

Combined exposures to stable and radioactive substances for ecological risk assessment

Rodolphe GILBIN, Laureline FÉVRIER, Catherine LECOMTE, Frédéric ALONZO, Béatrice GAGNAIRE, Jean-Marc BONZOM, Olivier ARMANT, Karine BEAUGELIN, Hugo LEPAGE, Jacqueline GARNIER-LAPLACE

Rodolphe GILBIN

rodolphe.gilbin@irsn.fr

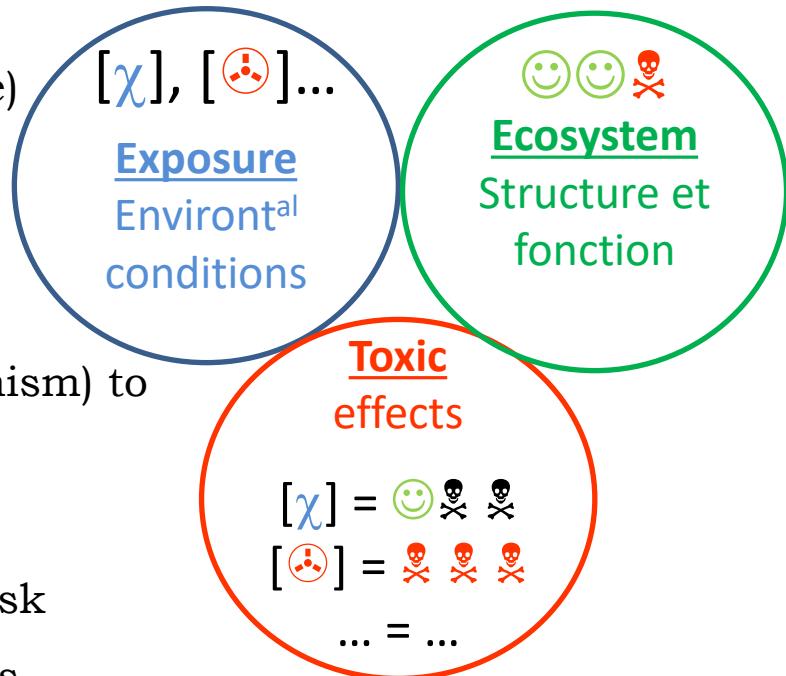
Research Department on Transfers and Effects of
Radionuclides on ecosystems
IRSN / PSE-ENV / SRTE
Cadarache, France



Introduction

1

- Exposure in ecosystems: Many substances (stable or radioactive) at potential risk
- Ecotoxicity of substances: approach by 1 (substance, organism) to determine toxic effects
- Protection target for ecological risk assessment: species assemblages, ecosystem structure and function sensitivity to all variables (contaminants and other environmental factors)



1- Introduction

Joint exposure to stable and radioactive substances

Example of stable and radioactive substances present in releases from nuclear fuel cycle facilities

Vandenbroucke *et al.* (2012) STAR D 4.1

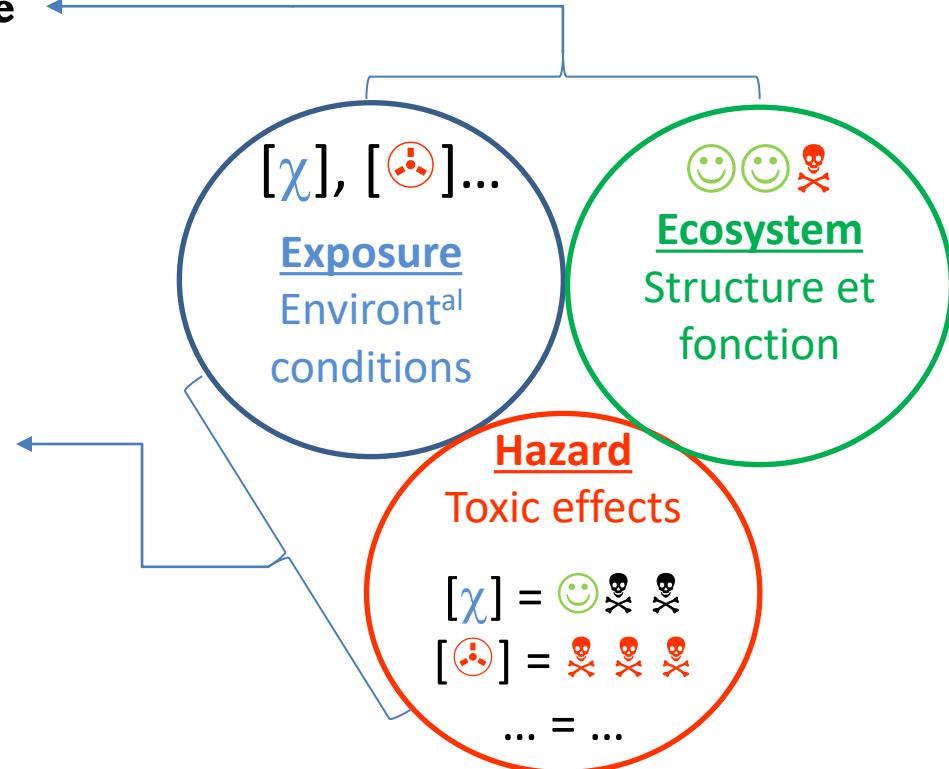
	Substances Stables	Radioactives
U mining (ex. Canada, Germany, Brazil, France)	As, Cd, Cr, Cu, Pb, Zn Ba, Fe, Al, F, Cl	U and daughters
Nuclear Fuel Production	F, NH ₄ , K, SO ₄ , As, Cl, Cr, Cu, Fe, Zn	¹⁴ C, ³ H, ⁹⁹ Tc, α, β
Electric power production	NH ₄ , B, Cu, Zn, LiOH Morpholine, hydrazine ...	^{110m} Ag, ¹⁴ C, ⁵⁸ Co, ^{134/7} Cs, ³ H, ¹³¹ I, ⁵⁴ Mn, ^{124/5} Sb, ^{123m} Te
Spent fuel reprocessing (ex. AREVA-NC La Hague)	TBP, NO ₃ , hydrazine, NH ₄ , SO ₄ , Fe, K, Ba, Al, F, Co, Cd, Ni, Cr, Hg, Pb, Zr	¹⁴ C, ³ H, ⁸⁵ K, ⁶⁰ Co, ^{134/7} Cs, I, ¹⁰⁶ Ru, ⁹⁰ Sr, ⁶⁰ Ni, ²³⁸⁻⁴¹ Pu, ¹⁰⁶ Rh, ¹²⁵ Sb, ²⁴¹ Am, ²²⁴⁴ Cm, U
Waste disposal
NORM (coal, petrol, gaz, minerals extraction...;) <i>Hospitals, R&D</i> <i>Accidents</i> <i>Other industries, agriculture, non-point sources...</i>	metals, hydrocarbons...	natural RNs ...

1- Introduction

Approaches developed at IRSN

Determination of the **ecological structure**
(species assemblages) **and function**
(integrated biomarker response)

Cumulative Risk Assessment (msPAF
approach for the prediction of chronological
impact to aquatic ecosystem)



Cumulative Risk Assessment

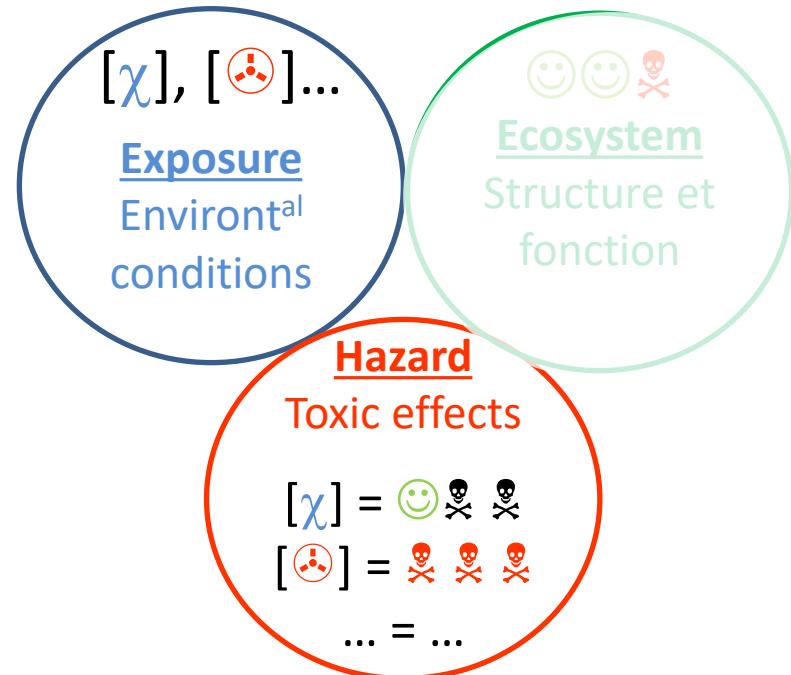
2



Example : routine releases
of NPPs in the Rhône river

Beaumelle et al. (2017), Environ. Poll. 231:1421-32

IPOTOMUS project (in progress)



2- Cumulative Risk Assessment

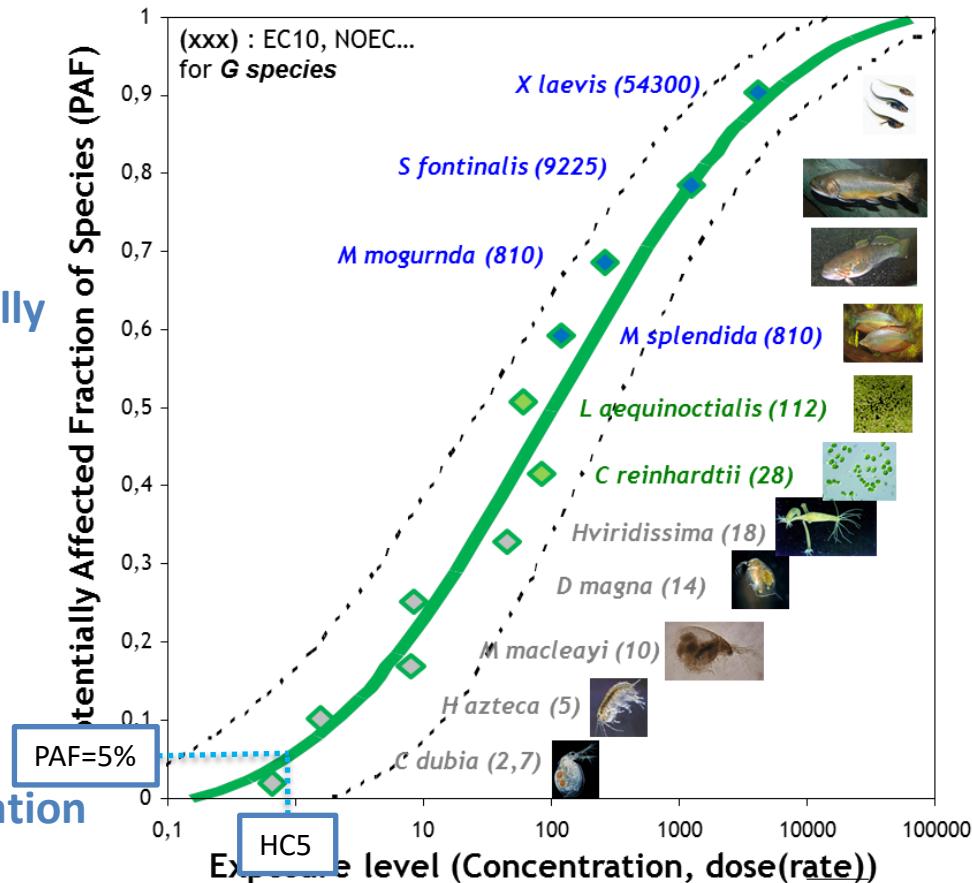
Methods : Species Sensitivity Distribution (SSD)

■ Cumulated Distribution Function (CDF) of chronic (or acute) ecotoxicological data (NOEC, EC10...) for *all** species

■ Expresses the **fraction of species potentially affected** (PAF) by the exposure to a given stressor

■ Widely used to account for the **variation in species sensitivity** (De Zwart & Posthuma, 2005)

& to derive **maximum acceptable concentration** of a substance
(e.g. HC5 : Hazardous Concentration that affect 5% of species)



↗ Consistent benchmark derivation for substances

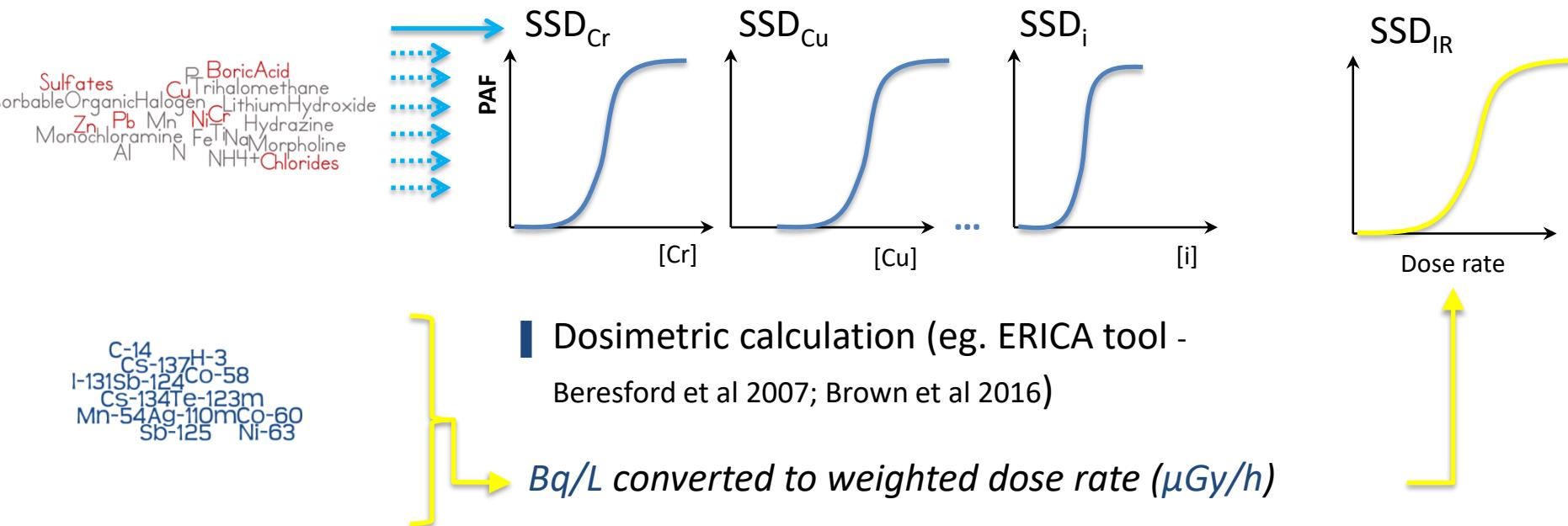
2- Cumulative Risk Assessment

Methods : Multi-Substances SSDs (msPAF)

- SSDs recommended to provide an **integrated indicator** of the potential ecological risk of a mixtures: **msPAF** (multi-substance Potentially Affected Fraction of species)

(De Zwart and Posthuma, 2005)

SSD derivation: every stable substance ionizing radiation



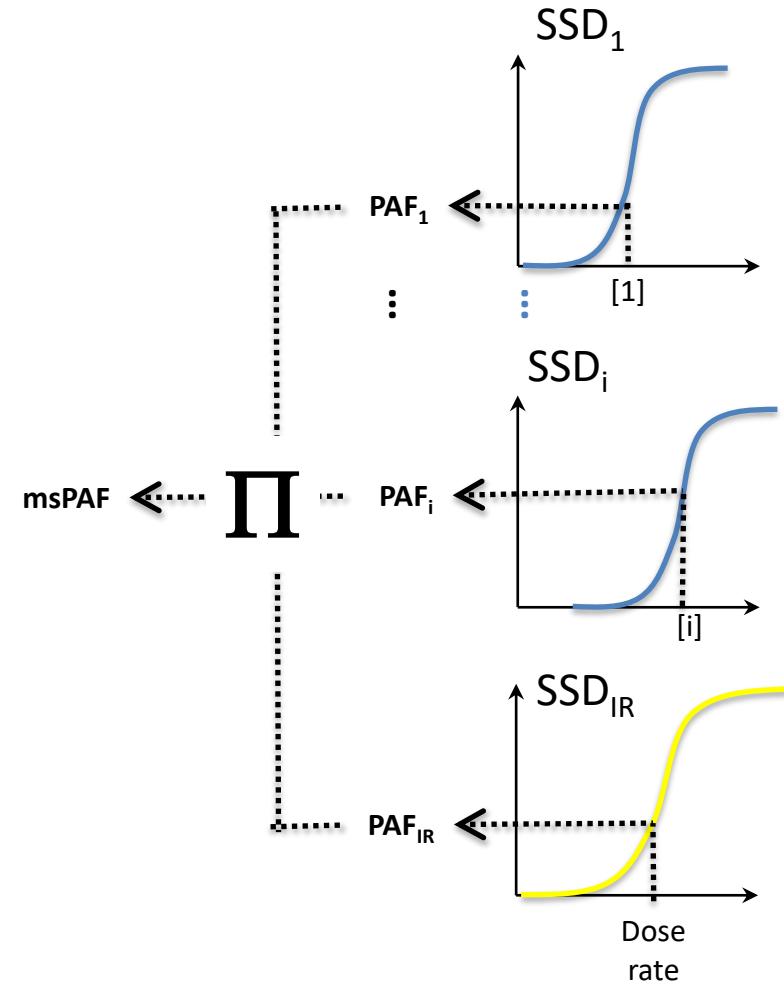
2- Cumulative Risk Assessment

Methods : SSDs combination (1/2)

Independent Action (IA)

- stressors considered independently
- PAF calculation for each stressor
- msPAF computed as following :

$$msPAF_{IA} = 1 - \prod_{i=1}^n (1 - PAF_i)$$



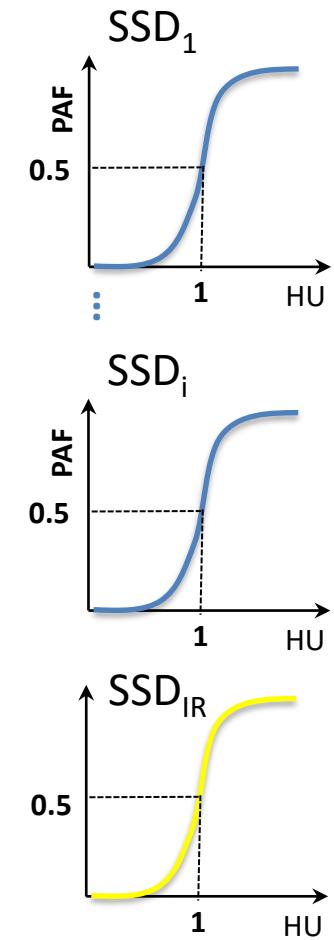
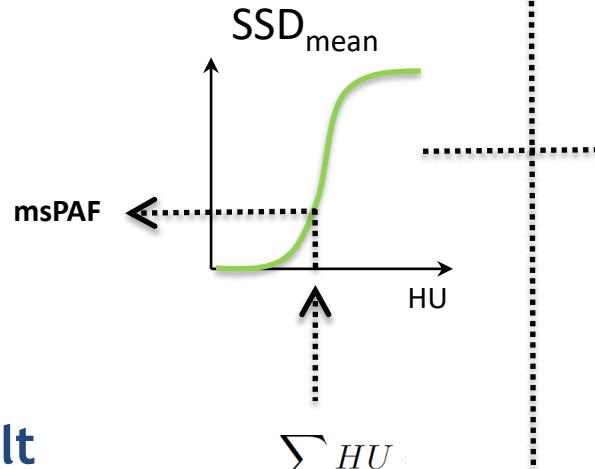
2- Cumulative Risk Assessment

Methods : SSDs combination (2/2)

Concentration Addition (CA)

- all SSDs normalized to their toxic potential (HC50)
- exposure expressed as Hazard unit (HU)
- 1 single average SSD for all stressor
- msPAF computed as following :

$$\sum HU = \sum_{i=1}^n \frac{C_i}{HC50_i}$$



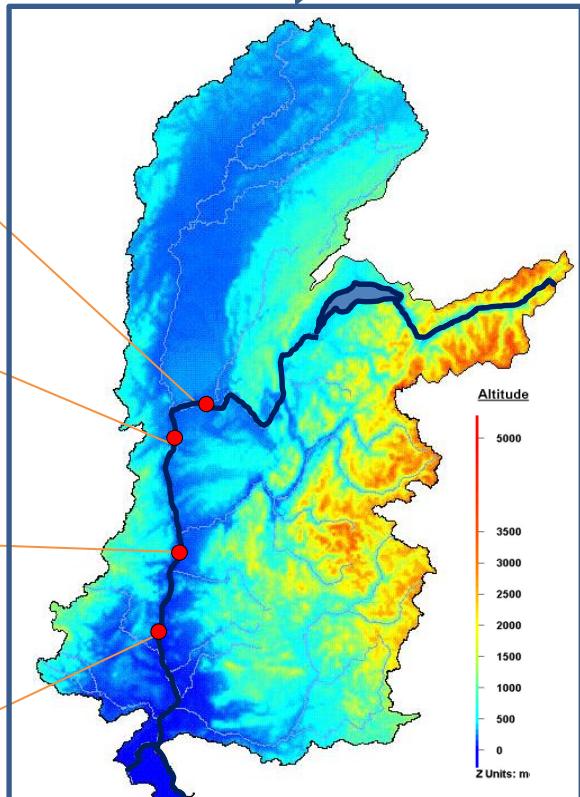
➤ recommended as the default (conservative) approach for multi-substance Risk Assessment

(Backaus et al., 2000)

2- Cumulative Risk Assessment

Case study : NPPs routine releases

Rhône river
catchment
(SE France)



4 NPPs

Bugey

4x900 MW
130-457 m³/s

St Alban

2x1300 MW
255-1040 m³/s

Cruas

4x900 MW
400-1500 m³/s

Tricastin

4x900 MW
300-1500 m³/s

Annual released quantities (Q_i)
under normal operating conditions

(EDF, 2013)

Q_i (kg or GBq)	Bugey	Tricastin	Cruas	St-Alban
Cr				1
Cu	17	2	15 100	1
Ni				1
Pb		1		1
Zn	7	9	6 600	4
Boric acid	9 000	9 050	13 700	3 730
Chlorides	44 200	13 900		33 000
Sulfates	142 000	98 000	3 967 000	
Ag-110m	0.47	0.05	0.13	0.01
C-14	38.00	45.00	47.00	31.00
Co-58	0.45	0.06	0.13	0.04
Co-60	0.10	0.16	0.18	0.07
Cs-134	0.02	0.03	0.04	0.01
Cs-137	0.02	0.04	0.04	0.01
H-3	44 300.00	51 300.00	58 300.00	55 600.00
I-131	0.02	0.03	0.03	0.01
Mn-54	0.02	0.04	0.04	0.01
Ni-63	0.13	0.07	0.19	0.14
Sb-124	0.05	0.03	0.03	0.01
Sb-125	0.08	0.09	0.09	0.03
Te-123m	0.01	0.04	0.04	0.01

2- Cumulative Risk Assessment

Case study : NPPs routine liquid releases

average annual conc. (C_i) ←

4 NPP considered

2 dilution scenarios

- mean flow rate
- minimum flow rate

$$C_i = \frac{Q_i F_{rel}}{V_{tot} F_{riv}}$$

C_i	concentration/activity in river (Bq/L)
Q_i	cumulated annual release (Bq or kg)
V_{tot}	volume of liquid effluents (L)
F_{rel}	flow rate of the releases (m^3/s)
F_{riv}	flow rate of the Rhône river (m^3/s)

■ Annual released quantities (Q_i) under normal operating conditions
(EDF, 2013)

Q_i (kg or GBq)	Bugey	Tricastin	Cruas	St-Alban
Cr			1	
Cu	17	2	15 100	1
Ni			1	
Pb		1	1	
Zn	7	9	6 600	4
Boric acid	9 000	9 050	13 700	3 730
Chlorides	44 200	13 900		33 000
Sulfates	142 000	98 000	3 967 000	
Ag-110m	0.47	0.05	0.13	0.01
C-14	38.00	45.00	47.00	31.00
Co-58	0.45	0.06	0.13	0.04
Co-60	0.10	0.16	0.18	0.07
Cs-134	0.02	0.03	0.04	0.01
Cs-137	0.02	0.04	0.04	0.01
H-3	44 300.00	51 300.00	58 300.00	55 600.00
I-131	0.02	0.03	0.03	0.01
Mn-54	0.02	0.04	0.04	0.01
Ni-63	0.13	0.07	0.19	0.14
Sb-124	0.05	0.03	0.03	0.01
Sb-125	0.08	0.09	0.09	0.03
Te-123m	0.01	0.04	0.04	0.01

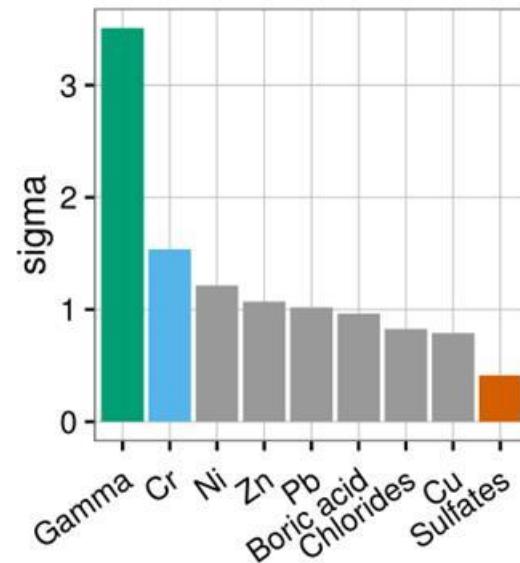
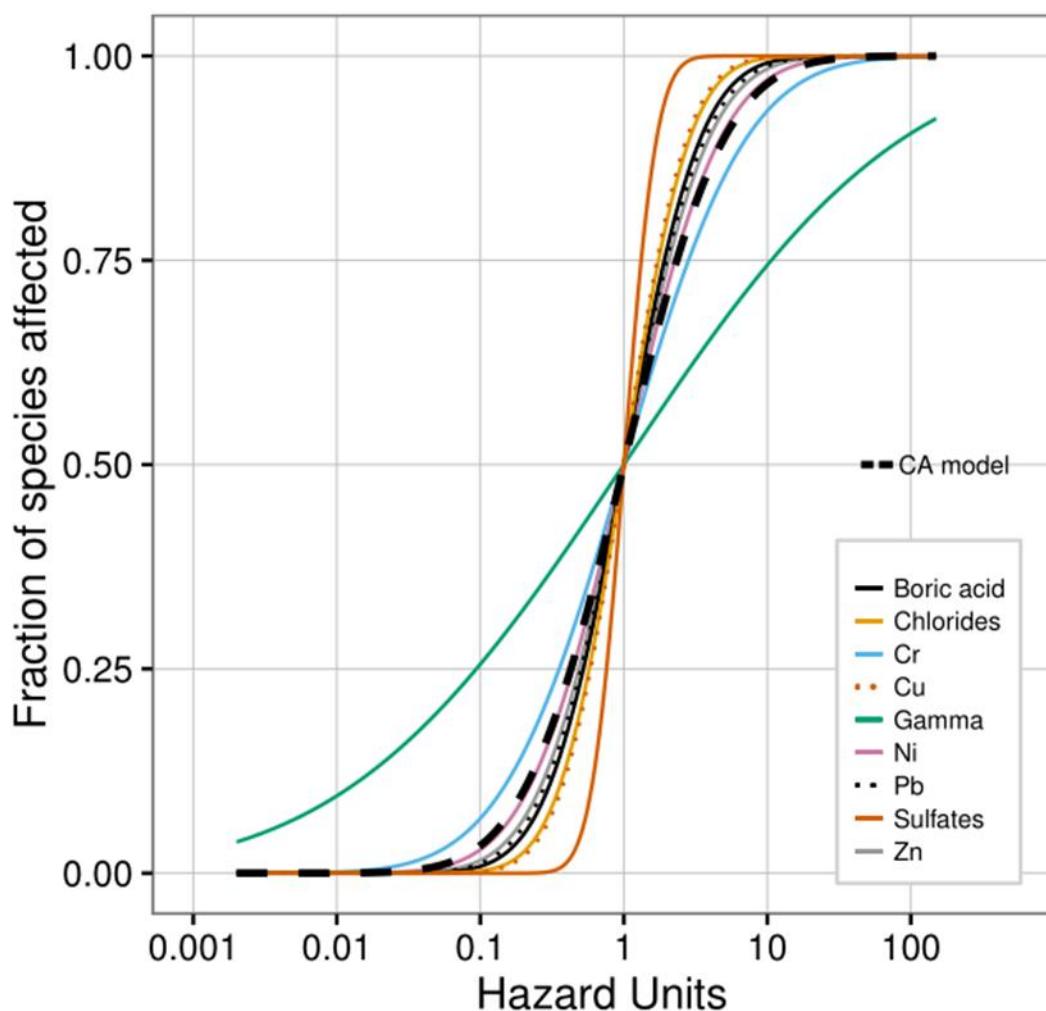
stable substances

radioactive substances

↗ 8 contrasted scenarios
(dilution, mixture composition)

2- Cumulative Risk Assessment

Average SSD derivation (for CA approach)



Statistically relevant groups = similar slopes (Bartlett test)

- ↗ 3 msPAF calculations
 - CA Conc. Addition
 - IA Independent Action
 - CA&IA 4 CA groups, then IA

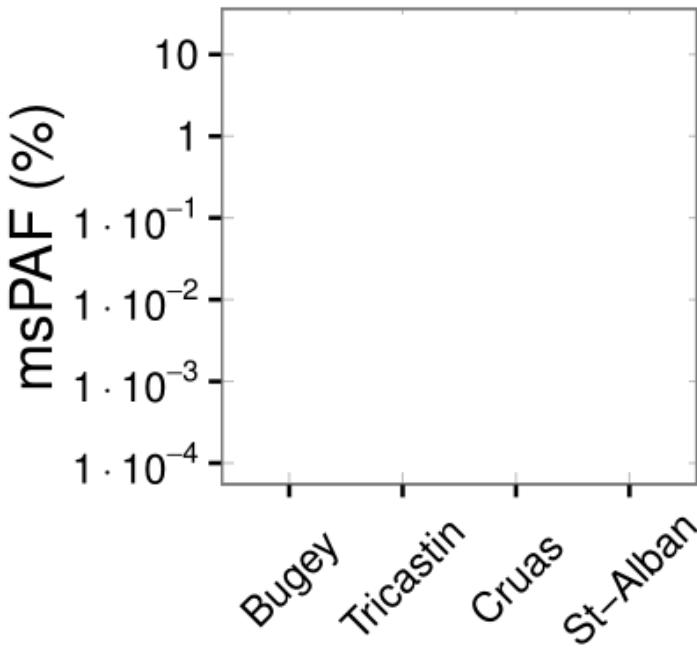
2- Cumulative Risk Assessment

msPAF calculation results

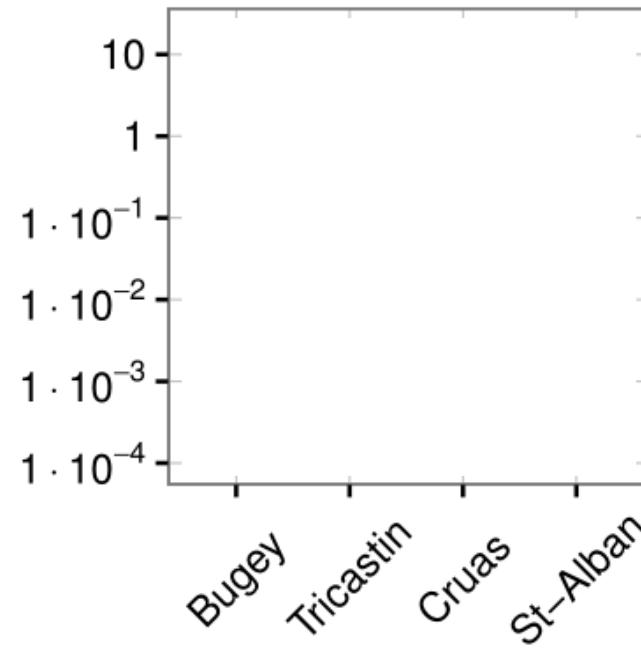
msPAFs associated with different release scenarios using



Mean-flow scenario



Lowest-flow scenario



- very low risk level : msPAF << 5% except 1 case (Cruas with CA)
- No difference between CA and CAIA
- Surprisingly, CA not always conservative

due to the differences in SSD slopes

↗ Use of both concepts (CA and IA) to provide a “*prediction window*”

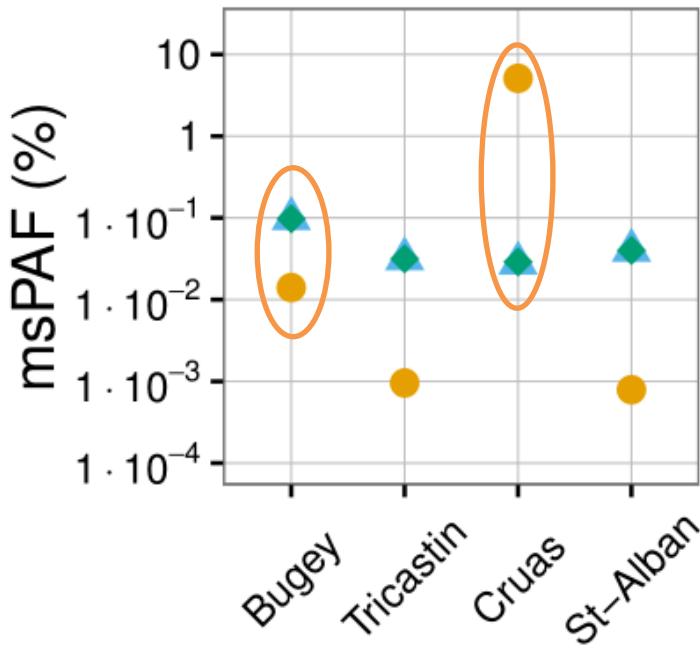
2- Cumulative Risk Assessment

msPAF calculation results

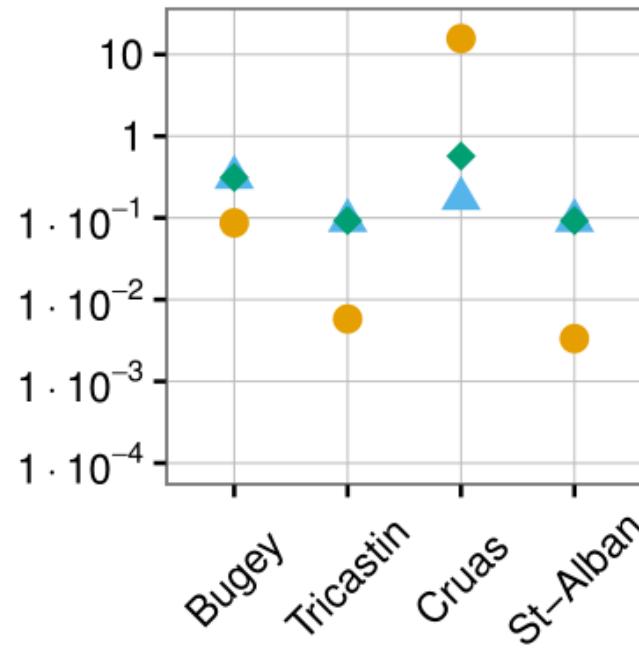
msPAFs associated with different release scenarios using



Mean-flow scenario



Lowest-flow scenario



- ↗ Contribution of the different stressors to the overall msPAF ?
eg. contrasted cases of Bugey and Cruas (mean flow)

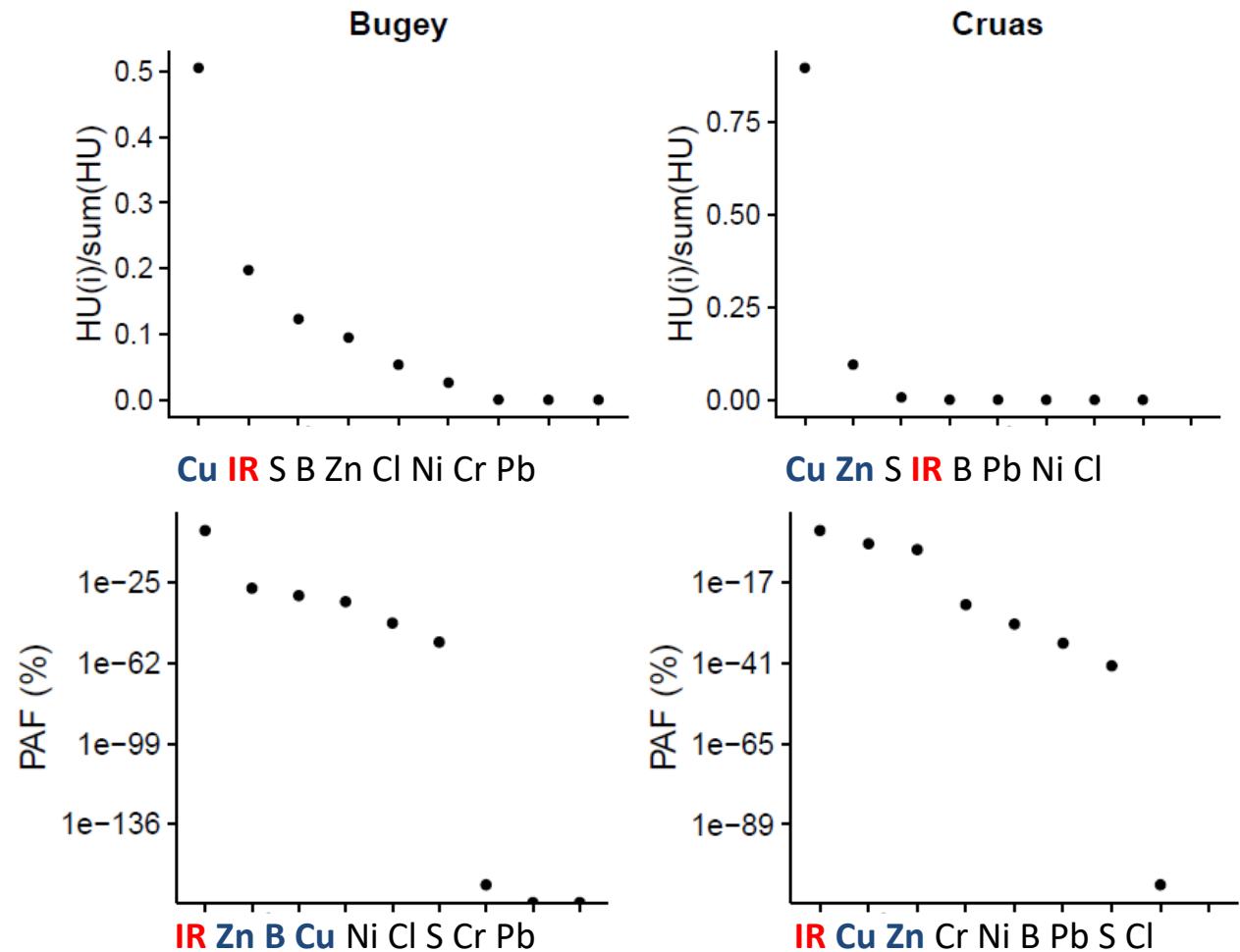
2- Cumulative Risk Assessment

Stressors combination and ranking

- Concentration Addition

$$HU_i = \frac{C_i}{HC_{50,i}}$$

- ▲ Independant Action
individual PAF
for each
stressor/group



↗ Different ranking as a function of the considered model (CA or IA)

2- Cumulative Risk Assessment

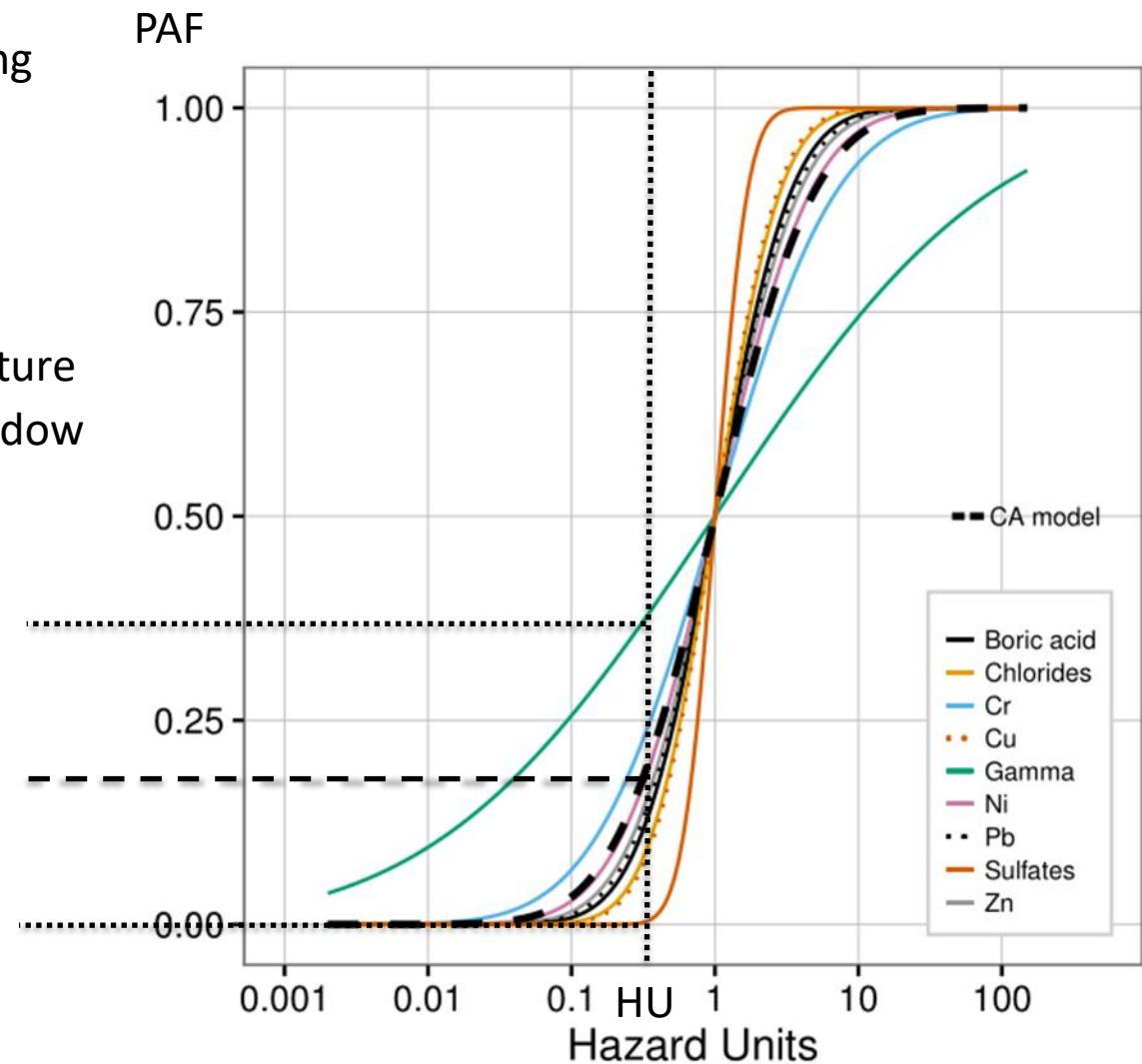
Conclusion (1/2)

Very different SSD shapes (ionising radiation vs. stable chemicals)

CA is not the most conservative

→ Considering both CA and IA mixture models to provide a prediction window

Ionizing rad.
underestimation
Average (CA)
overestimation
 SO_4



2- Cumulative Risk Assessment

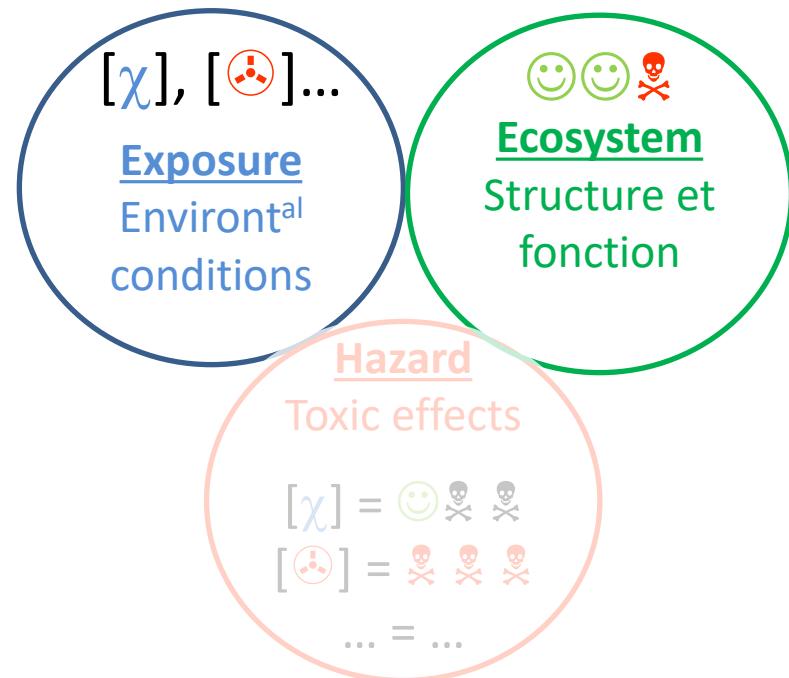
Conclusion (2/2)

- **Negligible overall potential ecological risk** in the context of routine effluents releases in aquatic ecosystems (except msPAF exceeding 5% in 1 case)
- **msPAF does not provides directly information about the contribution / ranking** of each stressor
 - ionizing radiation dominates the contribution to PAF (IA)
 - stable chemicals (mainly Cu, Zn) dominates contribution to HU (CA)
- Main limitations (and perspectives)
 - **Unsufficient ecotoxicity datasets** for some substances cannot allow a robust SSDs derivation (only 8 among 22 stable chemicals considered)
→ In progress (**Ipotomus Project**): use of monitoring data (chemical, radioactive) and the Δ PAF approach (> chemicals considered: 84 among 1046)
 - **Limited ecological relevance** (needs validation with field data)
 - Do not consider **realistic exposure conditions** (bioavailability, competitions...)
 - Do not consider **potential interactions** (synergism, antagonism)

Mixture Exposure and Ecological status

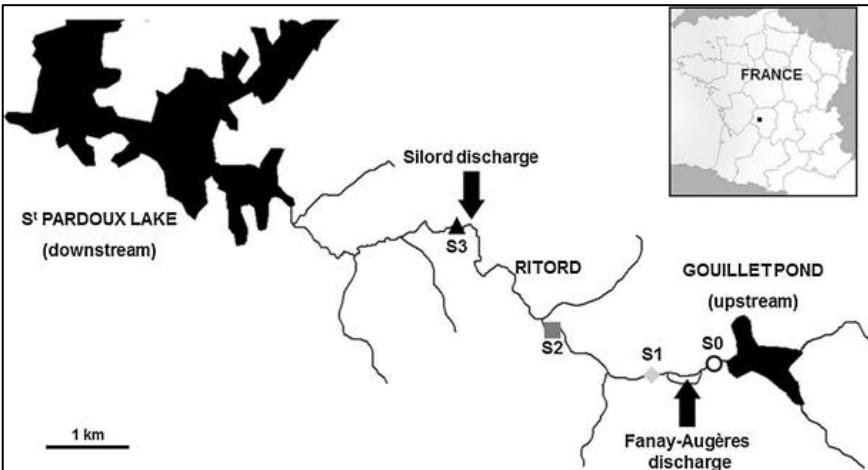


Example : releases from
former U mines in France



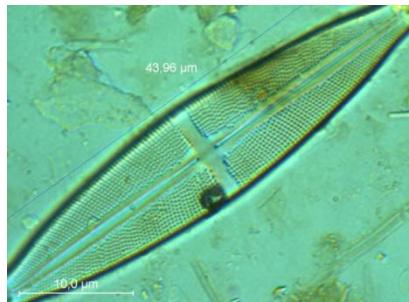
3- Mixture exposure and diatom community diversity

Example : Ritord, releases from former uranium mine sites



Herlory et al. (2013) Ecotoxicology

- Objective: to determine the **diversity and quality of species assemblages**
- Methodology: **Benthic diatoms** (community distribution and ecological preferences)
- Data set: **7-month monitoring** (July-January), monthly sampling
 - environmental conditions, contaminants
 - biomass, photosynthesis, specific composition of diatom assemblies



3- Mixture exposure and diatom community diversity

Example : Ritord, releases from former uranium mine sites

▪ Environmental conditions

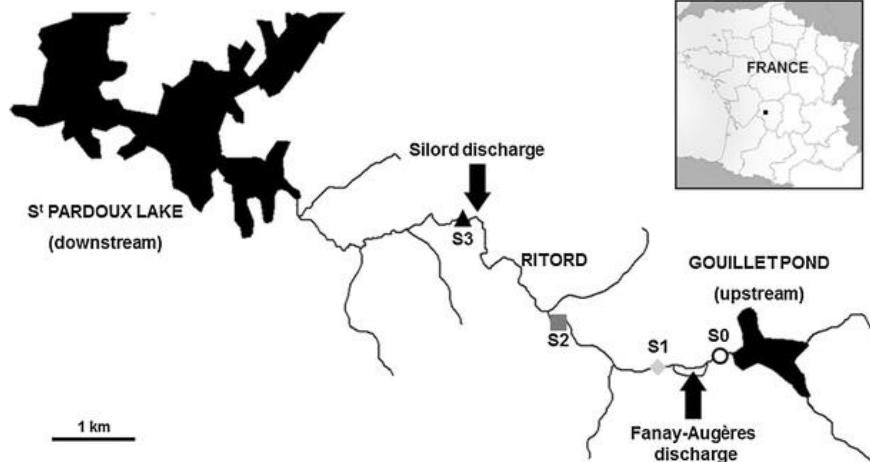
T°, pH, conductivity, O₂, ORP (redox)
Nutrients (Si, NO₂, NH₄, NO₃, SO₄, Cl)
Carbon inorg. (TIC) et organique (TOC)
Cations (Ca, K, Mg, Na)
Metals diss and bioacc (U, Al, Ba, Fe, Mn)

▪ Specific composition of diatom assemblies



→ Specific identification and abundance (microscopy after cleaning)

Herlory et al. (2013) Ecotoxicology



12km (1m prof.), BV 17km²
connects 2 ponds (Gouillet → St Pardoux)

7 former mines, collect and treatment of effluents
(Fanay-Augères, Silord) : BaCl₂ (²²⁶Ra↓), Al₂(SO₄)₃
(Fe, U ↓)

4 stations :

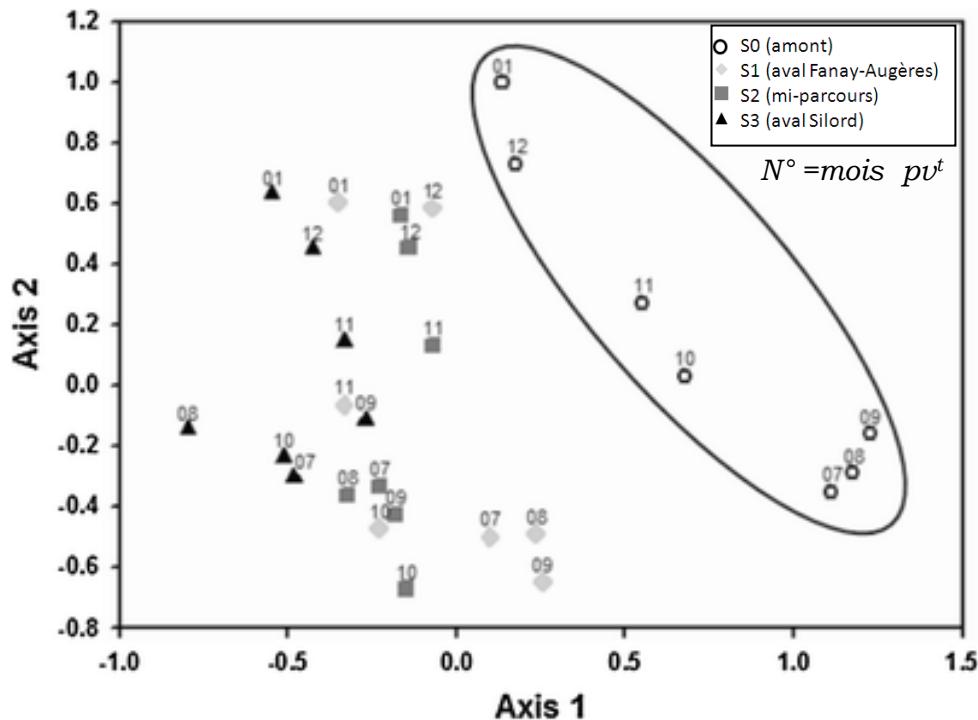
- S0 (upstream)
- ◆ S1 (downs. Fanay)
- S2 (mid-course)
- ▲ S3 (downstream Silord)

3- Mixture exposure and diatom community diversity

Example : Ritord, releases from former uranium mine sites

Herlory et al. (2013) Ecotoxicology

▪ Environmental conditions



Axis 1		Axis 2	
Variable	ρ_s	Variable	ρ_s
Al accumulated	-0.82	TIC	-0.85
Cl	-0.77	Conductivity	-0.83
pH	-0.67	Temperature	-0.79
Ba accumulated	-0.59	O ₂ dissolved	0.44
Mn accumulated	-0.46	Stream water flow	0.64
Na	-0.45	ORP	0.82
Ba dissolved	-0.41		
U dissolved	-0.39		
NH ₄	0.62		
Fe dissolved	0.65		
Fe accumulated	0.77		

NMDS (Non-metric multidimensional scaling)

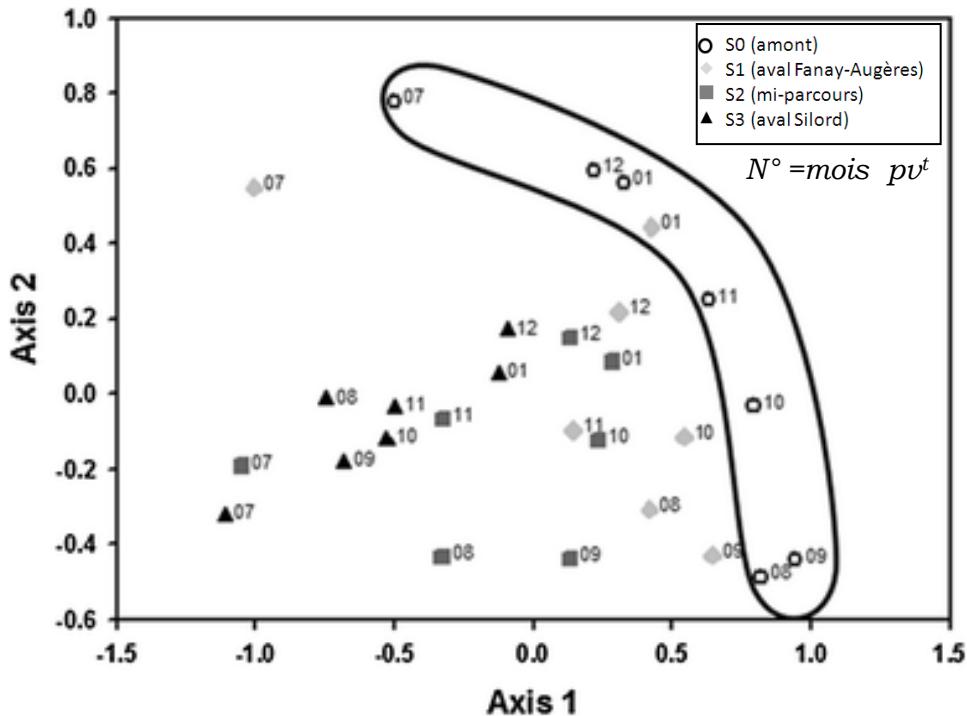
ρ_s : corrélation de Spearman (rang) entre les coordonnées de l'échantillon et les variables

- **S0 (upstream) significantly different** from other stations (PERMANOVA)
- Effect of **contamination (Axis 1)**: U and treatment effect (Ba, Al, Mn, pH)
- 'Season' effect (Axis 2): ICT, Cond, redox, T °, flux, O₂

3- Mixture exposure and diatom community diversity

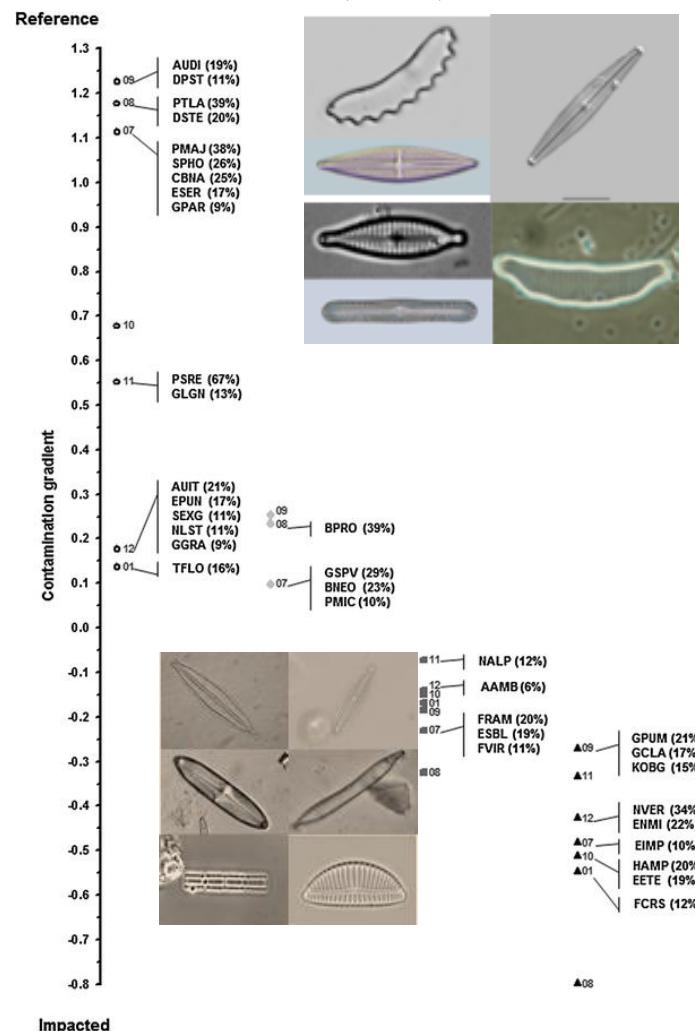
Example : Ritord, releases from former uranium mine sites

Diatom Species composition



- No effect on specific wealth** ($S = 55$ to 74), diversity ($H' = 2,5$ to $3,2$), biomass or photosynthesis
- Differences in community composition** (S0 differ from other stations, "indicator species")

Herlory et al. (2013) Ecotoxicology



3- Mixture exposure and diatom community diversity

Example : Ritord, releases from former uranium mine sites

- Diatom community composition differences attributed to the variables characterizing the **effluents 'as a whole'**
- U, but also** Cl, Ba, Al, Mn ...
- Role of uranium vs. other environmental variables on Diatom community composition ?
- Interactions between substances?
- Actual exposure and bioavailability conditions (pH changes, conductivity, redox ...)?

pS : Spearman correlation (rank) between similarity matrices 'Specified composition' and 'Environmental conditions'

Herlory et al. (2013) Ecotoxicology

Variable	p _s
<i>Corrélations individuelles</i>	
Cl	0.466***
U _{diss}	0.377***
Ba _{diss}	0.321***
ORP (redox)	0.294***
Conductivity	0.273**
Al _{acc}	0.258**
Mn _{diss}	0.270**
Ba _{acc}	0.250**
TIC	0.151**
Na	0.165**
Fe _{acc}	0.249**
NH ₄	0.176**
Temperature	0.168*
Mn _{acc}	0.188*
Fe _{diss}	0.211*
pH	0.165*
Flow	0.088*
TOC, SPM, O ₂ , Al	NS

BIO-ENV procedure, (Clarke and Ainsworth, 1993)

*** p ≤ 0.001; ** p ≤ 0.01; * p ≤ 0.05

Mixture Exposure and Ecological status



[χ], [☢]...

Exposure
Environmental
conditions



Ecosystem
Structure et
fonction

Hazard
Toxic effects

[χ] = ☺ ☠ ☠

[☢] = ☠ ☠ ☠

... = ...

Other examples on fish :

Integrated Biomarker Response

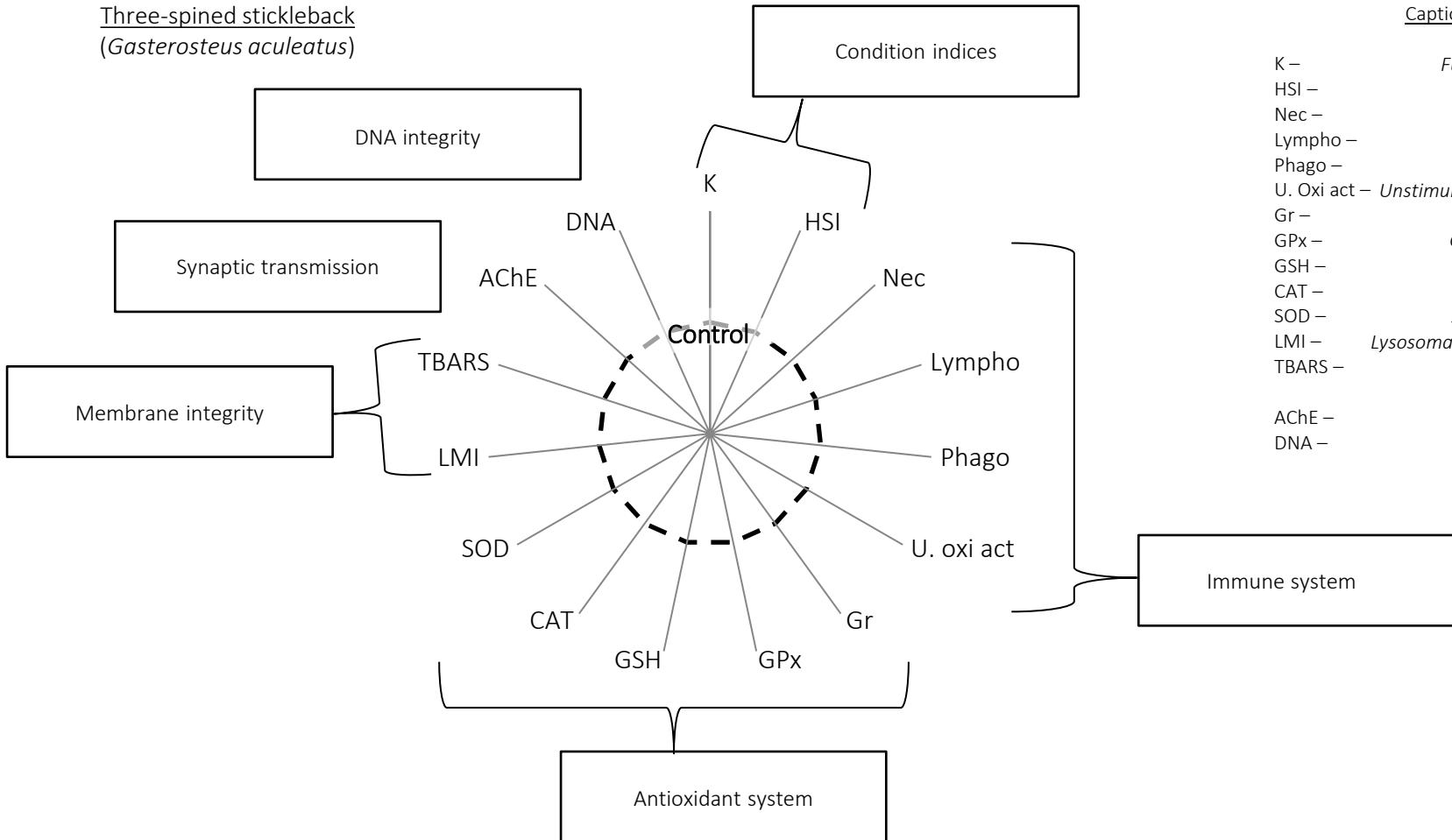
Approach

3- Integrated Biomarker Response approach

Responses on stickleback health



Three-spined stickleback
(*Gasterosteus aculeatus*)



Le Guernic et al. (2016),
Ecotoxicology 25(8):1478-99

Caption

K –	Fulton's condition index
HSI –	Hepatosomatic index
Nec –	Leucocyte necrosis
Lympho –	Leucocyte distribution
Phago –	Phagocytic capacity
U. Oxi act –	Unstimulated oxidative activity
Gr –	Glutathione reductase
GPx –	Glutathione peroxidase
GSH –	Glutathione total
CAT –	Catalase
SOD –	Superoxide dismutase
LMI –	Lysosomal membrane integrity
TBARS –	Thiobarbituric acid reactive substances
AChE –	Acetylcholinesterase
DNA –	DNA damage

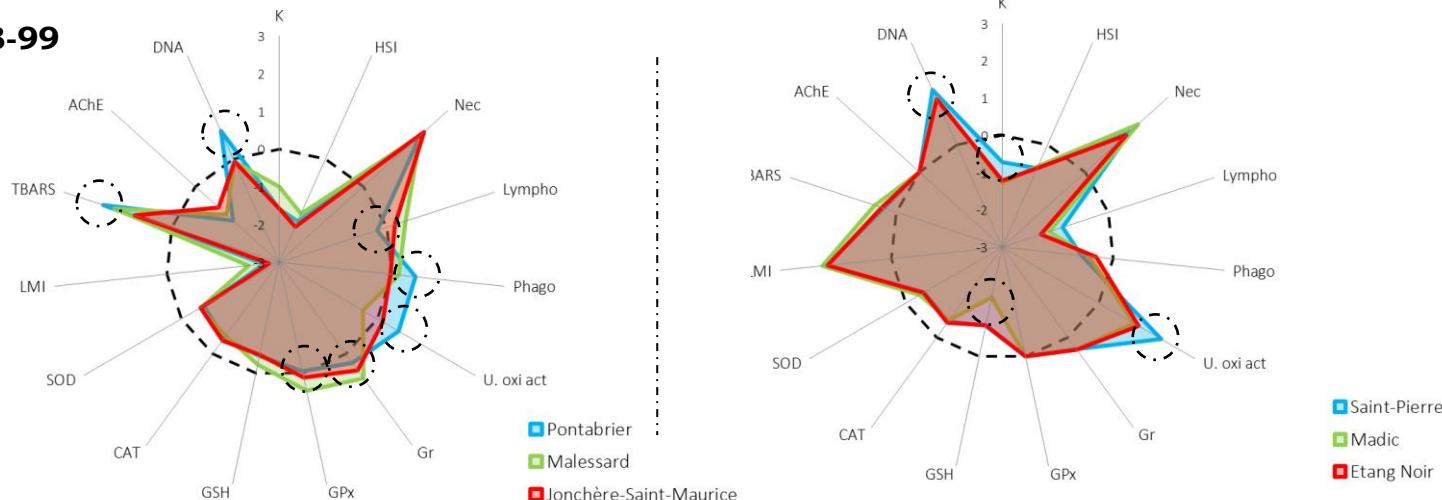
3- Integrated Biomarker Response approach

Responses on stickleback health

		U	AI	Fe µg/L	Mn	Ba
Haute Vienne	Pontabrier	23.5	82.7	263.6	52.2	18.5
	Malessard	0.3	44.9	82.0	16.1	6.4
	JSM	0.1	48.7	396.6	27.1	22.1
Cantal	Saint-Pierre	5.9	44.1	165.7	218.1	16.5
	Madic	0.1	137.0	466.4	44.3	40.5
	Etang Noir	0.2	202.2	295.0	28.7	21.2

Le Guernic et al. (2016),

Ecotoxicology 25(8):1478-99



Polymetallic contamination

Oxidative stress

DNA damage

Other effects

3- Integrated Biomarker Response approach

Responses on fathead minnow



Fathead minnow
(*Pimephales promelas*)

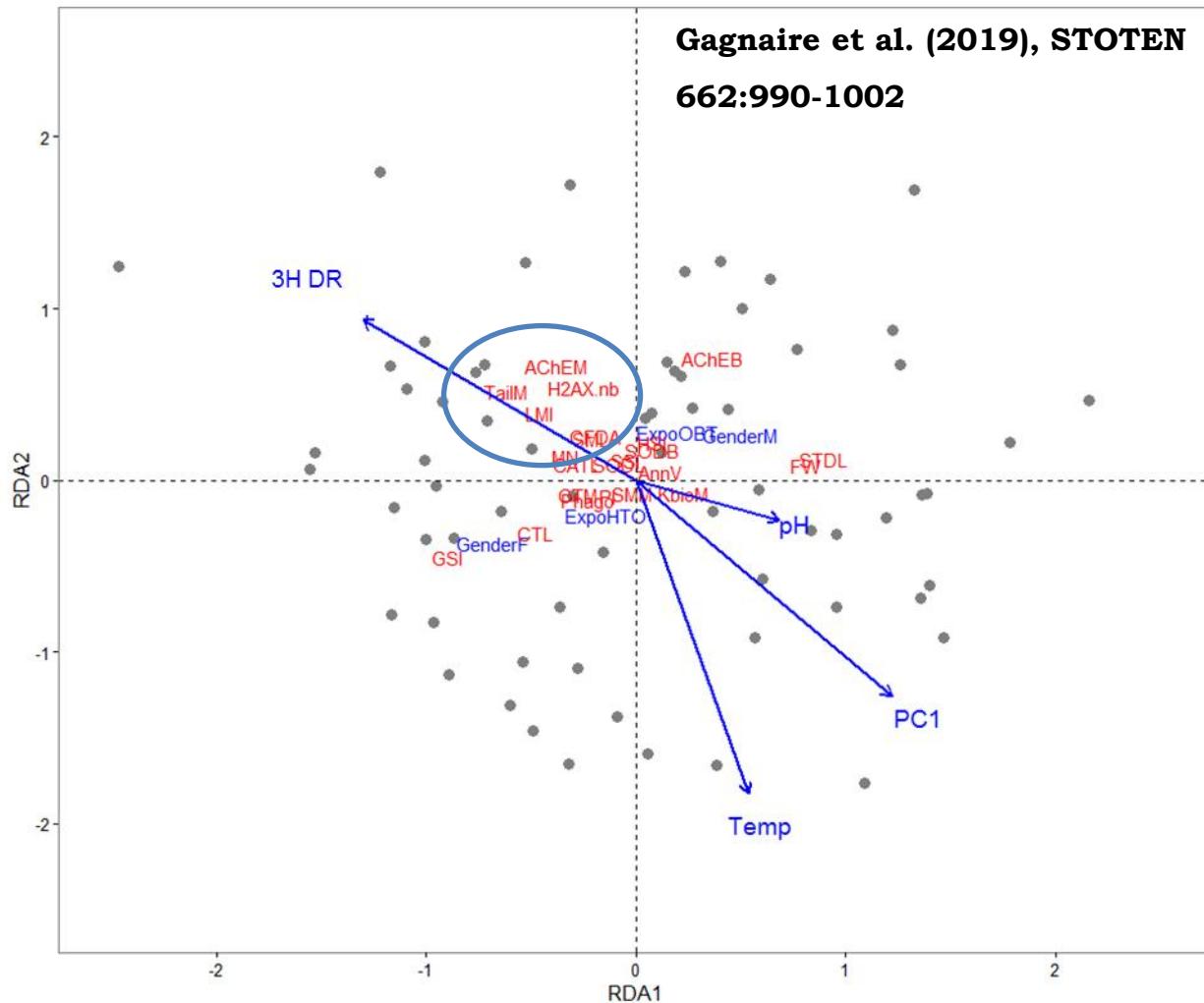
[Tritium] from 4 to 12000 Bq/L (max=0.15 μ Gy/h) - Chalk River Laboratory (CNL)
60 days contamination + 60 days depuration

■ Redundancy Discriminant

Analysis (RDA) with all biomarkers data

→ Positive correlation of responses (LMI, μ N, comet, H2AX, muscle AChE, SOD, CAT, fatty acids) with tritium exposure (despite very low dose-rate estimates)

■ Lower correlation (few biomarkers) linked with metals (PC1)



3- Mixture exposure and ‘ecological’ survey (diatom, fish)

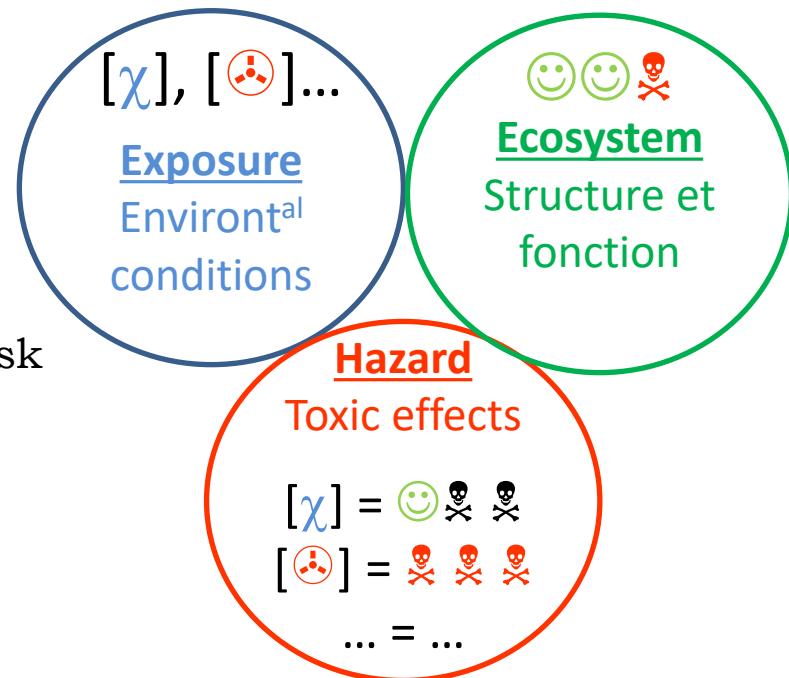
Conclusion

- | Provides relevant (and direct) information on structure and/or function of communities/organisms
- | However, results interpretation is not straightforward:
 - how to identify the contribution of stressors?
presented examples (former U mines, 3H contamination):
ionizing radiation may not be the main stressor
 - How to link changes (diversity, biomarkers...) with ecological consequences?

How to go beyond a descriptive approach?



- Exposure in ecosystems:
 - Chemical speciation?
 - Bioavailability?
- Protection target for ecological risk assessment:
 - Relevancy of the SSD?
- Ecotoxicity of substances in mixture:
 - CA or IA?
 - Interactions between substances?



4- Beyond a descriptive approach

Exposure : Speciation and Bioavailability

Tipping & Lofst (2013) Aquat Tox 142-3:114

- Generalization of BLM approach : WHAM-FTOX
- metal-organism binding on non-specific sites; analogy with humic acids (WHAM)
- Advantage: model already parameterized and thermodynamically coherent

Toxic Function: $F_{TOX} = \sum \alpha_M v_m$

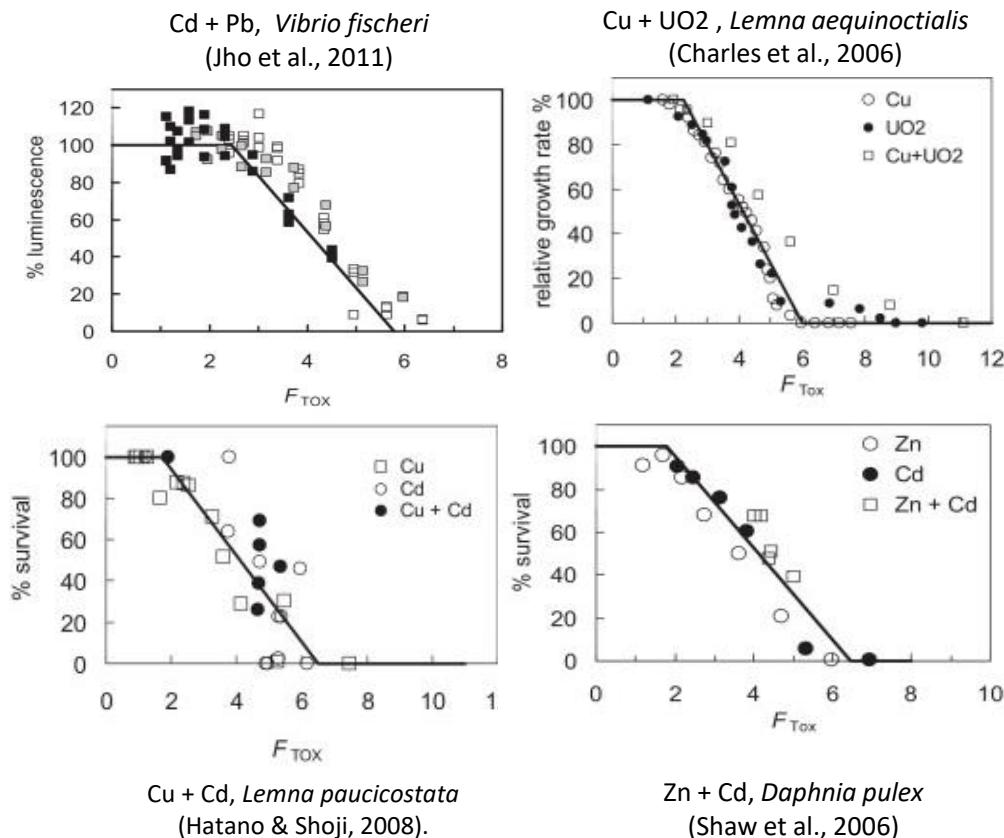
α_M : quantité de métal M accumulé ($mmol.g^{-1}$)

v_m : coefficient de toxicité

Toxic Response: $TR = \frac{F_{TOX} - F_{TOX,LT}}{F_{TOX,UT} - F_{TOX,LT}}$

$F_{TOX,LT}$: lower threshold (pas d'effet)

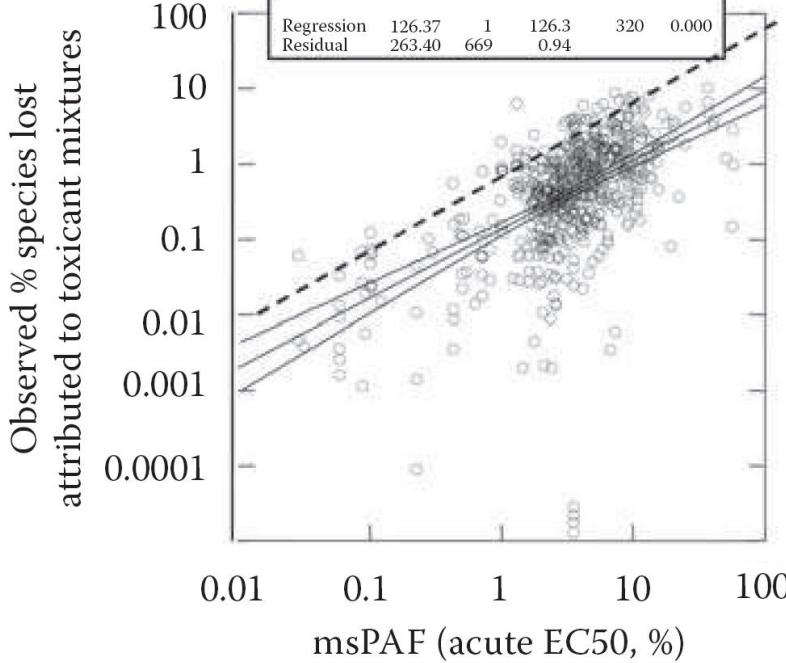
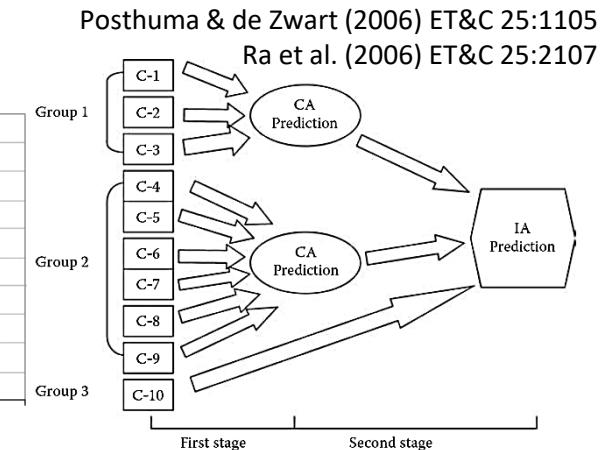
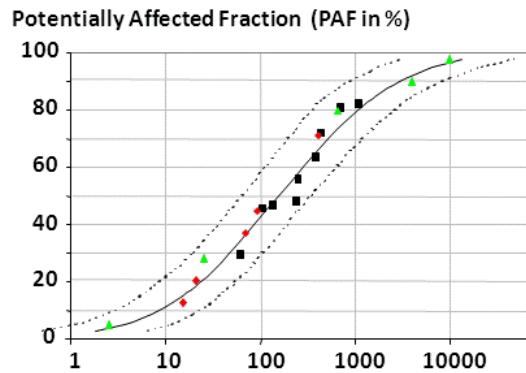
$F_{TOX,UT}$: upper threshold (effet maximum)



4- Beyond a descriptive approach

SSD and msPAF: Integrated Approach for Ecological Risk Assessment

- Prediction of effects from laboratory data base (SSD)
 - Measurement of the contamination pressure (Cd, Cu, Pb, Ni, Zn, ammonium, triclosan, ...)
 - Bioavailability (available fraction, hardness correction)
- Mixed mixture model (CA ou IA)
- Fish species diversity (espèces de poissons, 700 sites, Ohio)
- msPAF vs. species abundance correlation (GLM)



4- Beyond a descriptive approach

Ecotoxicity: Interactions between stable and radioactive substances

from Vanhoudt et al. (2012) Env Poll 168:177

- 16 reported studies (binary mixtures, 11 organisms)
- 13 Negative Interactions
U + Se, Cd
 γ + Cd, Pb, Al
- 10 Positive Interactions
 β + Cu
 γ + Cd, Hg
 γ + organiques
Pu + γ , Amiante, Be
- Exploratory studies (partial experimental design: doses, ratios, effect types considered)
- Not very informative about the mechanisms (toxicoco-kinetic and toxic-dynamic interaction)

	RN	Ch.	-	=	+	
<i>U + métaux/métalloïdes</i>						
<i>Lemna aequinoctialis</i>	U	Cu	X			Charles et al. (2006)
<i>Arabidopsis thaliana</i>	U	Cd		X		Vanhoudt et al. (2010)
<i>Caenorhabditis elegans</i>	U	Cd	X			Margerit et al (in prep)
<i>Daphnia magna</i>	U	Se	X			Zeman et al (2007)
<i>Rayonnement + métaux</i>						
<i>Chlamydomonas reinhardtii</i>	H-3	Cu		X		Réty et al (2012)
<i>Hordeum vulgare</i>	Cs-137	Cd	X	X		Geras'kin et al. (2005)
<i>Souris</i>	X-ray	Cd	X	X		Michel & Fritz-Niggli (1986)
<i>Salmo salar</i>	Gamma-rays	Cd		X		Mothersill et al. (2007)
<i>Salmo salar</i>	Co-60	Cd	X			Salbu et al. (2008)
<i>Salmo salar</i>	Co-60	Cd	X			Olsvik et al. (2010)
<i>Oryzias latipes</i>	Cs-137	Cd		X		Shimada et al. (1985)
<i>Souris</i>	X-ray	Cd	X			Michel & Balla (1987)
<i>Hordeum vulgare</i>	Cs-137	Pb	X	X		Geras'kin et al. (2005)
<i>Salmo salar</i>	Gamma-rays	Al		X		Mothersill et al. (2007)
<i>Salmo salar</i>	Co-60	Al		X		Salbu et al. (2008)
<i>Salmo salar</i>	Co-60	Al		X		Olsvik et al. (2010)
<i>Oryzias latipes</i>	Cs-137	As		X		Shimada et al. (1985)
<i>Souris</i>	X-ray	Hg			X	Michel & Balla (1987)
<i>Rayonnement + organiques</i>						
<i>Hordeum vulgare</i>	Cs-137	2,4-D	X	X		Geras'kin et al. (2005)
<i>Tradescantia</i>	X-ray	PDA	X	X		Xiao & Ichikawa (1998)
<i>Tradescantia</i>	X-ray	DMN	X	X		Xiao & Ichikawa (1998)
<i>Oryzias latipes</i>	Cs-137	Cycloheximide		X		Shimada et al. (1985)
<i>Pu en mélange</i>						
<i>Rat</i>	X-ray	Pu-239	X	X		Ballou et al. (1962)
<i>Rat</i>	Pu-239	Amiante	X			Sanders (1975)
<i>Rat</i>	Pu-239	Be		X		Finch et al. (1991)

4- Beyond a descriptive approach

Ecotoxicity: Interactions between stable and radioactive substances

Conclusions from WP4 (2015)

- general pattern of the combined effects of UxCd and yxCd across species
 - needed to examine a wide range of endpoints, timepoints and exposure conditions (bioavailability, bioaccumulation)

➔ consistent with the use of SSDs derivations?

- some interactions were explained (mainly antagonistic) by bioavailability and toxicokinetics (rarely toxicodynamic processes)
 - needs robust exposure characterization (speciation, bioavailability & bioacc.)
 - toxicokinetic/dynamics (eg. DEBtox) helps for an overall conclusion

➔ no evidence of large (but some) synergism: additivity assumption sufficient?

- we used huge, but mainly 'bioassay-derived' experimental designs
 - robust but low sensitive species, endpoints, timepoints (=high exposure needed)
- need more ecological and biological realism to identify interactions!

➔ interactions & mode of action at cell/molecular level?

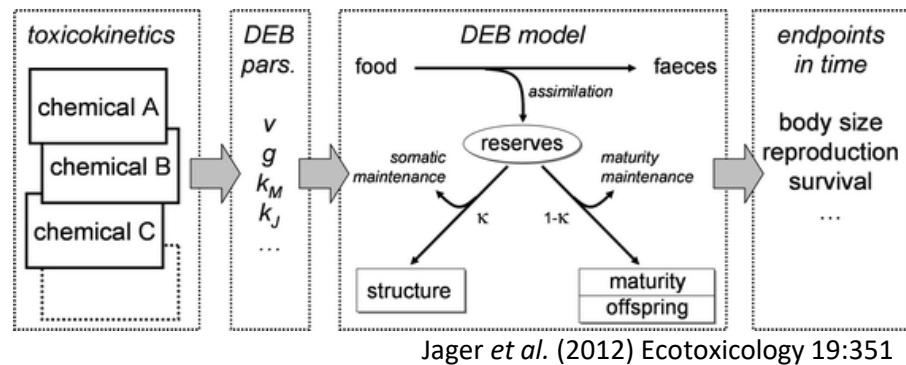
➔ interactions at low doses and effect?

➔ interactions at higher level (populations) and long-term (trans-generational)?

4- Beyond a descriptive approach

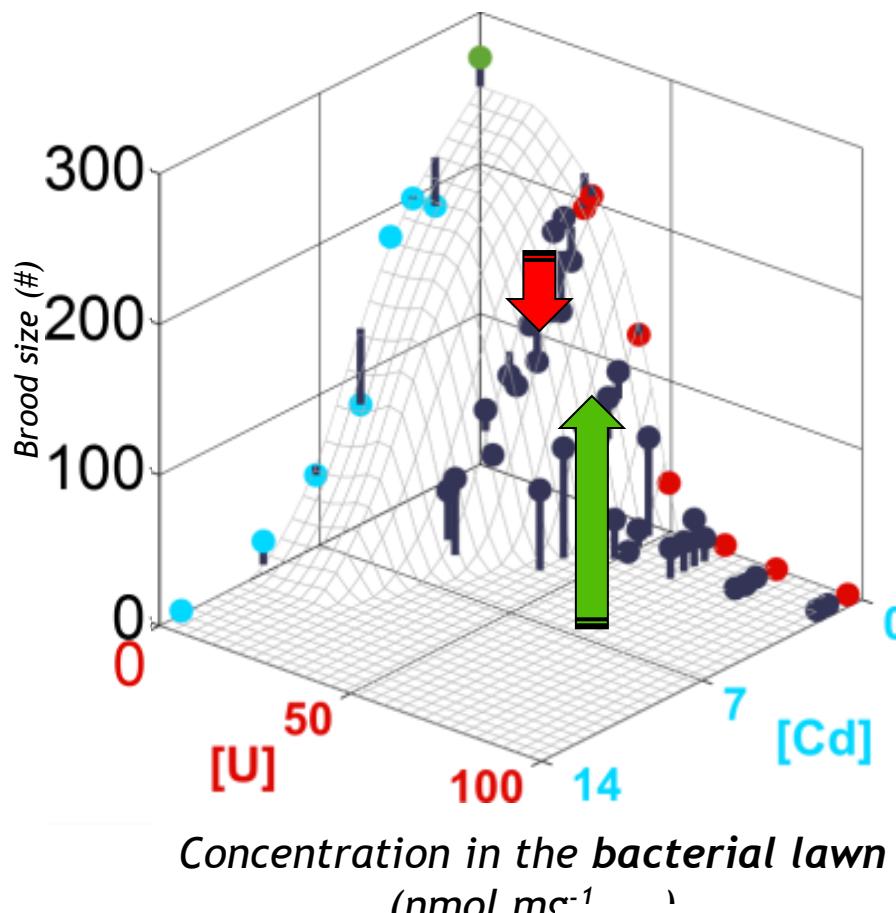
Ecotoxicity: Interactions between stable and radioactive substances

DEBtox: Toxicokinetic & dynamic model



ex. Toxicity of U+Cd to *C elegans*

DEBtox of individual substance (U and Cd)
+ use of additivity model (ex. IA, 11-day)

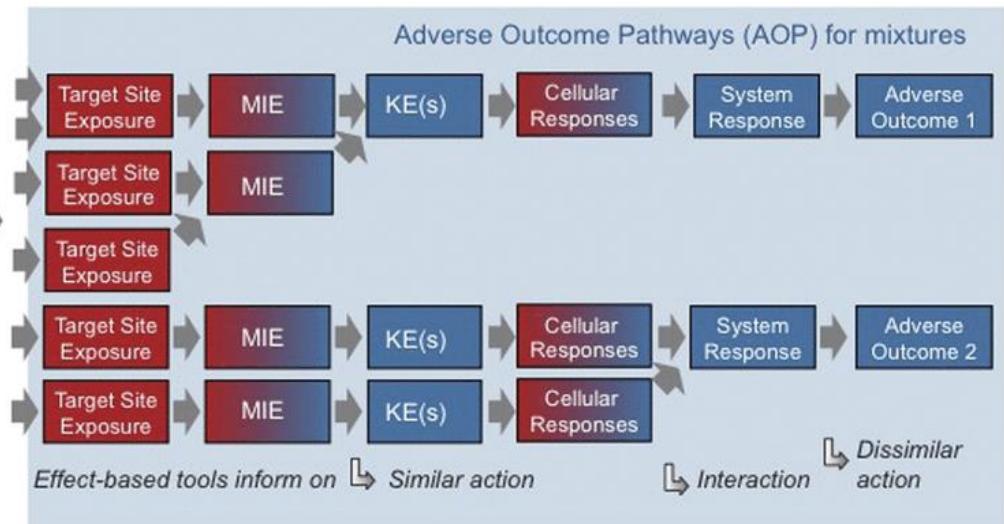


4- Beyond a descriptive approach

Strong potential of AOP approaches

'Extension' of the DEBtox approach

→ From linear AOPs to
AOP networks



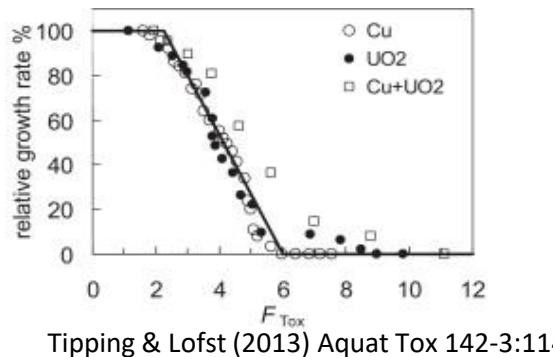
Usefulness :

- **Low doses:** undetectable individual exposure / effect levels. combined exposure with detectable answers
- Identification of the **main risk factors** in the exposome
- **Cause and effect relationships:** Similar AOs may be triggered by different MIEs (affect jointly the same KE)
limitation: complex interactions (eg, complex role of low-dose mixtures in carcinogenesis) - Goodson et al., 2015)

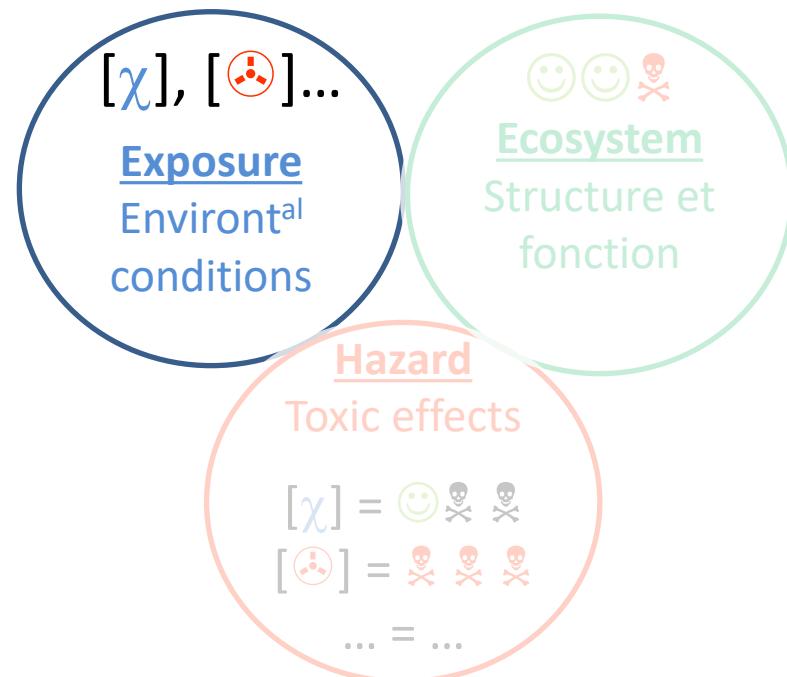
Conclusion / Take home message (1/3)

- **Ecosystem Exposure**

- Generalization of the BLM approach to consider interactions in the medium (speciation) and bioavailability (eg WHAM-Ftox)



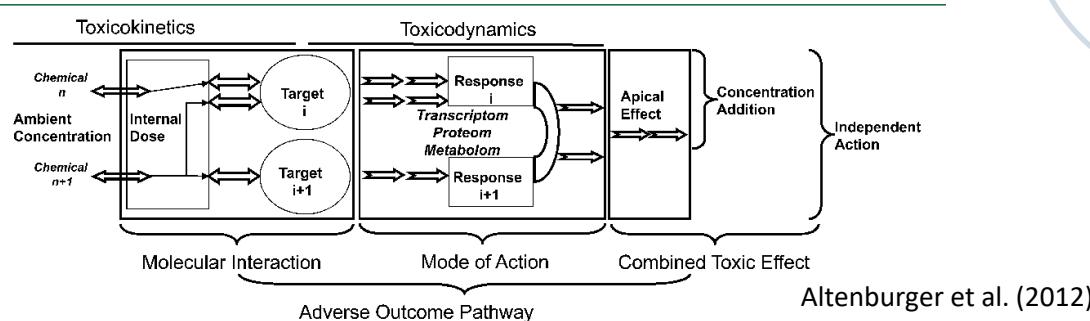
- Co-exposure with organic substances?
- integration of other stress factors (temperature, biotope changes, UVs...)?



Conclusion / Take home message (2/3)

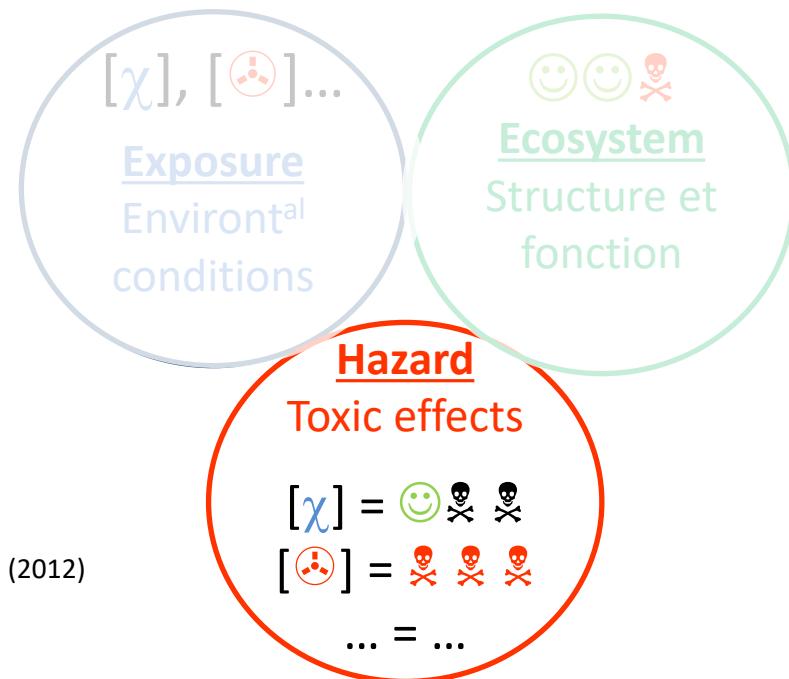
▪ Ecotoxicity

- Explicit modeling coupling toxicoco-kinetic and dynamic (eg DEBtox, AOP) to generalize the observations (mixtures, species, exposure)



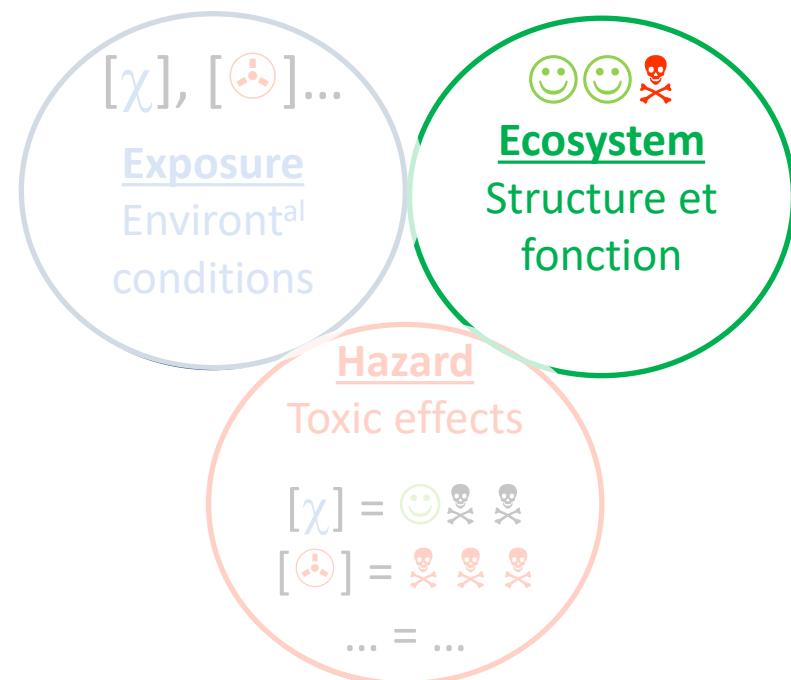
▪ Experimental evidences of interactions

Synergisms and/or Antagonisms may be explained (at least partially) with mechanistic approaches interactions at different steps from exposure, initial events up to Adverse Outcomes



Conclusion / Take home message (3/3)

- **Ecological Risk Analysis** needs a better prognostic power.
 - May be (partly) solved by a **proper exposure characterisation** (eg. taking into account bioavailability,... exposome) and description of *interactions* (TDTK models such as DEBtox; AOP)
 - **msPAF** (based on SSD) seems to be satisfactory to describe the response of communities, by taking into account bioavailability, modes of action (CA / IA) and lack of specific interaction
 - However, the **real ecological status** under mixture stress **needs more field survey** of ecologically-relevant endpoints (communities structure and function)



Combined exposures to stable and radioactive substances for ecological risk assessment

Rodolphe GILBIN, Laureline FÉVRIER, Catherine LECOMTE, Frédéric ALONZO, Béatrice GAGNAIRE, Jean-Marc BONZOM, Olivier ARMANT, Karine BEAUGELIN, Hugo LEPAGE, Jacqueline GARNIER-LAPLACE

Rodolphe GILBIN

rodolphe.gilbin@irsn.fr

Research Department on Transfers and Effects of
Radionuclides on ecosystems
IRSN / PSE-ENV / SRTE
Cadarache, France

