

1 **Estimating the potential overdiagnosis and overtreatment of acute appendicitis in**
2 **Thailand using a secondary data analysis of service utilization before, during and after**
3 **the COVID-19 lockdown policy**

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23 **Abstract (273 words)**

24 **Introduction:** Acute appendicitis is one of the most common surgical emergencies; however,
25 optimal diagnosis and treatment of acute appendicitis remains challenging. We used the
26 coronavirus disease 2019 (COVID-19) lockdown policy as a natural experiment to explore
27 potential overdiagnosis and overtreatment of acute appendicitis in Thailand. The aim of this
28 study was to estimate the potential overdiagnosis and overtreatment of acute appendicitis in
29 Thailand by examining service utilization before, during, and after the COVID-19 lockdown
30 policy.

31 **Methods:** A secondary data analysis of patients admitted with acute appendicitis under the
32 Universal Coverage Scheme (UCS) in Thailand over a 6-year period between 2016 and 2021
33 was conducted. The trend of acute appendicitis was plotted using a 14-day rolling average of
34 daily cases. Patient characteristics, clinical management, and outcomes were descriptively
35 presented and compared among three study periods, namely pre-pandemic, lockdown, and
36 post-lockdown

37 **Results:** The number of overall acute appendicitis cases decreased from 25,407 during pre-
38 pandemic to 22,006 during lockdown (13.4% reduction) and 21,245 during post-lockdown
39 (16.4% reduction). This reduction was mostly due to a lower incidence of uncomplicated
40 acute appendicitis, whereas cases of generalized peritonitis were scarcely affected by the
41 pandemic. There was an increasing trend towards the usage of diagnostic computerized
42 tomography for acute appendicitis but no significant difference in treatment modalities and
43 complication rates.

44 **Conclusion:** The stable rates of generalized peritonitis and complications during the COVID-
45 19 lockdown, despite fewer admissions overall, suggest that there may have been
46 overdiagnosis and overtreatment of acute appendicitis in Thailand. Policy makers could use

- 47 these findings to improve clinical practice for acute appendicitis in Thailand and support the
- 48 efficient utilization of surgical services in the future, especially during pandemics.

49 **Introduction**

50 Acute appendicitis is one of the most common surgical emergencies [1]. Diagnosis of acute
51 appendicitis is based on history and physical, laboratory evaluation, and imaging [2];
52 however, it can be difficult to diagnose due to similar presentation to other acute abdomen
53 pathologies and the poor predictive value of associated laboratory testing [3]. Rates of
54 negative appendectomy (appendectomy for uninflamed appendix) around 20% have been
55 reported in Lithuania and Israel [4,5]. Such overdiagnosis and overtreatment not only impact
56 the healthcare budget but also quality of life, as unnecessary appendectomy is associated with
57 higher mortality [6].

58 The coronavirus disease 2019 (COVID-19) pandemic has affected healthcare systems
59 across the world since the virus was first detected in December 2019 [7]. In Thailand, the
60 peak of the first wave was reached in March 2020, resulting in an announcement of full-scale
61 national lockdown from March 26 to May 3, 2020 [8]. Although these measures were
62 effective in containing the pandemic [9,10], they also had significant impact on the delivery
63 of health services. For instance, surgical services were affected by a reduced surgical
64 workforce, infection control measures, and elective cancellations [11].

65 During the COVID-19 pandemic, countries such as China, Egypt and Saudi Arabia
66 observed an overall decrease in total number of patients with acute appendicitis alongside a
67 concurrent increase in complications, such as gangrene or perforation [12–14]. We therefore
68 hypothesise that diagnosis and treatment of cases during the pandemic was restricted to
69 essential procedures only. Thus, we used the lockdown policy as a natural experiment to
70 explore the potential overdiagnosis and overtreatment of acute appendicitis in Thailand. This
71 information can be used to improve clinical practice for acute appendicitis in Thailand as well
72 as to assist in the planning on how to support surgical services during future pandemics. This
73 study aimed to estimate the potential overdiagnosis and overtreatment of acute appendicitis in

74 Thailand by examining service utilization before, during and after the COVID-19 lockdown
75 policy.

76 **Methods**

77 This study was a secondary data analysis of inpatient data from patients covered under the
78 Universal Coverage Scheme (UCS) in Thailand, comprising approximately 80% of the
79 population. Data were obtained from the National Health Security Office (NHSO) over a 6-
80 year period between 2016 and 2021. Patient-level data were anonymized and de-identified.
81 All patients diagnosed with acute appendicitis were identified using the 10th revision
82 International Classification of Diseases (ICD-10) code K35. Acute appendicitis was classified
83 into acute appendicitis with generalized peritonitis (ICD-10: K35.2), acute appendicitis with
84 localized peritonitis (ICD-10: K35.3), and uncomplicated acute appendicitis (ICD-10:
85 K35.9). The study time frame was divided into three periods: 1) the first period was defined
86 as March-June of 2020 which captured national lockdown in Thailand; 2) for comparison,
87 March-June in 2021 was assigned as a post-lockdown; and 3) March-June in 2019 was used
88 as a pre-pandemic comparison.

89 All statistical analyses were performed using R version 4.1.3 (R Foundation for
90 Statistical Computing, Vienna, Austria) [15]. The overall trend of acute appendicitis was
91 plotted from 2016 to 2021 and stratified by age group (children aged under 18 and adults).
92 Rolling averages of daily cases were computed using the ‘rollmean’ function of the zoo
93 package with a 14-day centered rolling window [16]. Descriptive statistics, namely frequency
94 and percentage, were used to explore patients’ characteristics (age, sex, hospital type, health
95 region, and comorbidities), clinical management (diagnostic imaging and treatment
96 modalities), and outcome in the three study periods (pre-pandemic, lockdown, and post-
97 lockdown). Diagnostic imaging included computerized tomography of abdomen (ICD-9-CM:
98 8801) and ultrasound of abdomen (ICD-9-CM: 8876). Treatment modalities in this study

99 covered open appendectomy (ICD-9-CM: 47.09), laparoscopic appendectomy (ICD-9-CM:
100 47.01), injection of antibiotics (ICD-9-CM: 99.21), and drainage of appendiceal abscess
101 (ICD-9-CM: 47.2). Patient outcomes were length of stay, cost of hospitalisation, and
102 complications (e.g., infection following a procedure (ICD-10: T81.4), in-hospital death, and
103 30-day readmission). A p-value of 0.05 or lower was considered statistically significant.

104 **Results**

105 A total of 25,407 (pre-pandemic), 22,006 (lockdown), and 21,245 (post-lockdown) patients
106 were admitted under the UCS with acute appendicitis (Table 1). The number of overall cases
107 decreased by 13.4% during lockdown and 16.4% post-lockdown in comparison to the pre-
108 pandemic period. There was a significant change in hospital type and health region of
109 admission. The proportion of referred patients slightly declined over time, while the
110 prevalence of hypertension, diabetes mellitus, and chronic kidney disease marginally
111 increased.

112 **Table 1. Characteristics of patients with acute appendicitis during three study periods.**

Characteristics	Pre-pandemic, 2019	Lockdown, 2020	Post-lockdown, 2021	P-value
	(N = 25407)	(N = 22006)	(N = 21245)	
Age (years), median (IQR)	29 (15,52)	30 (16,53)	30 (16,53)	0.003
Children (<18 years), n (%)	8132 (32.0)	6723 (30.6)	6538 (30.8)	0.001
Male, n (%)	12191 (48.0)	10753 (48.9)	10361 (48.8)	0.105
Hospital type, n (%)				< 0.001
Central hospital	9000 (35.4)	7309 (33.2)	6862 (32.3)	
General hospital	10304 (40.6)	9247 (42.0)	9002 (42.4)	
Community hospital	4046 (15.9)	3769 (17.1)	3798 (17.9)	
Private hospital	475 (1.9)	408 (1.9)	264 (1.2)	
Hospital outside MOPH	1575 (6.2)	1265 (5.7)	1314 (6.2)	
Health service centers and private clinics	7 (0)	8 (0)	5 (0)	
Health region, n (%)				< 0.001
1	3032 (11.9)	2671 (12.1)	2576 (12.1)	

2	1333 (5.2)	997 (4.5)	1113 (5.2)	
3	955 (3.8)	937 (4.3)	892 (4.2)	
4	1485 (5.8)	1226 (5.6)	1247 (5.9)	
5	1743 (6.9)	1547 (7.0)	1402 (6.6)	
6	2216 (8.7)	1786 (8.1)	1599 (7.5)	
7	2222 (8.7)	2053 (9.3)	2092 (9.8)	
8	2418 (9.5)	2200 (10.0)	2136 (10.1)	
9	2932 (11.5)	2456 (11.2)	2434 (11.5)	
10	2159 (8.5)	1853 (8.4)	1763 (8.3)	
11	1808 (7.1)	1637 (7.4)	1515 (7.1)	
12	1703 (6.7)	1459 (6.6)	1416 (6.7)	
13	1394 (5.5)	1175 (5.3)	1051 (4.9)	
14	7 (0)	9 (0)	9 (0)	
Referred patients, n (%)	2266 (8.9)	1866 (8.5)	1734 (8.2)	0.013
Comorbidities, n (%)				
Hypertension	1774 (7.0)	1690 (7.7)	1695 (8.0)	< 0.001

Diabetes mellitus	790 (3.1)	808 (3.7)	747 (3.5)	0.002
Cardiac disease	123 (0.5)	116 (0.5)	104 (0.5)	0.778
Chronic kidney disease	308 (1.2)	334 (1.5)	306 (1.4)	0.012
Diagnosis, n (%)				< 0.001
Acute appendicitis with generalized peritonitis	2424 (9.5)	2213 (10.1)	1989 (9.4)	
Acute appendicitis with localized peritonitis	11597 (45.6)	10728 (48.8)	11096 (52.2)	
Uncomplicated acute appendicitis	11386 (44.8)	9065 (41.2)	8160 (38.4)	

114 Fig 1 shows the overall trend of acute appendicitis from 2016 to 2021. The number of
115 patients with acute appendicitis with localized peritonitis (green line) and uncomplicated
116 acute appendicitis (blue line) significantly decreased during late-March to early-April 2020,
117 then started to climb back in late-April 2020. Another significant drop in the number of acute
118 appendicitis with localized peritonitis and uncomplicated acute appendicitis was observed in
119 early-May 2021. In comparison, the trend of acute appendicitis with generalized peritonitis
120 was quite stable. In the age-stratified analysis, similar trends were observed in adults;
121 however, no significant decrease in number of cases was observed among children (Fig 2).

122 **Fig 1. The overall trend of acute appendicitis from 2016 to 2021.**

123 **Fig 2. The trend of acute appendicitis from 2016 to 2021, stratified by age group.**

124 Clinical management and outcomes for patients with acute appendicitis during the three
125 study periods are presented in Table 2. There was an increasing trend towards the use of
126 computerized tomography during the pandemic from 7.4% pre-pandemic to 9.9% during
127 lockdown, and 12.6% post-lockdown, whereas the usage of ultrasonography remained steady
128 (~4%). During the three study periods, open appendectomy was the most performed
129 treatment modality (~88%). The proportions of open appendectomy, laparoscopic
130 appendectomy, and antibiotics were small (< 1%) and remained unchanged over time. The
131 median of length of stay was 3 days (interquartile range; IQR = 2 to 4 days) across the three
132 study periods. The median cost of hospital stay paid by NHSO marginally increased from
133 10,682 Thai Baht (THB) (IQR = 8,675 to 11,386 THB) during the pre-pandemic period to
134 11,112 THB (IQR = 9,095 to 12,260 THB) during lockdown, and 12,196 THB (IQR = 9,861
135 to 14,295 THB) in the post-lockdown period. The rates of in-hospital mortality, infection
136 following a procedure, and 30-day readmission over the three study periods were
137 approximately 0.7%, 0.2%, and 7.0%, respectively.

138 **Table 2. Clinical management and patient outcomes for acute appendicitis during the three study periods.**

	Pre-pandemic, 2019	Lockdown, 2020	Post-lockdown, 2021	P-value
	(N = 25407)	(N = 22006)	(N = 21245)	
Diagnostic imaging, n (%)				
Computerized tomography	1870 (7.4)	2180 (9.9)	2686 (12.6)	< 0.001
Ultrasonography	984 (3.9)	880 (4.0)	897 (4.2)	0.157
Treatment modalities, n (%)				
Open appendectomy	22223 (87.5)	19416 (88.2)	18642 (87.7)	0.039
Laparoscopic appendectomy	256 (1.0)	243 (1.1)	257 (1.2)	0.114
Antibiotics	60 (0.2)	41 (0.2)	29 (0.1)	0.047
Drainage of appendiceal abscess	57 (0.2)	64 (0.3)	56 (0.3)	0.356
Length of stay (days), median (IQR)	3 (2,4)	3 (2,4)	3 (2,4)	0.472
Hospitalisation costs paid by NHSO (THB), median (IQR)	10682.2 (8675.2,11385.8)	11112.2 (9094.6,12260.5)	12195.7 (9861.0,14295.4)	< 0.001

Complications, n (%)

Infection following a procedure	145 (0.6)	170 (0.8)	140 (0.7)	0.026
In-hospital death	66 (0.3)	43 (0.2)	54 (0.2)	0.297
30-day readmission	1899 (7.5)	1508 (6.8)	1500 (7.1)	0.027

139 Abbreviation: NHSO, National Health Security Office

140 **Discussion**

141 In this study, we observed a significant decrease in the number of uncomplicated acute
142 appendicitis cases during the COVID-19 lockdown. On the other hand, rates of acute
143 appendicitis with generalized peritonitis and acute appendicitis in children were not affected
144 by the pandemic. We further observed an increasing trend in the use of diagnostic
145 computerized tomography for acute appendicitis, but no significant change in treatment
146 modality, during the COVID-19 pandemic.

147 During the COVID-19 lockdown, there was a decrease in the number of admissions due
148 to acute appendicitis without generalized peritonitis, which aligns with the findings of
149 previous studies. The reduction in other studies varied from 12.9% to 40.7% depending on
150 study setting (single hospital, multiple centers, or population-based) and country (China,
151 Croatia, Germany, Israel, and United States) [12,17–20]. Although additional measures to
152 visit hospitals during lockdown and fear of being infected with COVID-19 from a hospital
153 visit may have led to a delay in diagnosis and more severe presentation, we did not find any
154 significant increase in the rate of acute appendicitis with generalized peritonitis or
155 complications either during or after the lockdown period. This finding may reflect the effort
156 to maintain effective communication between the Thai government and the public, as well as
157 continued access to health services, even during the lockdown [10]. This finding contrasts
158 with previous findings from China, Nepal, and the United States, which reported a higher
159 incidence of acute complex appendicitis (e.g., suppurative, perforated, or gangrenous
160 appendix) [13,20,21], which could be due to differences in settings and healthcare systems.

161 With the finding of stable complication rates overtime despite a decrease in hospital
162 admissions for acute appendicitis, we conclude that there may be significant overdiagnosis
163 and overtreatment for acute appendicitis in Thailand. By subtracting the number of cases in
164 2020 (an ideal scenario) from those in 2019 and multiplying by the median cost of hospital

165 stay for acute appendicitis paid for by NHSO in 2020, we estimate the medical care costs to
166 the UCS of potential overdiagnosis of acute appendicitis to be approximately 60 million THB
167 annually. We hypothesise that overdiagnosis and unnecessary treatment may result from the
168 pressure on higher level hospitals (central and general hospitals) to perform surgery for
169 patients referred with suspected acute appendicitis. Surgeons at higher level hospitals may
170 risk being sued if they miss a case of acute appendicitis cases; hence, opting for patient
171 observation either at home or in hospital may not be worth the risk. The hospital experiences
172 no financial loss from overdiagnosis and overtreatment under the UCS scheme, as inpatient
173 services are reimbursed based on case-mix classifying patient conditions into groups
174 according to resources consumed [22].

175 To the best of our knowledge, this is the first study to explore potential overdiagnosis
176 and overtreatment of acute appendicitis in Southeast Asia using COVID-19 lockdown as a
177 natural experiment. Given that potential overdiagnosis and overtreatment not only impact the
178 healthcare budget but can also lead to adverse health outcomes from unnecessary surgery, our
179 results can be used to improve clinical practice for acute appendicitis in Thailand in the
180 future. Nevertheless, there are some limitations in data availability in this study as we used
181 claims data from hospitals. The UCS database accounts for only approximately 80% of the
182 Thai population and does not cover the private sector. We used only data available from
183 existing administrative database, which did not include clinical information. For example,
184 there was no information regarding pathological confirmation of diagnosis and the number of
185 cases might have been overestimated. Additionally, diagnoses of acute appendicitis with or
186 without perforation or rupture, or with peritoneal abscess, were recorded using the same ICD-
187 10 code version 2016 (K35.3). Thus, we could not explore the trends of each diagnosis,
188 which have different degrees of severity. Finally, only descriptive analysis was performed in

189 this study; therefore, risk factors for overdiagnosis and overtreatment and other clinical
190 information should be explored in future research.

191 This study demonstrated a significant reduction in the number of admissions with
192 uncomplicated acute appendicitis during and after the COVID-19 lockdown in Thailand,
193 whereas the number with more severe generalized peritonitis and complications remained
194 stable over time. These findings suggest that there was potential overdiagnosis and
195 overtreatment of acute appendicitis cases in Thailand before the pandemic. With a better
196 understanding of emergency surgical service utilization during the COVID-19 pandemic,
197 policy makers could improve clinical practice for acute appendicitis in Thailand and optimize
198 the utilization of surgical services in future.

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220 **References**

- 221 1. Cervellin G, Mora R, Ticinesi A, Meschi T, Comelli I, Catena F, et al. Epidemiology
222 and outcomes of acute abdominal pain in a large urban Emergency Department:
223 retrospective analysis of 5,340 cases. *Ann Transl Med.* 2016;4: 362.
224 doi:10.21037/atm.2016.09.10
- 225 2. Moris D, Paulson EK, Pappas TN. Diagnosis and management of acute appendicitis in
226 adults: a review. *JAMA.* 2021;326: 2299–2311. doi:10.1001/jama.2021.20502
- 227 3. Pittman-Waller VA, Myers JG, Stewart RM, Dent DL, Page CP, Gray GA, et al.
228 Appendicitis: why so complicated? Analysis of 5755 consecutive appendectomies. *Am*
229 *Surg.* 2000;66: 548–554.
- 230 4. Zoarets I, Poluksht N, Halevy A. Does selective use of computed tomography scan
231 reduce the rate of “white” (negative) appendectomy? *Isr Med Assoc J IMAJ.* 2014;16:
232 335–337.
- 233 5. Kryzauskas M, Danys D, Poskus T, Mikalauskas S, Poskus E, Jotautas V, et al. Is acute
234 appendicitis still misdiagnosed? *Open Med.* 2016;11: 231–236. doi:10.1515/med-2016-
235 0045
- 236 6. Andersson MN, Andersson RE. Causes of short-term mortality after appendectomy: a
237 population-based case-controlled study. *Ann Surg.* 2011;254: 103–107.
238 doi:10.1097/SLA.0b013e31821ad9c4
- 239 7. World Health Organization. Archived: WHO Timeline - COVID-19. 27 Apr 2020 [cited
240 24 Mar 2022]. Available: [https://www.who.int/news/item/27-04-2020-who-timeline---](https://www.who.int/news/item/27-04-2020-who-timeline---covid-19)
241 [covid-19](https://www.who.int/news/item/27-04-2020-who-timeline---covid-19)
- 242 8. Rajatanavin N, Tuangratananon T, Suphanchaimat R, Tangcharoensathien V.
243 Responding to the COVID-19 second wave in Thailand by diversifying and adapting

- 244 lessons from the first wave. *BMJ Glob Health*. 2021;6: e006178. doi:10.1136/bmjgh-
245 2021-006178
- 246 9. Haddawy P, Lawpoolsri S, Sa-ngamuang C, Yin MS, Barkowsky T, Wiratsudakul A, et
247 al. Effects of COVID-19 government travel restrictions on mobility in a rural border
248 area of Northern Thailand: A mobile phone tracking study. *PLOS ONE*. 2021;16:
249 e0245842. doi:10.1371/journal.pone.0245842
- 250 10. Marome W, Shaw R. COVID-19 response in Thailand and its implications on future
251 preparedness. *Int J Environ Res Public Health*. 2021;18: 1089.
252 doi:10.3390/ijerph18031089
- 253 11. Rausei S, Ferrara F, Zurleni T, Frattini F, Chiara O, Pietrabissa A, et al. Dramatic
254 decrease of surgical emergencies during COVID-19 outbreak. *J Trauma Acute Care*
255 *Surg*. 2020;89: 1085–1091. doi:10.1097/TA.0000000000002923
- 256 12. Zheng Z, Bi JT, Liu YQ, Cai X. The impact of COVID-19 pandemic on the treatment of
257 acute appendicitis in China. *Int J Colorectal Dis*. 2022;37: 215–219.
258 doi:10.1007/s00384-021-04031-4
- 259 13. Yang Y, Li Y, Du X. Acute complex appendicitis during the COVID-19 epidemic: A
260 single-institution retrospective analysis based on real-world data. *Am J Emerg Med*.
261 2021;46: 74–77. doi:10.1016/j.ajem.2021.03.022
- 262 14. El Nakeeb A, Emile SH, AbdelMawla A, Attia M, Alzahrani M, ElGamdi A, et al.
263 Presentation and outcomes of acute appendicitis during COVID-19 pandemic: lessons
264 learned from the Middle East—a multicentre prospective cohort study. *Int J Colorectal*
265 *Dis*. 2022;37: 777–789. doi:10.1007/s00384-022-04108-8
- 266 15. R Core Team. *R: A language and environment for statistical computing*. Vienna,
267 Austria: R Foundation for Statistical Computing; 2021. Available: [https://www.R-](https://www.R-project.org/)
268 [project.org/](https://www.R-project.org/)

- 269 16. Zeileis A, Grothendieck G. zoo: S3 infrastructure for regular and irregular time series. J
270 Stat Softw. 2005;14: 1–27. doi:10.18637/jss.v014.i06
- 271 17. Tankel J, Keinan A, Blich O, Koussa M, Helou B, Shay S, et al. The decreasing
272 incidence of acute appendicitis during COVID-19: A retrospective multi-centre study.
273 World J Surg. 2020;44: 2458–2463. doi:10.1007/s00268-020-05599-8
- 274 18. Köhler F, Acar L, van den Berg A, Flemming S, Kastner C, Müller S, et al. Impact of
275 the COVID-19 pandemic on appendicitis treatment in Germany—a population-based
276 analysis. Langenbecks Arch Surg. 2021;406: 377–383. doi:10.1007/s00423-021-02081-
277 4
- 278 19. Bosak Veršić A, Šestan M, Čepić I, Nikolić H, Bukvić N, Sršen Medančić S, et al.
279 Characteristics of acute appendicitis before and during the COVID-19 pandemic: single
280 center experience. Cirocchi R, editor. Emerg Med Int. 2022;2022: 1–5.
281 doi:10.1155/2022/4541748
- 282 20. Orthopoulos G, Santone E, Izzo F, Tirabassi M, Pérez-Caraballo AM, Corriveau N, et
283 al. Increasing incidence of complicated appendicitis during COVID-19 pandemic. Am J
284 Surg. 2021;221: 1056–1060. doi:10.1016/j.amjsurg.2020.09.026
- 285 21. Baral S, Chhetri RK, Thapa N. Comparison of acute appendicitis before and within
286 lockdown period in COVID-19 era: A retrospective study from rural Nepal. Chen RJ,
287 editor. PLOS ONE. 2021;16: e0245137. doi:10.1371/journal.pone.0245137
- 288 22. Tangcharoensathien V, Suphanchaimat R, Thammatacharee N, Patcharanarumol W.
289 Thailand's Universal Health Coverage Scheme. Econ Polit Wkly. 2012;47: 53–57.

290 **List of tables and figures**

291 **Table 1** Characteristics of patients with acute appendicitis during the three study periods

292 **Figure 1** The overall trend of acute appendicitis from 2016 to 2021

293 **Figure 2** The trend of acute appendicitis from 2016 to 2021, stratified by age group

294 **Table 2** Clinical management and patient outcomes for acute appendicitis during the three

295 study periods