

## **C-Reactive Protein as a Prognostic Biomarker for Advanced Renal Cell Carcinoma Treated with Sunitinib**

**Introduction and Objective:** Since the introduction of sunitinib for patients with advanced renal cell carcinoma (RCC), significant objective responses have been reported. However, a prognostic marker is needed for selecting patients who will benefit most from sunitinib. Previous studies have shown that an elevated C-reactive protein (CRP) level predicts poor survival in patients with metastatic RCC treated with immunotherapy. In this study, we focus on non-tumor characteristics, including CRP, that can predict sunitinib effectiveness.

**Materials and Methods:** Between December 2008 and August 2011, 41 consecutive patients with advanced clear-cell RCC treated with sunitinib were enrolled in this study. Non-tumor variables were selected from pre-treatment characteristics and adverse events that occurred during the study. Logistic regression analysis estimated the relative importance of non-tumor variables, including CRP, and selected adverse events as predictive factors for sunitinib responses.

**Results:** Overall, 11 patients (26.8%) demonstrated a partial response and 10 patients (24.4%) had stable disease. On univariate analyses, Memorial Sloan-Kettering Cancer Center non-poor risk, normal CRP, hand-foot skin reaction, altered taste, fatigue, and leukopenia were significantly correlated with objective responses ( $P = 0.0206, 0.0011, 0.0069, 0.0064, 0.0238$ , and  $0.0377$ , respectively). On multivariate analysis, normal CRP was independently associated with objective response ( $P = 0.0163$ ). Based on this result, patients were grouped into two cohorts: those with normal CRP levels ( $\leq 0.30$  mg/dL) and those with elevated CRP levels ( $> 0.30$  mg/dL). Patients with a normal level of CRP had a significantly higher objective response rate (84.6% vs 35.7%,  $P = 0.0012$ ) and significantly longer progression-free survival time (median 19.0 vs 6.0 months,  $P = 0.0361$ ) than patients with an elevated level of CRP.

**Conclusions:** CRP is a significant independent prognostic indicator for patients with advanced RCC treated with sunitinib. Pre-treatment CRP level could be a useful biomarker for response to sunitinib treatment. Further study is needed to clarify the prognostic role of CRP with not only sunitinib but also other targeted agents.