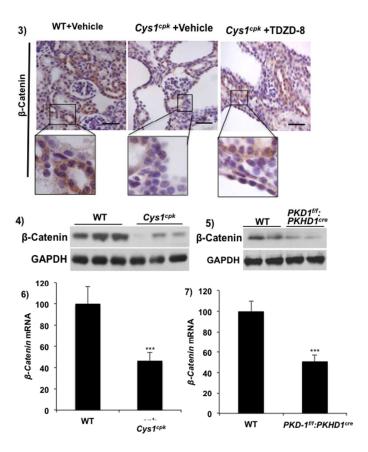


**Supplemental-1:** Bodyweight of WT and  $Cys1^{cpk}$  mice treated with vehicle or TDZD-8 from P3 to P14 showed no significant change.

Supplemental-2: Renal GSK3 $\beta$  expression (red) in proximal tubules (green) of  $Cys^{1cpk}$  + $GSK3\beta^{cD-KO}$  and  $PKD1^{lff}:GSK3\beta^{lff}:PKHD1^{cre}$  mice in which  $GSK3\beta$  is knocked out in the renal collecting duct. (Scale bar=50 $\mu$ m).

60x81mm (300 x 300 DPI)



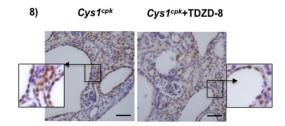
**Supplemental 3:** β-Catenin immunostaining (brown) was reduced in  $Cys1^{cpk}$ +vehicle compared to WT+vehicle kidneys. In the  $Cys1^{cpk}$ +TDZD-8 kidneys, more cytoplasmic staining for β-Catenin was observed. (Scale bar=25μm)

Supplemental 4:  $\beta$ -catenin protein levels at P14 are lower in the  $Cys1^{cpk}$  kidneys than in WT kidneys. Samples same as used in Fig-1a.

**Supplemental 5:** β-catenin protein levels at P21 are lower in the  $PKD1^{t/t}:PKHD1^{cre}$  kidneys than in WT kidneys. Samples same as used in Fig-1a.

**Supplemental 6:** β-catenin mRNA levels in WT and  $Cys1^{cpk}$  mice. (\*\*\*P<0.001, n=7/ group). Samples same as used in Fig-1d.

Supplemental 7: β-catenin mRNA levels in WT and PKD1<sup>th</sup>:PKHD1<sup>cre</sup> mice (\*\*\*P<0.001, n= 7/ group). Samples same as used in Fig-1e.



 $\textbf{Supplemental-8: } \textit{Cys1}^\textit{cpk} \text{ mice treated with vehicle or TDZD-8 from P3 to P14 showed increased immunostaining for Cyclin D1 in the cyst lining epithelium. (Scale bar=25µm).}$ 

60x81mm (300 x 300 DPI)