



This workshop is a satellite event of EUROTOX 2024, Copenagen, 8-11-09-2024 \*\*eurotox

#### Workshop background and objectives

The ECETOC Thyroid (T4) Task Force has worked on the question of how to assess and investigate compounds, which show effects in the thyroid of laboratory animals, with particular emphasis on the neurodevelopment of offspring following in utero exposure to compounds affecting thyroid hormone balance. In our publications (Sauer et al., 2020¹, Marty et al., 2021², Marty et al., 2022³, Melching-Kollmuss et al., 2023⁴)a both human evidence for maternal thyroid hormone related neurodevelopmental toxicity as well as an extensive evaluation of rat data (comprising thyroid hormone data in adults and offspring, brain histopathological and observational neurodevelopmental effects of compounds with different thyroid molecular initiating events), using Adverse Outcome Pathway (AOP) approaches, was conducted.

This allowed us to identify relevant neurodevelopmental parameters, potential thresholds for offspring thyroid hormone levels below which no neurodevelopmental toxicity is to be assumed and correlations between levels of thyroid hormone decrements, brain histopathological outcomes and neurobehavioural findings. Moreover, a scheme to assess thyroid active compounds was developed.

ECETOC and RSA would like to propose a workshop involving regulatory toxicologists, pathologists and clinical pathologists together with experts from European regulatory authorities to review the state of the science and its current application in regulatory decision-making, and to propose future options for improving the evaluation of thyroid hormone related neurodevelopmental toxicity, including the proposed ECETOC testing and assessment scheme.

A satellite meeting back-to-back with the EUROTOX Congress is considered to be the ideal opportunity to provide a forum for debate amongst regulatory and experimental toxicologists from authorities, academia and industry. The satellite Symposium will be independently organized by ECETOC and RSA, including venue, meeting rooms, speaker selection and invitation, and participant invitation.

<sup>&</sup>lt;sup>a</sup> References can be found at the end of the programme





This workshop is a satellite event of EUROTOX 2024, Copenagen, 8-11-09-2024 **eurotox** 

### Draft Outline of the Symposium programme

Item	Start	End	Agenda item	Who
1	11:30 AM	12:00 PM	Arrival and registration	
2	12:00 PM	12:30 PM	Welcome Lunch (Registration to continue through	lunch)
3	12:30 PM	12:40 PM	Welcome, introduction and symposium objectives	Bennard van Ravenzwaay (ECETOC) and Christine Walter (RSA)
4	12:40 PM	1:00 PM	ECETOC Testing and Assessment Scheme	Stephanie Melching- Kollmuss (BASF)
5	1:00 PM	1:20 PM	Experiences with Thyroid ED assessment - EFSA	Martina Panzarea (EFSA)
6	1:20 PM	1:40 PM	Experiences with Thyroid ED assessment & criteria for the ED Guidance – ECHA	Niklas Andersson (ECHA)
7	1:40 PM	2:00 PM	In vitro and in vivo investigation of pesticide effects on the T-axis	Philip Marx-Stoelting (BfR)
8	2:00 PM	2:20 PM	Coffee break	
9	2:20 PM	2:40 PM	Examples collected by industry in context of the ECHA ED Guidance	Helen Tinwell (Bayer)
10	2:40 PM	3:00 PM	How to identify adverse neurodevelopmental toxicity in laboratory animals	Heike-Antje Marxfeld (BASF)
11	3:00 PM	3:20 PM	Decisive new endpoints for neurodevelopment	Katie O'Shaugnessy (US EPA)
12	3:20 PM	3:40 PM	Assessment of thyroid hormone alterations in rat brain and plasma	Christiane Hindrichs (BASF Metabolome Solutions)
13	3:40 PM	4:00 PM	Quantitative and species-specific aspects	Lysiane Richert and Laure Asselin (Kaly-Cell)
14	4:00 PM	4:15 PM	Summarise and closing remarks	Bennard van Ravenzwaay and Christine Walter
15	4:15 PM	4:45 PM	30 min to EUROTOX opening ceremony	
16	4:45 PM	6:45 PM	EUROTOX Congress Opening Ceremony (Tivoli Hotel & Congress Center)	





This workshop is a satellite event of EUROTOX 2024, Copenagen, 8-11-09-2024 **eurotox** 

#### About the organisers

ECETOC is a not-for-profit scientific association that provides a collaborative space for leading scientists from academia, governments and industry to develop and promote trusted and practical scientific solutions which ensure a safe, sustainable and healthy world. For more information please visit https://www.ecetoc.org/

RSA (Regulatory Science Associates) was formed in 2007 with a group of experts delivering human health toxicology and regulatory affairs consultancy services. Further information can be found at <a href="https://www.regulatoryscience.com">www.regulatoryscience.com</a>

ECETOC and RSA have significant experience organising workshops and conferences to facilitate the communication of science. Both organisations are working jointly to organise this thyroid hormone related symposium for 2024.





This workshop is a satellite event of EUROTOX 2024, Copenagen, 8-11-09-2024 **Leurotox** 

#### References

<sup>1</sup> Sauer, U. G., Asiimwe, A., Botham, P. A., Charlton, A., Hallmark, N., Jacobi, S., . . . Swaen, G. (2020). Toward a science-based testing strategy to identify maternal thyroid hormone imbalance and neurodevelopmental effects in the progeny - part I: which parameters from human studies are most relevant for toxicological assessments? *Crit Rev Toxicol*, *50*(9), 740-763. doi:10.1080/10408444.2020.1839380

<sup>&</sup>lt;sup>2</sup> Marty, S., Beekhuijzen, M., Charlton, A., Hallmark, N., Hannas, B. R., Jacobi, S., . . . van Ravenzwaay, B. (2021). Towards a science-based testing strategy to identify maternal thyroid hormone imbalance and neurodevelopmental effects in the progeny - part II: how can key events of relevant adverse outcome pathways be addressed in toxicological assessments? *Crit Rev Toxicol*, *51*(4), 328-358. doi:10.1080/10408444.2021.1910625

<sup>&</sup>lt;sup>3</sup> Marty, M. S., Sauer, U. G., Charlton, A., Ghaffari, R., Guignard, D., Hallmark, N., . . . van Ravenzwaay, B. (2022). Towards a science-based testing strategy to identify maternal thyroid hormone imbalance and neurodevelopmental effects in the progeny-part III: how is substance-mediated thyroid hormone imbalance in pregnant/lactating rats or their progeny related to neurodevelopmental effects? *Crit Rev Toxicol*, *52*(7), 546-617. doi:10.1080/10408444.2022.2130166

<sup>&</sup>lt;sup>4</sup> Melching-Kollmuss, S., Bothe, K., Charlton, A., Gangadharan, B., Ghaffari, R., Jacobi, S., . . . van Ravenzwaay, B. (2023). Towards a science-based testing strategy to identify maternal thyroid hormone imbalance and neurodevelopmental effects in the progeny – Part IV: the ECETOC and CLE Proposal for a Thyroid Function-Related Neurodevelopmental Toxicity Testing and Assessment Scheme (Thyroid-NDT-TAS). *Critical Reviews in Toxicology, 53*(6), 339-371. doi:10.1080/10408444.2023.2231033