## Epigenetic regulation of AURKA by miR-4715-3p in upper gastrointestinal cancers

Ahmed Gomaa<sup>1</sup>, Dunfa Peng<sup>1</sup>, Zheng Chen<sup>1, 2</sup>, Mohammed Soutto <sup>1, 2</sup>, Khaled Abouelezz<sup>3</sup>
Alejandro Corvalan<sup>4</sup>, Wael El-Rifai <sup>1, 2</sup>

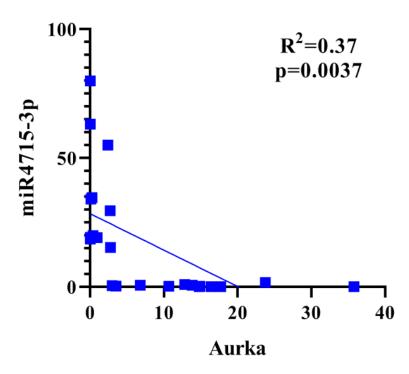
<sup>&</sup>lt;sup>1</sup> Department of Surgery, University of Miami, Miami, Florida, USA.

<sup>&</sup>lt;sup>2</sup> Department of Veterans Affairs, Miami Healthcare System, Miami, Florida, USA.

<sup>&</sup>lt;sup>3</sup> Department of Medicine, University of Miami, Miami, Florida, USA.

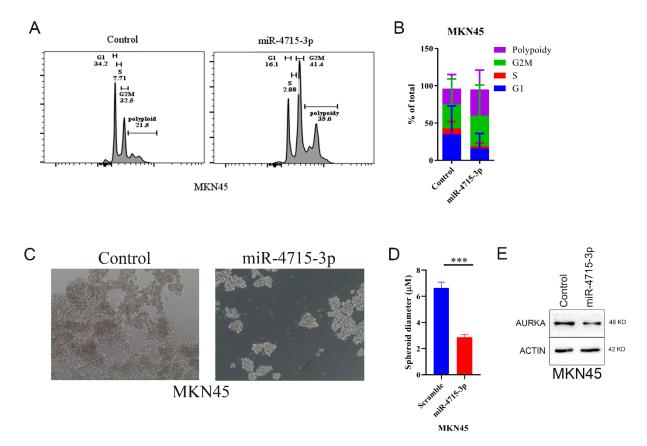
<sup>&</sup>lt;sup>4</sup> Advanced Center for Chronic Disease, School of Medicine, Pontificia Universidad Catolica de Chile, Portugal, Santiago, Chile

## **Supplementary Figure 1**

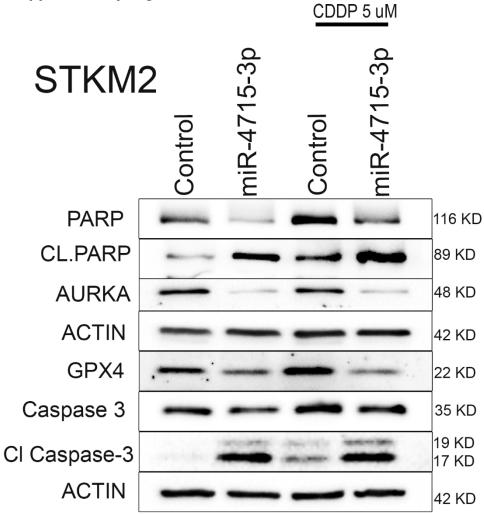


Supplementary Figure 1. Inverse correlation of expression AUKRA and miR4715-3p.

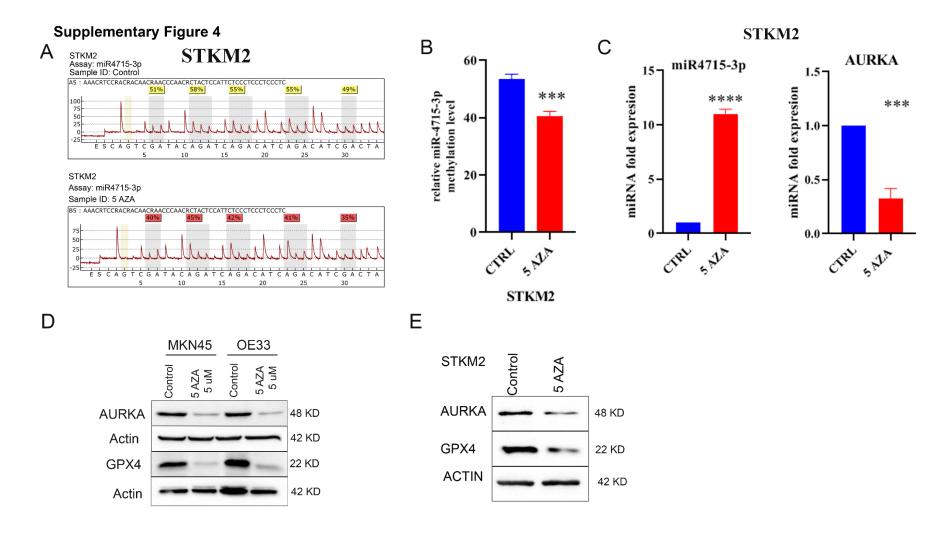
Normal gastric and tumor human tissues were analyzed for the level of AURKA and miR-47-15-3p using qPCR.



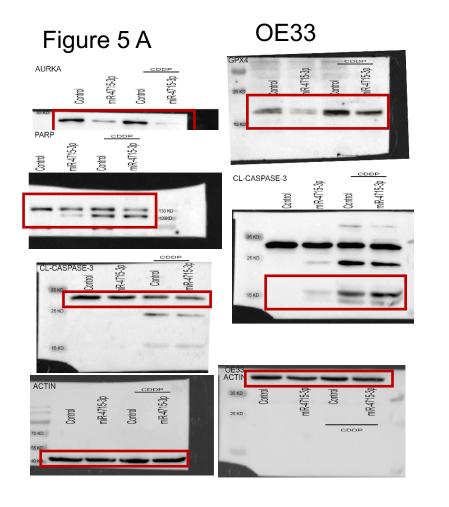
Supplementary Figure 2. miR-4715-3p suppresses cellular proliferation, and spheroid growth, enhances polyploidy, and alters cell cycle progression. (A & B) MKN45 cells were transfected with miR-4715-3p mimic 48h, and cell cycle progression was analyzed by flow cytometry. After 48h, miR-4715-3p reconstitution significantly enhanced polyploidy in MKN45 cells. (C & D) Reconstitution of miR-4715-3p was established by using lentiviral infection followed by establishing spheroid cultures. (E) Western blot analysis of MKN45 spheroids shows significant downregulation of AURKA after miR-4715-3p reconstitution.



Supplementary Figure 3. miR-4715-3p overexpression increases gastric cancer cell line, STKM2, death through inhibition of GPX4. STKM2 cells were transiently transfected with miR-4715-3p. After 48 h, cells were treated with or without Cisplatin 5  $\mu$ M 18h



**Supplementary Figure 4. (A, B)** Representative pyrosequencing profiles of STKM2 cells. **(C)** AURKA mRNA and miR-4715-3p expression levels in STKM2 cells treated with 5-Aza-2'-deoxycytidine (Decitabine) 5μM for 72 h. **(D,E)** Western blot analysis of AURKA and GPX4 protein expression levels in OE33 , MKN45 and STKKM2 cells, following treatment with 5μM of 5-Aza-2'-deoxycytidine (Decitabine) for 72h.



MKN45 Figure 5 B Control CDDP CL-Caspase -3

Figure 5 C

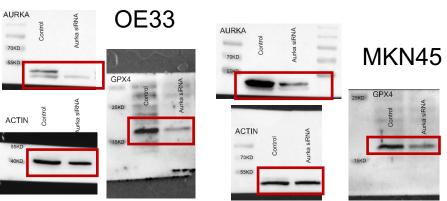
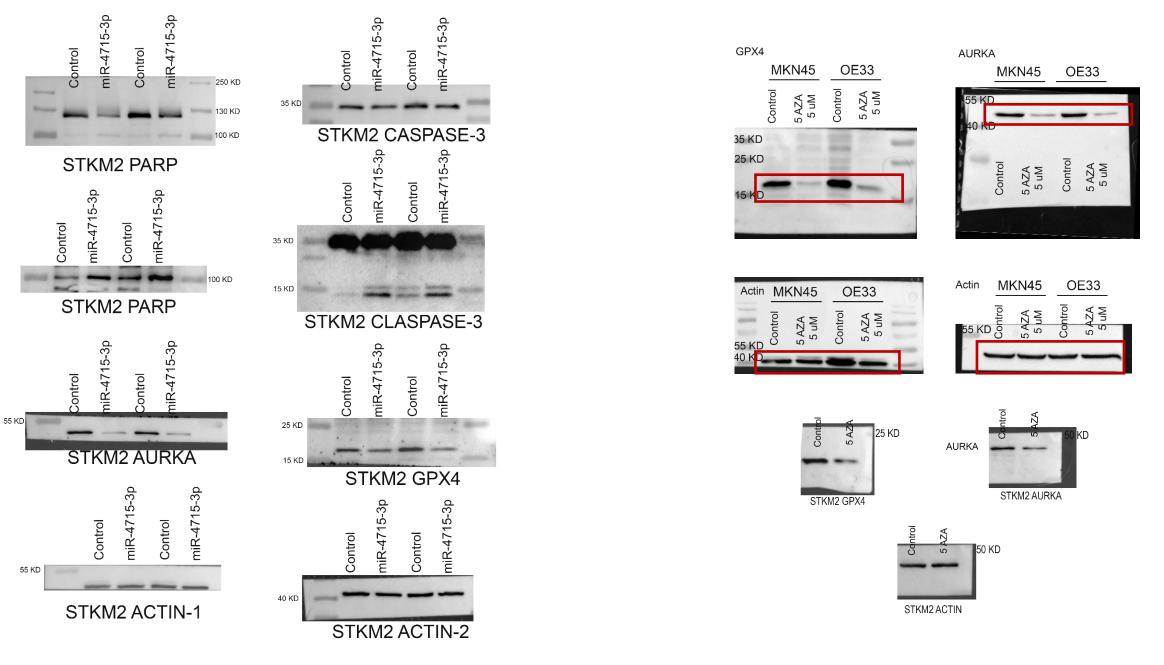


Figure 5 Raw data



Supplementary Figure 3 Raw data

Supplementary Figure 4 Raw data