Fig. 1A

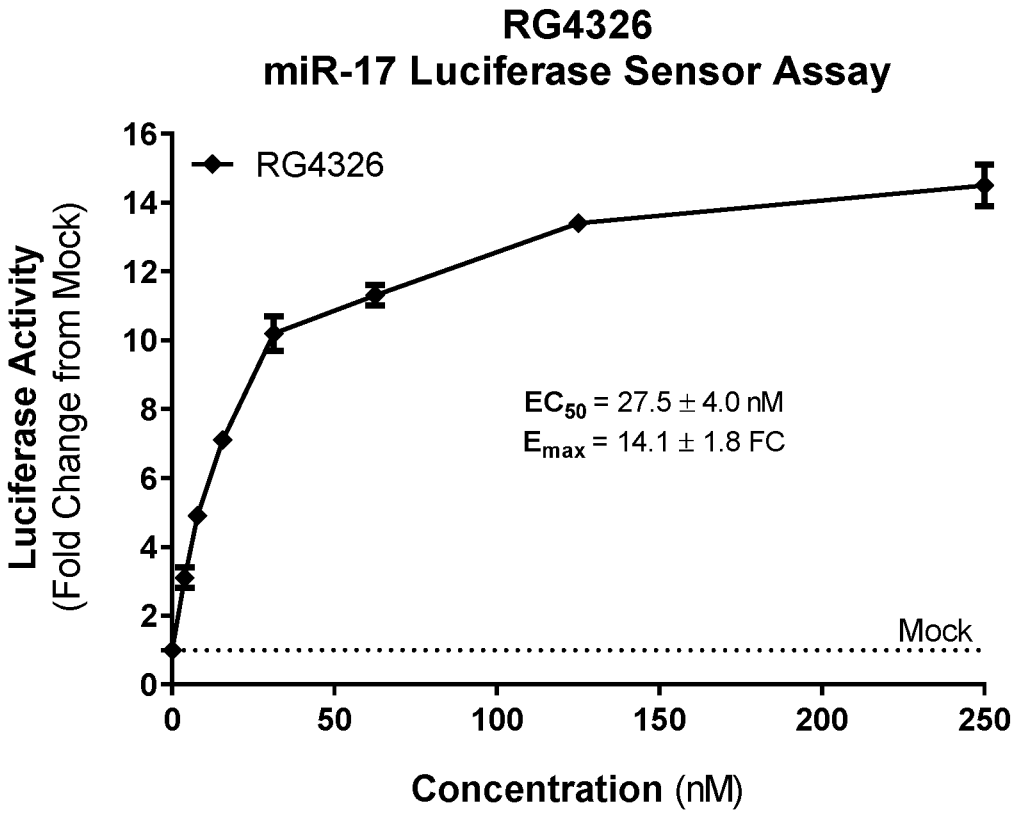

RG4326
miR-17 Luciferase Sensor Assay
RG4326
Luciferase Activity
(Fold Change from Mock)
16
14
12
10
8
6
4
2
0
EC50 = 27.5 ± 4.0 nM
Emax = 14.1 ± 1.8 FC
Mock
0
50
100
150
200
250
Concentration (nM)

Fig. 1B

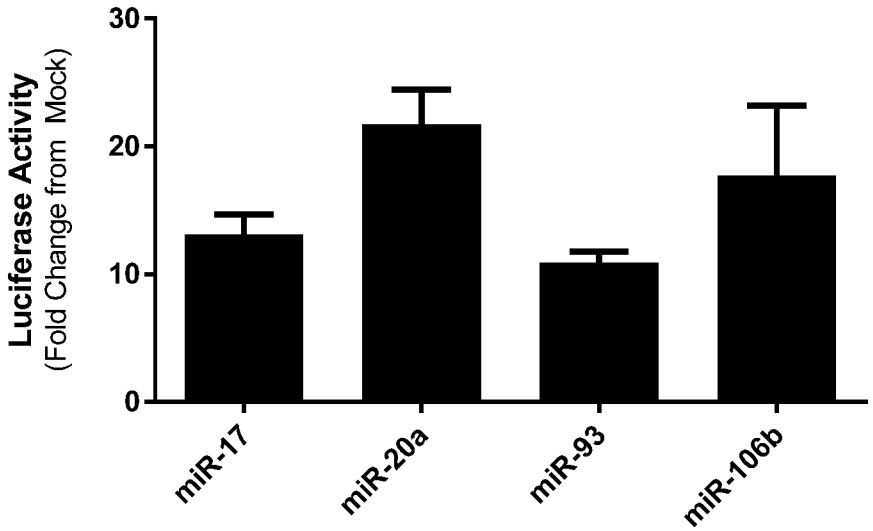

Luciferase Activity
(Fold Change from Mock)
30
20
10
0
miR-17
miR-20a
miR-93
miR-106b

Fig. 2

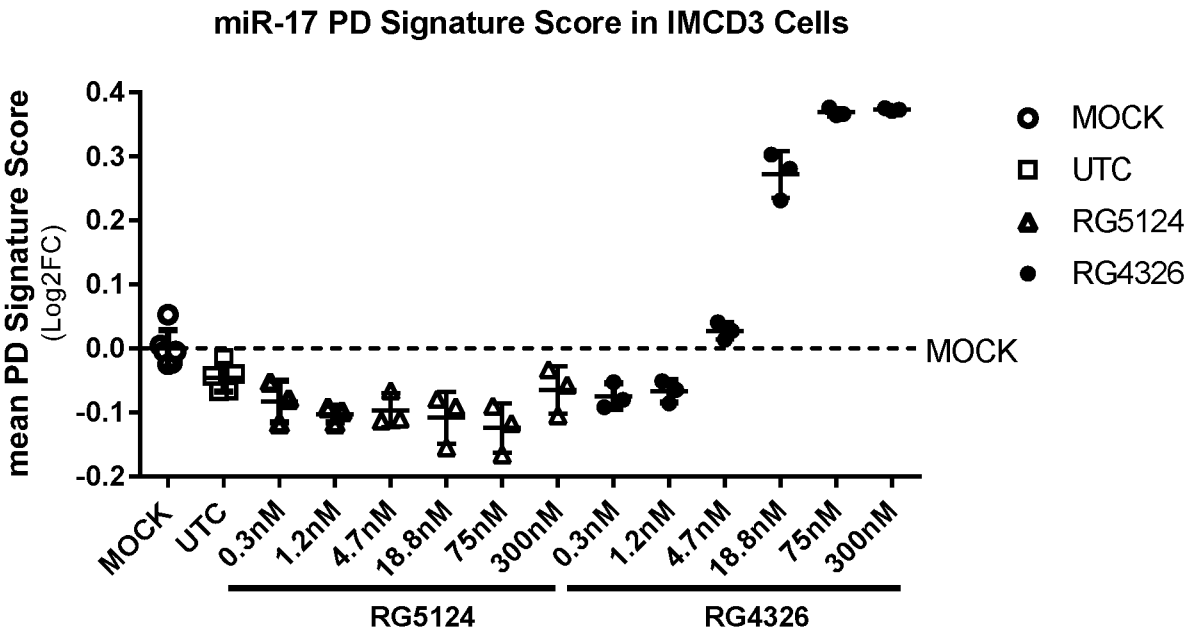

miR-17 PD Signature Score in IMCD3 Cells
mean PD Signature Score
(Log2FC)
0.4
0.3
0.2
0.1
0.0
-0.1
-0.2
MOCK
MOCK
UTC
RG5124
RG4326
MOCK
UTC
0.3nM
1.2nM
4.7nM
18.8nM
75nM
300nM
0.3nM
1.2nM
4.7nM
18.8nM
75nM
300nM
RG5124
RG4326

Fig. 3A

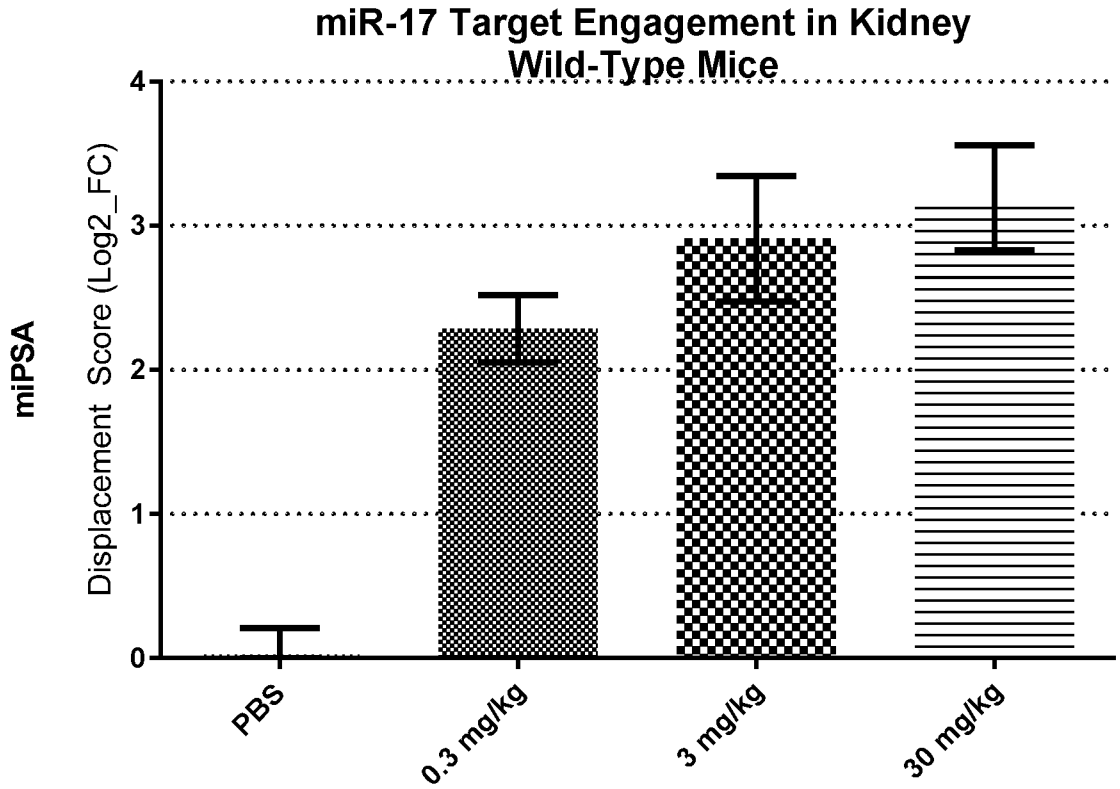

miR-17 Target Engagement in Kidney
Wild-Type Mice
miPSA
Displacement Score (Log2_FC)
0
1
2
3
4
PBS
0.3 mg/kg
3 mg/kg
30 mg/kg

Fig. 3B

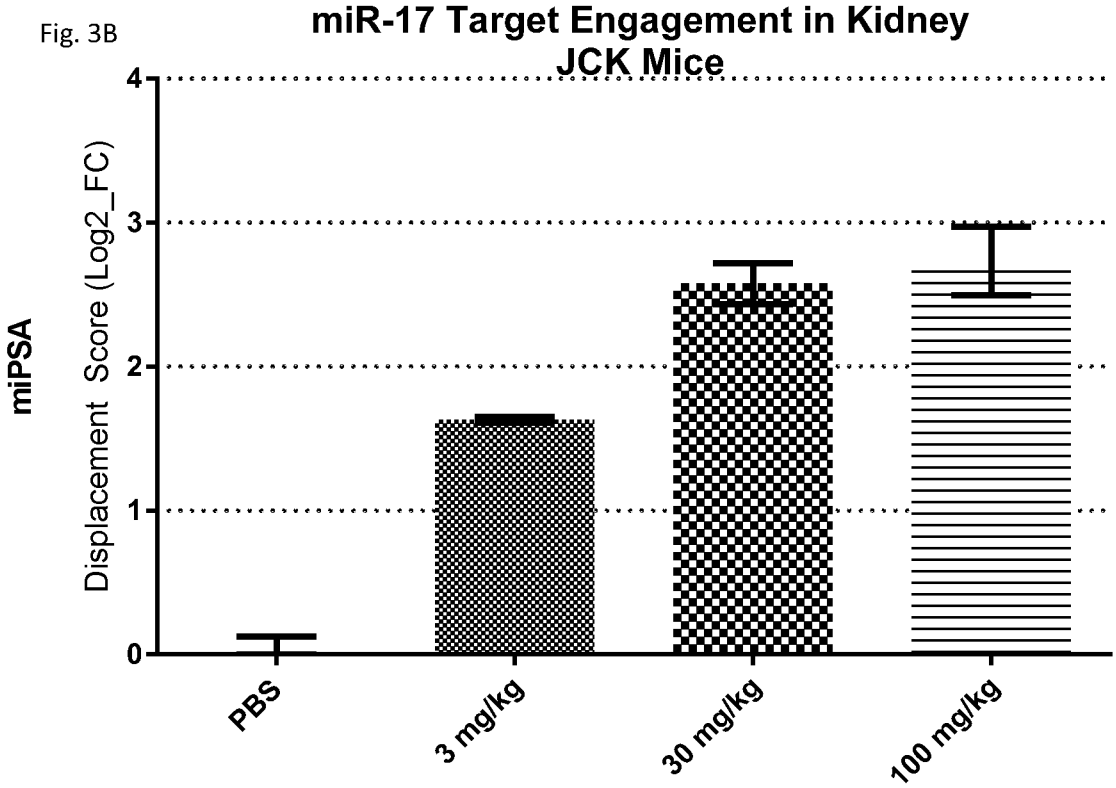

miR-17 Target Engagement in Kidney
JCK Mice
miPSA
Displacement Score (Log2_FC)
0
1
2
3
4
PBS
3 mg/kg
30 mg/kg
100 mg/kg

Fig. 4A

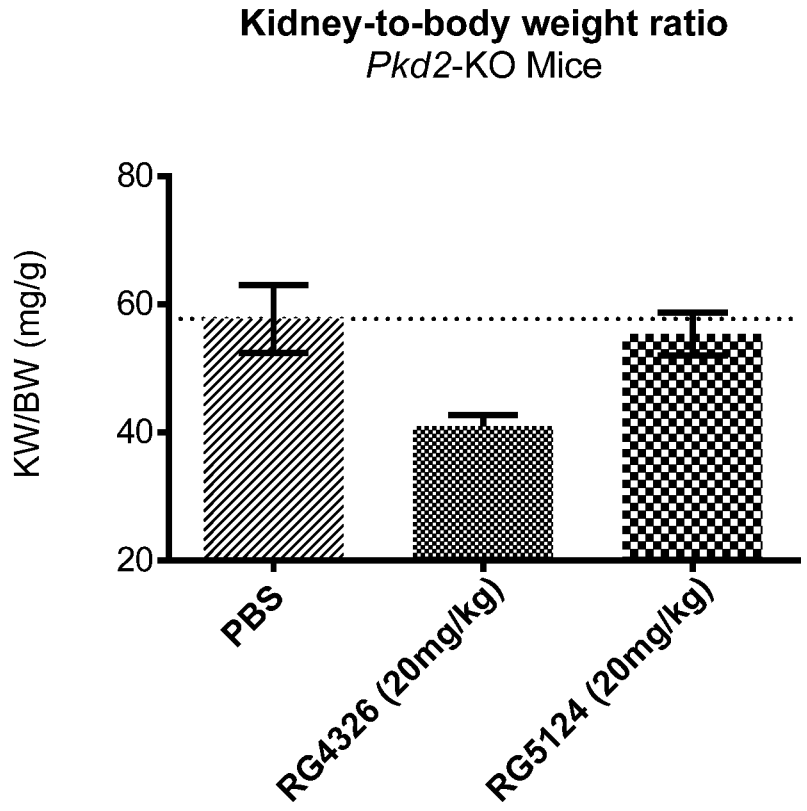

Kidney-to-body weight ratio
Pkd2-KO Mice
KW/BW (mg/g)
80
60
40
20
PBS
RG4326 (20mg/kg)
RG5124 (20mg/kg)

Fig. 4B

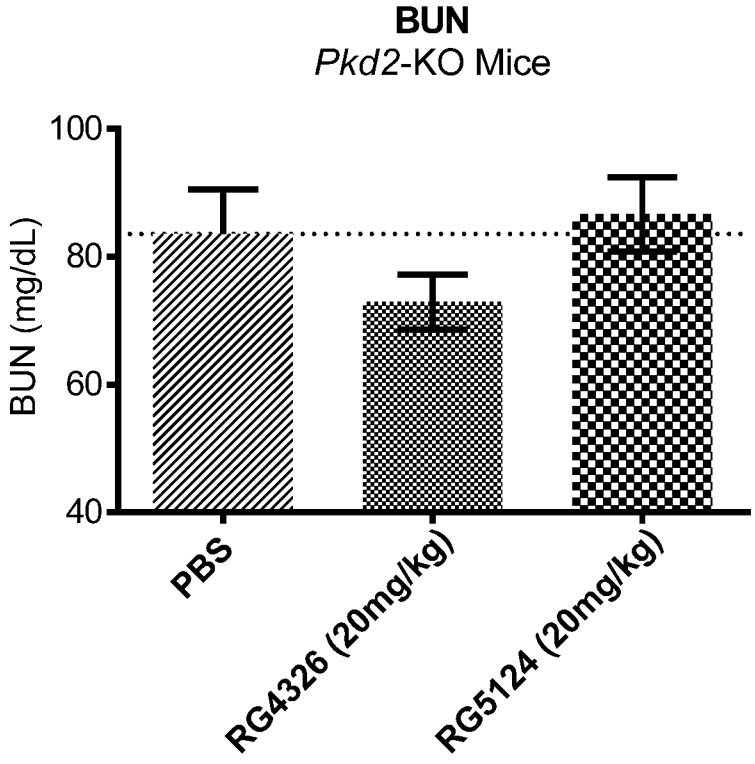

BUN
Pkd2-KO Mice
BUN (mg/dL)
100
80
60
40
PBS
RG4326 (20mg/kg)
RG5124 (20mg/kg)

Fig. 4C

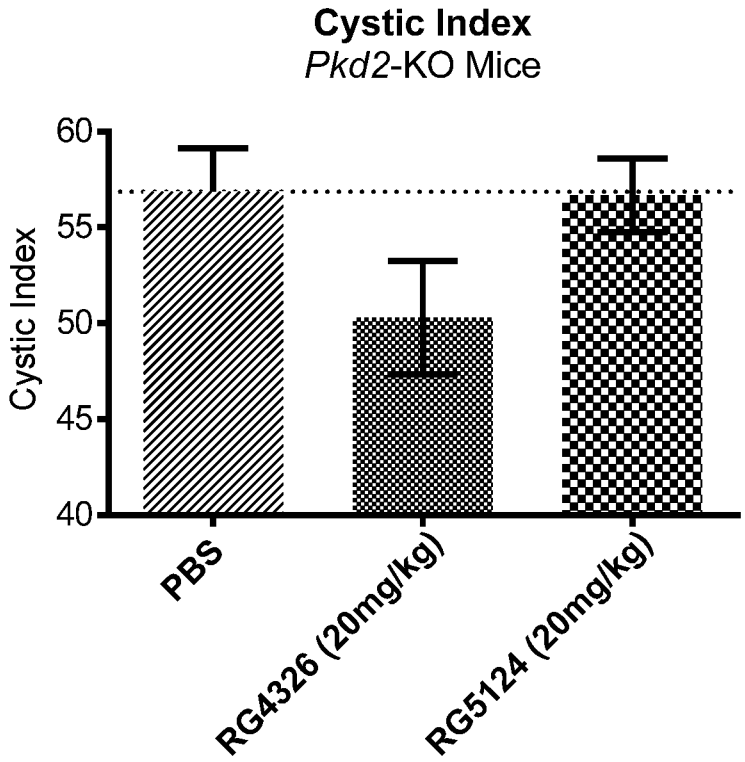

Cystic Index
Pkd2-KO Mice
Cystic Index
60
55
50
45
40
PBS
RG4326 (20mg/kg)
RG5124 (20mg/kg)

Fig. 5A

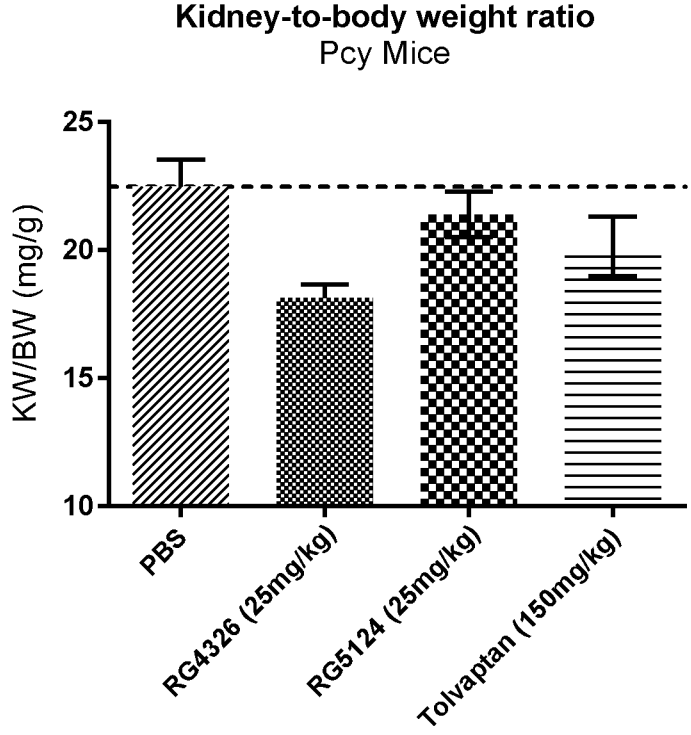

Kidney-to-body weight ratio
Pcy Mice
KW/BW (mg/g)
25
20
15
10
PBS
RG4326 (25mg/kg)
RG5124 (25mg/kg)
Tolvaptan (150mg/kg)

Fig. 5B

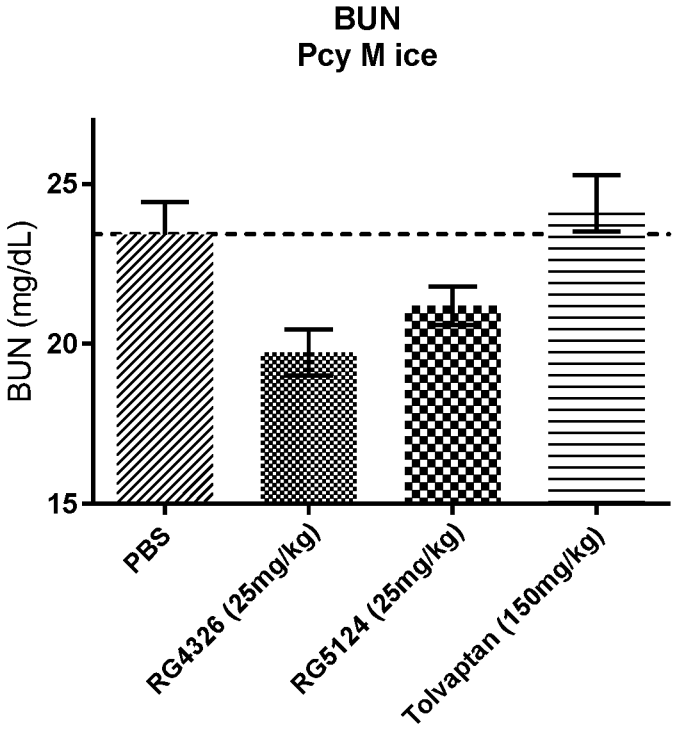
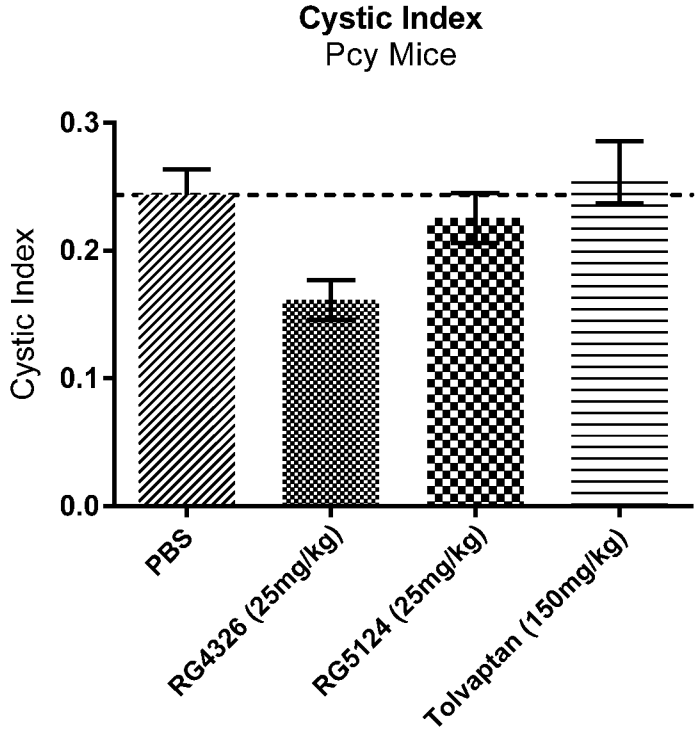

BUN
Pcy M ice
BUN (mg/dL)
25
20
15
PBS
RG4326 (25mg/kg)
RG5124 (25mg/kg)
Tolvaptan (150mg/kg)

Fig. 5C


Cystic Index
Pcy Mice
Cystic Index
0.3
0.2
0.1
0.0
PBS
RG4326 (25mg/kg)
RG5124 (25mg/kg)
Tolvaptan (150mg/kg)