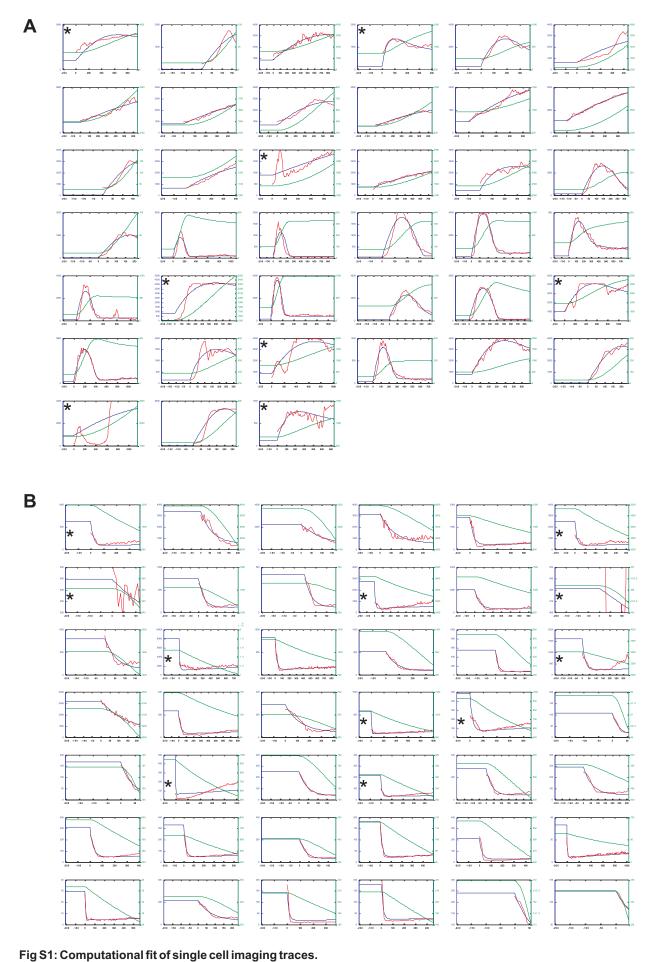
## Supplemental material S1



Single cells traces of HIF dynamics (blue line) that were fitted computationally using the model described in Fig 2 of the main manuscript  $(HIF, red \, line; PHD, green \, line). \, The \, ^* \, highlight \, the \, cells \, that \, could \, not \, be \, fitted \, appropriately \, with \, the \, 2 \, component \, model \, (see \, materiel \, and \, could \, not \, be \, fitted \, appropriately \, with \, the \, 2 \, component \, model \, (see \, materiel \, and \, could \, not \, be \, fitted \, appropriately \, with \, the \, 2 \, component \, model \, (see \, materiel \, and \, could \, not \, be \, fitted \, appropriately \, with \, the \, 2 \, component \, model \, (see \, materiel \, and \, could \, not \, be \, fitted \, appropriately \, with \, the \, 2 \, component \, model \, (see \, materiel \, and \, could \, not \, be \, fitted \, appropriately \, with \, the \, 2 \, component \, model \, (see \, materiel \, and \, could \, not \, be \, fitted \, appropriately \, with \, the \, 2 \, component \, model \, (see \, materiel \, and \, could \, not \, be \, fitted \, appropriately \, with \, the \, 2 \, component \, model \, (see \, materiel \, and \, could \, not \, be \, fitted \, appropriately \, with \, the \, 2 \, component \, model \, (see \, materiel \, and \, could \, not \, be \, fitted \, appropriately \, with \, the \, 2 \, component \, model \, (see \, materiel \, and \, could \, not \, be \, fitted \, appropriately \, with \, the \, 2 \, component \, model \, (see \, materiel \, and \, could \, not \, be \, fitted \, appropriately \, with \, could \, not \, be \, fitted \, appropriately \, approp$ methods section) A. Cells are initially at equilibrium in normoxia (h=1) and are de-oxygenated into hypoxia (h=0.14) at t=0. B. Cells are initially at equilibrium in hypoxia (h=0.14) and are re-oxygenated back into normoxia (h=1) at t=0.

## Supplemental movies

**movie 1:** HeLa cells were plated on a glass bottom dish and transfected with HIF-1 $\alpha$ -EGFP 24h hours before imaging . Cells were then placed onto the microscope stage and imaged every 5 min as indicated in the main manuscript. Cells were initially imaged at 20.8%O<sub>2</sub> for several frames before switching to 1%O<sub>2</sub> for 20h. The movie is accelerated to 30 frames/sec.

**movie 2:** HeLa cells were plated on a glass bottom dish and transfected with EGFP-HIF- $2\alpha$  24h hours before imaging . Cells were then placed onto the microscope stage and imaged every 5 min as indicated in the main manuscript. Cells were initially imaged at  $20.8\%O_2$  for several frames before switching to  $1\%O_2$  for 15h. The movie is accelerated to 30 frames/sec.