

Article



Hepatoblastoma in adults with clinical symptoms and histopathological evaluation

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Abstract: Introduction: Three instances of the HBs' encapsulation with a nodular, grey-white exterior surface were noted. The sliced surface was feshy, tan to grey-white, and there were patches of necrosis and haemorrhage.

Material and Methods: Names, ages, genders, and other information were recorded. Alpha fetoprotein levels (APLs), complete blood counts, liver function tests, and other laboratory data were noted. Contrast computed tomography was used to assign PRETEXT staging (CT).

Results: Patients frequently reported vomiting in 25 cases, 43 cases of abdominal distension, 21 cases of jaundice, and 19 cases of an abdomen mass. The distinction was noteworthy (P 0.05).

Conclusion: Researchers discovered that the majority of cases were epithelial as opposed to mixed epithelial and mesenchymal, and that women predominated.

Keywords: Histopathological; Hepatoblastoma; HB.

1. Introduction

R isk factors include things like prematurity, low birth weight, maternal smoking, alcohol usage, oral contraceptive use, and assisted reproductive technologies. Moreover, familial adenomatous polyposis, Li-Fraumeni syndrome, trisomy 18, Beckwith- Wiedemann syndrome, and other metabolic disorders have been linked to it [1]. Three instances of the HBs' encapsulation with a nodular, grey-white exterior surface were noted. The sliced surface was feshy, brown to grey-white, and there were necrotic and hemorrhagic patches throughout. One tumor had translucent regions that were observed. Review of the neoplasms' histological sections stained with haemotoxylin and eosin (H & E). Tumor subtype, necrosis, and mitotic activity were all analyzed. Weinberg and Finegold have summarized the histological criteria used to diagnose HB [2].

Histology is crucial since it is used in the Children's Oncology Group (COG) protocols for treatment planning as a risk stratification criterion [3]. Each histological characteristic is observed to have unique clinical correlations. Only a few studies have properly addressed the risk status, the pretreatment extent of the tumor (PRETEXT) staging, and the histologic favorability required by College of American Pathologists (CAPs) protocols [4].

The most frequent primary malignant liver tumor in children is hepatoblastoma (HB) [5]. Most frequently, it is connected to high alpha-fetoprotein (AFP), which is useful for diagnosis, assessing therapy response, and follow-up. Currently, HB is linked to very good survival rates of between 70 and 80 percent, whereas before the development of chemotherapy (CHT), these rates were under 30 percent [6]. This is a result of advancements in contemporary imaging, surgical methods like liver transplantation (LTX), and effective CHT regimens [7,8].

2. Material and Methods

The current investigation involved 68 cases of hepatoblastoma in both sexes. Participants were told of the study and given the opportunity to provide their permission. Names, ages, genders, and other information were recorded. Alpha fetoprotein levels (APLs), complete blood counts, liver function tests, and other laboratory data were noted. Contrast computed tomography was used to assign PRETEXT staging (CT). Histological information on biopsy and/or resection specimens as well as follow-up information on survival and treatment response were gathered and examined. P value 0.05 was regarded as significant.

3. Results

Table 1 shows that out of 68 cases, males were 38 and females were 30.

Table 1. Distribution of cases

Total- 68			
Gender	Males	Females	
Number	38	30	

Table 2 shows that common complaints of patients was vomiting in 25, abdominal distension in 43, jaundice in 21 and abdomen mass in 19 cases. The difference was significant (P < 0.05).

Table 2.	Clinical	sym	ptoms
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Clinical symptoms	Number	P value
Vomiting	25	
Abdominal distension	43	0.012
Jaundice	21	0.012
Abdomen mass	19	

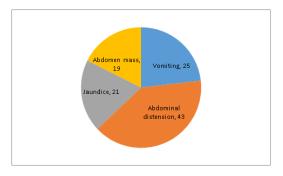


Figure 1. Symptoms

Table 3 shows that out of 48 cases, epithelial type was seen in 42 and mixed epithelial and mesenchymal in 26. Epithelial type 8 was pure fetal in 23, fetal mitotically active in 11, embryonal in 4 and mixed epithelial in 9 cases. The difference was significant (P < 0.05).

Clinical symptoms	Number	P value
Vomiting	25	
Abdominal distension	43	0.012
Jaundice	21	0.012
Abdomen mass	19	

Table 3. Histo	pathological	features
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4. Discussion

INI positive/negative small cell histology types have been included to the classification system. A favourable prognosis is linked to [9] INI1 positive tumors. As opposed to less favorable and unfavorable histological grades, favorable histology corresponds to a favourable prognosis, hence it must be included in the final report. While being relatively uncommon in children, liver tumors present a significant therapeutic and diagnostic challenge [10]. Primary liver tumors are the tenth most common tumor in children, accounting for 0.5% to 2% of all pediatric malignancies. Nonetheless, primary liver tumors account for 15% of all abdominal tumors in children; 66% of these are malignant, with hepatoblastoma being the most prevalent (HB). Comprehensive surgical removal of the tumor combined with adjuvant chemotherapy is linked to a good prognosis [11]. The goal of the current investigation was to histopathologically evaluate hepatoblastoma cases. Males made up 28 of the 48 cases in the current study, while females made up 20. Vomiting was a prevalent complaint for patients in 20, abdominal distension in 38, jaundice in 16 and abdominal mass in 14 cases, according to our research. The median age of these 10 infants upon diagnosis was 11 months, and only 1 child was born prematurely, according to Archana et al. [12] retrospective analysis on 10 children who had been diagnosed with hepatoblastoma. Abdominal distension was found in the majority of children. At presentation, one kid had lung metastases. 90% of the kids had elevated levels of alpha fetoprotein. The children were categorized according to their histological kinds, which included fetal, embryonal, macrotrabecular, and mixed epithelial-mesenchymal types. PRETEXT 1, 2, and 4 were present in three different children.

Out of 48 instances, we discovered that 32 had epithelial types and 16 had mixed epithelial and mesenchymal types. In 18 cases, epithelial type 8 was exclusively fetal, in 6, it was embryonal in 4, and in 4 cases, it was mixed epithelial. Between 1976 and 1995, Bhattacharya et al. [13] observed nine instances, of which eight were hepatoblastomas and one was a mesenchymal hamartoma. There was a 2:1 male to female ratio found, and 78% of the cases included infants under 2 years old. Five hepatoblastomas were exclusively epithelial, while cartilage, bone, and spindle sarcomatous cells were seen in the mesenchymal components of mixed hepatoblastomas.

An ongoing investigation has been conducted for potential molecular prognostic variables that could aid in the detection and management of this neoplasm. The relationship between biology and pathology is still not obvious, though. There are two primary histological forms of HB: mixed epithelial/mesenchymal and epithelial. The subtypes of the epithelial type include macrotrabecular and tiny cell type, fetal, embryonal, combined fetal and embryonal, and embryonal. The presence of some extra-mesenchymal components, like cartilage or osteoid, defines the mixed type [14,15]. The cellular basis of this neoplasm may account for variances in histology amongst different subgroups of HB, as well as the fact that 40% of tumor samples have both epithelial and mesenchymal components. HB can develop from less differentiated cells as well as primordial hepatoblasts. Poorly differentiated hepatic stem cells and human fetal liver multipotent progenitor cells (hFLMPCs) can differentiate into a wide range of tissues, including hepatocytes, bone, fat, or bile ducts. As a result, the variability of HB may be explained by its probable hFLMPC origin [16].

HB is known to be linked to conditions like Wilms' tumor, Beckwith-Wiedemann syndrome, familial adenomatous polyposis, hemihypertrophy, cleft palate, and glycogen storage diseases. In rare instances, human chorionic gonadotropin (HCG) elaboration by the tumor results in isosexual precocity [17]. Loss of heterozygosity on chromosome 11p has been observed in tumor tissue, supporting the hypothesis that aberrations on chromosome 11 play a role in the recognized link between HB and Beckwith-Wiedemann syndrome. The increased prevalence of HB in families with familial adenomatous polyposis suggests that chromosome 5q abnormalities may be significant. HB has also been linked to fetal alcohol syndrome, gonadotropins, and the consumption of contraceptives by mothers [18].

The study's flaw is its limited sample size.

5. Conclusion

Researchers discovered that the majority of cases were epithelial as opposed to mixed epithelial and mesenchymal, and that women predominated.

Author Contributions: All authors contributed equally to the writing of this paper. All authors read and approved the final manuscript.

Conflicts of Interest: The authors declare that they do not have any conflict of interests.

References

- Roebuck, D. J., Aronson, D., Clapuyt, P., Czauderna, P., de Ville de Goyet, J., Gauthier, F., ... & Perilongo, G. (2007).
 2005 PRETEXT: a revised staging system for primary malignant liver tumours of childhood developed by the SIOPEL group. *Pediatric radiology*, *37*, 123-132.
- [2] Herzog, C. E., Andrassy, R. J., & Eftekhari, F. (2000). Childhood cancers: hepatoblastoma. The oncologist, 5(6), 445-453.
- [3] López-Terrada, D., Alaggio, R., De Dávila, M. T., Czauderna, P., Hiyama, E., Katzenstein, H., ... & Finegold, M. J. (2014). Towards an international pediatric liver tumor consensus classification: proceedings of the Los Angeles COG liver tumors symposium. *Modern pathology*, 27(3), 472-491.
- [4] Stocker JT. (2001). Hepatic tumors in children. *Clin Liver Dis* 5,259-81.
- [5] Czauderna, P., Haeberle, B., Hiyama, E., Rangaswami, A., Krailo, M., Maibach, R., ... & Meyers, R. (2016). The Children's Hepatic tumors International Collaboration (CHIC): novel global rare tumor database yields new prognostic factors in hepatoblastoma and becomes a research model. *European Journal of cancer*, 52, 92-101.
- [6] Chopra, A., Iyer, V. K., Agarwala, S., Mathur, S. R., Aron, M., Gupta, S. D., & Verma, K. (2010). Apoptotic protein expression, glycogen content, DNA ploidy and cell proliferation in hepatoblastoma subtyping and their role in prognostication. *Pediatric surgery international*, 26, 1173-1178.
- [7] Baheti, A. D., Stanescu, A. L., Li, N., & Chapman, T. (2017). Contrast-enhanced CT features of hepatoblastoma: can we predict histopathology?. *Clinical imaging*, 44, 33-37.
- [8] Aronson, D. C., Czauderna, P., Maibach, R., Perilongo, G., & Morland, B. (2014). The treatment of hepatoblastoma: Its evolution and the current status as per the SIOPEL trials. *Journal of Indian Association of Pediatric Surgeons*, 19(4), 201.
- [9] Fasano, M., Theise, N. D., Nalesnik, M., Goswami, S., Garcia de Davila, M. T., Finegold, M. J., & Greco, M. A. (1998). Immunohistochemical evaluation of hepatoblastomas with use of the hepatocyte-specific marker, hepatocyte paraffin 1, and the polyclonal anti-carcinoembryonic antigen. *Modern pathology: an official journal of the United States and Canadian Academy of Pathology, Inc*, 11(10), 934-938.
- [10] Arora, R. S., Ramandeep, D., & Arora, S. (2012). Outcomes of hepatoblastoma in the Indian context. *Indian pediatrics*, 49(4), 307.
- [11] Mesquita, M. C. D. O., Ferreira, A. R., Veloso, L. F., Roquete, M. L. V., Lima, A. S. D., Pimenta, J. R., ... & Penna, F. J. (2008). Pediatric liver transplantation: 10 years of experience at a single center in Brazil. *Jornal de pediatria*, 84, 395-402.
- [12] Archana, B., Thanka, J., Sneha, L. M., Scott, J. J. X., Arunan, M., & Agarwal, P. (2019). Clinicopathological profile of hepatoblastoma: An experience from a tertiary care center in India. *Indian Journal of Pathology and Microbiology*, 62(4), 556.
- [13] Bhattacharya, S., Lobo, F. D., Pai, P. K., & Pai, G. K. (1998). Hepatic neoplasms in childhood–a clinicopathologic study. *Pediatric surgery international*, 14, 51-54.
- [14] Rowland, J. M. (2002). Hepatoblastoma: assessment of criteria for histologic classification. Medical and pediatric oncology, 39(5), 478-483.
- [15] Adesina, A. M., Lopez-Terrada, D., Wong, K. K., Gunaratne, P., Nguyen, Y., Pulliam, J., ... & Finegold, M. J. (2009). Gene expression profiling reveals signatures characterizing histologic subtypes of hepatoblastoma and global deregulation in cell growth and survival pathways. *Human pathology*, 40(6), 843-853.
- [16] Dan, Y. Y., Riehle, K. J., Lazaro, C., Teoh, N., Haque, J., Campbell, J. S., & Fausto, N. (2006). Isolation of multipotent progenitor cells from human fetal liver capable of differentiating into liver and mesenchymal lineages. *Proceedings of the National Academy of Sciences*, 103(26), 9912-9917.
- [17] Schnater, J. M., Aronson, D. C., Plaschkes, J., Perilongo, G., Brown, J., Otte, J. B., ... & Vos, A. (2002). Surgical view of the treatment of patients with hepatoblastoma: Results from the first prospective trial of the International Society of Pediatric Oncology Liver Tumor Study Group (SIOPEL-1). *Cancer*, 94(4), 1111-1120.
- [18] Luo, J. H., Ren, B., Keryanov, S., Tseng, G. C., Rao, U. N., Monga, S. P., ... & Michalopoulos, G. K. (2006). Transcriptomic and genomic analysis of human hepatocellular carcinomas and hepatoblastomas. *Hepatology*, 44(4), 1012-1024.



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