# Pesticides Peer Review (PREV) Unit 11<sup>th</sup> meeting of the Working Group on Neurotoxicity



9 December 2025 14:00-18:00 MINUTES - Agreed on 18 December 2025

Location: Webconference

#### Attendees:

#### Working Group Members:

Adeline Cavelier, Andrea Terron (Chair), Antonio Hernández-Jerez, Carsten Kneuer, Emily Mcvey, Helena Hogberg-Durdock, Kevin Crofton.

#### o European Commission and/or Member States representatives:

None

#### o EFSA:

Pesticides Peer Review (PREV) Unit: Iris Mangas, Anna Federica Castoldi, Martina Panzarea.

Methodological and Scientific Support (MESE) Unit: Irene Cattaneo

#### Others:

Tanja Burgdorf for agenda point 4.

Oddvar Myhre, Anna Forsby, Marie Gabrielle Zurich Fontanellaz, Katharina Koch participated as observers for agenda point 5, via web conference.

ECHA: Kati Hellsten (absent), Kai Craenen (absent).

# I. Welcome and apologies for absence

The Chair welcomed the participants. Apologies were received from Kati Hellsten and Kai Craenen (ECHA).

# II. Adoption of agenda

The agenda was adopted with no changes.

# III. Declarations of Interest of Working Groups members

In accordance with EFSA's Policy on Independence<sup>1</sup> and the Decision of the Executive Director on Competing Interest Management<sup>2</sup>, EFSA screened the Annual Declarations of Interest filled out by the Working Group members invited to the present meeting. No Conflicts of Interest related to the issues discussed in this meeting have been identified during the screening process, and no interests were declared orally by the members at the beginning of this meeting.

Emily McVey indicated a new DOI was submitted and is current under-assessment. No

<sup>&</sup>lt;sup>1</sup> http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/policy\_independence.pdf

<sup>2</sup> http://www.efsa.europa.eu/sites/default/files/corporate publications/files/competing interest management 17.pdf

# MEETING MINUTES – 9 December 2025 11<sup>th</sup> Working Group meeting on Neurotoxicity



Interest was declared orally.

# IV. Agreement of the minutes of the 10<sup>th</sup> Working Group meeting held on 8 December 2025, via web-conference

The minutes of the  $10^{th}$  Working Group meeting were agreed by written procedure on  $8^{th}$  of December 2025.

#### V. Scientific topic(s) for discussion

#### 1.1. Tebufenpyrad IATA: Recommendation and future activities section

The Working Group discussed responses to selected comments received from the OECD review and possible amendments to be included in the IATA. The WG agreed that the section on lessons learned and recommandations should reflect the outline of the IATA, which is based on three distinct pillars, and i.e. identification of in vitro PoD, exposure assessment and modelling, and identification of margins of internal exposure. The WG appreciated the scientific contribution given by Tanja Burgdorf for the revision and update of the IATA.

# 1.2. Update on the ongoing Brain Health Grant Agreement (GP/EFSA/ED/2022/01)

The Working Group was informed about the recent updates and status of Brain Health project **GP/EFSA/ED/2022/01** under the IRMA program. The Working Group was asked to review and advice on the deliverable 2 of the Brain Health project via written procedure after the meeting.

# **VI.** Any Other Business

None.

# VII. Next meeting

The dates for the next meetings will be decided via written procedure.

# Pesticides Peer Review (PREV) Unit 10<sup>th</sup> Working Group meeting on Neurotoxicity



01 December 2025 14:00-18:00 MINUTES - Agreed on 5 December 2025

Location: Webconference

#### Attendees:

#### Working Group Members:

Andrea Terron (Chair), Carsten Kneuer, Emily Mcvey, Helena Hogberg-Durdock, Kevin Crofton, Adeline Cavelier

Hearing Experts<sup>1</sup>:

None

European Commission and/or Member States representatives:

None

o EFSA:

Pesticides Peer Review (PREV) Unit: Iris Mangas, Anna Federica Castoldi, Martina Panzarea MESE Unit: Irene Cattaneo, Laura Martino, Alicia Paini

Others:

Kati Hellsten and Kai Craenen (ECHA) participated as observers for the entire meeting, via web conference.

# I. Welcome and apologies for absence

The Chair welcomed the participants.

# II. Adoption of agenda

The agenda was adopted without changes

# III. Declarations of Interest of Working Groups members

In accordance with EFSA's Policy on Independence2 and the Decision of the Executive Director on Competing Interest Management3, EFSA screened the Annual Declarations of Interest filled out by the Working Group members invited to the present meeting and the preparatory meetings listed under agenda item "Preparatory meeting/s". No Conflicts of Interest related to the issues discussed in these meetings have been identified during the screening process, and no interests were declared orally by the members at the beginning of this meeting.

# IV. Agreement of the minutes of the 9<sup>th</sup> Working Group meeting held on 17 November 2025, via web-conference

As defined in Article 34 of the Decision of the Executive Director concerning the selection of members of the Scientific Committee, the Scientific Panels, and the selection of external experts to assist EFSA with its scientific work: <a href="http://www.efsa.europa.eu/en/keydocs/docs/expertselection.pdf">http://www.efsa.europa.eu/en/keydocs/docs/expertselection.pdf</a>

<sup>&</sup>lt;sup>2</sup> http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/policy\_independence.pdf

<sup>3</sup> http://www.efsa.europa.eu/sites/default/files/corporate publications/files/competing interest management 17.pdf

# MEETING MINUTES – 1 December 2025 10<sup>th</sup> Working Group meeting on Neurotoxicity



The minutes of the 9<sup>th</sup> Working Group meeting were agreed by written procedure on 5 December 2025.

#### V. Scientific topic(s) for discussion

- 1.1. IATA tebufenpyrad update from OECD and plan
- 1.2. Comments on PBK update
- 1.3. Comments on UA and EKE process update
- 1.4. Comments to be discussed by the WG

The Working Group (WG) experts were updated by EFSA Staff on items 4, 5, 6 and 7. After that the WG members were asked to provide advice on the draft answers to OECD delegation of IATA Cs project. The answers were drafted in the compiled document to be shared with OECD IATA Cs project.

### VI. Preparatory Meeting/s

Since the last 9th WG meeting, no preparatory meetings were held

### VII. Next meeting

The next meetings will be held on:

• 9<sup>th</sup> December 2025, via webconference.

# Pesticides Peer Review (PREV) Unit 9<sup>th</sup> Working Group meeting on Neurotoxicity

efsa EUROPEAN FOOD SAFETY ALITHOPITY

17 November 2025 14:00-18:00 MINUTES - Agreed on 5 December 2025

Location: Webconference

#### Attendees:

#### Working Group Members:

Andrea Terron (Chair), Antonio Hernández-Jerez, Carsten Kneuer, Emily Mcvey, Helena Hogberg-Durdock, Kevin Crofton, Adeline Cavelier

#### Hearing Experts<sup>1</sup>:

None

European Commission and/or Member States representatives:

None

EFSA:

Pesticides Peer Review (PREV) Unit: Anna Federica Castoldi, Martina Panzarea

o Others:

Martin Paparella and Oddvar Myhre participated as observers for agenda point 4, via web conference.

Barbara Viviani participated as observers for agenda point 5, via web conference.

Kati Hellsten and Kai Craenen participated as observers for the entire meeting, via web conference.

#### I. Welcome and apologies for absence

The Chair welcomed the participants.

# II. Adoption of agenda

The agenda was adopted without changes.

# III. Declarations of Interest of Working Groups members

In accordance with EFSA's Policy on Independence2 and the Decision of the Executive Director on Competing Interest Management3, EFSA screened the Annual Declarations of Interest filled out by the Working Group members invited to the present meeting and the preparatory meetings listed under agenda item "Preparatory meeting/s". No Conflicts of Interest related to the issues discussed in these meetings have been identified during the screening process, and no interests were declared orally by the members at the beginning of this meeting.

As defined in Article 34 of the Decision of the Executive Director concerning the selection of members of the Scientific Committee, the Scientific Panels, and the selection of external experts to assist EFSA with its scientific work: <a href="http://www.efsa.europa.eu/en/keydocs/docs/expertselection.pdf">http://www.efsa.europa.eu/en/keydocs/docs/expertselection.pdf</a>

<sup>&</sup>lt;sup>2</sup> http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/policy\_independence.pdf

<sup>3</sup> http://www.efsa.europa.eu/sites/default/files/corporate publications/files/competing interest management 17.pdf

### MEETING MINUTES – 17 November 2025 9<sup>th</sup> Working Group meeting on Neurotoxicity



# IV. Agreement of the minutes of the 8<sup>th</sup> Working Group meeting held on 30 September 2025, via web-conference

The minutes of the 8<sup>th</sup> Working Group meeting were agreed by written procedure on 16 October 2025.

# V. Scientific topic(s) for discussion

# 1.1. Scientific advice on the on-going environmental neurotoxicant project (OC/EFSA/PREV/2023/01)

The Working Group (WG) experts were asked to provide revision and advice on the risk of bias assessment done by the contractor of LOT 4 on the in vivo DNT Test-Guideline studies included in deliverable 3 of LOT4. The outcome will be considered for inclusion in the LOT 4 deliverables.

One of the WG expert was asked to provide revision and advice on the assessment of human observational studies included in deliverable 2 of LOT 4. The outcome was presented to the other WG experts and will be considered for inclusion in the LOT 4 deliverables.

# a. NP/EFSA/PREV/2024/02 Development of an AOP network for Parkinsonian motor symptoms. Deliverables

The WG experts were asked to provide revision and advice on the AOP network for Parkinsonian motor symptoms. The advice will be considered for inclusion in the AOP deliverable.

# VI. Preparatory Meeting/s

Since the last 8th WG meeting, a preparatory meeting chaired by EFSA staff was held as follows: **VIII.1 Preparatory meeting to the 9<sup>th</sup> WG meeting on Neurotoxicity**, 30 October 2025

VIII.1a Andrea Terron

VIII.1b Preparation for the WG meeting discussion.

# VII. Next meeting

The next meetings will be held on:

- 1<sup>st</sup> December 2025, via webconference.
- 9<sup>th</sup> December 2025, via webconference.

# Pesticides Peer Review (PREV) Unit 8<sup>th</sup> Working Group meeting on Neurotoxicity

efsa EUROPEAN FOOD SAFETY AUTHORITY

30 September 2025 9:30-13:30 MINUTES - Agreed on 16 October 2025

Location: Webconference

Attendees:						
o Andrea	Working Group Members: a Terron (Chair), Antonio Hernández-Jerez					
o None	Hearing Experts <sup>1</sup> :					
o None	Applicants participating in a technical hearing <sup>2</sup> :					
o None	European Commission and/or Member States representatives:					

o EFSA:

Pesticides Peer Review (PREV) Unit: Martina Panzarea, Rafaela Rodrigues de Jesus

o Others:

None

# I. Welcome and apologies for absence

The Chair welcomed the participants.

# II. Adoption of agenda

The agenda was adopted without changes.

# **III. Declarations of Interest of Working Groups members**

<sup>&</sup>lt;sup>1</sup> As defined in Article 34 of the document "Implementing Rule of the Management Board of the European Food Safety Authority laying down the rules on the selection, appointment and operations of the Scientific Committee, Scientific Panels and of their Working Groups": <a href="https://www.efsa.europa.eu/sites/default/files/paneloperation.pdf">https://www.efsa.europa.eu/sites/default/files/paneloperation.pdf</a>

<sup>&</sup>lt;sup>2</sup> As defined in the "EFSA's Catalogue of support initiatives during the life-cycle of applications for regulated products": https://efsa.onlinelibrary.wiley.com/doi/epdf/10.2903/sp.efsa.2021.EN-6472

### MEETING MINUTES – 30 September 2025 8<sup>th</sup> Working Group meeting on Neurotoxicity



In accordance with EFSA's Policy on Independence<sup>3</sup> and the Decision of the Executive Director on Competing Interest Management<sup>4,</sup> EFSA screened the Annual Declarations of Interest filled out by the Working Group members invited to the present meeting. No Conflicts of Interest related to the issues discussed in this meeting have been identified during the screening process, and no interests were declared orally by the members at the beginning of this meeting.

# IV. Agreement of the minutes of the 7<sup>th</sup> Working Group meeting held on 29<sup>th</sup> September 2025, via web-conference

The minutes of the 7<sup>th</sup> Working Group meeting were agreed by written procedure on 15<sup>th</sup> October 2025.

### V. Scientific topic for discussion

# 1.1. Scientific advice on the on-going Environmental Neurotoxicant project OC/EFSA/PREV/2023/01

The Working Group discussed and finalised the advice on the assessment of human observational studies included in the deliverable 2 of LOT 4 of the Environmental Neurotoxicants project.

# VI. Any Other Business

Nothing was indicated during the meeting.

# VII. Next meeting

The next meeting will be held on 17<sup>th</sup> November 2025, via webconference.

<sup>3</sup> http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/policy\_independence.pdf

<sup>4</sup> https://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/decision-ed-on-competing-interest-management-2024.pdf

# Pesticides Peer Review (PREV) Unit 7th Working Group meeting on Neurotoxicity

ersa European FOOD SAFETY AUTHORITY

29 September 2025 14:00-18:00 MINUTES - Agreed on 15 October 2025

Location: Webconference

#### Attendees:

#### O Working Group Members:

Adeline Cavelier, Andrea Terron (Chair), Antonio Hernández-Jerez, Carsten Kneuer, Emily Mcvey, Helena Hogberg-Durdock, Kevin Crofton

#### Hearing Experts<sup>1</sup>:

Bertrand Desprez, Jaroslav Legath, Marcel Falis, Vladimir Petrovic (absent), Ahmed-Chaouki Zerouala (absent)

#### European Commission and/or Member States representatives:

None

#### o EFSA:

Pesticides Peer Review (PREV) Unit: Anna Federica Castoldi, Martina Panzarea, Rafaela Rodrigues de Jesus

#### o Others:

ECHA: Kati Hellsten (absent), Kai Craenen (absent)

# I. Welcome and apologies for absence

The Chair welcomed the participants. Apologies were received for Kati Hellsten.

# II. Adoption of agenda

The agenda was adopted with changes. An additional agenda point was added to discuss the Working Group meeting dates for December.

# III. Declarations of Interest of Working Groups members

In accordance with EFSA's Policy on Independence<sup>2</sup> and the Decision of the Executive Director on Competing Interest Management<sup>3</sup>, EFSA screened the Annual Declarations of Interest filled out by the Working Group members invited to the present meeting. No Conflicts of Interest related to the issues discussed in this meeting have been identified during the screening

As defined in Article 34 of the Decision of the Executive Director concerning the selection of members of the Scientific Committee, the Scientific Panels, and the selection of external experts to assist EFSA with its scientific work: <a href="http://www.efsa.europa.eu/en/keydocs/docs/expertselection.pdf">http://www.efsa.europa.eu/en/keydocs/docs/expertselection.pdf</a>

<sup>&</sup>lt;sup>2</sup> http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/policy\_independence.pdf

<sup>&</sup>lt;sup>3</sup> http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/competing\_interest\_management\_17.pdf

# MEETING MINUTES – 29 September 2025 7<sup>th</sup> Working Group meeting on Neurotoxicity



process, and no interests were declared orally by the members at the beginning of this meeting.

One of the experts, Kevin Crofton, declared that, while working at US EPA, his previous employment, he might have been consulted for the evaluation of the developmental neurotoxicity study of boscalid. However, it is noted that the interest is more than 5 years-old and (i.e. conducted by US EPA, in 2002/2003), therefore not subject to declaration.

### IV. Hearing Expert(s)

The experts Jaroslav Legath (SK), Marcel Falis (SK), Vladimir Petrovic (SK), Ahmed-Chaouki Zerouala (FR) and Bertrand Desprez (FR) participated as hearing experts being Rapporteur Member State (RMS) and co-RMS for boscalid, respectively, to give clarifications, where needed, on studies and data available.

# V. Agreement of the minutes of the 6<sup>th</sup> Working Group meeting held on 3<sup>rd</sup> June 2025, via web-conference

The minutes of the 6<sup>th</sup> Working Group meeting were agreed by written procedure on 25<sup>th</sup> of September 2025.

### VI. Scientific topic(s) for discussion

# 1.1. Boscalid developmental neurotoxicity and neurotoxicity in the context of <a href="EFSA-0-2015-00522">EFSA-0-2015-00522</a>

The available data for evaluating the Developmental Neurotoxicity and Neurotoxicity potential of the active substance boscalid were presented and discussed. The outcome of the discussion held during the Working Group meeting will form the basis of the Working Group advice, which will be shared with the Member States experts ahead of the Peer Review Expert meeting in November 2025 and presented during that meeting, as needed.

#### 1.2. Meeting dates Dec 2025

The Working Group meeting days for November and December were discussed and agreed by the Experts.

# VII. Any Other Business

 Update on the ongoing Environmental Neurotoxicant project <u>OC/EFSA/PREV/2023/01</u>

The Working Group experts were also informed about the recent updates and status of LOT 3 (To Generate Pesticides/Chemicals Data for Neurotoxicity Using NAMs) and LOT 4 (To Produce Collaborative AOP Informed IATA for DNT Risk Assessment of Pesticides Active Substances). The Working Group experts were asked to review and provide comments to the deliverable 3 from LOT 4

# MEETING MINUTES – 29 September 2025 7<sup>th</sup> Working Group meeting on Neurotoxicity



# VIII. Next meeting

	The nex	ct meeting	will be	held o	on the	30 <sup>th</sup>	September.	via w	eb conference
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# **Pesticides Peer Review (PREV) Unit** 6<sup>th</sup> Working Group meeting on Neurotoxicity



3 September 2025 9:00-13:00 MINUTES - Agreed on 25 September 2025

Locati	on: Webconference				
Attendees:					
o Andrea	Working Group Members: a Terron (Chair), Antonio Hernández-Jerez				
o None	Hearing Experts <sup>1</sup> :				
o None	Applicants participating in a technical hearing <sup>2</sup> :				
o None	European Commission and/or Member States representatives:				
o Pesticion Jesus	<b>EFSA:</b> des Peer Review (PREV) Unit: Anna Federica Castoldi, Martina Panzarea, Rafaela Rodrigues				
o None	Others:				

# I. Welcome and apologies for absence

The Chair welcomed the participants.

# II. Adoption of agenda

The agenda was adopted without changes.

# III. Declarations of Interest of Working Groups members

<sup>&</sup>lt;sup>1</sup> As defined in Article 34 of the document "Implementing Rule of the Management Board of the European Food Safety Authority laying down the rules on the selection, appointment and operations of the Scientific Committee, Scientific Panels and of their Working Groups": <a href="https://www.efsa.europa.eu/sites/default/files/paneloperation.pdf">https://www.efsa.europa.eu/sites/default/files/paneloperation.pdf</a>

<sup>&</sup>lt;sup>2</sup> As defined in the "EFSA's Catalogue of support initiatives during the life-cycle of applications for regulated products": https://efsa.onlinelibrary.wiley.com/doi/epdf/10.2903/sp.efsa.2021.EN-6472

### MEETING MINUTES – 3 September 2025 6<sup>th</sup> Working Group meeting on Neurotoxicity



In accordance with EFSA's Policy on Independence<sup>3</sup> and the Decision of the Executive Director on Competing Interest Management<sup>4</sup>, EFSA screened the Annual Declarations of Interest filled out by the Working Group members invited to the present meeting and the preparatory meetings listed under agenda item "Preparatory meeting/s". No Conflicts of Interest related to the issues discussed in these meetings have been identified during the screening process, and no interests were declared orally by the members at the beginning of this meeting.

# IV. Agreement of the minutes of the 5<sup>th</sup> Working Group meeting held on 14 July 2025, via web-conference

The minutes of the 5<sup>th</sup> Working Group meeting were agreed by written procedure on 29 July 2025.

#### V. Scientific topic for discussion

a. Scientific advice on the on-going Environmental Neurotoxicant project OC/EFSA/PREV/2023/01

The WG experts were asked to provide revision and advice on the assessment of human observational studies included in deliverable 2 of LOT 4 (To Produce Collaborative AOP Informed IATA for DNT Risk Assessment of Pesticides Active Substances). The outcome will be considered for inclusion in the LOT 4 deliverables.

# VI. Any Other Business

Nothing was indicated during the meeting.

# VII. Preparatory Meeting

Since the last WG meeting, 1 preparatory meeting chaired by EFSA staff was held as follows:

- 1.1 Preparatory meeting to the 6th WG meeting on Neurotoxicity, 1 September 2025
- 1.1a Andrea Terron
- 1.1b Update on the LOT 4 of the Environmental Neurotoxicants project

# VIII. Next meeting

The next meeting will be held on 29 September 2025, via webconference.

<sup>&</sup>lt;sup>3</sup> http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/policy\_independence.pdf

<sup>4</sup> https://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/decision-ed-on-competing-interest-management-2024.pdf

# Pesticides Peer Review (PREV) Unit 5th meeting of the Working Group on Neurotoxicity

ersa European FOOD SAFETY AUTHODITY

14 July 2025 14:00-18:00 MINUTES - Agreed on 29 July 2025

Location: Webconference

#### Attendees:

#### O Working Group Members:

Andrea Terron (Chair), Antonio Hernández-Jerez, Carsten Kneuer, Emily Mcvey, Helena Hogberg-Durdock, Kevin Crofton

#### Hearing Experts<sup>1</sup>:

Bettina Hrdina-Zoedl, Daniela Jölli, Nina Henriette Jorgensen

European Commission and/or Member States representatives:

None

o EFSA:

Pesticides Peer Review (PREV) Unit: Anna Federica Castoldi, Martina Panzarea, Rafaela Rodrigues Jesus

Others:

None

# I. Welcome and apologies for absence

The Chair welcomed the participants. Apologies were received from Adeline Cavelier.

# II. Adoption of agenda

The agenda was adopted without changes.

# III. Declarations of Interest of Working Groups members

In accordance with EFSA's Policy on Independence<sup>2</sup> and the Decision of the Executive Director on Competing Interest Management<sup>3,</sup> EFSA screened the Annual Declarations of Interest filled out by the Working Group members invited to the present meeting and the preparatory meetings listed under agenda item "Preparatory meeting/s". No Conflicts of Interest related to the issues discussed in these meetings have been identified during the screening process, and no interests were declared orally by the members at the beginning of this meeting.

Andrea Terron indicated a new DOI was submitted and is currently under assessment. No interest was declared orally.

As defined in Article 34 of the Decision of the Executive Director concerning the selection of members of the Scientific Committee, the Scientific Panels, and the selection of external experts to assist EFSA with its scientific work: <a href="http://www.efsa.europa.eu/en/keydocs/docs/expertselection.pdf">http://www.efsa.europa.eu/en/keydocs/docs/expertselection.pdf</a>

<sup>&</sup>lt;sup>2</sup> http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/policy\_independence.pdf

<sup>3</sup> http://www.efsa.europa.eu/sites/default/files/corporate publications/files/competing interest management 17.pdf

# MEETING MINUTES – 14 July 2025 5th Working Group meeting on Neurotoxicity



#### IV. Hearing Expert(s)

The experts Bettina Hrdina-Zoedl (AT), Daniela Jölli (AT), Nina Henriette Jorgensen (DK) participated as hearing experts being Rapporteur Member State (RMS) and co-RMS for Fluazinam, respectively, to give clarifications, where needed, on studies and data available.

# V. Agreement of the minutes of the 4<sup>th</sup> Working Group meeting held on 23<sup>rd</sup> June 2025, via web-conference

The minutes of the 4<sup>th</sup> Working Group meeting were agreed by written procedure on 11<sup>th</sup> of July 2025.

# VI. Scientific topic(s) for discussion

#### 1.1. Fluazinam DNT and brain weights in the context of <a href="EFSA-Q-2016-00313">EFSA-Q-2016-00313</a>

The available data for evaluating the developmental neurotoxicity potential of fluazinam were discussed. The outcome of the discussion held during the working group meeting will form the basis of a WG advice on data interpretation. This advice will be shared with the Member States experts ahead of the Peer Review Expert meeting in September 2025 and presented during that meeting, as needed.

# VII. Any Other Business

 Scientific advice on the on-going Environmental Neurotoxicant project <u>OC/EFSA/PREV/2023/01</u>

The WG experts were asked to finalise the proof-read of the deliverables of LOT 2 (To Produce Transferability for the DNT IVB) and LOT 4 (To Produce Collaborative AOP Informed IATA for DNT Risk Assessment of Pesticides Active Substances), presented and discussed in the previous WG meeting.

The WG experts were also informed about the recent updates and status of LOT 3 (To Generate Pesticides/Chemicals Data for Neurotoxicity Using NAMs).

# VIII. Preparatory Meeting/s

Since the last 4<sup>th</sup> WG meeting, a preparatory meeting chaired by EFSA staff was held as follows:

VIII.1 Preparatory meeting to the 5<sup>th</sup> WG meeting on Neurotoxicity, 9<sup>th</sup> July 2025 VIII.1a Andrea Terron

**VIII.1b** Preparation for the active substance fluazinam discussion. Update on LOT 2, 3 and 4 of the Environmental Neurotoxicants project.

# MEETING MINUTES – 14 July 2025 5th Working Group meeting on Neurotoxicity



# IX. Next meeting

The ne	yt meeting	will he	held on	the 3rd	September.	via weh	conference
THE HE	xt meeuna	will be	neia on	tile 5	September,	via web	connenence.

# Pesticides Peer Review (PREV) Unit 4<sup>th</sup> Working Group meeting on Neurotoxicity

ersa European Food Safety AUTHORITY

23 June 2025 14:00-18:00 MINUTES - Agreed on 11 July 2025

Location: Webconference

#### Attendees:

#### Working Group Members:

Adeline Cavelier, Andrea Terron (Chair), Antonio Hernández-Jerez, Carsten Kneuer, Emily Mcvey, Kevin Crofton

Hearing Experts<sup>1</sup>:

None

Applicants participating in a technical hearing<sup>2</sup>:

None

o European Commission and/or Member States representatives:

None

#### o EFSA:

Pesticides Peer Review (PREV) Unit: Annetta Grillo, Anna Federica Castoldi, Martina Panzarea, Rafaela Rodrigues Jesus

#### Others:

Kristina Bartmann and Joerdis Klose participated as observers for agenda point 5, via web conference

Martin Paparella and Oddvar Myhre participated as observers for agenda point 6, via web conference

# I. Welcome and apologies for absence

The Chair welcomed the participants. Apologies were received from Helena Hogberg-Durdock.

# II. Adoption of agenda

The agenda was adopted without changes.

<sup>&</sup>lt;sup>1</sup> As defined in Article 34 of the document "Implementing Rule of the Management Board of the European Food Safety Authority laying down the rules on the selection, appointment and operations of the Scientific Committee, Scientific Panels and of their Working Groups": <a href="https://www.efsa.europa.eu/sites/default/files/paneloperation.pdf">https://www.efsa.europa.eu/sites/default/files/paneloperation.pdf</a>

<sup>&</sup>lt;sup>2</sup> As defined in the "EFSA's Catalogue of support initiatives during the life-cycle of applications for regulated products": https://efsa.onlinelibrary.wiley.com/doi/epdf/10.2903/sp.efsa.2021.EN-6472

### MEETING MINUTES - 23rd June 2025 4<sup>th</sup> Working Group meeting on Neurotoxicity



#### **Declarations of Interest of Working Groups members** III.

In accordance with EFSA's Policy on Independence<sup>3</sup> and the Decision of the Executive Director on Competing Interest Management<sup>4,</sup> EFSA screened the Annual Declarations of Interest filled out by the Working Group members invited to the present meeting and the preparatory meetings listed under agenda item "Preparatory meeting/s". No Conflicts of Interest related to the issues discussed in these meetings have been identified during the screening process, and no interests were declared orally by the members at the beginning of this meeting.

Andrea Terron indicated a new DOI was submitted and is current under-assessment. No interests were declared orally.

#### Agreement of the minutes of the 3<sup>rd</sup> Working Group IV. meeting held on 13-24th May 2025, via web-conference

The minutes of the 3<sup>rd</sup> Working Group meeting were agreed by written procedure on 28<sup>th</sup> May 2025.

#### V. Scientific topic(s) for discussion

#### a. Introduction of the 4th WG meeting

EFSA introduced the aim of the meeting and informed the WG members that the discussion will be captured and considered by EFSA during the assessment of the deliverables.

#### b. Deliverable 1a of LOT 2

An overview of the deliverable 1a of LOT 2 for the Environmental Neurotoxicants project was presented by EFSA to the WG members.

#### c. Deliverable 2 of LOT 4

An overview of the deliverable 2 of LOT 4 for the Environmental Neurotoxicants project was presented by EFSA to the WG members.

#### VI. **Any Other Business**

#### Chemical space analysis on the DNT IVB reference chemicals and pesticides

Carsten Kneuer informed the WG members about a work conducted by BfR on Chemical Space Analysis on the DNT IVB reference chemicals and pesticides. The WG agreed to include the analysis as topic of discussion in another WG meeting.

#### Update on LOT 3 of the Environmental Neurotoxicity project

The WG was informed on the status of the deliverable expected at the end of July 2025.

http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/policy\_independence.pdf

https://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/decision-ed-on-competing-interest-management-2024.pdf

# MEETING MINUTES – 23rd June 2025 4<sup>th</sup> Working Group meeting on Neurotoxicity



# VII. Preparatory Meeting

Since the last 3<sup>rd</sup> WG meeting, 1 preparatory meeting chaired by EFSA staff was held as follows:

- 1.1 Preparatory meeting to the 4th WG meeting on Neurotoxicity, 19th June
- **1.1a** Andrea Terron
- 1.1b Update on the LOT 1, 2, 3 and 4 of the Environmental Neurotoxicants project

### VIII. Next meeting

The next meeting will be held on 14th July 2025 via web conference.

# Pesticides Peer Review (PREV) Unit 3rd Working Group meeting on Neurotoxicity

efsa EUROPEAN FOOD SAFETY AUTHORITY

13 - 14 May 2025 14:00-18:00 / 14:00-18:00 MINUTES - Agreed on 28 May 2025

Location: Webconference

#### Attendees:

#### Working Group Members:

Adeline Cavelier, Andrea Terron (Chair), Antonio Hernández-Jerez, Carsten Kneuer, Emily Mcvey, Helena Hogberg-Durdock, Kevin Crofton

#### Hearing Experts<sup>1</sup>:

Bertrand Desprez (attended only 13th May)

European Commission and/or Member States representatives:
 Not applicable

#### o EFSA:

Pesticides Peer Review (PREV) Unit: Iris Mangas, Rafaela Rodrigues De Jesus Methodology and Scientific Support (MESE) Unit: Irene Cattaneo, Laura Martino

#### Others:

Not applicable

# Welcome and apologies for absence

The Chair welcomed the participants.

# II. Adoption of agenda

The agenda was adopted with changes. The first part of item 6 in the agenda was addressed prior to item 5.

# III. Declarations of Interest of Working Groups members

In accordance with EFSA's Policy on Independence<sup>2</sup> and the Decision of the Executive Director on Competing Interest Management<sup>3,</sup> EFSA screened the Annual Declarations of Interest filled out by the Working Group members invited to the present meeting. No Conflicts of Interest related to the issues discussed in this meeting have been identified during the screening process, and no interests were declared orally by the members at the beginning of this meeting.

<sup>&</sup>lt;sup>1</sup> As defined in Article 34 of the document "Implementing Rule of the Management Board of the European Food Safety Authority laying down the rules on the selection, appointment and operations of the Scientific Committee, Scientific Panels and of their Working Groups": <a href="https://www.efsa.europa.eu/sites/default/files/paneloperation.pdf">https://www.efsa.europa.eu/sites/default/files/paneloperation.pdf</a>

<sup>&</sup>lt;sup>2</sup> http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/policy\_independence.pdf

<sup>3</sup> http://www.efsa.europa.eu/sites/default/files/corporate publications/files/competing interest management 17.pdf



# IV. Agreement of the minutes of the 2<sup>nd</sup> Working Group meeting held on 23<sup>rd</sup> April 2025, in Parma and via web-conference

The minutes of the 2<sup>nd</sup> Working Group meeting were agreed by written procedure on 23<sup>rd</sup> April 2025.

#### V. Scientific topic(s) for discussion.

#### a. Introduction to the 3<sup>rd</sup> WG meeting

EFSA presented the aim for this meeting and the status of the IATA case study. The WG was informed that the submission deadline to the OECD was anticipated to be the end of May 2025, rather than the previously reported date of June 2025.

#### b. EKE of uncertainties for tebufenpyrad IATA for PoD

#### KE4

The WG was informed about the additional analysis performed by EFSA on the point of departure (PoD) derivation. It was agreed that these findings will be considered in the Expert Knowledge Elicitation (EKE).

The most probable lower concentration of tebufenpyrad that may affect neurite outgrowth in LUHMES cells, used as a proxy for key event (KE) 4 in AOP3, was estimated by EKE methodologies. This estimation considered the additional analysis mentioned above, the in vitro assays and modelling approaches used for dose-response translation and the associated uncertainties.

#### **KE1 and KE2**

This was not performed. The WG considered appropriate to perform the EKE only for KE4.

#### c. Proposed cluster of uncertainties, weight and impact

The WG discussed the evidence dossier, focussing on cluster 4, for the EKE and the feedback was incorporated into a draft document that will be submitted to the OECD by the end of May 2025.

#### d. Hearing expert view on proposed uncertainties and EKE results for cluster 2, 3 and 4

The hearing expert opinion on the uncertainties proposed by the WG was considered.

# e. Discussion and consensus on how to express the MoIE based on the uncertainty distribution

The WG discussed and agreed that the margin of internal exposure (MoIE) should be expressed as the probability of the MoIE being less than 10. If less than 10, additional indicators such as the probability of being less than 1 or the mean or median of the distribution were considered to be potentially useful.

### MEETING MINUTES – 13-14 May 2025 3<sup>rd</sup> Working Group meeting on Neurotoxicity



# f. Independent uncertainty analysis on tebufenpyrad combined uncertainty distribution for MoIE

The combined uncertainty distribution for MoIE of tebufenpyrad was calculated by EFSA and presented to the WG. In addition, based on the WG recommendation for future case studies, refinements to the external exposure estimation, the modelling of the resulting concentration in the brain, and/or the intracellular concentration in the in vitro test system should be considered, due to the high level of associated uncertainty for tebufenpyrad.

#### g. Actions and next steps Neurotox WG

The WG agreed to publish the tebufenpyrad IATA case study as an EFSA Scientific Report after the OECD commenting phase.

### VI. Any other business

Nothing was indicated during the meeting.

#### VII. Next meeting

The next meeting will be held on 23<sup>rd</sup> June 2025, via web conference.

# Pesticides Peer Review (PREV) Unit 2nd Working Group meeting on Neurotoxicity

3 - 4 April 2025 09:00-18:00 / 08:30-16:00

MINUTES - Agreed on 23 April 2025



**Location**: EFSA – Parma / Webconference

#### Attendees:

#### Working Group Members:

Adeline Cavelier, Andrea Terron (Chair), Antonio Hernández-Jerez (attended only 3<sup>rd</sup> April), Carsten Kneuer, Emily Mcvey, Kevin Crofton

#### o Hearing Experts¹:

Not applicable

#### European Commission and/or Member States representatives:

Not applicable

#### o EFSA:

Pesticides Peer Review (PREV) unit: Anna Lanzoni (attended only morning 3<sup>rd</sup> April), Iris Mangas, Jochem Louisse, Rafaela Rodrigues De Jesus

Methodology and Scientific Support (MESE) Unit: Alicia Paini, Irene Cattaneo (attended only afternoon 3<sup>rd</sup> April and full day on 4<sup>th</sup> April), Laura Martino (attended only afternoon 3<sup>rd</sup> April and full day on 4<sup>th</sup> April)

#### Others:

Not applicable

# I. Welcome and apologies for absence

The Chair welcomed the participants. Apologies were received for Helena Hogberg-Durdock absence. Apologies were received for Antonio absence on the 4<sup>th</sup> of April. Irene Cattaneo and Laura Martino did not participate in agenda points 1-6. Anna Lanzoni did not participate in agenda points 7-12.

# II. Adoption of agenda

The agenda was adopted without changes

# III. Declarations of Interest of Working Groups members

As defined in Article 34 of the document "Implementing Rule of the Management Board of the European Food Safety Authority laying down the rules on the selection, appointment and operations of the Scientific Committee, Scientific Panels and of their Working Groups": <a href="https://www.efsa.europa.eu/sites/default/files/paneloperation.pdf">https://www.efsa.europa.eu/sites/default/files/paneloperation.pdf</a>

### MEETING MINUTES – 3 - 4 April 2025 2<sup>nd</sup> Working Group meeting on Neurotoxicity



In accordance with EFSA's Policy on Independence<sup>2</sup> and the Decision of the Executive Director on Competing Interest Management<sup>3</sup>, EFSA screened the Annual Declarations of Interest filled out by the Working Group members invited to the present meeting and the preparatory meetings listed under agenda item "Preparatory meeting/s". No Conflicts of Interest related to the issues discussed in these meetings have been identified during the screening process, and no interests were declared orally by the members at the beginning of this meeting.

Carsten Kneuer indicated that Penconazole dossier was curated by BfR team as Co-RMS of which he is manager and declared that he did not directly participate in the RAR preparation.

# IV. Agreement of the minutes of the 1<sup>st</sup> Working Group meeting held on 4-5<sup>th</sup> March 2025, via web-conference

The minutes of the  $1^{st}$  Working Group meeting were agreed by written procedure on  $18^{th}$  March 2025.

### V. Scientific topic(s) for discussion

#### a. Advice on Penconazole (and triazoles) neurotoxicity

The available data for the Neurotoxicity (NT) assessment of the active substance (a.s.) was discussed and the WG advice was agreed and drafted during the WG meeting. This advice will be presented during the Peer Review Meeting of the a.s..

#### b. Tebufenpyrad AOP informed IATA, Introduction to the uncertainty analysis

The WG was informed of the proposed approach for the uncertainty analysis. This will include an Expert Knowledge Elicitation (EKE) using the roulette method to determine the margin of internal exposure (MoIE). All experts agreed with the approach proposed by EFSA.

#### c. Proposed cluster of uncertainties, weight and impact

The WG discussed the evidence dossier for the EKE and the feedback was incorporated in a draft document that will be submitted to OECD in June 2025.

#### d. EKE training

The WG was trained on the elicitation method selected for the EKE - Roulette Method, using an exemplary exercise.

#### e. Tebufenpyrad AOP information IATA: EKE on exposure and PBK modelling

The highest tebufenpyrad concentration in the brain of operators with and without personal protection equipment (PPE) after life-time exposure was estimated by using a WG EKE methodology, taking into account the external exposure, the proposed PBK modelling and uncertainties therein.

http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/policy\_independence.pdf

<sup>&</sup>lt;sup>3</sup> http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/competing\_interest\_management\_17.pdf

# MEETING MINUTES – 3 - 4 April 2025 2<sup>nd</sup> Working Group meeting on Neurotoxicity



# **VI.** Any Other Business

Nothing was indicated during the meeting.

# VII. Next meeting

The next meeting will be held on  $13^{\text{th}}$ - $14^{\text{th}}$  May 2025, via web-conference.

# Pesticides Peer Review (PREV) Unit 1st Working Group meeting on Neurotoxicity



4 and 5 March 2025 14:00-18:00 / 14:00-18:00 MINUTES - Agreed on 18 March 2025

Location: EFSA - Parma / Webconference

#### Attendees:

#### O Working Group Members:

Adeline Cavelier, Andrea Terron (Chair), Antonio Hernández-Jerez, Carsten Kneuer, Emily Mcvey, Kevin Crofton

#### Hearing Experts<sup>1</sup>:

Bertrand Desprez (attended only on March 5<sup>th</sup>), Helena Hogberg-Durdock, Jerome Henri (attended only on March 5<sup>th</sup>)

#### European Commission and/or Member States representatives:

Not applicable

#### o EFSA:

Pesticides Peer Review (PREV) unit: Iris Mangas, Rafaela Rodrigues De Jesus

Methodology and Scientific Support (MESE) Unit: Irene Cattaneo (attended only on March 5<sup>th</sup>), Laura Martino (attended only on March 5<sup>th</sup>)

#### o Others:

Not applicable

# Welcome and apologies for absence

The Chair welcomed the participants.

Hearing experts, Bertrand Desprez and Jerome Henri from ANSES and Irene Cattaneo and Laura Martino from EFSA MESE Unit, did not participate the meeting on Day 1 (discussion points a, b and c - agenda points V, VII and VIII).

# II. Adoption of agenda

The agenda was adopted without changes.

# III. Declarations of Interest of Working Groups members

In accordance with EFSA's Policy on Independence<sup>2</sup> and the Decision of the Executive Director on Competing Interest Management<sup>3</sup>, EFSA screened the Annual Declarations of Interest filled

As defined in Article 34 of the Decision of the Executive Director concerning the selection of members of the Scientific Committee, the Scientific Panels, and the selection of external experts to assist EFSA with its scientific work: <a href="http://www.efsa.europa.eu/en/keydocs/docs/expertselection.pdf">http://www.efsa.europa.eu/en/keydocs/docs/expertselection.pdf</a>

<sup>&</sup>lt;sup>2</sup> http://www.efsa.europa.eu/sites/default/files/corporate\_publications/files/policy\_independence.pdf

<sup>3</sup> http://www.efsa.europa.eu/sites/default/files/corporate publications/files/competing interest management 17.pdf

# MEETING MINUTES - 4-5 March 2025 1st Working Group meeting on Neurotoxicity



out by the Working Group members invited to the present meeting and the preparatory meetings listed under agenda item "Preparatory meeting/s". No Conflicts of Interest related to the issues discussed in these meetings have been identified during the screening process, and no interests were declared orally by the members at the beginning of this meeting.

# IV. Agreement of the minutes of the 1<sup>st</sup> Working Group meeting held on 4<sup>th</sup>-5<sup>th</sup> March 2025 via web-conference

The minutes of the 1<sup>st</sup> Working Group meeting were agreed by written procedure on 18 March 2025.

### V. Scientific topic(s) for discussion

#### a. Presentation to the WG, ToRs and workflow

EFSA presented the current neurotoxicity (NT) and developmental neurotoxicity (DNT) projects – In particular, the IRMA programme projects "Environmental Neurotoxicants" and "Brain Health", along with their tender specifications and status were presented to the WG.

The terms of reference (ToRs) and preliminary plan for the timeline of the WG and were agreed by the WG.

# b. Hearing expert review on the current protocol in Toxtemp format for Mitochondrial complex inhibition assay

The WG discussed a preliminary deliverable of the Environmental Neurotoxicants project—"Screening of pesticides for parkinsonian motor deficits", with emphasis on experimental critical points. The discussion was captured during the meeting to be shared later with the contractor in a meeting with EFSA.

c. The DNT Compendium list, the DNT Reference Chemical list and the Brain Weight recent publications (<u>EFSA-Q-2024-00685</u>, <u>EFSA-Q-2024-00684</u>, <u>EFSA-Q-2024-00634</u>). Discussion/Possible endorsement by the WG of the DNT Reference Chemical list.

The WG was updated on the 2024 EFSA publications in the DNT area. The WG agreed that the published DNT Reference Chemical list represents the best available list of DNT substances with available human and in vivo rodents' data. The WG agreed on using this list for the DNT in vivo-in vitro concordance analysis.

#### d. Tebufenpyrad IATA project overview and Neurotoxicity WG

The EFSA case study on Tebufenpyrad was introduced by the Chair. This project aimed at identifying the neurotoxicity potential of this pesticide by using NAMs contextualized in an AOP informed IATA framework. The WG will perform an uncertainty analysis to conclude on the selected in vitro point of departure (aka reference point) and on the internal margin of safety.

# MEETING MINUTES - 4-5 March 2025 1st Working Group meeting on Neurotoxicity



e. EFSA Pilot Project on New Approach Methodologies (NAMs) for Tebufenpyrad Risk Assessment. Part 2. Hazard characterization and identification of the reference point (<u>EFSA-Q-2022-00875</u>)

The EFSA pilot project on NAMs for Tebufenpyrad Risk Assessment Part 2. report, available in the EFSA journal, was presented and discussed with the aim to map out the already identified uncertainties and to uncover new ones identified by the WG members.

1.1. EFSA Pilot Project on New Approach Methodologies (NAMs) for Tebufenpyrad Risk Assessment. Part 1. Development of a Physiologically based Kinetic (PBK) Model Coupled with Pulmonary and Dermal Exposure (EFSA-Q-2022-00872)

The EFSA pilot project on NAMs for Tebufenpyrad Risk Assessment Part 1. report, available in the EFSA journal, was presented and discussed with the aim to map out the already identified uncertainties and to uncover new ones identified by the WG members.

#### VI. Any Other Business

Nothing was indicated during the meeting.

### VII. Next meeting

The next meeting (2<sup>nd</sup> WG meeting) will be held on the 3<sup>rd</sup> and 4<sup>th</sup> of April 2025, in Parma.