

Exploring the mechanisms of metastatic onset for novel treatment strategies

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Propositions

Accompanying the thesis

Exploring the mechanisms of metastatic onset for novel treatment strategies

- 1. Upon detecting changes in the stiffness of the extracellular matrix, intracellular YAP/TAZ molecules translocate into the nucleus, where they oversee the regulation of intracellular stemness genes. *Chapter 2, this thesis*
- 2. Prostate cancer cells undergo metabolic rewiring through mTOR-AMPK during the process of metastasis to counteract cell death induced by environmental pressures within the circulatory system. *Chapter 3, this thesis*
- 3. Cyclic ruthenium-RGD conjugated prodrugs can penetrate the zebrafish's blood-brain barrier to target glioblastoma and kill tumor cells upon photoactivation. *Chapter 4, this thesis*
- 4. Nanoparticles modified with A6 peptides can specifically target tumor cells with high CD44 expression and deliver YAP/TAZ siRNA into the tumor cells, thereby achieving a highly effective anti-cancer effect. *Chapter 5, this thesis*
- 5. Circulating tumor cells in the bloodstream adapt to changes in the external environment through alterations in their intrinsic pathways, enabling tumor cells to achieve long-distance metastasis.
- 6. Exploring the aberrant pathways shared by cancer stem cells and circulating tumor cells with the initial tumor cells is of significant importance for the development of new cancer treatment strategies.
- 7. Gene therapy based on nanoparticle delivery systems is an advanced therapeutic approach that has demonstrated potential efficacy in various diseases, including cancer.
- 8. Modifying anticancer drugs with peptides that target particular proteins can effectively enhance the specificity of the drug in treating cancer.
- 9. Science demands independent thought, relentless exploration, and the audacity to question.
- 10. Interest and passion are the greatest motivators.

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Leiden, 5th December 2023