

# **POST-RENAL TRANSPLANT LEUCOPENIA AND ITS IMPACT ON GRAFT AND PATIENT OUTCOME**

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## *Abstract:*

**INTRODUCTION AND AIMS:** Post-renal transplant leucopenia is a common clinical challenge which needs fine dose adjustment of precipitating drugs, proper management of complications and use of granulocyte colony-stimulating factor (G-CSF). Serious infections, chemo-prophylactic and immunosuppressive drug reduction may affect patient and graft outcome. **Aim:** To study incidence and management of posttransplant leucopenia and its impact on graft and patient outcome after one year.

**METHODS:** We studied renal transplant patients operated during 2010 in our center who received immunosuppression and chemoprophylaxis according to our protocol, valgancyclovir 900mg and septrin ½ D/S tablet daily for 6 months. Significant leucopenia ( $<4000 \times 10^9$ ) was managed by reduction of valgancyclovir and mycophenolate mofetil (MMF) and giving G-CSF according to the response. All patients were screened for CMV infection by CMV-PCR titers at time of transplant and at 3, 6, 9 and 12 months after transplant.

**RESULTS:** Over one year, 79 patients were transplanted and divided into leucopenia and non-leucopenia group (group 1 and 2 respectively). Twenty seven patients were in group 1 (34.17%) and had at least one attack of significant leucopenia ( $p < 0.02$ ). Mean total leucocytic count of the whole year of follow up period was significantly lower in group 1 ( $4294 \pm 1488 \times 10^9$  versus  $8205 \pm 2123 \times 10^9$ ,  $p < 0.0001$ ). Valgancyclovir and septrin were stopped completely in 76.9% while MMF was reduced to  $\geq 50\%$  in 85.7% in group 1 ( $p < 0.0001$ ). Mean neutrophil count was  $964.3 \pm 192.7 \times 10^9$  in group 1 with significant positive correlation with total leucocytic count ( $p < 0.009$ ). G-CSF injections were given to all patients in group 1 with a mean dose of 146.6 megaunit/patient without significant side effects except for mild to moderate low back pain. There were no significant differences in demographic data including induction immunosuppression ( $p < 0.327$ ), maintenance immunosuppression ( $p < 0.12$ ), cases with delayed graft function ( $p < 0.994$ ), BK viremia (0.601), and incidence of associated infections other than CMV ( $p < 0.15$ ). Four cases of CMV infection were detected in group 1 during the first 6 months while none were in group 2 ( $p < 0.012$ ). There was higher number of posttransplant diabetes mellitus (PTDM) in group 1 ( $p < 0.037$ ) most likely due to higher maintenance doses of steroids and tacrolimus to compensate for the MMF dose reduction. Mean rejection episode/patient was significantly higher in group 1 ( $0.62 \pm 0.852$  versus  $0.28 \pm 0.49$ ,  $p < 0.03$ ). There was no difference in patient outcome at 12 months (100% in both groups). Graft failure was 3.7% in group 1 versus 7.7% in group 2 without significant difference ( $p < 0.44$ ).

**CONCLUSIONS:** Significant reduction of MMF and valgancyclovir due to leucopenia resulted in significantly higher rate of rejection episodes, CMV infection and PTDM. High doses of G-CSF were used safely to treat neutropenia without significant side effects. Prospective studies using smaller dose of prophylactic valgancyclovir are required.