Supplementary Materials

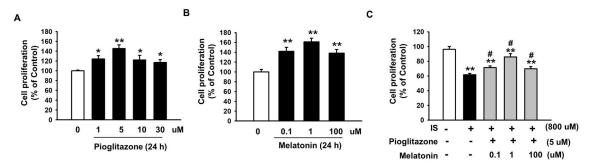


Figure S1. The effect of pioglitazone and melatonin on MSC proliferation. (**A**) After treatment of MSCs with pioglitazone (0, 1, 5, 10, and 30 μ M) for 24 h, cell proliferation was assessed. The values represent the mean \pm SEM. * p < 0.05 and ** p < 0.01 vs. non-treatment; (**B**) After treatment of MSCs with melatonin (0, 0.1, 1, 100 μ M) for 24 h, cell proliferation was assessed. The values represent the mean \pm SEM. ** p < 0.01 vs. non-treatment; (**C**) After exposure to indoxyl sulfate (IS; 800 μ M), cell proliferation was assessed after pretreatment with pioglitazone (5 μ M) and melatonin (0, 0.1, 1, 100 μ M) for 24 h. The values represent the mean \pm SEM. ** p < 0.01 vs. non-treatment, * p < 0.05 vs. MSCs treated IS.

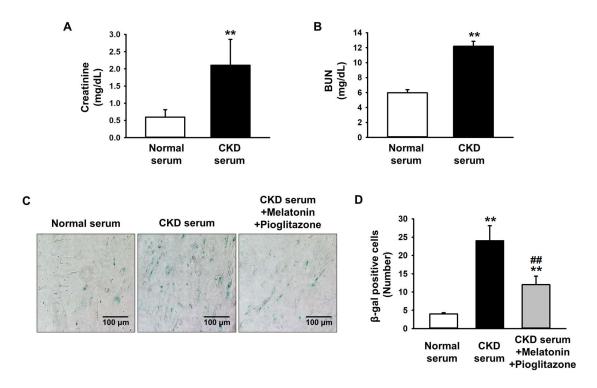


Figure S2. Effect of serum isolated from chronic kidney disease (CKD) mouse model on MSC senescence. (**A**) The concentration of creatinine in serum isolated from normal (normal serum) and CDK (CKD serum) mice. The values represent the mean \pm SEM. ** p < 0.01 vs. normal serum; (**B**) The concentration of blood urea nitrogen (BUN) in normal and CDK serum. The values represent the mean \pm SEM. ** p < 0.01 vs. normal serum; (**C**) Images of senescence-associated beta-galactosidase (SA-β-gal) staining in MSCs pretreated with melatonin and pioglitazone after exposure to CKD serum; (**D**) The number of SA-β-gal positive cells. The values represent the mean \pm SEM. ** p < 0.01 vs. normal serum, ** p < 0.01 vs. CKD serum.

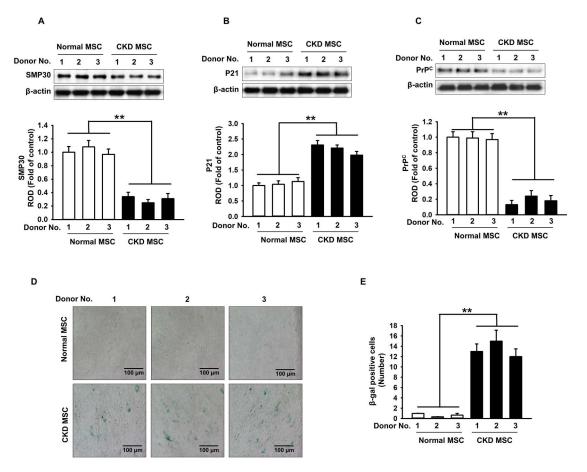


Figure S3. Assessment of senescence in bone marrow-derived MSCs isolated from CKD mice. (**A–C**) The expressions of SMP30 (**A**), P21 (**B**), and PrP^C (**C**) in bone marrow-derived MSCs isolated from normal (normal MSC; n = 3) and CKD (CKD MSC; n = 3) mice. The values represent the mean \pm SEM. ** p < 0.01 vs. Normal MSC; (**D**) Images of senescence-associated beta-galactosidase (SA-β-gal) staining in normal (n = 3) and CKD (n = 3) MSCs; (**E**) The number of SA-β-gal positive cells. The values represent the mean \pm SEM. ** p < 0.01 vs. normal MSCs.