

04 SEPT
2025

Thursday

LECTURE

MEET & EAT *

Light lunch provided
Salle Barbara Mc CLINTOCK

11.00am - 12.00pm

12.00pm - 1.00pm



miR-146a-5p increases melanoma brain metastasis development by elevating Notch signaling

ABSTRACT

Melanoma brain metastases (MBM) are a major clinical challenge due to the protective blood-brain barrier (BBB) and the brain's unique microenvironment. We demonstrate that extracellular vesicle (EV)-associated miR-146a promotes MBM by activating astrocytes and enhancing Notch signaling through downregulation of NUMB. This creates a tumor-supportive inflammatory niche that facilitates metastatic growth.

Knockdown of miR-146a or treatment with deserpentine significantly reduces tumor burden and improves survival in mouse models. In parallel, miR-138, also enriched in tumor EVs, increases BBB permeability, potentially supporting metastatic seeding. Multimodal imaging and spatial transcriptomics reveal tumor-induced vascular changes, hypoxia, and altered neuronal signaling in EV-exposed normal brain tissue.

Key transcriptomic responses involve VEGFB and hypoxic markers, suggesting exosome-driven reprogramming of the brain niche. These findings support a model where EV-delivered miRNAs establish a permissive environment for brain metastasis and highlight miR-146a as a promising therapeutic target.



SPEAKER

Prof Frits Thorsen

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HOST:

Luxembourg Institute of Health

RESPONSIBLE SCIENTIST:

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*Please note that registration for Meet and Eat is mandatory via the following link:

<https://lh1v-limsrvey01.lih.lu/index.php/191953?lang=en>

Location:

Lecture:

CHL - Centre

Room: Amphitheatre

4, rue Ernest Barblé
L-1210 Luxembourg

JOIN

Event number: 2794 070 6947

Event password: w3jTMf2Zz6w