# CYTOKINES: A POSSIBLE INDICES OF PERITONEAL ADEQUANCY

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#### **OBJECTIVES**

The Inflammation is highly prevalent in chronic kidney disease (CKD) patients. Renal replacement therapy itself may also promotes inflammation and long-term peritoneal dialysis (PD) is related to chronic inflammatory response. Systemic and local inflammatory mediators induce histopathological alterations in the peritoneal membrane. Furthermore, the most important problem during long-term PD is the preservation of peritoneal membrane function. PD adequacy is monitored primarily by indices of small solute clearance, Kt/Vurea and creatinine clearance.

Several systemic and local inflammation-related biomarkers have been associated with membrane failure and mortality in PD patients. Inflammation often coexists with malnutrition and there is a relationship between nutritional indices, as serum Albumin (Alb), and mortality. C-Reactive Protein (CRP) is an index of inflammatory activity. IL-6 and IL-1 $\beta$  are proinflammatory cytokines and modulate inflammation.

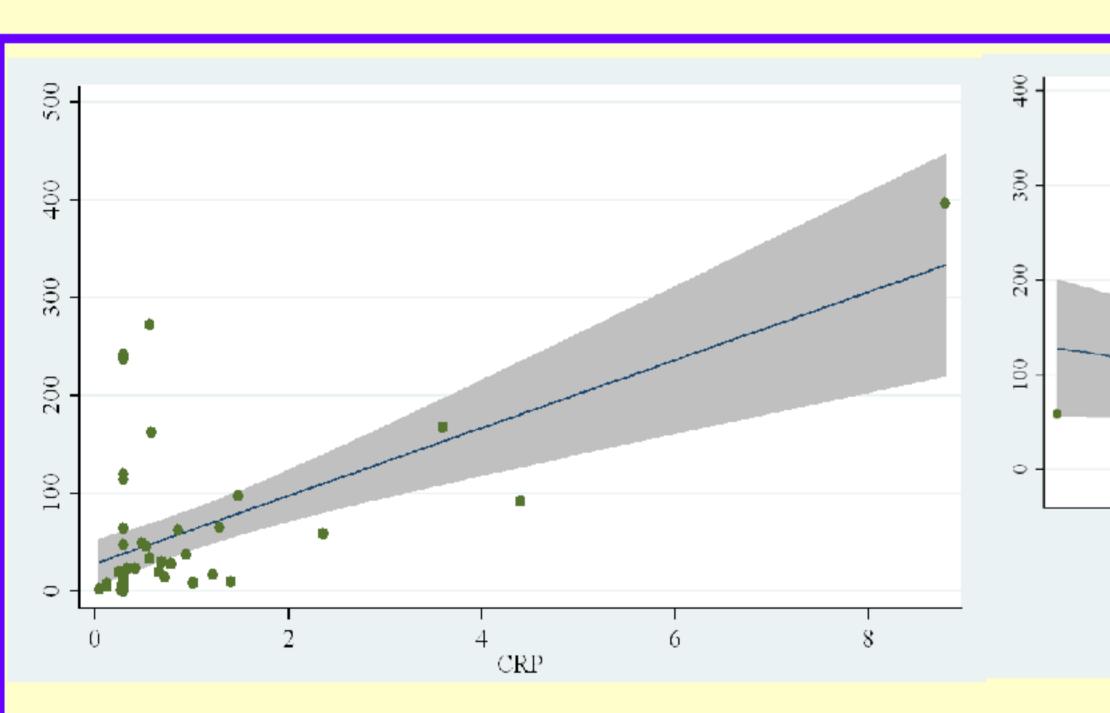
#### **METHODS**

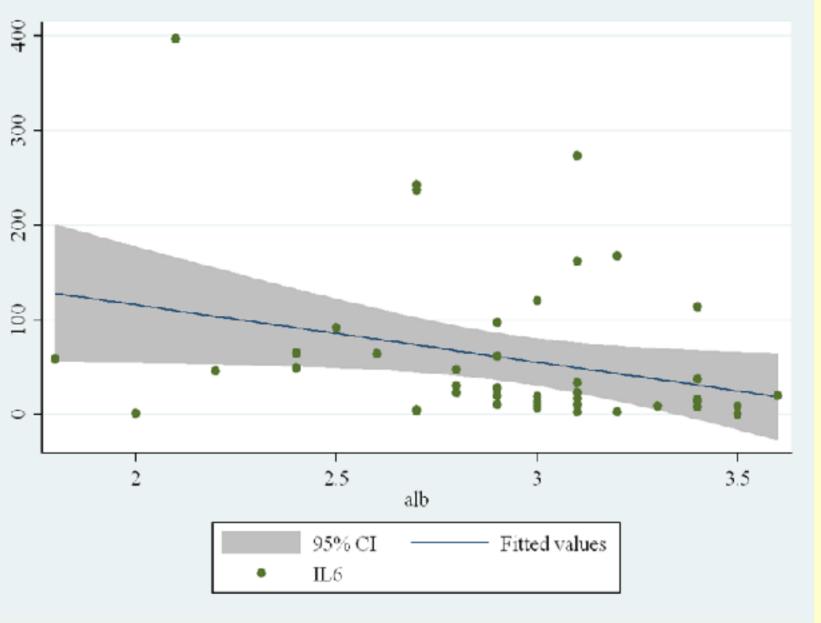
We enrolled 46 PD patients (25M/21 F, age 61.5±16.4 years) with CKD undergoing maintenance PD for a minimum of 3 months were enrolled in this study. Patients were in a stable condition and free from intercurrent illness and infection for at last 3 months. 31/46 PD patients were treated with CAPD and 15 with APD.

The average length of treatment was 21 months and the range was: minimum: 3.6 – maximum: 132.9 months. Plasma levels of Alb (g/dL), CRP (mg/dL) and cytokines (IL-6 and IL-1β, pg/mL) were measured in these patients. Quantitative determination of cytokines (IL-1β and IL-6) in plasma were performed by Human Instant ELISA kit (eBioscience, San Diego, CA, USA) according to manufacturer's instructions. Optical density was read by using a VICTORX4 Multilabel Plate Reader (PerkinElmer Life Sciences, Waltham, MA, USA) at 450 nm.

We used weekly Kt/Vurea and Creatinine Clearance (wCc) as estimates of PD adequacy. Statistical analysis was performed by STATA Software. A p<0.05 was considered statistically significant.

#### RESULTS





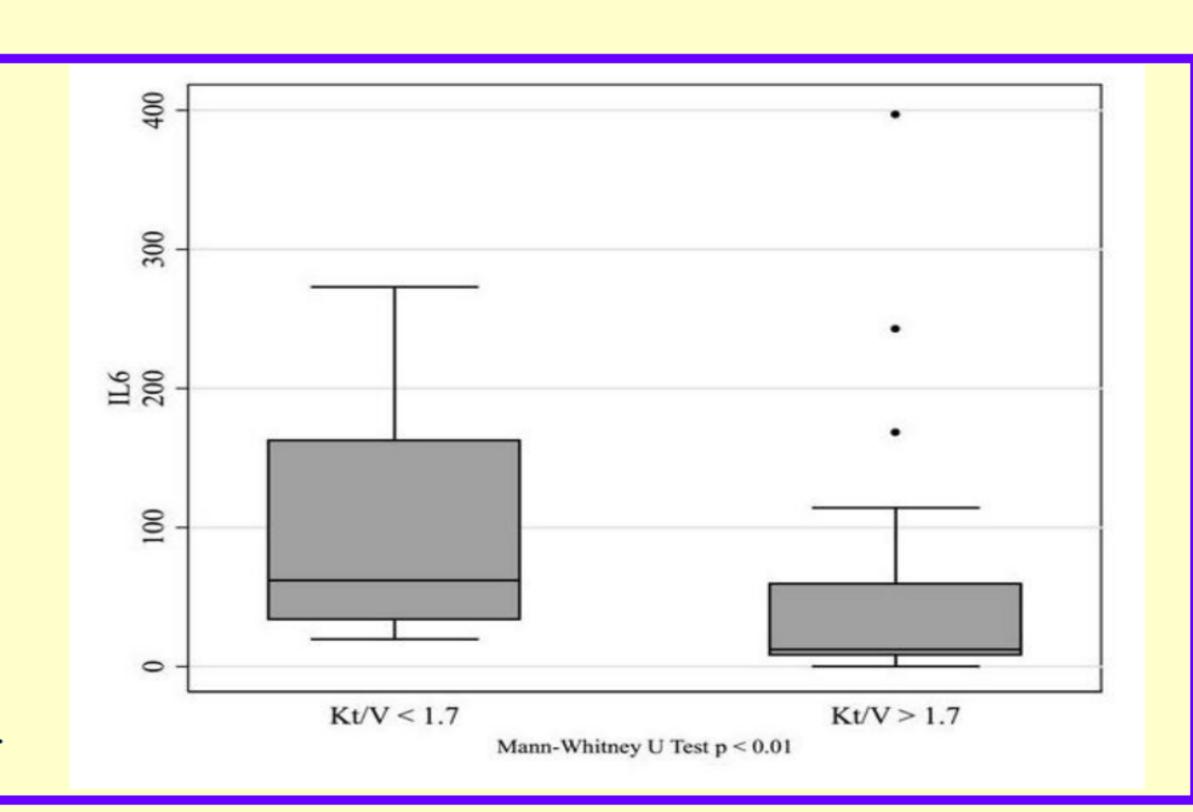


Figure 1. Correlation IL-6 and CRP and Alb

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Figure 2.

The median values of Alb, CPR, IL-6 and IL-1 $\beta$  showed no significant differences between CAPD and APD patients. IL-6 levels showed a positive correlation with CPR (p<0.001) and it correlated negatively with Alb (p=0.01) (figure 1). There is no s statistically significant relationship between IL-1 $\beta$  and CRP or Alb. Subsequently, PD patients were divided into 2 groups based on Kt/Vurea value: 1.7 was the cut-off value as recommended by K/DOQI guidelines. PD patients with Kt/V <1.7 had significantly higher IL-6 compared to PD patients with Kt/V>1.7(p=0.015) (Figure 2). The median value of IL-6 in PD patients with Kt/V<1.7 was 12 ng/ml (IQR 33-162). The median value of IL-6 in PD patients with Kt/V>1.7 was 12 ng/ml (IQR 8-59). No statistically significant relationship between IL-6 and the wCc was observed. There is no difference in IL-1 $\beta$  levels in PD patients with Kt/V <1.7 and with Kt/V<1.7 [.82 (0.88-5.2) versus 1.82 (0.95-2.7)].

### CONCLUSIONS

In conclusion, we observed the correlation between IL-6 and inflammation and nutritional markers. We reported a lower levels of IL-6 in patients with better peritoneal dialysis adequacy indices, but no difference in IL-1 $\beta$  levels. This study suggested that, unlikely from IL-1 $\beta$ , IL-6, as a marker of inflammation state, may be considered a specific index of PD adequacy.

It is necessary to increase the sample size of PD subjects enrolled to validate our hypothesis.

## References

Clinical practice guidelines for peritoneal adequacy, update 2006. American journal of kidney diseases: the official journal of the National Kidney Foundation. 2006; 48:S91-

S97.

Borazan A et al. The effects of peritoneal dialysis and hemodialysis on serum tumor necrosis factor-alpha, interleukin-6, interleukin-10 and c-reactive-protein levels. Mediators Inflamm. 2004; 13:201-4. Pecoits-Filho R et al. Systemic and intraperitoneal interleukin-6 system during the first year of peritoneal dialysis. Perit Dial Int. 2006; 26:53-63.