

## HbA<sub>1</sub>C Level in Late Trimester with Maternal Outcome in Diabetic Patients

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### ABSTRACT

**Objective:** The present study was undertaken to determine the association of maternal serum HbA<sub>1</sub>C level with maternal outcome.

**Materials and Methods:** It was prospective observational study. The study was carried out in the department of Obstetrics and Gynaecology in BIRDEM Hospital during the period of September 2006 to August 2007. During this study period, 100 pregnant patients with diabetes who attended or admitted at BIRDEM Hospital were studied. Estimation of serum HbA<sub>1</sub>C level was done in all patients in each trimester. From each patient 5 cc blood was taken & HbA<sub>1</sub>C level was measured with the help of enzymatic method. HbA<sub>1</sub>C level < 6 was considered as normal. The maternal complications in antenatal period, in postpartum period, during labour outcome were studied in both cases of controlled & uncontrolled HbA<sub>1</sub>C level.

**Result:** In this study serum HbA<sub>1</sub>C level was found raised in uncontrolled diabetic patients. The incidence of vulvovaginitis, preterm delivery and polyhydramnios were significantly high in 2nd and 3rd trimester in raised HbA<sub>1</sub>C level. The rate of normal vaginal delivery was higher inpatient with normal HbA<sub>1</sub>C level than uncontrolled HbA<sub>1</sub>C level (17.59% V/s 10.84%, p = 0.01), which statistically significant. Postpartum haemorrhage (PPH) was significantly higher in raised HbA<sub>1</sub>C level than normal (0.00% V/s 22.20%, p=0.01) in NVD and (0.00% V/s 16.22%, p = 0.01) in Caesarean section.

**Conclusion:** There is increasing evidence that the raised level of maternal serum HbA<sub>1</sub>C in antenatal period is associated with maternal & neonatal complications. By investigating HbA<sub>1</sub>C level in each trimester, blood sugar control can be done. This study was taken out to evaluate the usefulness of HbA<sub>1</sub>C for good glycaemic control in diabetic pregnancy.

**KEYWORDS:** Diabetes, HbA<sub>1</sub>C, Maternal outcome.

### INTRODUCTION

Diabetes Mellitus is an important medical disorder in pregnancy, which creates substantial risk for the mother and fetus during current pregnancy and it also has serious implication for their long time well being<sup>1</sup>. Pregnancy and preconception period are of particular importance to people with diabetes as pregnancy challenges to the metabolic management in diabetes and, at the same time it increases risk of diabetes related complications in mother (e.g. pre eclampsia, infection, postpartum haemorrhage, increase incidence of caesarian section, traumatic delivery. The discovery of glycosylated Hb has opened new horizon in all aspect of diabetic research and management<sup>2</sup>. HbA is the major component of adult Hb, comprising approximately 90%

of Hb. This Hb when combines with glucose becomes glycosylated (HbA<sub>1</sub>C). This glycosylated Hb (HbA<sub>1</sub>C) are negatively charged and thus migrate quickly than HbA on cation exchange chromatography. Glycosylated HbA<sub>1</sub>C is increased in diabetes as a consequence of chronic hyperglycemia<sup>3,4</sup> and co-relate closely with their blood level and urinary excretion of glucose<sup>5,6</sup>. Poor glycaemic control is associated with an increased risk of maternal complications, suggesting that strict glycaemic control may reduce the rate of maternal morbidity<sup>7</sup>. Glycosylated haemoglobin HbA<sub>1</sub>C levels were higher in the spontaneous preterm delivery group.<sup>8</sup>

There is increasing evidence that raised level of maternal serum HbA<sub>1</sub>C in antenatal period can cause maternal

complications<sup>9</sup>. HbA<sub>1c</sub> proved to be a useful indicator of average long term blood glucose level in diabetic and non-pregnant subjects<sup>10</sup>. Thus by investigation of HbA<sub>1c</sub> in each trimester, blood sugar control in each trimester can be done. Thus, adequate screening, strict control of hyperglycaemia and careful planning for pregnant diabetic women ensure a happy outcome. We had undertaken this study to see the maternal outcome in diabetic pregnancy in case of controlled and uncontrolled serum HbA<sub>1c</sub> level.

**OBJECTIVES**

- To evaluate the usefulness of HbA<sub>1c</sub> for good glycemic control for diabetes mellitus in pregnancy.
- To measure HbA<sub>1c</sub> in the 2<sup>nd</sup> and 3<sup>rd</sup> trimester of pregnancy in women with diabetes.
- To assess the pregnancy outcome in women with raised HbA<sub>1c</sub>.
- To assess the pregnancy outcome in women with normal HbA<sub>1c</sub>.

**MATERIALS AND METHODS**

It was a prospective observational study carried out from September 2006 to August 2007 in Department of Obstetrics and Gynaecology at Bangladesh Institute of Research and Rehabilitation in Diabetes, Endocrine and Metabolic disorder (BIRDEM). Hundred pregnant women with diabetes who attended or admitted to

BIRDEM Hospital during the study period were recruited.

Pregnant women with preexisting diabetes and with gestational diabetes were included in this study. Multiple pregnancies, pregnancy with other metabolic disorders, heart disease and chronic hypertension were excluded from this study. The variables included in the proposed study were age, status of glycemic control, complications in 2<sup>nd</sup> and 3<sup>rd</sup> trimester of pregnancy, mode of delivery, intrapartum and postpartum complications. Data collection sheet has formed which include all the variables of interest. Cases were collected from outdoor and in patient department of Obstetrics and Gynaecology, BIRDEM Hospital, Dhaka. Purpose and procedure of the study were discussed with the patients who fulfill the inclusion criteria. All the variables of interest were collected from history; clinical examination and biochemical investigation were recorded on the predesigned data collection sheet. Pregnancy was dated by early ultrasonography. Some patients were managed initially only by dietary advice and some needed injection Insulin also. From each patient 5c.c.blood was taken and HbA<sub>1c</sub> was measured by laboratory method. HbA<sub>1c</sub> level < 6% was considered as normal. Data were processed by Computer and analyzed by using SPSS (Statistical Package for Social Science). For statistical analysis Student ‘t’ test was used. It was considered statistically significant if p < 0.05.

**Table 1: Age distribution of the study subject.**

| Age (in years) | Case (n=100) | Percentage (%) |
|----------------|--------------|----------------|
| 20-25          | 12           | 12.00          |
| 26-30          | 48           | 48.00          |
| 31-35          | 26           | 26.00          |
| >35            | 14           | 14.00          |

Mean age ±SD = 29.77±4.52

**Table 2: Status of Glycemic control in late trimester of pregnancy in respect with HbA<sub>1c</sub> level**

**Controlled HbA<sub>1c</sub> level (n = 100)**

| Controlled HbA <sub>1c</sub> level |       | Uncontrolled HbA <sub>1c</sub> |       |
|------------------------------------|-------|--------------------------------|-------|
| No.                                | (%)   | No.                            | (%)   |
| 17                                 | 17.00 | 83                             | 83.00 |

**Table 3: Maternal outcome in controlled and uncontrolled HbA<sub>1c</sub> in 2<sup>nd</sup> & 3<sup>rd</sup> trimester of pregnancy.**

|                  | Controlled HbA <sub>1c</sub> |       | Uncontrolled HbA <sub>1c</sub> |       | p value |
|------------------|------------------------------|-------|--------------------------------|-------|---------|
|                  | (n=17)                       | (%)   | (n=83)                         | (%)   |         |
| Vulvovaginitis   | 1                            | 5.88  | 18                             | 21.67 | 0.02    |
| UTI              | 2                            | 11.76 | 15                             | 18.07 | 0.24    |
| Preterm delivery | 3                            | 17.64 | 20                             | 24.10 | 0.04    |
| Polyhydramnios   | 2                            | 11.76 | 18                             | 21.69 | 0.03    |

Unpaired ‘t’ test, p < 0.05 was considered significant.

**Table 4: Mode of delivery in respect with status of HbA<sub>1c</sub> in 2<sup>nd</sup> & 3<sup>rd</sup> trimester of pregnancy.**

|      | Controlled HbA <sub>1c</sub> |       | Uncontrolled HbA <sub>1c</sub> |       | p value |
|------|------------------------------|-------|--------------------------------|-------|---------|
|      | (n=17)                       | (%)   | (n=83)                         | (%)   |         |
| NVD  | 12                           | 70.59 | 9                              | 10.84 | 0.01    |
| LUCS | 5                            | 29.41 | 74                             | 89.16 | 0.23    |

Unpaired ‘t’ test, p < 0.05 was considered significant.

**Table 5: Intrapartum complications in study group who delivered vaginally.**

|                        | Controlled HbA <sub>1</sub> C |       | Uncontrolled HbA <sub>1</sub> C |       | p value |
|------------------------|-------------------------------|-------|---------------------------------|-------|---------|
|                        | (n=12)                        | (%)   | (n=9)                           | (%)   |         |
| Shoulder dystocia      | 0                             | 0.00  | 3                               | 33.33 | 0.01    |
| Complete perineal tear | 0                             | 0.00  | 4                               | 44.44 | 0.13    |
| Cervical tear          | 3                             | 25.00 | 6                               | 66.66 | 0.23    |

Unpaired 't' test,  $p < 0.05$  was considered significant.

**Table 6: Postpartum complications in study group who delivered vaginally.**

|              | Controlled HbA <sub>1</sub> C |      | Uncontrolled HbA <sub>1</sub> C |       | p value |
|--------------|-------------------------------|------|---------------------------------|-------|---------|
|              | (n=12)                        | (%)  | (n=9)                           | (%)   |         |
| PPH          | 0                             | 0.00 | 2                               | 22.20 | 0.02    |
| UTI          | 1                             | 8.30 | 2                               | 22.20 | 0.28    |
| Endometritis | 0                             | 0.00 | 1                               | 11.10 | 0.24    |

Unpaired 't' test,  $p < 0.05$  was considered significant.

**Table 7: Postpartum complications in study group who delivered by caesarean section.**

|                      | Controlled HbA <sub>1</sub> C |       | Uncontrolled HbA <sub>1</sub> C |       | p value |
|----------------------|-------------------------------|-------|---------------------------------|-------|---------|
|                      | (n=15)                        | (%)   | (n=74)                          | (%)   |         |
| PPH                  | 0                             | 0.00  | 12                              | 16.22 | 0.01    |
| UTI                  | 1                             | 20.00 | 10                              | 13.51 | 0.24    |
| Abd. Wound infection | 0                             | 0.00  | 4                               | 5.40  | 0.03    |
| Endometritis         | 0                             | 0.00  | 4                               | 5.40  | 0.08    |

Unpaired 't' test,  $p < 0.05$  was considered significant

## RESULTS

Table 1 shows the age distribution of the study objects. Age range was 20-38 years in study group. The highest incidence (48%) was found in age group 26 to 30 years. Table 2 shows the status of glycemic control in 2nd & 3rd trimester in which 17% women have controlled HbA<sub>1</sub>C level and 83% of women have uncontrolled level of HbA<sub>1</sub>C.

Table 3 shows maternal outcome in 2nd and 3rd trimester of pregnancy. Vulvovaginitis (5.88% V/s 21.67%,  $p = 0.02$ ), preterm delivery (17.64% V/s 24.10%,  $p = 0.04$ ) and polyhydramnios (11.76% V/s 21.69%,  $p = 0.03$ ) in controlled and uncontrolled HbA<sub>1</sub>C level respectively. These differences are statistically significant whereas UTI (11.76% V/s 18.07%,  $p = 0.24$ ) in controlled and uncontrolled HbA<sub>1</sub>C level which is statistically not significant.

Table 4 shows statistically higher rate of Normal vaginal delivery (NVD) in women with controlled HbA<sub>1</sub>C level than uncontrolled (70.59% V/s 10.84%,  $p = 0.01$ ). Rate of LUCS was high in women with uncontrolled HbA<sub>1</sub>C level (29.41% V/s 89.16%,  $p = 0.23$ ) which is statistically not significant.

Table 5 shows Shoulder dystocia in women with uncontrolled HbA<sub>1</sub>C level was statistically higher than women with controlled HbA<sub>1</sub>C level (0.00% V/s 33.33%,  $p = 0.01$ ), whereas complete perineal tear (0.00% V/s 44.44%,  $p=0.13$ ) and cervical tear (25.00% V/s 66.66%,  $p = 0.23$ ). These differences are not statistically significant.

Table 6 shows postpartum complications in women with uncontrolled HbA<sub>1</sub>C level higher than controlled HbA<sub>1</sub>C level. PPH (0.00% V/s 22.20%,  $p=0.02$ ) which is statistically significant whereas UTI (8.30% V/s 22.20%,

$p= 0.28$ ) and Endometritis (0.00% V/s 11.10%,  $p = 0.24$ ) are statistically not significant.

Table 7 shows PPH (0.00% V/s 16.22%,  $p = 0.01$ ), Abdominal wound infection (0.00% V/s 5.40%,  $p = 0.03$ ) are statistically higher in women with uncontrolled HbA<sub>1</sub>C level than controlled HbA<sub>1</sub>C level whereas UTI (20.00% V/s 13.51%,  $p=0.24$ ) and Endometritis (0.00% V/s 5.40%  $p=0.08$ ) are statistically not significant.

In this study serum HbA<sub>1</sub>C level was found raised in uncontrolled diabetic patients. The incidence of vulvovaginitis, preterm delivery and polyhydramnios were significantly high in 2nd and 3rd trimester in raised HbA<sub>1</sub>C level. The rate of normal vaginal delivery was higher inpatient with normal HbA<sub>1</sub>C level uncontrolled HbA<sub>1</sub>C level (17.59% V/s 10.84%,  $p = 0.01$ ), which statistically significant. Post-partum haemorrhage (PPH) was significantly higher in raised HbA<sub>1</sub>C level than normal (0.00% V/s 22.20%,  $p=0.01$ ) in NVD and (0.00% V/s 16.22%,  $p = 0.01$ ) in Caesarean section.

## DISCUSSION

Proper screening, diagnosis and management of diabetes in pregnancy can reduce maternal morbidity<sup>7</sup>. Diabetes and pregnancy may mutually affect each other over a range of interaction from conception to delivery, and possibly even later<sup>11</sup>. The highest incidence (48%) was found in age group 26 to 30 years. HbA<sub>1</sub>C level d" 6% was considered normal. In 2nd and 3rd trimester of pregnancy controlled and uncontrolled HbA<sub>1</sub>C level was (17% Vs 83%).

The incidence of vulvovaginitis 21.67% in uncontrolled HbA<sub>1</sub>C level similar to the finding in the study of Mangala R. et al<sup>12</sup> (19.8%). Incidence of UTI 18.07% in women with uncontrolled HbA<sub>1</sub>C which is similar to the

finding in the study of Khatun F<sup>13</sup> (17.8%). The incidence of Preterm delivery was 24.10% among uncontrolled HbA<sub>1</sub>C group which higher than the study done by Kovilam O et al<sup>7</sup> (10%). The incidence of polyhydramnios was 21.69% among uncontrolled HbA<sub>1</sub>C level which similar to the study of EskandarM<sup>14</sup> (23%) but higher than the study done by Metal S etal<sup>15</sup> (3.7%). In this study rate of Caesarean section in uncontrolled HbA<sub>1</sub>C group was 89.16% which is equivalent to the study done by Shikdar K et al<sup>16</sup> (87.33%), Ivy R<sup>17</sup> (88.32%), but higher than that reported by Metal S et al<sup>15</sup> (52%), Mangala R et al<sup>12</sup>(48.32%).

In this study the incidence of PPH who delivered Caesarean section was 16.22% in uncontrolled HbA<sub>1</sub>C group which is lower than the study done by Mangala et al<sup>12</sup>(29%). The incidence of Abdominal wound infection 5.40% and Endometritis 5.40% are equivalent to the study done by Mangala et al<sup>12</sup> (4.20% & 5.50%).

## CONCLUSION

Diabetes Mellitus in pregnancy is one of the leading cause of maternal morbidity and mortality. But it is preventable by controlling blood sugar level during pregnancy. There is increasing evidence that the raised level of maternal serum HbA<sub>1</sub>C in antenatal period is associated with maternal complications. HbA<sub>1</sub>C is proved to be a useful indicator of average long term blood glucose level in diabetic subjects. By investigating HbA<sub>1</sub>C level in each trimester blood sugar control can be done. Thus strict control of hyperglycemia in diabetic pregnant woman ensures a happy outcome.

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