Expanded View Figures



Figure EV1. Synaptic activity and BDNF stabilize HIF-1a.

A–D Representative Western blots of cortical neurons stimulated with (A) Bic+4-AP at the indicated time points or the hypoxia-mimetic CoCl₂ (200 μ M) for 4 h and (B) BDNF (25 ng/ml) for the indicated time points (n = 4–5 independent experiments) and (C and D) densitometric analysis. Values represent mean \pm s.e.m. *P < 0.05, two-tailed Student's *t*-test.

E Neurons were stimulated with Bic+4-AP for the indicated times and HIF-2 α was analyzed by Western blot.

F Densitometric analysis (n = 4 independent experiments). Values represent mean \pm s.e.m.

Source data are available online for this figure.



Figure EV2. Glut3 and Siah2 expression is regulated by CREB.

- A Luciferase-based MEF2 activity in neurons expressing a control plasmid (globin) or the CREB inhibitor ICER and stimulated with Bic+4-AP for 8 h (n = 7 independent experiments). Values represent mean \pm s.e.m. *P < 0.05, one-way ANOVA followed by Tukey's *post hoc* test.
- B Lactate released into the medium by neurons stimulated with 10 μ M forskolin (for 24 h; n = 4 independent experiments). Values represent mean \pm s.e.m. *P < 0.05, two-tailed Student's t-test.
- C ¹⁴C-U-glucose incorporation into lipids in neurons treated with 10 μ M forskolin for 48 h (n = 4 independent experiments). Values represent mean \pm s.e.m.

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Figure EV3. HIF-1 α is highly expressed in the brain of 10-day-old mice.

A, B (A) Representative Western blot of HIF-1 α expression in protein extracts of the cortex of 10-day and 3-month-old mice and (B) densitometric analysis (n = 4 mice per condition). Values represent mean \pm s.e.m. *P < 0.05, two-tailed Student's t-test.

C Immunofluorescence detection of HIF-1 α in coronal sections from 10-day-old mice (n = 3 mice). Scale bar, 100 μ m.

Source data are available online for this figure.