WORK PROGRAM

FOR THE EU REFERENCE LABORATORY FOR PESTICIDE RESIDUES REQUIRING SINGLE RESIDUE METHODS

Time: January-December 2015

LEGAL FUNCTIONS AND DUTIES

The functions and duties of the EU Reference Laboratory are described in Article 32 of the EC Regulation No 882/2004.

Contents:

- A. General tasks
- B. Development and validation of analytical methods
- C. Quality assurance and quality control including the organisation and implementation of proficiency tests
- D. Technical and scientific support to NRLs, OfLs and third country labs

A. General Tasks

A.1 EURL coordination

Tasks: Inter-EURL-meetings (eligible as missions), in some cases in presence of DG-SANCO representatives, will be carried out with the aim to discuss, plan, coordinate or evaluate EURL-activities such as the preparation of work programs, EUPTs or web-applications. In certain cases online-meetings or tele-conferences will be carried out. Date and place of these events will be decided later.

Period: To be decided later following consultations with the other EURLs and/or DG-SANCO. **Deliverables:** Minutes of meetings prepared by one of the participating EURLs

A.2 Preparation of annual (financial, technical and PI ex-post) reports for 2014, Preparation of and negotiations for fixing the work program and PI exante table for 2016

Period: March 2015 and July-December 2015

A.3 Cooperation with International Organizations

Tasks: Cooperation with Int. organizations such as EFSA, the Codex Committee on Pesticide Residues (CCPR), CEN, FAO/IAEA will continue as necessities arise. If requested by DG-SANCO, documents will be revised or missions to attend meetings will be conducted.

A.4 Joint EURL-Website: upgrading, maintenance, programming (HORIZONTAL task for the benefit of all 4 EURLs) Tasks/Period/Deliverables: see below

Background: Following an agreement between the COM and the other 3 EURLs on pesticides the EURL-SRM has introduced a Joint EURL-Website for the four pesticide EURLs (www.eurl-pesticides.eu). The Joint EURL-Website aims to facilitate the dissemination of information to NRLs and OfLs in an efficient, timely and transparent way. It consists of a joint portal-website that is administered by the EURL-SRM as well as by 4 individual websites that are administered by the 4 respective EURLs.

Tasks: In 2015 the joint portal-website and the individual web-sites of the EURLs will be further upgraded, gradually filled with new information and existing links, overview-sites as well as documents will be updated. Missing features will be gradually implemented considering the needs and suggestions by DG-SANCO, the 4 EURLs and the lab-Network.

Period: Throughout 2015

A.5 EURL-DataPool Service

Background: Following an agreement between the COM and the 4 EURLs, the EURL-SRM has installed an "**EURL-DataPool Service**" entailing numerous interlinked databases with information of practical interest to the network-laboratories (<u>www.eurl-pesticides-datapool.eu</u>).

The EURL-DataPool is administered by the EURL-SRM as a horizontal activity and aims to store information about the entire network of laboratories working in the area of pesticides, to illustrate, and at the same time, strengthen the laboratory network, to facilitate the conservation of knowledge about pesticides, and to offer COM, EFSA, EURLs, NRLs and OfLs fast access to valuable information that can be used to assist decision-making and strategic planning. Special focus is being placed on the generation, collection and evaluation of experimentally-obtained data generated by various laboratories including the EURLs (e.g. MS/MS-transitions; validation data, stability data of compounds).

Tasks / Deliverables: see table below;

Period: throughout 2015 at www.eurl-pesticides-test.eu and/or www.eurl-pesticides-test.eu

List of databases within the EURL-DataPool-Service

Database/Website	Task	Deliverables	Examples where DB is		
EUDI D . D .	T ((DD ()))		used for/interlinked with		
EURL-DataPool	Transfer of DBs (see below) from www.eurl-pesticides-test.eu (based on the state-of-the-art .NET 4.0-Framework). This activity involves extensive redesign and programming of databases and web-applications.	Gradual upgrades and updates throughout 2015 and beyond	See below		
Method Validation DB	Data collection on experimental details and recovery rates achieved from various labs using various methods (e.g. QuEChERS, QuPPe, QuOil, SweEt) and its import into the DB. This includes data submitted by EURL-SRM, -FV, -CF and -AO.	Gradual updates/upgrades throughout 2015 and beyond	a) "Art. 12" activities,b) Pesticides DB,c) Pesticide Ranking List (PeRL)		
Analytical Methods DB	Data collection on various methods and its import into the DB (needed in the background for Stability DB, Method Validation DB)	Gradual updates/upgrades throughout 2015 and beyond	Method Validation DB		
Pesticides DB	Generation or collection of further data for the characterization of pesticides (e.g. GC-, LC-amenability, analytical behavior, GC-MS-spectra, GC-MS/MS-transitions; exact masses, solubility in acetonitrile) and import into the DB. This includes the creation of new entries for pesticides and metabolites not yet in the DB.	Gradual updates/upgrades throughout 2015 and beyond	 a) "Art. 12" activities, b) Pesticide Ranking List (PeRL), c) Method Validation DB, d) Stability of Compounds DB 		
Stability of	Collection of more stability data on	Gradual updates/upgrades	Pesticides DB		
Compounds DB Pesticide Authorizations DB	pesticides/metabolites and import into DB Data collection and updating as well as import into the DB of information about the authorization of pesticides in the EU and some third countries	throughout 2015 and beyond Gradual updates/upgrades throughout 2015 and beyond	Pesticide Ranking List (PeRL)		
Commodities DB	Data collection on commodities and import into the DB	Gradual updates/upgrades throughout 2015 and beyond	Validation DB		
Lab-Network DB	Permanent updating of lab-specific information (lab contact data, lab-functions, fields of work, email-addresses of contact persons, instrumentation available, tasks within the frame of official controls (import controls, scope, etc.)	- up-to-date email-lists of lab contact persons of all NRLs and OfLs - only for EURL-members: lab-list reflecting the individual EURL-lab-network; - only for NRL-members: lab-list reflecting the individual NRL-lab-network for each country - export-function (csv-format) of network-related data (e.g. lab-functions; emails of contact persons) - "List of obliged labs" for participation in EUPTs in 2015	Convenient compilation of customized lab-contact lists (e.g. for invitations to EURL-activities such as EUPTs, Surveys, for the dissemination of information). Compilation of the "List of obliged labs" for participation in EUPTs		
EUPT-Archive DB	Import of EUPT-data (FV16; CF8; AO9; SRM9) into the DB; design and programming of new online views for EURLs and NRLs allowing them to view the EUPT results of the labs within their network. Evaluations, that are necessary for the planning of future PTs or the establishment of general evaluation rules, will be discussed and elaborated in close cooperation with the other 3 EURLs.	Gradual updates/upgrades throughout 2015 and beyond, new online views	Evaluation of long-term PT- performance of labs or countries, "SRM-PinBoard list"		
MRL Residue Definitions DB	- Updating of EU and Codex MRL residue definitions; - Updating of conversion factors within DB	Gradual updates/upgrades throughout 2015 and beyond	a) "Art. 12" activities, b) Pesticide Ranking List (PeRL), c) Pesticides DB		

B. Development and Validation of Analytical Methods

B.1 Quick Polar Pesticides Method (QuPPe Method)

Background: The EURL-SRM has developed a method for the simultaneous analysis of several highly polar pesticides not amenable to multiresidue procedures. The method, which is employed by various OfLs involves a common extraction followed by LC-MS/MS analysis.

Task: Further method development activities will be conducted with the aim to expand, where possible, the scope of the method by additional highly polar pesticides and metabolites such as e.g. Bromide, Propamocarb-N-desmethyl, Propamocarb-N-oxide. Main focus will be to find ways to improve chromatographic behavior for some compounds already in the method or to accommodate as many compounds as possible within the same LC-runs. Alternative LC-separation approaches will be tested. Activities to pursue CEN method standardization will continue.

Period: throughout the year

B.2 Solutions for pesticides requiring modified MRMs

Background: Several pesticides and legally relevant metabolites are known to pose problems in analysis and are thus considered "difficult" or non-amenable to multiresidue methods. In many cases analysis is possible following certain modifications of traditional multiresidue methods. Such modifications may entail adjustment of pH or addition of chemicals to prevent degradation or improve extractability, temperature control, special measurement conditions or cleavage reactions to release conjugates etc..

Task: Studies will be conducted to improve the analysis of captan, folpet, dichlofluanid, tolylfluanid and dicofol in commodities of plant origin (in the light of the new residue definitions including the metabolites) as well as of ethoxyquin in commodities of animal origin considering its metabolites as far as they are available.

Period: throughout the year

¹⁾Note/disclaimer: This activity may be shifted to 2016 in case of insufficient budget

B.3 Studies on the analysis of triazole metabolites in fruits and vegetables

Background: Triazole-pesticides are one of the largest groups of pesticides entailing more than 30 compounds. These pesticides have in common that they can metabolize to the following 1,2,4-triazole derivatives:1,2,4-triazole (T), triazole alanine (TA), triazolyl acetic acid (TAA) and triazolyl lactic acid. These metabolites are of high interest for risk assessment and EFSA repeatedly requested their inclusion in the EU-multiannual Control Program (MACP). Their analysis is however extremely difficult due to their high polarity and the very low molecular weight.

Task: The EURL-SRM has established a sensitive method based on ion-mobility mechanisms in 2014. In 2015 further samples will be analyzed to check for residues, especially for the newly introduced triazolyl lactic acid.

Period: throughout the year

B.4 Further studies on the analysis of selected volatile pesticides in F&V by QuMFu1)

Background: Various highly volatile pesticides are employed in agriculture and crop storage as fumigants. Such pesticides are typically non-amenable to multiresidue methods and they are thus rarely analyzed by official control laboratories. Simple methodologies are thus required. In 2013 and 2014 the EURL-SRM has established a method for phosphine by applying Headspace-GC-MSD and a <u>quick multi-method</u> for <u>fumigants</u> (QuMFu) for the simultaneous analysis of 15 different fumigants in cereals and dried fruits by applying GC-MS/MS (see EURL-Portal-website for reports).

Task: The EURL-SRM will conduct experiments to check the influence of sample processing on fumigant losses and if an up-scaling of the QuMFu method is necessary when applied to fruits and vegetables (e.g. increase of the sample weight from 10 g to 50 g). The resulting method will then be validated for fruits and vegetables. Priority will be given to commodities which are included in the EU coordinated control program (e.g. bananas), several fruits and vegetable samples will be analyzed by the developed method.

Period: throughout the year

B.5 Testing the stability of pesticides in stock solutions by quantitative NMR (qNMR)

Background: Checking the purity and stability of neat standards and stock solutions is time consuming and expensive. There is a need to investigate alternatives to the current approaches (mass spectrometry techniques) to simplify and speed-up work in the future. NMR technology has experienced major advancements in recent years and can be applied for purity assessment of neat standards and used to characterize their compositions. Quantitative proton NMR (qNMR) has been gaining popularity e.g. in quantitative drug analysis and quality control as it produces qualitative and quantitative information simultaneously. One major advantage is that the reference standard does not need to be the identical material, but just one universal standard unrelated to the target analyte. This considerably reduces the costs and logistic efforts of a pesticide laboratory. Additionally, qNMR has been reported to be highly precise and accurate, thus reducing the number of replicate measurements required. The non-destructive nature of this technique makes it possible that the same samples are kept for measurements over the course of several years, e.g. in flame sealed NMR tubes.

Task: The EURL-SRM will perform experiments by qNMR to check the purity and/or stability of stock solutions (solvent: acetonitrile) over a time course of 5 years for 50 pesticides. The qNMR-measurements will be performed at least twice a year.

Period: throughout the year

¹⁾ Note/disclaimer: This activity may be shifted to 2016 in case of budget cuts

B.6 Experiments to be conducted within the frame of the "Art 12"

Background: One of the highest priorities of DG-SANCO is the re-evaluation of MRLs and residue definitions within the frame of Art. 12 / Reg. 396/2005. In 2015 the EURLs will be frequently consulted to evaluate pesticides under review in order to express their views as regards LOQs and residue definitions. Circa 50 pesticides will be expectedly reviewed in 2015 by EFSA. In cases where analytical data is not sufficient or not available it has to be generated as far as practicable. Where residue definitions contain metabolites, they often require SRM-methods, thus falling within the responsibility of the EURL-SRM.

Where no standards of pesticides or metabolites are available, they have to be requested from pesticide companies.

Task: Based on the experiences acquired in Art. 12 evaluations from 2013 and 2014 H1 metabolites of pesticides, which are part of residue definitions are in many cases SRM-compounds. The EURL-SRM will order missing analytical standards of pesticides and metabolites and conduct experiments to check their analytical behavior, the MRM amenability and the chromatographic behavior, perform basic validation experiments and determine the achievable LOQs always taking into account the proposed residue definition and the capabilities of OfLs.

Estimated man-days for activity B.6:

	Expected	Lab activities involved					
Type of compound	No. of	NONE	SOME	EXTENSIVE	VERY EXTENSIVE	Sum (Working	
Estimated man-days for 10 compounds	compounds	0	15	25	80	days)	
requiring NO Lab Activities	25					0	
requiring SOME Lab Activities*	10		15			15	
requiring EXTENSIVE Lab Activities**	10			25		25	
requiring VERY EXTENSIVE Lab Activities***	5				40	40	
	<u> </u>			•	SUM	80	

^{*} e.g. for analytes requiring minor modifications of MRM-methods with few matrix groups being involved

Period: throughout the year

C. QA/QC (Quality Assurance and Quality Control)

C.1 QA/QC

Task: Jointly with the other EURLs the EURL-SRM will, assist the Commission by revising the Document N° SANCO/12571/2013 der EU, implemented by 01/01/2014

Period: Q1-Q3 in 2015

^{**} e.g. for analytes requiring minor modifications of MRM-methods with <u>many</u> matrix groups being involved OR non-MRM-amenable analytes (parent or metabolites) with <u>few</u> matrix groups being involved

^{***} e.g. for challenging non-MRM-amenable analytes (parent or metabolites), with many matrix groups being involved

C.2 Preparation of a list showing which labs are obliged to participate in EUPTs in 2015 (horizontal task for the benefit of all 4 EURLs)

Task: based on the information provided by the NRLs, concerning the commodity scope covered by each lab within its network a comprehensive list of all labs obliged to participate in pesticide-related EUPTs in 2015 will be published. Prior to generating this list the NRLs will be asked to update the commodity-scope profiles of the labs within their network and the EURL-DataPool will be updated accordingly. This list will be distributed to all NRLs which are responsible to check its correctness/completeness and report any errors. The pesticide scope of the labs will not be taken into account in this list. Labs not participating in an EUPT, due to limitations in the pesticide scope will have to give explanations during the registration period of the respective EUPT.

Period: Early in 2015

C.3 EU Proficiency Test SRM 10 (in cooperation with EURL-CF)

Task: A proficiency test covering single residue methods (SRM) will be performed. The intention is to use a commodity of plant origin that is also relevant as feed. Maize has been chosen as commodity, and this PT will be conducted in collaboration with the EURL-CF. In case of Ad-hoc needs to focus on a different commodity, the commodity might have to be changed following consultation with DG-SANCO.

All relevant documents and instructions will be distributed to the participants through the EURL website. Participant registration and data collection will be conducted using an online tool provided by the EURL-CF. Each participant will receive a detailed report summarizing the PT-scope, results, data treatment and additional information of the methods employed by the participants.

Prior to, during and after the EUPT, the EURL-SRM will furthermore address any PT-related requests of participating labs. Underperforming NRLs will be directly assisted and at request provided with information regarding OfLs within their network.

Certain tasks associated with the EUPT-SRM10 will or may be subcontracted to other parties e.g. purchase and preparation of test material, standards, shipping material and dry ice.

Period: H1 of 2015

C.4 Attend joint meeting to discuss and evaluate EUPT results, Establish criteria to assess PT-underperformance and overall PT-performance (horizontal activity) **Background:** EUPTs are a very valuable tool to assess the performance of laboratories in pesticide residue analysis. In the case of individual PT-results underperformance is well defined by the absolute z-score. When looking at multiple results generated within one PT or throughout many PTs, suitable underperformance criteria are yet to be set, considering both analytical results and scope. In case of underperformance corrective and follow-up actions are indicated.

Task: In cooperation with the other three EURLs the criteria to evaluate whether a laboratory was underperforming in EUPTs will be further revised.

Period: H2 of 2015

C.5 Update the "SRM-Pinboard" and promote concept of sub-contracting analyses within the Lab-Network

Background: Within the frame of official controls, SRM analytes are less frequently analyzed compared to MRM analytes. OfLs often complain that limitations in the available resources prevent them from establishing suitable methods for the analysis of SRM-analytes or applying such methods in case they are established. Lab-cooperation and subcontracting of analyses will help to reduce the overall number of labs, that will have to establish or apply SRMs thus improving overall efficiency and frequency of analysis of SRM compounds.

Task: Following the established performance criteria, the list of laboratories considered as proficient for the analysis of individual SRM-compounds will be updated ("SRM-Pinboard" = Pool of Proficient SRM-Labs) as soon as new PT results become officially available or whenever a lab wishes to enter the list or change its status. The updated list will be published in the EURL-portal.

Period: H2 of 2015

D. Technical and scientific support to DG-SANCO, EFSA, NRLs, OfLs and third country labs

The technical support to DG-SANCO and to EFSA (on behalf of DG-SANCO) is of high priority to the EURL-SRM and is among others accomplished through e-mail communication, participation in meetings, revision of documents, and drafting of opinions. Upon invitation EURL-SRM staff may participate in various meetings organized by COM or EFSA. NRLs, OfLs and third country labs are technically supported through EURL-Web-Portal and the EURL-DataPool (A.2). Further assistance is also provided via personal communication, presentations in conferences, workshops and trainings (see D.2). The comprehensive Lab-Network-DB further contributes in strengthening the network (see A.2).

D.1 Technical and scientific support to DG-SANCO and EFSA

Tasks/Period/Deliverables: See table below:

Task	Activities	When	Deliverables
Support DG-SANCO in	- Participate in meetings (e.g. in Brussels);	As	Prepare/Provide
drafting the MACP 2015-17	- Collect and evaluate data and prepare a new "Pesticide Priority	arranged/agreed	"Pesticide Priority List"
	List";	with or scheduled	opinions, assessments,
	- Read/Revise documents;	by DG-SANCO	advices and revisions as
	- Communicate with DG-SANCO, EFSA and other stakeholders	_	required
Support EFSA within the frame	- Participate in one or more meetings (e.g. in Parma);	As	Prepare/Provide
of the of the Networking	- Read/Revise documents;	arranged/agreed	opinions, assessments,
Group on Pesticide	- Communicate with DG-SANCO, EFSA and other stakeholders	with or scheduled	advices and revisions as
Monitoring		by EFSA	required
Support to EFSA and DG-	- Prepare an expected time schedule and coordinate the division	As	Prepare coordinated
SANCO in activities	of tasks among the 4 EURLs	arranged/agreed	positions of the 4 EURLs
concerning re-evaluation of	- Collect/Evaluate existing information from EFSA, DataPool and	with or scheduled	as regards the residue
pesticide MRLs according to	other sources;	by DG-SANCO	definitions and the LOQs
Article 10 and 12 of Reg.	- Compile the available data in a summarized form to facilitate	and EFSA	of pesticides proposed by
396/2005/EC* . 50 compounds	decision-making by the other 4 EURLs as regards LOQs and		EFSA and DG-SANCO.
will have to be expectedly	residue definitions;		
evaluated by the EURLs in	- Collect the positions of the 4 EURLs and finalize document with		Provide opinions,
2015 on behalf of EFSA, plus	joint position (coordinate among the 4 EURLs in case of		assessments, advices
some additional pesticides on	disagreements);		and revisions as required
behalf of DG-SANCO.	- Communicate with DG-SANCO, EFSA and other stakeholders		
General assistance to DG-	- E-mails and oral communications	Throughout 2015	Opinions, advices,
SANCO and EFSA	- Missions to Brussels, Parma or elsewhere	as required	revisions as required

*Note: This task does not include non-laboratory activities. These are covered by B.2.

D.2 Joint EURL Workshop for Pesticide Residues in Food and Feed

Task: A joint workshop (FV, CF, AO and SRM) will take place in Stuttgart in the second half of 2015. NRLs from all MS will be invited to attend, with the main objective to facilitate the interaction between them and the EURL-FV, -CF, -AO and -SRM. The workshop will be held during two days, and will entail technical and scientific communications regarding new activities of the EURLs and other developments in the field of pesticide residues analysis. The EURL-SRM will take over the full organization and administration of the workshop.

Period: Q3 2015

D.3 Trainings

Task: 8 NRLs will be invited to attend a training in Fellbach. The training will cover technical aspects as regards the analysis of SRM-pesticides and exchange of experiences. Special needs and problems of the laboratories selected to participate will be considered in the design of the training program. Additional ad-hoc trainings will be conducted as required.

Period: To be decided

D.4 Visit of 1 NRL

Task: The NRL-SRM of one selected country will be visited by representatives from the EURL-SRM. The country will be selected in agreement with DG-SANCO giving emphasis on poor EUPT scope, performance or participation over last years as well as on poor cooperation with the EURL.

Prior to the inspection a detailed study of the EUPT results during the last years as well as the current analytical scope of all OfLs will be carried out. During the visit the possible reasons for bad PT-performance will be discussed, where applicable and advices will be given to improve performance and expand the scope.

Period: To be decided

D.5 Webinars and Tutorials

Background: Webinars and Video-Tutorials provide the possibility to disseminate information to NRLs and OfLs in a cost effective way.

Task: In 2015 the EURL-SRM will organize/publish at least one webinar or one video-tutorial either individually or in collaboration with other EURLs.

Period: To be decided

D.6 Analysis of official samples, counter analysis (if required)

The EURL will ask DG-SANCO for approval of any activity this concerning and request for additional eligible budget, if required.

Task: Sample analyses if required and only after consultation with DG-SANCO

Period: unpredictable