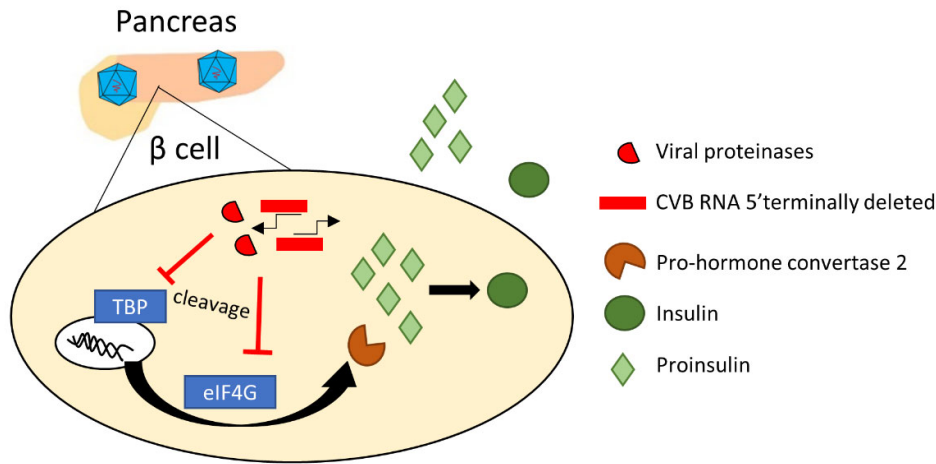
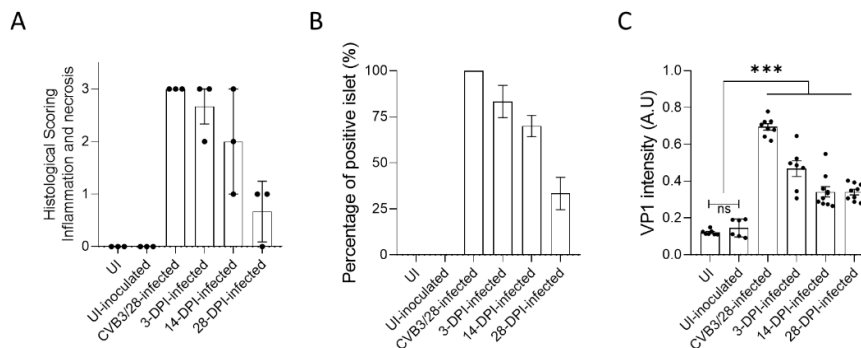


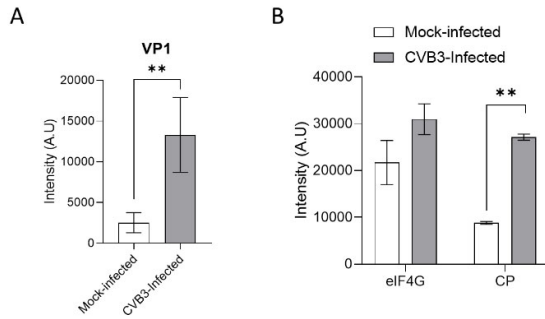
## Supplementary material



**Figure S1: Schematic representation of the scientific hypothesis.** Replication activities of CVB-TD RNA forms produced low levels of viral proteins, sufficient to cleave host cells factors such as TBP or eIF4G. These viral proteinase cleavage activities could result in a decrease in PCSK2 enzymatic activity, with the accumulation of proinsulin and a decrease in insulin secretion.



**Figure S2: Histological scoring and Viral Protein 1 positivity in pancreas of DBA/2J mice at 7 days post infection.** (A) Histological scoring for inflammation and necrosis in HES stained slices of pancreas, from uninfected or infected DBA/2J mice, or inoculated with homogenized pancreas from 3-DPI, 14-DPI or 28-DPI and uninfected mice (n=3). (B) Percentages of islets positive for VP1 in immunohistochemistry (n=3). (C) VP1 intensity on slices of pancreas (n=3, with two to three ROI per slice). Data presented as mean +/- SEM. Mann-Whitney test: \*\*\*= p<0.001. ns: non-significant. ROI: Region of interest. VP1: viral protein 1. UI: uninfected. CVB3: coxsackievirus B3. AU: arbitrary unit. DPI: days post infection.



**Figure S3: Western blot intensity analyses.** (A) Viral protein 1 intensity after 24 hours of incubation (mock-infected) or CVB3/28 infection, at a Multiplicity of Infection of 1 (n=3). (B) eIF4G and its cleavage product intensities after 24 hours of incubation (mock-infected) or CVB3/28 infection, at a Multiplicity of Infection of 1 (n=2). Data presented as mean +/- SD. Mann-Whitney test: \*\*= p<0.01. VP1: viral protein 1. UI: uninfected. CVB3: Coxsackievirus B3. AU: arbitrary unit. CP: cleavage product. eIF4G: Eukaryotic translation initiation factor 4 G.