Characteristics and outcome of elderly patients admitted for acute Cholecystitis to medical or surgical wards

Itamar Feldman¹, Lena Feldman¹, Dvorah S. Shapiro¹, Gabriel Munter², Amos M. Yinnon²* and Reuven Friedman¹

Abstract

Background: Elderly patients admitted because of acute cholecystitis are usually not operated during their initial admission and receive conservative treatment. To help formulate a new admission policy regarding elderly patients with acute cholecystitis we compared the demographic and clinical characteristics and outcome of patients > 65 with acute cholecystitis admitted to medical or surgical wards.

Methods: This retrospective study included all patients > 65 years admitted for acute cholecystitis between January, 2009 and September, 2016. Data were retrieved from the electronic health records.

Results: A total of 187 patients were detected, 54 (29%) in medical departments and 133 (71%) in surgical wards. The mean age (±SD) was 80 ± 7.5 and was higher among those in medical than surgical wards (84 ± 7 versus 79 ± 7, p < 0.05). Patients hospitalized in medical departments had more comorbidity, disability and mental impairment. However, there was no difference in mortality between the two groups, 1 (2%) and 6 (4%) respectively. Independent predictors for hospitalization in medical departments were chronic obstructive pulmonary disease (OR = 9.8, 95% C.I 1.6–59) and the Norton Scale score (NSS)(OR = 0.7, 95% C.I 0.7–0.8). Impaired mental condition was the only predictor for hospitalization > 1 week. The strongest predictor for having cholecystostomy was admission to the surgical department (OR = 14.7, 95% C.I 3.9–56.7). Linear regression showed a negative correlation between NSS and length of hospitalization (LOH; Beta = −0.5).

Conclusion: Elderly patients with acute cholecystitis who require conservative management, especially those with severe functional and mental impairment can be safely hospitalized in medical departments. Outcome was not inferior in terms of mortality and LOH. These results have practical policy implications for the placement of elderly patients with acute cholecystitis in medical rather than surgical departments.

Keywords: Cholecystitis, Medical department, Geriatric department, Outcome, Health policy
Introduction
Acute cholecystitis is a common clinical condition usually precipitated by cystic duct obstruction by a stone [1]. Advanced age is a risk factor for acute cholecystitis and most cases occur in older adults (50–70%) [2, 3]. These patients are usually admitted to a general surgical ward for medical treatment which includes intravenous fluids, antibiotics, restriction of oral intake and analgesics. The majority of patients treated medically experience remission of their symptoms within 2–7 days following hospitalization, although seriously ill or debilitated patients may be managed with cholecystostomy and tube drainage of the gallbladder. Cholecystectomy is the definitive treatment for cholecystitis and early cholecystectomy (i.e. operation within 72 h after 7 days of symptoms) is the preferred treatment in young patients and even in carefully selected elderly patients [4–7]. Nevertheless, many medical centers choose to treat elderly patients conservatively and delay surgery for an elective procedure. In Israel, due to logistical limitations, delayed surgery is the rule rather than the exception in many medical centers (even for young patients) [8–10].

There are several studies suggesting that conservative treatment in elderly patients is feasible and safe and is not associated with a poorer clinical outcome [9, 11, 12]. In our institution, the common practice is to discharge patients for subsequent elective laparoscopic cholecystectomy; operation during the initial admission is exceptional. Nevertheless, in other hospitals urgent or early laparoscopic cholecystectomy is the common practice [13].

Because older patients (> 65 years) often suffer from multiple comorbidities [14], it is not uncommon in our hospital to encounter older patients with acute cholecystitis in the medical or acute geriatric wards, although a prior these patients are supposed to be cared for in surgical wards.

We conducted the present study in order to formulate the most appropriate admission policy for elderly patients admitted for acute cholecystitis. Although there are data on the necessity of holistic treatment in elderly patients with surgical disease, this is the first study that evaluates management of such patients in medical wards.

We retrospectively reviewed the course and outcome of elderly patients admitted for acute cholecystitis over a 7.5 year period (2009–2016). The baseline expectation was that older patients admitted to medical departments would harbor more co-morbidities and functional disabilities [13], and therefore, would benefit from care in medical rather than in surgical departments.

Methods
Study design and participant selection
This study was conducted in Shaare Zedek Medical Center, a 1000-bed university affiliated, general hospital. The hospital provides all services except organ transplantations. The medical division consists of 4 medical departments and one acute geriatric department for a total of ±200 beds. The department of general surgery has ±60 beds.

We conducted a retrospective cohort review of all patients > 65 years of age with an admitting diagnosis of acute cholecystitis during a 7.5 year period (January 2009–September 2016). Patients were divided according to the department to which they were admitted after acute cholecystitis was diagnosed in the emergency department. Relevant demographic, clinical, laboratory and outcome data were compared between the two groups.

We used the Norton scale score (NSS) to evaluate the general condition of the patients. This scale, ranging from a maximal of 20 points to a minimal of 4, facilitates assessment of physical and mental conditions, activity, mobility and incontinence [15]. The NSS also enables prediction of the risk for pressure sores. The major endpoints selected for this study were (i) length of hospitalization in days (LOH), (ii) mortality, and (iii) percentage of patients treated with a cholecystostomy.

Statistical analysis
Demographic, clinical and laboratory data were compared according to department of admission (medicine versus surgery). Categorical variables were compared using the χ2 test, and continuous variable means were compared using the Student t-test. The cut-off level for statistical analysis for difference between the groups was \( p < 0.05 \). Univariate linear regression was performed to explore the correlation between two continuous variables. For finding the predicting factors for outcome and admission to medical departments, a multivariate logistic regression was performed. All statistical data were calculated using the SPSS Statistic program, version 17.0 (SPSS Inc).

Results
The study included all 187 conservatively managed patients older than 65, diagnosed with acute cholecystitis in the emergency department and admitted to either medicine (\( N = 54, 29\% \)) or surgery (\( N = 133, 71\% \)). The mean age of the patients was 80 ± 7. Of the total cohort, 52 (29%) were older than 85 years, 22 (13%) lived in nursing homes and the mean Norton Scale Score (NSS) was 16 ± 4. Patients hospitalized in medical departments were more likely to be older, nursing home residents, have lower NSS and more comorbidity (Table 1). The predictors for admission to medical departments were chronic obstructive pulmonary disease (COPD) (OR = 9.8, 95%CI 1.6–59, \( p < 0.05 \)) and NSS (OR = 0.7, 95%CI 0.7–0.8, \( p < 0.05 \)) (Table 2). Respiratory distress was more common among those hospitalized in medical
The mean LOH was 8.9 ± 6 days and was not significantly different between the two groups (Table 4). Impaired mental condition was found to be a predictor for hospitalization longer than 1 week (OR = 3.7) (Table 2). NSS was found to have an indirect and inverse correlation with LOH (β = -0.5).

The percentage of patients who underwent cholecystostomy was higher among patients hospitalized in surgery (45, 35%) than in medical departments (8, 14%) (p < 0.05). Hospitalization in surgery, nursing home residency, hyponatremia, leukocytosis and having COPD or chronic renal failure (CRF) were all found to be predictors for patients who went on to have a cholecystostomy (Table 2).

**Discussion**

Traditional admission practices are changing in the current era, often driven by scarcity of beds in certain disciplines and differential reimbursement policies for various specialties. In addition, there is emerging data on geriatric co-management of elderly patients with surgical problems, such management has the advantages of shorter LOH, less mortality and lower readmission rate [16]. A combination of these and other factors has lead to a gradual increase in admission to medical departments of patients with diagnoses that previously were considered as requiring care in surgical departments (Table 5). These patients are usually elderly, with considerable functional and cognitive impairment. On the one hand, these patients are considered poor surgical candidates, while on the other hand, their multiple comorbidities can be expected to be superiorly managed in surgical departments (24% versus 5%, p < 0.05) (Table 3). Laboratory values were also significantly different between patients hospitalized in medical as compared to those in surgical departments (Table 3).

Mortality was not significantly different between the two groups (Table 4). The cause of death in all cases was sepsis. Of the cohort initially admitted to the surgical departments, six died: one died in the surgical ward and five in the medical wards (N = 5) or ICU (N = 1), where they were transferred after their initial admission to surgery. The only parameter that showed a trend toward predicting mortality was the NSS; a decrease of one point in the NSS doubled the risk of death (\( P = 0.07 \); Table 2). Compared to patients who survived, the NSS was lower among patients who died (16.2 vs. 9.5, \( P < 0.05 \)). In addition, among the patients who died, two patients underwent cholecystostomy and the youngest patient was 75 years old (mean 84 ± 8 years).

**Table 1** Demographic, comorbidity and daily living characteristics of older adult patients admitted for acute cholecystitis

| Characteristics   | Total n = 187 (%) | Surgery n = 133 (%) | Medicine n = 54 (%) | P   |
|-------------------|-------------------|---------------------|--------------------|-----|
| Gender - N (%)    |                   |                     |                    | 0.29|
| Male              | 89 (48)           | 60 (45)             | 29 (54)            |     |
| Female            | 98 (52)           | 73 (55)             | 25 (46)            |     |
| Age - Mean ± SD   | 80 ± 7.5          | 79 ± 7              | 84 ± 7             | < 0.05|
| Age Group - N (%) |                   |                     |                    |     |
| 65–74             | 38 (22)           | 33 (26)             | 5 (10)             | < 0.05|
| 75–85             | 87 (49)           | 67 (52)             | 20 (42)            |     |
| > 85              | 52 (29)           | 29 (22)             | 23 (48)            |     |
| Nursing home      | 22 (13)           | 9 (7)               | 13 (25)            | < 0.05|
| Morbidity – Disease N (%) | | | | |
| CRF               | 20 (11)           | 12 (9)              | 8 (15)             | 0.2 |
| CHF               | 27 (14)           | 13 (10)             | 14 (26)            | < 0.05|
| DM                | 51 (27)           | 31 (23)             | 20 (37)            | 0.06|
| HTN               | 72 (38)           | 82 (62)             | 33 (61)            | 0.7 |
| CVA               | 30 (16)           | 11 (8)              | 19 (35)            | < 0.05|
| IHD               | 62 (33)           | 38 (29)             | 24 (44)            | < 0.05|
| COPD              | 11 (6)            | 3 (2)               | 8 (15)             | < 0.05|
| Norton Scale Score (NSS) | 16 (4) | 17 (3) | 13 (4) | < 0.05|
| NSS Criterion*    |                   |                     |                    |     |
| Mentally Alert    | 97 (66)           | 78 (82)             | 19 (37)            | < 0.05|
| Full Mobility     | 54 (38)           | 47 (51)             | 7 (14)             | < 0.05|
| Continent         | 87 (61)           | 71 (78)             | 16 (31)            | < 0.05|
| Good Daily Activity | 56 (42) | 49 (60) | 7 (14) | < 0.05|

**Table 2** Multivariate factors associated with hospitalization in internal medicine/geriatric departments and clinical outcome

| Risk factors                  | Odds ratio (OR) (95% CI) | P   |
|-------------------------------|--------------------------|-----|
| Risk factor for admission to medical wards |                         |     |
| Chronic obstructive pulmonary disease | 9.8 (1.6–59)             | < 0.05|
| Norton scale score            | 0.7 (0.7–0.8)            | < 0.05|
| Risk factors for mortality    |                          |     |
| Norton scale score            | 0.5 (0.3–1)              | 0.07 |
| Hospitalization duration > 1 week |                      |     |
| Impaired mental condition     | 3.7 (1.7–7.9)            | < 0.05|
| Cholecystostomy               |                          |     |
| Hospitalization in surgical ward | 14.7 (3.9–56.7)         | < 0.05|
| Chronic renal failure         | 3.9 (1.23–13.5)          | < 0.05|
| Chronic obstructive pulmonary disease | 16.5 (2.4–116)        | < 0.05|
| Institutional residency       | 6.3 (1.6–24.9)           | < 0.05|
| Serum sodium                  | 0.9 (0.8–1)              | < 0.05|
| White blood count             | 1.1 (1.0–1.2)            | < 0.05|

CRF chronic renal failure, CHF congestive heart failure, DM diabetes mellitus, HTN hypertension, CVA cerebrovascular accident, IHD ischemic heart disease, COPD chronic obstructive pulmonary disease

*Due to missing data, \( n = 146 \)
medical departments [15, 17]. Nonetheless, both internists and surgeons questioned the clinical validity of this ensuing reality and the current study was a direct result of our quest for data. Although the current study focused on acute cholecystitis, we believe it serves as a template for further studies to assess elderly patients with other, traditionally surgical diagnoses admitted to medical versus surgical departments.

Even without an official policy, we found that 29% of elderly patients admitted for acute cholecystitis in this study were managed in the medical department, whereas only 26% were managed in the surgical department. This finding suggests that, despite the higher mortality rate in the medical department, elderly patients with acute cholecystitis may benefit from surgical intervention. Table 3 shows the vital signs and laboratory values of the study population, and Table 4 provides the clinical outcome of patients > 65 with acute cholecystitis.

### Table 3 Vital signs and laboratory values of the study population

| Vital signs/Laboratory values | Reference Interval | All Surgery n = 133 | Medicine n = 54 | p |
|-----------------------------|-------------------|----------------------|----------------|---|
| **Vital Signs ±SD**         |                   |                      |                |
| Systolic BP                 | 143 ± 32          | 144 ± 32             | 140 ± 332      | 0.6 |
| Diastolic BP                | 74 ± 15           | 74 ± 14              | 74 ± 17        | 0.9 |
| Pulse                       | 84 ± 18           | 82 ± 17              | 90 ± 20        | < 0.05 |
| Respiratory Distress, N (%) | 13 (10)           | 4 (5)                | 9 (24)         | < 0.05 |
| **Laboratory Values ±SD**   |                   |                      |                |
| ALT (IU/L)                  | 0–55              | 126 ± 330            | 99 ± 211       | 0.5 |
| AST (IU/L)                  | 5–34              | 166 ± 550            | 112 ± 134      | 0.4 |
| GGT (IU/L)                  | 12–43             | 150 ± 217            | 195 ± 251      | 0.1 |
| ALP (IU/L)                  | 38–150            | 147 ± 116            | 192 ± 161      | 0.05 |
| Total Bilirubin (mg/dL)     | 0.2–1.3           | 1.5 ± 1.3            | 1.7 ± 1.5      | 0.4 |
| LDH (IU/L)                  | 125–220           | 917 ± 2214           | 730 ± 427      | 0.3 |
| Creatinine (mg/dL)          | 0.52–104          | 1.4 ± 3.1            | 1.3 ± 1.2      | 0.9 |
| BUN (mg/dL)                 | 9–120             | 24 ± 16              | 32 ± 23        | < 0.05 |
| BUN/CRT Ratio               | 22 ± 8            | 20 ± 7               | 25 ± 10        | < 0.05 |
| Sodium, mEq/L               | 135–145           | 136 ± 4              | 136 ± 6        | 0.9 |
| Potassium, mEq/L            | 3.6–5             | 4 ± 0.5              | 4.3 ± 0.8      | < 0.05 |
| WBC 10³/uL                  | 3.6–10            | 14 ± 6               | 14.0 ± 7       | 0.7 |
| Neutrophils (%)             | 50–75             | 80 ± 12              | 81 ± 12        | 0.7 |
| Platelets 10³/uL            | 150–450           | 229 ± 79             | 236 ± 92       | 0.7 |
| Hemoglobin (g/dL)           | 12–16             | 13 ± 2               | 12 ± 2         | < 0.05 |
| Amylase (IU/L)              | 30–125            | 208 ± 625            | 201 ± 713      | 0.9 |

BP: blood pressure, BUN: blood urea nitrogen, CRT: creatinine, WBC: white blood cells

### Table 4 Clinical outcome of patients > 65 with acute cholecystitis

| Clinical outcome of all patients > 65 | Total n = 186 (%) | Surgery n = 133 (%) | Medicine n = 54 (%) | p |
|--------------------------------------|-------------------|---------------------|---------------------|---|
| Days of Hospitalization (±SD)        | 8.9 ± 6           | 8.4 ± 6             | 10.2 ± 7            | 0.09 |
| Cholecystostomy                      | 53 (29)           | 45 (35)             | 8 (15)              | < 0.05 |
| Bacteremia                           | 13 (7)            | 10 (7)              | 3 (6)               | 0.9 |
| Death                                | 7 (4)             | 6 (4)               | 1 (2)               | 0.4 |

Clinical outcome of the subset of patients > 75 years

| Outcome marker | Total n = 137 (%) | Surgery n = 91 (%) | Medicine n = 46 (%) | p |
|---------------|-------------------|-------------------|---------------------|---|
| Days of Hospitalization (±SD)        | 9.2 ± 7           | 8.8 ± 7           | 10.0 ± 6            | 0.3 |
| Cholecystostomy                      | 41 (28)           | 34 (45)           | 7 (15)              | < 0.05 |
| Bacteremia                            | 11 (8)            | 9 (9)             | 2 (4)               | 0.6 |
| Death                                  | 7 (5)             | 6 (6)             | 1 (2)               | 0.7 |
Table 5 Several classic surgical diagnoses which could be treated in medical departments, especially in elderly patients with co-morbidities, functional and cognitive impairments who are poor surgical candidates

| No. | Diagnoses                                      |
|-----|-----------------------------------------------|
| 1   | Acute cholecystitis                           |
| 2   | Ascending cholangitis                         |
| 3   | Pancreatitis, not gallstone-related           |
| 4   | Gastro-intestinal bleeding, not life-threatening |
| 5   | Liver abscess, including gallstone-related    |
| 6   | Diverticulitis                                |

7.5 year retrospective study were admitted to medical departments. As expected, they had higher rates of co-morbidities, disabilities as well as mental impairment. The percent of patients living in a nursing home in our study (12%) was much higher than that for adults over 65 years of age in Israel’s general population, which was 2% in 2015 [14]. Rates were much higher for those admitted to medicine compared to surgery (25% vs 7%, \( p < 0.05 \); Table 1). These as well as the higher rates of cardiovascular, cerebrovascular and chronic respiratory disease and lower NSS indicate that the medical group had significantly more comorbidity, disability and immobility.

Patients hospitalized in medical departments presented with higher pulse rates. This could not be explained by the higher age of the population in medicine per se [18]. Possible explanations for this relative tachycardia were atrial fibrillation, the incidence of which increases with age [19], or more severe inflammatory responses among patients in the medical group. In younger patients with acute cholecystitis the mean value of liver enzymes are usually within normal range, unless the diagnosis is both acute cholecystitis and choledocholithiasis [20, 21]. In our study the mean value of all liver enzymes was above normal. This could have been due to a more serious presentation, with fatty liver disease [22] or concomitant choledocholithiasis [20], both of which are more common in the elderly [3]. Hemoglobin values, which continuously decrease in older patients [23], were found to be lower in the medical patients, who were on average older. Indeed, in our study, for the portion of the population above 75 years old, there was a negative correlation between age and hemoglobin level, as demonstrated by linear regression (beta = -0.053, \( P = 0.06 \)). Mean blood urea nitrogen, which also tends to increase with age [24], was found to be higher in patients admitted to medical wards. In addition, the blood urea nitrogen/creatinine ratio was higher among those in the medical group (25 vs 20), the difference implying that dehydration and hypo-perfusion were more common in this group [25].

The mortality rate of the total study population (4%) was higher than the mortality rate associated with conservative treatment of acute cholecystitis (0.8%) [11]. However, among high risk patients there are reports of rates of mortality as high as 17.5% [26]. The only parameter that showed a trend in predicting mortality was NSS (Table 2). The mean NSS among patients who died was a low 9.2, whereas the mean NSS of the entire cohort was 16. This is consistent with previous data that demonstrated the association between low NSS and mortality during hospitalization [27, 28]. In addition to a very low NSS, patients who died were very old, and five of the seven were transferred to the intensive care unit or medical department after being initially hospitalized in surgery.

The mean LOH of 8.9 ± 6 days was comparable to previous reports of elderly patients with acute cholecystitis who were managed conservatively [29]; however, it was longer than that for the general patient population (3–6 days) [30–33]. There was no statistically significant difference in LOH between the two groups. Notably, despite the absence of statistical significance, COPD and an impaired mental condition, the two independent predictors for long hospitalization (Table 2), were more common in the medical group. There was a clear negative correlation between NSS and LOH for both groups of patients, though the correlation was weaker for patients hospitalized in medical wards.

Although the definitive treatment for acute cholecystitis is cholecystectomy, such intervention when done early has a higher rate of mortality in the elderly and other high risk patients [34, 35]. The alternative bridging treatment for high risk patients is percutaneous cholecystostomy. In our study, the cholecystostomy rate was 29%, similar to figures reported in other studies; this suggests an increasing trend of using this modality in the treatment of elderly patients diagnosed with acute cholecystitis (24–54%) [35–38].

The strongest predictor of receiving a cholecystostomy was being hospitalized in surgery (OR 15, 95% C. I 4–57), a finding which may be attributable to an increased availability of invasive procedures and a more invasive approach among surgeons. This finding suggests that physicians in medical wards should be more aware of the cholecystostomy procedure as adjunct treatment for patients with acute cholecystitis. However, not being treated with cholecystostomy was not associated with increased mortality or LOH (Table 2).

Limitations

Our study was retrospective and this is probably its major drawback. As management of elderly patients with acute cholecystitis in medical departments has surreptitiously become an established practice in our hospital,
we elected to retrospectively investigate this practice. If care in medical departments was found associated with worse outcome, this would lead to immediate reversal of the norm – and these patients would all be directed to the surgical department. However, as expected, outcome of these elderly patients in medical departments turned out to be not inferior to that in surgical departments, and was possibly associated with shorter LOH for patients with low NSS. We believe these findings facilitate embracing a new policy of admission of elderly patients with acute cholecystitis to medical departments, especially for those who are poor surgical candidates on account of multiple co-morbidities. However, in order that these data and insights lead to a generally accepted change of policy there is need for a prospective, long-term study to evaluate readmissions, morbidity and mortality and surgical interventions that may occur after the initial hospitalization.

Conclusion
This is the first study to compare the outcomes of elderly patients with a surgical diagnosis admitted to medical and surgical wards. We found that conservative management of elderly patients with acute cholecystitis in medical departments is not inferior to treatment in surgery. Internists and geriatricians should be aware of the availability and efficacy of the percutaneous cholecystostomy for high-risk patients in medical departments. The data in this study indicate that elderly patients with acute cholecystitis, especially those with comorbidity and, functional and cognitive impairments, can and probably should be admitted to medical wards, both for the benefit of the patients themselves and because the surgical departments are not proficient in the management of these patients. However, a major concern is the already limited availability of medical beds in many hospitals, leading to frequent hallway admissions in medical departments. Our data add to other arguments to increase medical bed availability in general hospitals. Alternatively, additional medical or geriatric physicians should be made available to consult on geriatric patients in the surgical wards.

Abbreviations
NSS: Norton scale score; SD: Standard deviation; COPD: Chronic obstructive pulmonary disease; LOH: Length of hospitalization; CRF: Chronic renal failure; CHF: Congestive heart failure; OR: Odds ration; CI: Confidence interval

Acknowledgments
None to declare.

Authors’ contributions
All authors discussed the policy issues involved, the methodology of the study and results and read and agree with the final manuscript. If retrieved and analyzed the data. RF, LF, DSS and GM made significant contributions on methodology, data analysis and design of tables; AYM oversaw the project and edited the manuscript. The author(s) read and approved the final manuscript.

Funding
The authors declare that they did not receive funding for the preparation of the manuscript.

Availability of data and materials
Excel file is available upon request.

Ethics approval and consent to participate
The study was approved by the hospital’s internal review board (Helsinki committee) and, having a retrospective and non-interventional design, received a waiver for obtaining informed consent.

Consent for publication
All authors gave consent for publication.

Competing interests
The authors declare that they have no competing interests.

Author details
1 Department of Geriatrics, affiliated with the Hebrew University-Hadassah Medical School, Jerusalem, Israel. 2 Division of Internal Medicine, Shaare Zedek Medical Center, affiliated with the Hebrew University-Hadassah Medical School, P.O. Box 3235, 91031 Jerusalem, Israel.

Received: 22 August 2019 Accepted: 27 April 2020
Published online: 03 August 2020

References
1. Telfer S, Fenyo G, Holt PR, de Dombal FT. Acute abdominal pain in patients over 50 years of age. Scand J Gastroenterol Suppl. 1988;144:47–50.
2. Cho JY, Han HS, Yoon YS, Ahn KS. Risk factors for acute cholecystitis and a complicated clinical course in patients with symptomatic cholelithiasis. Arch Surg. 2010;145(4):329–33 discussion 33.
3. Kelly SG, Rice JC, Benson M, Lucey MR. Biliary and pancreatic disease. In: Halter JB, Ouslander JG, Studenski S, High KP, Asthana S, Supiano MA, et al., editors. Hazzard’s geriatric medicine and gerontology, 7e. New York: McGraw-Hill Education; 2017.
4. Loozen CS, van Santvoort HC, van Duijvenbalk P, Beselink MG, Gouma DJ, Nieuwenhuijzen GA, et al. Laparoscopic cholecystectomy versus percutaneous catheter drainage for acute cholecystitis in high risk patients (CHOCOLATE): multicentre randomised clinical trial. BMJ. 2018;363:k3965.
5. Loozen CS, van Ramshorst B, van Santvoort HC, Boerma D. Early cholecystectomy for acute Cholecystitis in the elderly population: a systematic review and meta-analysis. Dig Surg. 2017;34(5):371–9.
6. Halpin V. Acute cholecystitis. BMJ clinical evidence. Clin Evid. 2014;411.
7. Mayumi T, Okamoto K, Takada T, Strasberg SM, Solomkin JS, Schlossberg D, et al. Tokyo guidelines 2018: management bundles for acute cholangitis and cholecystitis. J Hepato-Bil-Pancreat Sci. 2018;25(1):96–100.
8. Rosin D. Outcome of delayed cholecystectomy after percutaneous Cholecystostomy for acute Cholecystitis. IMAJ. 2018;20(10):651.
9. Sakran N, Kopelman D, Dar R, Abaya N, Mokary SE, Handler C, et al. Outcome of delayed cholecystectomy after percutaneous Cholecystostomy for acute Cholecystitis. IMAJ. 2018;20(10):627–31.
10. Yuval JB, Mizrahi I, Mazeh H, Weiss DJ, Almogy G, Bala M, et al. Delayed laparoscopic cholecystectomy for acute Calculous Cholecystitis: is it time for a change? World J Surg. 2017;41(7):1762–8.
11. Loozen CS, Ooi JE, van Ramshorst B, van Santvoort HC, Boerma D. Conservative treatment of acute cholecystitis: a systematic review and pooled analysis. Surg Endos. 2017;31(2):504–15.
12. Janssen ERI, Hendriks T, Natarshvil T, Bremer A.J.A. Retrospective Analysis of Non-Surgical Treatment of Acute Cholecystitis. Surg Infect. 2019. https://doi.org/10.1089/sur.2019.261 [Epub ahead of print].
13. Ferrucci L, Fabbri E, Walston JD. Frailty. In: Halter JB, Ouslander JG, Studenski S, High KP, Asthana S, Supiano MA, et al., editors. Hazzard’s geriatric medicine and gerontology. New York: McGraw-Hill Education; 2017. p. 7e.
14. Institute M-J-B. 65 years old in Israel, annual data, 2015 2015. Available from: http://mashav.jdc.org.il/?CategoryID=233&ArticleID=162.
15. Norton D. Calculating the risk: reflections on the Norton scale. Decubitus. 1989;2(4):10.
16. Van Grootven B, Mendelson DA, Deschodt M. Impact of geriatric co-management programmes on outcomes in older surgical patients: update of recent evidence. Curr Opin Anaesthesiol. 2020;33(1):114–21.

17. Wessoric D, Flanders SA, Hall KE, Blaum CS. Acute Hospital Care. In: Halter JB, Outslender JS, Studencki S, High KP, Asharana S, Supiano MA, et al., editors. Hazzard’s geriatric medicine and gerontology, 7e. New York: McGraw-Hill Education; 2017.

18. Kitzman DW, Upadhye B, Haykowsky M, Taffet GE. Effects of aging on cardiovascular structure and function. In: Halter JB, Outslender JS, Studencki S, High KP, Asharana S, Supiano MA, et al., editors. Hazzard’s geriatric medicine and gerontology, 7e. New York: McGraw-Hill Education; 2017.

19. Go AS, Hylek EM, Phillips KA, Chang Y, Hentault LE, Selby JV, et al. Prevalence of diagnosed atrial fibrillation in adults: national implications for rhythm management and stroke prevention: the Anticoagulation and risk factors in atrial fibrillation (ATRIA) study. JAMA. 2001;285(18):2370–5.

20. Thapa PB, Maharjan DK, Suwal B, Byanjankar B, Singh DR. Serum gamma glutamyl transferase and alkaline phosphatase in acute cholecystitis. J Nepal Health Res Council. 2010;8(2):78–81.

21. Padda MS, Singh S, Tang SJ, Rockley DC. Liver test patterns in patients with acute calculous cholecystitis and/or choledocholithiasis. Alim Pharmacol Ther. 2009;29(9):1011–8.

22. Song SH, Kwon CI, Jin SM, Park HJ, Chung CW, Kwon SW, et al. Clinical characteristics of acute cholecystitis with elevated liver enzymes not associated with choledocholithiasis. Eur J Gastroenterol Hepatol. 2014;26(4):452–7.

23. Yamada M, Wong FL, Suzuki G. Study RESAH. Longitudinal trends of hemoglobin levels in a Japanese population—RERF’s adult health study subjects. Eur J Haematol. 2003;70(3):129–35.

24. Musch W, Verfaille I, Decaux G. Age-related increase in plasma urea level and decrease in fractional urea excretion: clinical application in the syndrome of inappropriate secretion of antidiuretic hormone. Clin J Am Soc Nephrol. 2006;1(5):909–14.

25. Morgan DB, Carver ME, Payne RB. Plasma creatinine and urea: creatinine ratio in patients with raised plasma urea. BMJ. 1977;2(6092):929–32.

26. Hatzidakis AA, Prassopoulos P, Petinarakis I, Sanidas E, Chrysos E, Chalkiadakis G, et al. Acute cholecystitis in high-risk patients: percutaneous cholecystostomy vs conservative treatment. Eur Radiol. 2002;12(7):1778–84.

27. Diez-Manglano J, Aramil-Longares MJ, Al-Cheikh-Felices P, Garces-Horna V, Pueyo-Tejedor P, Martinez-Rodes P, et al. Norton scale score on admission and mortality of patients hospitalised in internal medicine departments. Revista Cin Espanola. 2018;218(4):177–84.

28. Leshem-Rubinow E, Vaknin A, Sherman S, Justo D. Norton scale, cholecystostomy and interval laparoscopic cholecystectomy. Surg Endos. 2018;32(4):1858–66.

29. Yamada M, Wong FL, Suzuki G. Study RESAH. Longitudinal trends of hemoglobin levels in a Japanese population—RERF’s adult health study subjects. Eur J Haematol. 2003;70(3):129–35.

30. Musch W, Verfaille I, Decaux G. Age-related increase in plasma urea level and decrease in fractional urea excretion: clinical application in the syndrome of inappropriate secretion of antidiuretic hormone. Clin J Am Soc Nephrol. 2006;1(5):909–14.

31. Go AS, Hylek EM, Phillips KA, Chang Y, Hentault LE, Selby JV, et al. Prevalence of diagnosed atrial fibrillation in adults: national implications for rhythm management and stroke prevention: the Anticoagulation and risk factors in atrial fibrillation (ATRIA) study. JAMA. 2001;285(18):2370–5.

32. Rajcok M, Bak V, Danilov L, Kukucka M, Schnorrer M. Early versus delayed sphincterotomy in the initial management of acute cholecystitis in elderly hospitalised patients at high surgical risk. Endoscopy. 2006;38(8):773–8.

33. Go AS, Hylek EM, Phillips KA, Chang Y, Hentault LE, Selby JV, et al. Prevalence of diagnosed atrial fibrillation in adults: national implications for rhythm management and stroke prevention: the Anticoagulation and risk factors in atrial fibrillation (ATRIA) study. JAMA. 2001;285(18):2370–5.

34. Patterson EJ, McLoughlin RF, Mathison JR, Cooperberg PL, MacFarlane JK. An alternative approach to acute cholecystitis. Percutaneous cholecystostomy and interval laparoscopic cholecystectomy. Surg Endos. 1996;10(12):1185–8.

35. Ferrarese AG, Solej M, Enrico S, Falcone A, Catalano S, Pozzi G, et al. Elective and emergency laparoscopic cholecystectomy in the elderly: our experience. BMC Surg. 2013;13 Suppl 2:52.

36. Zarour S, Imam A, Kouniavsky G, Lin G, Zbar A, Mavor E. Percutaneous cholecystostomy in the management of high-risk patients presenting with acute cholecystitis: timing and outcome at a single institution. Am J Surg. 2017;214(3):456–61.

37. Cull JD, Velasco JM, Czubak A, Rice D, Brown EC. Management of acute cholecystitis: prevalence of percutaneous cholecystostomy and delayed cholecystectomy in the elderly. J Gastrointest Surg. 2014;18(2):328–33.

38. Barak O, Elazyari R, Appelbaum L, Rivkind A, Almogy G. Conservative treatment for acute cholecystitis: clinical and radiographic predictors of failure. IMAI. 2009;11(12):739–43.

Publisher’s Note
Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.