

**SCIENTIFIC PANEL ON
Plant Protection Products and their Residues (PPR)
3rd Working Group meeting on QIVIVE DNT IVB**



09 October 2024

13:00-18:00

MINUTES - Agreed on 25 October 2024

Location: Webconference

Attendees:

- Working Group Members:
Keyvin Darney, Antonio Hernández Jerez, Francesca Metruccio, Nynke Kramer, Emanuela Testai
- EFSA:
Jochem Lousse (PREV; chair), Alicia Paini (MESE)

I. Welcome and apologies for absence

The Chair welcomed the participants.
Apologies were received from Andrea Terron (PREV), Iris Mangas (PREV).

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 Independence¹ and the Decision of the Executive Director on Competing Interest Management², 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.

With regard to this meeting, Antonio Hernández Jerez declared the following interest: participation to an international expert group of the Danish Ministry of Environment and Gender Equality aimed to evaluate if the parametric value for nitrate in drinking water is sufficiently protective for the public with respect to the risk of developing colorectal cancer. In accordance with EFSA's Policy on Independence and the Decision of the Executive Director on Competing Interest Management and taking into account the specific matters discussed at the meeting in question, the interest above was not deemed to represent a Conflict of Interest for the expert concerned.

IV. Scientific topics for discussion ([EFSA-Q-2024-00299](#))

a. Cases literature QIVIVE DNT

Two case studies described in the literature on the QIVIVE of in vitro DNT data were presented. Critical aspects relevant for regulatory application were discussed.

b. Methods input parameters

An initial draft overview of methods to obtain chemical-specific model parameters was presented and discussed. Further information will be collected (point d).

¹ http://www.efsa.europa.eu/sites/default/files/corporate_publications/files/policy_independence.pdf

² http://www.efsa.europa.eu/sites/default/files/corporate_publications/files/competing_interest_management_17.pdf



c. In vitro distribution kinetics - chemical characteristics

In silico predictions of in vitro distribution kinetics in test systems of the DNT IVB were presented and discussed. Although currently available methods have large uncertainties (given the lack of test system specific information needs as input for the in silico models), insight can be obtained in the relation of chemical characteristics and expected in vitro biokinetics (and related in vitro exposure).

d. Discussion next steps and task division

Further steps were identified and tasks for the next meeting were divided among the WG members.

V. Next meeting

The next meeting will be held on 15 November 2024 (8.30-13.30), via web conference.

**SCIENTIFIC PANEL ON
Plant Protection Products and their Residues (PPR)
2nd Working Group meeting on QIVIVE DNT IVB**



05 September 2024
13:00-18:00 / 09:00-13:00
MINUTES - Agreed on 25 September 2024

Location: Webconference

Attendees:

- Working Group Members:
Keyvin Darney, Antonio Hernández Jerez, Nynke Kramer
- EFSA:
Jean-Lou Dorne (MESE), Jochem Louisse (PREV; chair), Alicia Paini (MESE)

I. Welcome and apologies for absence

The Chair welcomed the participants.
Apologies were received from Francesca Metruccio, Emanuela Testai, Andrea Terron (PREV), Iris Mangas (PREV).

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 Independence¹ and the Decision of the Executive Director on Competing Interest Management², 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. Scientific topic(s) for discussion ([EFSA-Q-2024-00299](#))

a. Finalisation protocol and interpretation of the ToRs

The updated protocol, including the assessment questions, was discussed and agreed.

b. Previous EFSA evaluations QIVIVE DNT IVB

DNT IVB QIVIVE cases previously evaluated by EFSA were presented and discussed.

c. PBK model parameters and methods

An overview on chemical-specific input parameters for PBK models was presented and discussed. Methods used to obtain values for such parameters will be identified.

d. Structure Scientific Opinion and task division

The first draft structure of the Scientific Opinion was discussed and tasks were allocated to EFSA staff and WG members.

¹ http://www.efsa.europa.eu/sites/default/files/corporate_publications/files/policy_independence.pdf

² http://www.efsa.europa.eu/sites/default/files/corporate_publications/files/competing_interest_management_17.pdf



V. Next meeting

The next meeting will be held on 9 October 2024 (13.00-18.00), via web conference.

**SCIENTIFIC PANEL ON
Plant Protection Products and their Residues (PPR)
1st Working Group meeting on QIVIVE DNT IVB**



04-05 July 2024
09:00-18:00 / 09:00-13:00
MINUTES - Agreed on 24 July 2024

Location: EFSA - Parma (Meeting Room 00/06b)/Webconference

Attendees:

- Working Group Members:
Keyvin Darney, Antonio Hernández Jerez, Nynke Kramer, Francesca Metruccio, Emanuela Testai
- Hearing Experts¹:
Ellen Fritsche (present only on 4 July)
- EFSA:
Jean-Lou Dorne (MESE), Jochem Louisse (PREV; chair), Alicia Paini (MESE)

I. Welcome and apologies for absence

The Chair welcomed the participants.
Apologies were received from Andrea Terron (PREV) and Iris Mangas (PREV).
Jean-Lou Dorne (MESE) participated online.
Ellen Fritsche participated as a hearing expert on the first day of the meeting.

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. 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. Introduction, ToRs and interpretation of the ToRs

Background information on the DNT IVB was presented. The terms of reference and interpretation of terms of reference were discussed. Based on the discussion, the text on the interpretation of the terms of reference will be updated by EFSA staff to be evaluated by WG members.

V. Scientific topic(s) for discussion ([EFSA-Q-2024-00299](#))

a. Protocol for the assessment, translation ToRs into assessment questions

¹ As defined in Article 17 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: <http://www.efsa.europa.eu/en/keydocs/docs/expertselection.pdf>

² http://www.efsa.europa.eu/sites/default/files/corporate_publications/files/policy_independence.pdf

³ http://www.efsa.europa.eu/sites/default/files/corporate_publications/files/competing_interest_management_17.pdf



The draft protocol, including the assessment questions, were discussed and adjustments were proposed. Based on the discussion, the assessment questions will be updated by EFSA staff to be evaluated by WG members.

b. Overview DNT IVB assays

The hearing expert provided a detailed overview of the 17 test methods that comprise the DNT IVB.

c. Considerations PBK model - input parameters

An overview was presented on PBK models and their input parameters of relevance for QIVIVE of DNT IVB data, and critical aspects were discussed.

d. Considerations in vitro/in vivo TK studies

An overview was presented on toxicokinetic studies for derivation of chemical-specific input parameter values, and critical aspects were discussed.

e. Considerations in vitro biokinetics/dose metrics

An overview was presented on the importance of consideration of in vitro chemical distribution kinetics for determining the in vitro dose metric to be applied for the QIVIVE.

f. Considerations on analytical methods for human biomonitoring (HBM) of pesticides

Principles of analytical methods and the importance of quality criteria relevant for analysis of HBM samples, but also for other kinetic studies were presented, and critical aspects were discussed.

g. In vitro and in vivo TK data in pesticide dossiers

An overview was presented on the typical in vitro and in vivo ADME/TK data available in the pesticide dossiers, and their possible use for PBK model parameterisation were discussed.

h. Previous EFSA evaluations QIVIVE DNT IVB

Due to time constraints, this agenda item has moved to the next WG meeting.

i. Definition and division of tasks, timeline of work

The draft timeline was presented and discussed. Based on the discussion the meeting schedule will be updated by EFSA staff. Tasks were defined for the next WG meeting.

VI. Next meeting

The next meeting will be held on 5 September 2024 (13.00-18.00), via web conference.
