Role of Imatinib in Achieving Euglycemic State in Patient with GIST

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Abstract
Gastrointestinal stromal tumor (GIST) are rare tumors arising from Interstitial Cells of Cajal and require immunohistochemical marker c-kit (CD 117) for diagnosis. Imatinib Mesylate is an orally administered drug which competitively inhibits tyrosine kinase associated with the KIT protein. We present a rare case report highlighting role of Imatinib Mesylate in achieving euglycemic state in patient with GIST. Imatinib Mesylate is an effective form of treatment for patients with KIT (CD 117) positive unresectable and/ or metastatic malignant GIST. Proper patient selection and adequate treatment may help in achieving euglycemic state in patient with GIST.

Key Words
Stromal, Immunohistochemical, Euglycemic, Unresectable

Introduction
Stromal tumor of the gastrointestinal tract are rare neoplasms accounting for <1% of all gastrointestinal tract malignancies. Gastrointestinal stromal tumor (GIST) are thought to arise from Interstitial Cells of Cajal (ICC) of the gastrointestinal tract (1). The immunohistochemical marker 'c-Kit' (CD117) identifies these cell and seems specific diagnostic marker currently available (2). Our case highlights dramatic improvement in clinical condition of patients with GIST on Imatininb Mesylate treatment.

Case Report
A 70 year old diabetic and hypertensive male presented to surgical outpatient department with complaint of pain in right upper abdomen for last six months and marked generalized weakness. Patient gave history of operation one year back for an abdominal mass. Computed Tomography (CT) abdomen at that time revealed mass arising from jejunum. Exploratory laparotomy was done along with resection of a 10 x 10 cm jejunal mass (Fig 1,4,5). No evidence of metastasis in liver or lymph nodes was noted. Histopathology report of resected jejunal mass was consistent with GIST (Fig 2,3). Patient had now presented with jaundice with total serum bilirubin 3 mg/dl. Patient has raised blood sugar profile (fasting blood sugar 126 mg/dl; postprandial blood sugar 175 mg/dl). Ultrasound abdomen and CT abdomen revealed target lesions suggestive of metastasis in bilateral lobes of liver. Ultrasound abdomen and CT abdomen revealed target lesions suggestive of metastasis in bilateral lobes of liver. Ultrasound guided FNAC from liver lesions revealed metastatic GIST. Treatment was started with Imatinib Mesylate 400 mg daily. Patient improved significantly within one month and his appetite increased. His liver function test results came within normal limits. CT abdomen at six months and at one year follow up period
revealed significant improvement in radiological findings. Patient had overall good functional status at one year. Also euglycemic status was achieved without oral hypoglycemic drugs (Graph-1).

**Discussion**

GIST are rare but are nevertheless the most common mesenchymal neoplasms of the gastrointestinal tract. Mazur and Clark first introduced the term GIST in 1983 (3).

GIST are most common in stomach (60-70%) followed by the small intestine (20-30%), the colorectum (10%) and the oesophagus (1-5%) (4,5). GIST are multicentric in fewer than 5% of cases (6). Imatinib Mesylate is an orally administered competitive inhibitor of the tyrosine kinase associated with the KIT protein (stem cell factor receptor), ABL protein and platelet derived growth factor receptors. The KIT tyrosine kinase is abnormally expressed in GIST (7). Imatinib Mesylate induces a

**Graph. 1 Showing Significant Improvement in Blood Sugar Levels with Imatinib Mesylate Treatment**
sustained objective response in more than half of patients with an advanced unresectable or metastatic GIST (8). Overall, Imatinib Mesylate was reported to be well tolerated. The most common serious events included unspecified hemorrhage and neutropenia. Skin rash, edema and periorbital edema were the common adverse events observed (9).

Review of literature revealed only two cases for Imatinib Mesylate and diabetes control (10,11). In our case, patient presented with metastatic lesions in liver one year after operation for jejunal GIST. Patient was treated with low dose Imatinib Mesylate (400 mg daily) which was well tolerated without any side effects. Patient achieved euglycemic status without oral hypoglycemics and showed progressive improvement in radiological findings.

**Conclusion**

Our case highlights Imatinib Mesylate as a effective form of treatment of patients with KIT (CD 117) positive unresectable and/or metastatic malignant GIST. Patients with GIST should be diagnosed early so that they are amenable to treatment with Imatinib Mesylate. Our case is in agreement with previous studies which highlight adjunctive role of Imatinib Mesylate with other forms of treatment.

**References**