

Modelling metastatic melanoma in zebrafish

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Propositions

Accompanying the PhD thesis

"Modelling metastatic melanoma in zebrafish"

- 1. Zebrafish models of metastatic uveal melanoma predict patient therapy response.
 - -This thesis, chapter 4
- 2. Melanin protects melanoma cells in circulation and therefore is a crucial, target in the treatment of metastatic melanoma.
 - -This thesis, chapter 5
- 3. Ferroptosis inducers are highly potent anti-cancer compounds *in vivo*, specifically targeting disseminating cancer cells.
 - -This thesis, chapter 5
- 4. Not only peer reviewed academical publications but also widely available (open access) dissemination of data/tools/findings are required to validate the contribution of science to the general public.
 - -This thesis, chapter 6
- 5. Despite recent therapeutic advances in cancer treatment, metastasis remains the principal cause of cancer death.
 - -Ganesh & Massagué Nat. Med. 2021
- 6. Inhibition of the cystine/glutamate antiporter (system x_c⁻) or glutathione peroxidase 4, create a void in the antioxidant defenses of the cell, ultimately leading to iron-dependent, oxidative death, namely ferroptosis.
 - -Dixon et al. Cell 2012 (paraphrasing)
- 7. Ferroptosis is induced in blood and inhibits metastasis of melanoma cells.
 - -Adapted from Ubellacker et al. Nature 2020
- 8. Induction of ferroptosis can be exploited to curb the spread of, and kill, responsive metastatic cancers.
 - -Adapted from Hadian & Stockwell, Cell, 2020
- 9. All models, and therefore all model species, are wrong (but some are useful).
 - adapted from George P. Box
- 10. Not only are zebrafish great for cancer biology, they are also beautiful, a powerful teaching aid and a great source of inspiration.

Arwin Groenewoud, Leiden, 7 June 2022