



## DECEASED DONOR KIDNEY TRANSPLANT WITH ATG INDUCTION AND TRIPLE IMMUNOSUPPRESSIVE AGENTS (TACROLIMUS, MMF, PREDNISONE)

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### ABSTRACT

**Aim:** To determine the outcomes of deceased donor kidney transplantation with ATG induction and triple immunosuppressive agents (Tacrolimus, MMF, and prednisone). **Materials and Methods:** A prospective observational study was conducted to determine the infection episodes, toxicity/drug induced effects of tacrolimus, mycophenolate, prednisone and ATG induction, graft function at the end of 6 months, graft loss / rejection episodes upto 6 months post transplantation in kidney transplant cases at Krishna Institute of Medical sciences (KIMS), Nephrology department. **Results:** The whole study was conducted on 30 patients out of which 16.66% (immediate graft functioning (GFR:  $\geq 77.56$ ), 60% (slow (GFR:  $\geq 56.21$ ), 23.33% (delayed (GFR: 23.33%). **Conclusion:** The current study helps in determining the improvement of graft functioning (immediate, slow, delayed) by taking triple immunosuppressants and ATG induction.

### INTRODUCTION

Chronic kidney disease (CKD) is defined as abnormalities in kidney structure or function, present for three months or longer, with implications for health [1-2]. CKD affects between 8% and 16% of the population worldwide and is often underrecognized by patients and clinicians [3-6]. Structural abnormalities include albuminuria of more than 30mg/day, presence of hematuria or red cell casts in urine sediment, electrolyte and other abnormalities due to tubular disorders, abnormalities detected by histology, structural abnormalities detected by imaging, or history of kidney transplant. CKD is classified by cause of kidney disease, glomerular filtration rate (GFR) category, and albuminuria level based on new recommendations from the kidney

Disease: Improving global outcomes (KDIGO) guidelines, referred to CGA staging (cause, GFR, albuminuria). CKD stage 5, previously referred to ESRD, occurs when the GFR falls below 15ml/min/1.73m<sup>2</sup> or in patients receiving renal replacement therapy (RRT) [7]. Hence the present study was designed to assess the effects of triple immunosuppressive agents and ATG induction in deceased donor kidney transplant patients in KIMS hospital.

### MATERIALS AND METHODS

A prospective, observational study was conducted in kidney transplant patients treated with triple immunosuppressive agents and ATG induction in Nephrology department at Krishna Institute of medical sciences over the period of six months. All CKD patients of

either sex with or without co-morbidities treated with immunosuppressive agents and ATG induction were included in the study. The patients case sheet of both gender with or without co-morbidities such as hypertension, diabetes, CKD (stage-5) were collected for this study.

### STATISTICAL ANALYSIS

Data was collected from the collected case sheets and entered in Microsoft Excel office (2007) spreadsheet and analyzed. Total number of patients and percentages were used to represent graft functioning (immediate, slow, delayed), drug toxicity, infection episodes.

### RESULTS

#### GRAFT FUNCTIONING

Graft functioning was showed in Table 1. The whole study was on 30 patients out of which 16.66% patients were showing immediate graft functioning, 60% patients were slow graft functioning, 23.33% patients were delayed graft functioning at discharge.

At the end of 6 months 16.66% of patients were having an average GFR of (people with immediate graft function)  $\geq 77.56$ , 60% of patients were having an average GFR of (people with slow graft function)  $\geq 56.21$ , 23.33% of patients were having an average GFR of (people with delayed graft function)  $\geq 31.5$ .

#### INFECTION EPISODES

Patients with infection episodes and without infections were presented in Table 2. Out of 30 patient cases, it was found that 3 patients

were under infection episodes and 27 were found without any infection episodes.

#### TOXICITY OR DRUG-INDUCED EFFECTS OF IMMUNOSUPPRESSANTS AND ATG INDUCTION

Toxicity or drug-induced effects of immunosuppressants and ATG induction were showed in Table 3. Out of 30 patients, no toxic effects were found in all the 30 patients.

#### AGE WISE CATEGORIZATION OF THE PATIENTS

Age wise categorization of the patients was presented in Table 4. Out of 30 cases, the patients belonging to age group of 18-30 years were found to be 9, 31-45 years were found to be 12 and 46-70 years were found to be 9.

#### GENDER WISE DISTRIBUTION OF THE PATIENTS

Gender wise distribution of the patients was given in Table 5. Out of 30 patients, 22 (73.33%) were male patients and 8 (26.66%) were female patients.

#### AGE AND GENDER WISE CATEGORISATION OF THE PATIENTS

Age and gender wise categorization of the patients was showed in Table 6. Out of 30 patients, 6 were found to be male patients (20%) and 3 were female patients (10%) belonging to the age group of 18-30 years, 9 were male patients (30%) and 3 were female patients (10%) belonging to the age group of 31-45 years, 7 were male patients (23.33%) and 2 were female patients (6.66%) belonging to the age group of 46-70 years.

**TABLE: 1**Graft functioning

S.NO.	GRAFT FUNCTIONING AT DISCHARGE	TOTAL NO. OF SUBJECTS	OVERALL% OF SUBJECTS	AVG. GFR AT END OF 6 MONTHS
1	Immediate	05	16.66%	$\geq 77.56$
2	Slow	18	60%	$\geq 56.21$
3	Delayed	07	23.33%	$\geq 31.5$

**TABLE: 2 - Infection episodes**

CATEGORY	FREQUENCY	PERCENTAGE
No.of patients	30	100%
No.of patients with infection episodes	3	10%
No.of patients without infection episodes	27	90%

**TABLE: 3 - Toxicity/Drug-Induced Effects of Immunosuppressants And Atg Induction**

CONDITION	NO.OF EFFECTED PATIENTS	OVERALL %
Toxic	0	0%
Non-toxic	30	100%
Total	30	100%

**TABLE: 4- Age wise categorization of the patients**

S.NO.	AGE DISTRIBUTION	NUMBER OF PATIENTS
1	18-30	09
2	31-45	12
3	46-70	09

**Table: 5- Gender wise categorization of the patients**

S.NO.	GENDER	NO. OF PATIENTS	PERCENTAGE
1	Male	22	73.33%
2	Female	08	26.66%

**TABLE: 6 - Age and gender wise categorization of the patients**

AGE	MALE		FEMALE		TOTAL NO. OF SUBJECTS	OVERALL PERCENTAGE (%)
	Frequency	%	Frequency	%		
18-30	6	20	3	10	9	30
31-45	9	30	3	10	12	40
46-70	7	23.33	2	6.66	9	30
<b>TOTAL</b>	22	73.33	8	26.66	30	100

## DISCUSSION

The current study helps in determining the improvement of graft functioning (immediate, slow, delayed) by taking triple immunosuppressants and ATG induction, by determining the percentage of slow, immediate and delayed graft functioning. During the six months study period, total number of 30 patients who visited the Nephrology department at Krishna Institute of medical science and undergone deceased donor kidney transplantation along with immunosuppressive therapy and ATG induction were selected. Total 30 deceased donor kidney transplant patients (both comorbid and non- co morbid) were distributed into different groups based on factors like graft functioning, GFR values, infection episodes and drug toxicity. Out of 30 patients, 16.66% of patients were showing immediate graft functioning ( $\geq 77.56$  GFR), 60% of patients were showing slow graft functioning ( $\geq 56.21$  GFR), 23.33% of patients were showing delayed graft functioning ( $\geq 31.5$  GFR). Out of 30 patients, it was found that 3 patients were under infection episodes and 27 were found non-infectious and no toxic effects were found in all the 30 patients. CKD is estimated to be more common in women than in men (16% vs 13%) [2,

8]. CKD is also estimated to be more common in non-Hispanic blacks than in non-Hispanic whites (18% vs 13%). 15% of Hispanics are estimated to have CKD. The main reported causes of new cases of ESRD are diabetes, high blood pressure and glomerulonephritis (inflammation of kidneys). **Antithymocyte globulin (ATG)** is produced by immunizing rabbits or horses with human thymocytes generating polyclonal antithymocyte antibodies. The major mode of action for ATG is the depletion of T cells. ATG also targets a wide range of antigens displayed on B cells, dendritic cells, NK cells, endothelial cells and disrupts cell trafficking. The goal of immunosuppressive therapy is to prevent the graft destructive immune response.

## CONCLUSION

The current clinical research study conducted at KIMS hospital, Secunderabad using 30 subjects who underwent kidney transplantation with deceased donor and the outcomes of triple immunosuppressants (tacrolimus, mycophenolate mofetil, prednisolone) and ATG induction (given before the transplantation to prevent the rejections) and GFR at discharge and at the end of 6 months were noticed to determine the graft functioning of transplanted kidney.

The current study illustrated improvement and stability in graft functioning of the recipients. The limitations that should be taken while planning this study, no formal calculation was done to estimate the number of patients required. The total number of patients was considered based on the feasibility and the limited time that was available to complete the study. The study was 6 months duration. This limited duration of the study might have not allowed reaching the maximum effect. Hence, future studies with long-term duration are recommended.

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