The four Poles of the compass to manage the challenges without losing sight of the north!





1st pole : Identify the intrinsic hazards

1st pôle : Identify the hazard(s)

- The hazard describes the undesirable properties intrinsically associated with the nature of an element: microbes which kill, a salt which corrosives, speed is dangerous, dioxin which is toxic;
- For biological agents this dangerous nature <u>is</u> not linked to the intensity of the exposure to this effect.
- For physical or chemical agents, it is a combination of the <u>dose</u> and <u>the duration</u> or frequency of exposure to this adverse effect.
- Technical and (eco)toxicological tests make it possible to determine for most chemical, biological and physical agents a "<u>no effect</u> <u>level</u>".





About acceptable limits and safety factors

- To define an acceptable exposure limit value for a dangerous biological agent and certain physical agents (eg radiation), the guidelines generally recommend safety distances and isolation measures;
- For chemicals, a safety margin is always included to take into account, for example, the possible differences between observations on animals and the reality and diversity of human exposures;
- These are generally between 100 and 1000, depending in particular on:
 - the type of effect : irreversible or not;

- the <u>degree of knowledge</u> of the hazardous properties: *number and types*

of test studies carried out, etc..



About the levels of safety factors

 By comparison, on some highways, the following warning signs appear about safety distances between two vehicles:



« One mark : danger ; two marks : safety » The safety factor applied there is 2.

 Given that a mark length is 25 m, should a safety factor of 100 be applied, <u>what would be the distance between two vehicles</u>?



Among major health hazards that appeared in the 20th century

- Influenza, leprosy, malaria, polio, tuberculosis and other "age-old" diseases are now often fought with vaccines but continue to cause millions of victims;
- Among the dangerous viral and bacteriological agents presenting major risks which appeared in the 20th century are those of AIDS, Ebola, Chykungunya;
- Others also affect plants, such as the Xylella Fastidiosa bacteria that destroy olive trees or Fomes annusus for pine;

The challenge is always to identify the *hazard*(s) for everyone, measure the *risks* and take the appropriate *safety* measures

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Methods for identifying infectious hazards

- <u>Traditional methods</u> of identifying a dangerous property for a <u>species of an infectious agent</u> rely on the multiplication in the laboratory of the potentially pathogenic agent in order to be able to identify it;
- These methods, such as giving birth to vaccines, are timeconsuming and sometimes very expensive;
- Genetic engineering now makes it possible to identify many pathogens in a faster and more reliable way;
- Other methods are based on the identification of <u>antibodies</u> produced in reaction to the pathogenic effect.



Multidisciplinary toxicology: to identify the dangerous properties of chemical substances

This implies, to collect data rom the multiple disciplines of human and veterinary medicine:

- <u>Pharmacology</u> including pharmacokinetics, (<u>histo)pathology</u>, <u>hematology</u>,...;
- <u>analytical toxicology</u>, "in vivo" and "in vitro"; <u>experimental toxicology and clinical toxicology</u>: toxicology of acute and repeated dose exposures: subacute, subchronic, chronic, multigenerational;
- clinical chemistry, cell toxicology, genetics (and "omics");
- <u>immunotoxicity</u> and toxicology of <u>reproduction</u>, <u>carcinogenicity</u>;
- <u>Mechanistic</u> toxicology and <u>epidemiological</u> toxicology;
- **Biostatistics** and mathematical modeling.
- ...



For hazards towards the environment, a complementary science: *ecotoxicology*

- For "generic" substances, the ecotoxicity is tested in the laboratory and in a reproducible manner on representative type <u>organisms of each trophic level</u>:
 - For plants: micro-algae (blue green algae, diatoms);
 - For crustaceans and molluscs: duckweed (daphnia);
 - For mammals: fish: (rainbow trout, exotic aquarium fish);
 - For the microorganisms in charge of biodegradation:
 - specific bacteria or sample of wastewater treatment plant.

