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| <p align="center">"RESPONSE" project Steering Committee Meeting combined by dissemination activities</p> <p align="center">at the EAFS-2015 meeting, Prague, 6th-11th September 2015</p> | | | |
| DOCUMENT TYPE : | Minutes | ISSUE DATE | 03/10/2015 |

1. Welcome, discussion and acceptance of agenda – S. Klemenc and all

Meeting took place in Corinthia hotel in Prague, at 8th September from 11 a.m. to 13 p.m. Ms Klemenc welcomed participants and asked for amendments to the proposed agenda. None was proposed. Agenda was accepted.

Participants list:

N. Meert (INCC, Belgium)
M. Johannsen (IFM, Aarhus University, Denmark)
L. Ask Reitzel (University Copenhagen Denmark)
L. Dujourdy (INPS, France)
A. Stemmelen (INPS, France - invited by SK)
T. Csesztregi (HIFS, Hungary)
S. Klemenc (Mol, NFL, Slovenia)

2. Project progress monitoring (WP action list checked, intermediate evaluation, adjustments, future activities - (all)

Outcome:

| WS | Work stream description | Status/conclusions/evaluation |
|------|---|---|
| WS0: | Management and coordination activities | tasks planned so far, timely completed and records available |
| WS1: | State of the art and procurement <ul style="list-style-type: none"> activity 1.1 (M6-M9) - new equipment in NFL, installed, validated and staff trained activity 1.2 CRM materials (M1-M24) – activity 1.3 net-NPS(M1-M24) - purchasing 1.4 (M1-M24) – other purchases | activities planned so far, timely completed and records available; <ul style="list-style-type: none"> - Activity 1.1 (closed) - (records available) - 1.2 running around 40/ compounds from Chiron purchased/ records available; - 1.3 running - around 70 purchased; (records available) - 1.4 Done or running |
| WS2 | Profiling methodology and competencies <ul style="list-style-type: none"> activity 2.1 (M1-M3) - SI tem visit to Lyon for heroin profiling activity 2.2 (M3-M10) - development of experimental methodology (heroin) in SI activity 2.6 – 1st Profiling workshop in Ljubljana June 2015 – | <ul style="list-style-type: none"> - activity 2.1 timely completed; practical exercising of experts successful, creation of databases tool in SI still remains opened - 2.2 currently running (one moth of delay is expected) - 2.6 timely completed; evaluation from attendees: excellent |



| | | |
|------|-------------------------------------|--|
| WS3: | Chemical characterizations (M1-M24) | <ul style="list-style-type: none">- running, characterizations in NFL timely completed (records available)- NMR structure conformation/ elucidation too slow (no results from July to September / P. Manager acted – urged/ because this slow down the reporting significantly and proactive component of the project is endangered |
| WS 4 | NPS FTIR and MS databases [M13-M24] | <ul style="list-style-type: none">- Activities (development of IR database, searchable IR libraries, guidelines) running on timely manner, MS spectra exchange accordingly in line with plans |

3. Discussion on sample acquisition, purchases, exchange, analytical procedures (all)

- a) Recent developments and challenges concerning the net-NPS purchasing have been discussed.
- b) Concerning the analytical procedures: the idea to “upgrade” NMR measurements to q-NMR or to employ elemental analysis in addition to NMR (for checking net-NPS purity as a quality criteria of the IR result) has been explained and discussed in June 2015 in Ljubljana (Mr. Csesztregi (HU), Mr. Košmrlj (SI2) and Ms Klemenc (SI1)). However, the Slovenian partner FKTT (SI2) who has instrumental resources is currently obviously not able (see point 1, WS3) to take over additional tasks.

Conclusions/ outcome:

a)

- I was agreed that partners will enhance information and samples exchange of novel NPS detected. /responsible all/
- It was agreed that Denmark laboratory (Copenhagen), who developed the strategy for searching novel NPS over internet will send information on detected NPS to Slovenian NFL, which will then try to find appropriate internet vendor and purchase samples net-NPS. Other partners can participate on the same manner or also by sending some novel - NET samples collected/seized to NFL to be included into “material data bank”.
- NFL has to enhance the purchasing velocity of CRM materials within the forthcoming months, because the public procurement procedures are quite time consuming;
- it was agreed that each partner will send a wish list of 10 CRMs (if appropriate with data on vendor) to be purchased from the RESPONSE project budget (newly appearing CRMs are the target materials, but list is not strictly limited to these).
- Enhanced cooperation (knowledge, information and samples exchange) with SPICE2 project is planned (on non formal level).

b)

At the moment quantification of net-NPS active ingredients by means of q-NMR or elemental analysis is obviously not possible /see point 2. WS 3/, despite the fact that Mr. Csesztregi provided description of measurement procedure, test amounts of two internal standards and excel worksheet for calculations (for q-NMR). Along Mr. Košmrlj, elemental analysis could be preferred and less time consuming option, however the tasks planned in the frame of the project (NMR structure conformation/ elucidation/ are the highest priority.



4. Dissemination activities (all)

a) Activities taken so far (M1 – M9) can be *evaluated* as very good. Short summary of main deliverables:

| Event/deliverable type | Where – when | description | comments |
|---|---|--|---|
| Training | INPS-Lyon, France March 2015 | exchange of knowledge, skills and good practice and profiling methodology development on the expert exchanged basis (3SI experts in Lyon) (FR/SI) | Records available |
| Oral/poster presentations | ENFSI DWG meeting, Dublin; Ireland, May 2015 (around 80 participants from all over the world) | Abstracts and oral (2), poster (1) presentations + 2x info included into oral presentations given of DWG subcommittees (profiling and databases) | records available |
| Oral presentation | EMCDDA, Lisbona, Portugal, June 2015 (No: attendees 66) | Oral (1) at 15th Annual meeting of the Reitox Early-warning system network | Records available |
| Report for EMCDDA and EUROPOL | March, 2015 | 1 x (not public opened) | Record available |
| Workshop on profiling/ | NFL, Ljubljana, Slovenia June 2015 (18 trainees and 5 trainers) | Knowledge, competencies, skills and tools: Training & training materials, presentations, tool for sample comparison | Records available/ evaluation from attendees: excellent |
| RESPONSE project web page | “Drugs monographs» at http://www.policija.si/en/g/index.php/generalpolicedirectorate/1669 ; lunched in August 2015 | Interactive, searchable table with NPS analytical reports and related chemical data /knowledge and data/ | Records are added continuously |
| Reporting forms EMCDDA/ EUROPOL | NFL, Sent in the frame of SI EWS sistem (January-September, 2015) (M1-M9) | Reports (through REITOX and/or ENU networks) + analytical reports on newly detected NPS (seized, collected, purchased), where applicable and raw analytical data | Records available – continuous activity |
| Reports on “anonymously” tested samples | NFL, Ljubljana, Slovenia (M1-M9) | Rising awareness: Analytical reports on active ingredients (NPS) collected by NGOs from anonymous users | Records available - continuous activity |
| Poster presentations | EAFS 2015 meeting; Prague No of attendees >1000 | Abstracts and 3 x posters | Records available |
| Shared – “identification reference materials” | April, 2015 | The first set of newly detected NPS samples was offered and delivered to interested project partners (April-July 2015). | Records available |
| Shared MS and FTIR raw data | April, 2015 | first sets of MS and FTIR spectra have been delivered to database managers and implemented into libraries | Records available |
| NPS materials shared with SPICE- | August, 2015 | cannabinoids-pure forms, herbal preparations and C-liquids - around 40 | Records available |



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| profiling project (BKA, Germany) | | samples | |
|-------------------------------------|--|---------|--|

b) Dissemination planing

| Dessimation activities | Comments and responsible |
|--|--|
| Active participation at the next DWG meeting, 2016 (see point 5)/ oral poster presentations scientific [M17] | all |
| Draft versions (for comments from DWG) of two planned FTIR documents [M17] | /HU and BE/ |
| Organization of 2 nd profiling workshop [M18] see point 6 for details | FR/SI |
| Further exchange of MS and FTIR data and implementation into libraries | SI/HR/DK |
| Ms Ask Reitzel proposed writing of an article for Forensic Science International with regard to the experiences obtained from internet purchasing, where the rate of false advertised products pose serious risk for NPS users [time frame not defined – see comments] | The proposal will be considered further (SI/DK1 and other) |
| Continuation of some permanent dissemination activities | SI and all |
| Share identification reference materials from internet purchases by partners | SI |

5. Planning of the ENFSI DWG 2016 – meeting May (May 10 to May 12, 2016) combined by the RESPONSE project SC final meeting (Bled, Slovenia, May 8th-13th, 2016) (all)

Info: In the frame of the “RESPONSE” project EU-grant (*Prevention of and fight against crime Programme of the European Union (grant agreement number JUST/2013/ISEC/DRUGS/AG/6413)*) financial support for the accommodation and partial reimbursement of travel costs for 28 attendees from EU member states ENFSI DWG members will be offered. (RESPONSE project steering committee and NFL staff members are excluded from this number). Details will be outlined on the meeting web page.

Plan:

| Event/activity | Dates /responsibles |
|--|---|
| RESPONSE project SC meeting | Monday, May 9 th , from 9 p.m. to 17 p.m; and optionally additional one – at May 12 afternoon or May 13 morning) - all |
| ENFSI DWG meeting & RESPONSE project dissemination activities | May 10 to May 12, 2016 (working days) – all |
| Communications with ENFSI DWG steering committee and program subcommittee, drafting program, abstracts collection, dissemination of all relevant meeting information | Klemenc/activity running [M8-M18] |
| Meeting web page (materials for web prepared – page currently under construction), forms: registration, abstract submission, meeting evaluation. | Klemenc/activity running [M8-M18] |
| Early registration | until February 19 th , 2016 |
| Late registration | after February 19 th , 2016 |
| Abstract submission | until February 19 th , 2016 |
| “technical” issues in Bled, Slovenia: meeting rooms, accommodation, meeting attendance certificates, final financial statement etc. | NFL staff and Albatros agency – activities running [M4-M20] local organizing committee: responsables: Klemenc, Gostič, Benčina, Godeša/ |



6. Planning of the 2nd profiling workshop, Ljubljana 2016 (L. Dujourdy and S. Klemenc)

Outcome:

- Workshop is tentatively scheduled within the last two weeks of June, 2016
- registration deadline /will be defined/
- duration of the workshop (tentatively planned for a day longer than the previous one)
- Registration deadline: April 8th, 2016

Activities/Responsibilities:

| Activity | responsible |
|--|--|
| check the availability of trainers from INPS, France and two from Lausanne University, Switzerland | Dujourdy [as soon as possible] |
| check the availability of the trainer from INCC, Belgium | Meert [as soon as possible] |
| Define workshop exact dates | FR/SI/BE |
| send out invitation and preliminary information | Besacier [as soon as possible]/Klemenc |
| Final program - detailed | Trainers [FR/BE/CH]/ Besacier |
| Registration process, technical logistic | SI /Klemenc/NFL staff/ |

Note: Meeting is opened for all ENFSI-DWG members, however only participants [up to 15, domestic (SI) are excluded from this number] from EU member states are eligible for the support from the RESPONSE project budget.

7. Planning of heroin profiling training in Ljubljana and planning the cocaine profiling training visit in Lyon

Ms Klemenc explained some not foreseen and unexpected difficulties concerning analytical method development and validation in NFL laboratory. Therefore, the heroin training & evaluation planned for week 44 was postponed to week 47.

Plan (dates, activities, personal involved):

a) Heroin

| Activity | Time frame/ responsables |
|---|--|
| heroin methodology training and statistical evaluation, in Ljubljana (5 days) | at week 47 , 2015 (2 FR experts: Dujourdy and Charvoz in Ljubljana and NFL staff Janežič, Benčina, Gostič, Koren) |
| respond to Ms Dujourdy's questionnaire received at | NFL staff (Benčina, Janežič) shall send together with other info about the current status of method development /ASAP/ |
| detailed daily working plan shall be agreed between two parties | /FR/SI/ before the training & evaluation start (at latest at week 46, 2015). |

b) Cocaine

- January/February 2016, 2 SI experts (training in Lyon) – exact dates will be defined latter on in Ljubljana (at week 47)



8. Networking with other projects and labs (as for example, CLEAN2SAND, Spice-profiling and I-SEE (S. Klemenc and all)

RESPONSE project is opened for networking (information, knowledge and samples exchange). Good communication has already been established by projects listed above.

9. Administrative issues

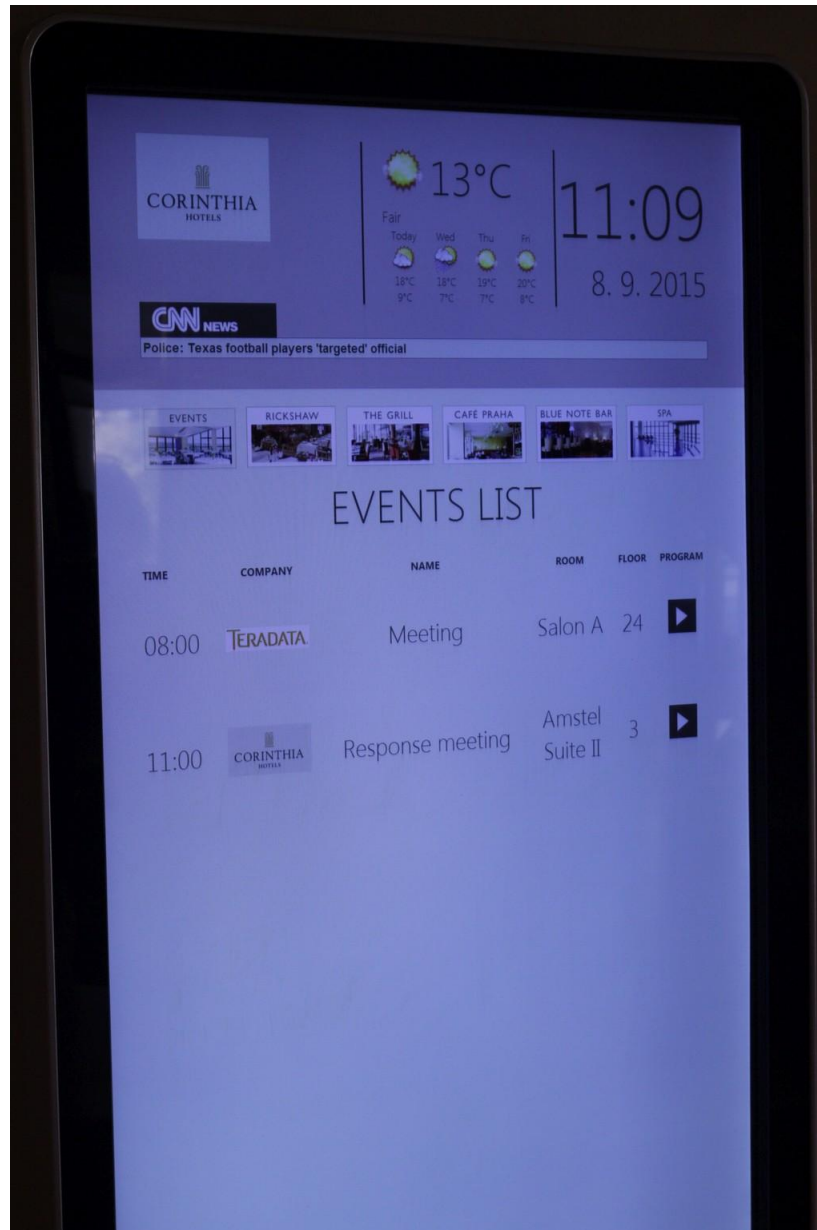
Ms Klemenc asked and reminded SC members to send reimbursement requests regularly (still missing some from already closed activities)! Please send reimbursement requests to Ms Lili Šipeč: lili.sipec@gov.si and Ms Melita Godeša: melita.godesa@gov.si, first by e-mail and after the approval originals by post mail.

10. Any other business
None.

Meeting was closed at 13 p.m.

Report prepared by:
Dr. Sonja Klemenc
PROJECT MANAGER

See some pictures bellow.





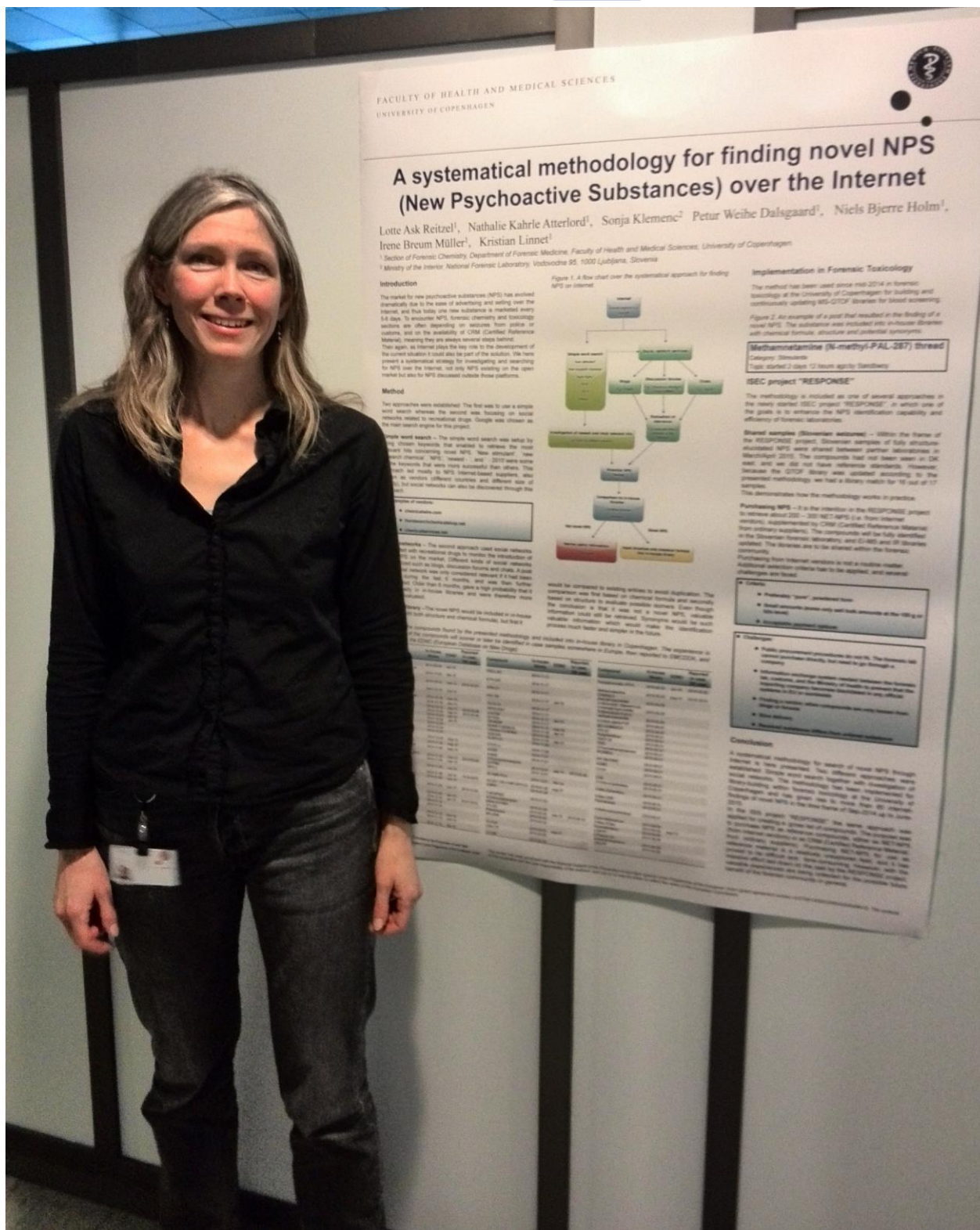
Steering committee meeting (photographer Tamas Csesztregi is missing on the picture)





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COLLECT, ANALYSE, ORGANIZE, EVALUATE, SHARE – A RESPONSE TO CHALLENGES IN FORENSIC DRUGS ANALYSES

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INTRODUCTION

Over the past few years Europe has seen an unexpected growth in the number, type and availability of new psychoactive substances (NPS). One of the main challenges to effectively respond to new psychoactive substances is the detection of these followed by the correct identification.

NPS are mainly detected in forensic and customs labs. The difficulty lies in identifying them. To identify new substance reference material (RM) or special techniques are needed. However, RM are mostly not available in real time. Therefore, most of laboratories have to rely on comparison of their results to available MS and/or FTIR spectra in searchable electronic libraries (not updated in real time) or existing information exchange channels and/or open source data.

This work presents the first impressions of a systematic (pro-active) tackling and identification of the new psychoactive substances (NPS) as applied in the frame of EU co-funded ISEC project "RESPONSE". The project is divided into several modules (see Table 1).

Table 1: Main response project modules

| Activity | Issue |
|-----------------------------|--|
| Collect/purchase NPS | Setting up a robust and efficient system for detection and collection/purchasing of the new compounds appearing on the market. |
| Analyses | Chemical characterizations – to define core analytical techniques necessary for NPS identification, structure elucidation, to provide reliable MS and FTIR spectra. To establish validation procedures which guarantee scientifically valid results. |
| Evaluation | Integration of all analytical data available for the substance in analytical reports, and development of databases and tools for implementation of acquired MS and FTIR spectra in ENFSI DWG databases. |
| Organize data & information | The main project objective is to share knowledge, information (A spectra) effectively through different communication platforms. |
| Share | project's identification reference materials* among partner's laboratories and wider and to raise the awareness among NPS users. |

SOURCES OF MATERIALS (PURCHASED/SEIZED/COLLECTED)

The main goal of the project is to provide numerous MS and FTIR spectral data on new or recently reported NPS. Several sources of materials will be applied (see Table 2). Undoubtedly, certified reference materials (CRM) are the most reliable source. The drawback is that CRMs are not available in real time, i.e. when laboratory detects a new compound. The producers (of CRMs) are usually several steps - months behind the "gray" internet market.

Thus as the main source of NPS the internet vendors of research chemicals (NET-NPS) are foreseen (the target number of NPS being purchased over the internet is 200-300). The strategy and challenges for finding and purchasing novel NPS over the internet are described at a separate poster presented at this conference (L. Dujourdy et al., Systematic methodology for finding novel NPS (New Psychoactive Substances) over the internet, EAFS-2015, Prague, 2015). So far around 50 samples have been purchased from 30 different web pages.

Other important sources have been the Police and Customs seizures, samples collected by SI NGO (nongovernmental organizations) or seized materials provided from some other forensic laboratories.



Figure 1: Purchased samples obtained from different web vendors (examples)

CHEMICAL CHARACTERIZATIONS (ANALYTICAL METHODS) FOLLOWED BY THE INTERPRETATION

The selection of methods depends on the sample type and information available about the sample (see Table 2). CRMs are mostly not problematic. The starting point is always GC-MS. If there are no matches in available libraries (NFI in house library, SWGDRUG, NIST, ENFSI, etc.), HPLC-TOF is applied to find exact mono-isotopic mass and proposed empirical formula. For newly detected compounds the structure is confirmed or elucidated by NMR, FTIR-ATR is scanned, when one compound is detected, and FTIR condensed phase when we have mixture of compounds or pills, tablets and similar.

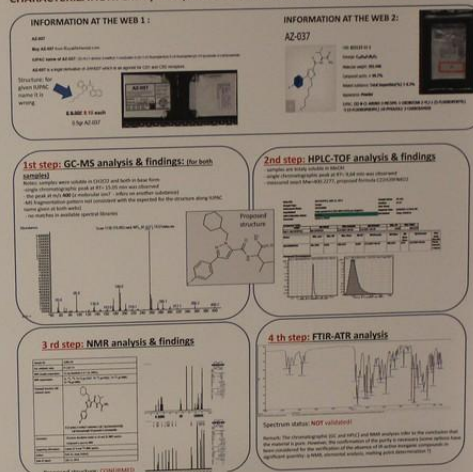
Table 2: Types of materials and analytical methods

| Type of materials | Analytical methods |
|---------------------|---|
| Reference materials | GC-MS, FTIR-ATR, GC-MS-FTIR (condensed phase) - optional |
| Test purchases | GC-MS, FTIR-ATR, GC-MS-FTIR (condensed phase), HPLC-TOF, NMR |
| Seized samples | GC-MS, FTIR-ATR and/or GC-MS-FTIR (condensed phase), HPLC-TOF and NMR for "unknowns" |
| Collected samples | GC-MS, FTIR-ATR and/or GC-MS-FTIR (condensed phase) and HPLC-TOF and NMR for "unknowns" |

This work has been produced with the financial support of the Prevention of and Fight against Crime Programme of the European Union (grant agreement number 81072013/SEC/DRUGS/AG/6413). The contents of this publication are the sole responsibility of the authors and can in no way be taken to reflect the views of the European Commission.

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CHARACTERIZATION: example of purchased samples AZ-037 (2 web pages):



EVALUATION OF ANALYTICAL DATA:

Validation procedures which guarantee scientifically valid results have been discussed at the first RESPONSE project steering committee meeting. For successful validation at available information shall be taken into account. The most challenging is the validation of analytical data (reliability of spectra) for new compounds, which has not yet been reported. When the structure is elucidated and confirmed by NMR mass spectrum fragmentation pattern has to be checked (by manual interpretation and/or by some supporting computerized tools).

Validation of FTIR-ATR spectra is even more demanding, however in depth explanation is beyond the scope of this poster. The idea of FTIR spectra validation was the comparison of FTIR-ATR vs FTIR condensed phase spectra, it was expected that for pure compounds (in base form) correlation coefficients will be in good agreement (correlation coefficient (based on cosine function) of 0.98 or better was expected). Some preliminary experiments did not confirm the expectations, at least not for all tested compounds (CRMs were applied).

Cross validation and detection of possible errors in the results reported by coordinators laboratory will be enhanced by sharing the collected/purchased samples among other project partners. "Retrospective" validation will be possible when more reports about the same compound from different sources (laboratories) will be available in "Data banks".

ORGANIZE DATA & SHARE THE INFORMATION

The project has the goal to disseminate the results of the project to the widest community. Therefore, to share gathered analytical information promptly, the public open RESPONSE project web page has recently been launched. Main analytical information on characterized NPS (CRMs, collected, seized) with several search tools is shown (and will be updated continuously) in "Drugs Monographs" section (see Figure 3). In addition the following connections have been enhanced: cooperation with ENFSI DWG (MS and FTIR spectra libraries managers) and membership in general, ENFSI, EUROPO, European Early Warning System including NGOs. At the non-formal level cooperation/communication and information exchange with several other complementary projects (SAFE, SPICE profiling, CLIN2SAR) was established as well.



Figure 3: Screenshot of the RESPONSE project web page

AWARENESS

The test purchases performed in the frame of the RESPONSE project so far, show that drugs users can never be sure of what they get, when buying from internet vendors. From 30 purchased samples the rate of "false advertised" compounds is at approximately 20%.

This poses a serious health risks for the population of NPS users.

Presented at EAFS 2015 conference, Prague, Czech Republic, September 8-11, 2015

