




The management of adult appendicitis during the COVID-19 pandemic: an interim analysis of a UK cohort study

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Abstract

Background Acute appendicitis (AA) is the most common general surgical emergency. Early laparoscopic appendectomy is the gold-standard management. SARS-CoV-2 (COVID-19) brought concerns of increased perioperative mortality and spread of infection during aerosol generating procedures: as a consequence, conservative management was advised, and open appendectomy recommended when surgery was unavoidable. This study describes the impact of the first weeks of the pandemic on the management of AA in the United Kingdom (UK).

Methods Patients 18 years or older, diagnosed clinically and/or radiologically with AA were eligible for inclusion in this prospective, multicentre cohort study. Data was collected from 23rd March 2020 (beginning of the UK Government lockdown) to 1st May 2020 and included: patient demographics, COVID status; initial management (operative and conservative); length of stay; and 30-day complications. Analysis was performed on the first 500 cases with 30-day follow-up.

Results The patient cohort consisted of 500 patients from 48 sites. The median age of this cohort was 35 [26–49.75] years and 233 (47%) of patients were female. Two hundred and seventy-one (54%) patients were initially treated conservatively; with only 26 (10%) cases progressing to an operation. Operative interventions were performed laparoscopically in 44% (93/211). Median length of hospital stay was significantly reduced in the conservatively managed group (2 [IQR 1–4] days vs. 3 [2–4], $p < 0.001$). At 30 days, complications were significantly higher in the operative group ($p < 0.001$), with no deaths in any group. Of the 159 (32%) patients tested for COVID-19 on admission, only 6 (4%) were positive.

Conclusion COVID-19 has changed the management of acute appendicitis in the UK, with non-operative management shown to be safe and effective in the short-term. Antibiotics should be considered as the first line during the pandemic and perhaps beyond.

Keywords Appendicitis · COVID-19 · Non-operative · Antibiotics · Appendectomy

Members of The COVID: HAREM (Had Appendicitis, Resolved/ Recurred Emergency Morbidity/Mortality) Collaborators Group are listed in the Acknowledgement section.

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Introduction

Acute appendicitis (AA) is the most common general surgical emergency worldwide [1]. The lifetime risk of developing AA is 6.7% and 8.6% in females and males respectively [2]. More than 30,000 appendicectomies are performed in England alone each year [3]. Mortality from uncomplicated AA is extremely low at 0.1%; however, mortality increases with delay in presentation [1]. The risk of appendix rupture increases significantly from 36 h after onset of symptoms [4]. Gangrenous appendicitis occurs in 10% of patients, and perforation or abscess is seen in up to one fifth, and both are associated with increased complications [5].

In the UK, operative intervention within 48 h of presentation is recommended for AA [6]. Laparoscopic appendicectomy offers clear advantages over open appendicectomy including less postoperative pain, fewer surgical site infections, decreased length of hospital stay (LOS), and quicker return to normal function [7], and accounts for around 94% and 98% of appendicectomies performed in males and females respectively [8].

There is a growing evidence base that AA not complicated by gangrene or perforation can be managed without surgery [9, 10], is associated with shorter time away from work or education, significantly lower overall complication rate at 5 years after the episode of AA and is cheaper [11, 12]. However, AA can return after successful non-operative management [9]. Despite these considerations, operative treatment remains the first line treatment for nearly all cases of AA in the UK; management with antibiotics usually reserved for those presenting with AA complicated by phlegmon or abscess [6, 8, 13].

SARS-CoV-2 (COVID-19) brought widespread concerns of the spread of infection during aerosol generating procedures (AGPs) such as surgery, and particularly, laparoscopic surgery [14, 15]. In addition, research into COVID positive patients having surgery reported high mortality rates even following minor procedures [16]. Compounding this, there was a lack of personal protective equipment (PPE) for surgeons at the start of the pandemic [17]. Consequently, conservative management with antibiotics was recommended early in the pandemic by UK surgical Royal Colleges as first-line treatment for acute uncomplicated AA. To minimise aerosol generation, open surgery was recommended over laparoscopic surgery when surgery was required [15, 17–20]. Computerised tomography (CT) scan was recommended for diagnosing AA and the exclusion of perforation or other pathology presenting with right iliac fossa pain [20].

The impact of COVID-19 and the impact of recommendations on surgical practice in the UK have not yet

been fully analysed. An early evaluation of any changes in standard UK practice should be performed to assess the safety of any move away from first line operative management of AA. This will inform practice during the rest of the COVID-19 pandemic, and potentially beyond. There is the opportunity to observe if the safe and efficient conservative management of AA previously seen in randomised controlled trials and meta-analysis in Europe and the USA is generalisable in the UK [21–26]. This interim analysis of our study aims to capture the management of AA during the first few weeks of the COVID-19 pandemic lockdown in the UK and to assess the 30-day outcomes [27].

Materials and methods

Study design

A prospective multicentre study on patients aged ≥ 18 years diagnosed either clinically and/or radiologically with AA in a secondary care setting was carried out. Data was collected from patients presenting from the date of the UK Government COVID-19 lockdown on 23rd March 2020. Study registration was delivered by the local principal investigator at each site as either a clinical audit or service evaluation. We collected routine, anonymised data that did not influence clinical care and published the protocol [27].

Outcomes

The primary aim of this interim analysis was to report on the initial management of patients diagnosed with AA from the start of the UK lockdown. Outcomes included conservative or operative management, surgical approach (open or laparoscopic), COVID status, Personal Protective Equipment (PPE) usage, imaging modality (CT scan; ultrasound[USS]), interventional radiology (IR) drain placement, admission to critical care [Level 2 (High Dependency Unit) or 3 (Intensive Care Unit)], 30-day complication rate, mortality, and length of stay (LOS).

Conservative management was defined as initial treatment with antibiotics and/or IR drainage (i.e., not straight to surgery). Acknowledging that the IR drainage group and patients with simple appendicitis are different patient populations, when statistical analysis was performed a comparison was made between conservative management group (IR drain and antibiotics) versus operative group, and antibiotics alone versus operative group. Failure of conservative management occurred when conservative management changed to surgery after ≥ 2 days after initial assessment. Patient demographics and outcomes were analysed in intention to treat by planned initial conservative management, even if they progressed to surgery later, and those started laparoscopically, even where

converted to open intraoperatively, were analysed in the laparoscopic group. LOS is the number of days in hospital during the primary admission and is expressed as median.

Site recruitment

All hospitals in the UK that provide emergency care for patients diagnosed with AA were eligible to participate. Publicity for the project was supported by The Association of Surgeons of Great Britain and Ireland (ASGBI) and the Royal College of Surgeons of England (<https://www.rcseng.ac.uk/coronavirus/rcs-covid-research-group/>). Regional research collaboratives and social media (@covidharem) aided trainee-led recruitment of sites.

Data collection

Patients presenting through the emergency department and surgical assessment units were screened by the local teams as per the inclusion criteria. Confirmation of local approval allowed collaborators to enter fully anonymised data to Research Electronic Data Capture (REDCap, www.project-redcap.org). The database was developed and maintained by the Major Trauma Team at Nottingham University Hospitals, UK.

In the interests of providing evidence to help clinicians with decision making going forward during the pandemic, we have performed an interim analysis of the first 500 patients submitted with 30-day follow-up.

Statistical analysis

The study was done according to Strengthening the Reporting of Observational studies in Epidemiology (STROBE) guidelines for observational studies. Descriptive data was reported as median [interquartile range (IQR)] or number/total (%) as appropriate. For all outcomes, proportions were reported as the number of events/total patients with data due to missing data for some outcomes. When comparing two groups with nominal outcomes Chi-squared or Fisher's exact test was used for inferential testing. For continuous, non-normal data the Mann–Whitney *U* test was used with $p < 0.05$ regarded as the level of statistical significance. Statistical analysis was performed using SPSS Version 26 (IBM, www.ibm.com/uk-en/analytics/spss-statistics-software) and Stata Version 16.1 (StataCorp, www.stata.com).

Results

Patient cohort

From 23rd March 2020 to 1st May 2020, 539 patients were entered from 50 sites across the UK (Fig. 1). Participating

sites were included in this analysis if their data achieved 95% completion, leaving 500 patients from 48 sites. The median age of this cohort was 35 [26–49.75] years and 233 (47%) of patients were female. Other demographics are displayed in Table 1.

Imaging

The majority of patients (425, 85%) had imaging to aid diagnosis. CT was the most commonly performed (353, 71%) throughout all ages in this study, including females under 40 years with no difference in frequency of use by gender (Table 2). An ultrasound scan was performed in 86 (17%), with females more likely to undergo ultrasound than males (76/233 vs. 10/267, $p < 0.001$).

Management approach

Two hundred seventy-one (54%) patients were initially managed conservatively, while the remaining 229 patients (46%) had a plan for surgery within the first day of admission (Fig. 1). Comparison of the initial conservative versus operative group found no difference between the two groups by sex, frailty score, comorbidities or smoking status (Table 1). Patients in the antibiotics alone group were younger than the operative group (median age 34 [25–47] years versus 37 [28.5–52] years, $p = 0.04$).

Within the initial conservative group, 263 (97%) received solely antibiotic therapy with 5 (2%) having an IR drain placed to manage an associated abscess on primary admission; 3 patients requiring an IR drain due to abscess following failed antibiotic management (5, 7 and 8 days after presentation). Failed conservative management occurred in 26 (10%), who went on to have surgery: 2 of these patients had previously had IR drainage. The decision that conservative management had failed and an operation was required, was made between 2–29-day post initial presentation, with 58% (15) of the decisions being made on the 2nd day after admission.

Within the operative group ($n = 211$), the majority had an open procedure 56% (118) versus 44% (93) laparoscopic, with a conversion to open reported in 11 patients (5%).

Figure 2 displays the week by week proportion of patients managed non-operatively and operatively (Fig. 2). During the first week of lockdown 35/74 (47%) of patients were initially managed conservatively, and of those who proceeded to operation, 19 (26%) had laparoscopic surgery and 22 (30%) had open surgery. By the third week of lockdown conservative management peaked at 64% (58/90).

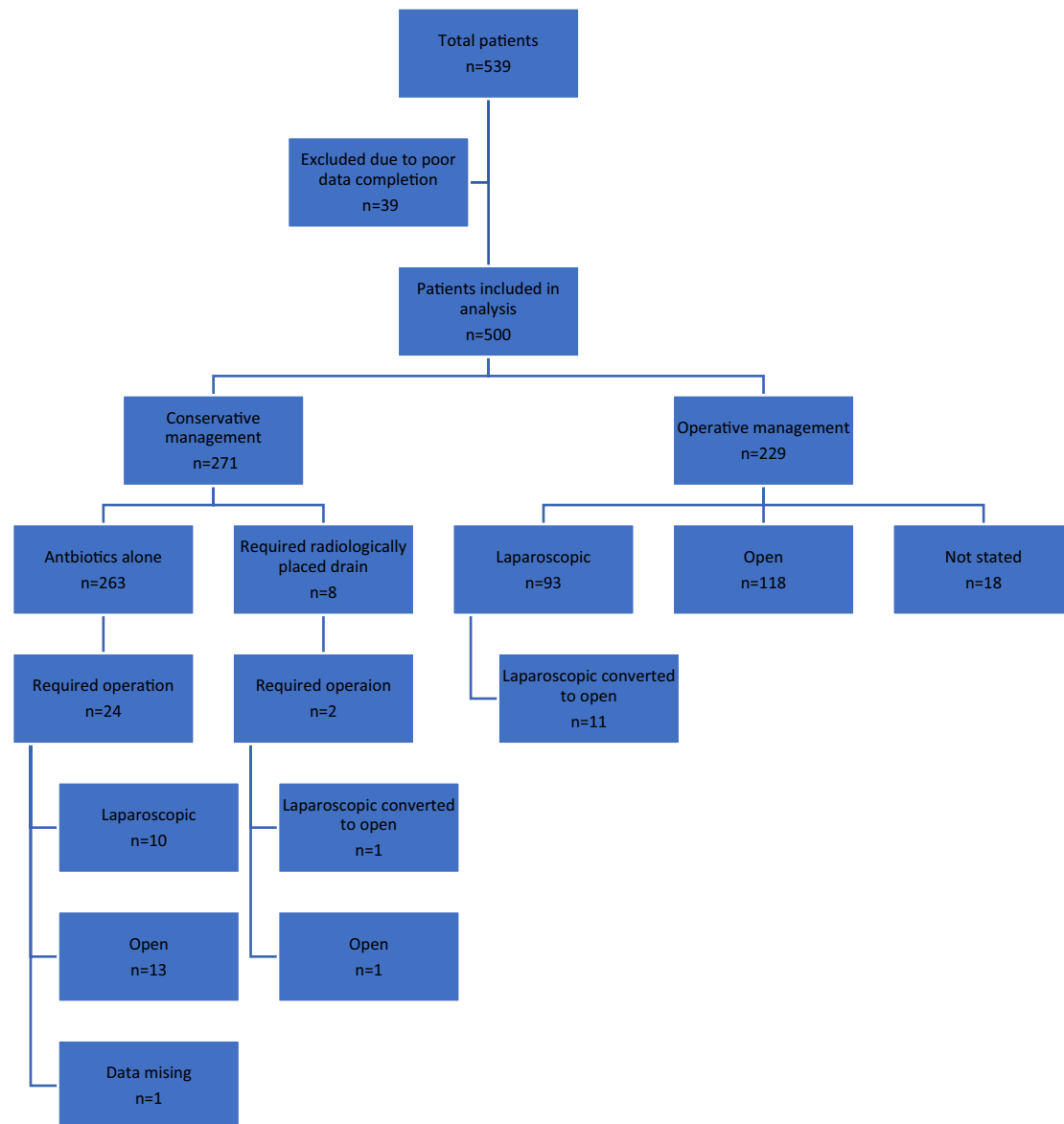


Fig. 1 Study flow chart

Patient outcomes and 30-day follow-up

LOS in the conservatively managed group, whether patients were given antibiotics alone or IR drain placed, was significantly less than in those who had an operation (2 [1–4] days vs. 3 [2–4] days, $p < 0.001$, Table 1). LOS was significantly lower in the laparoscopic appendectomy group than the open appendectomy group (2 [2–4] days vs. 3 [2–5] days, $p < 0.012$, Table 3).

At 30-day follow-up there were no reported deaths (Table 4). The majority of patients who developed complications were in the operative management group. Intra-abdominal collections were reported in 27 (6%) cases (19 in the operative group and 5 in the antibiotics only group,

$p = 0.001$). A reoperation, following initial operative intervention was required in a total of 8 patients (2%), while 6 patients (5 in the operative group), had unplanned admission to critical care. At 30 days, the overall complication rate (excluding readmissions) was 47/219 in the operative group compared to 11/242 in the conservatively managed group ($p < 0.001$).

Of those 26 (10%) managed conservatively that subsequently had an operation, 2 patients required a right hemicolectomy: one for complicated AA and the other for malignancy. Complications in those in the failed conservative management group included, 1 post-operative wound infection, 1 intra-abdominal collection, 1 unplanned and 2 planned admissions to critical care. One patient required

Table 1 Participant characteristics at time of diagnosis and initial management of acute appendicitis

Event/total (%)*	Total (n = 500)	Operative manage- ment (n = 229)	Initial conservative management (n = 271)			
			Antibiotics and IR drain (n = 271)	P value	Antibiotics alone (n = 263)	P value
Age (years) median, range	35 [26–49.75]	37 [28.5–52]	34 [25–48]	0.08	34 [25–47]	0.04
Female	233/500 (47)	99/229 (43)	134/271 (49)	0.18	132/263 (50)	0.13
Body Mass Index kg/m ²						
< 20	22/479 (5)	11/222 (6)	11/257 (3)	0.07	11/249 (4)	0.09
20–25	186/479 (39)	82/222 (37)	104/257 (41)		100/249 (40)	
25–30	165/479 (34)	68/222 (31)	97/257 (38)		94/249 (38)	
30–35	74/479 (15)	45/222 (20)	29/257 (11)		29/249 (12)	
35+	32/479 (7)	16/222 (8)	16/257 (6)		15/249 (6)	
Rockwood score						
Not frail (1–3)	478/500 (96)	220/229 (96)	258/271 (95)	0.87	250/263 (95)	0.84
Pre-frail (4)	14/500 (3)	6/229 (3)	8/271 (3)		8/263 (3)	
Frail (5–9)	8/500 (2)	3/229 (1)	5/271 (2)		5/263 (2)	
Comorbidities						
None reported	450/499 (90)	207/229 (90)	243/270 (90)	1	236/262 (90)	1
Diabetes	18/499 (3)	9/229 (4)	9/270 (3)	0.81	9/262 (3)	0.81
COPD	9/498 (2)	4/228 (2)	5/270 (2)	1	5/262 (2)	1
Myocardial infarction	15/498 (3)	8/313 (3)	7/270 (3)	0.61	7/262 (3)	0.61
Immunosuppressed	11/499 (2)	3/229 (1)	8/270 (3)	0.24	7/262 (3)	0.35
Active cancer	4/499 (0.7)	3/313 (1)	1/270 (0)	0.3	1/262 (0.4)	0.34
Dementia	4/499 (0.7)	2/229 (0.6)	2/270 (0.8)	0.87	2/262 (0.8)	1
Imaging						
No Imaging	77/500 (15)	35/229 (15)	42/271 (16)	0.95	42/263 (16)	0.9
CT scan	353/500 (71)	174/229 (76)	179/271 (66)	0.018	171/263 (65)	0.01
USS	86/500 (17)	26/229 (11)	60/271 (22)	0.002	60/263 (23)	0.001
CT scan and USS	16/500 (3)	6/229 (2)	10/271 (3)	0.61	10/263 (4)	0.61
Admission COVID swab						
Positive	6/159 (4)	3/91 (3)	3/68 (4)	1	3/64 (5)	0.69
Negative	153/159 (96)	88/91 (97)	65/68(96)		61/64 (90)	
Not performed	341/500 (68)	138/229 (60)	203/271(75)	< 0.001	199/263 (76)	< 0.001
Length of stay (days) median, range	3 [1–4]	3 [2–4]	2 [1–4]	< 0.001	2 [1–4]	< 0.001
Managed without admission	57/483 (12)	9/223 (4)	48/260 (18)			< 0.001

Analysis is separated into operative group versus antibiotics and interventional radiological placed drain (IR drain) and operative group versus antibiotics alone. Continuous data is presented as median [IQR]; *p* values calculated by Mann–Whitney–*U*. Categorical data are presented as number/denominator (percentage); *p* values calculated by χ^2 or Fisher's exact test as appropriate

The bold typeset in the table highlights statistical significance

COPD Chronic obstructive pulmonary disease; *CT* Computed tomography, *USS* Ultrasound scan, *IR* Interventional radiology placement of

*Unless otherwise indicated

a second operation. LOS was longer in this group than in the straight to operation group (4 days [2–5.5] vs. 3 [2–4], *p* = 0.03).

Histology demonstrated that 214/241 (89%) of patients operated upon had acute appendicitis. Only 6 patients (3%) had a histologically normal appendix removed, with the

remaining 21 (9%) having alternative pathology found at operation.

Table 2 Computed tomography scans performed by age group

Age group	Computed Tomography (CT) scan	Females	Males	<i>P</i> value
18–39 years	165/300 (55)	74/144 (51)	91/156 (58)	0.23
40–59 years	118/128 (92)	46/52 (88)	72/76 (95)	0.07
60 years and above	70/72 (97)	36/37 (97)	34/35 (97)	1

There was a significant difference between age groups in CT scans performed ($p < 0.001$). Number and total (%) within each age group category; p values calculated by χ^2

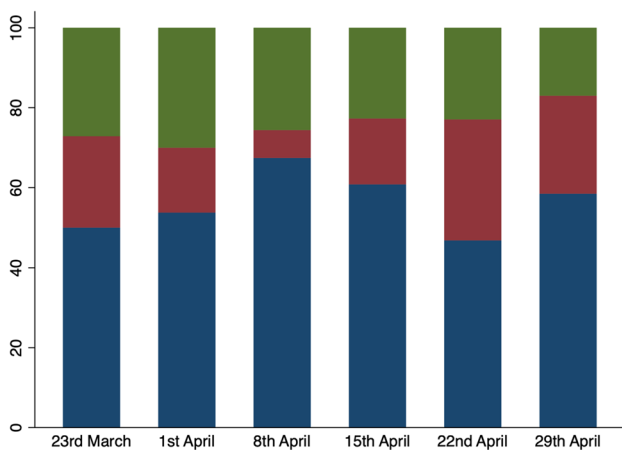


Fig. 2 Percentage of patients treated nonoperatively (blue), with laparoscopic appendicectomy (red), and open appendicectomy (green) on a week by week basis during the pandemic

COVID-19 and PPE

At presentation, 159 (32%) of patients were swabbed for COVID-19 with 6 (4%) positive results. Three of these patients had an operation, and 1 required reoperation. Patients managed conservatively initially were less likely to have a COVID swab (68/271 (25%), $p < 0.001$). Post-presentation, a further 4 patients that had all been swab negative on presentation, were found to have COVID. Of these, 2 had an operation. None of the 10 COVID positive patients required critical care, and there was no mortality at 30 days.

Through the study period use of filtering facepiece assigned protection factor 3 (FFP3) during appendicectomy increased from use in less than 60% of cases to nearly 100% (Fig. 3).

Discussion

Disruption caused by COVID-19 has rapidly changed the management of AA in the UK, with a clear shift to conservative management. Though previously rarely used, conservative management of AA, whether with antibiotics alone or IR drain, has been effectively applied in the majority of patients in this study. With only 10% failing such management and requiring surgery, a shorter LOS and fewer complications compared to those who had initial operations, this interim analysis supports previous studies that have reported non-operative management of AA as a safe and effective option [21–26]. This study reassures surgeons about their decision making during the pandemic, supports routine CT imaging to aid such decisions and demonstrates that this option is generalisable in UK practice and perhaps, signals a reconsideration for first line treatment of AA beyond the pandemic.

Evolution of AA management in the UK during the COVID-19 pandemic

On 20th March 2020, a leading UK surgical professional association issued COVID-19 guidance that stated “non-surgical solutions to be used to avoid surgery where possible” [17]. As a direct result of this, normal practice in the UK of early laparoscopic appendicectomy for adult AA immediately changed to over half of patients being conservatively managed and of those having an operation, the majority having an open procedure. Strengthened by greater patient numbers having imaging, the positive short-term outcomes from conservative management support this guidance. As the pandemic evolves, professional bodies continue to provide updated guidance. This may be reflected in further management shifts that will be seen in the planned longer term follow-up of this study.

Imaging in the diagnosis of appendicitis

Recently published work from the UK reported that almost 40% of patients had no imaging to support a clinical diagnosis of AA [8]. In contrast, our data shows 85% of patients had imaging with CT the favoured modality. Age and sex were not barriers, with CT scan very commonly used in younger patients and more frequently in women of reproductive age than before the pandemic [8]. Comparison with the recent UK observational study on right iliac fossa pain finds only 15% of their female patients had a CT scan, compared to 70% in this study [8]. In addition to aiding a decision for conservative management of AA, it is possible that clinicians wanted a definitive diagnosis of AA before embarking on an operative procedure that carried potentially higher

Table 3 Comparison of operative management for adult acute appendicitis: open versus laparoscopic appendectomy

Event/Total (%)*	Open (n = 133)	Laparoscopic (n = 104)	P value
Age (years) median, range	35 [29–51.5]	37.5 [25–53]	0.934
Female	47/133 (35)	56/104 (54)	0.006
No comorbidity	119/133 (90)	95/104 (91)	0.665
Time of day of operation			
8 am–6 pm	87/133 (66)	62/102 (61)	0.135
6 pm–10 pm	32/133 (24)	20/102 (20)	
10 pm–8 am	14/133 (11)	20/102 (20)	
Operative time			
< 30 min	3/133 (2)	5/102 (5)	0.612
30–60 min	53/133 (40)	44/102 (43)	
60–90 min	57/133 (43)	35/102 (34)	
90–120 min	16/133 (16)	14/102 (14)	
> 120 min	4/133 (3)	4/102 (4)	
Consultant performed procedure	49/133 (37)	33/103 (32)	0.173
Consultant performing or assisting	72/133 (54)	42/103 (41)	0.04
Postoperative care			
Level 2 High Dependency Unit	4/133 (3)	3/103 (3)	0.749
Level 3 Intensive Care Unit	3/133 (2)	1/103 (1)	
Converted to open from laparoscopic	–	11/104 (11)	–
Length of stay (days) median, range	3 [2–5]	2 [2–4]	0.012

Laparoscopic converted to open included in the laparoscopic group. 19 cases excluded due to insufficient information. Continuous data is presented as median [IQR]; *p* values calculated by Mann–Whitney–*U*. Categorical data are presented as number/denominator (percentage); *p* values calculated by χ^2 or Fisher's exact test as appropriate

The bold typeset in the table highlights statistical significance

*Unless otherwise indicated

risks for both patient and operating room staff [15]. CT is highly sensitive at detecting AA and is known to reduce the rate of negative appendectomies significantly, making routine CT imaging perhaps with a lower radiation dose a future consideration [28]. It is not apparent why patients in this study who had an USS were significantly more likely to have conservative management. It may be that they were clinically less convincing and the USS was used to exclude other pathology (especially gynaecological) rather than confirm appendicitis. Alternatively, the sonographers may have preferred USS to make the diagnosis of AA, rather than CT scan, an approach that is recommended by the recent WSES Jerusalem Guidelines [9].

Operative and non-operative management of AA

There were no differences between initial operative and conservative groups in the patient characteristics recorded other than age, highlighting that clinical decision-making for these patients may be multifactorial, including individual interpretation of the new guidelines, local surgical set-up for imaging and theatre access, surgical experience and patient

preference. However, 9 out of 10 of those treated with antibiotics alone needed no operative intervention and stayed for a shorter time in hospital, supporting previous evidence for a non-operative strategy in AA [9, 11, 12]. For those initially conservatively managed who came to surgery, reassuringly this was a small number (10%) with the majority of decisions for surgery taken early during the admission (58% at day 2). Overall, those who failed conservative management did not have significantly poorer outcomes when compared to those having initial surgery. Although a longer LOS was reported in this group, this is to be expected as total LOS will include the conservative management days. Of the 2 right hemicolectomies performed in this group, one was for complicated appendicitis, and the other for malignancy.

The observations in this study are in keeping with previous reports of non-operative management being safe and effective for the majority of patients, with other studies reporting similar success rates, reduced social costs to the patient and less financial cost to the healthcare system if conservative management is considered as a treatment option for AA [9, 11, 12, 21–26]. However, many of these studies were conducted in a research setting, where specific

Table 4 Thirty-day outcome data of initial operative versus conservative management with separate analyses for operative management versus antibiotics and interventional radiology drain placement and operative management versus antibiotics alone

Event/Total (%)	Total (n = 470)	Operative manage- ment (n = 219)	Initial conservative management (n = 251)			
			Antibiotics and IR drain (n = 251)	P value	Antibiotics alone (n = 244)	P value
Collection*	27/470 (6)	19/219 (8)	8/251 (4)	0.01	5/244 (2)	0.001
Collection requiring IR drain	3/470 (1)	0/219 (0)	3/251 (1)	0.252	0/244 (0)	
Collection requiring re-operation	6/470 (1)	5/219 (2)	1/251 (0)	0.102	0/244 (0)	
Ileus**	14/470 (4)	14/219 (8)	0/251 (0)	< 0.001	0/244 (0)	< 0.001
Wound infection***						
Antibiotics	18/470 (3)	16/219 (7)	2/251 (1)	< 0.001	2/242 (0.8)	< 0.001
Wound Opened	9/470 (2)	8/219 (4)	1/251 (1)	0.014	1/244 (0.4)	0.02
Hospital Acquired Pneumonia						
Oral Antibiotics	1/470 (0.2)	1/219 (0.5)	0/251 (0)	1	0/244 (0)	0.47
IV Antibiotics	6/470 (2)	6/219 (3)	0/251 (0)	0.01	0/244 (0)	0.01
Deep Vein Thrombosis/ Pulmonary Embolism	3/470 (0.2)	1/219 (0.5)	2/251 (1)	1	2/244 (1)	1
Reoperation****	8/470 (2)	7/219 (3)	1/251 (0)	0.03	0/244 (0)	0.005
Death	0/470 (0)	0/219 (0)	0/251 (0)	1	0/244 (0)	1
Unplanned level 2/3 care	6/470 (2)	5/219 (4)	1/251 (0)	0.1	1/243 (0.4)	0.11
Post presentation COVID	7/470 (1)	4/219 (2)	3/251 (1)	0.71	3/244 (1)	0.71
Failed conservative management	–	–	26/251 (10)	–	24/244 (9)	–

Categorical data are presented as number/denominator (percentage); *p* values calculated by χ^2 or Fisher's exact test as appropriate

The bold typeset in the table highlights statistical significance

IR Interventional radiolog placement of, IV Intravenous

*Collection refers to an infected fluid collection or intra-abdominal abscess requiring treatment **Ileus was defined as a partial or complete non-mechanical blockade of the small intestine ***Wound infection includes both superficial and deep incisional surgical site infection, defined according to Center for Disease Control criteria as superficial "involving only skin and subcutaneous tissues" and deep as "involving deep structures such as fascia or muscle".**** Reoperation is defined as a return to theatre

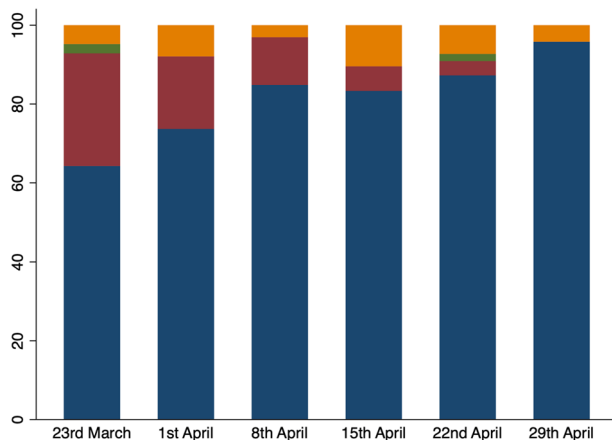


Fig. 3 Type of Personal Protective Equipment (PPE) used when performing appendicectomy by week of the study. FFP3 mask (blue), standard surgical mask (red), none (green), and other (orange)

inclusion criteria were applied. This contrasts sharply with our study, where all adults irrespective of their presentation, age, or CT scan findings were included.

Laparoscopic versus open surgery

Within this study cohort, 56% (133/237) patients having an operation had an open procedure. This is at significant odds with UK practice prior to the pandemic, where only a small number of patients were having open procedures (0.4%) [8]. This is likely due to guidance issues that suggested laparoscopic surgery should be avoided due to concerns about AGPs [17, 29].

COVID-19

The UK has been one of the hardest hit countries during the global COVID-19 pandemic, with the second highest reported mortality rate as of the beginning of June 2020 [30]. Despite this, only 32% of patients were swabbed for COVID-19 at presentation. Local policies for swabbing

were not recorded and are highly likely to have varied across the UK during the early phase of the pandemic. Follow-up work may show greater rates of testing.

Only 4% tested positive for the virus with poorer outcomes not reported, irrespective of whether having surgery or conservative management. This is in contrast to recent reports of an increase in mortality after even minor to moderate surgery [16]. However, most patients with AA are young and otherwise fit, which would predispose them to better outcomes than those reported for a broad variety of operative interventions [16].

Strengths and weaknesses

This is a large multicentre study of current AA practice in the UK with high data completion. Early analysis has allowed prompt reporting of a change in surgical practice to be demonstrated as safe in the short-term, reassuring surgeons in the current complex clinical climate and aiding decision-making in the case of a second viral wave; these findings are likely to be generalisable to other populations who normally defer to operative intervention, but whose practice has been disrupted by COVID-19, such as the United States and Europe.

The authors acknowledge that this is a pragmatic study, where bias may exist in the decision for initial management. Such bias may be a consequence of the initial conservative management group presenting with less severe symptoms compared to the initial operative group. We also appreciate that patients having laparoscopy without appendicectomy may not have been included in this data set, leading to an underestimation of the negative appendicectomy rate. One further limitation of the study is the lack of assessment of confounding variables on outcome data; this will be explored in future work with a larger dataset and maturation of follow-up to assess for longer term outcomes.

Conclusions

The COVID-19 pandemic markedly disrupted the usual surgical management of acute appendicitis in the UK with conservative management favoured. In this setting non-operative management of AA appears to be an effective first line treatment regardless of sex, co-morbidity or frailty, with only a minority requiring surgery as a second line. Disruption allows change, and these early findings should inform the continued management of AA during the COVID-19 pandemic and perhaps beyond.

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Authors contributions Study conception, protocol development and publication, data collection, data interpretation, and writing and revision of this manuscript were performed by HJ-E, MH, SJM and GMT. AA was involved in data collection, database design and build, and data interpretation and review of the manuscript. LD and RC contributed to data collection proforma, data collection and writing of the manuscript. BD and JNL were involved in data analysis and writing of the manuscript. LP, JC, SR, SH and SL contributed to study conception, data collection and review of the manuscript. All authors approved the final manuscript. GMT is the guarantor.

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Compliance with ethical standards

Conflict of interest The authors declare that they have no conflict of interest.

Ethics approval This is an observational study; therefore, no ethics approval was required.

Informed consent For this type of study, formal consent is not required.

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References

1. Collaborative GlobalSurg (2006) Mortality of emergency abdominal surgery in high-, middle- and low-income countries. *Br J Surg* 103:971–988
2. Addiss DG, Shaffer N, Fowler BS, Tauxe RV (1990) The epidemiology of appendicitis and appendectomy in the United States. *Am J Epidemiol* 132:910–925
3. NHS Digital. Hospital Admitted Patient Care Activity 2018–2019 [Internet]. 2019 [cited 26 May 2020]. <https://digital.nhs.uk/data-and-information/publications/statistical/hospital-admitted-patient-care-activity/2018-19#resources>
4. Bickell NA, Aufses AH, Rojas M, Bodian C (2006) How time affects the risk of rupture in appendicitis. *J Am Coll Surg* 202:401–406
5. De Wijkerslooth EML, De Jonge J, Van den Boom AL et al (2019) Postoperative outcomes of patients with nonperforated gangrenous appendicitis: a national multicenter prospective cohort analysis. *Dis Colon Rectum* 62:1363–1370
6. The Association of Surgeons of Great Britain and Ireland (2014) Commissioning guide: Emergency general surgery (acute abdominal pain)
7. Jaschinski T, Mosch CG, Eikermann M, Neugebauer EAM, Sauerland S (2018) Laparoscopic versus open surgery for suspected appendicitis. *Cochrane Database of Syst Rev*. <https://doi.org/10.1002/14651858.CD001546.pub4>
8. RIFT study group (2019) Evaluation of appendicitis risk prediction models in adults with suspected appendicitis. *Br J Surg* 107:73–86
9. Di Saverio S, Podda M, De Simone B et al (2020) Diagnosis and treatment of acute appendicitis: 2020 update of the WSES Jerusalem guidelines. *World J Emerg Surg* 2020:15
10. Di Saverio S, Birindelli A, Kelly MD et al (2016) WSES Jerusalem guidelines for diagnosis and treatment of acute appendicitis. *World J Emerg Surg* 11:1–25
11. Salminen P, Tuominen R, Paajanen H et al (2018) Five-year follow-up of antibiotic therapy for uncomplicated acute appendicitis in the APPAC randomized clinical trial. *JAMA J Am Med Assoc* 320:1259–1265

12. Sippola S, Gronroos J, Tuominen R et al (2017) Economic evaluation of antibiotic therapy versus appendicectomy for the treatment of uncomplicated acute appendicitis from the APPAC randomized clinical trial. *Br J Surg* 104:1355–1361
13. Andersson RE, Petzold MG (2007) Nonsurgical treatment of appendiceal abscess or phlegmon: a systematic review and meta-analysis. *Ann Surg* 246:741–748
14. Vigneswaran Y, Prachand VN, Posner MC, Matthews JB, Husain M (2020) What is the appropriate use of laparoscopy over open procedures in the current COVID-19 climate? *J Gastrointest Surg*. <https://doi.org/10.1007/s11605-020-04592-9>
15. De Simone B, Chouillard E, Di Saverio S et al (2020) Emergency surgery during the COVID-19 pandemic: what you need to know for practice. *Ann R Col Surg Eng* 102:323–332
16. COVIDSurg Collaborative (2020) Mortality and pulmonary complications in patients undergoing surgery with perioperative SARS-CoV-2 infection: an international cohort study. *Lancet*. [https://doi.org/10.1016/S0140-6736\(20\)31182-X](https://doi.org/10.1016/S0140-6736(20)31182-X)
17. Hettiaratchy S, Deakin D (2020) Joint guidance for surgeons: guidance for surgeons working during the COVID-19 pandemic from the Surgical Royal Colleges of the United Kingdom and Ireland. Royal College of Surgeons of England. <https://www.rcseng.ac.uk/coronavirus/joint-guidance-for-surgeons-v1/>
18. COVIDSurg Collaborative (2020) Global guidance for surgical care during the COVID-19 pandemic. *Br J Surg*. <https://doi.org/10.1002/bjs.11646>
19. Spinelli A, Pellino G (2020) COVID-19 pandemic: perspectives on an unfolding crisis. *Br J Surg*. <https://doi.org/10.1002/bjs.11627>
20. Association of Upper GI Surgery of Great Britain and Ireland (2020) AUGIS Guidelines: management algorithm for patients with clinically suspected appendicitis during Covid-19 pandemic. AUGIS. <https://www.augis.org/augis-guidelines/>
21. Prechal D, Damirov F, Grilli M, Ronellenfitch U (2019) Antibiotic therapy for acute uncomplicated appendicitis: a systematic review and meta-analysis. *Int J Colorectal Dis* 34:963–971
22. Yang Z, Sun F, Ai S, Wang J, Guan J, Liu S (2019) Meta-analysis of studies comparing conservative treatment with antibiotics and appendectomy for acute appendicitis in the adult. *BMC Surg* 19(1):110
23. Poon SHT, Lee JWY, Ng KM et al (2017) The current management of acute uncomplicated appendicitis: SHOULD there be a change in paradigm? A systematic review of the literatures and analysis of treatment performance. *World J Emerg* 12:46
24. Podda M, Cillara N, Di Saverios S et al (2017) Antibiotics-first strategy for uncomplicated acute appendicitis in adults is associated with increased rates of peritonitis at surgery. A systematic review with meta-analysis of randomized controlled trials comparing appendectomy and non-operative management with antibiotics. *Surgeon* 15:303–314
25. Harnoss JC, Zelenka I, Probst P et al (2017) Antibiotics versus surgical therapy for uncomplicated appendicitis: systematic review and meta-analysis of controlled trials (PROSPERO 2015:CRD42015016882). *Ann Surg* 265:889–900
26. Rollins KE, Varadhan KK, Neal KR, Lobo DN (2016) Antibiotics versus appendicectomy for the treatment of uncomplicated acute appendicitis: an updated meta-analysis of randomised controlled trials. *World J Surg* 40:2305–2318
27. The HAREM Steering Group (2020) The HAREM (Had Appendicitis and Resolved/Recurred Emergency Morbidity/Mortality) Study. *Br J Surg*. <https://doi.org/10.1002/bjs.11711>
28. Kim K, Kim YH, Kim SY et al (2012) Low-dose abdominal CT for evaluating suspected appendicitis. *N Engl J Med* 366:1596–1605
29. Griin M, Alderson D, Taylor J, Mealey K (2020) Joint guidance for surgeons: updated intercollegiate general surgery guidance on COVID-19. Royal College of Surgeons of England. <https://www.rcseng.ac.uk/coronavirus/joint-guidance-for-surgeons-v2/>
30. Burn-Murdoch J, Giles C (2020) UK suffers second-highest death rate from coronavirus. *Financial Times* (Internet). <https://www.ft.com/content/6b4c784e-c259-4ca4-9a82-648ffde71bf0>

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