DOI: 10.17957/TPMJ/17.4083

DIABETIC MOTHERS;

FREQUENCY OF MACROSOMIA AND HÝPOGLYCEMIA IN NEONATES OF (CONTROLLED VERSUS UNCONTROLLED).

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Article received on: 27/05/2017 Accepted for publication: 15/10/2017 Received after proof reading: 29/11/2017

INTRODUCTION

Diabetes is one of the commonest and important metabolic disorders that affect the health of pregnant women and infants.¹ About 3 – 10% of all pregnancies are complicated by diabetes.² Maternal diabetes is a significant cause of shortterm and long-term morbidity for the infant and the mother. Fetal complications are more frequent in patients with pre-existing diabetes than those with gestational diabetes. Early diagnosis of preexisting diabetes warrants early treatment and a strict clinical follow-up since early intervention has been shown to improve fetal and maternal outcomes in randomized controlled trials.¹

Women with diabetes seeking to conceive strive to achieve blood glucose and hemoglobin A1C (HbA1C) levels as close to normal as possible when they can be safely achieved without undue

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ABSTRACT... Objectives: To compare the frequency of macrosomia and hypoglycemia in neonates born to Pre-gestational diabetic mothers having controlled versus uncontrolled diabetes during pregnancy at Liaquat National Hospital Karachi. Study Design: Descriptive Cross-Sectional Study. Place and duration of Study: Ante-natal Clinic, Labor Room and Post Natal Unit of Pediatric Department, Liaguat National Hospital Karachi, from October 2013 to March 2014 over 376 neonates. Methodology: This study was carried out on 376 cases at ante-natal clinic, labor room and Post-natal ward. Her pregnancy record Checked for control of blood sugar level and brief history taken. Neonatal capillary blood taken at 30 minutes after birth to check the blood glucose level. Neonatal birth weight taken immediately after birth and plotted on growth chart for the assessment of macrosomia. Results: Mean age of the patients was 35.37 years, male prevalence was found higher, and that is 57.40%. Mean gestational age was 37.22 weeks. Mean duration of diabetes was 3.77 years. There were 57.70 patients with controlled diabetes. Frequency of hypoglycemia was found in 21.50% patients, while macrosomia was found in 36.20% patients. Conclusion: The frequency of hypoglycemia and macrosomia in infants of diabetic mother was 21.5% and 36.50% respectively. There was a significant difference in the frequency of macrosomia and hypoglycemia in infants of diabetic mothers among well controlled versus uncontrolled diabetics.

rords: Infant of Diabetic Mother, Hypoglycemia, Hyperglycemia, Macrosomia. Neonates.

Article Citation: Chohan MN, Ahmed I, Dass D, Shamim S. Diabetic mothers; frequency of macrosomia and hypoglycemia in neonates of (controlled versus uncontrolled).. Professional Med J 2017;24(12):1872-1877. DOI:10.17957/TPMJ/17.4083

hypoglycemia.³ There should be self-monitoring of blood glucose in all pregnant women with gestational or overt diabetes⁴ and testing before and either 1 or 2 hours after the start of each meal, as indicated, at bedtime and during the night.³ Pregnant women with overt or gestational diabetes strive to achieve a target pre-prandial blood glucose $\leq 95 \text{ mg/dL}^4$, and levels 1 hour after the start of a meal \leq 140 mg/dL (7.8 mmol/L) and 2 hours after the start of a meal ≤ 120 mg/ dL (6.7 mmol/L) when these targets can be safely achieved without undue hypoglycemia. Women with overt diabetes strive to achieve a HbA1C \leq 7% (ideally \leq 6.5%).³ Women with diabetes who receive preconception counseling have better preconception glycemic control and are more likely to have favorable pregnancy outcomes.⁵

Various previous studies shows association

between maternal diabetes and neonatal hypoglycemia and macrosomia but those were lacking the monitoring and management from very beginning of pregnancy, hence this study is done from initial days of pregnancy to make it more suitable. This study is also done to see the impact of blood glucose control during pregnancy in known pre-gestational diabetic women over blood glucose level and birth weight of newborn, as previous studies dose not touch this concept, hence to enforce the control of blood sugar level during pregnancy to prevent the neonatal hypoglycemia and macrosomia.

OBJECTIVE

To compare the frequency of macrosomia and hypoglycemia in neonates born to Pregestational diabetic mothers having controlled versus uncontrolled diabetes during pregnancy at Liaquat National Hospital Karachi

METHODOLOGY

It was a Descriptive and Cross- Sectional Study done with Non Probability Consecutive.

Technique, conducted at Gynecology OPD, the labor room and post natal unit of Liaquat National Hospital Karachi, from October 2013 to March 2014 over 376 neonates.

Pregnant women who came at Gynecology OPD for antenatal checkup and having Type 1 or Type 2 Diabetes were enrolled in the study. History was taken for the duration of diabetes and methods of its control. They were counseled for proper management to control their blood sugar level by life style modifications, dietary advice, insulin or oral medications. Blood Sugar documentation card was given to document the blood sugar level, both fasting and postprandial 3 times a week, while Hemoglobin C was recorded every 3 monthly. They were assessed again at post natal unit to check her pregnancy record, i.e. checked for control of blood sugar level and Hemoglobin A1C level.

Capillary blood was taken with the help of lancet prick and glucose level checked from the heel of the newborn using glucose reagent strips at 30 minutes after birth. In case of hypoglycemia mother feed was given and top feed was given in case of non-availability of mother feed.

Neonatal birth weight taken immediately after birth and plotted on growth chart for the assessment of macrosomia. All this data was collected and recorded by main researcher in predesigned proforma. Informed verbal consent was taken prior to asking questions and the information gathered from mothers was kept confidential. This study was done after taking the approval from Research Ethical Review Committee of Hospital.

The data entered and analyzed in statistical program SPSS version 16.0. Frequencies and percentages computed for qualitative variables like gender of baby, hypoglycemia (neonatal Glucose level), Macrosomia (weight of baby), Gravida, Para, whether diabetes controlled or uncontrolled, diabetes how controlled (Exercise, diet, Insulin), mode of delivery and duration of maternal diabetes.

The numerical variables such as age of mother (in years), gestational age of baby etc. presented as Mean \pm Standard deviation. Effect modifiers controlled by stratification of age of mother, duration of diabetes, gender of baby, gestational age of baby, parity and mode of delivery. Outcome compared among controlled and uncontrolled diabetic mothers applying chi square test. All the data calculated on 95% confidence interval. A p-value \leq 0.05 considered as statistically significant level.

Inclusion Criteria

All alive babies born to known Diabetic Mothers (whether controlled or uncontrolled diabetes) above 28 weeks of gestation (Assessed on Last Menstrual Period), Duration of Maternal Diabetes of more than 1 year.

Exclusion Criteria

Babies born before 28 weeks of gestation (Assessed on Last Menstrual Period), Babies born to Nondiabetic mothers and babies born to gestational diabetic mothers.

Operational Definitions

Diabetes: Diabetes may be defined as, that meeting the WHO criteria for diabetes that is Fasting plasma glucose level >126 mg/dl at 2m different occasions (24 hours apart)

Hypoglycemia:Any blood glucose levels of <45mg/dl (2.2mmol/l)

Macrosomia: Birth weight >90% for gestational age

Well controlled Diabetes: (Presence of either of the following)

Fasting blood glucose level = <100mg/dl,

HbA1C = <6.5%

Uncontrolled Diabetes: (Presence of either of the following)

Fasting blood glucose level = >100mg/dl,

HbA1C = >6.5%

RESULTS

Mean age of women was 35.37 years (Table-I). Most of the patient lies in age <35 years, that is 54.30% (Table-I). Male prevalence was found higher, that is 57.40% (Table-III). Mean gestational age was 37.22 weeks (Table-I). Most of the patient was present in <40 weeks of gestation that is 96.30% (Table-I). Mean duration of diabetes was 3.77 years (Table-IV). Most of the patients were present with duration of diabetes > 3 years that is 59.30% (Table-V). The majority of the patients were present with multi gravida status that is 74.50%, where as 77.70% patients were multiparous (Table-V & VI).

	Mean	S.D	Minimum	Maximum
Age of Mothers (Years)	35.37	1.49	34	37
Gestational age of Baby (Weeks)	37.22	1.13	37	43
Duration of Diabetes (Years)	3.77	1.47	2	5
Table-I. Age parameters				

Outcome		Diabetes	Diabetes Status		DValue
		Uncontrolled	Controlled	Total	P-Value
	Yes	66 (81.5)	15 (18.5)	81 (100)	
1 37	No	93 (31.5)	202 (68.5)	295 (100)	0.001
	Total	159 (42.3)	217 (57.7)	376 (100)	
Macrosomia Yes No Total	Yes	111 (81.6)	25 (18.4)	136 (100)	0.001
	No	48 (20)	192 (80)	240 (100)	
	Total	159 (42.3)	217 (57.7)	376 (100)	

 Table-II. Comparison of outcome and diabetes status n=376

Gender	Uuneghueemie	Diabetes Status		Tatal	D. Value
	Hypoglycemia	Uncontrolled	Controlled	Total	P- Value
Male	Yes	36 (92.3)	3 (7.7)	39 (100)	0.001
	No	17 (9.6)	160 (90.4)	177 (100)	
	Total	53 (24.5)	163 (75.5)	216 (100)	
	Yes	30 (71.4)	12 (28.6)	42 (100)	
Female	No	76 (64.4)	42 (35.6)	118 (100)	0.452
	Total	106 (66.2)	54 (33.8)	160 (100)	
Table-III. Comparison of hypoglycemia and diabetes status with gender of the baby					

Diabetes Status Gender Macrosomia Total **P-Value** Uncontrolled Controlled Yes 53 (94.6) 3 (5.4) 56 (100) Male No 0 (0) 160 (75.5) 160 (100) 0.001 Total 53 (24.5) 163 (75.5) 216 (100) Yes 58 (72.5) 22 (27.5) 80 (100) Female 0.132 No 48 (60) 32 (40) 80 (100) 106 (66.2) 160 (100) Total 54 (33.8) Table-IV. Comparison of macrosomia and diabetes status with gender of the baby

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Age Group	Live alve are in	Diabetes Status		Tetel	DVal	
(Years)	Hypoglycemia	Uncontrolled	Controlled	Total	P-Value	
	Yes	30 (81.1)	7 (18.9)	37 (100)	0.001	
≤ 35	No	50 (29.9)	117 (70.1)	167 (100)		
	Total	80 (39.2)	124 (60.8)	204 (100)		
	Yes	36 (81.8)	8 (18.2)	44 (100)		
>35-40	No	43 (33.6)	85 (66.4)	128 (100)	0.001	
	Total	79 (45.9)	93 (54.1)	172 (100)		
	Yes	4(80)	1 (20)	5 (100)		
>40	No	2 (22.2)	7 (77.8)	9 (100)	0.001	
	Total	6 (42.9)	8 (57.1)	14 (100)		
	Yes	11 (55)	9 (45)	20 (100)	0.001	
Diabetes Duration \leq 3 Years	No	23 (17.3)	110 (82.7)	133 (100)		
	Total	34 (22.3)	119 (77.8)	153 (100)		
	Yes	55 (90.2)	6 (9.8)	61 (100)	0.001	
Diabetes Duration > 3 Years	No	70 (43.2)	92 (56.8)	162 (100)		
	Total	125 (56.1)	98 (43.9)	223 (100)		
	Yes	36 (100)	0 (0)	36 (100)	0.001	
Primiparous	No	17 (35.4)	31 (64.6)	48 (100)		
	Total	53 (63.1)	31 (36.9)	84 (100)		
	Yes	30 (66.7)	15 (33.3)	45 (100)	0.001	
Multiparous	No	76 (30.8)	171 (69.2)	247 (100)		
	Total	106 (36.3)	186 (63.7)	292 (100)		

Table-V. Comparison of hypoglycemia and diabetes status with age, diabetes duration and parity n=376

Age Group	Maaraaamia	Diabetes Status		Tetel	
(Years)	Macrosomia	Uncontrolled	Controlled	Total	P-Value
	Yes	54 (78.3)	15 (21.7)	69 (100)	0.001
≤ 35	No	26 (19.3)	109 (80.7)	135 (100)	
	Total	80 (39.2)	124 (60.8)	204 (100)	
	Yes	57 (85.1)	10 (14.9)	67 (100)	
>35	No	22 (21)	83 (79)	105 (100)	0.001
	Total	79 (45.9)	93 (54.1)	172 (100)	
	Yes	4 (80)	1 (20)	5 (100)	
>40	No	2 (22.2)	7 (77.8)	9 (100)	0.001
	Total	6 (42.9)	8 (57.1)	14 (100)	
.	Yes	30 (66.7)	15 (33.3)	45 (100)	0.001
Diabetes Duration \leq 3 Years	No	4 (3.7)	104 (96.3)	108 (100)	
	Total	34 (22.3)	119 (77.8)	153 (100)	
	Yes	81 (89)	10 (11)	91 (100)	0.001
Diabetes Duration > 3 Years	No	44 (33.3)	88 (66.7)	132 (100)	
	Total	125 (56.1)	98 (43.9)	223 (100)	
	Yes	53 (100)	0 (0)	53 (100)	0.001
Primiparous	No	0 (0)	31 (64.6)	31 (100)	
	Total	53 (63.1)	31 (36.9)	84 (100)	
Multiparous	Yes	58 (69.9)	25 (30.1)	83 (100)	0.001
	No	48 (23)	161 (77)	209 (100)	
	Total	106 (36.3)	186 (63.7)	292 (100)	

Table-VI. Comparison of macrosomia and diabetes status with age, diabetes duration and parity n=376

Frequency of hypoglycemia was found in 21.50% patients, while macrosomia was found in 36.20% patients (Table-II). Stratification was done to see the effect of diabetes status on the outcome. Chi square test was applied and significant effect was observed as P-Value was found 0.001 in hypoglycemia and macrosomia (Table-V). Effect modifiers were controlled by stratification of age, gestational age, duration of diabetes, gender of the baby and parity.

DISCUSSION

Hypoglycemia is the most common cause of morbidity in the Infants of Diabetic Mothers and can be a challenging and protracted problem to manage. In our study the hypoglycemia was noted in (81) 21.5 %, which is supported by other similar studies conducted locally in Peshawar that showed 23.8% hypoglycemic neonates.²

Macrosomia is another major complication of the diabetic pregnancies. In this study total (136) 36.2% Infants of Diabetic Mothers were macrocosmic, (both uncontrolled and controlled diabetes during pregnancy), while a similar study from Hyderabad showed macrosomia was seen in 41.8% of fetuses.⁸

Hyderabad Another study from showed macrosomia 34% neonates.9 Another study from Saudi Arabia showed 8.2% macrocosmic neonates⁶, as 18.4% neonates in our study had macrosomia who born to mothers with controlled diabetes during pregnancy, that suggest the possibility of tight glycemic control in Saudi Arabian study. Another local study from Islamabad reported a relatively lower occurrence of macrosomia (28%), which may be due to a better glycemic control in mothers. As strict metabolic control with insulin in mothers with gestational diabetes mellitus has proven to attenuate the risk of LGA (macrosomia).

In our study Neonatal Hypoglycemia was most common in babies of uncontrolled diabetic mothers 66 (81.5), while 15 (18.5%) neonates had hypoglycemia who born to mothers with controlled diabetes during pregnancy, A similar study from Saudi Arabia showed neonatal hypoglycemia in 19 (24.4%) neonates¹⁰, although this study was done only on gestational diabetic mother but results are similar. A local study from Islamabad showed hypoglycemia in 28% of cases whose mother's blood sugar was not controlled during pregnancy.¹¹

Macrosomia was most common in neonates born to mothers having uncontrolled diabetes during pregnancy 111 (81.6), while 25 (18.4) neonates had hypoglycemia at birth, who born to mothers having controlled diabetes during pregnancy, while much better results were obtained in a study from Saudi Arabia that showed macrosomia in 9(11.5) neonates born to controlled diabetes during pregnancy.¹⁰

In our study hypoglycemia was more common in male neonates 36 (92.3%) as compared to female neonates 30 (71.4%), while male neonates of controlled diabetic mothers had less 3 (7.7%) neonatal hypoglycemia as compared to female neonates 12 (28.6%). There was no difference in macrosomia in males 53 (94.6%) and females 58 (72.5%), of uncontrolled diabetic mothers, while more females 22 (27.5%) were macrocosmic as compared to males 3 (5.4%).

Maternal age more than 40 years had less neonatal hypoglycemia 4 (80%) as compared to less than 40 years age 36 (81.8%) in babies of uncontrolled diabetic mothers, while babies of controlled diabetic mothers (Less than 40 years of age) has neonatal hypoglycemia in 1 (20%). Mothers whose diabetic control was >3 years before pregnancy and had uncontrolled diabetes during pregnancy born 55 (90.2%) newborns with hypoglycemia, while women who had diabetes <3 years before pregnancy had only 11 (55%) hypoglycemic neonates.

Parity had no association with neonatal hypoglycemia in uncontrolled diabetic mothers, while in controlled multiparous diabetic mothers had more neonates 15 (33.3%) neonates with hypoglycemia, while Primiparous had 0 (0%) neonatal hypoglycemia. There was no difference in in macrosomia in neonates of uncontrolled diabetic mothers 111 (81.6%) with age <40

years and age > 40 years 4 (80%). Women who had diabetes < 3 years before pregnancy and uncontrolled diabetes had 30 (66.7%) macrocosmic newborns, while women who had diabetes > 3 years had 81 (89.6%) macrocosmic newborns.

CONCLUSION

The frequency of hypoglycemia and macrosomia in infants of diabetic mother was 21.5% and 36.50% respectively. There was a significant difference in the frequency of macrosomia and hypoglycemia in infants of diabetic mothers among well controlled versus uncontrolled diabetics.

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AUTHORSHIP AND CONTRIBUTION DECLARATION

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2	Imran Ahmed	Analysis, Interpretation of data.	Carpe 2
3	Deve Dass	Critical revision of article, Statistical expertise.	Jung-
4	Samina Shamim	Final approval of analysis.	Stan Streen