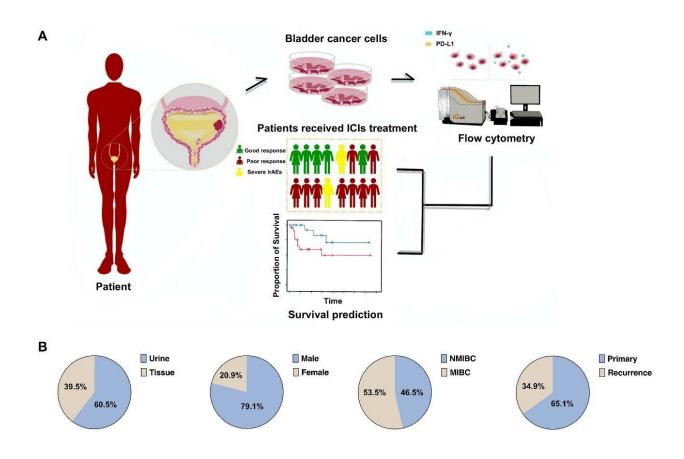


Novel urine-based test offers new hope for bladder cancer patients

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I Overview of establishment of BC-PD-L1. (A) Overview of the models of BCCs for potential applications in prediction of prognosis in bladder cancer. Urine/tumor samples were collected from BLCA patients for primary bladder cancer cell culturing. PD-L1 expression levels of BCCs before and after IFN- γ stimulation were measured by FCM and used for prognosis and ICI response prediction. (B) Pie charts showing the classification of all urine/tumor-provided patients based on sample source, gender, pathology group, and disease status. (C) Levels of PD-L1 expression on BCC cell surface before and after IFN- γ



treatment measured by FCM analysis. (MFI mean ± standard deviation). MFI, median fluorescence intensity. Credit: Jiaqi Wang, et al.

Bladder cancer, notorious for being one of the most costly malignancies to manage, often results in patient discomfort and frequent invasive and expensive procedures. Traditional treatments, though gradually being replaced by immune checkpoint inhibitors (ICIs), have yielded varied success rates with limited predictors for individual patient responses.

In a <u>new study</u> published in the journal *Genes & Diseases*, researchers from Fudan University, have developed a novel urine-based prognostic model that promises to transform the management and treatment of bladder cancer.

The research team collected urine and tumor samples from 43 bladder cancer patients, examining the expression levels of PD-L1—a key marker in <u>immune response</u>—on bladder cancer cells (BCCs). The study also focused on the variability of PD-L1 expression and its response to IFN- γ in bladder cancer cells (BCCs) as detected by the BC-PD-L1 platform.

From the 43 bladder cancer (BLCA) patients, urine and tumor samples were obtained and used for primary cell culturing. Intriguingly, PD-L1 expression, before and after treatment with IFN-γ, varied significantly among the BCCs. Moreover, the degree of PD-L1 upregulation post-IFN-γ treatment showed a negative correlation with baseline PD-L1 expression levels.

To ensure BC-PD-L1's reliability, the study employed RNA sequencing and correlated PD-L1 protein levels with mRNA levels, which validated its ability to detect PD-L1 regardless of its heavy glycosylation—a trait



that can potentially obstruct accurate detection. In evaluating the platform's clinical potential, the study delved into how BC-PD-L1, especially urine-derived BC-PD-L1 (UBC-PD-L1), can predict BLCA patient outcomes.

Although prior research on the prognostic significance of PD-L1 in bladder cancer has been mixed, this study's analysis revealed that higher surface PD-L1 levels were associated with longer disease-free survival and overall survival rates. Intriguingly, RNA sequencing highlighted differences between high and low PD-L1 expressing BCCs.

High PD-L1 BCCs showed upregulation of pathways related to the extracellular matrix (ECM)—a key component influencing cancer progression. Additionally, these high PD-L1 BCCs were more immuneactive, suggesting a better prognosis. This finding underscores the importance of PD-L1 as a potential biomarker and the BC-PD-L1's potential role in personalizing bladder cancer treatment strategies.

The study underscores the power of personalized medicine, highlighting the potential of urine-based tests in transforming <u>bladder</u> cancer treatments. Given <u>bladder cancer</u>'s propensity for recurrence and the invasive nature of current diagnostic and treatment methods, this innovation offers a beacon of hope.

More information: Jiaqi Wang et al, A novel in vitro prognostic model of bladder cancer based on urine-derived living tumor cells, *Genes & Diseases* (2023). DOI: 10.1016/j.gendis.2022.10.022

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