## Significance of glycemic status in patients of major depressive disorder with suicidal behaviour

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## **Abstract**

**Background:** Suicidal ideation is common in major depressive disorder, and it is thought to be a strong contributing factor for completed or attempted suicide. Hyperglycemia and hyperinsulinemia may increase the risk of depression and also suicide risk. Therefore this study was conceived to explore the possible relationship between suicidal behaviour and impaired glucose metabolism in major depressive disorders. **Materials and Method:** This study was undertaken in Department of Biochemistry and Psychiatry, Government Medical College, Nagpur. 60 diagnosed cases of major depressive disorder in the age range of 21-60 years of either sex were enrolled. The diagnosis was determined by DSM IV criteria. HAMD scale was used to assess the severity of depression. Fasting and postprandial blood glucose along with HbA1c was determined in the study participants. **Results:** Out of 60 patients, nearly more than half of (53.33%, n=32) patients were having suicidal ideation and attempt. The male depressive cases (73.33%, n=22) outnumbered the female depressive cases (33.33%, n=10) in having suicidal ideation and attempt. Statistically significant differences were found in fasting, postprandial blood glucose and HbAa1c of the major depressive cases with no suicidal ideation and behaviour, and with suicidal ideation and attempt. **Conclusion:** The observed association between blood glucose levels, HbA1c and suicidal behaviour suggests that disturbances in glucose metabolism are associated with suicidal ideation, Suicidal attempts. **Key Words:** Blood glucose levels, Major depressive disorder, Suicidal behaviour, Suicidal ideation, Suicidal attempt.

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### INTRODUCTION

Altered lifestyle of the modern society and tendency to excel has ultimately culminated in the increasing incidence of depression. Depression is major health concern, in that it leads to significant dysfunction<sup>1</sup>, disability2, and poor quality of life in sufferers<sup>3</sup> and pose

a significant burden on the care givers<sup>4</sup>. Being multicentric in origin, depression has always attracted the researchers and still continuing. Suicidal ideation is one of the most serious and commonly seen symptom of the depression which often remains undetected. Depression is thought to be a strong contributing factor for completed or attempted suicide. Approximately 50% of those who commit suicide, suffers from serious depressive disorder<sup>5</sup>.The relationship between depression and suicidal behaviour is complex. Little literature is available on the association between depression, suicidal behaviour, glucose levels or insulin resistance although some studies suggested that higher glucose levels are associated with dysthymia<sup>6</sup> and higher HbA1c concentrations with recurrent or psychotic depression.<sup>6</sup> Association between suicidal ideation and HbA1c levels in diabetic patients has been reported<sup>7</sup>. A recent metaanalysis reported an association between depression and

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insulin resistance. Hyperglycemiaand insulin resistance may increase the risk of depression and also the risk of suicide, therefore this increases our need to explore the possible relationship between suicidal behaviour and disturbances in glucose metabolism in depressive patients.

### MATERIAL AND METHODS

The present study has been carried out in Department of Biochemistry, Government Medical College and Hospital Nagpur, during 2011- 2012. The study protocol was approved by the Institutional Ethical Committee. Informed written consent was obtained from all the study participants enrolled in the study. Study sample consisted of 60 diagnosed cases of major depressive disorders in the age group of 21 -60 years, of either sex, attending psychiatry department. The diagnosis of major depressive disorder was determined by DSM-IV criteria. Hamilton's depressive rating scale was used to assess the severity of the depressive symptoms, <sup>10</sup> higher scores on the HAMD expressed more severe symptomatology. The following severity range for the HAMD score was used to classify the depressive symptom severity: mild depression (8-16); moderate depression (17-23); and severe depression  $(\geq 24)^{11}$  in the present study suicidal behaviour was defined as 'suicidal ideation during the past month' or 'suicide attempt during the past month or in a lifetime. The patients with a prior clinical and / or laboratory evidence of hypo or hyperthyroidism, somatic illness (diabetes and renal or hepatic disorders ), infections or autoimmune diseases, a recent surgical treatment or a significantly changed body weight, axis I and axis II diagnoses and pregnant or lactating female subjects were excluded from the study. We measured the fasting blood glucoselevels and HbA1c between 8 and 9 am after 10-12 hours of fasting. Postprandial blood glucose was measured 2 hours after the meal. Blood glucose was estimated by enzymatic (Glucose oxidase - peroxidase method) whereas HbA1c was determined by Latex agglutination method which is measured as turbidity.

### **Statistical Analysis**

All values are reported as mean  $\pm$  SD. Statistical analysis was done by using the chi – square test, the student's t-test. Differences were considered statistically significant at a probability value p<0.05. All statistical analyses were performed with SPSS statistics version 19.0.

### RESULTS

**Table 1**: Sexwise distribution of depressive cases according to suicidal ideation and behaviour

Sex	No suicidal ideation and behaviour (n= 28)	Suicidal ideation and attempt (n= 32)	Total
Male (n=30)	08 (26.66 %)	22 (73.33 %)	30 (100%)
Female ( n=30)	20 (66.66%)	10 (33.33%)	30 (100%)

Table 1 shows Sexwise distribution of depressive cases according to suicidal ideation and behaviour. Out of 60 patients, nearly more than half of (53.33%) patients were having suicidal ideation and attempt. The male depressive cases (73.33%, n=22) outnumbered the female depressive cases (33.33%, n= 10) in having suicidal ideation and attempt.

**Table 2**: Age wise distribution of depressive cases according to suicidal ideation and behaviour

Age group (In years)	No suicidal Ideation and behaviour (n= 28)	Suicidal Ideation and attempt (n= 32)	Total
21-30	06 (21.42 %)	08 (25%)	14 (23.33%)
31-40	07 (25 %)	06 (18.75%)	13 (21.66 %)
41-50	08 (28.57%)	14 (43.75%)	22 (36.66 %)
51-60	07 (25 %)	04 (12.5 %)	11 (18.33 %)
Total	28 (100%)	32 (100%)	60 (100%)

Table 2 shows Age wise distribution of depressive cases according to suicidal ideation and behaviour. There were 08 (25%) depressive case in the age group 21 -30 yrs, 06 (18.75%) depressive cases in the 31-40, 14 (43.75%) depressive cases in the age group 41-50 and 04 (12.5%) depressive cases in the age group of 51-60 seen.

**Table 3**: Comparison of Blood Sugar Levels in Controls and Major Depressive Cases With and Without Suicidal Ideation and

Behaviour						
Study participants and Biochemical parameters	No suicidal Ideation and behaviour (n= 28)	Suicidal Ideation and attempt (n= 32)	P value			
Fasting blood glucose (mg/dl)	102.78 ± 07.06	109.26 ± 08.24	<0.001			
Postprandial blood glucose (mg/dl)	105.30± 08.94	120.42 ± 11.18	<0.01			
HbA1c (%)	5.68 ± 1.07	6.4 ± 1.02	< 0.05			
HAMD Score	21± 7	26 ± 8	< 0.01			

Table 3 shows Comparison of Blood Sugar Levels in Controls and Major Depressive Cases with and without Suicidal Ideation and behaviour. The mean value of fasting blood glucose was  $102.78 \pm 07.06$  mg/dl in depressive patients with no suicidal ideation and behaviour, whereas it was  $109.26 \pm 08.24$  mg/dl in depressive patients with suicidal ideation and attempts and the difference was statistically significant (p<0.001). The mean value of Postprandial blood glucose was

 $105.30\pm~08.94~$  mg/dl in depressive patients with no suicidal ideation and behaviour, whereas mean value of Postprandial blood glucose was  $120.42\pm11.18~$  mg/dl in depressive patients with suicidal ideation and attempts, and the difference was statistically significant (p<0.01). Statistically significant difference (p<0.05) was seen in the mean values of HbA1c indepressive patients with no suicidal ideation and behaviour and depressive patients with suicidalideation and attempt. HAMD Score was  $21\pm7~$  and  $26\pm8~$  in depressive cases with no suicidal ideation and behaviour, and with suicidal ideation and attempt respectively. The difference was statistically significant (p<0.01).

### **DISCUSSION**

In our study, the observed association between higher blood glucose levels, HbA1c and suicidal behaviour in depressive cases is an interesting finding. More than half of the participants in our study reported suicidal ideation or previous attempts of suicide, a similar observation was made by Sokero PT et  $al^{12}$ . The male depressive cases outnumbered the female depressive cases in having suicidal ideation and attempt in our study. A similar preponderance of male suicides was also observed by Nandi etal<sup>13</sup> and Hedge *et al*<sup>14</sup>. In the present study 22 depressive cases (36.66 %) with suicidal ideation and attempt were found to be in the age group of 41-50 years. This is supported by the fact that 71% of suicides in India (National Crime Records Bureau. Ministry of Home Affairs, Government of India: 2005) are by persons below the age of 44 years. In the present study, higher fasting and postprandial blood glucoselevels were observed in patients with suicidal ideation or who had previously attempted suicide. A similar observation of three-fold higher suicidal ideation rates in insulin-treated type 2 diabetic patients was made by Ceretta et al (2012)<sup>6</sup>. Which suggested an association between suicidal tendencies and blood glucose levels. We also observed higher HbA1c values in patients with suicidal ideation or who had previously attempted suicide. Bot et al (2013)<sup>7</sup> reported an association between higher HbA1c values and depressive mood, sleep disturbances, appetite problems and suicidal ideation. Various studies 15 reported an increased risk for suicidal ideation in adults with diabetes and depression, but to our knowledge no study have specifically studied the association between glucose levels and suicidal behaviour in depressed patients. The association observed between glucose levels and suicidal behaviour may be related to the cytokine mediated inflammatory process that results in the activation. of indoleamine 2.3-dioxygenase, the depletion of tryptophan and suicidal behaviour related to emerging serotonergic hypofunction, depression and impulsivity 16,17,18, Patients

with suicidal behaviour in our study also had higher HAMD Scores which was also observed by Brent DA *et al* (1988). Subjects scoring high on depression rating scale tend to have lack of ability to overcome suicidal wishes and may have plan and wishes to commit suicide. Therefore depression may be regarded as potent risk factor for suicidal ideation and suicide risk itself. In addition, hyperactivation of the hypothalamic-pituitary-adrenal axis has been reported in 20 to 80 % of depressed individuals. Ithis hyperactivity has been considered as an important mechanism that explains the pathophysiology of depression. Higher cortisol levels are associated with age and more severe forms of depression, may thus also contribute to the association between higher HAMD score and risk of suicidal behaviour. 11, 22

### **CONCLUSION**

The observed association between blood glucose levels, HbA1c and suicidal behaviour suggests that disturbances in glucose metabolism are associated with suicidal ideation and attempts. However further studies are necessary to elaborate the pathophysiology behind these associations.

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