# **Supplemental Figure Legends**

### Supplemental Figure 1

Colony forming units (cfu)/g tissue of *Hif1a*<sup>IEC</sup> and *Hif1a*<sup>MC</sup> animals compared to their wildtype littermates 4 days post infection. Non normalized cfu counts in indicated organs of (**A**) *Hif1a*<sup>IEC</sup> (*n=7*) and (**B**) *Hif1a*<sup>MC</sup> (*n=6*) mice and wildtype littermates. Data represent means with SEM. Statistical significance was determined according to Mann-Whitney U test (A, B).

#### Supplemental Figure 2

Pathoscoring and macrophage infiltration in the healthy and *Salmonella* infected small intestine and colon of wildtypes and knock-outs utilized in this study.(A) Pathoscoring of small intestines and colons of healthy controls (Ctrl) and 4 days p.i. with *Salmonella* (*S*. Tm). Measured were edema/enlargement of the submucosa, neutrophil infiltration, goblet cell numbers as well as epithelial damage. Representative images of small intestines (**B**) and colons (**C**) 4 days post oral PBS administration (control) or *S*. Tm infection (p.i.) stained for F4/80 hematoxylin counterstaining of healthy control animals and 4 days p.i. (wildtype, *Hif1a*<sup>IEC</sup>, *Hif1a*<sup>MC</sup>.

#### Supplemental Figure 3

*Salmonella*-induced HIF-1α and HIF-2α stabilization in wildtype and knock-out IECs and BMDMs. (A) HIF-2α and (B) HIF-1α western blots of nuclear extracts of IECs of uninfected controls and infected (4 days p.i.) WT littermates and *Hif1a*<sup>IEC</sup> mice (*n*=3) and (C) WT and *Hif1a*<sup>IECind</sup> animals harboring a Tamoxifen-inducible HIF-1α knock-out in IECs (n=2). (D) *Hif1a* mRNA expression relative to β-*actin* upon *Salmonella* (*S*. Tm) infection in WT and *Hif1a*<sup>IECind</sup> IECs (*n*=3). (E) HIF-2α western blot of nuclear extracts of uninfected (*n=2*) and infected (4 hours; MOI 10; *n=3*) BMDMs isolated from WTs and *Hif1a*<sup>MC</sup> mice. Data represent means with SEM. <sup>\*</sup> P < 0.05 according to Mann-Whitney U test (D).

## Supplemental Figure 4

**FACS-based Leukocyte Quantification in lamina propria of uninfected WT littermates,** *Hif1a*<sup>IEC</sup> and *Hif1a*<sup>MC</sup> animals. (A) Gating strategy for DC and myeloid cell gating. Counts of Neutrophils, including P1 and P2 Neutrophils, Eosinophils and Macrophages in the small intestinal Lamina propria of (B) *Hif1a*<sup>IEC</sup> and (C) *Hif1a*<sup>MC</sup> mice and their WT littermates (*n*=3). Data represent means with SEM. \* P < 0.05; \*\* P < 0.01; \*\*\* P < 0.001 according to Mann-Whitney U test (B, C).

## Supplemental Figure 5

SPI-2 deficiency in *Salmonella* does not interfere with intracellular HIF-1-dependent bactericidal functions of Macrophages (A) Intracellular survival of *Salmonella* in bone marrow derived WT and HIF-1 $\alpha$  deficient (*Hif1a*<sup>MC</sup>) macrophages within 4 hours of infection utilizing wildtype *Salmonella*, a  $\Delta$ *sseB* mutant and the complemented strain  $\Delta$ *sseBpsseB* (*n=3*). Data represent means with SEM. \* P < 0.05; \*\* P < 0.01; \*\*\* P < 0.001 according to one-way analysis of variance followed by Tukey post hoc test (A, B).

# Hif1a<sup>IEC</sup>

Α



Hif1a<sup>™C</sup>





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Leukocytes in Lamina Propria of healthy WT and Hif1a<sup>IEC</sup> mice



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