Supplementary Figure.1. *KD-mTOR transgene is specifically express in \beta-cell but not in glucagon-positive cells.* (A) Immunofluorescence staining of insulin (blue), EGFP (green) and phosphorylated S6 (Ser 240, red) in RIPCre;KD-mTOR (Homo) mice and control neonates (postnatal day 1). (B) Immunofluorescence staining of glucagon (red), EGFP (green) in RIPCre;KD-mTOR (Homo) mice and control neonates. (C) Approximately 80% of insulin-positive cells express EGFP in RIPCre;KD-mTOR (Homo) mice. Image magnification in A is 40X and B is 20X.



° ₀⊥**u** RIPCre; KD-mTOR (Homo)

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Supplementary Figure.2. *KD-mTOR transgene islet morphology in RIPCre;KD-mTOR mice.* (A) Immunofluorescence staining of insulin (blue), phosphorylated S6 (red), and GFP (green) in adult male RIPCre;KD-mTOR mice (Het). (B) Immunofluorescence staining of GFP (green), insulin (blue) and phosphorylated S6 (red) in male RIPCre;KD-mTOR (Homo) mice and control. Image magnification is 20X.





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Supplementary Figure.3. *Normal \beta-cell mass in RIPCre;Raptor^{fl/fl} and RIPCre;Rictor^{fl/fl} neonates.* (A-F) Body weight, glucose level and in newborn β -cell area in RIPCre;Raptor^{fl/fl} neonates (A-C) or RIPCre;Rictor^{fl/fl} (D-F) neonates compare to controls, n=4-12. Statistical analyses were conducted using unpaired, non-parametric Mann-Whitney test (u test).



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Supplementary Figure.4. Fasting insulin and proliferation rate in RIPCre;KD-mTOR mice in high-fat diet.(A) Fasting insulin levels 6 weeks after high-fat diet among genotypes, n=10-12. No significant changes in proliferation rate (measured by Ki67 in insulin-positive cells) and ~50 islets per animal for Ki67 analysis, n=4=6. Statistical analyses were conducted using unpaired, non-parametric Mann-Whitney test (*u*-test).

