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## **Population-based cohort study of outcomes following cholecystectomy for benign gallbladder diseases**

CholeS Study Group, West Midlands Research Collaborative\*

\*Members of the CholeS Study Group and West Midlands Research Collaborative are co-authors of this study and may be found under the heading Collaborators

*Correspondence to:* Mr E. A. Griffiths, Department of Upper Gastrointestinal Surgery, University Hospitals Birmingham NHS Foundation Trust, Birmingham B15 2WB, UK (e-mail: ewen.griffiths@uhb.nhs.uk)

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**Background:** The aim was to describe the management of benign gallbladder disease and identify characteristics associated with all-cause 30-day readmissions and complications in a prospective population-based cohort

**Methods:** Data were collected on consecutive patients undergoing cholecystectomy in acute UK and Irish hospitals between 1 March and 1 May 2014. Potential explanatory variables influencing all cause 30-day readmissions and complications were analysed by means of

multilevel, multivariable logistic regression modelling using a two-level hierarchical structure with patients (level 1) nested within hospitals (level 2).

**Results:** Data were collected on 8909 patients undergoing cholecystectomy from 167 hospitals. Some 1451 cholecystectomies (16.3 per cent) were performed as an emergency, 4165 (46.8 per cent) as elective operations, and 3293 patients (37.0 per cent) had at least one previous emergency admission, but were operated on a delayed basis. The readmission and complication rates at 30 days were 7.1 per cent (633 of 8909) and 10.8 per cent (962 of 8909) respectively. Both readmissions and complications were independently associated with increasing ASA fitness grade, duration of surgery, and increasing numbers of emergency admissions with gallbladder disease before cholecystectomy. No identifiable hospital characteristics were linked to readmissions and complications.

**Conclusion:** Readmissions and complications following cholecystectomy are common and associated with patient and disease characteristics.

## **+A: Introduction**

Benign gallbladder disease is a major global health burden<sup>1</sup>. It is estimated that for every 100 000 of the world's population, 115 patients undergo cholecystectomy every year. In England alone, approximately 70 000 cholecystectomies are performed annually<sup>2</sup>. Among those fit for surgery, patients follow one of three pathways from presentation to definitive treatment linked to emergency admissions with gallbladder disease: emergency cholecystectomy during an emergency admission; elective cholecystectomy with no previous emergency admission; or delayed cholecystectomy following one or more previous emergency admissions with gallbladder pathology.

Variations in outcomes exist following these different patient pathways. When performed with no previous emergency admission, either as an index emergency or elective operation, cholecystectomy is associated with fewer gallbladder-specific complications, a shorter total length of hospital stay and similar operative complications compared with those among patients who have had one or more emergency admissions<sup>3-5</sup>. Many of these studies were conducted in specialized centres by enthusiasts. In contrast, population-level data from Hospital Episodes Statistics (HES)<sup>2</sup> and a retrospective study from Scotland<sup>6</sup> both suggested that emergency cholecystectomy may be associated with poorer surgical outcomes. The reasons for these differences are unclear, but may be due to patient, disease, surgical and hospital variables not fully realized in administrative data sets.

Patient outcomes are used to measure quality of healthcare, such as readmission, complication, reoperation and mortality rates<sup>6-13</sup>. The mortality rate is low following cholecystectomy, and therefore a poor measure of quality in this cohort<sup>14</sup>. In contrast, reducing hospital readmissions and complications after surgery can lower hospital costs and improve patient satisfaction<sup>15-17</sup>. The causes of readmission and complications after cholecystectomy have been poorly studied in large prospective series, but may vary according to patient pathways<sup>11</sup>.

Over the past 8 years, trainee-led networks in the UK have adopted a collaborative approach to deliver prospective population-level data collection, and measure patient, disease, surgical and hospital variables with short-term endpoints such as readmissions and complications<sup>18</sup>. Using these networks, a prospective, population-based cohort study was conducted to describe management of patients with benign gallbladder disease<sup>19</sup>, and identify patient, disease, surgical and hospital-related characteristics that might be associated with all-cause 30-day readmission and complications.

## **+A: Methods**

The study was carried out as described previously<sup>19</sup>. The study protocol did not require research registration as anonymized, observational data were collected. This was confirmed by the online National Research Ethics Service decision tool (<http://www.hra-decisiontools.org.uk/research/>), and further supported by written confirmation and advice from the Research and Development Director at University Hospitals Birmingham NHS Foundation Trust, UK. The study was registered as a clinical audit or service evaluation at each participating hospital under the supervision of a named senior investigator (consultant surgeon).

## **+B: Inclusion and exclusion criteria**

All patients undergoing cholecystectomy for benign gallbladder diseases in acute UK and Irish hospitals participating in this study between 1 March and 1 May 2014 were included. Patients were grouped according to the timing of cholecystectomy: emergency cholecystectomy, defined as cholecystectomy during an emergency admission; delayed cholecystectomy, defined as a scheduled cholecystectomy following an emergency admission with gallbladder disease in the preceding 12 months; or elective cholecystectomy, defined as a planned elective admission for cholecystectomy following referral by a family doctor and addition to the routine surgical waiting list from an outpatient department visit. Open, laparoscopic and laparoscopic procedures converted to open surgery were included. Cholecystectomies for a known gallbladder cancer or as a part of another surgical procedure. (such as pancreaticoduodenectomy) were excluded.

## **+B: Outcome measures**

Planned analyses comprised a description of current management of benign gallbladder disease with readmission and complication rates. Variations in 30-day complication and readmission rates were studied as well as predictive factors for these events. A list of recorded complications with their definitions is available in *Table S1* (supporting information).

#### **+B: Data quality**

To standardize data quality, a quality assurance programme was developed<sup>19</sup>. This included a detailed study protocol, pilot phase, and a requirement for a minimum of 95 per cent data completeness at submission. Case ascertainment, including readmission to a different hospital from where cholecystectomy was performed, and data accuracy, were validated by independent investigators at selected hospitals, who checked data from 10 per cent of patients against original medical records. These independent investigators were not involved in the original data collection.

#### **+B: Explanatory variables**

Patient, disease and hospital characteristics were considered as potential explanatory variables influencing the performance of emergency cholecystectomy. A full list including definitions has been published previously<sup>19</sup>. Briefly, patient characteristics included: age, sex, ASA fitness grade (I, normal healthy patient; II, mild systemic disease; III, severe systemic disease; IV, severe systemic disease that is a constant threat to life; V, moribund patient who is not expected to survive without the operation) and BMI (less than 17.9 kg/m<sup>2</sup>, underweight; 18.0–24.9 kg/m<sup>2</sup>, normal; 25.0–29.9 kg/m<sup>2</sup>, overweight; 30.0–34.9 kg/m<sup>2</sup>, moderate obesity; 35.0 kg/m<sup>2</sup> and above, severe or very severe obesity). The following disease characteristics were considered: indication (biliary colic, cholecystitis, pancreatitis, common bile duct (CBD) stones), ultrasound findings (gallbladder wall thickness, dilated

CBD), findings from other radiological investigations (CT, magnetic retrograde cholangiopancreatography (MRCP), endoscopic retrograde cholangiopancreatography (ERCP)) and total number of emergency admissions with biliary symptoms in the 12 months before cholecystectomy.

Surgical characteristics included degree of difficulty according to the Nassar score<sup>20</sup> and surgical approach (laparoscopic, open or laparoscopic converted to open surgery). Hospital characteristics, including type (university or not), specialist hepatobiliary centre and acute hospital status, were recorded along with the number of consultants within the reporting hospital performing cholecystectomy, hospital country, number of beds within the reporting hospital (less than 100, 101–500, 501–1000, more than 1000) and presence of an on-site ERCP service. The hospital's policy regarding the ease of performing intraoperative cholangiography and use of dedicated emergency gallbladder operating lists were considered. The consultant's presence, specialty and grade were also recorded at the time of cholecystectomy.

#### **+B: Statistical analysis**

Results are reported in accordance with the STROBE statement for observational studies<sup>21</sup>. Crude rates of emergency cholecystectomy at each hospital were calculated for all patients. Descriptive statistics were obtained for all variables. Median values with interquartile range (i.q.r.) are reported. The  $\chi^2$  test or Fisher's exact test, as appropriate, was used to identify differences between categorical variables.

To investigate the relationship between 30-day readmission and complication rates and the variables studied, data were analysed using multilevel, multivariable logistic regression modelling. Both 30-day readmission and complication rates were recorded as binary outcome measures, requiring a logistic analytical approach. These data comprise a

two-level hierarchical structure, with patients at level 1, nested within hospitals at level 2. Multilevel (hierarchical) modelling was selected owing to the structure of the data. Patient outcomes are not independent of each other as patients are clustered within hospitals. This clustering introduces multilevel dependency or correlation among patient observations that can have implications for model parameter estimates. Because of this dependency among observations, multilevel modelling has advantages over other approaches, such as logistic regression analysis, which assume that observations are independent.

Before model building, variables were investigated for potential confounding relationships using correlation and scatter plots. As the variables included in this study could be viewed as being on a causal path potentially linking each variable to the outcome, they were assessed during model building and none were excluded. Variable collinearity was also tested using variance inflation factor scores.

Variables were included in the models if they were found to be significant at the 5 per cent level. The forward and back Collett method<sup>22</sup> was used to select variables for inclusion.

To investigate between-hospital variation, the random intercept model was extended to a random coefficient model, including each variable in turn, allowing the factor to vary across hospitals. Results are expressed as adjusted odds ratios (ORs) with 95 per cent confidence intervals (c.i.).

All two-way interactions were assessed and no significant interactions were identified in the final models. Model testing was performed using likelihood ratio tests, Wald tests, and residual and deviance plots. Owing to the binary nature of the outcome data, the multilevel logistic regression model deviates from the normal assumptions that underlie regression models and as such these were not tested for. However, testing for the assumption of



independent co-variance structure between random effects was not possible as no explanatory variables were included at level 2.

Missing data accounted for 0.8 per cent of the data set. Models were fitted with missed data and therefore all available data were included.

All statistical analyses were performed using Stata® version 12 (StataCorp, College Station, Texas, USA). The multilevel, multivariable logistic regression modelling was carried out in MLwiN version 2.14 (<http://www.cmm.bristol.ac.uk/MLwiN>).

#### +A: **Results**

Data were collected on 8909 patients undergoing cholecystectomy from 167 hospitals. Case ascertainment and accuracy of collected data was above 95.3 and 99.2 per cent respectively, compared with information from 10 per cent of patients checked independently against the original medical records.

The median age of the cohort was 51 (38–64) years. The numbers of patients undergoing emergency, delayed and elective cholecystectomy were 1451 (16.3 per cent), 3293 (37.0 per cent) and 4165 (46.8 per cent) respectively. A total of 3713 patients (41.7 per cent) had at least one emergency admission with acute gallbladder disease before cholecystectomy. Elective cholecystectomy was mainly undertaken for biliary colic, in younger women with a low ASA fitness grade, who did not require additional radiological tests or interventions (such as MRCP or ERCP) compared with patients who had delayed and emergency cholecystectomy (*Table 1*). Patients with biliary colic underwent elective surgery (73.3 per cent) more frequently, whereas cholecystitis and pancreatitis were more frequently treated by delayed cholecystectomy (53.0 and 64.0 per cent respectively). Elective cholecystectomy was more likely to be completed laparoscopically, was less difficult as

assessed by the operating surgeon, and with fewer intraoperative complications (bile contamination, stones spilled, bleeding, bowel and significant biliary injury), compared with delayed and emergency cholecystectomy (*Table 2*). The difference in incidence of bile duct injury, however, did not reach statistical significance. The median duration of an elective procedure was 60 (45–79) min compared with 65 (50–90) and 80 (60–110) min for delayed and emergency cholecystectomy respectively.

#### **+B: Outcomes at 30 days**

Readmission and complication rates at 30 days were 7.1 per cent (633 of 8909) and 10.8 per cent (962 of 8909) respectively (*Table 3*). The rates of readmission, overall complications and individual complications at 30 days were higher in the delayed and emergency groups than in the elective group. Both readmission and complication rates differed according to the number of surgical admissions (*Fig. 1*). Readmission rates among patients with no, one or at least two surgical admissions before cholecystectomy were 6.1, 7.2 and 12.9 per cent respectively; corresponding complication rates were 8.9, 11.8 and 19.1 per cent.

Reoperations were more frequent following delayed cholecystectomy, although this difference was not statistically significant. The 30-day mortality rate was higher when cholecystectomy was performed as an emergency or delayed operation, but again this difference was not statistically significant.

#### *+C: Thirty-day readmission rate*

The null random intercept model containing no exploratory variables was used to test variation across the 167 hospitals, and demonstrated a statistically significant between-hospital variance (likelihood ratio statistic 11.57,  $P = 0.003$ ). Estimates of the hospital effects obtained from the null model showed the hospital effects in rank order together with 95 per cent confidence intervals (*Fig. 2*). Only two of 167 hospitals (1.2 per cent) had 30-day

readmission rates higher than the rest. When the model was extended to include explanatory variables, duration of surgery and greater number of surgical admissions before the index surgery were associated with a higher risk of readmission, whereas there was no difference in readmissions between emergency and delayed operation (*Table 4*). Elective cholecystectomy, younger patients and ASA score I–II were associated with a lower risk of readmission. The variables included in this multilevel model accounted for 97 per cent of the detected variation. BMI, indication for cholecystectomy and hospital-related variables were not independently associated with readmission.

#### *+C: Thirty-day complication rate*

The null random intercept model demonstrated significant between-hospital variance (likelihood ratio statistic 74.80,  $P = 0.003$ ). Eleven hospitals (6.6 per cent) had a significantly different 30-day complication rate from the others (*Fig. 3*). When the model was extended to include explanatory variables, age, ASA grade, surgical admission type, number of admissions and duration of surgery were again associated with complications (*Table 5*). In addition, having an open or a laparoscopic procedure converted to open surgery and a greater degree of operative difficulty were associated with a higher risk of complications. These variables accounted for 93 per cent of the detected variation. Other patient categories, disease and hospital factors were not independently associated with the outcomes measured.

#### **+A: Discussion**

This study evaluated the current management and outcomes of cholecystectomy for benign gallbladder disease in the UK and Ireland. Readmissions and complications were common and varied across the 167 hospitals studied. An emergency or delayed operation and greater numbers of surgical admissions before the index surgery were associated with a higher risk of readmission and complications. Intraoperative features, such as difficulty, duration and

operative method, were also associated with poorer outcomes across hospitals. Importantly, the hospital characteristics analysed did not influence these associations.

Readmission and complication rates at 30 days were similar to those reported previously. Other studies<sup>2,6,12,23–26</sup> have described patient and hospital factors, both independently and in combination, linked to complications and readmissions. Factors associated with readmission or complication in the present study were analysed by means of hierarchical modelling. The advantage of this type of analysis was that it allowed the effects of patient variables on certain outcomes to be considered independently of hospital variables that may also influence outcomes<sup>27</sup>. Age, ASA grade, duration of surgery, operative approach and degree of operative difficulty were all associated with worse outcomes, as demonstrated previously<sup>25,28</sup>.

Increasing numbers of emergency admissions before cholecystectomy were independently associated with both readmission and complication rates here. This supports a pathway in which definitive cholecystectomy should be performed during the first admission. Emergency cholecystectomy was mainly carried out in younger patients and those presenting with cholecystitis, whereas older patients and those requiring further investigations tended to be discharged and brought back for a delayed cholecystectomy.

Delayed cholecystectomy can result in emergency readmissions and poorer outcomes<sup>29–33</sup>. Logistical barriers to emergency cholecystectomy, such as lack of prompt access to specialist investigations and emergency theatre availability, appear to explain differences in service provision. Hospital policy and surgeon or patient preference may contribute to delaying cholecystectomy. Increasing the numbers of emergency cholecystectomies by addressing modifiable barriers, such as reducing the number of surgical

admissions with gallbladder pathology before cholecystectomy, would improve outcomes. This is generalizable to healthcare providers outside the UK and Ireland.

In contrast to previous studies, there was no statistical difference in readmissions and complications when emergency and delayed cholecystectomies were compared, suggesting that the delayed cohort was not homogeneous. Definitions of emergency and delayed cholecystectomy varied in previous studies. Some considered emergency or early cholecystectomy to comprise operations performed within 48–72 h of symptom onset<sup>5</sup>. A population-based study<sup>4</sup> of 14 200 patients in Canada showed that patients experienced fewer complications when cholecystectomy was performed within 7 days of hospital admission. A study<sup>3</sup> in Switzerland demonstrated that a 6-day delay in performing emergency cholecystectomy was associated with a higher conversion rate (12 per cent at day 0 *versus* 18 per cent at day 6), more complications (5.7 *versus* 13 per cent respectively) and the reoperation rate almost double in the delayed compared with the emergency group (0.8 *versus* 0.5 per cent).

There are limitations to the present study. The data represent a 2-month snapshot of practice and this may account for unexpected observations, such as the inverse association between age and risk of readmission. However, short intensive data collection allowed surgical teams to contribute meaningful numbers of patients with high levels of accuracy and without additional resources. The number of patients who did not undergo cholecystectomy is unknown. It seems intuitive that those with less co-morbidity are more likely to be offered cholecystectomy. The overall models accounted for nearly 90 per cent of the variations in outcomes seen across the hospitals studied, suggesting the presence of other factors not characterized in this study. Despite these limitations, the accuracy of the data set was validated independently and it contained variables not typically collected in similar studies.

Sweden<sup>34</sup>, Denmark<sup>35</sup> and Switzerland<sup>25</sup> use prospective registries of cholecystectomy for continual quality improvement. Although this is costly and time-consuming, the results of this prospective population-based cohort study demonstrate that cholecystectomy following multiple surgical admissions is linked to 30-day complications and readmissions. Measuring numbers of surgical admissions before cholecystectomy provides a simple quality improvement metric. There needs to be a focus on offering emergency rather than delayed cholecystectomy for patients presenting with acute benign gallbladder disease.

#### **+A: Collaborators**

Study management group: R. S. Vohra (Trent Oesophago-Gastric Unit, Nottingham University Hospitals NHS Trust, Nottingham, UK); S. Pasquali (Surgical Oncology Unit, Veneto Institute of Oncology IOV-IRCCS, Padova, Italy); A. J. Kirkham (Cancer Research UK Clinical Trials Unit, University of Birmingham, Birmingham, UK); P. Marriott, M. Johnstone, P. Spreadborough (West Midlands Research Collaborative, Academic Department of Surgery, University of Birmingham, Birmingham, UK); D. Alderson (Academic Department of Surgery, University of Birmingham, Birmingham, UK); E. A. Griffiths (Department of Upper Gastrointestinal Surgery, University Hospitals Birmingham NHS Foundation Trust, Birmingham, UK).

Other members of the CholeS Study Group and West Midlands Research Collaborative are as follows. England: S. Fenwick, M. Elmasry, Q. Nunes, D. Kennedy (Aintree University Hospital NHS Foundation Trust); R. Basit Khan, M. A. S. Khan (Airedale General Hospital); C. J. Magee, S. M. Jones, D. Mason, C. P. Parappally (Wirral University Teaching Hospital); P. Mathur, M. Saunders, S. Jamel, S. Ul Haque, S. Zafar (Barnet and Chase Farm Hospital); M. H. Shiwani, N. Samuel, F. Dar, A. Jackson (Barnsley District General Hospital); B. Lovett,

S. Dindyal, H. Winter, S. Rahman (Basildon University Hospital); K. Wheatley, T. Nieto, S. Ayaani (Sandwell and West Birmingham Hospitals NHS Trust); H. Youssef, R. S. Nijjar, H. Watkin, D. Naumann, S. Emeshi; P. B. Sarmah, K. Lee, N. Joji (Heart of England Foundation NHS Trust); J. Heath, R. L. Teasdale, C. Weerasinghe (Blackpool Teaching Hospitals NHS Foundation Trust); P. J. Needham, H. Welbourn, L. Forster, D. Finch (Bradford Teaching Hospitals NHS Foundation Trust); J. M. Blazeby, W. Robb, A. G. K. McNair, A. Hrycaiczuk (University Hospitals Bristol NHS Trust); A. Charalabopoulos, S. Kadiramanathan, C.-B. Tang, N. V. G. Jayanthi, N. Noor (Broomfield Hospital); B. Dobbins, A. J. Cockbain, A. Nilsen-Nunn, J. de Siqueira (Calderdale and Huddersfield NHS Trust); M. Pellen, J. B. Cowley, W.-M. Ho, V. Miu (Hull and East Yorkshire NHS Trust); T. J. White, K. A. Hodgkins, A. Kinghorn (Chesterfield Royal Hospital NHS Foundation Trust); M. G. Tutton, Y. A. Al-Abed, D. Menzies, A. Ahmad, J. Reed, S. Khan (Colchester Hospital University NHS Foundation Trust); D. Monk, L. J. Vitone, G. Murtaza, A. Joel (Countess of Chester NHS Foundation Trust); S. Brennan, D. Shier, C. Zhang, T. Yoganathan (Croydon Health Services NHS Trust); S. J. Robinson, I. J. D. McCallum, M. J. Jones, M. Elsayed, L. Tuck, J. Wayman, K. Carney (North Cumbria University Hospitals Trust); S. Aroori, K. B. Hosie, A. Kimble, D. M. Bunting (Plymouth Hospitals NHS Trust); A. S. Fawole, M. Basheer, R. V. Dave, J. Sarveswaran, E. Jones, C. Kendal (Mid Yorkshire NHS Trust); M. P. Tilston, M. Gough, T. Wallace, S. Singh, J. Downing, K. A. Mockford, E. Issa, N. Shah, N. Chauhan (Northern Lincolnshire and Goole NHS Foundation Trust); T. R. Wilson, A. Forouzanfar, J. R. L. Wild, E. Nofal, C. Bunnell, K. Madbak (Doncaster and Bassetlaw Hospitals NHS Foundation Trust); S. T. V. Rao, L. Devoto, N. Siddiqi, Z. Khawaja (Dorset County Hospital NHS Foundation Trust); J. C. Hewes, L. Gould, A. Chambers, D. Urriza Rodriguez (North Bristol NHS Trust); G. Sen, S. Robinson, K. Carney, F. Bartlett (Freeman Hospital); D. M. Rae, T. E. J. Stevenson, K. Sarvananthan (Frimley Park Hospital NHS Trust); S. J. Dwerryhouse, S. M. Higgs, O. J.

Old, T. J. Hardy, R. Shah, S. T. Hornby, K. Keogh, L. Frank (Gloucestershire Hospitals NHS Trust); M. Al-Akash, E. A. Upchurch (Great Western Hospitals NHS Foundation Trust); R. J. Frame, M. Hughes, C. Jelley (Harrogate and District NHS Foundation Trust); S. Weaver, S. Roy, T. O. Sillo, G. Galanopoulos (Wye Valley NHS Trust); T. Cuming, P. Cunha, S. Tayeh, S. Kaptanis (Homerton University Hospital NHS Trust); M. Heshaihi, A. Eisawi, M. Abayomi; W. S. Ngu, K. Fleming, D. Singh Bajwa (Tees Hospitals NHS Foundation Trust); V. Chitre, K. Aryal, P. Ferris (Paget University Hospitals NHS Foundation Trust); M. Silva, S. Lammy, S. Mohamed, A. Khawaja, A. Hussain, M. A. Ghazanfar, M. I. Bellini (Oxford University NHS Trust); H. Ebdewi, M. Elshaer, G. Gravante, B. Drake (Kettering General Hospital NHS Foundation Trust); A. Ogedegbe, D. Mukherjee, C. Arhi, L. Giwa Nusrat Iqbal (Barking, Havering and Redbridge University Hospitals NHS Trust); N. F. Watson, S. Kumar Aggarwal, P. Orchard, E. Villatoro (Kings Mill Hospital); P. D. Willson, K. Wa, J. Mok, T. Woodman, J. Deguara (Kingston Hospital NHS Foundation Trust); G. Garcea, B. I. Babu, A. R. Dennison, D. Malde, D. Lloyd (University Hospitals of Leicester NHS Trust); J. P. Slavin, R. P. Jones, L. Ballance, S. Gerakopoulos (Leighton Hospital, Mid Cheshire Hospitals NHS Foundation Trust); P. Jambulingam, S. Mansour, N. Sakai, V. Acharya (Luton and Dunstable University Hospital NHS Foundation Trust); M. M. Sadat, L. Karim, D. Larkin, K. Amin (Macclesfield District General Hospital); A. Khan, J. Law, S. Jamdar, S. R. Smith, K. Sampat, K. M O'shea (Central Manchester NHS Foundation Trust); M. Manu, F. M. Asprou, N. S. Malik, J. Chang, M. Johnstone (Royal Wolverhampton Hospitals NHS Trust); M. Lewis, G. P. Roberts, B. Karavadra, E. Photi (Norfolk and Norwich University Hospitals NHS Foundation Trust); J. Hewes, L. Gould, A. Chambers, D. Rodriguez (North Bristol NHS Trust); D. A. O'Reilly, A. J. Rate, H. Sekhar, L. T. Henderson, B. Z. Starmer, P. O. Coe, S. Tolofari, J. Barrie (Pennine Acute NHS Trust); G. Bashir, J. Sloane, S. Madanipour, C. Halkias, A. E. J. Trevatt (North Middlesex Trust); D. W. Borowski, J. Hornsby, M. J. Courtney, S. Virupaksha (North



Tees and Hartlepool NHS Foundation Trust); K. Seymour, S. Robinson, H. Hawkins, S. Bawa, P. V. Gallagher, A. Reid, P. Wood (Northumbria Healthcare NHS Foundation Trust); J. G. Finch, J. Parmar, E. Stirland (Northampton General Hospital NHS Trust); J. Gardner-Thorpe, A. Al-Muhktar, M. Peterson, A. Majeed (Sheffield Teaching Hospitals NHS Foundation Trust); F. M. Bajwa, J. Martin, A. Choy, A. Tsang (Peterborough City Hospital); N. Pore, D. R. Andrew, W. Al-Khyatt, C. Taylor S. Bhandari, A. Chambers, D. Subramaniam (United Lincolnshire Hospitals NHS Trust); S. K. C. Toh, N. C. Carter, S. J. Mercer, B. Knight (Portsmouth Hospitals NHS Trust); V. Vijay, S. Alagaratnam, S. Sinha, S. Khan (The Princess Alexandra Hospital NHS Trust); S. S. El-Hasani, A. A. Hussain (King's College Hospital NHS Foundation Trust); V. Bhattacharya, N. Kansal, T. Fasih, C. Jackson (Gateshead Health NHS Foundation Trust); M. N. Siddiqui, I. A. Chishti, I. J. Fordham, Z. Siddiqui (Lewisham and Greenwich NHS Trust); H. Bausbacher, I. Geogloma, K. G. (Queen Elizabeth Hospital NHS Trust); G. Tsavellas, P. Basynat, A. Kiran Shrestha, S. Basu, A. Chhabra Mohan Harilingam, M. Rabie, M. Akhtar (East Kent Hospitals University NHS Foundation Trust); P. Kumar, S. F. Jafferbhoy, N. Hussain, S. Raza (Burton Hospitals NHS Foundation Trust); M. Haque, I. Alam, R. Aseem, S. Patel, M. Asad (Royal Albert Edward Infirmary, Wigan Wroughton and Leigh NHS Trust); M. I. Booth, W. R. Ball, C. P. J. Wood, A. C. Pinho-Gomes (Royal Berkshire NHS Foundation Trust); A. Kausar, M. Rami Obeidallah (East Lancashire Hospital Trust); J. Varghase, J. Lodhia, D. Bradley, C. Rengifo, D. Lindsay (Royal Bolton Hospital NHS Foundation Trust); S. Gopalswamy, I. Finlay, S. Wardle, N. Bullen (Royal Cornwall NHS Trust); S. Y. Iftikhar, A. Awan, J. Ahmed, P. Leeder (Royal Derby NHS Foundation Trust); G. Fusai, G. Bond-Smith, A. Psica, Y. Puri (Royal Free, London); D. Hou, F. Noble, K. Szentpali, J. Broadhurst (Hampshire Hospital NHS Foundation Trust); R. Date, M. R. Hossack, Y. Li Goh, P. Turner, V. Shetty (Lancashire Teaching Hospitals NHS Foundation Trust); M. Riera, C. A. W. Macano, A. Sukha (Royal Shrewsbury Hospital); S. R. Preston, J. R. Hoban, D. J.

Puntis, S. V. Williams (Royal Surrey County Hospital NHS Foundation Trust); R. Krysztopik, J. Kynaston, J. Batt, M. Doe (Royal United Hospital Bath NHS Trust); A. Goscimski, G. H. Jones, S. R. Smith, C. Hall (Salford Royal NHS Foundation Trust); N. Carty, J. Ahmed, S. Panteleimonitis (Salisbury Hospital Foundation Trust); R. T. Gunasekera, A. R. G. Sheel, H. Lennon, C. Hindley (Southport and Ormskirk Hospital NHS Trust); M. Reddy, R. Kenny, N. Elkheir, E. R. McGlone (St George's Healthcare NHS Trust); R. Rajaganeshan, K. Hancorn, A. Hargreaves (St Helens and Knowsley Teaching Hospitals NHS Trust); R. Prasad, D. A. Longbotham, D. Vijayanand, I. Wijetunga (Leeds Teaching Hospitals); P. Ziprin, C. R. Nicolay, G. Yeldham, E. Read (Imperial College Healthcare NHS Trust); J. A. Gossage, R. C. Rolph, H. Ebied, M. Phull (St Thomas' Hospital, London); M. A. Khan, M. Popplewell, D. Kyriakidis, A. Hussain (Mid Staffordshire NHS Foundation Trust); N. Henley, J. R. Packer, L. Derbyshire, J. Porter (Stockport NHS Foundation Trust); S. Appleton, M. Farouk, M. Basra (Bucks Healthcare NHS Trust); N. A. Jennings, S. Ali, V. Kanakala (City Hospitals Sunderland NHS Foundation Trust); H. Ali, R. Lane, R. Dickson-Lowe, P. Zarsadias (Tunbridge Wells and Maidstone NHS Trust); D. Mirza, S. Puig, K. Al Amari, D. Vijayan, R. Sutcliffe, R. Marudanayagam (University Hospital Birmingham NHS Foundation Trust); Z. Hamady, A. R. Prasad, A. Patel (University Hospital Coventry and Warwickshire NHS Trust); D. Durkin, P. Kaur, L. Bowen (University Hospital of North Staffordshire NHS Trust); J. P. Byrne, K. L. Pearson, T. G. Delisle, J. Davies (University Hospital Southampton NHS Foundation Trust); M. A. Tomlinson, M. A. Johnpulle, C. Slawinski (University Hospitals of Morecambe Bay); A. Macdonald, J. Nicholson, K. Newton, J. Mbuvi (University Hospital South Manchester NHS Foundation Trust); A. Farooq, B. Sidhartha Mothe, Z. Zafrani, D. Brett (Warrington and Halton Hospitals NHS Trust); J. Francombe, P. Spreadborough, J. Barnes, M. Cheung (South Warwickshire NHS Foundation Trust); A. Z. Al-Bahrani, G. Preziosi, T. Urbonas (Watford General Hospital); J. Alberts, M. Mallik, K. Patel, A. Segaran, T. Doulias (West Suffolk NHS

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Gibson, G. A. Nicholson, D. G. Vass (Crosshouse Hospital, Ayrshire and Arran); A. J. Grant, D. J. Holroyd, M. A. Jones, C. M. L. R. Sutton (Dr Gray's Hospital); P. O'Dwyer, F. Nilsson (Gartnavel General Hospital); B. Weber, T. K. Williamson, K. Lalla, A. Bryant (Gilbert Bain Hospital); C. R. Carter, C. R. Forrest, D. I. Hunter (Glasgow Royal Infirmary); A. H. Nassar, M. N. Orizu, K. Knight, H. Qandeel (Monklands Hospital); S. Suttie, R. Belding, A. McClarey (Ninewells Hospital); A. T. Boyd, G. J. K. Guthrie, P. J. Lim, A. Luhmann (Perth Royal Infirmary); A. J. M. Watson, C. H. Richards, L. Nicol, M. Madurska (Raigmore Hospital); E. Harrison, K. M. Boyce, A. Roebuck, G. Ferguson (Royal Infirmary of Edinburgh); P. Pati, M. S. J. Wilson, F. Dalgaty, L. Fothergill (Stracathro Hospital); P. J. Driscoll, K. L. Mozolowski, V. Banwell, S. P. Bennett (Victoria Hospital, Kirkcaldy); P. N. Rogers, B. L. Skelly, C. L. Rutherford, A. K. Mirza (Western Infirmary Glasgow). Wales: T. Lazim, H. C. C. Lim, D. Duke, T. Ahmed (Bronglais General Hospital); William D. Beasley, M. D. Wilkinson, G. Maharaj, C. Malcolm (Glangwili General and Prince Philip Hospital); T. H. Brown, G. M. Shingler, N. Mowbray, R. Radwan (Morrison and Singleton Hospitals); P. Morcous, S. Wood, A. Kadhim (Princess of Wales Hospital); D. J. Stewart, A. L. Baker, N. Tanner, H. Shenoy (Wrexham Maelor Hospital). Data validators: S. Hafiz, J. A. De Marchi, D. Singh-Ranger, E. Hisham, P. Ainley, S. O'Neill, J. Terrace, S. Napetti, B. Hopwood, T. Rhys, J. Downing, O. Kanavati, M. Coats, D. Aleksandrov, C. Kallaway, S. Yahya, B. Weber, A. Templeton, M. Trotter, C. Lo, A. Dhillon, N. Heywood, Y. Aawsaj, A. Hamdan, O. Reece-Bolton, A. McGuigan, Y. Shahin, A. Ali, A. Luther, J. A. Nicholson, I. Rajendran, M. Boal.

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**Supporting information**

Additional supporting information may be found in the online version of this article:

**Appendix S1** Definitions of complications (Word document)

**Typesetter: please refer to marked-up figures**

**Fig. 1** Thirty-day **a** readmission rate and **b** complication rate according to number of emergency admissions before cholecystectomy

**Fig. 2** Plot examining hospital effects (residuals) and corresponding 95 per cent confidence intervals for 30-day readmissions

**Fig. 3** Plot examining hospital effects (residuals) and corresponding 95 per cent confidence intervals for 30-day complications

**Table 1** Patient and disease characteristics, and radiological investigations according to performance of emergency, delayed or elective cholecystectomy

	All patients ( <i>n</i> = 8909)	Emergency cholecystectomy ( <i>n</i> = 1451)	Delayed cholecystectomy ( <i>n</i> = 3293)	Elective cholecystectomy ( <i>n</i> = 4165)	<i>P</i> *
Age (years)					< 0.001
< 40	2534 (28.4)	441 (30.4)	859 (26.1)	1234 (29.6)	
40–60	3531 (39.6)	510 (35.1)	1161 (35.3)	1860 (44.7)	
61–80	2602 (29.2)	435 (30.0)	1108 (33.6)	1059 (25.4)	
> 80	242 (2.7)	65 (4.5)	165 (5.0)	12 (0.3)	
Sex					< 0.001
F	6565 (73.7)	1000 (68.9)	2189 (66.5)	3376 (81.1)	
M	2344 (26.3)	451 (31.1)	1104 (33.5)	789 (18.9)	
BMI (kg/m <sup>2</sup> )					0.033
< 17.9	41 (0.5)	6 (0.4)	17 (0.5)	18 (0.5)	
18.0–24.9	1749 (20.6)	262 (19.4)	667 (21.1)	820 (20.5)	
25.0–29.9	3101 (35.7)	494 (36.6)	1108 (35.1)	1429 (35.8)	
30.0–34.9	2078 (24.4)	337 (25.0)	805 (25.5)	936 (23.4)	
≥ 35.0	1600 (18.8)	250 (18.5)	560 (17.7)	790 (19.8)	
Unknown	410	102	136	172	
ASA fitness grade					< 0.001
I–II	7897 (89.4)	1220 (85.0)	2873 (87.9)	3804 (92.1)	
≥ III	937 (10.6)	216 (15.0)	396 (12.1)	325 (7.9)	
Unknown	75	15	24	36	
Indication					< 0.001
Colic	4683 (52.6)	295 (20.3)	955 (29.0)	3433 (82.5)	
Cholecystitis	2581 (29.0)	795 (54.8)	1369 (41.6)	417 (10.0)	
Pancreatitis	851 (9.6)	268 (18.5)	545 (16.6)	38 (0.9)	
CBD stone	589 (6.6)	83 (5.7)	386 (11.7)	120 (2.9)	
Polyp	135 (1.5)	2 (0.1)	16 (0.5)	117 (2.8)	
Dyskinesia	31 (0.3)	1 (0.1)	9 (0.3)	21 (0.5)	
Acalculous	28 (0.3)	6 (0.4)	11 (0.3)	11 (0.3)	
Other	2 (0.0)	0 (0)	0 (0)	2 (0.0)	
Missing	9	1	2	6	
No. of admissions before surgery					< 0.001
0	5196 (58.3)	913 (62.9)	118 (3.6)	4165 (100.0)	
1	2859 (32.1)	361 (24.9)	2498 (75.9)	0 (0.0)	
2	623 (7.0)	121 (8.3)	502 (15.2)	0 (0.0)	
≥ 3	231 (2.6)	56 (3.9)	175 (5.3)	0 (0.0)	
Ultrasonography performed	8539 (96.0)*				
Ultrasound findings					
Thick-walled	2855 (32.8)†	729 (51.9)	1412 (43.6)	714 (17.6)	< 0.001
CBD dilated	1398 (16.0)‡	289 (20.6)	793 (24.5)	316 (7.8)	< 0.001
CT performed	1307 (14.8)§	290 (20.1)	680 (20.8)	337 (8.2)	< 0.001
MRCP performed	2301 (26.1)¶	417 (28.9)	1319 (40.2)	565 (13.8)	< 0.001

ERCP performed	960 (10.9)#	139 (9.6)	670 (20.5)	151 (3.7)	< 0.001
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Values in parentheses are percentages. CBD common bile duct; MRCP magnetic retrograde cholangiopancreatography; ERCP endoscopic retrograde cholangiopancreatography. Data missing for \*12, †219, ‡212, §101, ¶93 and #105 patients. \*\* $\chi^2$  test or Fisher's exact test, as appropriate.

**Table 2** Surgical and hospital-related factors in 8909 cholecystectomies according to performance of emergency, delayed or elective cholecystectomy

	All patients (n = 8909)	Emergency cholecystectomy (n = 1451)	Delayed cholecystectomy (n = 3293)	Elective cholecystectomy (n = 4165)	<i>P</i> ‡
Grade of operating surgeon					
Junior trainee	403 (4.5)	91 (6.3)	114 (3.5)	198 (4.8)	< 0.001
Senior trainee	1488 (16.7)	278 (19.2)	503 (15.3)	707 (17.0)	< 0.001
Consultant	7007 (78.7)	1080 (74.5)	2675 (81.3)	3252 (78.2)	< 0.001
Missing	11	2	1	8	
Consultant present	7755 (89.5)*	1178 (85.9)	2940 (90.9)	3637 (89.6)	< 0.001
Consultant specialty					< 0.001
Oesophagogastric	3416 (38.5)	560 (38.8)	1220 (37.1)	1636 (39.5)	
HPB	1918 (21.6)	314 (21.8)	634 (19.3)	970 (23.4)	
Colorectal	1958 (22.1)	352 (24.4)	831 (25.3)	775 (18.7)	
Breast	348 (3.9)	42 (2.9)	137 (4.1)	169 (4.1)	
Vascular	373 (4.2)	57 (3.9)	141 (4.3)	175 (4.2)	
Other	863 (9.7)	117 (8.1)	327 (9.9)	419 (10.1)	
Missing	33	9	3	21	
Operative method					< 0.001
Laparoscopic	8523 (95.7)	1343 (92.6)	3105 (94.3)	4075 (97.8)	
Converted	297 (3.3)	77 (5.3)	152 (4.6)	68 (1.6)	
Open	89 (1.0)	31 (2.1)	36 (1.1)	22 (0.5)	
Hospital type					< 0.001
Non-university	4843 (54.4)	687 (47.3)	1915 (58.2)	2241 (53.8)	
University	4066 (45.6)	764 (52.7)	1378 (41.8)	1924 (46.2)	
Tertiary HPB centre					< 0.001
No	6602 (74.1)	941 (64.9)	2588 (78.6)	3073 (73.8)	
Yes	2307 (25.9)	510 (35.1)	705 (21.4)	1092 (26.2)	
Acute hospital					< 0.001
No	561 (6.3)	13 (0.9)	222 (6.7)	326 (7.8)	
Yes	8348 (93.7)	1438 (99.1)	3071 (93.3)	3839 (92.2)	
Day case					< 0.001
No	1297 (97.1)	1297 (97.1)	958 (29.2)	822 (19.8)	

Yes	39 (2.9)	39 (2.9)	2324 (70.8)	3337 (80.2)	
Missing	132	115	11	6	
Emergency cholecystectomy list					< 0.001
No	6036 (67.8)	993 (68.4)	2225 (67.6)	2818 (67.7)	
<i>Ad hoc</i>	984 (11.0)	152 (10.5)	403 (12.2)	429 (10.3)	
Once per week	646 (7.3)	162 (11.2)	212 (6.4)	272 (6.5)	
More than once per week	788 (8.8)	138 (9.5)	292 (8.9)	358 (8.6)	
Elective surgery only at hospital	455 (5.1)	6 (0.4)	161 (4.9)	288 (6.9)	
Nassar operative difficulty score					< 0.001
1	3554 (40.2)	360 (25.1)	1146 (35.1)	2048 (49.5)	
2	2644 (29.9)	374 (26.1)	942 (28.9)	1328 (32.1)	
3	1814 (20.5)	423 (29.5)	775 (23.8)	616 (14.9)	
4	821 (9.3)	278 (19.4)	397 (12.2)	146 (3.5)	
Missing	76	16	33	27	
IOC					< 0.001
Not performed	7770 (87.9)	1121 (78.0)	2869 (87.6)	3780 (91.6)	
Planned	965 (10.9)	295 (20.5)	354 (10.8)	316 (7.7)	
Unplanned	105 (1.2)	22 (1.5)	54 (1.6)	29 (0.7)	
Missing	69	13	16	40	
CBD exploration	282†	99 (6.9)	113 (3.4)	70 (1.7)	< 0.001

Values in parentheses are percentages. HPB hepatobiliary; IOC, intraoperative

cholangiography; CBD, common bile duct. Data missing for \*244 and †75 patients. ‡ $\chi^2$  test.

**Table 3** Thirty-day readmission and complications according to performance of emergency delayed or elective cholecystectomy.

	Emergency cholecystectomy ( <i>n</i> = 1451)	Delayed cholecystectomy ( <i>n</i> = 3293)	Elective cholecystectomy ( <i>n</i> = 4165)	<i>P</i> ‡
Readmissions	138 (9.5)	270 (8.2)	225 (5.4)	< 0.001
All complications	223 (15.4)	420 (12.8)	319 (7.7)	< 0.001
Intraoperative complications				
Stones spilt	222 (15.3)	400 (12.1)	224 (5.4)	< 0.001
Bleeding	195 (13.4)	302 (9.2)	257 (6.2)	< 0.001
Bowel injury	8 (0.6)	22 (0.7)	19 (0.5)	0.479
CBD injury	5 (0.3)	12 (0.4)	7 (0.2)	0.232
Postoperative complications				
Collections	44 (3.0)	88 (2.7)	57 (1.4)	< 0.001
Surgical-site infection	38 (2.6)	78 (2.4)	76 (1.8)	0.114
Pancreatitis	13 (0.9)	19 (0.6)	4 (0.1)	< 0.001
CBD stone	30 (2.1)	28 (0.9)	25 (0.6)	< 0.001
Bile leak	32 (2.2)	52 (1.6)	37 (0.9)	< 0.001
Respiratory	29 (2.0)	69 (2.1)	29 (0.7)	< 0.001
Reimaging	172 (11.8)	300 (9.1)	206 (4.9)	< 0.001
Radiological drain	16 (1.1)	37 (1.1)	16 (0.4)	< 0.001
Relaparoscopy or laparotomy	7 (0.5)	27 (0.8)	27 (0.6)	0.399
30-day mortality	3 (0.2)	6 (0.2)	2 (0.05)	0.160

Values in parentheses are percentages. CBD, common bile duct. \* $\chi^2$  test.

**Table 4** Multilevel random intercept logistic regression analysis of association between patient and surgery characteristics with all-cause 30-day readmission following cholecystectomy

	Odds ratio	<i>P</i>
Random intercept	0.07 (0.06, 0.09)	< 0.001
Patient factors		
Age (years)		
< 40	1.00 (reference)	
40–60	0.78 (0.63, 0.95)	0.014
61–80	0.72 (0.58, 0.91)	0.005
> 80	0.56 (0.33, 0.94)	0.028
ASA fitness grade		
I–II	1.00 (reference)	
II–IV	1.47 (1.14, 1.90)	0.003
Surgical factors		
Surgery admission type		
Delayed	1.00 (reference)	
Emergency	0.91 (0.68, 1.21)	0.519
Elective	0.48 (0.35, 0.66)	< 0.001
No. of surgical admissions		
0	1.00 (reference)	
1	1.50 (1.11, 2.03)	0.009
≥ 2	1.59 (1.18, 2.14)	0.002
Duration of surgery (per min)	1.01 (1.00, 1.01)	0.043

Values in parentheses are 95 per cent confidence intervals.



**Table 5** Multilevel random intercept logistic regression of the association between patient and surgery characteristics with all-cause 30-day complications following cholecystectomy

	Odds ratio	<i>P</i>
Random intercept	0.06 (0.04, 0.07)	< 0.001
Patient factors		
Age (years)		
< 40	1.00 (reference)	
40–60	0.78 (0.65, 0.95)	0.012
60–80	0.98 (0.80, 1.19)	0.838
> 80	1.52 (1.08, 2.16)	0.018
ASA fitness grade		
I–II	1.00 (reference)	
II–IV	1.44 (1.17, 1.78)	0.001
Surgical factors		
Surgical admission type		
Delayed	1.00 (reference)	
Emergency	1.01 (0.80, 1.29)	0.902
Elective	0.61 (0.46, 0.81)	< 0.001
No. of surgical admissions		
0	1.00 (reference)	
1	1.36 (1.05, 1.75)	0.021
≥ 2	1.28 (0.99, 1.65)	0.055
Method of operation		
Laparoscopic	1.00 (reference)	
Open	1.77 (1.04, 3.01)	0.034
Laparoscopic converted to open	2.59 (1.91, 3.52)	< 0.001
Duration of surgery (per min)	1.01 (1.01, 1.01)	0.003
Level of operation difficulty (per Nassar level)	1.22 (1.12, 1.33)	< 0.001

Values in parentheses are 95 per cent confidence intervals.

