Original Research Article

A clinical epidemiological study of gallbladder carcinoma: a retrospective study

Shweta Sahai*

Department of Medicine, Gajra Raja Medical College, Gwalior, Madhya Pradesh, India

Received: 04 January 2019
Accepted: 29 January 2019

*Correspondence:
Dr. Shweta Sahai,
E-mail: sahay.shweta2@gmail.com

Copyright: © the author(s), publisher and licensee Medip Academy. This is an open-access article distributed under the terms of the Creative Commons Attribution Non-Commercial License, which permits unrestricted non-commercial use, distribution, and reproduction in any medium, provided the original work is properly cited.

ABSTRACT

Background: Incidence of gallbladder carcinoma (GBC) is high among the north Indian population. It may be due to environmental, dietary and reproductive factors. Evaluation of GBC epidemiology will provide important insights into determining causes and risk factors for gallbladder cancer. The aims and objectives were to study the epidemiology, clinical presentation and etiological factors of GBC.

Methods: Fifty-one cases of GBC were studied retrospectively at Cancer Hospital and Research Institute (CHRI) Gwalior, Madhya Pradesh, India from January to December 1998. All the patients underwent a detailed history taking and physical examination followed by routine laboratory tests and other investigations including ultrasonographic and computed tomography scan of whole abdomen. Histopathological evaluation of gallbladder specimen was also performed after surgical resection of the GB mass.

Results: Incidence of GBC at this centre was 1.49%. GBC was more prevalent among old women (mean age 52.62±12.14 years) belonging to low socio-economic status (SES) (52.94%). Most common presenting symptom of GBC was pain in right hypochondrium (70.4%) followed by anorexia (42.8%), pruritus (35.7%), weight loss (31.2), nausea and vomiting (30.4%). Histopathological examination revealed majority of the case to have adenocarcinoma (84.31%). Gallstones were present in 35.29% patients.

Conclusions: GBC is more prevalent in older women belonging to low SES. Gallstones remain the chief risk factor for this malignancy. Histopathological examination of all cholecystectomy specimens for early pre cursor lesion could serve as an important means of early detection of this carcinoma.

Keywords: Adenocarcinoma, Gallbladder cancer, Gallstones, Malignancy

INTRODUCTION

Although the overall prevalence of gallbladder carcinoma (GBC) is low, it is still the most common cancer of the biliary tree. GBC is a highly malignant tumor with poor prognosis. After colon cancer, GBC is the 5th most common GIT related malignancy.1

GBC is 2-6 times more common in females compared to males. Incidence of GBC increases with increasing age and more than 75% of the patients suffering from GBC are older than 65 years.2 The etiology of GBC is still a source of speculation. Incidence of GBC is high among patients with gallstone disease, especially patients with large and longstanding gallstones are at higher risk.3 Reports have shown an association between a porcelain gallbladder, pancreaticobiliary duct junction and other biliary disorders including choledochal cyst, primary sclerosing cholangitis, Miririzi’s syndrome and gall bladder cancer.4 In present study, author tried to evaluate incidence and clinicopathological aspect of the disease in patients of GBC.
METHODS

An institution based retrospective study was performed on 51 patients having GBC at Cancer Hospital and Research Institute (CHRI) Gwalior, Madhya Pradesh, India from January to December 1998.

Institutional Ethics Committee approval was obtained before starting study.

Inclusion criteria

The records of all patients of either sex of all age groups admitted to (CHRI) Gwalior, Madhya Pradesh, India diagnosed with GBC using cytological examination and by radiological method were included.

Exclusion criteria

Patients with pancreatitis, liver cirrhosis, hepatitis, other co-existing malignancy and patients with common bile duct obstruction due to stone were excluded from the present study.

Methodology

During the study period, all selected records were scanned specially regarding the detailed history and physical examination followed by routine laboratory tests and other investigations including ultrasonographic (US) and computed tomography (CT) scan of whole abdomen.

The histopathological diagnoses for all gall bladder specimens received in its Department of Pathology were noted. The histopathological report with description of gross appearance of the gall bladder together with the Haematoxylin and eosin stained slides were reviewed.

Statistical analysis

All the data were analyzed using IBM SPSS version 20 software. Frequency distribution and cross tabulation was used to prepare the tables. Microsoft office 2010 and PRISM was used to prepare the graphs. Data was expressed as number, percentage and mean±standard deviation.

RESULTS

Out of total 3421 cases of malignancies at the centre, 51 (1.49%) were of GBC. Mean age of study cohort was 52.62±12.14 years. Majority of the subjects belonged to 51-60 years (32 (62.74%)) age group and low socio-economic status (27 (52.94%)) (Table 1). There were only 11 (21.56%) male and 41 female subjects (80.39%).

Table 2 highlights the histopathological findings in 51 surgical specimens where adenocarcinoma was most common. Most common presenting symptom of GBC was pain in right hypochondrium (70.4%) followed by anorexia (42.8%), pruritus (35.7%), weight loss (31.2), nausea and vomiting (30.4%) (Figure 1). Out of 51 GBC patients, 18 (35.29%) patients had gallstones whereas remaining had no gallstone. Secondary metastasis was seen in 56.86%.

Table 1: Distribution of study cohort according to demographic characteristics.

| Characteristics          | No. of subjects | % (n=51) |
|--------------------------|-----------------|----------|
| Age (years)              |                 |          |
| <40                      | 7               | 13.73    |
| 41-50                    | 8               | 15.69    |
| 51-60                    | 32              | 62.75    |
| 61-70                    | 5               | 9.80     |
| SES                      |                 |          |
| Low                      | 27              | 52.94    |
| Medium                   | 16              | 31.37    |
| High                     | 9               | 17.65    |

Table 2: Histopathological findings in 51 surgical specimens.

| Histopathological variants          | No. of patients | %     |
|-------------------------------------|-----------------|-------|
| Adenocarcinoma                      | 42              | 82.35 |
| Papillary adenocarcinoma            | 5               | 9.80  |
| Mucinous adenocarcinoma             | 2               | 3.92  |
| Squamous cell carcinoma             | 1               | 1.96  |
| Adenosquamous carcinoma             | 1               | 1.96  |

Figure 1: Distribution according to presenting symptoms in GBC.

DISCUSSION

GBC is one of the most common culprits of death of biliary related malignancies. Due to rapid progression, GBC is usually detected at an advanced stage. Among the Indian population, GBC has varying geographic distribution. Reports of Tyagi BB et al, has revealed that incidence is higher in north Indians compared to South Indian population. The incidence of GBC varies in different parts of the world. India, Chile and some parts of Japan reported a very high incidence of gall bladder...
carcinoma while other countries reported a lower incidence. Khoo JJ et al, studied 1122 cholecystectomies cases out of that author found 9 (0.80%) cases to have GBC which is lower than the incidence (1.49%) revealed in present study.2 Study from Lucknow reported 358 patients with GBC out of 2600 cholecystectomies performed for a period of 11 years (13.77%) which higher compared to present study findings, while studies from other parts of the world, reported a range of incidence from 0.38% to 1.15%.3-11 This is comparable to the findings of present study. A recent study from China reported that out of 86609 of all biliary tract diseases, GBC accounted for 2.7%.

In the present study, GBC was more prevalent among older women. The higher incidence of GBC in women appears to be due to higher estrogen levels.12 Reports of Pandey M et al, has shown that women with multiple pregnancies are at higher risk of developing gall bladder cancer which may be due to the higher level of sex hormones like progesterone and endogenous estrogen during pregnancy.9 Khoo JJ et al, also reported that there were more females undergoing cholecystectomies than male patients with mean age of 56.7 years which is in agreement to present study finding where mean age of study cohort was 52.6±12.14 years.7 In agreement to present study, Shen HX et al, also reported that GBC was more prone to occur in elderly women.13

In this study, the mean age was also comparable with most of the other reported studies, although in some studies it had been suggested that there was higher incidence of GBC with an older age.14-18 A Chinese survey, has also reported higher mean age (63.7±11.3 years) as compared to present study mean age of 52.6±12.14 years.13

In the present study, most common presenting symptom of GBC was pain in right hypochondrium followed by anorexia, pruritus, weight loss and nausea and vomiting. In agreement to present study, study from Nepal including 47 patients with GBC also reported abdomen pain (93.6%) as the chief presenting symptoms followed by weight loss (51.1%) and jaundice (46.8%).19

In this study, gallstones were present in 35.29% cases which is in agreement with the study done by Khoo JJ et al, which reported 55.6% cases who were having gallstone.7 Shen HX et al, studied 2379 patients with GBC and found that 57.2% had gallstones.13 Gallstones are reported to induce mechanical damage and carcinogens in bile that may trigger epithelial proliferation in the mucous membrane of the gall bladder which can result in malignancy.20 Therefore, it is important to advice for surgery or regular follow-up for patients having long term history of gallbladder stones.

In comparison to Western countries, this correlation of GB stones with GBC is much lower. The correlation of GB stones with GBC is nearly 94-98% in western countries whereas in this study the correlation is much lower. This suggests another etiological factor for GBC.

Gross examination of gallbladder alone is not enough for exclusion of malignancy, histopathological examination is needed for final diagnosis. Moreover, histopathological examination of gallbladder is recommended by several studies to diagnose GBC.21 In present study, histopathological examination revealed majority of the case to have adenocarcinoma (84.31%) which is consistent with other studies.22-25 Khoo JJ et al, also reported that majority of the tumours were adenocarcinoma (77.8%).7

**CONCLUSION**

In the Indian scenario, GBC remains common a clinical entity predominantly affecting females of older age. Therefore, any women especially in fourth and fifth decade of life with symptoms of recent onset constant pain in the right hypochondrium should be thoroughly evaluated for GBC. Present study has also found the association between gallstones and GBC. Despite of advancement in the imaging techniques for gallbladder, prognosis of GBC is still poor. Therefore, this study recommends routine histopathological examination for early diagnosis of gallbladder carcinoma for early intervention.

**Funding:** No funding sources

**Ethical sources:** None declared

**Conflict of interest:** None declared

**Ethical approval:** The study was approved by the Institutional Ethics Committee

**REFERENCES**

1. Pavlidis TE, Pavlidis ET, Symeonidis NG, Psarras K, Sakantamis AK. Current curative surgical management of gallbladder cancer: a brief review. J Current Surg. 2012;2(3):81-3.

2. Mishra R, Goda C, Arora M, Sood M, Siddiqui AA, Husain A, Rashid M, Mishra S. Treatment of gall bladder cancer: a review. Indo Global J Pharm Sci. 2011;2:54-62.

3. Tyagi BB, Manoharan N, Raina V. Risk factors for gallbladder cancer: a population based case-control study in Delhi. Ind J Med Paed Oncol. 2008;29(1):16.

4. Pandey M, Shukla VK. Lifestyle, parity, menstrual and reproductive factors and risk of gallbladder cancer. Euro J Cancer Prev. 2003;12(4):269-72.

5. Khan RA, Wahab S, Khan MA, Siddiqui S, Maheshwari V. Advanced presentation of gallbladder cancer: epidemiologicopathological study to evaluate the risk factors and assess the outcome. JPM. 2010;60(3):217.

6. Le MD, Henson D, Young H, Albores-Saavedra J. Is gallbladder cancer decreasing in view of increasing laparoscopic cholecystectomy?. Ann Hepatol. 2016;10(3):306-14.
7. Khoo JJ, Misron NA. A clinicopathological study of nine cases of gallbladder carcinoma in 1122 cholecystectomies in Johor, Malaysia. Malaysian J Pathol. 2008;30(1).
8. Kapoor VK. Incidental gallbladder cancer. Am J Gastroenterol. 2001; 3:627-9.
9. Dix FP, Bruce IA, Krypezyk A, Ravi S. A selective approach to histopathology of the gallbladder is justifiable. Surg. 2003;1(4):233-5.
10. Samad A. Gall bladder carcinoma in patients undergoing cholecystectomy for cholelithiasis. J Pak Med Assoc. 2005;55(11):497.
11. Barcia JJ, Rodríguez A, Siri L, Maslorenes A, Szwebel P, Acosta G. Gallbladder carcinoma in the "Hospital de Clinicas" of Uruguay: 1998-2002. A clinicopathologic study of five cases in 802 cholecystectomies. Ann Diagn Pathol. 2004;8(1):1-5.
12. Randi G, Franceschi S, La Vecchia C. Gallbladder cancer worldwide: geographical distribution and risk factors. Int J Cancer. 2006;118(7):1591-602.
13. Shen HX, Song HW, Xu XJ, Jiao ZY, Ti ZY, Li ZY, et al. Clinical epidemiological survey of gallbladder carcinoma in north western China, 2009-2013: 2379 cases in 17 centers. Chronic Dis Translational Med. 2017;3(1):60-6.
14. Batra Y, Pal S, Dutta U, Desai P, Garg PK, Makharia G, et al. Gallbladder cancer in India: a dismal picture. J Gastroenterol Hepatol. 2005;20(2):309-14.
15. Kumar JR, Tewari M, Rai A, Sinha R, Mohapatra SC, Shukla HS. An objective assessment of demography of gallbladder cancer. J Surg Oncol. 2006;93(8):610-4.
16. Barcia JJ, Rodríguez A, Siri L, Maslorenes A, Szwebel P, Acosta G. Gallbladder carcinoma in the “hospital de clinicas” of Uruguay: 1998–2002. A clinicopatologic study of five cases in 802 cholecystectomies. Ann Diagn Pathol. 2004;8(1):1-5.
17. Henson DE, Albores-Saavedra J, Code D. Carcinoma of the gall bladder. Histologic types, stage of disease, grade, and survival rates. Cancer. 1992;70(6):1493-7.
18. Hidalgo LA, Feliú J, Admella C, Muns R, Suñol X. Gallbladder adenocarcinoma: tumoral staging, histological prognostic factors and survival. Cirugia Espanola. 2005;77(1):18-21.
19. Poudel R, Singh SK, Basnet S, Devkota H, Adhikari SK. Clinicopathological study of gall bladder cancer and its relationship with gall stones. J Soc Surg Nepal. 2015;18(3):46-.
20. Hundal R, Shaffer EA. Gallbladder cancer: epidemiology and outcome. Clin Epidemiol. 2014;6:99.
21. Taylor HW, Huang JK. ‘Routine’ pathological examination of the gallbladder is a futile exercise. Brit J Surg. 1998;85(2):208.
22. Zou S, Zhang L. Relative risk factors analysis of 3,922 cases of gallbladder cancer. Chinese J Surg. 2000;38(11):805-8.
23. Hamdani NH, Qadri SK, Aggarwalla R, Bhartia VK, Chaudhuri S, Debakshi S, Baj SJ, Pal NK. Clinicopathological study of gall bladder carcinoma with special reference to gallstones: our 8-year experience from eastern India. Asian Pacific J Cancer Prevention. 2012;13(11):5613-7.
24. Duffy A, Capanu M, Abou-Alfa GK, Huitzil D, Jarnagin W, Fong Y, et al. Gallbladder cancer (GBC): 10-year experience at memorial Sloan-Kettering cancer centre (MSKCC). J Surg Oncol. 2008;98(7):485-9.
25. Alexander S, Lemmens VE, Houterman S, Nollen L, Roumen R, Slooter GD. Gallbladder cancer, a vanishing disease?. Cancer Causes Control. 2012;23(10):1705-9.

Cite this article as: Sahai S. A clinical epidemiological study of gallbladder carcinoma: a retrospective study. Int J Adv Med 2019;6:404-7.