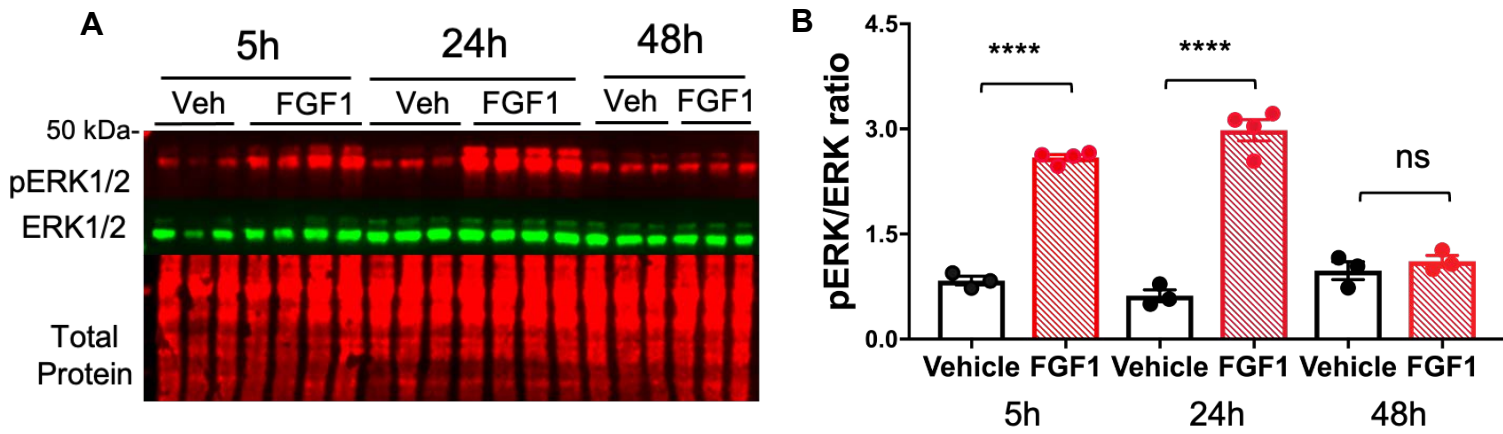


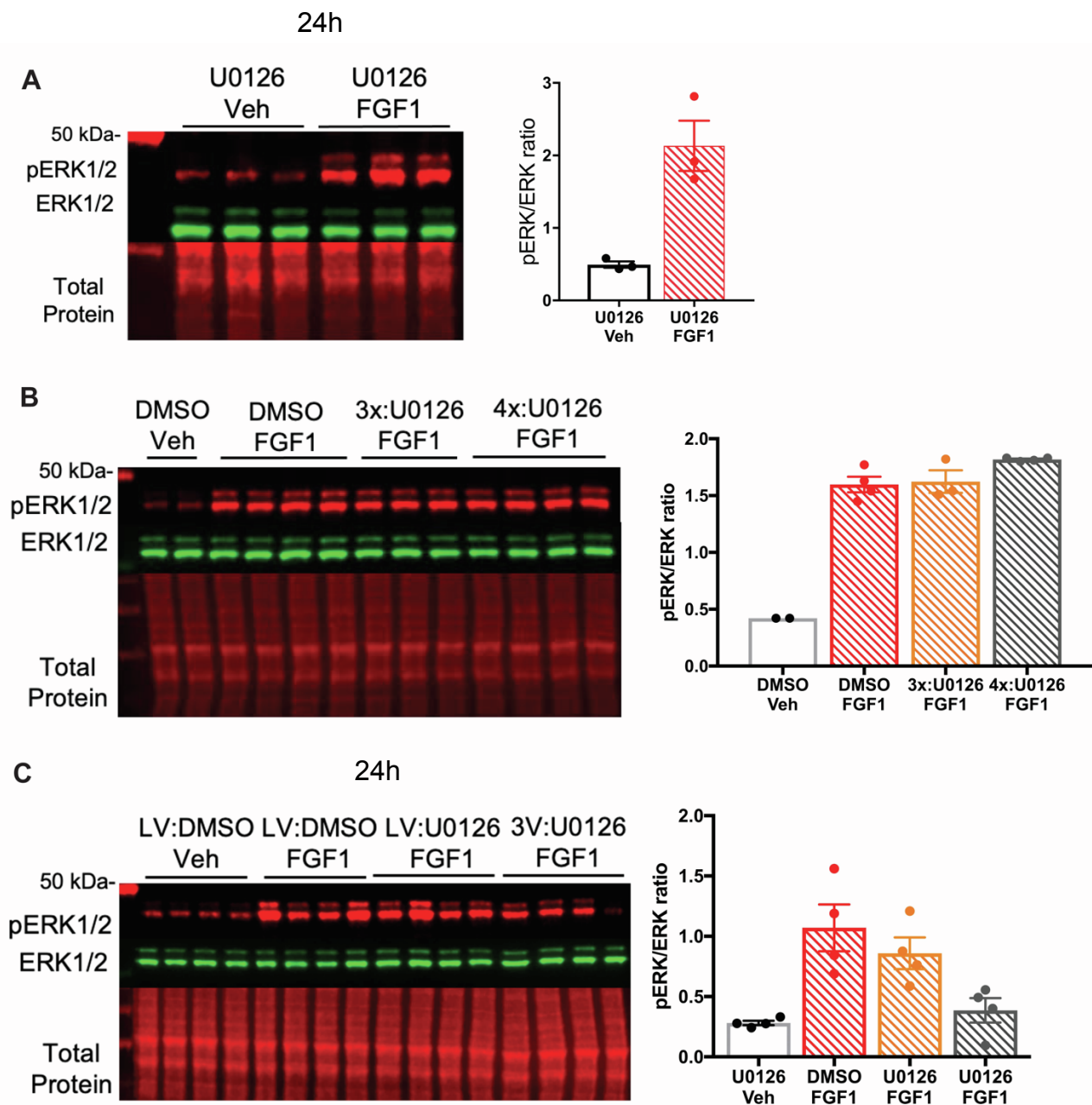
**Supplemental information**

**Role of hypothalamic MAPK/ERK signaling  
and central action of FGF1 in diabetes remission**

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**Supplementary Figure 1- Central Administration of FGF1 Induces Sustained ERK1/2 Signaling in the MBH of db/db mice. (Related to Figure 1)** A) Representative western blot (left panel) of phosphorylated (red) and total ERK1/2 (green) and total protein (red) from mediobasal hypothalamic punches from adult male db/db mice after a single icv injection of either vehicle or FGF1 (3 ug) and quantitative comparison at B) 5 hours (n=3-4/group,  $t=24.48$  df=5  $p=0.0001$ ), 24 hours (n=3-4/group,  $t=12.21$  df=5  $p=0.0001$ ), 48 hours (n=3/group,  $t=-0.0344$ , df=4,  $p=0.2121$ ) after injection. pERK1/2 ratio unpaired Welch two sample t-test (one sided) \*\*\*\* $P<0.0001$ .



**Supplementary Figure 2- Blockade of Prolonged MAPK/ERK signaling in the MBH induced by central FGF1. (Related to Figure 2)** **A**) Representative western blot (left panel) and quantitative comparison of phosphorylated (red) and total ERK1/2 (green) and total protein (red) (right panel) from mediobasal hypothalamic punches from adult male C57Bl6J mice 24 hours after a icv injection of an inhibitor of MAPK signaling U0126 (5  $\mu$ g) followed by vehicle or FGF1 into the lateral ventricle and **B**) 24 hours after 3 and 4 repeated injections of U0126 (5  $\mu$ g) 3h apart after a single icv injection of either vehicle or FGF1. **C**) a single icv injection of FGF1 into the lateral ventricle (LV) or 3rd ventricle (3V) followed by continuous infusion via osmotic pump of 30mM 1ul/hour U0126 or DMSO for 24 hours.