order of increasing difficulty but also increasing anticipated efficacy. Clinician education is straightforward and may use a variety of platforms from traditional didactics to 'just-in-time' digital platform-based training. Despite the ease of providing education, it is challenging to demonstrate robust practice change in its wake [50–52]. Providing an incentive to change practice is more appealing, especially when financially based, but requires institutional resources or insurer resources to realize; incentives may be especially problematic in resource limited spaces. Exacting a penalty for compliance failure may be effective but then requires data reporting, analysis, and a larger system to ensure accurate attribution and penalty application. This is especially true when the penalty is related to finances that flow to an individual clinician as opposed to an established institution. Public reporting of compliance and outcomes may be built as an outgrowth of data acquisition and works best when it is mandated by a state or national agency. Such is the case for the New York State Department of Health's mandatory reporting around sepsis care. There is substantial pressure for institutional performance when it is documented as lagging behind that of other institutions or practitioners. Finally, establishing a local champion who can access clinician specific data, review it with them, and provide peer-to-peer education and feedback holds the potential to be quite effective [53–55]. However, that individual must be credible, have sufficient time to do so, be compensated for time and effort, and work within a medical staff that is willing to have their practice examined by a peer. In the private sector, that peer may also compete for the same patients, relegating such 'counselling' to teaching institutions where such competition is less applicable. When such a system is feasible, the peer reviewer functions as a team member for the clinician, a post that may be particularly important for those in solitary practice.

Our observational study of acute appendicitis management offers a view of current practice while also demonstrating important limitations. First, due to time-bound simultaneous patient accrual, our data do not capture outcomes beyond 90 days after admission. Second, the original intention of the 90-day follow-up informed by anecdotal experience during the early phase of the COVID-19 pandemic was to capture early readmission with recrudescent appendicitis due to failure of nonoperative antibiotic therapy. However, very few patients undergoing nonoperative therapy were captured in our dataset. This is at least in part be explained using operating room logs to identify enrollable patients at centers without an EHR. Third, granular elements of patient

care were not captured including, but not limited to, anesthetic technique, culture data, and specific antibiotic prescription. However, those elements were not the focus of the study. Furthermore, due to the heterogeneity on a national level, socioeconomic status, race/ethnicity, and insurance status were not recorded. Fourth, the correlations presented are merely associative and should not be construed as indicating causation, especially regarding complications that were quite infrequent. Prospective correlative analysis of outcomes would require a population-level epidemiologic study. Fifth, we did not assess all the bundle elements from the SSC as that inquiry would require a granularity that exceeded the scope of this investigation. Finally, as these data were limited to adult patients admitted for appendicitis, they should not be extrapolated to the pediatric population.

Conclusions

The observed diagnostic and treatment practices of the participating institutions displayed variability in compliance with key recommendations from existing guidelines (2020 WSES Jerusalem and the 2021 Surviving Sepsis Campaign) [30, 31]. In general, practice was congruent with recommendations pertaining to preoperative antibiotic surgical site infection prophylaxis administration, time to surgery, and operative approach. However, there remains opportunities for improvement in the choice of diagnostic imaging modality, postoperative antibiotic stewardship to timely discontinue prophylactic antibiotics, and the implementation of ambulatory treatment pathways for uncomplicated appendicitis in the healthy young adult. Data from the SnapAppy are hypothesis generating and should optimally inform future investigations and implementation science initiatives specifically designed to close the evidence-to-practice gap.

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Karolinska Instituet, Stockholm, Sweden: Rebecka Ahl Hulme; Humanitas Clinical and Research Center, IRCCS, Milano, Italy: Hayato Kurihara; Clinical Epidemiology and Biostatistics, School of Medical Sciences, Orebro University, Orebro, Sweden: Yang Cao; Corporal Michael Crescenz Veterans Affairs Medical Center, Philadelphia, USA: Lewis J. Kaplan. Contributing Centers and Authors: Bahrain Defence Force-Royal Medical Services, Bahrain: Nayef Louri, Fatema Nedham, Thomas Noel Walsh, Jamal Hashem; King Hamad University Hospital, Bahrain: Martin Corbally, Abeer Farhan, Hamad Al Hamad, Rawan Elhennawy; Salmaniya Medical Complex, Bahrain: Mariam AlKooheji, Manar AlYusuf, Wissal Aknouche, Anas A. Zeidan, Yusuf S. Alsaffar; North Estonia Medical Center, Estonia: Edgar Lipping, Peep Talving, Sten Saar, Katrina Graumann, Liis Kibuspuu, Eduard Harkov; HUS Meilahden Tornisairaala, Finland: Gisele Aaltonen, Iines S. Sillman, Sami Haapanen; HUS Jorvin sairaala, Finland: Hanna Lampela, Henna Sammalkorpi, Sofia Eskola, Altti Laakso; Hyvinkää Hospital Area, Finland: Johan Back, Ulla Kettunen, Antti M. Nummi, Anika Szwedyc, Taina Nykänen, Rolle Rantala; Oulun Yliopistollinen Sairaala, Finland: Elisa J. Mäkäräinen-Uhlbäck, Sanna A. Meriläinen, Heikki I. Huhta, Jukka M. J. Rintala, Kirsi E. M. Laitakari; Turku University Hospital, Finland: Elina Lietzen, Paulina Salminen, Risto K. A. Rapola; Namazi Hospital, Shiraz University of Medical Sciences, Iran: Vahid Zangouri, Mohammad Y. Karami, Sedigheh Tahmasebi, Majid Akrami, Alireza Golchini, Faranak Bahrami; Tullamore General Hospital, Ireland: Sean M. Johnston, Sean T. Lim, Irele Ifijeh Ahonkhai, Eltahir Eltagani, Odhran K. Ryan; St Vincent's University Hospital, Ireland: Ailbhe O'Driscoll-Collins, Aine O'Neill, Zakiya Penny, Orlaith Kelly, Carolyn Cullinane, Ian Reynolds, Helen Heneghan, Sean Martin, Des Winter; Galway University Hospitals, Ireland: Matthew Davey, Maha Alkhattab, Aoife J. Lowery, Michael J. Kerin, Aisling M. Hogan, Martin S. Davey, Ke En Oh; Letterkenny University Hospital, Ireland: Syed Mohammad Umar Kabir, Huilun Huan, Charlotte Aziz, Michael Sugrue; University Hospital Waterford, Ireland: Jessica M. Ryan, Tara M. Connelly, Mohammad Alhazmi, Youssef Al-Mukhaizeem, Fiachra Cooke, Peter M. Neary; Beaumont Hospital, Ireland: Arnold D. K. Hill, Michael R. Boland, Angus J. Lloyd, Frances Fallon, Eoin F. Cleere, James Toale; Mayo University Hospital, Ireland: Patrick A. Boland, Michael Devine, Conor Keady, Sarah Hunter, M. Kevin Barry; Tallaght University Hospital, Ireland: Michael E. Kelly, Aidan T. O'Dowling, Ben Creavin, Dara O. Kavanagh, Paul Neary, Paul F. Ridgway, Cathleen A. McCarrick; St James' University Hospital, Ireland: Jarlath Bolger, Barry Maguire, Cian Keogh, Surbhi Chawla; Mater Misericordiae University Hospital, Ireland: John Conneely, Emilie McCormack , Ben Shanahan, Nicola Raftery, Darragh Rice, Niall McInerney, Aine Stakelum, Jan Mares, Jonavan Tan, Mark Hanna, Ishwarya Balasubramanian, Christina Fleming; Soroka University Medical Center, Israel: Guy Barsky, Gad Shaked; Emergency Surgery and Trauma Section, Humanitas Research Hospital, Rozzano, Italy: Simone Giudici, Martina Ceolin, Simona Mei, Francesca Mazzarella; Trieste University Hospital, Italy: Annalisa Zucca, Susanna Terranova, Nicolo de Manzini; Azienda Ospedaliero-Universitaria Città della Salute e della Scienza di Torino, Italy: Diego Visconti, Emanuele Doria, Mauro Santarelli; San Maurizio Hospital, Bolzano, Italy: Giovanni Scotton, Francesca Notte, Giacomo Bertelli, Anna Malpaga, Giulia Armatura, Antonio Frena; Cisanello Hospital, University of Pisa, Italy: Dario Tartaglia, Federico Coccolini, Camilla Cremonini, Enrico Cicuttin, Alessio Mazzoni, Massimo Chiarugi; Centro Hospitalar Universitário da Cova da Beira, Portugal: Constança M. Azevedo, Filipa D. Mendes, Luis Q. Faria, Carlos Nazario, Daniela Machado, Miguel Semiao; Centro Hospitalar Tondela-Viseu, Portugal: Jorge Pereira, Carlos Casimiro, Jose Pinto, Tiago Pavão, Raquel Pereira, Bruno Barbosa; Centro Hospitalar de Trás-os-Montes e Alto Douro, Portugal: Nadia Tenreiro, Catia Ferreira, Goncalo Guidi, Daniela C. Martins, Clara Leal, Bruno B. Vieira; North Lisbon University Hospital Center, Portugal: Luís S. Castro, Aldara Faria, Alberto Figueira, Mauro Sousa, Pedro Rodrigues, Rodrigo Roquette; Centro Hospitalar Universitário do Algarve—Hospital de Faro, Portugal: Ricardo Ribeiro, Paulo Cardoso, Joana Domingues, Maria Isabel Manso, Rute Pereira, Tatiana Revez; Ponderas Academic Hospital, Bucharest, Romania: Bogdan D. Dumbrava, Florin Turcu, Ionut Hutopila, Bogdana Banescu, Gerald Filip, Catalin Copaescu; Hospital Universitario Juan Ramón Jiménez, Spain: Marcos Alba Valmorisco, Isabel Manzano Martín, Rocio, José Ortega Seda, Pablo Rodríguez González, Jose Antonio Becerra Toro, Enrique Rodríguez Lara, Jose Antonio González Minchón; Hospital Universitario Son Espases, Spain: Juan José Segura-Sampedro, Sebastián Jerí-McFarlane, Alejandro Gil-Catalán, Andrea Craus-Miguel, Laura Fernández-Vega, Xavier González-Argenté; Hospital General Universitario de Ciudad Real, Spain: Mercedes Estaire-Gómez, Borja Camacho Fernández-Pacheco, Rebeca Vitón-Herrero, Elisa Jimenez-Higuera, Alejandro Barbero, José M. Valverde; Hospital Universitario Son Llàtzer, Spain: Enrique Colás-Ruiz, Maria del Mar Escales-Oliver, Olga Claramonte-Bellmunt, Marta Castro-Suárez, Naila Pagés-Valle, José Andrés Cifuentes-Ródenas; Hospital Universitario Central de Asturias, Spain: Marta Merayo Alvarez, Jose Luis Michi Campos, Luis Alejandro, Beatriz, Jaime, Jose Luis; Infanta Sofía University Hospital, Spain: Carmen Rodríguez Haro, Sara Núñez O'Sullivan, Mariana García Virosta, María Hernández O'Reilly; Hospital Universitario de La Ribera, Spain: Izaskun Balciscueta-Coltell, Javier Lorenzo-Perez, Sonia Martinez-Alcaide, Susana Martinez-Ramos, Maria Sebastian-Fuertes, Laura Gomez-Romer; Hospital Universitario de Gran Canaria Dr Negrín, Spain: Maria M. Pelloni, Aida Cristina Rahy-Martín, Andrés Felipe Yepes-Cano; Hospital Universitario Virgen Macarena, Spain: Julio Reguera-Rosal, Jose A. Lopez-Ruiz, Beatriz Marenco, Marina Retamar-Gentil, Estela Romero-Vargas, Angeles Gil-Olarte; Urduliz Alfredo Espinosa Hospital, Spain: Aitor Landaluce-Olavarria, Begoña Estraviz-Mateos , Jose-Mario De Francisco-Rios, Aitor Sainz-Lete , Ane Emaldi-Abasolo, Manolo Leon-Valarezo; Donostia University Hospital, Spain: Claudia C. Lopes Moreira, Aintzane Lizarazu Perez, Araceli Rodriguez Gonzalez, Iñigo Augusto Ponce, Ignacio Maria Goena Iglesias; Hospital Universitario de Burgos, Spain: Cristina González-Prado , Guillermo Cabriada, Beatriz López, Michelle C Otero, Nerea Muñoz-Plaza, Alberto Palomo; Hospital Universitario Príncipe de Asturias, Spain: Fernando Mendoza-Moreno, Manuel Díez-Alonso, Francisca García-Moreno-Nisa, Belén Matías-García, Enrique Ovejero-Merino, Ana Quiroga-Valcárcel; Elche University General Hospital, Alicante, Spain: Luis Sánchez-Guillén, Inmaculada Oller-Navarro, Álvaro Soler-Silva, Antonio Francisco Sanchís-López; Complejo Asistencial Universitario de Salamanca, Spain: Francisco Blanco-Antona, Luis Muñoz-Bellvis, Jaime López-Sánchez, Sonsoles Garrosa-Muñoz, Beatriz Barón-Salvador, Juan Manuel Nieto-Arranz; Hospital Universitari Parc Taulí, Spain: Andrea Campos-Serra, Raquel Gràcia-Roman, Anna Muñoz-Campaña, Carla Zerpa-Martin, Andrea Torrecilla-Portoles, Tessa Landa; Virgen del Rocío University Hospital, Spain: Virginia Durán Muñoz-Cruzado, Felipe Pareja-Ciuró, Daniel Aparicio-Sánchez, Eduardo Perea del Pozo, Sandra Dios-Barbeito, Carlos García-Sánchez, Antonio Jesús García-Moriana; Hospital Clinic Barcelona, Spain: Victor Turrado-Rodriguez, Roser Termes-Serra, Paula Gonzalez-Atienza, Xavier Morales-Sevillano, Alba Torroella, César Ginestà Hospital Universitario Arnau de Vilanova, Spain: Alfredo Escartín, Ferney Gomez, Ana Pinillos, Jaume Ortega, Guillermo Lopez, Eric Gutierrez; Hospital Del Mar de Barcelona , Spain: Estela Membrilla-Fernandez, Francisco Ocho-Segarra, Ana María González-Castillo, Amalia Pelegrina-Manzano, Juan Guzmán-Ahumada, Juan Jose Sancho-Insenser; Complejo Hospitalario Universitario de A Coruña, Spain: María Lourdes García-Jiménez, Laura Castro-Diez, Manuel González-Bermúdez, Mónica Torres-Díaz, Carla Madarro Pena, Angélica Blanco Rodríguez; Örebro University Hospital, Sweden: Dhanisha Trivedi, Souheil Reda; Capio S:t Göran Hospital, Sweden: Hans Edvardsson, Lovisa Strömmer; Sahlgrenska University Hospital, Sweden: Eva-Corina Caragounis, Karin Sillén, Sofia Warfvinge; Sahlgrenska University Ostra Hospital, Sweden: Fredrik Bergstedt, Philip Enström, Harald Olsson, Anders Rosemar; Karolinska University Hospital,



Sweden: Nathalie Young, Agnieszka Popowicz, Johanna Lerström, Johanna Jäderbo, Folke Hammarqvist; Danderyds Hospital, Sweden: Hanna Zacharias; Karlstad Hospital, Sweden: Maria B. Wikström, Anna Stene Hurtsén; Östersund County Hospital, Sweden: Haytham Bayadsi, Emma Jansson, Nils Brunstrom, Ellen B. Malers; Linköping University Hospital, Sweden: Per I. Loftås, Anders Möller, Elena Atanasova; Bern University Hospital, University of Bern, Switzerland: Simone N. Zwicky, Beat Schnüriger; Aintree University Hospital, UK: Olga Rutka, Arjun T. Kattakayam, Mushfique Alam, John V. Taylor; Tameside and Glossop Integrated Care NHSFT, UK: Andrei Mihailescu, Eszter T. Karip, Ehtisham Zeb, Adam O'Connor, Goran Pokusevski; Brighton and Sussex University Hospitals, Brighton, UK: Mansoor Khan, Charlotte Florance, Christie Swaminathan, Shameen Jaunoo, Mohammed Sajid; University of Pennsylvania Hospital System, Philadelphia, USA: Caoimhe C. Duffy, John Rees, Mark J. Seamon, Niels D. Martin, Ian J. McCurry, Emily A. Vail, Bradford C. Bormann; Maine Medical Center, USA: Daniel C. Cullinane, Jaswin S. Sawhney, Jonathan Dreifus, Forest R. Sheppard; Riverside University Health System Medical Center, USA: Raul Coimbra, Paul Albini, Sara Edwards.

Author contributions Authors are parsed into the following groups at the end of the manuscript, according to the CREDIT taxonomy: Manuscript Writing Group, SnapAppy Steering Committee, and Study Collaborators and their affiliations. All are listed in PubMed.

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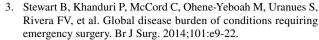
Declarations

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Authors and Affiliations

Gary Alan Bass^{1,2,3,4} · Shahin Mohseni² · Éanna J. Ryan⁵ · Maximilian Peter Forssten² · Matti Tolonen⁶ · Yang Cao⁷ · Lewis J. Kaplan^{1,8} · ESTES SnapAppy Group

- Division of Traumatology, Surgical Critical Care & Emergency Surgery, Perelman School of Medicine at the University of Pennsylvania, Philadelphia, USA
- Division of Trauma & Emergency Surgery, Orebro University Hospital and School of Medical Sciences, Orebro University, Örebro, Sweden
- Center for Perioperative Outcomes Research and Transformation (CPORT), University of Pennsylvania, Philadelphia, USA
- Leonard Davis Institute of Health Economics (LDI), University of Pennsylvania, Philadelphia, USA
- 5 Tallaght University Hospital, Dublin, Ireland

- Helsinki University Hospital HUS Meilahden Tornisairaala, Helsinki, Finland
- Department of Clinical Epidemiology and Biostatistics, School of Medical Sciences, Orebro University, Orebro, Sweden
- Orporal Michael Crescenz Veterans Affairs Medical Center, Philadelphia, USA

