

TO STUDY THE ASSESSMENT OF LIPID ABNORMALITIES IN PATIENTS SUFFERING FROM GALLSTONES

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ABSTRACT

BACKGROUND

80 percent of gallstones are made of cholesterol, whereas 20 percent of gallstones are made of calcium salts and bilirubin. Gallstone diseases being common disorders might have an association with abnormal lipids.

Aims and Objectives- This study is done to assess the lipid abnormalities in patients suffering from gallstone diseases.

MATERIALS AND METHODS

This is a case control study, the sample size was taken for convenience during the study, and was carried out in the Department of Biochemistry among the healthy individuals and patients of gallstones attending the OPD of General Surgery, Rama Medical College Hospital and Research Centre, Mandhana, Kanpur (UP). The gallstones were collected after cholecystectomy. 5 mL of blood sample was collected after overnight fasting from each of the participants to measure the serum lipid parameters. Student's independent t-test was used to examine the degree of significance. P values less than 0.05 were considered significant.

Limitations- Since the calculated sample size was too high and thereby not feasible to include in this limited period of study, we had to limit the sample size for convenience.

RESULTS

Levels of serum triglyceride and levels of VLDL are statistically significant ($p < 0.05$), while the levels of serum cholesterol, HDL and LDL are not statistically significant ($p > 0.05$). The results are depicted in different tables to compare the groups individually.

CONCLUSION

Although, this small sample size study concluded that there is correlation between presence of gallstones and TGs as well as VLDL, but a larger sample size study is required to further strengthen these risk factors.

KEY WORDS

Gallstones, Lipid Profile, Cholesterol (C), Low-Density Lipoprotein (LDL), High-Density Lipoprotein (HDL), Triglycerides, Very Low-Density Lipoprotein (VLDL).

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BACKGROUND

80 percent of gallstones are made of cholesterol, whereas 20 percent of gallstones are made of calcium salts and bilirubin¹ and are known as pigment stones. Cholesterol stones may develop when there is too much cholesterol in the bile secreted by liver. Bile usually dissolves or breaks down cholesterol² and bilirubin is a chemical produced when liver destroys old red blood cells. Stones form when gallbladder cannot break down the excess bilirubin. Gallstone diseases being common disorder might have an association with abnormal lipids. Blood lipid profile is a panel of blood tests that serves as an initial broad medical screening tool for abnormalities in lipids such as cholesterol and triglycerides.³

The lipid profile typically includes Low-density lipoprotein (LDL), High-density lipoprotein (HDL), Triglycerides and Total cholesterol.⁴⁻⁷ Using their values, we may also calculate Very low-density lipoprotein (VLDL) and Cholesterol: HDL ratio. Our study is aimed in finding the lipid abnormalities in patients suffering from gallstone diseases.

MATERIALS AND METHODS

The case control study was used in the study and the sample size was taken for convenience during the study, which was carried out in the Department of Biochemistry among the healthy individuals and patients of gallstones attending to the OPD of General Surgery, Rama Medical College Hospital and Research Centre, Mandhana, Kanpur (UP). This study was conducted from 1st April 2016 to 31st December 2016.

Gallstones are solid particles that develop in the gallbladder. Stones are formed from the crystallisation of bile, a fluid made by the liver and secreted into the bowel through the bile ducts to help digest fats. The gallstones were collected after cholecystectomy.

Total subjects taken were 50 cases and 50 controls. These cases and controls were selected by using a standard questionnaire. It was used as a study tool to collect the data including basic profile of participants and the proforma also included the different types of investigations related to the

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study, i.e. lipid profile assessment, ultrasound of the abdomen. A verbal or a written consent was obtained from participants before the sample collection.

These cases and controls were selected from the patients who visited the Department of Biochemistry, Rama Medical College Hospital and Research Centre, Mandhana, Kanpur (UP).

All the patients were in the age group of 20 to 70 years with the mean age in cases was 43.2 ± 13.69 and in controls was 46.2 ± 14.39 . Comparison of serum lipid profile between patients and control groups showed that the levels of serum total cholesterol (147.22 ± 38.71) in patients were slightly higher than that of the control group (138.32 ± 49.54), but there was no significant variation in total cholesterol ($p > 0.05$) between patient and control groups.

A standard questionnaire was used as study tool to collect the data including basic profile of participants and the proforma also included the different types of investigations related to the study, i.e. lipid profile assessment, ultrasound of the abdomen. A verbal or a written consent was obtained from participants before the sample collection. 5 mL of blood sample was collected after overnight fasting from each of the participants to measure the serum lipid parameters. Serum cholesterol levels were measured with Cholesterol Reagent (CHOD-PAP). Serum triglycerides were measured with Triglycerides DES Reagent (Dynamic Extended Stability with Lipid Clearing Agent GPO- Trinder Method, End Point). HDL Cholesterol were measured with ERBA Cholesterol Reagent. (Phosphotungstic Acid Method, End Point). LDL Cholesterol was measured by Friedewald equation and the beta-quantification method. $VLDL \text{ Cholesterol} = TGs/5$. The

gallstones were collected after cholecystectomy. Statistical Software namely SPSS 21.0 version was used for the analysis of the data and Microsoft Word and Excel had been used to generate the graphs and tables. Descriptive statistics like mean, standard deviation and student's independent t-test was used to examine the degree of significance. P values less than 0.05 was considered significant.

RESULTS

Total patients taken were 50 cases and 50 controls, patients participated in this study Cases i.e. patients with gallstone disease and Controls i.e. patients without gallstone disease. Out of these 26 (52%) males and 24 (48%) females were in case group and had gallstone disease whereas 33 (66%) were males and 17 (34%) were females in the control group. All the patients were in the age group of 20 to 70 years with mean age in cases was 43.2 ± 13.69 and in controls was 46.2 ± 14.39 . Comparison of serum lipid profile between patients and controls group showed that the levels of serum total cholesterol (147.22 ± 38.71) in patients were slightly higher than that of the control group (138.32 ± 49.54), but there was no significant variation in total cholesterol ($p > 0.05$) between patients and controls group. LDL-C was even lower in patients (74.74 ± 28.84) as compared to control group (76.40 ± 37.00). HDL-C was slightly higher in patients (47.78 ± 15.47) than the control group (44.06 ± 15.65), but it was also not significant. In our study triglycerides (126.34 ± 64.83) and very low-density lipoprotein (25.50 ± 12.56) concentration had significant ($p < 0.05$) increase in patients with gallstone formation compared with control (105.66 ± 38.55 and 21.26 ± 7.96 respectively).

Age Group (in yrs.)	Cases				Total		Controls				Total	
	Male		Female				Male		Female			
	No.	%	No.	%	No.	%	No.	%	No.	%	No.	%
20-30	5	19	6	25	11	22	5	15	4	24	9	18
30-40	4	15	7	29	11	22	6	18	4	24	10	20
40-50	6	23	6	25	12	24	5	15	5	29	10	20
50-60	7	27	5	21	12	24	9	27	4	24	13	26
60-70	4	15	-	-	4	8	8	24	-	-	8	16
Total	26		24		50		33		17		50	
Mean \pm S.D.	43.2 ± 13.69						46.2 ± 14.39					

Table 1. Age Wise distribution of Cases and Controls

Sex	Cases		Control	
	No.	%	No.	%
Male	26	52%	33	66%
Female	24	48%	17	34%
Total	50		50	

Table 2. Sex Wise distribution of Cases and Controls

Range (mg/dL)	Cases				Total		Controls				Total	
	Male		Female				Male		Female			
	No.	%	No.	%	No.	%	No.	%	No.	%	No.	%
80-100	4	15	2	8	6	12	5	15	4	24	9	18
100-120	6	23	4	17	10	20	6	18	3	18	9	18
120-140	4	15	2	8	6	12	5	15	5	29	10	20
140-160	6	23	5	21	11	22	8	24	3	18	11	22
160-180	4	15	6	25	10	20	5	15	2	12	7	14
180-200	2	8	0	0	2	4	1	3	0	0	1	2
200-220	0	0	4	0	4	8	1	3	0	0	1	2
220 and above	0	0	1	0	1	2	2	6	0	0	2	4
Total	26		24		50		33		17		50	
Mean \pm S.D.	147.22 ± 38.71						138.32 ± 49.54					

Table 3. Serum Cholesterol Level in Cases and Controls

*p > 0.05 not significant

Range (mg/dL)	Cases				Total		Control				Total	
	Male		Female				Male		Female			
	No.	%	No.	%	No.	%	No.	%	No.	%	No.	%
0-50	1	4	1	4	2	4	1	3	0	0	1	2
50-100	11	42	9	38	20	40	19	58	9	53	28	56
100-150	8	31	7	29	15	30	7	21	8	47	15	30
150-200	4	15	5	21	9	18	5	15	0	0	5	10
200-250	0	0	1	4	1	2	0	0	0	0	0	0
250-300	2	8	-	-	2	4	1	3	0	0	1	2
300 and above	0	-	1	-	1	2	0	0	0	0	0	0
Total	26		24		50		33		17		50	
Mean ± S.D.	126.34 ± 64.83						105.66 ± 38.55					

Table 4. Serum Triglyceride Level in Cases and Control Group

*p < 0.05. Comparison of cases and control group shows that p < 0.05 was moderately significant.

Range (mg/dL)	Cases				Total		Controls				Total	
	Male		Female				Male		Female			
	No.	%	No.	%	No.	%	No.	%	No.	%	No.	%
0-25	2	8	0	0	2	4	0	0	1	6	1	2
25-50	5	19	1	4	6	12	7	21	3	18	10	20
50-75	8	31	7	29	15	30	8	24	5	29	13	26
75-100	9	35	11	46	20	40	12	36	8	47	20	40
100-125	2	8	3	13	5	10	4	12	0	0	4	8
125-150	0	0	1	4	1	2	1	3	0	0	1	2
150-175	0	0	0	0	0	0	0	0	0	0	0	0
175-200	0	0	1	4	1	2	0	0	0	0	0	0
200 and above	0	0	0	0	0	0	1	3	0	0	1	2
Total	26		24		50		33		17		50	
Mean ± S.D.	74.74 ± 28.84						76.40 ± 37.00					

Table 5. LDL Level in Cases and Controls

*p > 0.05 Not Significant.

Range (mg/dL)	Cases				Total		Controls					
	Male		Female				Male		Female			
	No.	%	No.	%	No.	%	No.	%	No.	%		
10-20	13	50	11	0	24	48	20	61	10	59		
20-30	8	31	7	29	15	30	9	27	6	35		
30-40	3	12	4	17	7	14	3	9	1	6		
40-50	0	0	1	4	1	2	0	0	0	0		
50-60	2	8	0	0	2	4	1	3	0	0		
60 and above	0	0	1	4	1	2	0	0	0	0		
Total	26		24		50		33		17			
Mean ± S.D.	25.50 ± 12.56						21.26 ± 7.96					

Table 6. VLDL Level in Cases and Controls

*p < 0.05 Significant.

DISCUSSION

Gallstones have higher incidence in females⁸ due to the presence of endogenous and exogenous oestrogens (Given as oral contraceptive pills or hormone replacement therapy), which induce chenodeoxycholic acid and increased total bile acid pool with consequent saturation (Table 1). Also, progesterone causes smooth muscle relaxation and impairs emptying of gall bladder.⁹ Saadeldin A et al¹⁰ reported the high incidence of gallstones in the age group of 31 - 50 years (68.08%), while it was rare in patients < 30 and elderly patients > 70 years. A probable explanation for the higher prevalence of gallstones in these age groups was given by David C¹¹ that with increasing age biliary cholesterol saturation increases due to decline in the activity of cholesterol 7α hydroxylase, the rate limiting enzyme for bile acid synthesis (Table 2). In the elderly, bile acid synthesis is

reduced, biliary cholesterol output is increased and cholesterol saturation of bile increases.

Stones are usually classified into 3 groups depending upon their colours: pale yellow and whitish stones as cholesterol stones, black and blackish brown as pigment calculi and brownish yellow or greenish with laminated features as mixed calculi. There are studies which demonstrate the percentage of the types of gallstones present in patients. Ibtisam BM et al¹² in their study found 53.85% as pigment stones, 29.23% mixed and 16.92% as cholesterol stones and these results were in agreement with Saadeldin A et al¹³ who considered that the most common type of gallstone in Sudanese patients was the pigment calculi 48 (51.07%) followed by mixed stone (31.9%) and then cholesterol type (17%). Although, in our study we have not classified the stones according to the colours, but they have been classified according to their presence and absence in patients.

In this study the levels of serum lipid profile; total cholesterol, triglycerides, high density lipoprotein cholesterol and low-density lipoprotein cholesterol were measured in patients and controls group. Comparison of serum lipid profile between patients and controls group showed that the levels of serum total cholesterol (147.22 ± 38.71) in patients were slightly higher than that of the control group (138.32 ± 49.54), but there was no significant variation in total cholesterol ($p > 0.05$) between patients and controls group (Table 3). These results were in agreement with Narjis et al⁸ and Naseem A et al¹⁴ who compared serum lipid profile between patients and controls group and showed that the levels of serum total cholesterol and LDL-C in patients were slightly higher than that of the control group, but there was no significant variation in total cholesterol and LDL-C ($p > 0.05$) between patients and controls group. However, in our study LDL-C was even lower in patients (74.74 ± 28.84) as compared to control groups (76.40 ± 37.00) (Table 5). Similarly, HDL-C was slightly higher in patients (47.78 ± 15.47) than the control groups (44.06 ± 15.65), but it was also not significant. In our study, triglycerides (126.34 ± 64.83) and very low-density lipoprotein (25.50 ± 12.56) concentration had significant ($p < 0.05$) increase in patients with gallstone formation compared with control (105.66 ± 38.55 and 21.26 ± 7.96 respectively) (Table 4 and 6). These results were in confirmation with the earlier reports of Atman¹⁵ and Narjis HA et al in their studies between lipid profile and gallstone formation. Although, gallstones are formed from supersaturation of cholesterol in the bile, but high total cholesterol levels themselves are not necessarily associated with gallstones as suggested by Dennis L¹⁶ and Portincasa.¹⁷ Some evidence suggests that high levels of triglycerides may impair the emptying actions of the gallbladder. The mechanism by which cholesterol stones form is not fully understood, but are likely as the result of a complex alteration in hepatobiliary function.¹⁸

CONCLUSION

Gallstone diseases being common disorders, might have an association with abnormal lipids. This study is done to compare the serum lipid abnormalities in patients who have cholelithiasis with controls. The results showed moderate significance in regard to TGs and VLDL in formation of gallstones. Although, this small sample size study concluded that there is correlation between presence of gallstones and TGs as well as VLDL, but a larger sample size study is required to further strengthen the findings of this study. The understanding of various other risk factors is also essential to establish the formation of gallstones.

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