## Loss of selenoprotein W1 inhibits skeletal muscle growth and development in zebrafish (*Danio rerio*) by promoting protein ubiquitination degradation

Chenlei Liao<sup>1, 2</sup>, Li Wang<sup>1</sup>, Hongyun Li<sup>1</sup>, Yiwen Tong<sup>1</sup>, Danqiu Wei<sup>1</sup>, Hai Huang<sup>2</sup>, Xuezhen Zhang<sup>1, \*</sup>

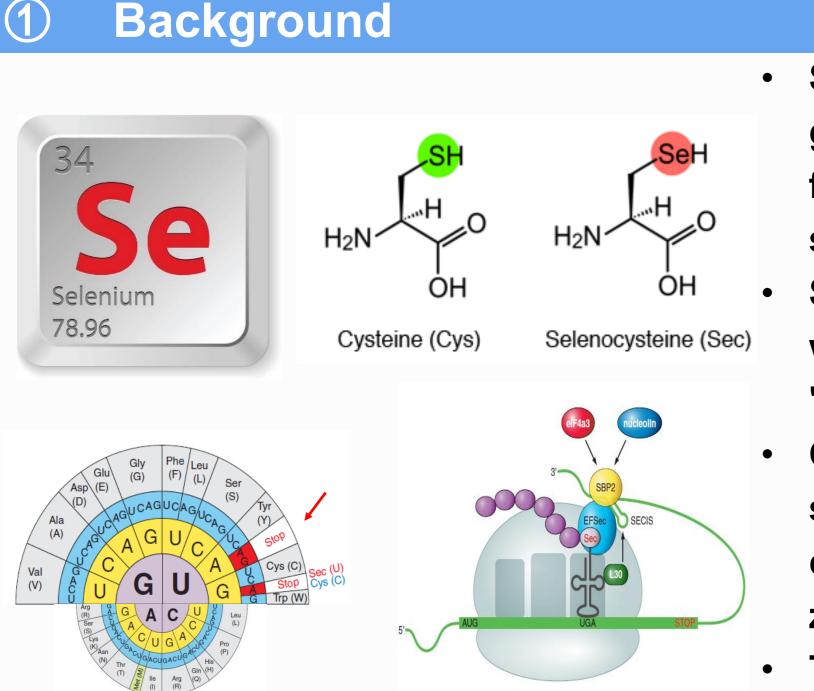
College of Fisheries, Huazhong Agricultural University, Hubei Hongshan Laboratory, Hubei Wuhan 430070, PR China

College of Fisheries and Life Sciences, Hainan Tropical Ocean University, Hainan Sanya 572022, PR China









Translation-of-selenoprotein-mRNA

- Selenium (Se) plays an essential role in the growth of fish and performs its physiological functions mainly through incorporation into selenoproteins.
- Sec is known as the 21<sup>st</sup> amino acid in protein, which is encoded by the termination codon "UGA".
- Our previous studies suggested that the selenoprotein W (selenow) is sensitive to changes in dietary Se in rainbow trout and zebrafish.
- The mechanism of *selenow1* gene in zebrafish muscle is unknown.

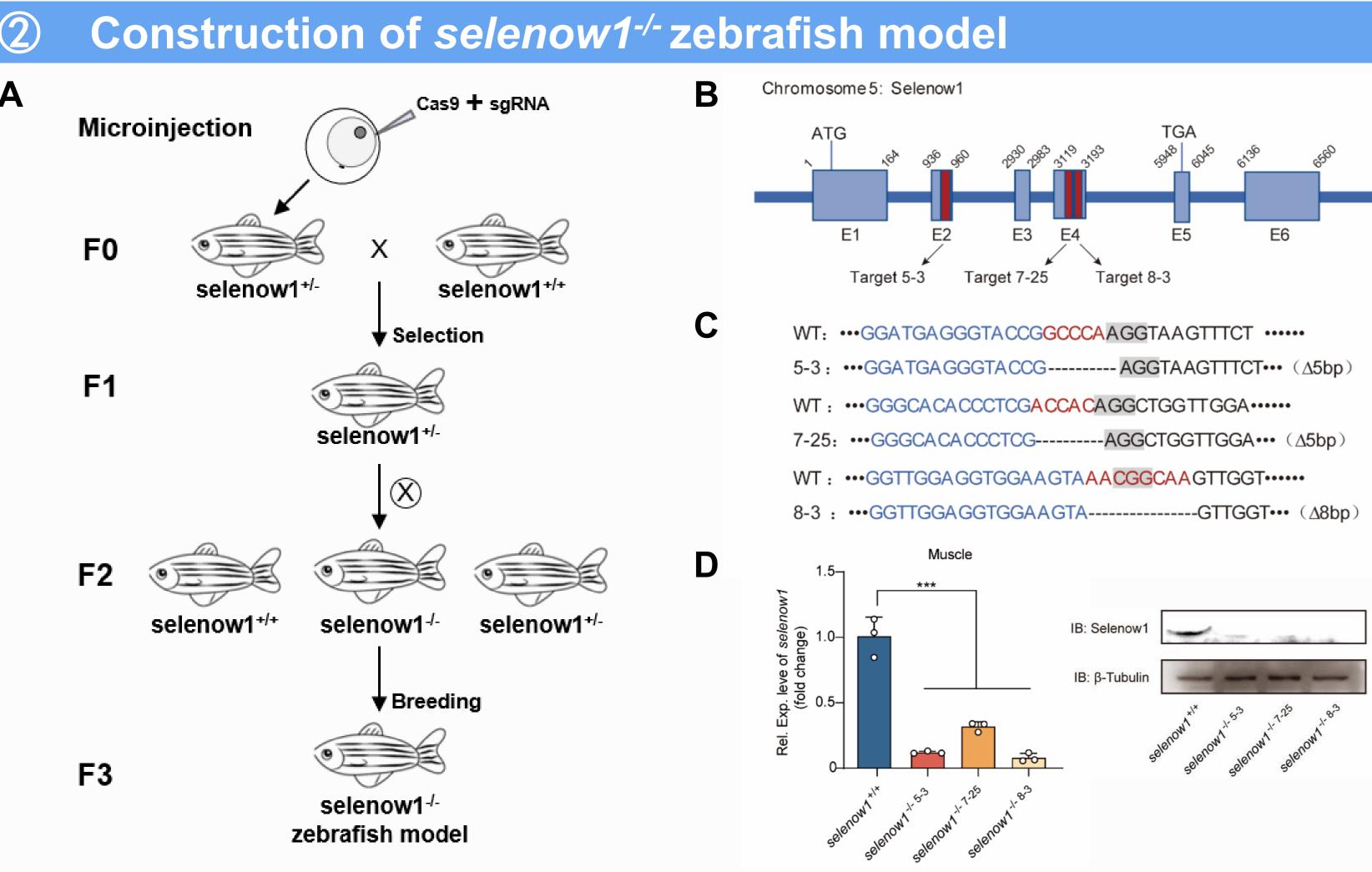


Fig. 1 A. Schematic diagram of *selenow1-/-* zebrafish model construction. B. Genomic structure of zebrafish *selenow1* and genotypes of the mutation. C. Sequencing results of the targeted region from the WT and three kinds of homozygous zebrafish. D. The mRNA and protein level of *selenow1-/-* zebrafish.

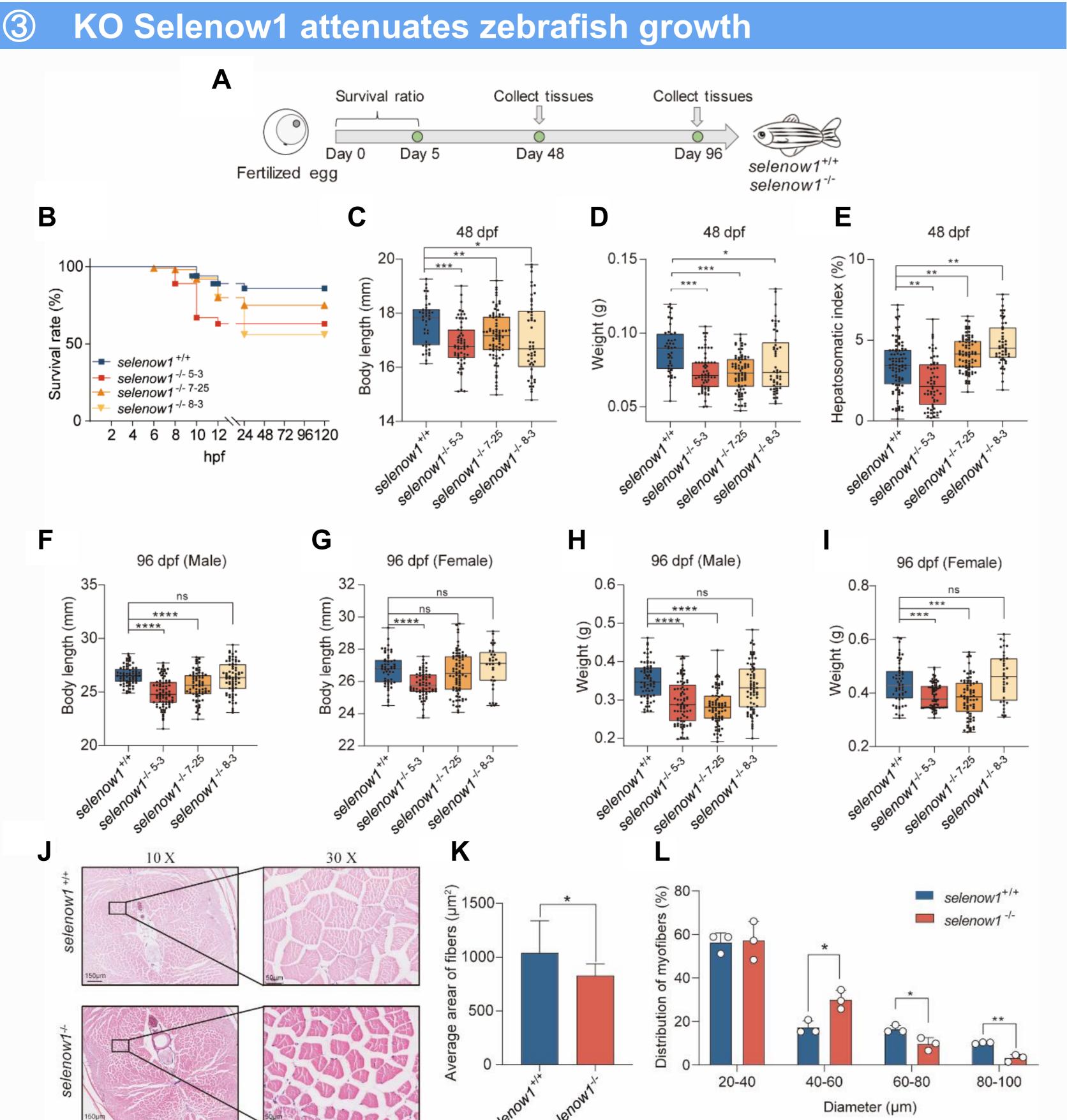


Fig. 2 A. Schematic diagram of zebrafish growth and development experiment. B. Survival curve of zebrafish embryos. C to I. Growth performance and somatic parameter of zebrafish at 48d (C to E) and 96d (F to I). J to L. Effect of knockout selenow1 on zebrafish skeletal muscle.

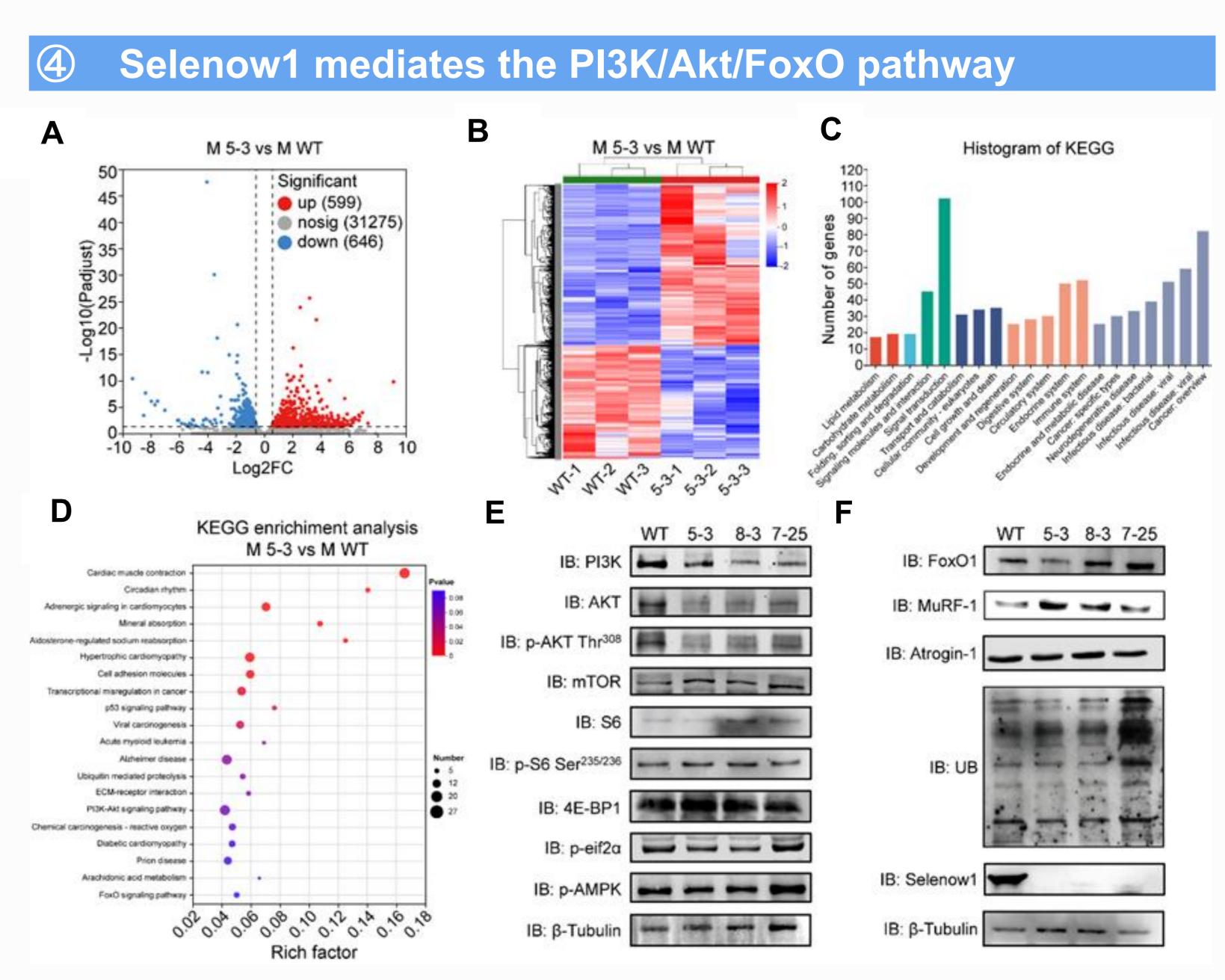


Fig. 3 Volcano map (A), Heat map (B), Kyoto Encyclopedia of Genes and Genomes (KEGG) pathway (C, D) categories of the differentially expressed genes (DEGs) in WT and *selenow1-/- 5-3* muscle. E and F. The expression of key factors of PI3K/Akt/FoxO pathway in *selenow1-/-* muscle.

## **Selenow1 stabilizes the AKT protein**

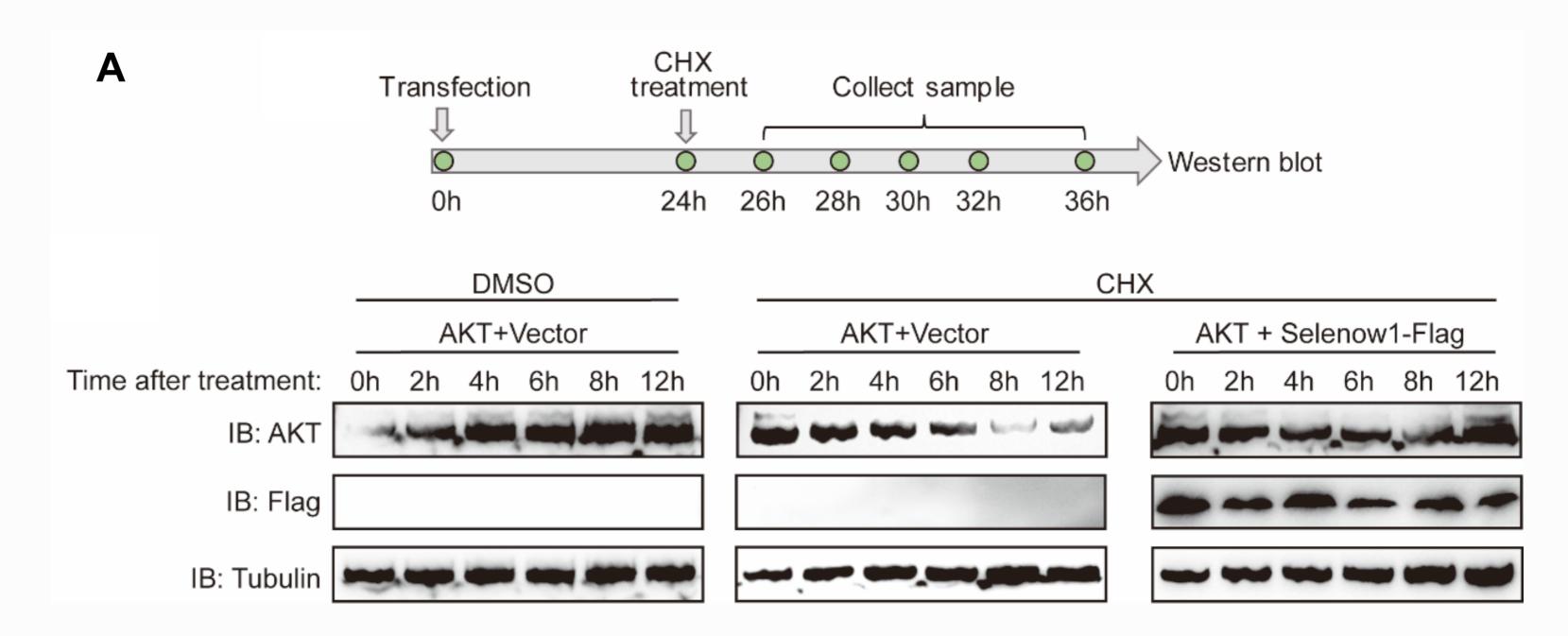
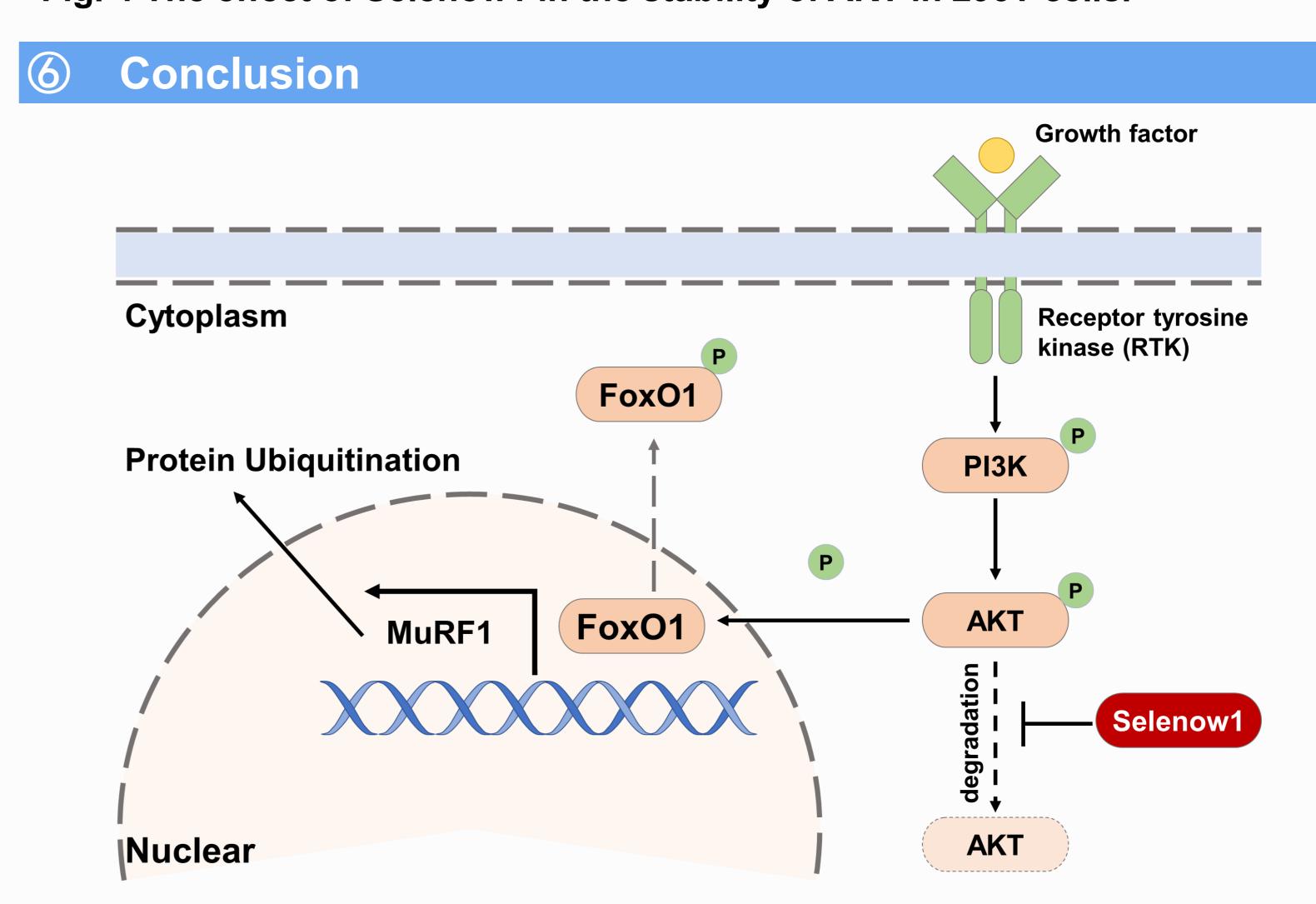


Fig. 4 The effect of Selenow1 in the stability of AKT in 293T cells.



- 1. Successfully constructed the *selenow1-/-* zebrafish model.
- 2. Selenow1 deficiency significantly attenuate the embryo development and growth of zebrafish.
- 3. Selenow1 deficiency significantly decreased the muscle development of zebrafish
- 4. Selenow1 mainly function by activating the PI3K/Akt/FoxO pathway.
- 5. Selenow1 contributes to the stabilization of the AKT protein.