

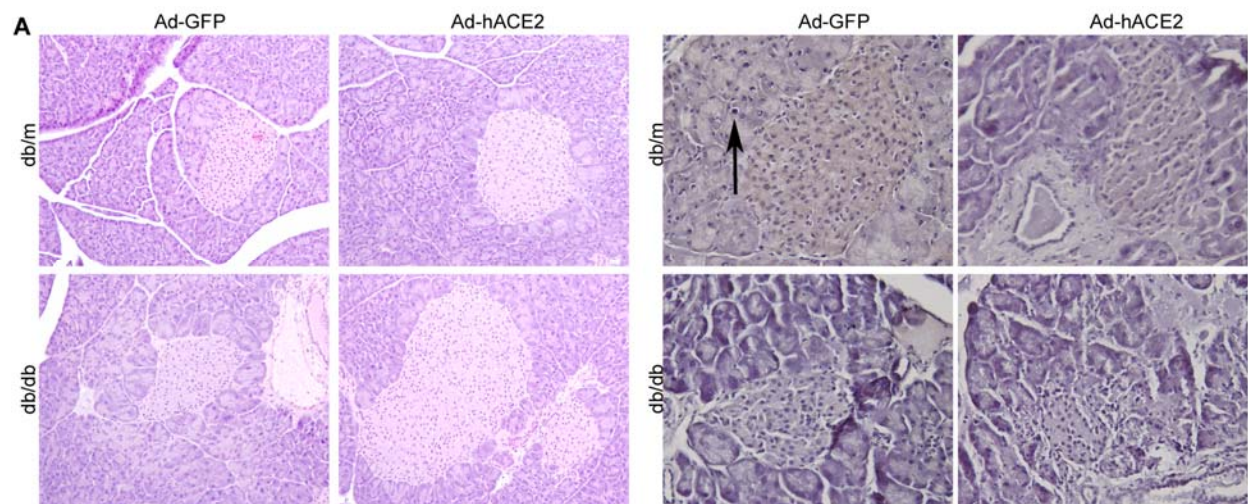
ONLINE APPENDIX

Supplemental Table 1. Metabolic parameters in Ad-hACE2 and Ad-GFP infected db/db and db/m mice.

	Ad-GFP db/m	Ad-GFP db/db	Ad-hACE2-eGFP db/m	Ad-hACE2-eGFP db/db
Body Weight (g)	18.18 ± 0.56	24.70 ± 1.14*	18.82 ± 0.57	23.2 ± 1.42*
Fasting Blood Glucose (mg/dl)	85.75 ± 6.5	123.8 ± 13.2	126.00 ± 10.3 [†]	144.4 ± 13.38
IPGT AUC (mg/dl/min)	12.68 ± 0.4	18.48 ± 1.8*	13.35 ± 0.5	18.05 ± 1.2*
Insulin sensitivity AUC (mg/dl/min)	8.05 ± 0.9	10.9 ± 2.5	5.3 ± 0.1	9.06 ± 0.8

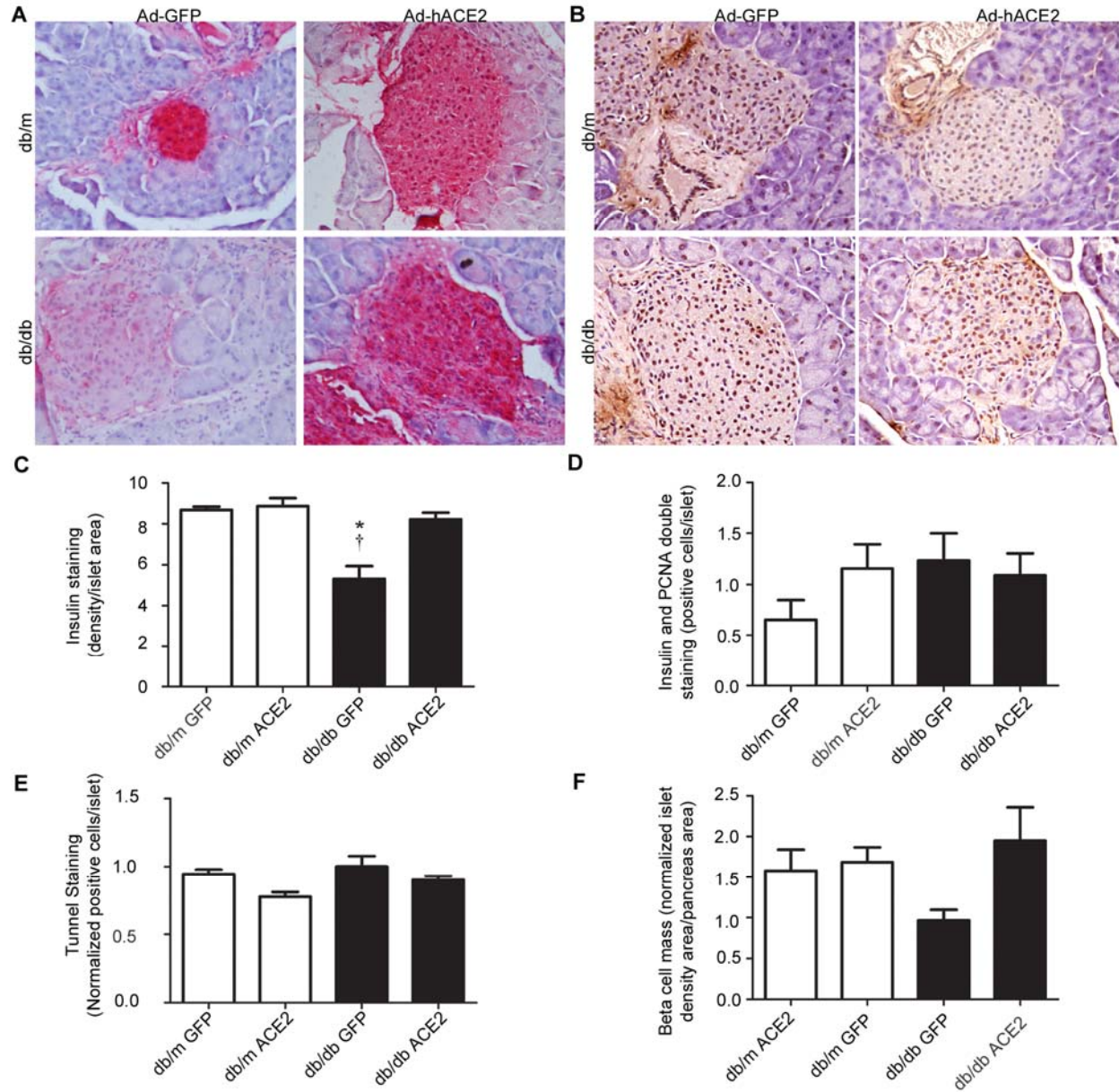
Data represent metabolic parameters of 4 week-old db/m and db/db mice (n=6 per group) following infection with Ad-hACE2-eGFP or Ad-eGFP viruses. Statistical significance: * $P < 0.001$ vs. treatment- matched db/m, [†] $P < 0.05$ vs. GFP treated controls. Abbreviation: IPGT AUC, intra-peritoneal glucose tolerance area under the curve.

Supplemental Figure 1



Adenoviral delivery via direct injection does not cause pancreatic inflammation. **A:** H&E staining revealed preservation of pancreatic architecture following direct viral injection. **B:** Immunostaining for CD3⁺ revealed only occasional lymphocytes present in pancreatic tissue.

Supplemental Figure 2



ACE2 overexpression enhanced islet insulin content in db/db mice in comparison to Ad-eGFP-expressing db/db mice (A,C). There was no significant change in pancreatic β -cell proliferation (D) apoptosis (C,E) or pancreatic β -cell mass (F). Values expressed as mean \pm SEM.