

## **SIGNIFICANCE STATEMENT**

Adiponectin is an adipokine secreted by adipocytes known to exert favorable effects in the milieu of diabetic and metabolic syndrome through its anti-inflammatory, antifibrotic, and antioxidant effects. In human and animal diabetic nephropathy, the expression of adiponectin receptors is reduced. This manuscript reveals a favorable effect of AdipoRon, an orally active synthetic adiponectin receptor agonist, in protection, even reversal, of diabetic nephropathy produced by the activation of adiponectin receptors and downstream targets through increased intracellular  $\text{Ca}^{++}$ /AMPK-LKB1/PPAR $\alpha$  pathway. AdipoRon may be a promising drug for restoration of diabetic nephropathy in type 2 diabetes.