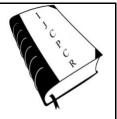
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TO DIAGNOSIS OF ASCETIC FLUID CHOLESTEROL AND SERUM ALBUMIN IN DIFFERENTIAL VALUES OF ASCITES

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ABSTRACT

Ascites is determined as presences of pathological changes in peritoneal cavity. It's mainly accumulation of free fluid. **Aims and Objectives:** In prospective study ascitic fluid cholesterol levels in various types of ascites, to compare diagnostic values of ascitic fluid cholesterol levels v/s SAAG, to study SAAG in various types of ascites. To find out specificity & sensitivity of ascitic fluid cholesterol levels in differential diagnosis of ascites. The study was conducted on 100 patients of ascites of different etiologies admitted in Bhaarath Medical College and Hospital, chennai of 15-65 years of either sex. The following study groups were made viz.:-

Group A 29 Cases of ascites caused by cirrhosis of liver

Group B 30 cases of ascites caused by Tuberculosis

Group C 25 cases of ascites caused by Malignancies

Group D, E, F25 cases of ascites caused by others (e.g. CHF, CRF, Nephrotic Syndrome, Anemia-hypoproteinemia, Bacterial Peritonitis, etc. The paracentesis was performed after proper positioning of patient .150 ml. of ascitic fluid was drawn and examined for, total protein, albumin, adenosine deaminase (ADA) (when indicated), sugar, cholesterol, total cell count, cell type, malignant cells, micrbiological tests (AFB stain , Gram staining, aerobic and anaerobic culture, sputum culture,) Liver function test, Serum cholesterol, Sputum for AFB and culture (When indicated), ECG, Chest X-ray, X-ray abdomen, Ultrasonography of abdomen: Histopathological examination. Exclusion Criteria- Haemodynamically unstable patients: Arterial hypotension and bleeding abnormality. **Conclusion:** Ascitic fluid cholesterol is an effortless, despicable and consistent biochemical parameter to differentiate (cirrotic) transudative and other ascites from malignant ascites. It is not useful in differentiating transudative cirrhotic from other tubercular ascites.

Key words Nephrotic Syndrome, Anemia-hypoproteinemia, Bacterial Peritonitis.

INTRODUCTION

Ascites is determined as presences of pathological changes in peritoneal cavity. It's mainly accumulation of free fluid. Clinically highest cases of ascites are a most complications diseases such as cirrhosis, congestive heart failure, tubercular peritonitis, nephrosis, disseminated malignancy, ect.As the development of ascites in cirrhotic patients is related with a mortality of 15%-44% for long periods. ^{1, 2}Ascites can only be treated by correction of essential cause.

Therefore, to valuation of patient with ascites is imperfect without cause of ascites is recognized. If not, a positive diagnosis of ascites patients in malignancy/any infection is proved by cytology or culture, a final cause cannot be absolutely recognized by predictable analysis of ascitic fluid. Some time malignant tumors can be produce ascites without shedding malignant cells into ascitic fluids, that is, by setting up inflammation of the peritoneum and blocking lymphatic or blood vessels. These cells are rarely found in patients with diffuse hepatic metastasis in the absence of peritoneal implants and in patients with hepatocellular carcinoma superimposed on cirrhosis with portal hypertension. Little amount of ascitic fluid has minute defer and poor preservation of cell, at the same time, benign "mesotheliosis.

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It is impossible to differentiate from malignant cells by routine morphology studies alone.

Recently, malignant and tubercular are both ascites are transuding in nature with lymphocytic predominance. Serum-ascites albumin gradient (SAAG) showed less value and It is not differentiated easily from each other. Ascitic fluid adenosine deaminase activity (ADA) is notable elevated in tubercular peritonitis than other ascites diseases. Many studies have shown ascitic fluid fibronectin and cholesterol are establish superior to the conventional methods of ascitic fluid analysis in differentiating ascites caused by malignancies from others.³⁻⁵ Estimation of cholesterol levels in ascitic fluid has been initiate useful in discriminate various types of maligant ascites from tubercular ascites .

The cholesterol may initiate in cell membrane can be degeneration of tumor cells or surrounding benign cell.Which can enter the cavity from the interstitial space because of obstructed lymph vessels or be related to raised permeability of the carcinomatous serous membrane or due to enhanced movement of plasma liporproteins into the cavity of peritoneal cavity. Elevated cholesterol concentrations have also been reported in inflammatory conditions involving the peritoneum, acute pancreatitis and chronic cardiac congestion. Usually, Ascitic fluid cholesterol is not a reflection of serum cholesterol concentration.⁶Ascitic fluid cholesterol alone due to higher concentration of cholesterol in effusion is more specifically related to tumor involvement of the serosal cavity. Now a days, fibronectin and cholesterol levels of peritoneal fluid had been found to differentiate among ascites of various aetiologies. The estimation of fibronectin stages in fluid is sensitive but complicated. Therefore the present study we have taken. To diagnosis of ascetic fluid cholesterol and serum albumin in differential values of Ascites.

Aims and Objectives

- To determine the ascitic fluid cholesterol levels in various types of ascites.
- To evaluated SAAG in various types of ascites.
- To compare diagnostic values of ascitic fluid cholesterol levels v/s SAAG.
- To compare clinical diagnosis with biochemical and cytological findings.
- To study specificity & sensitivity of ascitic fluid cholesterol levels in differential diagnosis of ascites. Material and methods

This prospective study was conducted. One hundred patients with ascites attending Bhaarath Medical College and Hospital,chennai were selected for the study. The study was approved by Ethics Committee and informed consent was obtained.

100 cases of ascites of different etiology were selected in the age group of 15-65 years of either sex. Group A: 29 Cases of ascites caused by cirrhosis of liver, Group B 30 cases of ascites caused by Tuberculosis, Group C 25 cases of ascites caused by Malignancies, Group D 16 cases of ascites caused by others e.g. CHF, CRF, Nephrotic Syndrome, Aneia-hypoproteinemia, Bacterial Peritonitis.

Paracentesis procedure:

- Proper positioning of patients lower left abdominal quadrant.
- Aseptic precautions taken.
- 150 ml samples collected in two vials- one in EDTA for cytology& another one in plain vial for biochemistry.
- Samples were processed as soon as possible.

Ascitic fluid was drawn and examined like complete blood picture, liver profile, serum amylase and lipase, serum creatinine Alfa fetoprotein, CA-125, Prothrombin Time, Lipid profile, Serum CRP, Urine albumin, 24 hours urinary protein. Biochemical analysis of Ascitic fluid for Albumin, Total Protein, Cholesterol, Glucose, Amylase, LDH, CRP and ADA, Cytological examination for Cell counts and Differential count. Ascitic fluid with microbiological Analysis for gram stain, ZN stain, malignant cells and bacterial culture were performed. Other examination such as USG abdomen, upper GI Endoscopy, CT scan abdomen and Pelvis. Echocardiogram, Thyroid Profile, ECG, 2D echo, X ray chest. Upper GI endoscopy, FNAC of the peritoneal nodules and liver biopsy other tumor marker were done in selected cases where it was needed.

Inclusion criteria:

- Age >15 years
- Clinical features suggestive ascites.

Exclusive criteria:

- Haemodynamically unstable patients:
- Those patients who are already on treatment of chemotherapy and or radiotherapy.
- Bleeding abnormality.

Data analysis:

The results were statistically analyzed and their normal distribution, range, the mean, and standard deviation were calculated. Two tailed probability values of <0.05 was taken as representing significance.

Results:

The frequently cirrhosis and tuberculosis are 3rd and 4th decade of ascites. In the 4th and 5th decade showed malignancy.In female, more common in tuberculosis congestive cardiac failure and anaemia. While cirrhosis of liver, nephrotic syndrome & malignancy were frequently results in males.

Sl.no	Aetiological	Cases	Percentage							
Group A	Cirrhosis of liver	29	29%							
Group B	Peritoneal Tuberculosis	30	30%							
Group C	Ascites in malignancy	25	25%							
Group D	Congestive Cardic failure	6	6%							
Group E	Nephrotic syndrome	05	5%							
Group F	Anemia hypoprotienaemia	05	5%							

Table:1 Aetiological distribution of ascites cases.

Table 2. Age wise distribution of ascites cases

Groups	Age ranges							Ages mean ±S.D.				
	23-40)	41-50)	51-60		>61		percentage			
	Μ	F	Μ	F	М	F	Μ	F	Total	Total	Male	Female
									Male%	Female%		
Α	5	4	9	3	6	1	1	0	21(72.4%)	8(27.5%)	5.25 ±3.30	2 ±1.82
В	6	4	5	3	8	2	1	1	20(66.6%)	10(33.3%)	5 ±2.94	2.5±1.29
С	4	2	6	3	4	5	1	-	15(60%)	10(40%)	3.75±2.06	2.5±2.08
D	0	1	2	1		2	-	-	2(33.3%)	4(80%)	0.5 ±1	1 ±0.81
E	-	-	3	1	-	-	1	-	4(80%)	1(20%)	1± 1.41	0.25± 0.5
F	-	1	2	-	1	-	1	-	4(80%)	1(20%)	1±0.81	0.25± 0.5

Out of 29 patients in group (cirrhotic), 21 (72.4%) were male and 8 (27.5%) were female.

Amongst 30 patients of group (tubercular ascites), 20(66.6%) were male and 10 (33.3%) were female. In malignant ascites group , out of 25 patients, 15 (60%) were male and 10 (40%)were females. Congestive Cardic failure Out of 6 patients 2 (33.3%) were male and 4(80%) were female. The maximum number of cases in group and were in the age group of 41-50 years (11& 8 respectively). In group, the significant numbers of 29 cases were in age group of more than 51 years. While in group, age group of 20-40 years had maximum cases (15). The mean age among patients of group was 43.68.89 years, in group, 39.9 8.14 years, in group, 52.710.76 years, while in group it was 38.47.8 years

Table: 3 Gross appearances of Ascites fluids

Colure of fluid	Cirrhosis	Peritoneal	Ascites in	Congestive	Nephrotic	Anemia	
	of liver	Tuberculosis	malignancy	Cardic failure	syndrome	hypoprotienaemia	
Yellow	69%	-	-	100%	100%	100%	
Amber	17.2%	-	-	-	-	-	
Turbid	10.3%	90%	64%	-	-	-	
Haemorragic	3.4%	10%	36%	-	-	-	

In present study, turbid appearance mostly present in the tuberculous peritonitis and in Malignancy .while in the cases of cirrhosis of liver, congestive cardiac failure, nephrotic syndrome and anaemia mostly clear yellow/amber coloured. In tuberculosis (90% cases) and ascites in Malignancy (64 % cases) haemorrhagic fluid was established.

Table: 4 Distribution of SAAG

SAAG(mg/dl)	Group-A	Group-B	Group-C	Group-D	Group-E	Group-F
	No of patients					
0-1.1	19(65.5%)	24(80%)	6(24%)	4(66.6%)	4(80%)	3(60%)
>1.1	10(34.4%)	6(20%)	19(76.0%)	2(33.3%)	1(20%)	2(40%)

Ascitic fluid cholesterol (mg/dl)ranges	Group-A N=29	Group-B N=30	Group-C N=25	Group-D N=6	Group-E N=5	Group-F N=5	Mean±S.D.
	No. of patients						
<25	9(31.0%)	12(40%)	4(16%)	1(16.6%)	1(20%)	-	4.5± 4.92
26-47	10(34.4%)	16(53.3%)	5(25%)	1(16.6%)	3(60%)	3(60%)	6.33± 5.64
48-60	6(20.6%)	1(3.3%)	13(52%)	4(66.6%)	-	2(40%)	4.33 ±4.76
>60	4(13.7%)	1(3.3%)	2(8%)	-	1(20%)	-	1.33 ±1.50

 Table: 5 Distribution of ascites cholesterol

Discussion:

Nowadays one of the most frequent challenges in the treatment of ascites patients that is cirrhosis, congestive heart failure, tubecular perionitis, nephrosis, disseminated carcinomas. This fluid can be exudates of tuberculous peritonitis and malignancy is common and also transudate, form that play vital role various cirrhosis, renal problem and other pathological problem. Many authors identified for new biochemical markers in the serum and ascitic fluid is still under identification In present study cholesterol have shown to be capable marke. In our study we found most of our cases (51%) in age group 41to 50 yrs., which matches with study of yogesh kumar . et al2017⁷. i.e. (72%).

In the present study 66 % patients were male and 34 % (1) were female that correlated with Nath et al 1966⁸ in which male and female percentage was 70% and 30% respectively. 42% cases of Jain S.C. et al 1966⁹ and Mehrotra M.P¹⁰1964 (30%).(50%) cases of ascites were seen in tuberculosis that are comparable with 30% our observation due to more social campaigning, early identification & treatment of tuberculosis we found .

Our study observation 06 % cases of congestive cardiac failure as a cause of ascites. which is correlated with Mehrotra M. P. and Mangal R.P. 1964 (6%) study. For unexplained reason CCF was found to be the cause of ascites in a study by Sikka et al 1967^{11} . (21.7%). Similar to the finding of Tito L.et al 1988^{12} (56.7%) and Ljubi Ci. C.N. et al 1993 (59.2%) ¹³ cases.

In present study male and female had about equal rate of malignant and other ascites , which was correlated to Garg R et al ¹⁴and Sood et al in cirrhotic and tubercular ascites ,while elevated frequency 15, 12 of malignant ascites in female than male. Predominant cell type in ascitic fluid was lymphocyte in all groups, except in group –B where it was RBC which was similar to the study by Simon B et al.¹⁵

Similarly cytological examination for malignant cells in group –C was positive in 12out of 25(48%) patients whereas sensitivity, specificity, positive predictive value, negative predictive value and diagnostic accuracy was 48% 100%, 100%, 64. 51%, and 72. 5% respectively. Even

though, the specificity was very high (100%) but the sensitivity was very low (48%). Cytological examination cannot be used as a good screening diagnostic tool. These results are similar with studies of Gerber AL et al ¹⁶. Rommette et al ¹⁷, Castardo et al¹⁸.

In present study, in serum total protein of cirrhotic with tubercular and malignant ascites showed low diagnostic accuracy and sensitivity. The serum ascites albumin rise was >1.1gm / in cirrhotic ascites and other ascites while it was < 1.1 gm/ dl in tubercular and malignant ascites this results correlated by Pierre Pare et al.¹⁹

Using cut off, value of ascitic fluid cholesterol 48 mg/dl had lowest sensitivity (15%) and diagnostic accuracy (50.50%) there is no diagnostic importance in differentiating cirrhotic from tubercular ascites, while at same cut off value of 48 mg/ dl it appeared highest sensitivity (89.5%) and diagnostic accuracy (95%) in differentiating cirrhotic from malignant ascites. This findings is similar to the observation made by Garg R et al and Gupta R et al. There was no significant difference in the serum cholesterol level in group A, group B, group C and group D,E,F and In ascites due to malignancy a mean ascitic fluid cholesterol level of 61 mg/dL was found. These values specify that cholesterol levels in ascites due to malignancy are significantly higher than those found in non - malignant lesion these results correlated with Martin Prieto et al²⁰ have found high concentration of serum cholesterol in malignant ascites group than non malignant ascites group.

Conclusion:

In conclusion of our study, Ascitic fluid cholesterol having high specificity, it can be used for differentiating between non-malignant and malignant ascites and also help to differentiate tubercular ascites from malignant ascites. Therefore, this parameter is being clearcut and cost efficient can be widely used to differentiate non-malignant and malignant ascites but is not helpful in differentiating cirrhotic and others from tubercular ascites.

REFERENCE:

- 1. Biecker E. Diagnosis and therapy of ascites in liver cirrhosis. World J Gastroenterol 2011;17:1237–1248.
- 2. Planas R, Montoliu S, Balleste B, Rivera M, Miquel M, Masnou H, et al. Natural
- 3. history of patients hospitalized for management of cirrhotic ascites. ClinGastroenterol Hepatol 2006;4:1385-1394.
- Colli A, Guglielmo B et al: Diagnostic accuracy of fibronectin in differential diagnosis of ascites. Cancer . 1986;58:2489-2493.
- 5. Deverbizer G et al: Fibronectin a marker for malignant ascites. Lancet 1989; 11:1104.
- 6. Martin P et al.: Comparison of ascitic fibronectin cholesterol and SAAG. Diag. Dis.and Sci. 1988; 33:833-838.
- 7. Jungst D, Xie et al.: Pathophysiology of elevated ascitic fluid cholesterol in malignant ascites. J. Hepatol. 1992, March,14(23) 244-8.
- 8. Yogesh kumardhandh et al: To study the role of ascitic fluid cholesterol in differntial diagnosis of ascites. indian journal of applied research volume 7/issue 2/february 2017/issn 2249-555x / if : 3.919 /ic value : 79.96
- 9. Nath K, Mital HS, Mishra SD, et al. Diagnostic value of ascitic fluid examination. J Assoc Physicians India 1968;16(12):991 6
- 10. Jain SC, Misra SM, Misra NP, et al. Diagnostic value of ascitic fluid examination. J Asso Physicians India 1966;14(1):59 69.
- 11. Mehrotra MP, Mangal RP, Biochemical and cytological findings of ascitic fluid in different disorders a preliminary report. J Assoc Physicians India 1964;12:53 -61.
- 12. Sikka KK, Nath K, Agarwal SN, et al. Study of serum proteins in health and in ascites of varying aetiology. JAssoc Physicians India 1967;15(12):591 4.
- 13. Tito L, Rimola A, Gines P, et al. Recurrence of spontaneous bacterial peritonitis in cirrhosis: frequency & predicative factors. Hepatology 1988;8(1):27 31.
- 14. Ljubicic N, Bilic A, Babic Z, et al. The significance of low levels of total proteins, albumin, globulins and compliment factors, in ascitic fluid and the development of spontaneous bacterial peritonitis in patients with liver cirrhosis. Lijec Vjesn 1992;114(9 12):213 5.
- 15. Garg R, Sood A, Arora S et al.: Ascitic fluid cholesterol in differential diagnosis of ascites. JAPI. 1993;41(10):644-46.
- 16. Simon Bar- Meir et al.: Analysis of ascitic fluid in cirrhosis . Dig.Dis.Sci. 1979:24 (2): 136144.
- 17. Garbes AL et al.: Recent aspects in diagnosis and therapy of ascites .J. Gastroenterol.1987, 25 (10): 672-82.
- 18. Romett J et al.: ascitic fluid analysis in malignancy related ascites. Hepatology. 1989; 8:1104-9.
- 19. Castaldo G, Ciminol et al.: Total discrimination of peritoneal malignant ascites from cirrhosis and hepatocarcinoma associated ascites by assay of ascitic fluid cholesterol and LDH. Clin. Chem. 1994;40(3):478-83.
- 20. Pierre Pare et al:Serum ascites albumin concentration gradient. A physiologic approach to the differential diagnosis of ascites. Gastroenterology. 1983;85:240-4.
- 21. Prieto M, Gómez Lechón MJ, Hoyos M, et al. Diagnosis of malignant ascites. Comparison of ascitic fibronectin, cholesterol, and serum ascites albumin difference. Dig Dis Sci 1988:33(7):833 8.