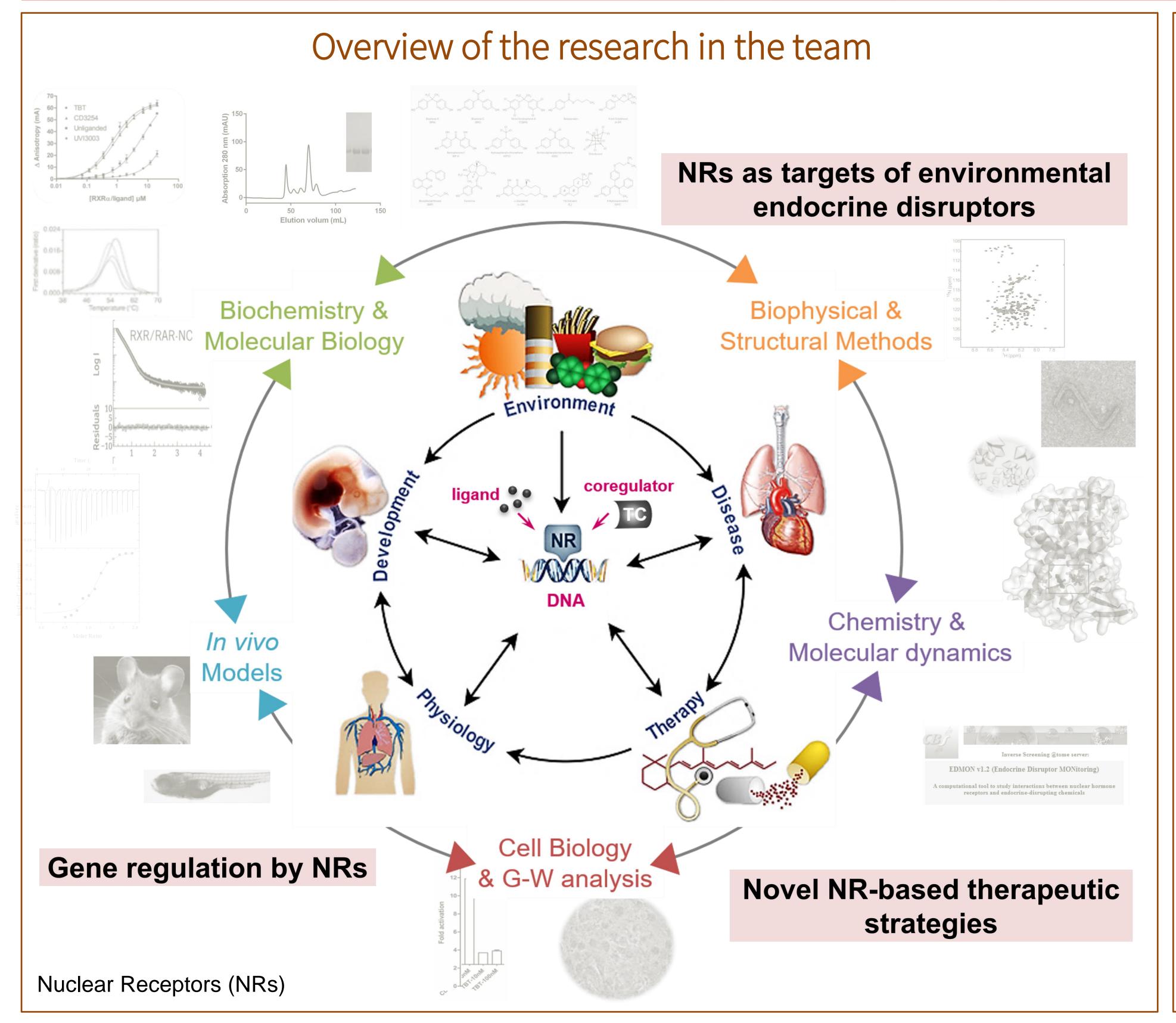


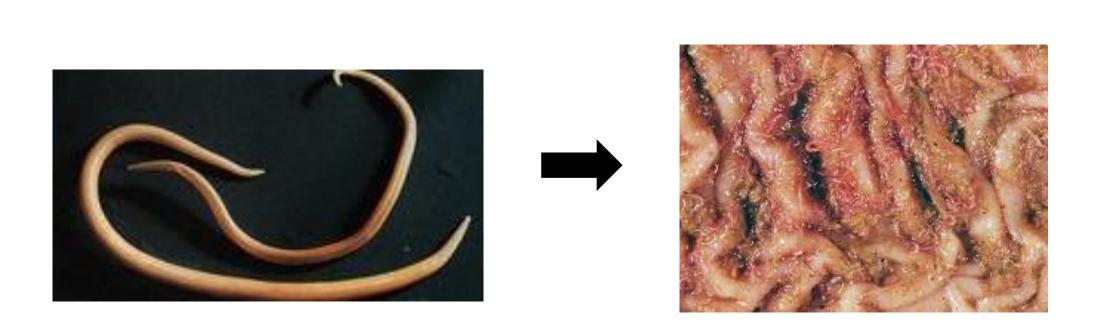
Development of an inhibitor against the nuclear receptor DAF-12 from nematodes by structural and interaction studies

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The scientific context of the M1 project



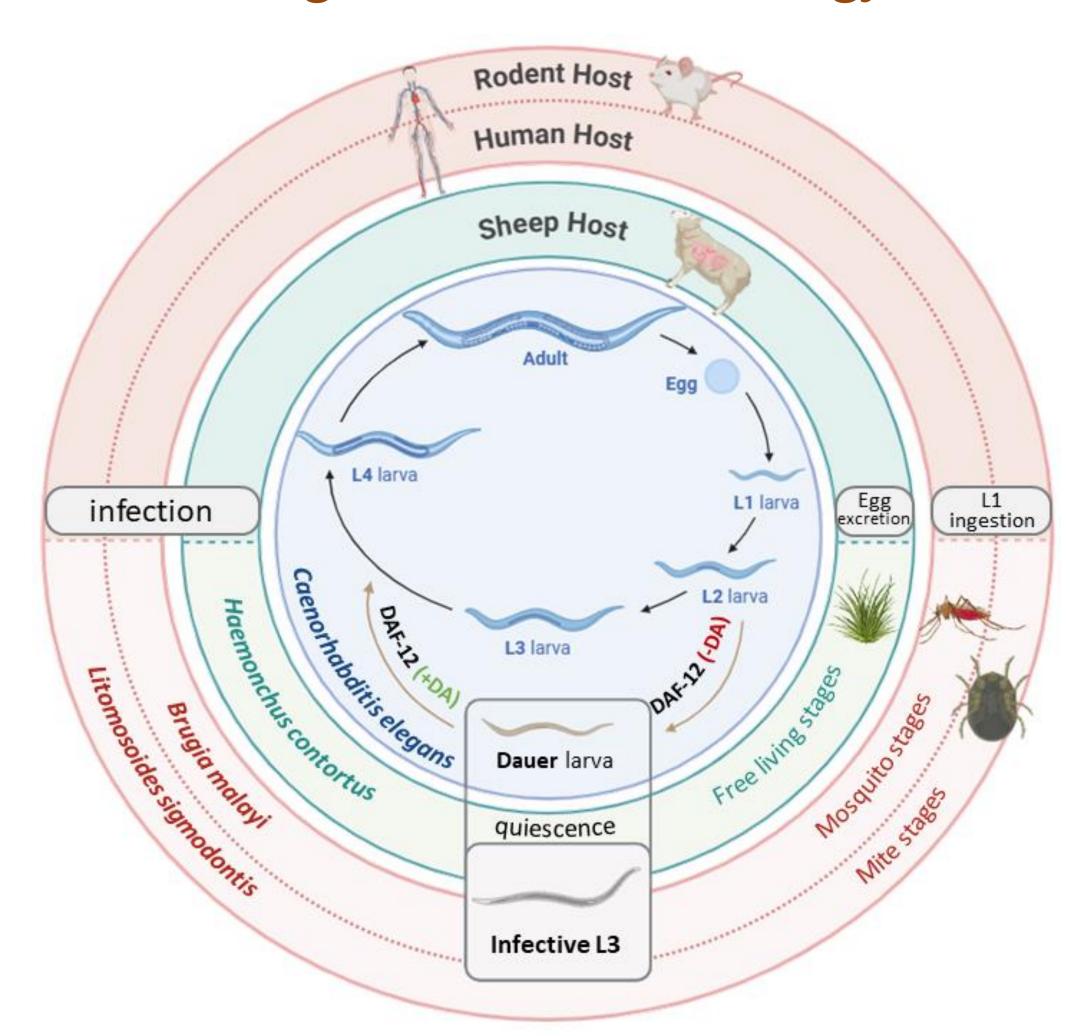
- Neglected Tropical Diseases
 Livestock and domesticated animals diseases
 Plant infection
 - ➤ RESISTANCE due to extensive use of drugs
 ➤ MODIFICATION OF PREVALENCE due to
 climate change



A global challenge in human and animal health and agriculture

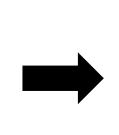
Novel therapeutic strategies are needed

The goal and methodology of the M1 project



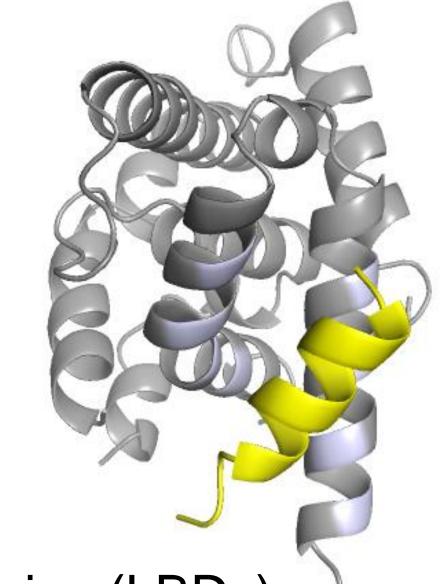
Inhibiting DAF-12 to block the infective L3 larval stage

Targeting NRs regulating infectious processes in parasitic nematodes



No drugs specifically target this critical stage making NRs a promising therapeutic target distinct from current anthelmintics

Developping an inhibitor molecule from our newly discovered lead compound



- Production and purification of DAF-12 ligand binding domains (LBDs)
- > Biochemical and biophysical interaction studies with project-developed molecules
- > Crystallographic studies of the LBDs in complex with the molecules of interest

Main collaborations and financial supports

P. Balaguer (Cell biology, IRCM, Montpellier)

M. Amblard (Chemistry, IBMM, Montpellier)

A. Lespine (Pharmacology, INTHERES, Toulouse)

C. Martin (Parasitology, MNHN, Paris)











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