Figure S1: WT = Ezh2 fl/fl  KO = LysM x Ezh2 fl/fl

A

|       | PECS |       | BMDM |       | Alveolar Macrophages |
|-------|------|-------|------|-------|----------------------|
| **Ezh2 (RQ)** | **Ezh2 (RQ)** | **Ezh2 (RQ)** |       |       |                      |
| WT    | **   | **    | **   | **    | **                   |
| KO    | **   | **    | **   | **    | **                   |

B

|       | TNF-α |        | Mcp-1 |        | Il-12p40 |       | Il-1b |        |
|-------|--------|--------|-------|--------|----------|-------|-------|--------|
| DMSO  | -      | -      | -     | -      | -        | -     | -     | -      |
| 10ng/ml 2 hours | - | + | - | + | - | + | - | + |
| 10ng/ml 6 hours | - | + | - | + | - | + | - | + |
| 100ng/ml 2 hours | - | - | - | + | - | + | - | + |
| 100ng/ml 6 hours | - | - | - | + | - | + | - | + |

*** *** ***
Figure S2

EZH2 inhibitor concentration (nM)

IL-6 (RQ)

- Vehicle
- GSK126
- GSK343

***
Figure S4. Influenza. WT = Ezh2 fl/fl  KO = CCSP-icre x Ezh2 fl/fl

A

- % initial body weight vs. Days

B

- Histology scoring

| Score  | WT | KO |
|--------|----|----|
| 1 = None | 0  | 0  |
| 2 = Mild   | 2  | 2  |
| 3 = Moderate | 2 | 1  |
| 4 = Severe  | 1  | 0  |
| 5 = Very severe | 0 | 2  |

- Total number of animals: 5
- Average score +/- SD: 2.8 +/- 0.83, 3.4 +/- 1.5

D

- Blood Neutrophils vs. Day 9
- BAL Neutrophils vs. Day 9
- BAL Macrophages vs. Day 9

E

- Blood Neutrophils vs. Day 21
- BAL Neutrophils vs. Day 21
- BAL Macrophages vs. Day 21